

Stroke rehabilitation in adults (update)

[N] Evidence reviews for the clinical and cost-effectiveness of music therapy for adults after a stroke

NICE guideline NG236

Evidence reviews underpinning recommendations for research in the NICE guideline

October 2023

Final

*These evidence reviews were developed
by NICE*

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1 Music therapy

1.1 Review question

In people after stroke, what is the clinical and cost effectiveness of music therapy to improve mood and activities of daily living?

1.1.1 Introduction

Music activates a wide range of regions within the brain including networks involved in speech, motor function and cognition. Music therapy aims to facilitate recovery mechanisms in the brain to enhance rehabilitation and overall improvements.

Usually trained music therapists deliver it with an individual or in a group. Music is used in a number of different ways; for example listening to music, actively participating in music or writing and composing music.

1.1.2 Summary of the protocol

Table 1: PICO characteristics of review question

| | |
|----------------------|--|
| Population | <p>Inclusion:</p> <ul style="list-style-type: none">Adults (age ≥ 16 years) who have had a first stroke or recurrent stroke (including people after a subarachnoid haemorrhage) <p>Exclusion:</p> <ul style="list-style-type: none">Children (age < 16 years)People who have had a transient ischaemic attack |
| Interventions | <ul style="list-style-type: none">Neurologic music therapy delivered by trained music therapistsMusic therapy delivered by trained music therapistsMusic interventions delivered by healthcare professionalsMusic interventions delivered by non-healthcare professionals <p>The interventions will be analysed as separate stratifications.</p> |
| Comparisons | <ul style="list-style-type: none">Compared to each otherPassive music listening (for example: music played in the background)Placebo music therapyNo treatment <p>Each comparator will be analysed in separate stratifications by different types of comparators.</p> |
| Outcomes | <p>All outcomes are considered equally important for decision making and therefore have all been rated as critical:</p> <p>At time period</p> <ul style="list-style-type: none">< 6 months≥ 6 months <ul style="list-style-type: none">Person/participant generic health-related quality of life (continuous outcomes will be prioritised [validated measures])Carer generic health-related quality of life (continuous outcomes will be prioritised [validated measures]) |

| | |
|---------------------|---|
| | <ul style="list-style-type: none">• Activities of daily living (continuous outcomes will be prioritised)• Psychological distress (continuous outcomes will be prioritised)<ul style="list-style-type: none">○ Depression○ Anxiety○ Distress• Stroke-specific Patient-Reported Outcome Measures (continuous outcomes will be prioritised)• Wellbeing scores (continuous outcomes will be prioritised)• Participation in leisure activities/social groups scores (continuous outcomes will be prioritised)• Withdrawal due to adverse events (dichotomous outcome) |
| Study design | <ul style="list-style-type: none">• Systematic reviews of RCTs• Parallel RCTs• Cluster randomised crossover trials (unit of randomisation = unit [for example: hospital, community location]) <p>If there is insufficient randomised trial evidence, non-randomised studies (prospective and retrospective cohort trials) will be considered after discussion with the committee.</p> |

For full details see the review protocol in Appendix A.

1.1.3 Methods and process

This evidence review was developed using the methods and process described in [Developing NICE guidelines: the manual](#). Methods specific to this review question are described in the review protocol in Appendix A and the methods document.

Declarations of interest were recorded according to [NICE's conflicts of interest policy](#).

1.1.4 Effectiveness evidence

1.1.4.1 Included studies

Twenty one randomised controlled trials (twenty three papers) were included in the review;^{2-7, 9-12, 15-27} these are summarised in Table 2 below. Evidence from these studies is summarised in the clinical evidence summary below (Table 3).

The studies included the following comparisons:

- Neurologic music therapy delivered by trained music therapists compared to no treatment^{17, 26}
- Music therapy delivered by trained music therapists compared to no treatment^{3, 4, 9, 10, 15, 16, 19}
- Music intervention delivered by healthcare professionals compared to passive music listening²
- Music intervention delivered by healthcare professionals compared to placebo music therapy²
- Music intervention delivered by healthcare professionals compared to no treatment^{5-7, 11, 12, 18, 22-25, 27}
- Music intervention delivered by non-healthcare professionals compared to no treatment^{20, 21}

While comparisons were available for each intervention category compared to no treatment, there was limited or no evidence comparing therapies to:

- Each other
- Passive music listening
- Placebo music therapy

See also the study selection flow chart in Appendix C, study evidence tables in Appendix D, forest plots in Appendix E and GRADE tables in Appendix F.

1.1.4.1.1 Types of intervention

The types of interventions delivered in the studies varied. They included:

- Rhythmic auditory cueing^{3, 6, 7, 12, 22, 23, 25}
- Interventions where musical instruments are played (including clinical improvisation)^{4, 5, 10, 16, 19}
- Receptive interventions in which participants listen to music^{2, 11, 17, 27}
- Singing a music-based voice interventions^{20, 26}
- Sonofication¹⁸
- Combinations of the above^{9, 15}

Where heterogeneity was present there was an insufficient number of studies in each group representing different types of intervention, and so the heterogeneity was not resolved by subgroup analysis by these groups.

1.1.4.2 Excluded studies

One Cochrane review was identified that included relevant information for this review, Magee 2017¹³. This review was excluded as it included people with conditions other than stroke (including any acquired brain injury). While the review was excluded, the references were checked for studies relevant for this review.

A significant number of studies were excluded as they did not report outcomes relevant to the protocol, the majority of these reporting outcomes relevant to individual impairments (for example: motor function, communication). These outcomes were considered of a lower priority than functional outcomes (for example: activities of daily living) and were considered through other outcomes (for example: health-related quality of life).

See the excluded studies list in Appendix J.

1.1.5 Summary of studies included in the effectiveness evidence

1.1.5.1 Neurologic music therapy delivered by trained music therapists

Table 2: Summary of studies included in the evidence review

| Study | Intervention and comparison | Population | Outcomes | Comments |
|---------------------------------------|---|---|--|---|
| Pocwierz-Marciniak 2017 ¹⁷ | <p>Neurologic music therapy delivered by trained music therapists (n=30) One-to-one sessions using mainly a receptive approach based on cognitive music therapy and guided imagery and music. Twice a week for 10 meetings.</p> <p>Group/individual sessions: Individual Hospital/community : Hospital Type of intervention: Receptive interventions in which participants listen to music</p> <p>No treatment (n=31) No listening materials</p> <p>Concomitant therapy All people were undergoing an inpatient neurological rehabilitation in hospital and receiving standard care, including physiotherapy, ergotherapy,</p> | <p>Adults who have had a first stroke Mean age (range): 64 (44 to 84) years N=61</p> <p>Time after stroke: Acute (72 days - 7 days) – not explicitly stated Severity: Not stated/unclear</p> <p>Type of stroke: Ischaemic = 49 Haemorrhagic = 12</p> | <p>Person/participant generic health-related quality of life at <6 months Stroke-specific Patient-Reported Outcome Measures at <6 months</p> | <p>Setting: Inpatient neurological rehabilitation in hospital in Gdynia (Northern Poland).</p> <p>Funding: No additional information.</p> |

| Study | Intervention and comparison | Population | Outcomes | Comments |
|--------------------------|---|---|---|--|
| Zhang 2021 ²⁶ | <p>psychological diagnosis and maintenance psychotherapy.</p> <p>Neurologic music therapy delivered by trained music therapists (n=20) Melodic intonation therapy for 30 minutes per session, five sessions a week for 8 weeks.</p> <p>Group/individual sessions: Individual Hospital/community : Hospital Type of intervention: Singing and music-based voice interventions</p> <p>No treatment (n=20) Usual care by a speech therapist for the same amount of time.</p> <p>Concomitant therapy All patients underwent routine treatment during the study period, including taking medication and other care and support.</p> | <p>Adults who have had a first stroke Mean age (SD): 53.5 (10.0) years N=61</p> <p>Time after stroke: 2.27 (1.59) months Severity: Mixed</p> <p>Type of stroke: Ischaemic = 24 Haemorrhagic = 16</p> | <p>Psychological distress (depression) at <6 months Psychological distress (anxiety) at <6 months</p> | <p>Setting: Inpatient care in China.</p> <p>Funding: Funded by a governmental/non-for-profit research grant.</p> |

1.1.5.2 Music therapy delivered by trained music therapists

Table 3: Summary of studies included in the evidence review

| Study | Intervention and comparison | Population | Outcomes | Comments |
|-----------------------|---|---|--|---|
| Cha 2014 ³ | <p>Music therapy delivered by trained music therapists (n=10) Intensive gait training with rhythmic auditory</p> | <p>Adults who have had a first or recurrent stroke Mean age (SD): 61.4 (13.1) years N = 20</p> | <p>Stroke-specific Patient-Reported Outcome Measures at <6 months</p> | <p>Setting: Outpatient follow up in the Republic of Korea</p> <p>Funding: This study was supported by S University (2013)</p> |

| Study | Intervention and comparison | Population | Outcomes | Comments |
|---------------------------|--|---|---|---|
| | <p>stimulation for 30 minutes, five days per week.</p> <p>Group/individual sessions: Individual Hospital/community : Hospital Type of intervention: Rhythmic auditory cueing</p> <p>No treatment (n=10) Intensive gait training only.</p> <p>Concomitant therapy All participants received general physical therapy, including Bobath approach and proprioceptive neuromuscular facilitation for 30 minutes per day, five days per week.</p> | <p>Time after stroke (SD): 14.6 (5.5) years Severity: Not stated/unclear</p> <p>Type of stroke: Not stated/unclear</p> | <p>Withdrawal due to adverse events at <6 months</p> | |
| Fujioka 2018 ⁴ | <p>Music therapy delivered by trained music therapists (n=14) Music-supported therapy added to support motor, cognitive and psychosocial functions combined with an existing conventional physical training programme (GRASP)</p> <p>Group/individual sessions: Individual Hospital/community : Hospital Type of intervention: Interventions where musical instruments are played (including</p> | <p>Adults who have had a first or recurrent stroke Mean age (SD): 59.4 (11.5) years N = 29</p> <p>Time after stroke (SD): 5.4 (6.7) years Severity: Not stated/unclear</p> <p>Type of stroke: Not stated/unclear</p> | <p>Psychological distress (depression and other) at <6 months Stroke-specific Patient-Reported Outcome Measures at <6 months Withdrawal due to adverse events at <6 months</p> | <p>Setting: Outpatient follow up in Canada</p> <p>Funding: This research was supported by the Canadian Institutes of Health Research and the Heart and Stroke Foundation Ontario.</p> |

| Study | Intervention and comparison | Population | Outcomes | Comments |
|-----------------------|---|---|---|---|
| | <p>clinical improvisation)</p> <p>No treatment (n=15) Conventional physical training programme (GRASP) without music</p> <p>Concomitant therapy No additional information</p> | | | |
| Jun 2013 ⁹ | <p>Music therapy delivered by trained music therapists (n=20) Music movement therapy including 20 minutes of preparatory activities, 30 minutes of main activities and 10 minutes of finishing activities delivered 3 times per week for 8 weeks.</p> <p>Group/individual sessions: Group Hospital/community : Hospital Type of intervention: Any combination of the above (interventions where musical instruments are played and receptive interventions in which participants listen to music)</p> <p>No treatment (n=20) Routine treatment only</p> <p>Concomitant therapy</p> | <p>Adults who have had a first or recurrent stroke Mean age (SD): 57.9 (13.9) years N = 40</p> <p>Time after stroke: Acute (72 hours – 7 days) – not explicitly stated Severity: Not stated/unclear</p> <p>Type of stroke: Infarction = 25 Haemorrhage = 5</p> | <p>Activities of daily living at <6 months Psychological distress (depression) at <6 months</p> | <p>Quasi-experimental trial but states it was randomised. Has been included but downgraded for risk of selection bias.</p> <p>Setting: Hospital inpatients in South Korea</p> <p>Funding: This work was supported by Dong-eui University Foundation Grant (2011).</p> |

| Study | Intervention and comparison | Population | Outcomes | Comments |
|--------------------------|---|--|--|--|
| | No additional information | | | |
| Kim 2011 ¹⁰ | <p>Music therapy delivered by trained music therapists (n=9) Music therapy program following a standard 40-minute format carried out in accordance with the physical strength and individual characteristics of patients for 4 weeks</p> <p>Group/individual sessions: Not stated/unclear Hospital/community : Hospital Type of intervention: Interventions where music instruments are played (including clinical improvisation)</p> <p>No treatment (n=9) No music intervention</p> <p>Concomitant therapy All people received comprehensive rehabilitation treatment including physiotherapy, occupational therapy or speech therapy. All people received regular counselling by a licensed psychotherapist.</p> | <p>Adults who have had a first or recurrent stroke Mean age (SD): 49.5 (12.8) years N = 18</p> <p>Time after stroke: Subacute (7 days – 6 months) – not explicitly stated Severity: Not stated/unclear</p> <p>Type of stroke: Not stated/unclear</p> | <p>Psychological distress (depression and anxiety) at <6 months Withdrawal due to adverse events at <6 months</p> | <p>Setting: Inpatient setting in South Korea</p> <p>Funding: The authors have no financial conflicts of interest</p> |
| Nayak 2000 ¹⁵ | <p>Music therapy delivered by trained music therapists (n=10) Music therapy 3 treatments per</p> | <p>Adults with acute brain injury (including people after a first or recurrent stroke)</p> | <p>Psychological distress (depression) at <6 months Participation in leisure</p> | <p>Setting: Inpatient facility at the Kessler Institute for Rehabilitation in the United States of America</p> |

| Study | Intervention and comparison | Population | Outcomes | Comments |
|----------------------------|--|---|--|---|
| | <p>week for up to 10 treatments (treatment two or three times per week). A variety of procedures were used based on the needs of the person.</p> <p>Group/individual sessions: Group Hospital/community : Hospital Type of intervention: Any combination of the above (combination of musical instruments, singing and songwriting)</p> <p>No treatment (n=8) Standard rehabilitation.</p> <p>Concomitant therapy Conventional rehabilitation was provided to both groups.</p> | <p>Mean age (SD): 59.9 (16.3) years N = 18</p> <p>Time after stroke: Acute (72 hours – 7 days) – not explicitly stated Severity: Not stated/unclear</p> <p>Type of stroke: Not stated/unclear</p> | <p>activities/social groups at <6 months Withdrawal due to adverse events at <6 months</p> | <p>Funding: Support for this research was provided by National Institutes of Health Grant U24-HD32994</p> <p>The proportion of participants after stroke was unclear. Due to this the study was included but downgraded for population indirectness.</p> |
| Palumbo 2022 ¹⁶ | <p>Music therapy delivered by trained music therapists (n=15) Music therapy integrated with upper limb exercise using liver, interactive music making using a Nordoff-Robbins approach. For 45 minutes, twice a week for 6 weeks.</p> <p>Group/individual sessions: Group Hospital/community : Hospital Type of intervention: Interventions where</p> | <p>Adults with acute brain injury (including people after a first or recurrent stroke) Mean age (SD): 61.5 (11.1) years N = 30</p> <p>Time after stroke: Subacute (7 days - 6 months) Severity: Not stated/unclear</p> <p>Type of stroke: Not stated/unclear</p> | <p>Psychological distress (depression) at <6 months Stroke-specific Patient-Reported Outcome Measures at <6 months</p> | <p>Setting: Outpatient follow-up in the United States of America.</p> <p>Funding: Supported by New York University Clinical Translational Science Award UL1TR000038 from the National Center for Advancing Translational Sciences (NCATS) and U54NS081765, National Institutes of Health, and in part by grants from the GRAMMY Foundation and the John and Jennifer Clay Foundation.</p> |

| Study | Intervention and comparison | Population | Outcomes | Comments |
|---------------------------------|--|--|--|--|
| | <p>musical instruments are played (including clinical improvisation)</p> <p>No treatment (n=15) Home exercise programme for a matched amount of time.</p> <p>Concomitant therapy No additional information.</p> | | | <p>The control group intervention may have received exercise that was of greater intensity than that received by the intervention group and so was downgraded for comparator indirectness.</p> |
| <p>Raglio 2017¹⁹</p> | <p>Music therapy delivered by trained music therapists (n=19) Relational active music therapy approach using rhythmical-melodic instrumentation. 20 sessions lasting 30 minutes each, three weekly.</p> <p>Group/individual sessions: Individual Hospital/community : Hospital</p> <p>Type of intervention: Interventions where musical instruments are played (including clinical improvisation)</p> <p>No treatment (n=19) No music intervention</p> <p>Concomitant therapy The standard of care treatment consisted of daily sessions of physiotherapy (passive/assisted</p> | <p>Adults who have had a first or recurrent stroke Mean age (SD): 63.6 (13.1) years N = 38</p> <p>Time after stroke (median [IQR]): Intervention – 29.5 (31.8) days Control – 34.5 (38.3) days Severity (NIHSS – mean [SD]): 4.8 (2.3)</p> <p>Type of stroke: Ischaemic = 35 Haemorrhagic = 3</p> | <p>Patient/participant generic health-related quality of life at <6 months Activities of daily living at <6 months Psychological distress (depression and anxiety) at <6 months</p> | <p>Setting: Inpatients in Italy</p> <p>Funding: The authors received no financial support for the research, authorship, and/or publication of this article.</p> |

| Study | Intervention and comparison | Population | Outcomes | Comments |
|-------|---|------------|----------|----------|
| | active mobilization and neurorehabilitative techniques of paretic upper limbs, coordination and balance exercises, and gait training) and occupational therapy (exercises improving fine motor skills and recovering activities of daily living). | | | |

1.1.5.3 Music interventions delivered by healthcare professionals

Table 4: Summary of studies included in the evidence review

| Study | Intervention and comparison | Population | Outcomes | Comments |
|--------------------------|---|--|--|---|
| Baylan 2016 ² | <p>Music interventions delivered by healthcare professionals (n=23)</p> <p>Mindful music listening. People were given an iPod Nano (7th Generation, Apple Inc.) and asked to listen to their material daily on their own for at least an hour during the intervention phase (target 56 hours over 8 consecutive weeks).</p> <p>Group/individual sessions: Individual Hospital/community : Hospital</p> <p>Type of intervention: Receptive interventions in which participants listen to music</p> <p>Passive music listening (n=24)</p> | <p>Adults who have had a first or recurrent stroke</p> <p>Mean age (SD): 64.0 (11.8) years N = 72</p> <p>Time after stroke (median): 19 days Severity: Not stated/unclear</p> <p>Type of stroke: Cortical = 46 Subcortical = 26</p> | <p>Psychological distress (depression and anxiety) at <6 months</p> <p>Participation in leisure activities/social groups scores at <6 months</p> <p>Withdrawal due to adverse events at <6 months</p> | <p>Setting: Acute Stroke Units within NHS Greater Glasgow and Clyde, United Kingdom</p> <p>Funding: This work was supported by the Dunhill Medical Trust, grant R432/0214. Additional support from Scottish Executive Chief Scientist Office (TQ/BC), Stroke Association (TQ) and The Dr Mortimer and Theresa Sackler Foundation (BC/SB).</p> |

| Study | Intervention and comparison | Population | Outcomes | Comments |
|--------------------------------------|--|--|--|--|
| | <p>Listening to music without any mindfulness components.</p> <p>Placebo music therapy (n=25) Audiobook listening instead of music listening.</p> <p>Concomitant therapy No additional information</p> | | | |
| <p>Grau-Sanchez 2018⁵</p> | <p>Music interventions delivered by healthcare professionals (n=20) Music supported therapy delivered by an occupational therapists and physiotherapist. 20 individual sessions (5 sessions per week, 30 minutes each).</p> <p>Group/individual sessions: Individual Hospital/community : Hospital Type of intervention: Interventions where musical instruments are played (including clinical improvisation)</p> <p>No treatment (n=20) Additional individual training of the upper extremity. 20 individual sessions (5 sessions per week, 30 minutes each) consisting of passive mobilization,</p> | <p>Adults who have had a first or recurrent stroke Mean age (range): 61.3 (45-74) years N = 40</p> <p>Time after stroke (range): 65.4 (28-162) days Severity (NIHSS – mean [range]): 5.4 (2-14)</p> <p>Type of stroke: Not stated/unclear</p> | <p>Patient/participant generic health-related quality of life at <6 months Psychological distress (depression and other) at <6 months Stroke-specific Patient-Reported Outcome Measures at <6 months Withdrawal due to adverse events at <6 months</p> | <p>Setting: Outpatient follow up. Rehabilitation delivered in hospital setting in Spain.</p> <p>Funding: This work was supported by the Spanish Government (Ministerio de Economía y Competitividad, PSI2015-69178-P, Fondo Europeo de Desarrollo Regional (FEDER)).</p> |

| Study | Intervention and comparison | Population | Outcomes | Comments |
|------------------------|---|---|--|---|
| | <p>stretch and progressive resistance exercises, and task-specific training.</p> <p>Concomitant therapy Both groups received an outpatient rehabilitation program that consisted of two 1 hour group sessions of occupational therapy and physiotherapy a day (5 days per week, 10 hours in total per week).</p> | | | |
| Hill 2011 ⁶ | <p>Music interventions delivered by healthcare professionals (n=6) Interactive metronome intervention delivered by an occupational therapist. Completing the same exercises as the occupational therapy only group, with an additional 30 minute interactive metronome session. 1 hour treatment, 3 times a week for 10 weeks.</p> <p>Group/individual sessions: Individual Hospital/community : Hospital Type of intervention: Rhythmic auditory cueing</p> | <p>Adults who have had a first or recurrent stroke Mean age (SD): 60.0 (9.2) years N = 10</p> <p>Time after stroke (SD): 3.3 (2.3) years Severity: Not stated/unclear</p> <p>Type of stroke: Not stated/unclear.</p> | <p>Activities of daily living at <6 months Stroke-specific Patient-Reported Outcome Measures at <6 months Withdrawal due to adverse events at <6 months</p> | <p>Setting: Outpatient follow up. Delivered in hospital setting in the United States of America.</p> <p>Funding: Interactive Metronome equipment and software provided for the study.</p> |

| Study | Intervention and comparison | Population | Outcomes | Comments |
|-------------------------|--|--|---|--|
| | <p>No treatment (n=4) Usual occupational therapy only. 1 hour treatment, 3 times a week for 10 weeks.</p> <p>Concomitant therapy Usual occupational therapy including prefunctional activities, functional activities and COPM (Canadian Occupational Performance Measure) tasks.</p> | | | |
| Jeong 2007 ⁷ | <p>Music interventions delivered by healthcare professionals (n=16) Rhythmic auditory stimulation music-movement program for 2 hours/week for 8 weeks</p> <p>Group/individual sessions: Group Hospital/community : Community Type of intervention: Rhythmic auditory cueing</p> <p>No treatment (n=17) Received referral information about available usual care services (available to both groups)</p> <p>Concomitant therapy No additional information.</p> | <p>Adults who have had a first or recurrent stroke Mean age (SD): 60.2 (8.0) years N = 33</p> <p>Time after stroke (SD): 6.4 (5.0) years Severity: Not stated/unclear</p> <p>Type of stroke: Not stated/unclear</p> | <p>Psychological distress (other) at <6 months Stroke-specific Patient-Reported Outcome Measures at <6 months</p> | <p>Setting: A neighbourhood community health center located in the metropolitan area in Seoul, South Korea.</p> <p>Funding: This study was supported by the BK21 project (Grant No. 0522-20010002), the Korea Science and Engineering Foundation (Grant No. R04-2001-000-00197-0), and the Research Institute of Nursing Science at Seoul National University.</p> |
| Lin 2017 ¹¹ | <p>Music interventions delivered by</p> | <p>Adults who have had a first or recurrent stroke</p> | <p>Activities of daily living at <6 months</p> | <p>Setting: Inpatients and China</p> |

| Study | Intervention and comparison | Population | Outcomes | Comments |
|-------|--|---|---|---|
| | <p>healthcare professionals (n=30) Acupuncture with a five phase music therapy intervention. Music therapy was administered twice daily, once in the morning and once in the afternoon for 20 minutes per session. All treatments were administered in a 5 day cycle, for three continuous cycles with an interval between two cycles of 1 day (17 days in total).</p> <p>Group/individual sessions: Individual Hospital/community : Hospital Type of intervention: Receptive interventions in which participants listen to music</p> <p>No treatment (n=30) Acupuncture needling only for the same timings.</p> <p>A third arm was reported (n=32) that received 50mg sertraline hydrochloride only. This group was not included in the analysis as it did not fulfil the criteria of the protocol.</p> <p>Concomitant therapy No additional information</p> | <p>Mean age (SD): 70.3 (11.3) years N = 92</p> <p>Time after stroke: Subacute (7 days – 6 months) – not explicitly stated Severity: Not stated/unclear</p> <p>Type of stroke: No additional information</p> | <p>Psychological distress (depression) at <6 months Withdrawal due to adverse events at <6 months</p> | <p>Funding: State Administration of Traditional Chinese Medicine of the People's Republic of China, State Clinical Research Base of Traditional Chinese Medicine, The Second Batch of Professional Skill Scientific and Research Special Project (No. JDZX2015127); Jiangsu Natural Science Foundation Youth Project (No. BK20171070); Nanjing Scientific Development Planned Project (No. 201402057)</p> |

| Study | Intervention and comparison | Population | Outcomes | Comments |
|---------------------------|--|--|---|---|
| Luft 2004 ¹² | <p>Music interventions delivered by healthcare professionals (n=11) Bilateral arm training with rhythmic auditory cueing in hour long therapy sessions (four 5 minute movement periods with 10 minute rest periods between) 3 times per week for 6 weeks.</p> <p>Group/individual sessions: Individual Hospital/community : Hospital Type of intervention: Rhythmic auditory cueing</p> <p>No treatment (n=15) Dose matched therapeutic exercise (same timings).</p> <p>Concomitant therapy No additional information</p> | <p>Adults who have had a first or recurrent stroke Mean age (SD): 61.2 (12.9) years N = 26</p> <p>Time after stroke (median [IQR]): Intervention – 75 (37.9-84.5) months Control: 45.5 (22.6-66.3) months Severity: Not stated/unclear</p> <p>Type of stroke: Cortical = 12 Subcortical = 6 Brainstem = 3</p> | <p>Activities of daily living at <6 months</p> | <p>Setting: Outpatient rehabilitation care in hospital in the United States of America</p> <p>Funding: This study was funded by National Institutes of Health grants from the National Institute on Aging (P60AG 12583); University of Maryland Claude D. Pepper Older Americans Independence Center, National Institute on Disability and Rehabilitation Research (H133G010111); the Baltimore Department of Veterans Affairs Geriatrics Research, Education and Clinical Center (GRECC); National Institute of Neurological Disorders and Stroke 1R01 NS 24282-08; the France-Merrick Foundation; the Johns Hopkins GCRC (NCRR MO1-00052); and the Eleanor Naylor Dana Charitable Trust, Deutsche Forschungsgemeinschaft (Lu 748/2, 748/3).</p> |
| Raglio 2021 ¹⁸ | <p>Music interventions delivered by healthcare professionals (n=33) Sonofication. Synthesized sounds/musical texture and their parameters (mainly rhythm, pitch/melody, intensity/dynamics, harmony and</p> | <p>Adults who have had a first or recurrent stroke Mean age (SD): 63.6 (13.1) years N = 65</p> <p>Time after stroke (median [IQR]): Intervention: 29.5 (31.8) days Control: 34.5 (38.3) days</p> | <p>Person/participant generic health-related quality of life at <6 months Withdrawal due to adverse events at <6 months</p> | <p>Setting: Rehabilitation units (outpatient follow up) in Italy</p> <p>Funding: This work was partially supported by the "Ricerca Corrente" funding provided by the Italian Ministry of Health.</p> |

| Study | Intervention and comparison | Population | Outcomes | Comments |
|-------------------------|--|---|--|---|
| | <p>timbre) are used to represent movements characteristics. Delivered by physiotherapists or occupational therapists 5 days a week for 4 weeks, for a total of 20 sessions.</p> <p>Group/individual sessions: Individual Hospital/community : Hospital Type of intervention: Other (Sonofication)</p> <p>No treatment (n=32) No music intervention.</p> <p>Concomitant therapy All people received usual care for people with subacute strokes (such as occupational therapy, speech therapy, psychological support, lower extremity rehabilitation ect.). All people received standard motor exercises (with or without sonofication). The first phase included passive treatment (15 minutes) while the second phase included active movements (20 minutes).</p> | <p>Severity: Not stated/unclear</p> <p>Type of stroke: No additional information</p> | | |
| Tian 2020 ²² | <p>Music interventions delivered by healthcare professionals (n=16)</p> | <p>Adults who have had a first or recurrent stroke Mean age (SD): 65.5 (13.6) years N = 32</p> | <p>Activities of daily living at <6 months Withdrawal due to adverse events at <6 months</p> | <p>Setting: Inpatient in China Funding: Supported by the Project fund of Shanghai Science</p> |

| Study | Intervention and comparison | Population | Outcomes | Comments |
|---|---|--|--|---|
| | <p>Rhythmic auditory stimulation 30 minutes every day, 5 days per week for 4 weeks.</p> <p>Group/individual sessions: Individual Hospital/community : Hospital Type of intervention: Rhythmic auditory cueing</p> <p>No treatment (n=16) An additional 15 minutes regular physical therapy and 15 minutes of regular occupational therapy per day.</p> <p>Concomitant therapy Everyone received 30 minutes individualised physical therapy and 30 minutes individualised occupational therapy per day, 5 days per week for 4 weeks.</p> | <p>Time after stroke (SD): 4.4 (8.3) months Severity: Not stated/unclear</p> <p>Type of stroke: Frontal temporal lobe = 1 Frontal lobe = 1 Corona radiate = 1 Capsule externa = 1 Thalamus = 3 Basal ganglia = 17 Brainstem = 4 Paraventricular = 1 Cerebellum = 1</p> | | <p>and technology commission. The project number was 18411962300</p> |
| <p>van Delden 2009²³</p> <p>ULTRA-Stroke</p> <p>Subsidiary studies: Van Delden 2013²⁴</p> | <p>Music interventions delivered by healthcare professionals (n=19)</p> <p>Modified bilateral arm training rhythmic auditory cueing group applied by physiotherapists and/or occupational therapists working at the rehabilitation centre. Treatment was given as 60 minute sessions, 3 days a week for 6 weeks.</p> | <p>Adults who have had a first or recurrent stroke Mean age (SD): 59.8 (11.7) years N = 38</p> <p>Time after stroke (SD): 9.5 (6.2) weeks Severity: Not stated/unclear</p> <p>Type of stroke: No additional information</p> | <p>Stroke-specific Patient-Reported Outcome Measures at <6 months</p> <p>Withdrawal due to adverse events at <6 months</p> | <p>Setting: A rehabilitation centre (outpatient follow up) in the Netherlands</p> <p>Funding: This study was funded by the Dutch Scientific College of Physiotherapy of the Royal Dutch Society for Physical Therapy.</p> |

| Study | Intervention and comparison | Population | Outcomes | Comments |
|----------------------------|---|--|---|---|
| | <p>Group/individual sessions: Individual Hospital/community : Hospital</p> <p>Type of intervention: Rhythmic auditory cueing</p> <p>No treatment (n=19) Dose matched control treatment</p> <p>Concomitant therapy No additional information</p> | | | |
| Whitall 2011 ²⁵ | <p>Music interventions delivered by healthcare professionals (n=55) Bilateral arm training with rhythmic auditory cueing for 1 hour, 3 times a week for 6 weeks (for a total of 18 sessions)</p> <p>Group/individual sessions: Individual Hospital/community : Hospital</p> <p>Type of intervention: Rhythmic auditory cueing</p> <p>No treatment (n=56) Dose matched therapeutic exercises</p> <p>Concomitant therapy No additional information</p> | <p>Adults who have had a first or recurrent stroke Mean age (SD): 58.7 (11.3) years N = 111</p> <p>Time after stroke (SD): 4.3 (4.7) years Severity: Not stated/unclear</p> <p>Type of stroke: Brainstem = 6 Cerebellar = 2 Cortex = 39 Multiple = 3 Subcortical = 19</p> | <p>Stroke-specific Patient-Reported Outcome Measures at <6 months (at 6 weeks only) Withdrawal due to adverse events at <6 months (at 4 months)</p> | <p>Setting: Outpatient follow up in the United States of America</p> <p>Funding: P60AG12583; PI AG, NIDDR H H133G010111, the Baltimore Veterans Administration Geriatrics Research, Education and Clinical Center (GRECC). Andreas Luft was supported by DFG SFB 550, C 12.</p> |
| Zhao 2022 ²⁷ | <p>Music interventions delivered by</p> | <p>Adults who have had a first or recurrent stroke</p> | <p>Psychological distress</p> | <p>Setting: Inpatients in China.</p> |

| Study | Intervention and comparison | Population | Outcomes | Comments |
|-------|--|---|---------------------------|---|
| | <p>healthcare professionals (n=32) Musicokinetic therapy for 30 minutes, twice a day for 8 weeks.</p> <p>Group/individual sessions: Individual Hospital/community : Hospital Type of intervention: Receptive interventions in which participants listen to music</p> <p>No treatment (n=33) Dose matched exercise without music.</p> <p>Concomitant therapy No additional information</p> | <p>Mean age (SD): 81.14 (8.33) years N = 66</p> <p>Time after stroke: Subacute (7 days - 6 months) Severity: Not stated/unclear</p> <p>Type of stroke: Not stated/unclear</p> | (depression) at <6 months | Funding: Supported partially by the Tianjin Social Science Foundation of China (TJJX21-011) and the Developmental Program of Liberal and Social Sciences of Nankai University (ZB22BZ0109). |

1.1.5.4 Music interventions delivered by non-healthcare professionals

Table 5: Summary of studies included in the evidence review

| Study | Intervention and comparison | Population | Outcomes | Comments |
|---|--|--|---|--|
| <p>Tarrant 2021²⁰</p> <p>Subsidiary studies: Tarrant 2018²¹</p> | <p>Music interventions delivered by non-healthcare professionals (n=20) Singing for people with aphasia intervention consisting of 10 weekly sessions delivered in a community facility across three sites in the South-West of England, with each session lasting 90 minutes. Group/individual sessions: Group</p> | <p>Adults who have had a first or recurrent stroke Mean age (SD): 66.5 (10.5) years N = 41</p> <p>Time after stroke (SD): 5.1 (5.5) years Severity – Aphasia severity: Mild = 27 Moderate = 7 Severe = 7</p> <p>Type of stroke: No additional information</p> | <p>Patient/participant generic health-related quality of life at <6 months and ≥6 months Carer generic health-related quality of life at ≥6 months Stroke-specific Patient-Reported Outcome Measures at <6 months and ≥6 months Wellbeing scores at <6 months and ≥6 months Participation in leisure</p> | <p>Setting: Three community settings: a church hall, a community centre and a dedicated music venue in the United Kingdom</p> <p>Funding: The trial is funded by the Stroke Association (QQ12/TSA 2016/14). Excess treatment costs have been covered by South Devon and Torbay Clinical Commissioning Group, North East and West Devon</p> |

| Study | Intervention and comparison | Population | Outcomes | Comments |
|-------|--|------------|--|--|
| | <p>Hospital/community : Community</p> <p>Type of intervention: Singing and music-based voice interventions</p> <p>No treatment (n=21) No additional treatment</p> <p>Concomitant therapy All people received a resource pack in aphasia-friendly format, constructed for the purpose of the study, which provided information on living with aphasia and the available local community services.</p> | | <p>activities/social groups scores at <6 months and ≥6 months</p> | <p>Clinical Commissioning group and the University of Exeter Medical School. The report is independent research supported by the National Institute for Health Research Applied Research Collaboration South West Peninsula.</p> |

1.1.5.5 Summary matrix

Table 6: Summary matrix of the protocol interventions compared to no treatment

| | | Neurologic music therapy delivered by trained healthcare professionals | Music therapy delivered by trained music therapists | Music interventions delivered by healthcare professionals | Music interventions delivered by non-healthcare professionals |
|---|-----------|---|--|--|--|
| Person/participant generic health-related quality of life | <6 months | 8 outcomes 1 study (n=61) Low-very low quality | 1 outcome 1 study (n=38) Very low quality | 9 outcomes 2 studies (n=66) Very low quality | 1 outcome 1 study (n=36) Very low quality |
| | ≥6 months | No outcomes identified | No outcomes identified | No outcomes identified | 1 outcome 1 study (n=34) Very low quality |
| Carer generic health-related quality of life | <6 months | No outcomes identified | No outcomes identified | No outcomes identified | No outcomes identified |
| | ≥6 months | No outcomes identified | No outcomes identified | No outcomes identified | 1 outcome 1 study (n=34) Very low quality |
| Activities of daily living | <6 months | No outcomes identified | 2 outcomes 2 studies (n=68) Very low quality | 2 outcomes 4 studies (n=119) Low-very low quality | No outcomes identified |
| | ≥6 months | No outcomes identified | No outcomes identified | No outcomes identified | No outcomes identified |
| Psychological distress – Depression | <6 months | 1 outcome 1 study (n=40) Low quality | 2 outcomes 6 studies (n=154) Low-very low quality | 2 outcomes 4 studies (n=195) Very low quality | No outcomes identified |
| | ≥6 months | No outcomes identified | No outcomes identified | No outcomes identified | No outcomes identified |
| Psychological distress – Anxiety | <6 months | 1 outcome 1 study (n=40) Low quality | No outcomes identified | No outcomes identified | No outcomes identified |

| | | Neurologic music therapy delivered by trained healthcare professionals | Music therapy delivered by trained music therapists | Music interventions delivered by healthcare professionals | Music interventions delivered by non-healthcare professionals |
|--|-----------|---|--|--|--|
| | ≥6 months | No outcomes identified | No outcomes identified | No outcomes identified | No outcomes identified |
| Psychological distress – Distress | <6 months | No outcomes identified | No outcomes identified | No outcomes identified | No outcomes identified |
| | ≥6 months | No outcomes identified | No outcomes identified | No outcomes identified | No outcomes identified |
| Psychological distress (other) | <6 months | No outcomes identified | 1 outcome 1 study (n=28) Moderate quality | 1 outcome 1 study (n=37) Very low quality | No outcomes identified |
| | ≥6 months | No outcomes identified | No outcomes identified | No outcomes identified | No outcomes identified |
| Stroke-specific Patient-Reported Outcome Measures | <6 months | 1 outcome 1 study (n=61) Very low quality | 10 outcomes 3 studies (n=73) Moderate-very low quality | 11 outcomes 5 studies (n=192) Very low quality | 1 outcome 1 study (n=36) Very low quality |
| | ≥6 months | No outcomes identified | No outcomes identified | No outcomes identified | 1 outcome 1 study (n=34) Very low quality |
| Wellbeing scores | <6 months | No outcomes identified | 1 outcome 1 study (n=25) Very low quality | No outcomes identified | 1 outcome 1 study (n=36) Very low quality |
| | ≥6 months | No outcomes identified | No outcomes identified | No outcomes identified | 1 outcome 1 study (n=34) Very low quality |
| Participation in leisure activities/social groups scores | <6 months | No outcomes identified | 1 outcome 1 study (n=18) Very low quality | No outcomes identified | 1 outcome 1 study (n=36) Very low quality |
| | ≥6 months | No outcomes identified | No outcomes identified | No outcomes identified | 1 outcome 1 study (n=34) |

| | | Neurologic music therapy delivered by trained healthcare professionals | Music therapy delivered by trained music therapists | Music interventions delivered by healthcare professionals | Music interventions delivered by non-healthcare professionals |
|----------------------------------|-----------|---|--|--|--|
| | | | | | Very low quality |
| Withdrawal due to adverse events | <6 months | No outcomes identified | 1 outcome 4 studies (n=84) Very low quality | 1 outcome 7 studies (n=355) Very low quality | 1 outcome 1 study (n=41) Very low quality |
| | ≥6 months | No outcomes identified | No outcomes identified | No outcomes identified | No outcomes identified |

See Appendix D for full evidence tables.

1.1.6 Summary of the effectiveness evidence

1.1.6.1 Neurologic music therapy delivered by trained music therapists compared to no treatment

Table 7: Clinical evidence summary: neurologic music therapy delivered by trained music therapists compared to no treatment

| Outcomes | No of participants (studies) Follow up | Certainty of the evidence (GRADE) | Relative effect (95% CI) | Anticipated absolute effects | | Comments |
|--|--|--------------------------------------|--------------------------|--|---|-----------------------------------|
| | | | | Risk with no treatment | Risk difference with neurologic music therapy delivered by trained music therapists | |
| Person/participant generic health-related quality of life (SF-36 physical functioning, 0-100, higher values are better, final values) at <6 months | 61 (1 RCT) follow up: 5 weeks | ⊕○○○ ○ Very low _{a,b} | - | The mean person/participant generic health-related quality of life was 16 | MD 1.43 higher (1.11 lower to 3.97 higher) | MID = 3 (SF-36 established value) |
| Person/participant generic health-related quality of life (SF-36 bodily pain, 0-100, higher values are better, final values) at <6 months | 61 (1 RCT) follow up: 5 weeks | ⊕⊕○○ Low _a | - | The mean person/participant generic health-related quality of life was 8.16 | MD 0.47 higher (0.78 lower to 1.72 higher) | MID = 3 (SF-36 established value) |
| Person/participant generic health-related quality of life (SF-36 role physical, 0-100, higher values are better, final values) at <6 months | 61 (1 RCT) follow up: 5 weeks | ⊕⊕○○ Low _a | - | The mean person/participant generic health-related quality of life was 4.71 | MD 0.29 higher (0.21 lower to 0.79 higher) | MID = 3 (SF-36 established value) |
| Person/participant generic health-related quality of life (SF-36 vitality, 0-100, higher values are better, final | 61 (1 RCT) follow up: 5 weeks | ⊕⊕○○ Low _a | - | The mean person/participant generic health-related quality of life was 14.52 | MD 4.11 higher (2.34 higher to 5.88 higher) | MID = 2 (SF-36 established value) |

| Outcomes | № of participants (studies) Follow up | Certainty of the evidence (GRADE) | Relative effect (95% CI) | Anticipated absolute effects | | Comments |
|--|---------------------------------------|--------------------------------------|--------------------------|--|---|-----------------------------------|
| | | | | Risk with no treatment | Risk difference with neurologic music therapy delivered by trained music therapists | |
| values) at <6 months | | | | | | |
| Person/participant generic health-related quality of life (SF-36 general health, 0-100, higher values are better, final values) at <6 months | 61 (1 RCT) follow up: 5 weeks | ⊕○○○ ○ Very low _{a,b} | - | The mean person/participant generic health-related quality of life was 15.97 | MD 1.46 higher (0.25 lower to 3.17 higher) | MID = 2 (SF-36 established value) |
| Person/participant generic health-related quality of life (SF-36 role emotional, 0-100, higher values are better, final values) at <6 months | 61 (1 RCT) follow up: 5 weeks | ⊕⊕○○ Low _a | - | The mean person/participant generic health-related quality of life was 5 | MD 0.83 higher (0.34 higher to 1.32 higher) | MID = 4 (SF-36 established value) |
| Person/participant generic health-related quality of life (SF-36 mental health, 0-100, higher values are better, final values) at <6 months | 61 (1 RCT) follow up: 5 weeks | ⊕○○○ ○ Very low _{a,b} | - | The mean person/participant generic health-related quality of life was 20.48 | MD 3.75 higher (1.32 higher to 6.18 higher) | MID = 3 (SF-36 established value) |
| Person/participant generic health-related quality of life (SF-36 social functioning, 0-100, higher values are better, final values) at <6 months | 61 (1 RCT) follow up: 5 weeks | ⊕⊕○○ ○ Low _a | - | The mean person/participant generic health-related quality of life was 6.81 | MD 0.56 higher (0.54 lower to 1.66 higher) | MID = 3 (SF-36 established value) |
| Psychological distress - depression (Hamilton) | 40 (1 RCT) follow-up: 8 weeks | ⊕⊕○○ ○ Low _{b,c} | - | The mean psychological distress - depression at <6 | MD 1.95 lower (3.07 lower to | MID = 1.1 (0.5 x median) |

| Outcomes | № of participants (studies) Follow up | Certainty of the evidence (GRADE) | Relative effect (95% CI) | Anticipated absolute effects | | Comments |
|---|---------------------------------------|-------------------------------------|--------------------------|--|---|---------------------------------------|
| | | | | Risk with no treatment | Risk difference with neurologic music therapy delivered by trained music therapists | |
| Depression Scale, 0-56, lower values are better, final value) at <6 months | | | | months was 10.9 | 0.83 lower) | baseline SD) |
| Psychological distress - anxiety (Hamilton Anxiety Rating Scale, 0-56, lower values are better, final value) at <6 months | 40 (1 RCT) follow-up: 8 weeks | ⊕⊕○ ○ Low _{b,c} | - | The mean psychological distress - anxiety at <6 months was 9.65 | MD 1.05 lower (2.46 lower to 0.36 higher) | MID = 1.40 (0.5 x median baseline SD) |
| Stroke-specific Patient-Reported Outcome Measures (Stroke adjusted Sickness Impact Profile 30, 0-68, lower values are better, final values) at <6 months | 61 (1 RCT) follow up: 5 weeks | ⊕○○ ○ Very low _{a,b} | - | The mean stroke-specific Patient-Reported Outcome Measures was 16.42 | MD 1.52 lower (3.84 lower to 0.8 higher) | MID = 2.1 (0.5 x median baseline SD) |
| <p>a. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias arising from the randomisation process, bias due to deviations from the intended interventions and bias in measurement of the outcome)</p> <p>b. Downgraded by 1 increment if the confidence interval crossed one MID or by 2 increments if the confidence interval crossed both MIDs</p> <p>c. Downgraded by 1 increment as the majority of the evidence was of high risk of bias (due to bias arising from the randomisation process)</p> | | | | | | |

1.1.6.2 Music therapy delivered by trained music therapists compared to no treatment

Table 8: Clinical evidence summary: music therapy delivered by trained music therapists compared to no treatment

| Outcomes | No of participants (studies) Follow-up | Certainty of the evidence (GRADE) | Relative effect (95% CI) | Anticipated absolute effects | | Comments |
|--|--|--------------------------------------|--------------------------|---|--|--|
| | | | | Risk with no treatment | Risk difference with music therapy delivered by trained music therapists | |
| Patient/participant generic health-related quality of life (McGill Quality of Life, 0-10, higher values are better, final value) at <6 months | 38 (1 RCT) follow-up: 2 months | ⊕○○○ ○ Very low _{a,b} | - | The mean patient/participant generic health-related quality of life at <6 months was 7.49 | MD 0.27 higher (0.7 lower to 1.24 higher) | MID = 0.78 (0.5 x median baseline SD) |
| Activities of daily living (Korean-Modified Barthel Index, 0-100, higher values are better, change score) at <6 months | 30 (1 RCT) follow-up: 8 weeks | ⊕○○○ ○ Very low _{b,c} | - | The mean activities of daily living at <6 months was 7.2 | MD 2 higher (13.25 lower to 17.25 higher) | MID = 1.85 (Barthel index established MID) |
| Activities of daily living (Functional independence measure, 18-126, higher values are better, final value) at <6 months | 38 (1 RCT) follow-up: 2 months | ⊕⊕○○ ○ Low _a | - | The mean activities of daily living at <6 months was 106.89 | MD 3.58 higher (5.2 lower to 12.36 higher) | MID = 22 (Functional independence measure established MID) |
| Psychological distress - Depression (HADS-D, BDI, Faces scale, PANAS - negative affect [different scale ranges], lower values are better, final values) at <6 months | 124 (5 RCTs) follow-up: mean 8 weeks | ⊕⊕○○ ○ Low _d | - | - | SMD 0.03 SD lower (0.39 lower to 0.32 higher) | MID = 0.5 SD (SMD) |
| Psychological distress - | 30 (1 RCT) | ⊕○○○ ○ | - | The mean psychological | MD 3.21 higher | MID = 6.1 (0.5 x |

| Outcomes | № of participants (studies) Follow-up | Certainty of the evidence (GRADE) | Relative effect (95% CI) | Anticipated absolute effects | | Comments |
|---|---------------------------------------|--|--------------------------|---|--|--------------------------------------|
| | | | | Risk with no treatment | Risk difference with music therapy delivered by trained music therapists | |
| Depression (Center for Epidemiologic Studies Depression Scale, 0-60, lower values are better, change score) at <6 months | follow-up: 8 weeks | Very low _{b,c} | | distress - Depression at <6 months was - 9.67 | (6.56 lower to 12.98 higher) | median baseline SD) |
| Psychological distress - Anxiety (HADS-A, BAI [different scale ranges], lower values are better, final values) at <6 months | 56 (2 RCTs) follow-up: mean 6 weeks | ⊕○○○ ○ Very low _{b,e} | - | - | SMD 0.18 SD lower (0.7 lower to 0.35 higher) | MID = 0.5 SD (SMD) |
| Psychological distress (PANAS - positive affect, 10-50, higher values are better, final value) at <6 months | 28 (1 RCT) follow-up: 4 months | ⊕⊕⊕○ e _b Moderate | - | The mean psychological distress at <6 months was 32.64 | MD 4.15 higher (2.01 lower to 10.31 higher) | MID = 4.7 (0.5 x median baseline SD) |
| Stroke-specific Patient-Reported Outcome Measures (Stroke specific quality of life, 49-245, higher values are better, final value) at <6 months | 20 (1 RCT) follow-up: 6 weeks | ⊕○○○ ○ Very low _{b,e} | - | The mean stroke-specific Patient-Reported Outcome Measures at <6 months was 159.2 | MD 24.5 higher (7.36 higher to 41.64 higher) | MID = 8.9 (0.5 x median baseline SD) |
| Stroke-specific Patient-Reported Outcome Measures (SIS physical strength, 0-100, higher values are better, final | 25 (1 RCT) follow-up: 6 weeks | ⊕○○○ ○ Very low _{b,f,g} | - | The mean stroke-specific Patient-Reported Outcome Measures at <6 months was 49.1 | MD 10.7 lower (23.83 lower to 2.43 higher) | MID = 8.5 (0.5 x median baseline SD) |

| Outcomes | No of participants (studies) Follow-up | Certainty of the evidence (GRADE) | Relative effect (95% CI) | Anticipated absolute effects | | Comments |
|--|--|--|--------------------------|--|--|--------------------------------------|
| | | | | Risk with no treatment | Risk difference with music therapy delivered by trained music therapists | |
| value) at <6 months | | | | | | |
| Stroke-specific Patient-Reported Outcome Measures (SIS activities, 0-100, higher values are better, final value) at <6 months | 25 (1 RCT) follow-up: 6 weeks | ⊕○○○ ○ Very low _{b,f,g} | - | The mean stroke-specific Patient-Reported Outcome Measures at <6 months was 53.3 | MD 6.4 lower (20.13 lower to 7.33 higher) | MID = 7.9 (0.5 x median baseline SD) |
| Stroke-specific Patient-Reported Outcome Measures (SIS hand use, 0-100, higher values are better, final value) at <6 months | 25 (1 RCT) follow-up: 6 weeks | ⊕○○○ ○ Very low _{b,f,g} | - | The mean stroke-specific Patient-Reported Outcome Measures at <6 months was 31 | MD 12 lower (33.74 lower to 9.74 higher) | MID = 9.9 (0.5 x median baseline SD) |
| Stroke-specific Patient-Reported Outcome Measures (SIS mobility, 0-100, higher values are better, final value) at <6 months | 53 (2 RCTs) follow-up: mean 11 weeks | ⊕⊕⊕○ Moderate _b | - | The mean stroke-specific Patient-Reported Outcome Measures at <6 months was 60.9 | MD 4.96 lower (13.36 lower to 3.44 higher) | MID = 8.8 (0.5 x median baseline SD) |
| Stroke-specific Patient-Reported Outcome Measures (SIS communication, 0-100, higher values are better, final value) at <6 months | 53 (2 RCTs) follow-up: mean 11 weeks | ⊕⊕⊕○ Moderate _b | - | The mean stroke-specific Patient-Reported Outcome Measures at <6 months was 82.8 | MD 1.89 lower (8.05 lower to 4.27 higher) | MID = 7.6 (0.5 x median baseline SD) |
| Stroke-specific Patient-Reported Outcome Measures (SIS memory, 0-100, higher values are better, final | 53 (2 RCTs) follow-up: mean 11 weeks | ⊕⊕○○ ○ Very low _{b,f,g} | - | The mean stroke-specific Patient-Reported Outcome Measures at <6 | MD 0.65 lower (9.93 lower to 8.64 higher) | MID = 9.1 (0.5 x median baseline SD) |

| Outcomes | No of participants (studies) Follow-up | Certainty of the evidence (GRADE) | Relative effect (95% CI) | Anticipated absolute effects | | Comments |
|---|--|-----------------------------------|--------------------------|---|--|--------------------------------------|
| | | | | Risk with no treatment | Risk difference with music therapy delivered by trained music therapists | |
| value) at <6 months | | | | months was 77.1 | | |
| Stroke-specific Patient-Reported Outcome Measures (SIS emotion, 0-100, higher values are better, final value) at <6 months | 53 (2 RCTs) follow-up: mean 11 weeks | ⊕⊕○○ Low _{b,g} | - | The mean stroke-specific Patient-Reported Outcome Measures at <6 months was 67.9 | MD 1.01 lower (7.7 lower to 9.72 higher) | MID = 8.7 (0.5 x median baseline SD) |
| Stroke-specific Patient-Reported Outcome Measures (SIS social/participation, 0-100, higher values are better, final value) at <6 months | 53 (2 RCTs) follow-up: mean 11 weeks | ⊕⊕○○ Low _{b,g} | - | The mean stroke-specific Patient-Reported Outcome Measures at <6 months was 57.2 | MD 2.38 higher (12.84 lower to 8.08 higher) | MID = 8.8 (0.5 x median baseline SD) |
| Stroke-specific Patient-Reported Outcome Measures (SIS recovery, 0-100, higher values are better, final value) at <6 months | 25 (1 RCT) follow-up: 6 weeks | ⊕○○○ Very low _{b,f,g} | - | The mean stroke-specific Patient-Reported Outcome Measures at <6 months was 62.7 | MD 1.2 lower (13.66 lower to 11.26 higher) | MID = 7.5 (0.5 x median baseline SD) |
| Wellbeing scores (WHO five item well-being index, 0-25, higher values are better, final value) at <6 months | 25 (1 RCT) follow-up: 6 weeks | ⊕○○○ Very low _{b,f,g} | - | The mean wellbeing scores at <6 months was 19 | MD 2.4 higher (1.71 lower to 6.51 higher) | MID = 3.1 (0.5 x median baseline SD) |
| Participation in leisure activities/social groups (Sickness Impact Profile Social | 18 (1 RCT) follow-up: 4 months | ⊕○○○ Very low _{e,h} | - | The mean participation in leisure activities/social groups at <6 months was 42.88 | MD 13.28 lower (19.7 lower to 6.86 lower) | MID = 3.7 (0.5 x median baseline SD) |

| Outcomes | No of participants (studies) Follow-up | Certainty of the evidence (GRADE) | Relative effect (95% CI) | Anticipated absolute effects | | Comments |
|---|--|-----------------------------------|--------------------------|------------------------------|--|--|
| | | | | Risk with no treatment | Risk difference with music therapy delivered by trained music therapists | |
| Interaction subscale, 0-102, lower values are better, final value) at <6 months | | | | | | |
| Withdrawal due to adverse events at <6 months | 84 (4 RCTs) follow-up: 7 weeks | ⊕⊕○○ Low _{i,j} | RD 0.00 (-0.09 to 0.09) | 0 per 1,000 | 0 fewer per 1,000 (90 fewer to 90 more) ^k | Sample size used to determine precision: 75-150 = serious imprecision, <75 = very serious imprecision. |

- a. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias arising from the randomisation process and bias due to deviations from the intended interventions)
- b. Downgraded by 1 increment if the confidence interval crossed one MID or by 2 increments if the confidence interval crossed both MIDs
- c. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias arising from the randomisation process, bias due to deviations from the intended interventions, bias due to missing outcome data and bias in measurement of the outcome)
- d. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to a mixture of bias arising from the randomisation process, bias due to deviations from the intended interventions, bias due to missing outcome data and bias in measurement of the outcome)
- e. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to a mixture of bias arising from the randomisation process, bias due to deviations from the intended interventions and bias in measurement of the outcome)
- f. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias arising from the randomisation process and bias due to missing outcome data)
- g. Downgraded by 1 increment due to comparator indirectness (due to the comparator group in 1 study receiving an exercise intervention that may have been more intense than that received than the intervention group, however it was unclear as to whether this was the case from the information provided in the study)
- h. Downgraded by 1 increments due to population indirectness (10-20% of people in a study having had a traumatic brain injury rather than a stroke)
- i. Downgraded by 1 increment as the majority of the evidence was of high risk of bias (due to bias arising from the randomisation process)
- j. Downgraded by 1 to 2 increments for imprecision due to zero events and small sample size
- k. Absolute effect calculated by risk difference due to zero events in at least one arm of one study

1.1.6.3 Music interventions delivered by healthcare professionals compared to passive music listening

Table 9: Clinical evidence summary: music intervention delivered by healthcare professionals compared to passive music listening

| Outcomes | No of participants (studies) Follow up | Certainty of the evidence (GRADE) | Relative effect (95% CI) | Anticipated absolute effects | | Comments |
|---|--|-----------------------------------|--------------------------|-----------------------------------|---|---------------------------------|
| | | | | Risk with passive music listening | Risk difference with music intervention delivered by healthcare professionals | |
| Withdrawal due to adverse events at <6 months | 48 (1 RCT) follow up: 3 months | ⊕○○○ Very low a,b | RR 4.35 (0.52 to 36.11) | 40 per 1,000 | 134 more per 1,000 (19 fewer to 1,404 more) | MID (precision): RR = 0.8-1.25. |
| Withdrawal due to adverse events at ≥6 months | 48 (1 RCT) follow up: 6 months | ⊕○○○ Very low a,b | RR 4.35 (0.52 to 36.11) | 40 per 1,000 | 134 more per 1,000 (19 fewer to 1,404 more) | MID (precision): RR = 0.8-1.25. |

a. Downgraded by 1 increment as the majority of the evidence was of high risk of bias (due to bias due to missing outcome data)

b. Downgraded by 1 increment if the confidence interval crossed one MID or by 2 increments if the confidence interval crossed both MIDs

1.1.6.4 Music interventions delivered by healthcare professionals compared to placebo music therapy

Table 10: Clinical evidence summary: music intervention delivered by healthcare professionals compared to placebo music therapy

| Outcomes | No of participants (studies) Follow up | Certainty of the evidence (GRADE) | Relative effect (95% CI) | Anticipated absolute effects | | Comments |
|--|--|-----------------------------------|--------------------------|---------------------------------|---|---|
| | | | | Risk with placebo music therapy | Risk difference with music intervention delivered by healthcare professionals | |
| Psychological distress - Depression (HADS-D, 0-42, higher values are better, mean difference) at <6 months | 48 (1 RCT) follow up: 3 months | ⊕○○○ Very low a,b | - | - | MD 0.7 higher (1.65 lower to 3.05 higher) | MID = 2.1 (0.5 x SD calculated from mean difference standard error) |
| Psychological distress - Depression (HADS-D, 0-42, higher values are better, mean | 48 (1 RCT) follow up: 6 months | ⊕○○○ Very low a,b | - | - | MD 1.02 higher (1.36 lower to 3.4 higher) | MID = 2.1 (0.5 x SD calculated from mean difference |

| Outcomes | № of participants (studies) Follow up | Certainty of the evidence (GRADE) | Relative effect (95% CI) | Anticipated absolute effects | | Comments |
|--|---------------------------------------|-----------------------------------|--------------------------|---------------------------------|---|--|
| | | | | Risk with placebo music therapy | Risk difference with music intervention delivered by healthcare professionals | |
| difference) at ≥6 months | | | | | | standard error) |
| Psychological distress - Anxiety (HADS-A, 0-42, higher values are better, mean difference) at <6 months | 48 (1 RCT) follow up: 3 months | ⊕○○○ Very low a,b | - | - | MD 0.69 higher (1.47 lower to 2.85 higher) | MID = 1.9 (0.5 x SD calculated from mean difference standard error) |
| Psychological distress - Anxiety (HADS-A, 0-42, higher values are better, mean difference) at ≥6 months | 48 (1 RCT) follow up: 6 months | ⊕○○○ Very low a,b | - | - | MD 2 higher (0.28 lower to 4.28 higher) | MID = 2.0 (0.5 x SD calculated from mean difference standard error) |
| Participation in leisure activities/social group scores (Mayo-Portland Adaptability Inventory 4 participation, 0-30, higher values are better, mean difference) at <6 months | 48 (1 RCT) follow up: 3 months | ⊕⊕○○ Low a | - | - | MD 1.72 higher (11.75 lower to 15.19 higher) | MID = 11.9 (0.5 x SD calculated from mean difference standard error) |
| Withdrawal due to adverse events at <6 months | 48 (1 RCT) follow up: 3 months | ⊕○○○ Very low b,c | RR 4.35 (0.52 to 36.11) | 40 per 1,000 | 134 more per 1,000 (19 fewer to 1,404 more) | MID (precision) = RR 0.8-1.25. |
| Withdrawal due to adverse events at ≥6 months | 48 (1 RCT) follow up: 6 months | ⊕○○○ Very low b,c | RR 4.35 (0.52 to 36.11) | 40 per 1,000 | 134 more per 1,000 (19 fewer to 1,404 more) | MID (precision) = RR 0.8-1.25. |

a. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias due to deviations from the intended interventions and bias due to missing outcome data)

b. Downgraded by 1 increment if the confidence interval crossed one MID or by 2 increments if the confidence interval crossed both MIDs

c. Downgraded by 1 increment as the majority of the evidence was of high risk of bias (due to bias due to missing outcome data)

1.1.6.5 Music interventions delivered by healthcare professionals compared to no treatment

Table 11: Clinical evidence summary: music intervention delivered by healthcare professionals compared to no treatment

| Outcomes | № of participants (studies) Follow-up | Certainty of the evidence (GRADE) | Relative effect (95% CI) | Anticipated absolute effects | | Comments |
|--|---------------------------------------|--------------------------------------|--------------------------|--|---|-----------------------------------|
| | | | | Risk with no treatment | Risk difference with music intervention delivered by healthcare professionals | |
| Person/participant generic health-related quality of life (SF-36 physical function, 0-100, higher values are better, final value) at <6 months | 37 (1 RCT) follow-up: 4 weeks | ⊕○○○ ○ Very low _{a,b} | - | The mean person/participant generic health-related quality of life at <6 months was 45 | MD 10.3 higher (4.31 lower to 24.91 higher) | MID = 3 (SF-36 established value) |
| Person/participant generic health-related quality of life (SF-36 bodily pain, 0-100, higher values are better, final value) at <6 months | 37 (1 RCT) follow-up: 4 weeks | ⊕○○○ ○ Very low _{a,b} | - | The mean person/participant generic health-related quality of life at <6 months was 50.4 | MD 6.3 higher (13.48 lower to 26.08 higher) | MID = 3 (SF-36 established value) |
| Person/participant generic health-related quality of life (SF-36 role physical, 0-100, higher values are better, final value) at <6 months | 37 (1 RCT) follow-up: 4 weeks | ⊕○○○ ○ Very low _{a,b} | - | The mean person/participant generic health-related quality of life at <6 months was 23.7 | MD 6.1 lower (29.41 lower to 17.21 higher) | MID = 3 (SF-36 established value) |
| Person/participant generic health-related quality of life (SF-36 vitality, 0-100, higher values are better, final value) at <6 months | 37 (1 RCT) follow-up: 4 weeks | ⊕○○○ ○ Very low _{a,b} | - | The mean person/participant generic health-related quality of life at <6 months was 53.2 | MD 6.8 higher (8.74 lower to 22.34 higher) | MID = 2 (SF-36 established value) |
| Person/participant generic health-related quality of life (SF-36 general | 37 (1 RCT) follow-up: 4 weeks | ⊕○○○ ○ Very low _{a,b} | - | The mean person/participant generic health-related quality of life at | MD 3.9 higher (9.57 lower to 17.37 higher) | MID = 2 (SF-36 established value) |

| Outcomes | № of participants (studies) Follow-up | Certainty of the evidence (GRADE) | Relative effect (95% CI) | Anticipated absolute effects | | Comments |
|--|---------------------------------------|--|--------------------------|--|---|---------------------------------------|
| | | | | Risk with no treatment | Risk difference with music intervention delivered by healthcare professionals | |
| health, 0-100, higher values are better, final value) at <6 months | | | | <6 months was 57.2 | | |
| Person/participant generic health-related quality of life (SF-36 role emotional, 0-100, higher values are better, final value) at <6 months | 37 (1 RCT) follow-up: 4 weeks | ⊕○○○ ○ Very low _{a,b} | - | The mean person/participant generic health-related quality of life at <6 months was 70.9 | MD 5.6 higher (21.41 lower to 32.61 higher) | MID = 4 (SF-36 established value) |
| Person/participant generic health-related quality of life (SF-36 mental health, 0-100, higher values are better, final value) at <6 months | 37 (1 RCT) follow-up: 4 weeks | ⊕○○○ ○ Very low _{a,b} | - | The mean person/participant generic health-related quality of life at <6 months was 66.3 | MD 4.8 higher (9.05 lower to 18.65 higher) | MID = 3 (SF-36 established value) |
| Person/participant generic health-related quality of life (SF-36 social function, 0-100, higher values are better, final value) at <6 months | 37 (1 RCT) follow-up: 4 weeks | ⊕○○○ ○ Very low _{b,c} | - | The mean person/participant generic health-related quality of life at <6 months was 64.7 | MD 11 higher (7.93 lower to 29.93 higher) | MID = 3 (SF-36 established value) |
| Person/participant health-related quality of life (McGill Quality of Life, 0-10, higher values are better, final value) at <6 months | 29 (1 RCT) follow-up: 2 months | ⊕⊕○○ ○ Low _d | - | The mean person/participant health-related quality of life at <6 months was 7.34 | MD 0.49 lower (1.65 lower to 0.67 higher) | MID = 0.78 (0.5 x median baseline SD) |
| Activities of daily living (Canadian Occupational Performance Measure, | 89 (3 RCTs) follow-up: 6 weeks | ⊕○○○ ○ Very low _{b,e,f} | - | - | SMD 0.64 SD higher (0.15 lower to 1.42 higher) | MID = 0.5 SD (SMD) |

| Outcomes | № of participants (studies) Follow-up | Certainty of the evidence (GRADE) | Relative effect (95% CI) | Anticipated absolute effects | | Comments |
|---|---------------------------------------|--------------------------------------|--------------------------|--|---|--|
| | | | | Risk with no treatment | Risk difference with music intervention delivered by healthcare professionals | |
| University of Maryland Arm Questionnaire for Stroke, Activities of daily living score [different scale ranges], higher values are better, change scores) at <6 months | | | | | | |
| Activities of daily living (Barthel index, 0-100, higher values are better, final value) at <6 months | 30 (1 RCT) follow-up: 4 weeks | ⊕⊕○○ ○ Low _g | - | The mean activities of daily living at <6 months was 69.67 | MD 10.66 higher (4.85 higher to 16.47 higher) | MID = 1.85 (Barthel index established MID) |
| Psychological distress - Depression (Hamilton's Depression Scale-17, 0-56, lower values are better, change score) at <6 months | 60 (1 RCT) follow-up: 17 days | ⊕○○○ ○ Very low _{b,e} | - | The mean psychological distress - Depression at <6 months was -4.9 | MD 1.3 lower (2.22 lower to 0.38 lower) | MID = 1.9 (0.5 x median baseline SD) |
| Psychological distress - Depression (BDI, Hamilton Depression Rating Scale-24, profile of mood states [different scale ranges], lower values are better, final values) at <6 months | 135 (3 RCTs) follow-up: mean 6 weeks | ⊕○○○ ○ Very low _{b,h} | - | - | SMD 0.68 SD lower (1.03 lower to 0.33 higher) | MID = 0.5 SD (SMD) |
| Psychological distress (PANAS positive affect, 10-50, higher values are better, change score) at <6 months | 37 (1 RCT) follow-up: 4 weeks | ⊕○○○ ○ Very low _{a,b} | - | The mean psychological distress at <6 months was 30.3 | MD 3.4 higher (1.95 lower to 8.75 higher) | MID = 4.3 (0.5 x median baseline SD) |

| Outcomes | № of participants (studies) Follow-up | Certainty of the evidence (GRADE) | Relative effect (95% CI) | Anticipated absolute effects | | Comments |
|--|---------------------------------------|--|--------------------------|--|---|--------------------------------------|
| | | | | Risk with no treatment | Risk difference with music intervention delivered by healthcare professionals | |
| Stroke-specific Patient-Reported Outcome Measures (SIS [different scale ranges], higher values are better, change scores) at <6 months | 89 (2 RCTs) follow-up: 8 weeks | ⊕○○○ ○ Very low _{b,f,h} | - | - | SMD 1.23 SD lower (2.57 lower to 0.1 higher) | MID = 0.5 SD (SMD) |
| Stroke-specific Patient-Reported Outcome Measures (SIS strength subscale, 0-100, higher values are better, change score) at <6 months | 32 (1 RCT) follow-up: 12 weeks | ⊕○○○ ○ Very low _{b,i} | - | The mean stroke-specific Patient-Reported Outcome Measures at <6 months was 11.7 | MD 12.8 lower (21.9 lower to 3.7 lower) | MID = 7.4 (0.5 x median baseline SD) |
| Stroke-specific Patient-Reported Outcome Measures (SIS memory subscale, 0-100, higher values are better, change score) at <6 months | 32 (1 RCT) follow-up: 12 weeks | ⊕○○○ ○ Very low _{b,i} | - | The mean stroke-specific Patient-Reported Outcome Measures at <6 months was -0.9 | MD 1.1 higher (5.07 lower to 7.27 higher) | MID = 8.1 (0.5 x median baseline SD) |
| Stroke-specific Patient-Reported Outcome Measures (SIS emotion subscale, 0-100, higher values are better, change score) at <6 months | 32 (1 RCT) follow-up: 12 weeks | ⊕○○○ ○ Very low _{b,i} | - | The mean stroke-specific Patient-Reported Outcome Measures at <6 months was 3.9 | MD 1.9 higher (8.54 lower to 12.34 higher) | MID = 7.6 (0.5 x median baseline SD) |
| Stroke-specific Patient-Reported Outcome Measures (SIS communication | 32 (1 RCT) follow-up: 12 weeks | ⊕○○○ ○ Very low _{b,i} | - | The mean stroke-specific Patient-Reported Outcome | MD 0.8 lower (9.4 lower to 7.8 higher) | MID = 7.0 (0.5 x median baseline SD) |

| Outcomes | № of participants (studies) Follow-up | Certainty of the evidence (GRADE) | Relative effect (95% CI) | Anticipated absolute effects | | Comments |
|---|---------------------------------------|--------------------------------------|--------------------------|---|---|---------------------------------------|
| | | | | Risk with no treatment | Risk difference with music intervention delivered by healthcare professionals | |
| subscale, 0-100, higher values are better, change score) at <6 months | | | | Measures at <6 months was 3.1 | | |
| Stroke-specific Patient-Reported Outcome Measures (SIS ADL subscale, 0-100, higher values are better, change score) at <6 months | 32 (1 RCT) follow-up: 12 weeks | ⊕○○○ ○ Very low _{b,i} | - | The mean stroke-specific Patient-Reported Outcome Measures at <6 months was 0 | MD 2.8 higher (6.28 lower to 11.88 higher) | MID = 9.1 (0.5 x median baseline SD) |
| Stroke-specific Patient-Reported Outcome Measures (SIS mobility subscale, 0-100, higher values are better, change score) at <6 months | 32 (1 RCT) follow-up: 12 weeks | ⊕○○○ ○ Very low _{b,i} | - | The mean stroke-specific Patient-Reported Outcome Measures at <6 months was 3.1 | MD 1.3 lower (7.6 lower to 5 higher) | MID = 12.8 (0.5 x median baseline SD) |
| Stroke-specific Patient-Reported Outcome Measures (SIS hand function subscale, 0-100, higher values are better, change score) at <6 months | 32 (1 RCT) follow-up: 12 weeks | ⊕○○○ ○ Very low _{b,i} | - | The mean stroke-specific Patient-Reported Outcome Measures at <6 months was 9.7 | MD 1.8 higher (13.96 lower to 17.56 higher) | MID = 13.5 (0.5 x median baseline SD) |
| Stroke-specific Patient-Reported Outcome Measures (SIS social participation subscale, 0-100, higher values are better, change score) at <6 months | 32 (1 RCT) follow-up: 12 weeks | ⊕○○○ ○ Very low _{b,i} | - | The mean stroke-specific Patient-Reported Outcome Measures at <6 months was 3.1 | MD 3.7 higher (10.15 lower to 17.55 higher) | MID = 10.0 (0.5 x median baseline SD) |

| Outcomes | № of participants (studies) Follow-up | Certainty of the evidence (GRADE) | Relative effect (95% CI) | Anticipated absolute effects | | Comments |
|---|---------------------------------------|--|--------------------------|------------------------------|---|---|
| | | | | Risk with no treatment | Risk difference with music intervention delivered by healthcare professionals | |
| Stroke-specific Patient-Reported Outcome Measures (Stroke Specific Quality of Life [different scale ranges], higher values are better, final values) at <6 months | 71 (2 RCTs) follow-up: mean 6 weeks | ⊕○○○ ○ Very low ^{b,f,h} | - | - | SMD 0.4 SD higher (0.28 lower to 1.09 higher) | MID = 0.5 SD (SMD) |
| Withdrawal due to adverse events at <6 months | 355 (7 RCTs) follow-up: mean 7 weeks | ⊕○○○ ○ Very low ^{j,k,l} | RD 0.01 (-0.05 to 0.07) | 79 per 1,000 | 10 fewer per 1,000 (70 fewer to 50 more) ^m | Precision calculated through Optimal Information Size (OIS) due to zero events in some studies. OIS determined power for the sample size = 0.04 (0.8-0.9 = serious, <0.8 = very serious). |

a. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias arising from the randomisation process, bias due to deviations from the intended interventions and bias in selection of the reported result)

b. Downgraded by 1 increment if the confidence interval crossed one MID or by 2 increments if the confidence interval crossed both MIDs

c. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias due to deviations from the intended interventions and bias in selection of the reported result)

d. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias due to deviations from the intended interventions and bias due to missing outcome data)

e. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to a mixture of bias arising from the randomisation process, bias due to deviations from the intended interventions and bias in measurement of the outcome)

f. Downgraded by 1 or 2 increments because heterogeneity, unexplained by subgroup analysis

| Outcomes | № of participants (studies) Follow-up | Certainty of the evidence (GRADE) | Relative effect (95% CI) | Anticipated absolute effects | | Comments |
|--|---------------------------------------|-----------------------------------|--------------------------|------------------------------|---|----------|
| | | | | Risk with no treatment | Risk difference with music intervention delivered by healthcare professionals | |
| g. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias arising from the randomisation process and bias due to deviations from the intended interventions) | | | | | | |
| h. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to a mixture of bias arising from the randomisation process, bias due to deviations from the intended interventions, bias due to missing outcome data, bias in measurement of the outcome and bias in selection of the reported result) | | | | | | |
| i. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias due to deviations from the intended interventions and bias due to missing outcome data) | | | | | | |
| j. Downgraded by 1 increment as the majority of the evidence was of high risk of bias (due to a mixture of bias arising from the randomisation process, bias due to missing outcome data, bias in measurement of the outcome and bias in selection of the reported result) | | | | | | |
| k. Downgraded for heterogeneity due to conflicting number of events in different studies (zero events in one or more studies) | | | | | | |
| l. Downgraded by 1 to 2 increments for imprecision due to zero events and small sample size | | | | | | |
| m. Absolute effect calculated by risk difference due to zero events in at least one arm of one study | | | | | | |

1.1.6.6 Music interventions delivered by non-healthcare professionals compared to no treatment

Table 12: Clinical evidence summary: music interventions delivered by non-healthcare professionals compared to no treatment

| Outcomes | № of participants (studies) Follow-up | Certainty of the evidence (GRADE) | Relative effect (95% CI) | Anticipated absolute effects | | Comments |
|---|---------------------------------------|--------------------------------------|--------------------------|---|---|------------------------------------|
| | | | | Risk with no treatment | Risk difference with music intervention delivered by non-healthcare professionals | |
| Person/participant generic health-related quality of life (EQ-5D 5L, -0.11-1, higher values are better, final value) at <6 months | 36 (1 RCT) follow-up: 3 months | ⊕○○○ ○ Very low _{a,b} | - | The mean person/participant generic health-related quality of life at <6 months was 0.727 | MD 0.04 lower (0.21 lower to 0.12 higher) | MID = EQ-5D 0.03 (established MID) |
| Person/participant generic health-related quality of life | 34 (1 RCT) follow-up: 6 months | ⊕○○○ ○ Very low _{b,c} | - | The mean person/participant generic health-related | MD 0.1 higher (0.06 lower) | MID = EQ-5D 0.03 (established MID) |

| Outcomes | № of participants (studies) Follow-up | Certainty of the evidence (GRADE) | Relative effect (95% CI) | Anticipated absolute effects | | Comments |
|---|---------------------------------------|--------------------------------------|--------------------------|---|---|---------------------------------------|
| | | | | Risk with no treatment | Risk difference with music intervention delivered by non-healthcare professionals | |
| (EQ-5D 5L, -0.11-1, higher values are better, final value) at ≥6 months | | | | quality of life at ≥6 months was 0.651 | to 0.26 higher) | |
| Carer generic health-related quality of life (CarerQoL-7D, 0-14, higher values are better, final value) at ≥6 months | 34 (1 RCT) follow-up: 6 months | ⊕○○○ ○ Very low _{b,c} | - | The mean carer generic health-related quality of life at ≥6 months was 9 | MD 0 (1.97 lower to 1.97 higher) | MID = 1.2 (0.5 x median baseline SD) |
| Stroke-specific Patient-Reported Outcome Measures (Stroke and Aphasia Quality of Life, 1-5, higher values are better, final value) at <6 months | 36 (1 RCT) follow-up: 3 months | ⊕○○○ ○ Very low _{a,b} | - | The mean stroke-specific Patient-Reported Outcome Measures at <6 months was 3.6 | MD 0 (0.49 lower to 0.49 higher) | MID = 0.35 (0.5 x median baseline SD) |
| Stroke-specific Patient-Reported Outcome Measures (Stroke and Aphasia Quality of Life, 1-5, higher values are better, final value) at ≥6 months | 34 (1 RCT) follow-up: 6 months | ⊕○○○ ○ Very low _{b,c} | - | The mean stroke-specific Patient-Reported Outcome Measures at ≥6 months was 3.4 | MD 0.3 higher (0.2 lower to 0.8 higher) | MID = 0.35 (0.5 x median baseline SD) |
| Wellbeing scores (ICEpop CAPability measure for adults, 0-1, higher values are better, final value) at <6 months | 36 (1 RCT) follow-up: 3 months | ⊕○○○ ○ Very low _{b,d} | - | The mean wellbeing scores at <6 months was 0.748 | MD 0.07 higher (0.3 lower to 0.44 higher) | MID = 0.07 (0.5 x median baseline SD) |

| Outcomes | № of participants (studies) Follow-up | Certainty of the evidence (GRADE) | Relative effect (95% CI) | Anticipated absolute effects | | Comments |
|---|---------------------------------------|--------------------------------------|-------------------------------|---|---|---------------------------------------|
| | | | | Risk with no treatment | Risk difference with music intervention delivered by non-healthcare professionals | |
| Wellbeing scores (ICEpop CAPability measure for adults, 0-1, higher values are better, final value) at ≥6 months | 34 (1 RCT) follow-up: 6 months | ⊕○○○ ○ Very low _{b,d} | - | The mean wellbeing scores at ≥6 months was 0.777 | MD 0.04 higher (0.06 lower to 0.13 higher) | MID = 0.07 (0.5 x median baseline SD) |
| Participation in leisure activities/social groups scores (modified Reintegration to Normal Living Index, 0-100, higher values are better, final value) at <6 months | 36 (1 RCT) follow-up: 3 months | ⊕⊕○○ ○ Low _{a,b} | - | The mean participation in leisure activities/social groups scores at <6 months was 22.1 | MD 1 higher (3.39 lower to 5.39 higher) | MID = 3.5 (0.5 x median baseline SD) |
| Participation in leisure activities/social groups scores (modified Reintegration to Normal Living Index, 0-100, higher values are better, final value) at ≥6 months | 34 (1 RCT) follow-up: 6 months | ⊕○○○ ○ Very low _{b,c} | - | The mean participation in leisure activities/social groups scores at ≥6 months was 22.1 | MD 1.8 higher (2.86 lower to 6.46 higher) | MID = 3.5 (0.5 x median baseline SD) |
| Withdrawal due to adverse events at <6 months | 41 (1 RCT) follow-up: 3 months | ⊕⊕○○ ○ Low _b | Peto OR 8.19 (0.49 to 135.71) | 0 per 1,000 | 100 more per 1,000 (50 fewer to 250 more) ^e | MID (precision) = Peto OR 0.8-1.25. |

a. Downgraded by 1 increment as the majority of the evidence was of high risk of bias (due to bias due to deviations from the intended interventions)

b. Downgraded by 1 increment if the confidence interval crossed one MID or by 2 increments if the confidence interval crossed both MIDs

c. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias due to deviations from the intended interventions and bias due to missing outcome data)

d. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias arising from the randomisation process, bias due to deviations from the intended interventions and bias due to missing outcome data)

| Outcomes | № of participants (studies) Follow-up | Certainty of the evidence (GRADE) | Relative effect (95% CI) | Anticipated absolute effects | | Comments |
|--|---------------------------------------|-----------------------------------|--------------------------|------------------------------|---|----------|
| | | | | Risk with no treatment | Risk difference with music intervention delivered by non-healthcare professionals | |
| e. Absolute effect calculated by risk difference due to zero events in at least one arm of one study | | | | | | |

See Appendix F for full GRADE tables.

1.1.7 Economic evidence

1.1.7.1 Included studies

One health economic study was included in this review.²⁰ This related to a music intervention delivered by non-healthcare professionals. This is summarised in the health economic evidence profile below (Table 13) and the health economic evidence table in **Error! Reference source not found..**

No health economic studies were included that related to neurologic music therapy delivered by trained music therapists, music therapy delivered by trained music therapists or music interventions delivered by healthcare professionals.

1.1.7.2 Excluded studies

No relevant health economic studies were excluded due to assessment of limited applicability or methodological limitations.

See also the health economic study selection flow chart in **Error! Reference source not found..**

1.1.8 Summary of included economic evidence

Table 13: Health economic evidence profile: music interventions delivered by non-healthcare professionals compared to no treatment

| Study | Applicability | Limitations | Other comments | Incremental cost | Incremental effects | Cost effectiveness | Uncertainty |
|---------------------------------|-------------------------------------|--|---|---------------------|---|--|--|
| Tarrant 2021 ²⁰ (UK) | Partially applicable ^(a) | Potentially serious limitations ^(b) | <ul style="list-style-type: none"> • Within-RCT analysis (Tarrant 2021²⁰) • Cost-utility analysis (CUA) (health outcome: QALYs) • Population: Adults with post-stroke aphasia • Comparators: <ol style="list-style-type: none"> 1. Control group (no treatment) 2. Singing groups for people with aphasia (SPA) • Time horizon: 6-months follow-up | £399 ^(c) | <p>0.05 QALYs</p> <p>From clinical review (2 vs 1) – same paper:</p> <p>EQ-5D-5L scores^(d) 3-month follow-up: -0.04 6-month follow-up: 0.10</p> <p>Carer generated health-related quality of life (CarerQoL-7D):^(d) 3-month follow-up: NR 6-month follow-up: 0.00</p> <p>Other outcomes were reported and can be seen in clinical evidence table.</p> | £7,980 per QALY gained. ^(e) | <p>Cost CI: not reported</p> <p>EQ-5D-5L CI: 3-month follow-up: -0.21 to 0.12 6-month follow-up: -0.06 to 0.26</p> <p>CarerQoL-7D CI: 6-month follow-up: -1.97 to 1.97</p> <p>No sensitivity analyses undertaken.</p> |

Abbreviations: CI = 95% confidence interval; ICER= incremental cost-effectiveness ratio; QALY= quality-adjusted life years; RCT= randomised controlled trial

- (a) Mean EQ-5D-5L scores (UK tariff) at 6-months were used to calculate the cost per QALY gained for this review: the NICE reference case currently prefers EQ5D-3L. It is not stated that an NHS and PSS perspective is taken however, the costs included are all considered relevant if the intervention is funded by the NHS.*
- (b) 6-month time horizon may not capture full health benefits of the intervention if these persist. Pilot feasibility RCT (n=41) that was not powered to test the effectiveness of the intervention or differences in healthcare resource use; the aim was to inform a future study where effectiveness and cost-effectiveness could be assessed. Within-trial analysis only reflects health outcomes and resource use from single trial; however, as this was the only trial included in the clinical review for music interventions delivered by non-healthcare professionals and so reflects the best currently available evidence. It is unclear if all relevant costs are included; interventions costs were included but other healthcare resource use was collected but not included; other healthcare resource used was numerically higher with the interventions but authors note the study was not powered to detect differences and did not present these as costs. Sensitivity analysis was not performed.*
- (c) 2019 UK pounds. Cost components incorporated: training costs, staff time during training and delivery of the intervention, travel costs for facilitators and singing champions, course materials (song books, percussion instruments, badges, and flip charts), venue costs and refreshments.*
- (d) Mean difference taken from figures 60, 65 and 66 of guideline clinical review.*
- (e) Cost per QALY gained not reported but was estimated using 6-month EQ-5D-5L scores collected within the study and assuming no difference in mortality.*

1.1.9 Economic model

This area was not prioritised for new cost-effectiveness analysis.

1.1.10 Unit costs

Music therapy and music interventions require additional resource use compared to not providing such interventions related to staff time and equipment. Studies included in the clinical review reported varied resource use (see Table 1 for details) due to:

- Variation in the delivery of therapy sessions: studies based on music therapists delivering therapy reported both individual and group-based sessions, while all but one of the studies⁷ that delivered therapy using health care professionals (HCPs) reported sessions on an individual basis. Group therapy will be lower cost per person.
- Significant variation in the frequency and duration of music therapy delivered, with sessions ranging from 20-90 minutes for 1-5 days per week. In the included clinical studies music therapy was generally delivered for between 5 and 10 weeks.
- Additional equipment required as part of the intervention, such as instruments (particularly keyboards, percussion/melodic instruments), metronomes, digital audio interface programs, iPod Nanos, music tapes and mindfulness audio materials.
- Clinical setting, as most studies were conducted in either an inpatient setting or as part of outpatient follow-up rehabilitation care in hospital. Jeong 2007⁷ and Tarrant (2018,²¹ 2021²⁰) were the only two studies that were conducted in a community setting. Baylan 2020¹ provided materials for participants to carry out sessions in their own time.
- Music interventions delivered by non-healthcare professionals may be funded by the NHS.

Relevant unit costs are provided below to aid consideration of cost effectiveness.

Table 14: Unit costs of health care professionals who may be involved in delivering music therapy interventions

| Resource | Cost per working hour (hospital/community) ^(a) | Source |
|---|---|-------------------------|
| Band 6 Music therapist (Arts therapist (entry level) ^(b) | £52/£54 | PSSRU 2021 ⁸ |
| Band 7 Music Therapist (Art therapist) ^(b) | £63/£65 | |
| Band 6 PT/OT | £53/£55 | |
| Band 7 PT/OT | £64/£67 | |

Abbreviations: OT= occupational therapist; PT= physiotherapist; PSSRU= personal and social services research unit

(a) Costs per working hour include salary, salary oncosts, overheads (management and other non-care staff costs including administration and estates staff) and capital overheads. For PT/OT costs it also includes qualification costs; qualification costs are not included in music therapist costs (qualification costs are not reported by PSSRU for art therapists and music therapy does not qualify for NHS bursaries).

(b) The British Association for Music Therapy (BAMT) states that a music therapist falls under the title of Arts therapist as listed in the PSSRU (2021).

Music therapists can complete additional specialist training to become 'neurologically trained' and thus provide neurologic music therapy (NMT). The committee advised that this training typically consists of a short course and people would typically still be employed at band 6 or 7. Health care professionals that delivered music interventions in the included clinical studies were either a physiotherapist or an occupational therapist. One study¹⁰ also provided counselling by a licensed psychotherapist for both trial arms, while another had therapy materials delivered by an assistant psychologist¹ however, such staff types were not mentioned in any of the other studies.

1.1.11 Evidence statements

Effectiveness/Qualitative

Economic

One cost-utility analysis found that a singing group intervention for people with aphasia was cost-effective compared to usual care (£7,980 per QALY gained). This study was assessed as partially applicable with potentially serious limitations.

1.1.12 The committee's discussion and interpretation of the evidence

1.1.12.1. The outcomes that matter most

The committee included the following outcomes: person/participant and carer generic health-related quality of life, activities of daily living, psychological distress, stroke-specific Patient-Reported Outcome Measures, wellbeing scores, participation in leisure activities/social group scores and withdrawal due to adverse events. All outcomes were considered equally important for decision making and therefore have all been rated as critical. The committee noted that music therapy may have benefits in other outcomes, such as physical function, communication and cognition. The committee considered that the outcomes included (namely health-related quality of life, activities of daily living and stroke-specific Patient-Reported Outcome Measures) would also encompass any such benefits. The committee considered wellbeing scores and participation in leisure activities/social groups score as important to capture the holistic benefits that could be experienced by people participating in music interventions. The committee chose to investigate these outcomes at less than 6 months and more than and equal to 6 months, as they considered that there could be a difference in the short-term and long-term effects of the intervention.

All outcomes were reported in at least 1 study but were not given in others. The limited evidence produced an element of uncertainty, and the committee agreed that there was insufficient evidence to make a recommendation.

1.1.12.2 The quality of the evidence

Twenty one randomised controlled trials were included in the review. Evidence was available for the following comparisons:

- Neurologic music therapy delivered by trained music therapists compared to no treatment (2 studies)
- Music therapy delivered by trained music therapists compared to no treatment (7 studies)
- Music intervention delivered by healthcare professionals compared to passive music listening (1 study)
- Music intervention delivered by healthcare professionals compared to placebo music therapy (1 study)
- Music intervention delivered by healthcare professionals compared to no treatment (11 studies)
- Music intervention delivered by non-healthcare professionals compared to no treatment (1 study)

There was limited evidence comparing any intervention to comparators other than no treatment (including comparisons to other music interventions, passive music listening and placebo music therapy).

The evidence varied from moderate to very low quality, with the majority being of very low quality. Outcomes were commonly downgraded for risk of bias and imprecision. Risk of bias was commonly due to selection, performance, attrition and measurement bias. A significant number of studies had different baseline values for outcomes between the intervention and comparator study groups. The majority of studies had very small sample sizes, which contributed to the imprecision in the outcomes. In most cases, it was not possible to conduct a meta-analysis on outcomes as there was limited outcome data reported by the studies that was comparable enough to be meta-analysed. Where meta-analysis was possible, outcomes often had heterogenous results within studies where there was an insufficient number of studies to form valid subgroups. In these cases, outcomes were downgraded for inconsistency. Indirect evidence was uncommon, although 1 study reported a population that may have included people who did not have a stroke and so was downgraded for population indirectness.

The type of music therapy or music intervention varied between studies. This included rhythmic auditory cueing; interventions where music instruments are played (including clinical improvisation); receptive interventions in which participants listen to music; singing and music-based voice interventions; sonification; and combinations of these interventions. the above. For the most part, these interventions were offered as a part of music therapy or as music interventions delivered by non-music therapists. However, the majority of studies reporting rhythmic auditory cueing were delivered by physiotherapists or occupational therapists rather than music therapists.

The no-treatment comparison varied. This included scenarios where no additional therapy was offered to participants who did not receive music interventions, but also included studies where usual care was offered to both study arms (which could include physiotherapy, occupational therapy, speech therapy and psychological support) and therefore the only difference in care was the music intervention.

The committee concluded that the evidence was of low quality. They acknowledged the effects that the heterogenous baseline values and small sample sizes had on the quality rating and took this into consideration while interpreting the evidence. They noted the potential bias introduced by the baseline values between intervention and control arms made it difficult to interpret the evidence. Consequentially, they found it difficult to interpret the effectiveness of music therapy and music interventions based on the evidence currently available.

1.1.12.3 Benefits and harms

1.1.12.3.1 Neurologic music therapy delivered by trained music therapists

The results showed that, when compared to no treatment, there were clinically important benefits in some subscales for person/participant generic health-related quality of life (namely SF-36 vitality and mental health) and psychological distress – depression at less than 6 months, but otherwise no clinically important difference in other subscales for health-related quality of life, psychological distress - anxiety and in stroke-specific Patient-Reported Outcome Measures at less than 6 months.

These outcomes were reported in 2 small studies with the outcomes being of low to very low quality. With this, the committee acknowledged that the evidence in this area was limited and insufficient to make a recommendation for neurologic music therapy. However, they noted the possible benefits in the intervention and made a research recommendation with the aim to gain more high-quality evidence. This should involve a large number of participants and where there were comparisons to active interventions that provide an equal intensity of therapy to those received from a music intervention, and placebo music therapy.

1.1.12.3.2 Music therapy delivered by trained music therapists

The results showed that, when compared to no treatment, there was a clinically important benefit in participation in leisure activities/social groups (based on 1 very small study with 18 participants). There were inconsistent effects seen in activities of daily living and stroke-specific Patient-Reported Outcome Measures with some outcomes showing clinically important benefits, others showing no clinically important difference and others showing clinically important harms. No clinically important difference was seen in person/participant generic health-related quality of life, psychological distress and withdrawal due to adverse events. No outcomes were reported at more than and equal to 6 months.

The evidence came from several small studies (the largest number of participants included in an outcome was 84) with the majority being of very low quality. With this, the committee acknowledged that the evidence in this area was limited. While there were more studies reporting music therapy than neurologic music therapy, the studies reported a range of different outcome measures in small trials that were probably not powered to show reliable changes in outcomes. Studies included intervention and control arms where the baseline values of outcomes were different at the start of the trial, making interpretation difficult. These trials were conducted comparing music therapy to no treatment (or usual care provided in both study arms), with no trials comparing music therapy to an intervention with equal contact with a professional to help show whether it is the music intervention that provides benefit or the interaction with the healthcare professional. These trials were less than 6 months duration with no long-term evidence being available. Based on this, the committee decided that the evidence was insufficient to make a recommendation for music therapy. However, they noted the possible benefits in the intervention and made a research recommendation with the aim to gain more high-quality research. This should involve a large number of participants and include comparisons to active interventions that provide an equal intensity of therapy to those received from a music intervention, or to placebo music therapy.

1.1.12.3.3 Music interventions delivered by healthcare professionals

Evidence was available comparing music interventions delivered by healthcare professionals to passive music listening, placebo music therapy and no treatment. All the outcomes comparing to passive music listening and placebo music therapy were reported in 1 study. When compared to passive music listening, there was a clinically important increase in withdrawal due to adverse events in those receiving a music intervention delivered by healthcare professionals at less than and more than and equal to 6 months (observed in one small study). This was also seen when compared to placebo music therapy. Otherwise, no clinically important difference was seen between music interventions delivered by healthcare professionals and placebo music therapy in psychological distress at less than and more than and equal to 6 months and participation in leisure activities/social group scores at less than 6 months only.

When compared to no treatment, clinically important benefits were seen in some subscales of person/participant health-related quality of life (namely SF-36 physical function, bodily pain, vitality, general health, role emotional, mental health and social function) while other measures showed no clinically important difference (McGill Quality of life) and other subscales showed clinically important harms (SF-36 role physical). Otherwise clinically important benefits were seen in activities of daily living and psychological distress (depression scores). No clinically important difference was seen in psychological distress (positive affect score), stroke-specific Patient-Reported Outcome Measures and withdrawal due to adverse events. Outcomes were only reported at less than 6 months for this comparison. The outcomes were reported in a range of different studies, with some outcomes including a larger number of participants while others had a very small number. However, the majority of evidence was of very low quality.

With this taken into account, committee acknowledged that the evidence in this area was limited. While there were more studies reporting music interventions delivered by healthcare

professionals than other interventions, the studies reported a range of different outcome measures in small trials that were likely not sufficiently powered to show reliable changes in outcomes. Studies included intervention and control arms where the baseline values of outcomes were different at the start of the trial, making interpretation difficult. While there was 1 trial comparing music interventions to placebo music therapy and music listening, this was limited evidence and most comparisons studied the effect compared to no treatment (or usual care provided to both study arms). These trials were mostly performed at less than 6 months with limited long-term evidence being available. The committee noted that clinically important harms were seen in some outcomes (in particular, withdrawal due to adverse events). However, they acknowledged that due to the small sample sizes the effect on dichotomous outcomes may be overemphasised and that trials with a larger number of participants were critical for understanding this further. Based on this, the committee decided that the evidence was insufficient to make a recommendation for music interventions delivered by healthcare professionals. However, they noted the possible benefits in the intervention and made a research recommendation with the aim to gain more high-quality research. This should involve a large number of participants and include comparisons to active interventions that provide an equal intensity of therapy to those received from a music intervention, or to placebo music therapy.

1.1.12.3.4 Music interventions delivered by non-healthcare professionals

The results showed that, when compared to no treatment, there are clinically important benefits in person/participant generic health-related quality of life at more than and equal to 6 months. There were no clinically important differences in stroke-specific Patient-Reported Outcome Measures, wellbeing scores and participation in leisure activities/social group scores at less than and more than and equal to 6 months, and carer generic health-related quality of life at more than and equal to 6 months only. There were clinically important harms in person/participant health-related quality of life at less than 6 months and withdrawal due to adverse events at less than 6 months. These outcomes were reported in 1 small study (with ≤ 41 participants) with the majority of outcomes being very low quality. Taking this into account, the committee agreed that the evidence was limited.

On examining the effect on person/participant generic health-related quality of life, the committee thought that the small study size considered that it could be possible that people with stroke may feel apprehensive at the start of the trial and may not engage more with the singing group until later on, which may have an effect on their initial quality of life results. However, the committee acknowledged the wide confidence intervals showing very serious imprecision in the outcomes, which affected their confidence in the results. Due to the limitations in the evidence, the committee decided not to recommend music interventions delivered by non-healthcare professionals. However, they noted the possible benefits in the intervention and made a research recommendation with the aim to gain more high-quality research. This should involve a large number of participants and include comparisons to active interventions that provide an equal intensity of therapy to those received from a music intervention, or to placebo music therapy.

1.1.12.4 Cost effectiveness and resource use

No health economic studies were included that related to either music therapy (including neurologic music therapy) delivered by trained music therapists or music interventions delivered by healthcare professionals.

The review identified one health economic analysis that compared a music intervention (singing for people with aphasia (SPA)) delivered by non-healthcare professionals to no treatment. This was a within-trial cost-utility analysis of a pilot feasibility RCT which was included in the clinical review. The intervention lasted 10 weeks and involved 1.5-hour group sessions once a week which were led by a music facilitator and assisted by an individual with post-stroke aphasia. The trial was designed to assess feasibility of a trial to assess

effectiveness and cost effectiveness and so had a small sample size (n=41) and was not powered to test the effectiveness of the SPA intervention.

Only intervention costs were considered. It was not stated that an NHS and PSS perspective was taken, however, the costs included are all considered relevant if the intervention is funded by the NHS. A micro-costing approach was adopted to estimate the intervention costs associated with SPA, taken from trial notes on staffing, purchases of equipment and venue costs charged by the sites. The authors then costed the staff at equivalent grades to the NHS PSSRU to show the costs that would be borne to the NHS if it were to provide these. The results found that the average cost of the intervention per participant was £399 including training costs, based on 2019-unit costs. Data on other healthcare resource use was collected but not included in cost calculations; other healthcare resource used was numerically higher with the intervention although the authors note the study was not powered to detect differences (as 1 of the objectives of the pilot study was to assess feasibility of collection).

A negative effect (-0.04) on quality of life (EQ-5D-5L) was found for participants at 3 months compared to the control group, and an improvement was found (0.10) at 6 months. Treatment effects beyond 6 months were not assessed. No change in carer quality of life (CarerQoL 7-D) was reported. Cost per QALY not reported but was estimated for this review to be £7,980 per QALY gained using 6-month EQ-5D-5L scores collected within the study and assuming no difference in mortality. This suggests that the intervention was cost-effective, however, there is uncertainty around these results as the confidence intervals for the quality-of-life follow-up estimates span across positive and negative values. As such, the committee were cautious in interpreting the results.

The study was assessed as partially applicable as mean EQ-5D-5L scores (UK tariff) at 6-months were used to calculate the cost per QALY gained for this review when the NICE reference case currently prefers EQ-5D-3L. Potentially serious limitations were noted for this study due to the small sample size and the fact that it was not powered to test the effectiveness of the music intervention or confirm differences in healthcare resource use between the groups, uncertainty around whether all relevant costs have been included, and uncertainty about long term treatment effects. Sensitivity analyses were also not performed. The committee felt that the study population (adults with post-stroke aphasia) was too specific to reflect the entire stroke population. Previous committee discussions noted that music therapy could potentially be useful to a broad range of people post-stroke but acknowledged that in practice it may be people with a higher level of disability it is used for.

In addition to this study, relevant unit costs were presented to the committee to aid consideration of cost effectiveness of neurologic or standard music therapy delivered by trained music therapists and music interventions delivered by healthcare professionals. Music therapy and interventions require additional resource use related to staff time and equipment. Studies included in the clinical review reported varied resource use, owing to a few factors such as the delivery of therapy sessions (either individual and group-based); the frequency and duration of music therapy delivered (with sessions ranging from 20 to 90 minutes for 1 to 5 days per week for between 5 and 10 weeks); additional equipment (for example, instruments) required as part of the intervention; clinical setting (most reported an inpatient setting or hospital-based outpatient follow-up) and interventions delivered by non-healthcare professionals. The heterogeneity of the interventions reported in the clinical evidence made it challenging for the committee to confidently assess the resource impact of providing these interventions nationwide. Staff costs, however, were found to be similar for healthcare professionals who may be involved in delivering music therapy or interventions. Although neurologic music therapists complete additional specialist training to become 'neurologically trained', the committee advised that this training typically consists of a short course and people would typically still be employed at band 6 or 7. Healthcare professionals who delivered music interventions in the included clinical studies were either a physiotherapist or an occupational therapist also be employed at band 6 or 7. One study

(Kim, 2011) also provided counselling by a licensed psychotherapist for both trial arms, while another had therapy materials delivered by an assistant psychologist (Baylan, 2020), however, such staff roles were not mentioned in any other studies. The committee noted that music interventions delivered by the rehabilitation team and non-music therapists are used in current practice, but that music therapy delivered by music therapists and neurologic music therapy are not widely available in the NHS.

The committee discussed the clinical and economic evidence and, based on the limitations described in the clinical evidence section and the uncertainty in the economic evidence, were not able to make recommendations about which music interventions may be appropriate for people following a stroke. A research recommendation has been made.

1.1.12.5 Other factors the committee took into account

The committee acknowledged that benefits may be seen in outcomes specific to impairments that were not included in the protocol (for example: physical function, communication, cognition). A Cochrane review¹³ investigating music therapy for people with acquired brain injury had identified additional evidence showing benefit for these outcomes. When designing the protocol, the committee prioritised functional outcomes over impairment-based outcomes which meant that those were not identified. The committee considered that there could be additional benefits during their deliberation.

The committee highlighted that additional therapies may be present that incorporate sound and could therefore be beneficial for people after stroke (for example: sound therapy). While this was not investigated during this guideline update, this was highlighted as a potential area that could be beneficial for people to consider.

Members of the committee also spoke about their own personal experiences of music therapy, stating it significantly improved both their quality of life and that of their family members. They highlighted this as an important area that required consideration in the future. The committee believe that future studies which are larger and more rigorous than those currently available, should be conducted in this area so that a complete understanding of the intervention can be obtained.

It was noted that music therapy interventions may be delivered outside of NHS services by third sector organisations, such as charities (either as outsourcing of services by the NHS or outside of formal care). The involvement of third sector organisations was emphasised by the committee as important for the delivery of interventions in this area.

1.1.13 Recommendations supported by this evidence review

This evidence review supports the research recommendation on music therapy in Appendix K. No recommendations were made for this review.

1.1.14 References

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Appendices

Appendix A – Review protocols

Review protocol for the clinical and cost-effectiveness of music therapy after a stroke

| ID | Field | Content |
|----|------------------------------|---|
| 0. | PROSPERO registration number | CRD42021249939 |
| 1. | Review title | In people after stroke, what is the clinical and cost effectiveness of music therapy to improve mood and activities of daily living? |
| 2. | Review question | 4.8 In people after stroke, what is the clinical and cost effectiveness of music therapy to improve mood and activities of daily living? |
| 3. | Objective | To determine the clinical and cost-effectiveness of music therapy in improving mood and activities of daily living for people after a stroke. |
| 4. | Searches | <p>The following databases (from inception) will be searched:</p> <ul style="list-style-type: none"> • Cochrane Central Register of Controlled Trials (CENTRAL) • Cochrane Database of Systematic Reviews (CDSR) • Embase • MEDLINE • BAMT Register of Surveys, Research and Evaluation Projects (ROSREP) • CAMbase • Music Therapy World Journal Index <p>Searches will be restricted by:</p> <ul style="list-style-type: none"> • English language studies • Human studies <p>Other searches:</p> <ul style="list-style-type: none"> • Inclusion lists of systematic reviews <p>The searches may be re-run 6 weeks before the final committee meeting and further studies retrieved for inclusion if relevant.</p> <p>The full search strategies will be published in the final review.</p> |

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| | | Medline search strategy to be quality assured using the PRESS evidence-based checklist (see methods chapter for full details). |
| 5. | Condition or domain being studied | Adults and young people (16 or older) after a stroke |
| 6. | Population | <p>Inclusion:</p> <ul style="list-style-type: none"> Adults (age ≥ 16 years) who have had a first stroke or recurrent stroke (including people after a subarachnoid haemorrhage) <p>Exclusion:</p> <ul style="list-style-type: none"> Children (age < 16 years) People who have had a transient ischaemic attack |
| 7. | Intervention | <ul style="list-style-type: none"> Neurologic music therapy delivered by trained music therapists Music therapy delivered by trained music therapists Music interventions delivered by healthcare professionals Music interventions delivered by non-healthcare professionals <p>The interventions will be analysed as separate stratifications.</p> |
| 8. | Comparator | <ul style="list-style-type: none"> Compared to each other Passive music listening (for example: music played in the background) Placebo music therapy No treatment <p>Each comparator will be analysed in separate stratifications by different types of comparators.</p> |
| 9. | Types of study to be included | <ul style="list-style-type: none"> Systematic reviews of RCTs Parallel RCTs Cluster randomised crossover trials (unit of randomisation = unit [for example: hospital, community location]) <p>If there is insufficient randomised trial evidence, non-randomised studies (prospective and retrospective cohort trials) will be considered after discussion with the committee.</p> <p>Published NMAs and IPDs will be considered for inclusion.</p> |

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| 10. | Other exclusion criteria | <ul style="list-style-type: none"> • Non-English language studies • Crossover RCTs (unit of randomisation = participant) <p>Conference abstracts will be excluded as it is expected there will be sufficient full text published studies available.</p> |
| 11. | Context | <p>People after a stroke. This may include people in an acute (<7 days), subacute (7 days – 6 months) or chronic (>6 months) time horizon.</p> |
| 12. | Primary outcomes (critical outcomes) | <p>All outcomes are considered equally important for decision making and therefore have all been rated as critical:</p> <p>At time period</p> <ul style="list-style-type: none"> • <6 months • ≥6 months <ul style="list-style-type: none"> • Person/participant generic health-related quality of life (continuous outcomes will be prioritised [validated measures]) <ul style="list-style-type: none"> ○ EQ-5D ○ SF-6D ○ SF-36 ○ SF-12 ○ Other utility measures (AQOL, HUI, 15D, QWB) • Carer generic health-related quality of life (continuous outcomes will be prioritised [validated measures]) <ul style="list-style-type: none"> ○ EQ-5D ○ SF-6D ○ SF-36 ○ SF-12 ○ Other utility measures (AQOL, HUI, 15D, QWB) • Activities of daily living (continuous outcomes will be prioritised) <ul style="list-style-type: none"> ○ Barthel Index ○ National Institutes of Health Stroke Scale ○ Orpington Prognostic Scale ○ Canadian Occupational Performance Measure ○ Extended activities of daily living • Psychological distress (continuous outcomes will be prioritised) <ul style="list-style-type: none"> ○ Depression <ul style="list-style-type: none"> – PHQ-9 – Hospital Anxiety and Depression scale - depression subscale – Beck Depression Inventory – Hamilton Depression Scale |

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| | | <ul style="list-style-type: none"> - Centre of Epidemiologic Studies Depression - GHQ-28 - Geriatric Depression Scale o Anxiety <ul style="list-style-type: none"> - GAD-7 - Hospital Anxiety and Depression scale - anxiety subscale - The Geriatric Anxiety Inventory - GHQ-28 - Beck Anxiety Inventory o Distress <ul style="list-style-type: none"> - The Distress Management System for Stroke (DMSS) • Stroke-specific Patient-Reported Outcome Measures (continuous outcomes will be prioritised) <ul style="list-style-type: none"> o Stroke-Specific Quality of Life (SS-QOL) o Stroke Impact Scale (SIS) o Stroke-specific Sickness Impact Profile (SA-SIP30) o Neuro-QOL o PROMIS-10 o Satisfaction with International Classification of Functioning, Disability and Health – Stroke (SATIS-Stroke) • Wellbeing scores (continuous outcomes will be prioritised) <ul style="list-style-type: none"> o Warwick-Edinburgh Mental wellbeing scale o WHO-5 World Health Organisation Wellbeing Index o Rosenberg Self Esteem Scale o ICEpop CAPability measure for adults • Participation in leisure activities/social groups scores (continuous outcomes will be prioritised) <ul style="list-style-type: none"> o Mayo-Portland Adaptability Inventory 4 (MPAI-4) part C (participation) o Frenchay Activities Index • Withdrawal due to adverse events (dichotomous outcome) <p>If not mentioned above, other validated scores will be considered and discussed with the committee to deliberate on their inclusion.</p> |
| 14. | Data extraction (selection and coding) | All references identified by the searches and from other sources will be uploaded into EPPI reviewer and de-duplicated. |

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| | | <p>10% of the abstracts will be reviewed by two reviewers, with any disagreements resolved by discussion or, if necessary, a third independent reviewer.</p> <p>The full text of potentially eligible studies will be retrieved and will be assessed in line with the criteria outlined above.</p> <p>A standardised form will be used to extract data from studies (see Developing NICE guidelines: the manual section 6.4).</p> <p>10% of all evidence reviews are quality assured by a senior research fellow. This includes checking:</p> <ul style="list-style-type: none"> • papers were included /excluded appropriately • a sample of the data extractions • correct methods are used to synthesise data • a sample of the risk of bias assessments <p>Disagreements between the review authors over the risk of bias in particular studies will be resolved by discussion, with involvement of a third review author where necessary.</p> <p>Study investigators may be contacted for missing data where time and resources allow.</p> |
| 15. | Risk of bias (quality) assessment | <p>Risk of bias will be assessed using the appropriate checklist as described in Developing NICE guidelines: the manual.</p> <ul style="list-style-type: none"> • Systematic reviews: Risk of Bias in Systematic Reviews (ROBIS) • Randomised Controlled Trial: Cochrane RoB (2.0) • Non randomised study, including cohort studies: Cochrane ROBINS-I |
| 16. | Strategy for data synthesis | <ul style="list-style-type: none"> • Pairwise meta-analyses will be performed using Cochrane Review Manager (RevMan5). Fixed-effects (Mantel-Haenszel) techniques will be used to calculate risk ratios for the binary outcomes where possible. Continuous outcomes will be analysed using an inverse variance method for pooling weighted mean differences. <p>Heterogeneity between the studies in effect measures will be assessed using the I^2 statistic and visually inspected. An I^2 value greater than 50% will be considered indicative of substantial heterogeneity. Sensitivity analyses will be conducted based on pre-specified subgroups using stratified meta-analysis to explore the heterogeneity in effect estimates. If this does not</p> |

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| | | <p>explain the heterogeneity, the results will be presented pooled using random-effects.</p> <ul style="list-style-type: none"> • GRADEpro will be used to assess the quality of evidence for each outcome, taking into account individual study quality and the meta-analysis results. The 4 main quality elements (risk of bias, indirectness, inconsistency and imprecision) will be appraised for each outcome. Publication bias is tested for when there are more than 5 studies for an outcome. <p>The risk of bias across all available evidence was evaluated for each outcome using an adaptation of the 'Grading of Recommendations Assessment, Development and Evaluation (GRADE) toolbox' developed by the international GRADE working group http://www.gradeworkinggroup.org/</p> <ul style="list-style-type: none"> • Where meta-analysis is not possible, data will be presented and quality assessed individually per outcome. • WinBUGS will be used for network meta-analysis, if possible given the data identified. |
| 17. | Analysis of sub-groups | <p>Subgroups that will be investigated if heterogeneity is present:</p> <p>Time after stroke at the start of the trial</p> <ul style="list-style-type: none"> • Hyperacute <72 hours • Acute 72 hours – 7 days • Subacute 7 days – 6 months • Chronic >6 months <p>Severity (as stated by category or as measured by NIHSS scale):</p> <ul style="list-style-type: none"> • Mild (or NIHSS 1-5) • Moderate (or NIHSS 5-14) • Severe (or NIHSS 15-24) • Very severe (or NIHSS >25) <p>Group compared to individual sessions of music therapy:</p> <ul style="list-style-type: none"> • Group sessions • Individual sessions • Mixed (programmes including group and individual) <p>Delivered in hospital compared to delivered in the community:</p> <ul style="list-style-type: none"> • Hospital sessions • Community sessions • Mixed (programmes including both hospital and community sessions) |

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| | | <p>Types of intervention:</p> <ul style="list-style-type: none"> • Interventions where musical instruments are played (including clinical improvisation) • Singing and music-based voice interventions • Rhythmic auditory cueing • Receptive interventions in which participants listen to music • Songwriting • Any combination of the above | | |
| 18. | Type and method of review | <input checked="" type="checkbox"/> | Intervention | |
| | | <input type="checkbox"/> | Diagnostic | |
| | | <input type="checkbox"/> | Prognostic | |
| | | <input type="checkbox"/> | Qualitative | |
| | | <input type="checkbox"/> | Epidemiologic | |
| | | <input type="checkbox"/> | Service Delivery | |
| | | <input type="checkbox"/> | Other (please specify) | |
| 19. | Language | English | | |
| 20. | Country | England | | |
| 21. | Anticipated or actual start date | 24/02/2021 | | |
| 22. | Anticipated completion date | 14/12/2022 | | |
| 23. | Stage of review at time of this submission | Review stage | Started | Completed |
| | | Preliminary searches | <input type="checkbox"/> | <input type="checkbox"/> |
| | | Piloting of the study selection process | <input type="checkbox"/> | <input type="checkbox"/> |
| | | Formal screening of search results against eligibility criteria | <input type="checkbox"/> | <input type="checkbox"/> |
| | | Data extraction | <input type="checkbox"/> | <input type="checkbox"/> |
| | | Risk of bias (quality) assessment | <input type="checkbox"/> | <input type="checkbox"/> |
| | | Data analysis | <input type="checkbox"/> | <input type="checkbox"/> |
| 24. | Named contact | <p>5a. Named contact National Guideline Centre</p> <p>5b Named contact e-mail</p> | | |

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| | | <p>StrokeRehabUpdate@nice.nhs.uk</p> <p>5e Organisational affiliation of the review National Institute for Health and Care Excellence (NICE) and National Guideline Centre</p> |
| 25. | Review team members | <p>From the National Guideline Centre:</p> <p>Bernard Higgins (Guideline lead)</p> <p>George Wood (Senior systematic reviewer)</p> <p>Madelaine Zucker (Systematic reviewer)</p> <p>Kate Lovibond (Health economics lead)</p> <p>Claire Sloan (Health economist)</p> <p>Joseph Runicles (Information specialist)</p> <p>Nancy Pursey (Senior project manager)</p> |
| 26. | Funding sources/sponsor | <p>This systematic review is being completed by the National Guideline Centre which receives funding from NICE.</p> |
| 27. | Conflicts of interest | <p>All guideline committee members and anyone who has direct input into NICE guidelines (including the evidence review team and expert witnesses) must declare any potential conflicts of interest in line with NICE's code of practice for declaring and dealing with conflicts of interest. Any relevant interests, or changes to interests, will also be declared publicly at the start of each guideline committee meeting. Before each meeting, any potential conflicts of interest will be considered by the guideline committee Chair and a senior member of the development team. Any decisions to exclude a person from all or part of a meeting will be documented. Any changes to a member's declaration of interests will be recorded in the minutes of the meeting. Declarations of interests will be published with the final guideline.</p> |
| 28. | Collaborators | <p>Development of this systematic review will be overseen by an advisory committee who will use the review to inform the development of evidence-based recommendations in line with section 3 of Developing NICE guidelines: the manual. Members of the guideline committee are available on the NICE website: https://www.nice.org.uk/guidance/indevelopment/gid-ng10175</p> |
| 29. | Other registration details | N/A |
| 30. | Reference/URL for published protocol | N/A |
| 31. | Dissemination plans | <p>NICE may use a range of different methods to raise awareness of the guideline. These include standard approaches such as:</p> <ul style="list-style-type: none"> • notifying registered stakeholders of publication |

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| | | <ul style="list-style-type: none"> • publicising the guideline through NICE's newsletter and alerts • issuing a press release or briefing as appropriate, posting news articles on the NICE website, using social media channels, and publicising the guideline within NICE. |
| 32. | Keywords | Adults; Intervention; Music therapy; Rehabilitation; Rhythmic auditory cueing; Stroke |
| 33. | Details of existing review of same topic by same authors | N/A |
| 34. | Current review status | <input type="checkbox"/> Ongoing |
| | | <input type="checkbox"/> Completed but not published |
| | | <input checked="" type="checkbox"/> Completed and published |
| | | <input type="checkbox"/> Completed, published and being updated |
| | | <input type="checkbox"/> Discontinued |
| 35.. | Additional information | N/A |
| 36. | Details of final publication | www.nice.org.uk |

Review protocol for health economic literature review

| Review question | All questions – health economic evidence |
|------------------------|---|
| Objectives | To identify health economic studies relevant to any of the review questions. |
| Search criteria | <ul style="list-style-type: none"> • Populations, interventions and comparators must be as specified in the clinical review protocol above. • Studies must be of a relevant health economic study design (cost–utility analysis, cost-effectiveness analysis, cost–benefit analysis, cost–consequences analysis, comparative cost analysis). • Studies must not be a letter, editorial or commentary, or a review of health economic evaluations. (Recent reviews will be ordered although not reviewed. The bibliographies will be checked for relevant studies, which will then be ordered.) • Unpublished reports will not be considered unless submitted as part of a call for evidence. • Studies must be in English. |
| Search strategy | <p>A health economic study search will be undertaken using population-specific terms and a health economic study filter – see appendix B below.</p> <p>Databases searched:</p> <ul style="list-style-type: none"> • Centre for Reviews and Dissemination NHS Economic Evaluations Database (NHS EED) – all years (closed to new records April 2015) • Centre for Reviews and Dissemination Health Technology Assessment database – all years (closed to new records March 2018) • International HTA database (INAHTA) – all years • Medline and Embase – from 2014 (due to NHS EED closure) |
| Review strategy | <p>Studies not meeting any of the search criteria above will be excluded. Studies published before 2006 (including those included in the previous guideline), abstract-only studies and studies from non-OECD countries or the USA will also be excluded.</p> <p>Each remaining study will be assessed for applicability and methodological limitations using the NICE economic evaluation checklist which can be found in appendix H of Developing NICE guidelines: the manual (2014)¹⁴</p> <p>Studies published in 2006 or later that were included in the previous guideline will be reassessed for inclusion and may be included or selectively excluded based on their relevance to the questions covered in this update and whether more applicable evidence is also identified.</p> <p>Inclusion and exclusion criteria</p> <ul style="list-style-type: none"> • If a study is rated as both ‘Directly applicable’ and with ‘Minor limitations’ then it will be included in the guideline. A health economic evidence table will be completed and it will be included in the health economic evidence profile. • If a study is rated as either ‘Not applicable’ or with ‘Very serious limitations’ then it will usually be excluded from the guideline. If it is excluded then a health economic evidence table will not be completed and it will not be included in the health economic evidence profile. • If a study is rated as ‘Partially applicable’, with ‘Potentially serious limitations’ or both then there is discretion over whether it should be included. <p>Where there is discretion</p> <p>The health economist will make a decision based on the relative applicability and quality of the available evidence for that question, in discussion with the guideline committee if required. The ultimate aim is to include health economic studies that are helpful for decision-making in the context of the guideline and the current NHS setting. If several studies are considered of sufficiently high applicability and methodological quality that they could all be included, then the health economist, in</p> |

discussion with the committee if required, may decide to include only the most applicable studies and to selectively exclude the remaining studies. All studies excluded on the basis of applicability or methodological limitations will be listed with explanation in the excluded health economic studies appendix below.

The health economist will be guided by the following hierarchies.

Setting:

- UK NHS (most applicable).
- OECD countries with predominantly public health insurance systems (for example, France, Germany, Sweden).
- OECD countries with predominantly private health insurance systems (for example, Switzerland).
- Studies set in non-OECD countries or in the USA will be excluded before being assessed for applicability and methodological limitations.

Health economic study type:

- Cost–utility analysis (most applicable).
- Other type of full economic evaluation (cost–benefit analysis, cost-effectiveness analysis, cost–consequences analysis).
- Comparative cost analysis.
- Non-comparative cost analyses including cost-of-illness studies will be excluded before being assessed for applicability and methodological limitations.

Year of analysis:

- The more recent the study, the more applicable it will be.
- Studies published in 2006 or later (including any such studies included in the previous guideline) but that depend on unit costs and resource data entirely or predominantly from before 2006 will be rated as 'Not applicable'.
- Studies published before 2006 (including any such studies included in the previous guideline) will be excluded before being assessed for applicability and methodological limitations.

Quality and relevance of effectiveness data used in the health economic analysis:

- The more closely the clinical effectiveness data used in the health economic analysis match with the outcomes of the studies included in the clinical review the more useful the analysis will be for decision-making in the guideline.

Appendix B – Literature search strategies

B.1 Clinical search literature search strategy

Searches were constructed using a PICO framework where population (P) terms were combined with Intervention (I) and in some cases Comparison (C) terms. Outcomes (O) are rarely used in search strategies as these concepts may not be indexed or described in the title or abstract and are therefore difficult to retrieve. Search filters were applied to the search where appropriate.

Table 15: Database parameters, filters and limits applied

| Database | Dates searched | Search filter used |
|--|---|---|
| Medline (OVID) | Inception – 08 January 2023 | Randomised controlled trials Systematic review studies Exclusions (animal studies, letters, comments, editorials, case studies/reports) English language |
| Embase (OVID) | Inception – 08 January 2023 | Randomised controlled trials Systematic review studies Exclusions (animal studies, letters, comments, editorials, case studies/reports, conference abstracts) English language |
| The Cochrane Library (Wiley) | Cochrane Reviews to 2023 Issue 1 of 12 CENTRAL to 2023 Issue 1 of 12 | Exclusions (clinical trials, conference abstracts) |
| CAMbase (Complementary and Alternative Medicine) | Inception – 08 January 2023 | English language |

Medline (Ovid) search terms

| | |
|----|--|
| 1. | exp Stroke/ |
| 2. | Stroke Rehabilitation/ |
| 3. | exp Cerebral Hemorrhage/ |
| 4. | (stroke or strokes or cva or poststroke* or apoplexy or "cerebrovascular accident").ti,ab. |
| 5. | ((cerebro* or brain or brainstem or cerebral*) adj3 (infarct* or accident*)).ti,ab. |
| 6. | "brain attack*".ti,ab. |

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| 7. | or/1-6 |
| 8. | letter/ |
| 9. | editorial/ |
| 10. | news/ |
| 11. | exp historical article/ |
| 12. | Anecdotes as Topic/ |
| 13. | comment/ |
| 14. | case report/ |
| 15. | (letter or comment*).ti. |
| 16. | or/8-15 |
| 17. | randomized controlled trial/ or random*.ti,ab. |
| 18. | 16 not 17 |
| 19. | animals/ not humans/ |
| 20. | exp Animals, Laboratory/ |
| 21. | exp Animal Experimentation/ |
| 22. | exp Models, Animal/ |
| 23. | exp Rodentia/ |
| 24. | (rat or rats or mouse or mice or rodent*).ti. |
| 25. | or/18-24 |
| 26. | 7 not 25 |
| 27. | limit 26 to English language |
| 28. | randomized controlled trial.pt. |
| 29. | controlled clinical trial.pt. |
| 30. | randomi#ed.ti,ab. |
| 31. | placebo.ab. |
| 32. | randomly.ti,ab. |
| 33. | Clinical Trials as topic.sh. |
| 34. | trial.ti. |
| 35. | or/28-34 |
| 36. | Meta-Analysis/ |
| 37. | exp Meta-Analysis as Topic/ |
| 38. | (meta analy* or metanaly* or metaanaly* or meta regression).ti,ab. |
| 39. | ((systematic* or evidence*) adj3 (review* or overview*)).ti,ab. |
| 40. | (reference list* or bibliograph* or hand search* or manual search* or relevant journals).ab. |
| 41. | (search strategy or search criteria or systematic search or study selection or data extraction).ab. |
| 42. | (search* adj4 literature).ab. |
| 43. | (medline or pubmed or cochrane or embase or psychlit or psyclit or psychinfo or psycinfo or cinahl or science citation index or bids or cancerlit).ab. |
| 44. | cochrane.jw. |
| 45. | ((multiple treatment* or indirect or mixed) adj2 comparison*).ti,ab. |
| 46. | or/36-45 |
| 47. | music/ or music therapy/ or singing/ or acoustic stimulation/ |
| 48. | (music* or rhythm* or melod* or harmon* or sonification).ti,ab. |

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| 49. | ((auditory or acoustic or sound*) adj5 (stimulat* or cue* or treatment* or rehab* or intervention*)).ti,ab. |
| 50. | (sing or sings or singing or singer or song* or chant* or compose or composing or improvis*).ti,ab. |
| 51. | ((vocal or voice) adj5 intonat*).ti,ab. |
| 52. | (play* adj2 instrument*).ti,ab. |
| 53. | ((passive or passively) adj2 listen*).ti,ab. |
| 54. | or/47-53 |
| 55. | 27 and 54 |
| 56. | 35 or 46 |
| 57. | 55 and 56 |

Embase (Ovid) search terms

| | |
|-----|--|
| 1. | exp Cerebrovascular accident/ |
| 2. | exp Brain infarction/ |
| 3. | Stroke Rehabilitation/ |
| 4. | (stroke or strokes or cva or poststroke* or apoplexy or "cerebrovascular accident").ti,ab. |
| 5. | ((cerebro* or brain or brainstem or cerebral*) adj3 (infarct* or accident*)).ti,ab. |
| 6. | "brain attack".ti,ab. |
| 7. | Intracerebral hemorrhage/ |
| 8. | or/1-7 |
| 9. | letter.pt. or letter/ |
| 10. | note.pt. |
| 11. | editorial.pt. |
| 12. | case report/ or case study/ |
| 13. | (letter or comment*).ti. |
| 14. | (conference abstract or conference paper).pt. |
| 15. | or/9-14 |
| 16. | randomized controlled trial/ or random*.ti,ab. |
| 17. | 15 not 16 |
| 18. | animal/ not human/ |
| 19. | nonhuman/ |
| 20. | exp Animal Experiment/ |
| 21. | exp Experimental Animal/ |
| 22. | animal model/ |
| 23. | exp Rodent/ |
| 24. | (rat or rats or mouse or mice or rodent*).ti. |
| 25. | or/17-24 |
| 26. | 8 not 25 |
| 27. | limit 26 to English language |
| 28. | random*.ti,ab. |
| 29. | factorial*.ti,ab. |
| 30. | (crossover* or cross over*).ti,ab. |
| 31. | ((doubl* or singl*) adj blind*).ti,ab. |

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| 32. | (assign* or allocat* or volunteer* or placebo*).ti,ab. |
| 33. | crossover procedure/ |
| 34. | single blind procedure/ |
| 35. | randomized controlled trial/ |
| 36. | double blind procedure/ |
| 37. | or/28-36 |
| 38. | systematic review/ |
| 39. | meta-analysis/ |
| 40. | (meta analy* or metanaly* or metaanaly* or meta regression).ti,ab. |
| 41. | ((systematic* or evidence*) adj3 (review* or overview*)).ti,ab. |
| 42. | (reference list* or bibliograph* or hand search* or manual search* or relevant journals).ab. |
| 43. | (search strategy or search criteria or systematic search or study selection or data extraction).ab. |
| 44. | (search* adj4 literature).ab. |
| 45. | (medline or pubmed or cochrane or embase or psychlit or psyclit or psychinfo or psycinfo or cinahl or science citation index or bids or cancerlit).ab. |
| 46. | cochrane.jw. |
| 47. | ((multiple treatment* or indirect or mixed) adj2 comparison*).ti,ab. |
| 48. | or/38-47 |
| 49. | music therapy/ or music/ or singing/ or auditory stimulation/ |
| 50. | (music* or rhythm* or melod* or harmon* or sonification).ti,ab. |
| 51. | ((auditory or acoustic or sound*) adj5 (stimulat* or cue* or treatment* or rehab* or intervention*)).ti,ab. |
| 52. | (sing or sings or singing or singer or song* or chant* or compose or composing or improvis*).ti,ab. |
| 53. | ((vocal or voice) adj5 intonat*).ti,ab. |
| 54. | (play* adj2 instrument*).ti,ab. |
| 55. | ((passive or passively) adj2 listen*).ti,ab. |
| 56. | or/49-55 |
| 57. | 27 and 56 |
| 58. | 57 and (37 or 48) |

Cochrane Library (Wiley) search terms

| | |
|------|---|
| #1. | MeSH descriptor: [Stroke] explode all trees |
| #2. | MeSH descriptor: [Stroke Rehabilitation] explode all trees |
| #3. | MeSH descriptor: [Cerebral Hemorrhage] explode all trees |
| #4. | (stroke or strokes or cva or poststroke* or apoplexy or "cerebrovascular accident"):ti,ab |
| #5. | ((cerebro* or brain or brainstem or cerebral*) near/3 (infarct* or accident*)):ti,ab |
| #6. | brain attack*:ti,ab |
| #7. | (or #1-#6) |
| #8. | conference:pt or (clinicaltrials or trialsearch):so |
| #9. | #7 not #8 |
| #10. | MeSH descriptor: [Music] explode all trees |
| #11. | MeSH descriptor: [Music Therapy] explode all trees |
| #12. | MeSH descriptor: [Singing] explode all trees |

| | |
|------|--|
| #13. | MeSH descriptor: [Acoustic Stimulation] explode all trees |
| #14. | (music* or rhythm* or melod* or harmon* or sonification):ti,ab |
| #15. | ((auditory or acoustic or sound*) near/5 (stimulat* or cue* or treatment* or rehab* or intervention*)):ti,ab |
| #16. | (sing or sings or singing or singer or song* or chant* or compose or composing or improvis*):ti,ab |
| #17. | ((vocal or voice) near/5 intonat*):ti,ab |
| #18. | (play* near/2 instrument*):ti,ab |
| #19. | ((passive or passively) near/2 listen*):ti,ab |
| #20. | (or #10-#19) |
| #21. | #9 and #20 |

CAMbase search terms

| | |
|----|---------------|
| 1. | Stroke |
| 2. | Music Therapy |
| 3. | 1 and 2 |

B.2 Health Economics literature search strategy

Health economic evidence was identified by conducting searches using terms for a broad Stroke Rehabilitation population. The following databases were searched: NHS Economic Evaluation Database (NHS EED - this ceased to be updated after 31st March 2015), Health Technology Assessment database (HTA - this ceased to be updated from 31st March 2018) and The International Network of Agencies for Health Technology Assessment (INAHTA). Searches for recent evidence were run on Medline and Embase from 2014 onwards for health economics, and all years for quality-of-life studies. Additional searches were run in CINAHL and PsycInfo looking for health economic evidence.

Table 2: Database parameters, filters and limits applied

| Database | Dates searched | Search filters and limits applied |
|----------------|--|--|
| Medline (OVID) | Health Economics 1 January 2014 – 08 January 2023 | Health economics studies Quality of life studies |
| | Quality of Life 1946 – 08 January 2023 | Exclusions (animal studies, letters, comments, editorials, case studies/reports,) English language |
| Embase (OVID) | Health Economics 1 January 2014 – 08 January 2023 | Health economics studies Quality of life studies |
| | Quality of Life 1974 – 08 January 2023 | Exclusions (animal studies, letters, comments, editorials, case studies/reports, conference abstracts) English language |

| Database | Dates searched | Search filters and limits applied |
|--|---|--|
| NHS Economic Evaluation Database (NHS EED) (Centre for Research and Dissemination - CRD) | Inception –31 st March 2015 | |
| Health Technology Assessment Database (HTA) (Centre for Research and Dissemination – CRD) | Inception – 31 st March 2018 | |
| The International Network of Agencies for Health Technology Assessment (INAHTA) | Inception - 08 January 2023 | English language |
| PsycINFO (OVID) | 1 January 2014 – 08 January 2023 | Health economics studies Exclusions (animal studies, letters, case reports) Human English language |
| Current Nursing and Allied Health Literature - CINAHL (EBSCO) | 1 January 2014 – 08 January 2023 | Health economics studies Exclusions (Medline records, animal studies, letters, editorials, comments, theses) Human English language |

Medline (Ovid) search terms

| | |
|-----|--|
| 1. | exp Stroke/ |
| 2. | exp Cerebral Hemorrhage/ |
| 3. | (stroke or strokes or cva or poststroke* or apoplexy or "cerebrovascular accident").ti,ab. |
| 4. | ((cerebro* or brain or brainstem or cerebral*) adj3 (infarct* or accident*)).ti,ab. |
| 5. | "brain attack*".ti,ab. |
| 6. | or/1-5 |
| 7. | letter/ |
| 8. | editorial/ |
| 9. | news/ |
| 10. | exp historical article/ |
| 11. | Anecdotes as Topic/ |
| 12. | comment/ |
| 13. | case report/ |
| 14. | (letter or comment*).ti. |

| | |
|-----|---|
| 15. | or/7-14 |
| 16. | randomized controlled trial/ or random*.ti,ab. |
| 17. | 15 not 16 |
| 18. | animals/ not humans/ |
| 19. | exp Animals, Laboratory/ |
| 20. | exp Animal Experimentation/ |
| 21. | exp Models, Animal/ |
| 22. | exp Rodentia/ |
| 23. | (rat or rats or mouse or mice or rodent*).ti. |
| 24. | or/17-23 |
| 25. | 6 not 24 |
| 26. | Economics/ |
| 27. | Value of life/ |
| 28. | exp "Costs and Cost Analysis"/ |
| 29. | exp Economics, Hospital/ |
| 30. | exp Economics, Medical/ |
| 31. | Economics, Nursing/ |
| 32. | Economics, Pharmaceutical/ |
| 33. | exp "Fees and Charges"/ |
| 34. | exp Budgets/ |
| 35. | budget*.ti,ab. |
| 36. | cost*.ti. |
| 37. | (economic* or pharmaco?economic*).ti. |
| 38. | (price* or pricing*).ti,ab. |
| 39. | (cost* adj2 (effective* or utilit* or benefit* or minimi* or unit* or estimat* or variable*)).ab. |
| 40. | (financ* or fee or fees).ti,ab. |
| 41. | (value adj2 (money or monetary)).ti,ab. |
| 42. | or/26-41 |
| 43. | quality-adjusted life years/ |
| 44. | sickness impact profile/ |
| 45. | (quality adj2 (wellbeing or well being)).ti,ab. |
| 46. | sickness impact profile.ti,ab. |
| 47. | disability adjusted life.ti,ab. |
| 48. | (qal* or qtime* or qwb* or daly*).ti,ab. |
| 49. | (euroqol* or eq5d* or eq 5*).ti,ab. |
| 50. | (qol* or hql* or hqol* or h qol* or hrqol* or hr qol*).ti,ab. |
| 51. | (health utility* or utility score* or disutilit* or utility value*).ti,ab. |
| 52. | (hui or hui1 or hui2 or hui3).ti,ab. |
| 53. | (health* year* equivalent* or hye or hyes).ti,ab. |

| | |
|-----|---|
| 54. | discrete choice*.ti,ab. |
| 55. | rosser.ti,ab. |
| 56. | (willingness to pay or time tradeoff or time trade off or tto or standard gamble*).ti,ab. |
| 57. | (sf36* or sf 36* or short form 36* or shortform 36* or shortform36*).ti,ab. |
| 58. | (sf20 or sf 20 or short form 20 or shortform 20 or shortform20).ti,ab. |
| 59. | (sf12* or sf 12* or short form 12* or shortform 12* or shortform12*).ti,ab. |
| 60. | (sf8* or sf 8* or short form 8* or shortform 8* or shortform8*).ti,ab. |
| 61. | (sf6* or sf 6* or short form 6* or shortform 6* or shortform6*).ti,ab. |
| 62. | or/43-61 |
| 63. | 25 and 42 |
| 64. | 25 and 62 |
| 65. | limit 63 to English language |
| 66. | limit 64 to English language |

Embase (Ovid) search terms

| | |
|-----|--|
| 1. | exp Cerebrovascular accident/ |
| 2. | exp Brain infarction/ |
| 3. | (stroke or strokes or cva or poststroke* or apoplexy or "cerebrovascular accident").ti,ab. |
| 4. | ((cerebro* or brain or brainstem or cerebral*) adj3 (infarct* or accident*)).ti,ab. |
| 5. | "brain attack".ti,ab. |
| 6. | Intracerebral hemorrhage/ |
| 7. | or/1-6 |
| 8. | letter.pt. or letter/ |
| 9. | note.pt. |
| 10. | editorial.pt. |
| 11. | case report/ or case study/ |
| 12. | (letter or comment*).ti. |
| 13. | or/8-12 |
| 14. | randomized controlled trial/ or random*.ti,ab. |
| 15. | 13 not 14 |
| 16. | animal/ not human/ |
| 17. | nonhuman/ |
| 18. | exp Animal Experiment/ |
| 19. | exp Experimental Animal/ |
| 20. | animal model/ |
| 21. | exp Rodent/ |
| 22. | (rat or rats or mouse or mice).ti. |
| 23. | or/15-22 |
| 24. | 7 not 23 |
| 25. | health economics/ |
| 26. | exp economic evaluation/ |

| | |
|-----|---|
| 27. | exp health care cost/ |
| 28. | exp fee/ |
| 29. | budget/ |
| 30. | funding/ |
| 31. | budget*.ti,ab. |
| 32. | cost*.ti. |
| 33. | (economic* or pharmaco?economic*).ti. |
| 34. | (price* or pricing*).ti,ab. |
| 35. | (cost* adj2 (effective* or utilit* or benefit* or minimi* or unit* or estimat* or variable*)).ab. |
| 36. | (financ* or fee or fees).ti,ab. |
| 37. | (value adj2 (money or monetary)).ti,ab. |
| 38. | or/25-37 |
| 39. | quality adjusted life year/ |
| 40. | "quality of life index"/ |
| 41. | short form 12/ or short form 20/ or short form 36/ or short form 8/ |
| 42. | sickness impact profile/ |
| 43. | (quality adj2 (wellbeing or well being)).ti,ab. |
| 44. | sickness impact profile.ti,ab. |
| 45. | disability adjusted life.ti,ab. |
| 46. | (qal* or qtime* or qwb* or daly*).ti,ab. |
| 47. | (euroqol* or eq5d* or eq 5*).ti,ab. |
| 48. | (qol* or hq1* or hqol* or h qol* or hrqol* or hr qol*).ti,ab. |
| 49. | (health utility* or utility score* or disutilit* or utility value*).ti,ab. |
| 50. | (hui or hui1 or hui2 or hui3).ti,ab. |
| 51. | (health* year* equivalent* or hye or hyes).ti,ab. |
| 52. | discrete choice*.ti,ab. |
| 53. | rosser.ti,ab. |
| 54. | (willingness to pay or time tradeoff or time trade off or tto or standard gamble*).ti,ab. |
| 55. | (sf36* or sf 36* or short form 36* or shortform 36* or shortform36*).ti,ab. |
| 56. | (sf20 or sf 20 or short form 20 or shortform 20 or shortform20).ti,ab. |
| 57. | (sf12* or sf 12* or short form 12* or shortform 12* or shortform12*).ti,ab. |
| 58. | (sf8* or sf 8* or short form 8* or shortform 8* or shortform8*).ti,ab. |
| 59. | (sf6* or sf 6* or short form 6* or shortform 6* or shortform6*).ti,ab. |
| 60. | or/39-59 |
| 61. | limit 24 to English language |
| 62. | 38 and 61 |
| 63. | 60 and 61 |

NHS EED and HTA (CRD) search terms

| | |
|-----|---|
| #1. | MeSH DESCRIPTOR Stroke EXPLODE ALL TREES |
| #2. | MeSH DESCRIPTOR Cerebral Hemorrhage EXPLODE ALL TREES |

| | |
|-----|---|
| #3. | (stroke* or cva or poststroke* or apoplexy or "cerebrovascular accident") |
| #4. | ((((cerebro* or brain or brainstem or cerebral*) adj3 (infarct* or accident*))) |
| #5. | ("brain attack*") |
| #6. | #1 OR #2 OR #3 OR #4 OR #5 |

INAHTA search terms

| | |
|----|--|
| 1. | (brain attack*) OR (((cerebro* or brain or brainstem or cerebral*) and (infarct* or accident*))) OR ((stroke or strokes or cva or poststroke* or apoplexy or "cerebrovascular accident")) OR ("Cerebral Hemorrhage"[mhe]) OR ("Stroke"[mhe]) |
|----|--|

CINAHL search terms

| | |
|-----|---|
| 1. | MH "Economics+" |
| 2. | MH "Financial Management+" |
| 3. | MH "Financial Support+" |
| 4. | MH "Financing, Organized+" |
| 5. | MH "Business+" |
| 6. | S2 OR S3 or S4 OR S5 |
| 7. | S1 not S6 |
| 8. | MH "Health Resource Allocation" |
| 9. | MH "Health Resource Utilization" |
| 10. | S8 OR S9 |
| 11. | S7 OR S10 |
| 12. | (cost or costs or economic* or pharmacoeconomic* or price* or pricing*) OR AB (cost or costs or economic* or pharmacoeconomic* or price* or pricing*) |
| 13. | S11 OR S12 |
| 14. | PT editorial |
| 15. | PT letter |
| 16. | PT commentary |
| 17. | S14 or S15 or S16 |
| 18. | S13 NOT S17 |
| 19. | MH "Animal Studies" |
| 20. | (ZT "doctoral dissertation") or (ZT "masters thesis") |
| 21. | S18 NOT (S19 OR S20) |
| 22. | PY 2014- |
| 23. | S21 AND S22 |
| 24. | MW Stroke or MH Cerebral Hemorrhage |
| 25. | stroke* or cva or poststroke* or apoplexy or "cerebrovascular accident" |
| 26. | (cerebro* OR brain OR brainstem OR cerebral*) AND (infarct* OR accident*) |
| 27. | "brain attack*" |
| 28. | S24 OR S25 OR S26 OR S27 |
| 29. | S23 AND S28 |

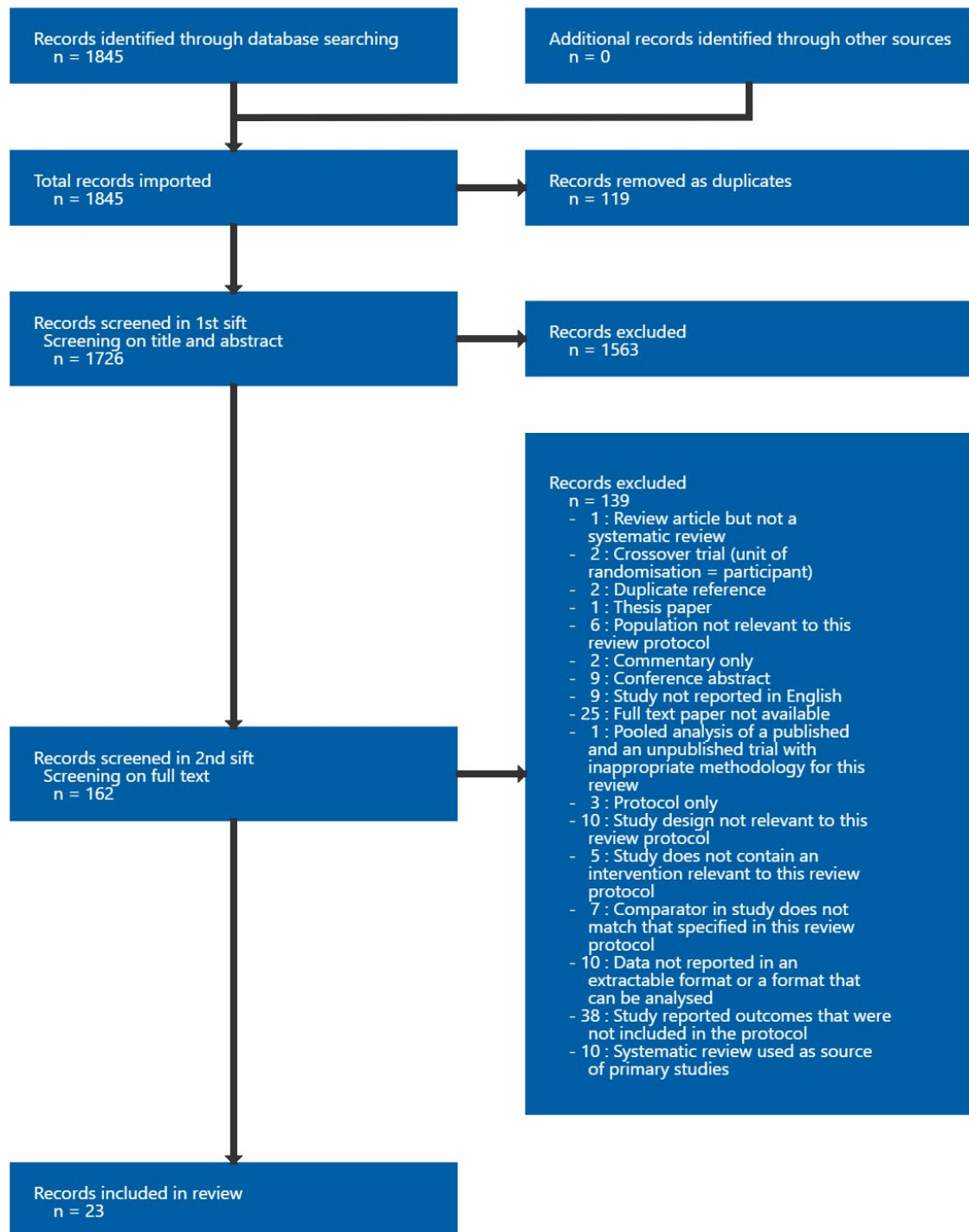
PsycINFO search terms

| | |
|----|--------------------------|
| 1. | exp Stroke/ |
| 2. | exp Cerebral hemorrhage/ |

| | |
|-----|---|
| 3. | (stroke or strokes or cva or poststroke* or apoplexy or "cerebrovascular accident").ti,ab. |
| 4. | ((cerebro* or brain or brainstem or cerebral*) adj3 (infarct* or accident*)).ti,ab. |
| 5. | "brain attack*".ti,ab. |
| 6. | Cerebrovascular accidents/ |
| 7. | exp Brain damage/ |
| 8. | (brain adj2 injur*).ti. |
| 9. | or/1-8 |
| 10. | Letter/ |
| 11. | Case report/ |
| 12. | exp Rodents/ |
| 13. | or/10-12 |
| 14. | 9 not 13 |
| 15. | limit 14 to (human and English language) |
| 16. | First posting.ps. |
| 17. | 15 and 16 |
| 18. | 15 or 17 |
| 19. | "costs and cost analysis"/ |
| 20. | "Cost Containment"/ |
| 21. | (economic adj2 evaluation\$).ti,ab. |
| 22. | (economic adj2 analy\$).ti,ab. |
| 23. | (economic adj2 (study or studies)).ti,ab. |
| 24. | (cost adj2 evaluation\$).ti,ab. |
| 25. | (cost adj2 analy\$).ti,ab. |
| 26. | (cost adj2 (study or studies)).ti,ab. |
| 27. | (cost adj2 effective\$).ti,ab. |
| 28. | (cost adj2 benefit\$).ti,ab. |
| 29. | (cost adj2 utili\$).ti,ab. |
| 30. | (cost adj2 minimi\$).ti,ab. |
| 31. | (cost adj2 consequence\$).ti,ab. |
| 32. | (cost adj2 comparison\$).ti,ab. |
| 33. | (cost adj2 identificat\$).ti,ab. |
| 34. | (pharmacoeconomic\$ or pharmaco-economic\$).ti,ab. |
| 35. | or/19-34 |
| 36. | (0003-4819 or 0003-9926 or 0959-8146 or 0098-7484 or 0140-6736 or 0028-4793 or 1469-493X).is. |
| 37. | 35 not 36 |
| 38. | 18 and 37 |

Appendix C – Effectiveness evidence study selection

Figure 1: Flow chart of clinical study selection for the review of music therapy



Appendix D – Effectiveness evidence

Baylan, 2020

Bibliographic Reference

Baylan, S.; Haig, C.; MacDonald, M.; Stiles, C.; Easto, J.; Thomson, M.; Cullen, B.; Quinn, T. J.; Stott, D.; Mercer, S. W.; Broomfield, N. M.; Murray, H.; Evans, J. J.; Measuring the effects of listening for leisure on outcome after stroke (MELLO): A pilot randomized controlled trial of mindful music listening; International Journal of Stroke; 2020; vol. 15 (no. 2); 149-158

Study details

| | |
|---|---|
| Secondary publication of another included study- see primary study for details | No additional information. |
| Other publications associated with this study included in review | No additional information. |
| Trial name / registration number | Clinicaltrials.gov - NCT02259062. |
| Study type | Randomised controlled trial (RCT) |
| Study location | United Kingdom. |
| Study setting | Acute Stroke Units within NHS Greater Glasgow and Clyde. |
| Study dates | 12th January 2015 to 28th January 2016. |
| Sources of funding | This work was supported by the Dunhill Medical Trust, grant R432/0214. Additional support from Scottish Executive Chief Scientist Office (TQ/BC), Stroke Association (TQ) and The Dr Mortimer and Theresa Sackler Foundation (BC/SB). |

| | |
|---|---|
| Inclusion criteria | Native English-speaking adults (aged 18, upper age limit of 80 for the first 11 months of recruitment) in the acute phase (no more than 14 days post-stroke) following clinically and/or radiologically (CT and/or MRI) confirmed diagnosis of ischemic stroke. |
| Exclusion criteria | Comorbid progressive neurological or neurodegenerative condition; major psychiatric disorder (pre-stroke history of mood disorder or stable antidepressant medication did not lead to exclusion); history of major substance abuse problems; clinically unstable; unable to give informed consent or unable to cooperate with the study protocol (e.g. due to severe aphasia, uncorrected impairment of hearing or vision); co-recruitment with intervention studies with potential impact on mood/cognition. |
| Recruitment / selection of participants | The Scottish Stroke Research Network nurses approached potential participants at the ward. |
| Intervention(s) | <p>Music intervention delivered by healthcare professionals N=23</p> <p>Mindful music listening. People were given an iPod Nano (7th Generation, Apple Inc.) and asked to listen to their material daily on their own for at least an hour during the intervention phase (target 56 hours over 8 consecutive weeks). They were introduced to the concept of mindfulness and were given a recording containing a brief mindfulness exercise (Body scan) to complete daily prior to music listening. The brief exercises focussed on key elements of mindfulness (e.g. paying attention to the present moment). If people were to notice any thoughts or sensations during the exercise and the music listening, they were to allow them to pass and gently bring their attention back to the exercise/music. Delivered by an assistant psychologist.</p> |
| Subgroup 1: Time after stroke at the start of the trial | Subacute (7 days - 6 months) |
| Subgroup 2: Severity (as stated by category or as measured by NIHSS scale) | Not stated/unclear |
| Subgroup 3: Group compared to individual sessions of music therapy | Individual sessions |

| | |
|---|--|
| Subgroup 4: Delivered in hospital compared to delivered in the community | Hospital sessions |
| Subgroup 5: Type of intervention | Receptive interventions in which participants listen to music With mindfulness exercise |
| Population subgroups | No additional information. |
| Comparator | Passive music listening N=24 Listening to music without any mindfulness components. Placebo music intervention N=25 Audiobook listening instead of music listening. |
| Number of participants | 72. |
| Duration of follow-up | 8 weeks of intervention, 3 months and 6 months follow up. |
| Indirectness | No additional information. |
| Additional comments | People were analysed on an intention-to-treat basis. Missing data were not imputed. |

Study arms

Music intervention delivered by healthcare professionals (N = 23)

Mindful music listening. People were given an iPod Nano (7th Generation, Apple Inc.) and asked to listen to their material daily on their own for at least an hour during the intervention phase (target 56 hours over 8 consecutive weeks). They were introduced to the concept of mindfulness and were given a recording containing a brief mindfulness exercise (Body scan) to complete daily prior to music listening. The brief exercises focussed on key elements of mindfulness (e.g. paying attention to the present moment). If people were to notice any thoughts or sensations during the exercise and the music listening, they were to allow them to pass and gently bring their attention back to the exercise/music. Delivered by an assistant psychologist.

Passive music listening (N = 24)

Listening to music without any mindfulness components.

Placebo music intervention (N = 25)

Audiobook listening instead of music listening.

Characteristics

Study-level characteristics

| Characteristic | Study (N = 72) |
|---------------------------------|-----------------------|
| Time after stroke (days) | 5 to 28 |
| Range | |
| Time after stroke (days) | 19 (NR to NR) |
| Median (IQR) | |

Arm-level characteristics

| Characteristic | Music intervention delivered by healthcare professionals (N = 23) | Passive music listening (N = 24) | Placebo music intervention (N = 25) |
|-----------------------|--|---|--|
| % Female | 10 | 7 | 10 |
| Nominal | | | |
| Mean age (SD) | 65.3 (11.13) | 61.1 (10.36) | 65.7 (12.97) |
| Mean (SD) | | | |
| Ethnicity | NR | NR | NR |
| Nominal | | | |
| Comorbidities | NR | NR | NR |
| Nominal | | | |
| Severity | NR | NR | NR |
| Nominal | | | |
| Type of stroke | NA | NA | NA |
| Nominal | | | |
| Cortical | 15 | 15 | 16 |
| Nominal | | | |
| Subcortical | 8 | 9 | 9 |
| Nominal | | | |

Outcomes

Study timepoints

- Baseline
- 6 month
- 3 month

Music intervention delivered by healthcare professional compared to placebo therapy at less than/equal to 6 months

| Outcome | Music intervention delivered by healthcare professionals, Baseline, N = 23 | Music intervention delivered by healthcare professionals, 6 month, N = 23 | Music intervention delivered by healthcare professionals, 3 month, N = 23 | Passive music listening, Baseline, N = 24 | Passive music listening, 6 month, N = 24 | Passive music listening, 3 month, N = 24 | Placebo music intervention, Baseline, N = 25 | Placebo music intervention, 6 month, N = 25 | Placebo music intervention, 3 month, N = 25 |
|--|--|---|---|---|--|--|--|---|---|
| Psychological distress - Depression (HADS depression) Scale range: 0-42. Adjusted mean difference (between music intervention and placebo). Mean (95% CI) | NR (NR to NR) | 1.02 (-1.36 to 3.4) | 0.7 (-1.65 to 3.04) | NR (NR to NR) | NR (NR to NR) | NR (NR to NR) | NR (NR to NR) | NA (NA to NA) | NA (NA to NA) |
| Psychological distress - Depression | 5 (2 to 7) | NR (NR to NR) | NR (NR to NR) | 5 (1.5 to 11.5) | NR (NR to NR) | NR (NR to NR) | 6.5 (2 to 8) | NR (NR to NR) | NR (NR to NR) |

| Outcome | Music intervention delivered by healthcare professionals, Baseline, N = 23 | Music intervention delivered by healthcare professionals, 6 month, N = 23 | Music intervention delivered by healthcare professionals, 3 month, N = 23 | Passive music listening, Baseline, N = 24 | Passive music listening, 6 month, N = 24 | Passive music listening, 3 month, N = 24 | Placebo music intervention, Baseline, N = 25 | Placebo music intervention, 6 month, N = 25 | Placebo music intervention, 3 month, N = 25 |
|---|---|--|--|--|---|---|---|--|--|
| (HADS depression) Scale range: 0-42. Adjusted mean difference (between music intervention and placebo). Median (IQR) | | | | | | | | | |
| Psychological distress - Anxiety (HADS anxiety) Scale range: 0-42. Adjusted mean difference (between music intervention and placebo). Median (IQR) | 7 (6 to 9.5) | NR (NR to NR) | NR (NR to NR) | 5 (1.5 to 11.5) | NR (NR to NR) | NR (NR to NR) | 7 (3 to 12) | NR (NR to NR) | NR (NR to NR) |
| Psychological distress - Anxiety (HADS) | NR (NR to NR) | 2 (-0.28 to 4.28) | 0.69 (-1.47 to 2.84) | NR (NR to NR) | NR (NR to NR) | NR (NR to NR) | NR (NR to NR) | NA (NA to NA) | NA (NA to NA) |

| Outcome | Music intervention delivered by healthcare professionals, Baseline, N = 23 | Music intervention delivered by healthcare professionals, 6 month, N = 23 | Music intervention delivered by healthcare professionals, 3 month, N = 23 | Passive music listening, Baseline, N = 24 | Passive music listening, 6 month, N = 24 | Passive music listening, 3 month, N = 24 | Placebo music intervention, Baseline, N = 25 | Placebo music intervention, 6 month, N = 25 | Placebo music intervention, 3 month, N = 25 |
|---|--|---|---|---|--|--|--|---|---|
| <p>anxiety) Scale range: 0-42. Adjusted mean difference (between music intervention and placebo).</p> <p>Mean (95% CI)</p> | | | | | | | | | |
| <p>Participation in leisure activities/social groups scores (Mayo-Portland Adapatability Inventory 4 participation) Scale range: 0-30. Adjusted mean difference (between music intervention and placebo).</p> <p>Mean (95% CI)</p> | NR (NR to NR) | 1.72 (-11.75 to 15.19) | NR (NR to NR) | NR (NR to NR) | NR (NR to NR) | NR (NR to NR) | NR (NR to NR) | NA (NA to NA) | NR (NR to NR) |

Psychological distress - Depression (HADS depression) - Polarity - Higher values are better

Psychological distress - Anxiety (HADS anxiety) - Polarity - Higher values are better

Participation in leisure activities/social groups scores (Mayo-Portland Adaptability Inventory 4 participation) - Polarity - Higher values are better

Only reports adjusted mean differences for music intervention compared to placebo, or passive music listening compared to placebo. The only comparator relevant to our protocol is music intervention compared to placebo.

Music intervention delivered by healthcare professional compared to placebo therapy and passive music listening at less than/equal to 6 months

| Outcome | Music intervention delivered by healthcare professionals, Baseline, N = 23 | Music intervention delivered by healthcare professionals, 6 month, N = 23 | Music intervention delivered by healthcare professionals, 3 month, N = 23 | Passive music listening, Baseline, N = 24 | Passive music listening, 6 month, N = 24 | Passive music listening, 3 month, N = 24 | Placebo music intervention, Baseline, N = 25 | Placebo music intervention, 6 month, N = 25 | Placebo music intervention, 3 month, N = 25 |
|--|---|--|--|--|---|---|---|--|--|
| Withdrawal due to adverse events Withdrew due to ill health. Mindful music listening: 4, Music group: 1, Audiobook: 1. Nominal | NA | 4 | 4 | NA | 1 | 1 | NA | 1 | 1 |

Withdrawal due to adverse events - Polarity - Lower values are better

Critical appraisal - Cochrane Risk of Bias tool (RoB 2.0) Normal RCT

Music intervention delivered by healthcare professional compared to placebo therapy at less than/equal to 6 months - Psychological distress - Depression (HADS depression) - Mean Nine Five Percent CI - Music intervention delivered by healthcare professionals - Passive music listening - Placebo music intervention - t6

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professional compared to placebo therapy at less than/equal to 6 months - Psychological distress - Anxiety (HADS anxiety) - Mean Nine Five Percent CI - Music intervention delivered by healthcare professionals - Passive music listening - Placebo music intervention - t6

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professional compared to placebo therapy at less than/equal to 6 months - Participation in leisure activities/social group scores (Mayo-Portland Adapatability Inventory 4 participation) - Mean Nine Five Percent CI - Music intervention delivered by healthcare professionals - Passive music listening - Placebo music intervention - t6

| Section | Question | Answer |
|-----------------------------|------------------------|--------|
| Overall bias and Directness | Risk of bias judgement | High |

| Section | Question | Answer |
|-----------------------------|--------------------|---------------------|
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professional compared to placebo therapy and passive music listening at less than/equal to 6 months- Withdrawal due to adverse events-Nominal-Music intervention delivered by healthcare professionals-Passive music listening-Placebo music intervention-t6

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | Some concerns |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professional compared to placebo therapy at less than/equal to 6 months-Psychological distress- Depression (HADS depression)-Mean Nine Five Percent CI-Music intervention delivered by healthcare professionals-Passive music listening-Placebo music intervention-t3

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professional compared to placebo therapy at less than/equal to 6 months - Psychological distress - Anxiety (HADS anxiety) - Mean Nine Five Percent CI - Music intervention delivered by healthcare professionals - Passive music listening - Placebo music intervention - t3

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professional compared to placebo therapy and passive music listening at less than/equal to 6 months - Withdrawal due to adverse events - Nominal - Music intervention delivered by healthcare professionals - Passive music listening - Placebo music intervention - t3

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | Some concerns |
| Overall bias and Directness | Overall Directness | Directly applicable |

Cha, 2014

Bibliographic Reference

Cha, Y.; Kim, Y.; Hwang, S.; Chung, Y.; Intensive gait training with rhythmic auditory stimulation in individuals with chronic hemiparetic stroke: a pilot randomized controlled study; Neurorehabilitation; 2014; vol. 35 (no. 4); 681-8

Study details

| | |
|---|----------------------------|
| Secondary publication of another included | No additional information. |
|---|----------------------------|

| | |
|---|---|
| study- see primary study for details | |
| Other publications associated with this study included in review | No additional information. |
| Trial name / registration number | No additional information. |
| Study type | Randomised controlled trial (RCT) |
| Study location | Republic of Korea. |
| Study setting | Outpatient follow up. |
| Study dates | No additional information. |
| Sources of funding | This study was supported by S University (2013). |
| Inclusion criteria | At least six months since a clinical diagnosis of ischaemic or haemorrhagic hemiparetic stroke; sufficient cognitive ability to participate, as indicated by a Mini-Mental State Examination score of 20 or higher; Brunnstrom stage III or IV in the proximal and distal parts of the lower extremity; no serious auditory or visual deficits. |
| Exclusion criteria | A symptomatic cardiac failure or unstable angina; uncontrolled hypertension; significant orthopaedic or chronic pain conditions affecting gait performance; any neurologic disease except for the initial stroke. |
| Recruitment / selection of participants | People were recruited from the H hospital. No additional information. |
| Intervention(s) | <p>Music therapy delivered by a music therapist N=10</p> <p>Rhythmic auditory stimulation training group. Intensive gait training with rhythmic auditory stimulation. People used a metronome and specifically prepared music tapes (based on personal preference of either pop or country music). A music specialist then played three songs emphasizing the rhythms based on selected music, using a synthesizer keyboard along with the MIDI Cuebase musical instrument digital interface program, and a KM Player version 3.3 to control the rhythmic tempo in each participant. A metronome was played over the beat of the music in order to enhance the rhythmic perception for the person. The metronome was set up so that it matched directly with the person's step pattern. Participants were</p> |

| | |
|---|---|
| | <p>asked to move around a track at a self-selected, comfortable pace for 2 minutes to establish a baseline cadence of rhythmic auditory stimulation. This was then progressively increased over the study duration.</p> <p>Concomitant therapy: All participants received general physical therapy, including Bobath approach and proprioceptive neuromuscular facilitation for 30 minutes per day, five days per week.</p> |
| Subgroup 1: Time after stroke at the start of the trial | Chronic (>6 months) |
| Subgroup 2: Severity (as stated by category or as measured by NIHSS scale) | Not stated/unclear |
| Subgroup 3: Group compared to individual sessions of music therapy | Individual sessions |
| Subgroup 4: Delivered in hospital compared to delivered in the community | Hospital sessions |
| Subgroup 5: Type of intervention | Rhythmic auditory cueing |
| Population subgroups | No additional information |

| | |
|-------------------------------|---|
| Comparator | No treatment N=10 Intensive gait training only (same progressive protocol, completed without music). Concomitant therapy: All participants received general physical therapy, including Bobath approach and proprioceptive neuromuscular facilitation for 30 minutes per day, five days per week. |
| Number of participants | 20 |
| Duration of follow-up | 6 weeks (end of intervention). |
| Indirectness | No additional information. |
| Additional comments | All participants completed the study in their intended arms (available case analysis, no loss of participants). |

Study arms

Music therapy delivered by a music therapist (N = 10)

Rhythmic auditory stimulation training group. Intensive gait training with rhythmic auditory stimulation. People used a metronome and specifically prepared music tapes (based on personal preference of either pop or country music). A music specialist then played three songs emphasizing the rhythms based on selected music, using a synthesizer keyboard along with the MIDI Cuebase musical instrument digital interface program, and a KM Player version 3.3 to control the rhythmic tempo in each participant. A metronome was played over the beat of the music in order to enhance the rhythmic perception for the person. The metronome was set up so that it matched directly with the person's step pattern. Participants were asked to move around a track at a self-selected, comfortable pace for 2 minutes to establish a baseline cadence of rhythmic auditory stimulation. This was then progressively increased over the study duration. Concomitant therapy: All participants received general physical therapy, including Bobath approach and proprioceptive neuromuscular facilitation for 30 minutes per day, five days per week.

No treatment (N = 10)

Intensive gait training only (same progressive protocol, completed without music). Concomitant therapy: All participants received general physical therapy, including Bobath approach and proprioceptive neuromuscular facilitation for 30 minutes per day, five days per week.

Characteristics**Arm-level characteristics**

| Characteristic | Music therapy delivered by a music therapist (N = 10) | No treatment (N = 10) |
|-----------------------------------|--|------------------------------|
| % Female | n = 4 ; % = 40 | n = 4 ; % = 40 |
| Sample size | | |
| Mean age (SD) (years) | 59.8 (11.7) | 63 (14.1) |
| Mean (SD) | | |
| Ethnicity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Comorbidities | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Time after stroke (Months) | 14.5 (5.5) | 14.7 (5.4) |
| Mean (SD) | | |
| Severity | n = NR ; % = NR | n = NR ; % = NR |
| No of events | | |

| Characteristic | Music therapy delivered by a music therapist (N = 10) | No treatment (N = 10) |
|----------------|---|-----------------------|
| Type of stroke | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |

Outcomes

Study timepoints

- Baseline
- 6 week (<6 months)

Music therapy delivered by music therapists compared to no treatment at <6 months - continuous outcomes

| Outcome | Music therapy delivered by a music therapist, Baseline, N = 10 | Music therapy delivered by a music therapist, 6 week, N = 10 | No treatment, Baseline, N = 10 | No treatment, 6 week, N = 10 |
|--|--|--|--------------------------------|------------------------------|
| Stroke-specific Patient-Reported Outcome Measures (Stroke specific quality of life scale) Scale range: 49-245. Final values. | 158.6 (18.3) | 183.7 (21.5) | 153 (17.1) | 159.2 (17.4) |
| Mean (SD) | | | | |

Stroke-specific Patient-Reported Outcome Measures (Stroke specific quality of life scale) - Polarity - Higher values are better

Music therapy delivered by music therapists compared to no treatment at <6 months - dichotomous outcome

| Outcome | Music therapy delivered by a music therapist, Baseline, N = 10 | Music therapy delivered by a music therapist, 6 week, N = 10 | No treatment, Baseline, N = 10 | No treatment, 6 week, N = 10 |
|---|--|--|--------------------------------|------------------------------|
| Withdrawal due to adverse events | n = NA ; % = NA | n = 0 ; % = 0 | n = NA ; % = NA | n = 0 ; % = 0 |
| No of events | | | | |

Withdrawal due to adverse events - Polarity - Lower values are better

Critical appraisal - Cochrane Risk of Bias tool (RoB 2.0) Normal RCT**Music therapy delivered by music therapists compared to no treatment at <6 months - continuous outcomes - Stroke-specific Patient-Reported Outcome Measures (Stroke specific quality of life scale) - Mean SD - Music therapy delivered by a music therapist - No treatment - t6**

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music therapy delivered by music therapists compared to no treatment at <6 months - dichotomous outcome - Withdrawal due to adverse events - No Of Events - Music therapy delivered by a music therapist - No treatment - t6

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | Some concerns |
| Overall bias and Directness | Overall Directness | Directly applicable |

Fujioka, 2018**Bibliographic Reference**

Fujioka, T.; Dawson, D. R.; Wright, R.; Honjo, K.; Chen, J. L.; Chen, J. J.; Black, S. E.; Stuss, D. T.; Ross, B.; The effects of music-supported therapy on motor, cognitive, and psychosocial functions in chronic stroke; Annals of the New York Academy of Sciences; 2018; vol. 24; 24

Study details

| | |
|---|---|
| Secondary publication of another included study- see primary study for details | No additional information. |
| Other publications associated with this study included in review | No additional information. |
| Trial name / registration number | NCT01721668. |
| Study type | Randomised controlled trial (RCT) |
| Study location | Canada |
| Study setting | Outpatient follow up |
| Study dates | No additional information. |
| Sources of funding | This research was supported by the Canadian Institutes of Health Research and the Heart and Stroke Foundation Ontario. |
| Inclusion criteria | Unilateral arm and hand impairment; English speaking; near-normal hearing as assessed by clinical audiometry (<40 dB, 250-2000 Hz) and corrected vision adequate for verbal communication; first-time stroke at least 6 months prior to |

| | |
|---|--|
| | enrolment; demonstrating voluntary movement in the arm or hand measured as passing at least one of the Stage 3 tasks on the Chedoke-McMaster Stroke Assessment Impairment Inventory of which we administered only arm and hand sections. |
| Exclusion criteria | Severe apraxia, aphasia, spatial neglect, or sensory loss in the paretic hand based on the self-report as well as observation during the screening visit; dementia or cognitive impairment based on the Montreal Cognitive Assessment; psychiatric disorders including depression based on the Center for Epidemiological Studies Depression Scale using a cut off score of 20; severe pain and/or fatigue (no more than 45 in any area of the Visual Analogue Scale for Pain with a straight horizontal line of 100-mm length regarding the affected arm, hand and shoulder); extensive prior musical experience (formal musical training >2 years within the past 10 years or >10 years in total). |
| Recruitment / selection of participants | No additional information. |
| Intervention(s) | <p>Music therapy delivered by music therapists N=14</p> <p>Music-supported therapy added to support motor, cognitive and psychosocial functions combined with an existing conventional physical training programme (GRASP). Adjustments included drum pads being assigned pitches on a scale from C to C'; drum sticks, brushes and mallets being used besides palm and fist; African djembe used for timbre and dynamic variations as well as joint music-making with the therapist. These were incorporated into activities.</p> <p>Concomitant therapy: No additional information.</p> |
| Subgroup 1: Time after stroke at the start of the trial | Chronic (>6 months) |
| Subgroup 2: Severity (as stated by category or as measured by NIHSS scale) | Not stated/unclear |
| Subgroup 3: Group compared to | Individual sessions |

| | |
|---|--|
| individual sessions of music therapy | |
| Subgroup 4: Delivered in hospital compared to delivered in the community | Hospital sessions |
| Subgroup 5: Type of intervention | Interventions where musical instruments are played (including clinical improvisation) |
| Population subgroups | No additional information. |
| Comparator | No treatment N=15 Conventional physical training programme (GRASP) without music. Including a variety of objects including tea pot and cups, table napkins and rings, scarves and fishing reels etc. Concomitant therapy: No additional information. |
| Number of participants | 28 |
| Duration of follow-up | 5-10 weeks (end of intervention, 1 participant chose to complete 15 sessions over 5 weeks, while the remainder chose 30 sessions over 10 weeks) and 4 months. |
| Indirectness | No additional information. |
| Additional comments | Intention to treat analysis (last follow up data carried forward). |

Study arms

Music therapy delivered by music therapists (N = 14)

Music-supported therapy added to support motor, cognitive and psychosocial functions combined with an existing conventional physical training programme (GRASP). Adjustments included drum pads being assigned pitches on a scale from C to C'; drum sticks, brushes and mallets being used besides palm and fist; African djembe used for timbre and dynamic variations as well as joint music-making with the therapist. These were incorporated into activities. Concomitant therapy: No additional information.

No treatment (N = 15)

Conventional physical training programme (GRASP) without music. Including a variety of objects including tea pot and cups, table napkins and rings, scarves and fishing reels etc. Concomitant therapy: No additional information.

Characteristics

Arm-level characteristics

| Characteristic | Music therapy delivered by music therapists (N = 14) | No treatment (N = 15) |
|------------------------------|---|------------------------------|
| % Female | n = 5 | n = 3 |
| Sample size | | |
| Mean age (SD) (years) | 64.2 (9.4) | 54.3 (11.3) |
| Mean (SD) | | |
| Ethnicity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Comorbidities | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |

| Characteristic | Music therapy delivered by music therapists (N = 14) | No treatment (N = 15) |
|---------------------------|--|-----------------------|
| Time after stroke (years) | 6.1 (6.6) | 4.7 (6.7) |
| Mean (SD) | | |
| Severity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Type of stroke | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |

Outcomes

Study timepoints

- Baseline
- 4 month (3 weeks after completing the training. <6 months.)

Music therapy delivered by a music therapy compared to no treatment at <6 months - continuous outcomes

| Outcome | Music therapy delivered by music therapists, Baseline, N = 14 | Music therapy delivered by music therapists, 4 month, N = 14 | No treatment, Baseline, N = 15 | No treatment, 4 month, N = 14 |
|--|---|--|--------------------------------|-------------------------------|
| Psychological distress (Positive affect) | 34.43 (7.38) | 36.79 (8.19) | 29.79 (11.56) | 32.64 (8.45) |
| Scale range: 10-50. Final values (from PANAS). | | | | |
| Mean (SD) | | | | |

| Outcome | Music therapy delivered by music therapists, Baseline, N = 14 | Music therapy delivered by music therapists, 4 month, N = 14 | No treatment, Baseline, N = 15 | No treatment, 4 month, N = 14 |
|--|--|---|---------------------------------------|--------------------------------------|
| Psychological distress (Negative affect) Scale range: 10-50. Final values (from PANAS). Mean (SD) | 20 (6.13) | 18 (6.82) | 18.5 (8.44) | 17.29 (8.56) |
| Stroke Specific Patient-Reported Outcome Measures (Stroke Impact Scale) Scale range: 0-100. Final values. Mean (SD) | NR (NR) | NR (NR) | NR (NR) | NR (NR) |
| SIS Mobility Mean (SD) | 59.89 (17.08) | 60.86 (9.68) | 58.11 (15.34) | 63.13 (16.61) |
| SIS Memory Mean (SD) | 81.14 (12.78) | 82.91 (12.59) | 81.88 (26.07) | 83.16 (22.55) |
| SIS Emotion Mean (SD) | 71.46 (20.22) | 76.8 (18.22) | 73.61 (22.86) | 77 (20.68) |
| SIS Communication Mean (SD) | 84.23 (14.46) | 90.3 (9.23) | 82.64 (26.56) | 92.11 (10.37) |
| SIS Social Mean (SD) | 59.64 (18.27) | 63.52 (14.36) | 65.45 (22.09) | 59.3 (24.95) |

Psychological distress (Positive affect) - Polarity - Higher values are better

Psychological distress (Negative affect) - Polarity - Lower values are better

Stroke Specific Patient-Reported Outcome Measures (Stroke Impact Scale) - Polarity - Higher values are better

Music therapy delivered by a music therapy compared to no treatment at <6 months - dichotomous outcome

| Outcome | Music therapy delivered by music therapists, Baseline, N = 14 | Music therapy delivered by music therapists, 4 month, N = 14 | No treatment, Baseline, N = 15 | No treatment, 4 month, N = 14 |
|---|--|---|---------------------------------------|--------------------------------------|
| Withdrawal due to adverse events | n = NA ; % = NA | n = 0 ; % = 0 | n = NA ; % = NA | n = 0 ; % = 0 |
| No of events | | | | |

Withdrawal due to adverse events - Polarity - Lower values are better

Critical appraisal - Cochrane Risk of Bias tool (RoB 2.0) Normal RCT

Music therapy delivered by a music therapy compared to no treatment at <6 months - continuous outcomes -

Psychological distress (Positive affect) - Mean SD - Music therapy delivered by music therapists - No treatment - t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | Low |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music therapy delivered by music therapy compared to no treatment at <6 months - continuous outcomes - Psychological distress (Negative affect) - Mean SD - Music therapy delivered by music therapists - No treatment - t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | Low |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music therapy delivered by music therapy compared to no treatment at <6 months - continuous outcomes - Stroke Specific Patient - Reported Outcome Measures (Stroke Impact Scale) - SIS Mobility - Mean SD - Music therapy delivered by music therapists - No treatment - t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | Low |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music therapy delivered by music therapy compared to no treatment at <6 months - continuous outcomes - Stroke Specific Patient - Reported Outcome Measures (Stroke Impact Scale) - SIS Memory - Mean SD - Music therapy delivered by music therapists - No treatment - t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | Low |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music therapy delivered by music therapy compared to no treatment at <6 months-continuous outcomes-Stroke Specific Patient-Reported Outcome Measures (Stroke Impact Scale)-SISEmotion-MeanSD-Music therapy delivered by music therapists-No treatment-t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | Low |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music therapy delivered by music therapy compared to no treatment at <6 months-continuous outcomes-Stroke Specific Patient-Reported Outcome Measures (Stroke Impact Scale)-SIS Communication-MeanSD-Music therapy delivered by music therapists-No treatment-t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | Low |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music therapy delivered by music therapy compared to no treatment at <6 months-continuous outcomes-Stroke Specific Patient-Reported Outcome Measures (Stroke Impact Scale)-SIS Social-MeanSD-Music therapy delivered by music therapists-No treatment-t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | Low |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music therapy delivered by music therapists compared to no treatment at <6 months - dichotomous outcome - Withdrawal due to adverse events - No Of Events - Music therapy delivered by music therapists - No treatment - t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | Low |
| Overall bias and Directness | Overall Directness | Directly applicable |

Grau-Sanchez, 2018

Bibliographic Reference Grau-Sanchez, J.; Duarte, E.; Ramos-Escobar, N.; Sierpowska, J.; Rueda, N.; Redon, S.; Veciana de Las Heras, M.; Pedro, J.; Sarkamo, T.; Rodriguez-Fornells, A.; Music-supported therapy in the rehabilitation of subacute stroke patients: a randomized controlled trial; Annals of the New York Academy of Sciences; 2018; vol. 01; 01

Study details

| | |
|---|-----------------------------------|
| Secondary publication of another included study- see primary study for details | No additional information |
| Other publications associated with this study included in review | No additional information |
| Trial name / registration number | NCT02208219 |
| Study type | Randomised controlled trial (RCT) |

| | |
|---|--|
| Study location | Spain. |
| Study setting | Outpatient follow up. Rehabilitation delivered in hospital setting. |
| Study dates | December 2013 to May 2017. |
| Sources of funding | This work was supported by the Spanish Government (Ministerio de Economía y Competitividad, PSI2015-69178-P, Fondo Europeo de Desarrollo Regional (FEDER)). |
| Inclusion criteria | Mild-to-moderate paresis of the upper extremity after a first-ever stroke; less than 6 months after the stroke; age between 30 and 75 years; able to speak Spanish and/or Catalan. |
| Exclusion criteria | Major cognitive deficits affecting comprehension (Mini-Mental State Examination >24); neurological or psychiatric comorbidity; previous formal musical education. |
| Recruitment / selection of participants | People involved in a program of outpatient rehabilitation at the Department of Physical Medicine and Rehabilitation at Hospitals de Mar i de L'Esperança. |
| Intervention(s) | <p>Music intervention delivered by a healthcare professional N=20</p> <p>Music supported therapy delivered by an occupational therapists and physiotherapist. 20 individual sessions (5 sessions per week, 30 minutes each) of music supported therapy involving playing a keyboard and an electronic drum set following a modular therapy regime with stepwise increase of complexity.</p> <p>Concomitant therapy: Both groups received an outpatient rehabilitation program that consisted of two 1 hour group sessions of occupational therapy and physiotherapy a day (5 days per week, 10 hours in total per week).</p> |
| Subgroup 1: Time after stroke at the start of the trial | Subacute (7 days - 6 months) |
| Subgroup 2: Severity (as stated by category or as measured by NIHSS scale) | <p>Moderate (or NIHSS 5-14)</p> <p>Mean 5.4 ranging from 2-14.</p> |

| | |
|---|---|
| Subgroup 3: Group compared to individual sessions of music therapy | Individual sessions |
| Subgroup 4: Delivered in hospital compared to delivered in the community | Hospital sessions |
| Subgroup 5: Type of intervention | Interventions where musical instruments are played (including clinical improvisation) |
| Population subgroups | No additional information. |
| Comparator | <p>No treatment N=20</p> <p>Additional individual training of the upper extremity. 20 individual sessions (5 sessions per week, 30 minutes each) consisting of passive mobilization, stretch and progressive resistance exercises, and task-specific training.</p> <p>Concomitant therapy: Both groups received an outpatient rehabilitation program that consisted of two 1 hour group sessions of occupational therapy and physiotherapy a day (5 days per week, 10 hours in total per week).</p> |
| Number of participants | 40 |
| Duration of follow-up | Intervention for 4 weeks, 3 months follow up in total. |
| Indirectness | No additional information. |
| Additional comments | Available case analysis (any people remaining at the end of the trial, cases where people did not fill out questionnaires excluded). |

Study arms

Music intervention delivered by a healthcare professional (N = 20)

Music supported therapy delivered by an occupational therapists and physiotherapist. 20 individual sessions (5 sessions per week, 30 minutes each) of music supported therapy involving playing a keyboard and an electronic drum set following a modular therapy regime with stepwise increase of complexity. Concomitant therapy: Both groups received an outpatient rehabilitation program that consisted of two 1 hour group sessions of occupational therapy and physiotherapy a day (5 days per week, 10 hours in total per week).

No treatment (N = 20)

Additional individual training of the upper extremity. 20 individual sessions (5 sessions per week, 30 minutes each) consisting of passive mobilization, stretch and progressive resistance exercises, and task-specific training. Concomitant therapy: Both groups received an outpatient rehabilitation program that consisted of two 1 hour group sessions of occupational therapy and physiotherapy a day (5 days per week, 10 hours in total per week).

Characteristics

Arm-level characteristics

| Characteristic | Music intervention delivered by a healthcare professional (N = 20) | No treatment (N = 20) |
|-----------------------|---|------------------------------|
| % Female | n = 8 ; % = 42.1 | n = 8 ; % = 40 |
| Sample size | | |
| Mean age (SD) | 45 to 74 | 49 to 72 |
| Range | | |
| Mean age (SD) | 60.1 (NR) | 62.5 (NR) |
| Mean (SD) | | |

| Characteristic | Music intervention delivered by a healthcare professional (N = 20) | No treatment (N = 20) |
|---------------------------------|---|------------------------------|
| Ethnicity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Comorbidities | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Time after stroke (days) | 32 to 162 | 28 to 136 |
| Range | | |
| Time after stroke (days) | 65.8 (NR) | 64.9 (NR) |
| Mean (SD) | | |
| Severity | 2 to 14 | 2 to 9 |
| Range | | |
| Severity | 5.8 (NR) | 5.3 (NR) |
| Mean (SD) | | |
| Type of stroke | n = NA ; % = NA | n = NA ; % = NA |
| Sample size | | |
| Ischaemic | n = 18 ; % = 94.7 | n = 14 ; % = 70 |
| Sample size | | |
| Haemorrhagic | n = 1 ; % = 5.3 | n = 6 ; % = 30 |
| Sample size | | |

Outcomes

Study timepoints

- Baseline
- 4 week (End of intervention (no values reported at 3 months for these outcomes). <6 months.)

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - continuous outcomes (Stroke-specific Patient Reported Outcome Measures)

| Outcome | Music intervention delivered by a healthcare professional, Baseline, N = 20 | Music intervention delivered by a healthcare professional, 4 week, N = 20 | No treatment, Baseline, N = 18 | No treatment, 4 week, N = 18 |
|---|---|---|--------------------------------|------------------------------|
| Stroke specific Patient-Reported Outcome Measures (Stroke Specific Quality of Life) Scale range: 49-245. Final values. Mean (SD) | 174.6 (32.9) | 185.9 (38.8) | 179.6 (27.7) | 183.5 (23.4) |

Stroke specific Patient-Reported Outcome Measures (Stroke Specific Quality of Life) - Polarity - Higher values are better

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - continuous outcomes

| Outcome | Music intervention delivered by a healthcare professional, Baseline, N = 20 | Music intervention delivered by a healthcare professional, 4 week, N = 20 | No treatment, Baseline, N = 17 | No treatment, 4 week, N = 17 |
|---|---|---|--------------------------------|------------------------------|
| Patient/participant generic health-related quality of life (SF-36) Scale range: 0-100. Final values. Mean (SD) | NR (NR) | NR (NR) | NR (NR) | NR (NR) |

| Outcome | Music intervention delivered by a healthcare professional, Baseline, N = 20 | Music intervention delivered by a healthcare professional, 4 week, N = 20 | No treatment, Baseline, N = 17 | No treatment, 4 week, N = 17 |
|--------------------------|--|--|---------------------------------------|-------------------------------------|
| Physical function | 46.2 (23.6) | 55.3 (23.4) | 39.2 (20.1) | 45 (21.9) |
| Mean (SD) | | | | |
| Role physical | 14.7 (28) | 17.6 (32.8) | 17.1 (27.7) | 23.7 (38.6) |
| Mean (SD) | | | | |
| Pain | 61.1 (34.3) | 56.7 (27.5) | 54 (36.5) | 50.4 (33) |
| Mean (SD) | | | | |
| General health | 55.3 (18.3) | 61.1 (18.4) | 59.1 (21.4) | 57.2 (22.7) |
| Mean (SD) | | | | |
| Vitality | 55.9 (21.2) | 60 (20.3) | 48.4 (21.9) | 53.2 (26.8) |
| Mean (SD) | | | | |
| Social function | 53.8 (26.1) | 75.7 (27.5) | 54.7 (31.2) | 64.7 (30.7) |
| Mean (SD) | | | | |
| Role emotional | 62.7 (48.4) | 76.5 (40.4) | 70.5 (43.1) | 70.9 (42.9) |
| Mean (SD) | | | | |
| Mental health | 69.6 (17.8) | 71.1 (21.2) | 61.7 (17.6) | 66.3 (21.6) |
| Mean (SD) | | | | |

| Outcome | Music intervention delivered by a healthcare professional, Baseline, N = 20 | Music intervention delivered by a healthcare professional, 4 week, N = 20 | No treatment, Baseline, N = 17 | No treatment, 4 week, N = 17 |
|--|--|--|---------------------------------------|-------------------------------------|
| Psychological distress - Depression (Beck Depression Inventory) Scale range: 0-63. Final values. Mean (SD) | 10.2 (6.9) | 8.8 (7.5) | 12.3 (8.5) | 10.6 (9.1) |
| Psychological distress (PANAS) Scale range: 10-50. Higher values are better for the positive affect, lower values are better for the negative affect. Final values. Mean (SD) | <i>empty data</i> | NR (<i>empty data</i>) | <i>empty data</i> | NR (NR) |
| Positive affect Mean (SD) | 33.5 (9.5) | 33.7 (9.3) | 31.5 (7.6) | 30.3 (7.3) |
| Negative affect Mean (SD) | 19.8 (9.2) | 15.8 (4.9) | 21.2 (9) | 20 (8.7) |

Patient/participant generic health-related quality of life (SF-36) - Polarity - Higher values are better

Psychological distress - Depression (Beck Depression Inventory) - Polarity - Lower values are better

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - dichotomous outcome

| Outcome | Music intervention delivered by a healthcare professional, Baseline, N = 20 | Music intervention delivered by a healthcare professional, 4 week, N = 19 | No treatment, Baseline, N = 20 | No treatment, 4 week, N = 20 |
|---|--|--|---------------------------------------|-------------------------------------|
| Withdrawal due to adverse events One person withdrew from the music intervention but not for adverse events | n = NA ; % = NA | n = 0 ; % = 0 | n = NA ; % = NA | n = 0 ; % = 0 |

| Outcome | Music intervention delivered by a healthcare professional, Baseline, N = 20 | Music intervention delivered by a healthcare professional, 4 week, N = 19 | No treatment, Baseline, N = 20 | No treatment, 4 week, N = 20 |
|--|---|---|--------------------------------|------------------------------|
| (because they transferred to a different clinic) | | | | |
| No of events | | | | |

Withdrawal due to adverse events - Polarity - Lower values are better

Critical appraisal - Cochrane Risk of Bias tool (RoB 2.0) Normal RCT

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - continuous outcomes - Stroke specific Patient-Reported Outcome Measures (Stroke Specific Quality of Life) - Mean SD - Music intervention delivered by a healthcare professional - No treatment - t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - continuous outcomes (Stroke-specific Patient Reported Outcome Measures) - Patient/participant generic health-related quality of life (SF-36) - Physical function - Mean SD - Music intervention delivered by a healthcare professional - No treatment - t4

| Section | Question | Answer |
|-----------------------------|------------------------|--------|
| Overall bias and Directness | Risk of bias judgement | High |

| Section | Question | Answer |
|-----------------------------|--------------------|---------------------|
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months-continuous outcomes (Stroke-specific Patient Reported Outcome Measures)-Patient/participant generic health-related quality of life (SF-36)-Role physical-Mean SD-Music intervention delivered by a healthcare professional-No treatment-t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months-continuous outcomes (Stroke-specific Patient Reported Outcome Measures)-Patient/participant generic health-related quality of life (SF-36)-Pain-Mean SD-Music intervention delivered by a healthcare professional-No treatment-t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months-continuous outcomes- Patient/participant generic health-related quality of life (SF-36)-General health-Mean SD-Music intervention delivered by a healthcare professional-No treatment-t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months-continuous outcomes- Patient/participant generic health-related quality of life (SF-36)-Vitality-Mean SD-Music intervention delivered by a healthcare professional-No treatment-t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months-continuous outcomes- Patient/participant generic health-related quality of life (SF-36)-Social function-Mean SD-Music intervention delivered by a healthcare professional-No treatment-t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months-continuous outcomes- Patient/participant generic health-related quality of life (SF-36)-Role emotional-Mean SD-Music intervention delivered by a healthcare professional-No treatment-t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months-continuous outcomes- Patient/participant generic health-related quality of life (SF-36)-Mental health-Mean SD-Music intervention delivered by a healthcare professional-No treatment-t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months-continuous outcomes-Psychological distress- Depression (Beck Depression Inventory)-Mean SD-Music intervention delivered by a healthcare professional-No treatment-t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - continuous outcomes - Psychological distress (PANAS) - Positive affect - Mean SD - Music intervention delivered by a healthcare professional - No treatment - t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - continuous outcomes - Psychological distress (PANAS) - Negative affect - Mean SD - Music intervention delivered by a healthcare professional - No treatment - t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - dichotomous outcome - Withdrawal due to adverse events - No Of Events - Music intervention delivered by a healthcare professional - No treatment - t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | Low |
| Overall bias and Directness | Overall Directness | Directly applicable |

Hill, 2011**Bibliographic Reference**

Hill, V.; Dunn, L.; Dunning, K.; Page, S. J.; A pilot study of rhythm and timing training as a supplement to occupational therapy in stroke rehabilitation; Topics in stroke rehabilitation; 2011; vol. 18 (no. 6); 728-737

Study details

| | |
|---|--|
| Secondary publication of another included study- see primary study for details | No additional information. |
| Other publications associated with this study included in review | No additional information. |
| Trial name / registration number | No additional information. |
| Study type | Randomised controlled trial (RCT) |
| Study location | United States of America. |
| Study setting | Outpatient follow up. Delivered in hospital setting. |
| Study dates | No additional information. |
| Sources of funding | Interactive Metronome equipment and software provided for the study. |
| Inclusion criteria | Right hemiparesis; active flexion of the affected wrist a minimum of 10 degrees and active flexion of the metacarpophalangeal joints of the thumb and at least 2 additional metacarpophalangeal joints in 2 additional fingers at least 10 degrees; stroke experience 1 year or more prior to study enrolment; a score of >23 on the Mini-Mental Status Examination; between the age 21 and 85 years; have experienced one stroke resulting in residual hemiparesis; discharged from all forms of physical rehabilitation; able to follow directions in order to participate in therapy. |

| | |
|--|--|
| Exclusion criteria | Excessive pain in the affected hand, arm, or shoulder, as measured by a score >4 on a 10-point visual analogue scale; excessive spasticity in the affected biceps, triceps, wrist or fingers, as defined by a score >2 at any upper extremity joint on the Modified Ashworth Spasticity Scale; unilateral neglect as measured by observing the subject's interaction with their environment (ie, the subject's lack of attention to one side of their body and environment); absent bilateral or unilateral hearing as indicated by subject report; currently participating in any other experimental rehabilitation or drug studies; passive range of motion <45 degrees for abduction, flexion, or external rotation at shoulder or pronation of forearm of >30 degrees flexion contracture at any finger joint. |
| Recruitment / selection of participants | People were recruited by local rehabilitation therapists and by subject inquiry regarding current studies. |
| Intervention(s) | <p>Music intervention delivered by a healthcare professional N=6</p> <p>Interactive metronome intervention delivered by an occupational therapist. Completing the same exercises as the occupational therapy only group, with an additional 30 minute interactive metronome session. This consisted of a computer-based rhythmic and auditory training program where people had to match the beat from the metronome via a repetitive limb movement lasting for 1-3 minutes. The number of sessions ranged from 10 to 30 dependent on tolerance. Audiovisual guidance was used to provide immediate feedback and help the person get closer to the beat. Movements included clapping hands together, tapping unaffected hand or leg, tapping affected hand on leg, tapping foot on floor mat, and tapping hand on leg and then contralateral foot on floor. All completed both sitting and standing as people progressed through the protocol. 1 hour treatment, 3 times a week for 10 weeks.</p> <p>Concomitant therapy: Usual occupational therapy including prefunctional activities, functional activities and COPM (Canadian Occupational Performance Measure) tasks.</p> |
| Subgroup 1: Time after stroke at the start of the trial | Chronic (>6 months) |
| Subgroup 2: Severity (as stated by category or as | Not stated/unclear |

| | |
|---|--|
| measured by NIHSS scale) | |
| Subgroup 3: Group compared to individual sessions of music therapy | Individual sessions |
| Subgroup 4: Delivered in hospital compared to delivered in the community | Hospital sessions |
| Subgroup 5: Type of intervention | Rhythmic auditory cueing |
| Population subgroups | No additional information. |
| Comparator | No treatment N=4 Usual occupational therapy only. 1 hour treatment, 3 times a week for 10 weeks. Concomitant therapy: Usual occupational therapy including prefunctional activities, functional activities and COPM (Canadian Occupational Performance Measure) tasks. |
| Number of participants | 10 |
| Duration of follow-up | 10 weeks (end of intervention) |
| Indirectness | No additional information. |
| Additional comments | Intention to treat (no people discontinued the study). |

Study arms

Music intervention delivered by a healthcare professional (N = 6)

Interactive metronome intervention delivered by an occupational therapist. Completing the same exercises as the occupational therapy only group, with an additional 30 minute interactive metronome session. This consisted of a computer-based rhythmic and auditory training program where people had to match the beat from the metronome via a repetitive limb movement lasting for 1-3 minutes. The number of sessions ranged from 10 to 30 dependent on tolerance. Audiovisual guidance was used to provide immediate feedback and help the person get closer to the beat. Movements included clapping hands together, tapping unaffected hand or leg, tapping affected hand on leg, tapping foot on floor mat, and tapping hand on leg and then contralateral foot on floor. All completed both sitting and standing as people progressed through the protocol. 1 hour treatment, 3 times a week for 10 weeks. Concomitant therapy: Usual occupational therapy including prefunctional activities, functional activities and COPM (Canadian Occupational Performance Measure) tasks.

No treatment (N = 4)

Usual occupational therapy only. 1 hour treatment, 3 times a week for 10 weeks. Concomitant therapy: Usual occupational therapy including prefunctional activities, functional activities and COPM (Canadian Occupational Performance Measure) tasks.

Characteristics

Arm-level characteristics

| Characteristic | Music intervention delivered by a healthcare professional (N = 6) | No treatment (N = 4) |
|-----------------------|--|-----------------------------|
| % Female | n = 4 ; % = 67 | n = 2 ; % = 50 |
| Sample size | | |
| Mean age (SD) | 62.67 (8.21) | 56 (9.02) |

| Characteristic | Music intervention delivered by a healthcare professional (N = 6) | No treatment (N = 4) |
|----------------------------------|--|-----------------------------|
| Mean (SD) | | |
| Ethnicity | n = NA ; % = NA | n = NA ; % = NA |
| Sample size | | |
| Caucasian | n = 4 ; % = 67 | n = 3 ; % = 75 |
| Sample size | | |
| Comorbidities | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Time after stroke (years) | 3.67 (2.58) | 2.75 (1.71) |
| Mean (SD) | | |
| Severity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Type of stroke | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |

Outcomes

Study timepoints

- Baseline
- 10 week (End of intervention. <6 months.)

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - continuous outcomes

| Outcome | Music intervention delivered by a healthcare professional, Baseline, N = 6 | Music intervention delivered by a healthcare professional, 10 week, N = 6 | No treatment, Baseline, N = 4 | No treatment, 10 week, N = 4 |
|---|--|---|-------------------------------|------------------------------|
| Activities of daily living (Canadian Occupational Performance Measure - Performance) Scale range: Unclear (5-50?). Change score. Mean (SD) | 14 (7.81) | 15 (5.57) | 15.6 (7.37) | 10.2 (16.18) |
| Stroke-specific Patient Reported Outcome Measure (Stroke Impact Scale) Scale range: 0-100. Change score. Mean (SD) | 76.5 (13.48) | 1.25 (6.9) | 70.8 (32.33) | 7.4 (20) |

Activities of daily living (Canadian Occupational Performance Measure - Performance) - Polarity - Higher values are better
Stroke-specific Patient Reported Outcome Measure (Stroke Impact Scale) - Polarity - Higher values are better

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - dichotomous outcome

| Outcome | Music intervention delivered by a healthcare professional, Baseline, N = 6 | Music intervention delivered by a healthcare professional, 10 week, N = 6 | No treatment, Baseline, N = 4 | No treatment, 10 week, N = 4 |
|---|--|---|-------------------------------|------------------------------|
| Withdrawal due to adverse events No of events | n = NA ; % = NA | n = 0 ; % = 0 | n = NA ; % = NA | n = 0 ; % = 0 |

Withdrawal due to adverse events - Polarity - Lower values are better

Critical appraisal - Cochrane Risk of Bias tool (RoB 2.0) Normal RCT

Music intervention delivered by healthcare professionals compared to no treatment at <6 months-continuous outcomes-Activities of daily living (Canadian Occupational Performance Measure-Performance)-Mean SD-Music intervention delivered by a healthcare professional-No treatment-t10

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months-continuous outcomes-Stroke-specific Patient Reported Outcome Measure (Stroke Impact Scale)-Mean SD-Music intervention delivered by a healthcare professional-No treatment-t10

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months-dichotomous outcome-Withdrawal due to adverse events-No Of Events-Music intervention delivered by a healthcare professional-No treatment-t10

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Jeong, 2007**Bibliographic Reference**

Jeong, S.; Kim, M. T.; Effects of a theory-driven music and movement program for stroke survivors in a community setting; Applied Nursing Research; 2007; vol. 20 (no. 3); 125-31

Study details

| | |
|---|--|
| Secondary publication of another included study- see primary study for details | No additional information. |
| Other publications associated with this study included in review | No additional information. |
| Trial name / registration number | No additional information. |
| Study type | Randomised controlled trial (RCT) |
| Study location | Seoul, South Korea. |
| Study setting | A neighbourhood community health center located in the metropolitan area. |
| Study dates | No additional information. |
| Sources of funding | This study was supported by the BK21 project (Grant No. 0522-20010002), the Korea Science and Engineering Foundation (Grant No. R04-2001-000-00197-0), and the Research Institute of Nursing Science at Seoul National University. |

| | |
|---|--|
| Inclusion criteria | Have had a stroke at least 6 months earlier; scores of 2-4 on a muscle strength test (indicating poor to moderate muscle strength); disability on one side of the body; intact auditory function; ability to communicate with the research team without problems. |
| Exclusion criteria | Previous participation in a rehabilitation program. |
| Recruitment / selection of participants | Recruitment was conducted by trained research assistants, with referrals from the public health nurses working in the centre. |
| Intervention(s) | <p>Music intervention delivered by healthcare professionals N=16</p> <p>Rhythmic auditory stimulation music-movement program consisted of four sections: preparatory activities, main activities, wrap-up activities and follow up. This group activity was conducted for 2 hours/week for 8 weeks at a public health center in Seoul, South Korea.</p> <p>Concomitant therapy: No additional information.</p> |
| Subgroup 1: Time after stroke at the start of the trial | Chronic (>6 months) |
| Subgroup 2: Severity (as stated by category or as measured by NIHSS scale) | Not stated/unclear |
| Subgroup 3: Group compared to individual sessions of music therapy | Group sessions |
| Subgroup 4: Delivered in hospital compared | Community sessions |

| | |
|---|---|
| to delivered in the community | |
| Subgroup 5: Type of intervention | Rhythmic auditory cueing |
| Population subgroups | No additional information. |
| Comparator | No treatment N=17 Received referral information about available usual care services (available to both groups). Concomitant therapy: No additional information. |
| Number of participants | 36 (data is only available for 33). |
| Duration of follow-up | 8 weeks (end of intervention). |
| Indirectness | No additional information. |
| Additional comments | Not intention to treat. Missing cases. No additional information. |

Study arms

Music intervention delivered by healthcare professionals (N = 16)

Rhythmic auditory stimulation music-movement program consisted of four sections: preparatory activities, main activities, wrap-up activities and follow up. This group activity was conducted for 2 hours/week for 8 weeks at a public health center in Seoul, South Korea. Concomitant therapy: No additional information.

No treatment (N = 17)

Received referral information about available usual care services (available to both groups). Concomitant therapy: No additional information.

Characteristics**Arm-level characteristics**

| Characteristic | Music intervention delivered by healthcare professionals (N = 16) | No treatment (N = 17) |
|----------------------------------|--|------------------------------|
| % Female | n = 5 ; % = 31.3 | n = 5 ; % = 29.4 |
| Sample size | | |
| Mean age (SD) | 58 (7.19) | 62.2 (8.16) |
| Mean (SD) | | |
| Ethnicity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Comorbidities | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Time after stroke (years) | 5.44 (4.53) | 7.29 (5.3) |
| Mean (SD) | | |
| Severity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Type of stroke | n = NA ; % = NA | n = NA ; % = NA |

| Characteristic | Music intervention delivered by healthcare professionals (N = 16) | No treatment (N = 17) |
|--------------------|---|-----------------------|
| Sample size | | |
| Infarction | n = 9 ; % = 56.3 | n = 11 ; % = 64.7 |
| Sample size | | |
| Haemorrhage | n = 7 ; % = 43.8 | n = 6 ; % = 35.3 |
| Sample size | | |

Outcomes

Study timepoints

- Baseline
- 8 week (End of intervention. <6 months.)

Music intervention delivered by healthcare professionals compared to no treatment at <6 months

| Outcome | Music intervention delivered by healthcare professionals, Baseline, N = 16 | Music intervention delivered by healthcare professionals, 8 week, N = 16 | No treatment, Baseline, N = 17 | No treatment, 8 week, N = 17 |
|---|--|--|--------------------------------|------------------------------|
| Psychological distress (Profile of Mood states) Scale range: 0-136. Final values (but looks too small, ?transformed data). Mean (SD) | 2.11 (1.02) | 1.56 (0.82) | 2.81 (0.9) | 2.29 (0.77) |
| Stroke-specific Patient Reported Outcome Measures (Stroke | 3.25 (1.08) | 3.58 (0.87) | 2.54 (0.8) | 2.92 (0.8) |

| Outcome | Music intervention delivered by healthcare professionals, Baseline, N = 16 | Music intervention delivered by healthcare professionals, 8 week, N = 16 | No treatment, Baseline, N = 17 | No treatment, 8 week, N = 17 |
|--|--|--|--------------------------------|------------------------------|
| Specific Quality of Life) Scale range: 49-245. Final values (but looks too small, ?transformed data). Mean (SD) | | | | |

Psychological distress (Profile of Mood states) - Polarity - Lower values are better

Stroke-specific Patient Reported Outcome Measures (Stroke Specific Quality of Life) - Polarity - Higher values are better

Critical appraisal - Cochrane Risk of Bias tool (RoB 2.0) Normal RCT

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - Psychological distress (Profile of Mood states) - Mean SD - Music intervention delivered by healthcare professionals - No treatment - t8

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months-Stroke-specific Patient Reported Outcome Measures (Stroke Specific Quality of Life)-Mean SD-Music intervention delivered by healthcare professionals-No treatment-t8

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Jun, 2013

Bibliographic Reference

Jun, E. M.; Roh, Y. H.; Kim, M. J.; The effect of music-movement therapy on physical and psychological states of stroke patients; Journal of Clinical Nursing; 2013; vol. 22 (no. 12); 22-31

Study details

| | |
|---|----------------------------|
| Secondary publication of another included study- see primary study for details | No additional information. |
| Other publications associated with this study included in review | No additional information. |
| Trial name / registration number | No additional information. |

| | |
|--|---|
| Study type | Quasi- randomised controlled trial States it is quasi-experimental, but also states it is randomised. Will be included but downgraded for selection bias. |
| Study location | South Korea. |
| Study setting | Hospital inpatients. |
| Study dates | No additional information. |
| Sources of funding | This work was supported by Dong-eui University Foundation Grant (2011). |
| Inclusion criteria | An acute ischaemic stroke in the left or right temporal, frontal, parietal or subcortical brain regions; hospitalised less than 2 weeks; were fully conscious without L tube or T tube; could communicate verbally; had a Korean Mini-Mental State Examination Score >20 points; were able and willing to participate in the study |
| Exclusion criteria | Prior neurological or psychiatric disease; hearing deficit |
| Recruitment / selection of participants | The total number of people from the neurology unit were invited to participate. |
| Intervention(s) | Music therapy delivered by a trained music therapist N=20 Music movement therapy including 20 minutes of preparatory activities, 30 minutes of main activities and 10 minutes of finishing activities delivered 3 times per week for 8 weeks. Exercises were completed while people were using wheelchairs while accompanied by meditational music, or when using different musical instruments. This was followed up by people being given a chance to reflect and express their experiences. Concomitant therapy: No additional information |
| Subgroup 1: Time after stroke at the start of the trial | Acute (72 hours - 7 days) |
| Subgroup 2: Severity (as stated by category or as | Not stated/unclear |

| | |
|---|--|
| measured by NIHSS scale) | |
| Subgroup 3: Group compared to individual sessions of music therapy | Group sessions |
| Subgroup 4: Delivered in hospital compared to delivered in the community | Hospital sessions |
| Subgroup 5: Type of intervention | Any combination of the above Interventions where musical instruments are played and receptive interventions in which participants listen to music |
| Population subgroups | No additional information. |
| Comparator | No treatment N=20 Routine treatment only Concomitant therapy: No additional information |
| Number of participants | 40. |
| Duration of follow-up | 8 weeks (end of intervention). |
| Indirectness | No additional information. |
| Additional comments | Not intention to treat - excludes participants who did not complete the study. |

Study arms

Music therapy delivered by a trained music therapist (N = 20)

Music movement therapy including 20 minutes of preparatory activities, 30 minutes of main activities and 10 minutes of finishing activities delivered 3 times per week for 8 weeks. Exercises were completed while people were using wheelchairs while accompanied by meditational music, or when using different musical instruments. This was followed up by people being given a chance to reflect and express their experiences. Concomitant therapy: No additional information

No treatment (N = 20)

Routine treatment only Concomitant therapy: No additional information

Characteristics

Arm-level characteristics

| Characteristic | Music therapy delivered by a trained music therapist (N = 20) | No treatment (N = 20) |
|------------------------------|--|------------------------------|
| % Female | n = 9 ; % = 60 | n = 6 ; % = 40 |
| Sample size | | |
| Mean age (SD) (years) | 60.7 (8.59) | 55.1 (17.23) |
| Mean (SD) | | |
| Ethnicity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Comorbidities | n = NR ; % = NR | n = NR ; % = NR |

| Characteristic | Music therapy delivered by a trained music therapist (N = 20) | No treatment (N = 20) |
|--------------------------|--|------------------------------|
| Sample size | | |
| Time after stroke | NR (NR) | NR (NR) |
| Mean (SD) | | |
| Severity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Type of stroke | n = NA ; % = NA | n = NA ; % = NA |
| Sample size | | |
| Infarction | n = 13 ; % = 86.7 | n = 12 ; % = 80 |
| Sample size | | |
| Haemorrhage | n = 2 ; % = 13.3 | n = 3 ; % = 20 |
| Sample size | | |

Outcomes

Study timepoints

- Baseline
- 8 week (End of intervention. <6 months.)

Music therapy delivered by trained music therapists compared to no treatment at <6 months - continuous outcomes

| Outcome | Music therapy delivered by a trained music therapist, Baseline, N = 20 | Music therapy delivered by a trained music therapist, 8 week, N = 15 | No treatment, Baseline, N = 20 | No treatment, 8 week, N = 15 |
|--|--|--|--------------------------------|------------------------------|
| Activities of daily living (Korean-Modified Barthel Index) Scale range: 0-100. Change scores. Mean (SD) | 52.53 (20.02) | 9.2 (4.81) | 60.66 (23.44) | 7.2 (29.75) |
| Psychological distress - Depression (Center for Epidemiologic Studies Depression Scale) Scale range: 0-60. Change scores. Mean (SD) | 32.06 (11.34) | -6.46 (11.82) | 40.4 (13.05) | -9.67 (15.27) |

Activities of daily living (Korean-Modified Barthel Index) - Polarity - Higher values are better

Psychological distress - Depression (Center for Epidemiologic Studies Depression Scale) - Polarity - Lower values are better

Critical appraisal - Cochrane Risk of Bias tool (RoB 2.0) Normal RCT**Music therapy delivered by trained music therapists compared to no treatment at <6 months - continuous outcomes -****Activities of daily living (Korean-Modified Barthel Index) - Mean SD - Music therapy delivered by a trained music therapist - No treatment - t8**

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music therapy delivered by trained music therapists compared to no treatment at <6 months-continuous outcomes-Psychological distress-Depression (Center for Epidemiologic Studies Depression Scale)-Mean SD-Music therapy delivered by a trained music therapist-No treatment-t8

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Kim, 2011

Bibliographic Reference

Kim, D. S.; Park, Y. G.; Choi, J. H.; Im, S. H.; Jung, K. J.; Cha, Y. A.; Jung, C. O.; Yoon, Y. H.; Effects of music therapy on mood in stroke patients; Yonsei Medical Journal; 2011; vol. 52 (no. 6); 977-81

Study details

| | |
|---|-----------------------------------|
| Secondary publication of another included study- see primary study for details | No additional information. |
| Other publications associated with this study included in review | No additional information. |
| Trial name / registration number | No additional information. |
| Study type | Randomised controlled trial (RCT) |

| | |
|--|---|
| Study location | South Korea. |
| Study setting | Inpatient setting. |
| Study dates | No additional information. |
| Sources of funding | The authors have no financial conflicts of interest. |
| Inclusion criteria | Post-stroke patients, within 6 months of onset and mini mental status examination score of over 20 |
| Exclusion criteria | No additional information |
| Recruitment / selection of participants | People who were admitted to the Department of Rehabilitation Medicine of Gangnam Severance Hospital. |
| Intervention(s) | <p>Music therapy delivered by music therapists N=9</p> <p>Music therapy program following a standard 40-minute format carried out in accordance with the physical strength and individual characteristics of patients. The sessions consisted of a hello song and sharing of events in their lives (5 minutes); planned musical activities (30 minutes) including respiration and phonation, improvised play, hand bell play, singing, songwriting, and expression in tune with music; and share of feelings and a goodbye song (6 minutes). Keyboards, hand bells, percussion instruments, flutes and other tools such as picture cards, flowers and fruit scents were used in accordance with the planned activities. People were encouraged to improvise depending on their feelings and sing children's and folk songs. For 4 weeks.</p> <p>Concomitant therapy: All people received comprehensive rehabilitation treatment including physiotherapy, occupational therapy or speech therapy. All people received regular counselling by a licensed psychotherapist.</p> |
| Subgroup 1: Time after stroke at the start of the trial | Subacute (7 days - 6 months) |
| Subgroup 2: Severity (as stated by category or as | Not stated/unclear |

| | |
|---|---|
| measured by NIHSS scale) | |
| Subgroup 3: Group compared to individual sessions of music therapy | Not stated/unclear |
| Subgroup 4: Delivered in hospital compared to delivered in the community | Hospital sessions |
| Subgroup 5: Type of intervention | Interventions where musical instruments are played (including clinical improvisation) |
| Population subgroups | No additional information. |
| Comparator | No treatment N=9 No music intervention. Concomitant therapy: All people received comprehensive rehabilitation treatment including physiotherapy, occupational therapy or speech therapy. All people received regular counselling by a licensed psychotherapist. |
| Number of participants | 18. |
| Duration of follow-up | 4 weeks (end of intervention). |
| Indirectness | No additional information. |
| Additional comments | ITT (no dropouts). |

Study arms

Music therapy delivered by music therapists (N = 9)

Music therapy program following a standard 40-minute format carried out in accordance with the physical strength and individual characteristics of patients. The sessions consisted of a hello song and sharing of events in their lives (5 minutes); planned musical activities (30 minutes) including respiration and phonation, improvised play, hand bell play, singing, songwriting, and expression in tune with music; and share of feelings and a goodbye song (6 minutes). Keyboards, hand bells, percussion instruments, flutes and other tools such as picture cards, flowers and fruit scents were used in accordance with the planned activities. People were encouraged to improvise depending on their feelings and sing children's and folk songs. For 4 weeks. Concomitant therapy: All people received comprehensive rehabilitation treatment including physiotherapy, occupational therapy or speech therapy. All people received regular counselling by a licensed psychotherapist.

No treatment (N = 9)

No music intervention. Concomitant therapy: All people received comprehensive rehabilitation treatment including physiotherapy, occupational therapy or speech therapy. All people received regular counselling by a licensed psychotherapist.

Characteristics

Arm-level characteristics

| Characteristic | Music therapy delivered by music therapists (N = 9) | No treatment (N = 9) |
|------------------------------|--|-----------------------------|
| % Female | n = 1 ; % = 11 | n = 0 ; % = 0 |
| Sample size | | |
| Mean age (SD) (years) | 51.7 (13.5) | 47.3 (11.7) |
| Mean (SD) | | |

| Characteristic | Music therapy delivered by music therapists (N = 9) | No treatment (N = 9) |
|--------------------------|--|-----------------------------|
| Ethnicity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Comorbidities | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Time after stroke | NR (NR) | NR (NR) |
| Mean (SD) | | |
| Severity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Type of stroke | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |

Outcomes

Study timepoints

- Baseline
- 4 week (End of intervention. <6 months.)

Music therapy delivered by music therapists compared to no treatment at <6 months - continuous outcomes

| Outcome | Music therapy delivered by music therapists, Baseline, N = 9 | Music therapy delivered by music therapists, 4 week, N = 9 | No treatment, Baseline, N = 9 | No treatment, 4 week, N = 9 |
|--|--|--|-------------------------------|-----------------------------|
| Psychological distress - Depression (Beck Depression Inventory) Scale range: 0-63. Final value. Mean (SD) | 14.8 (6.4) | 12.4 (4.6) | 10.9 (2.5) | 11.1 (2.5) |
| Psychological distress - Anxiety (Beck Anxiety Inventory) Scale range: 0-63. Final value. Mean (SD) | 9.2 (4) | 9 (4.3) | 9.2 (4) | 9.2 (2.6) |

Psychological distress - Depression (Beck Depression Inventory) - Polarity - Lower values are better

Psychological distress - Anxiety (Beck Anxiety Inventory) - Polarity - Lower values are better

Music therapy delivered by music therapists compared to no treatment at <6 months - dichotomous outcome

| Outcome | Music therapy delivered by music therapists, Baseline, N = 9 | Music therapy delivered by music therapists, 4 week, N = 9 | No treatment, Baseline, N = 9 | No treatment, 4 week, N = 9 |
|---|--|--|-------------------------------|-----------------------------|
| Withdrawal due to adverse events No of events | n = NA ; % = NA | n = 0 ; % = 0 | n = NA ; % = NA | n = 0 ; % = 0 |

Withdrawal due to adverse events - Polarity - Lower values are better

Critical appraisal - Cochrane Risk of Bias tool (RoB 2.0) Normal RCT**Music therapy delivered by music therapists compared to no treatment at <6 months - continuous outcomes - Psychological distress - Depression (Beck Depression Inventory) - Mean SD - Music therapy delivered by music therapists - No treatment - t4**

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music therapy delivered by music therapists compared to no treatment at <6 months - continuous outcomes - Psychological distress - Anxiety (Beck Anxiety Inventory) - Mean SD - Music therapy delivered by music therapists - No treatment - t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music therapy delivered by music therapists compared to no treatment at <6 months - dichotomous outcome - Withdrawal due to adverse events - No Of Events - Music therapy delivered by music therapists - No treatment - t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | Some concerns |
| Overall bias and Directness | Overall Directness | Directly applicable |

Lin, 2017**Bibliographic Reference**

Lin, F.; Gu, Y.; Wu, Y.; Huang, D.; He, N.; Effect of music therapy derived from the five elements in Traditional Chinese Medicine on post-stroke depression; Journal of Traditional Chinese Medicine; 2017; vol. 37 (no. 5); 675-680

Study details

| | |
|---|---|
| Secondary publication of another included study- see primary study for details | No additional information |
| Other publications associated with this study included in review | No additional information |
| Trial name / registration number | No additional information |
| Study type | Randomised controlled trial (RCT) |
| Study location | China |
| Study setting | Inpatients |
| Study dates | March 2014 to February 2015. |
| Sources of funding | State Administration of Traditional Chinese Medicine of the People's Republic of China, State Clinical Research Base of Traditional Chinese Medicine, The Second Batch of Professional Skill Scientific and Research Special Project (No. JDZX2015127); Jiangsu Natural Science Foundation Youth Project (No. BK20171070); Nanjing Scientific Development Planned Project (No. 201402057) |
| Inclusion criteria | Met the Chinese and western medicine diagnostic criteria of ischaemic cerebral infarction and were diagnosed with cerebral infarction by skull CT and MRI; met the diagnostic criteria for depression, with a self-rating depression scale and a |

| | |
|---|--|
| | Hamilton's depression scale (HAMD-17) score >7 points; experienced secondary ischaemic cerebral stroke in the acute stage (2 weeks) within 6 months, and had depression symptoms that lasted for >2 weeks; were aged 45-85 years; had stable vital signs and clear consciousness and cooperated with physical examination with adequate communication ability; signed informed consent and voluntarily participated in the study. |
| Exclusion criteria | Did not meet the diagnostic criteria; experienced ischaemic stroke in the acute stage within 2 weeks or in the sequelae stage >6 months; were >85 years old; had severe diabetes or severe hepatorenal diseases; had unstable vital signs or mental disease; had dementia, disturbance of consciousness and/or aphasia that might have influenced their expression of feelings; had taken antidepressants in the previous 1 month; were allergic to Erigeron breviscapus or sertraline. |
| Recruitment / selection of participants | People hospitalised in the Department of Neurology in Guangzhou Hospital of Traditional Chinese Medicine |
| Intervention(s) | <p>Music intervention delivered by healthcare professionals N=30</p> <p>Acupuncture needling at Baihui (GV20) and acupoint injection at Yanglingquan (GB34) with a five phase music therapy intervention. Music was selected based on the principle of treating emotional disturbance with hyperaffectivity. Acupuncture was conducted during one of the daily music therapy times. Music therapy was administered twice daily, once in the morning and once in the afternoon for 20 minutes per session. All treatments were administered in a 5 day cycle, for three continuous cycles with an interval between two cycles of 1 day (17 days in total).</p> <p>Concomitant therapy: No additional information.</p> |
| Subgroup 1: Time after stroke at the start of the trial | Subacute (7 days - 6 months) |
| Subgroup 2: Severity (as stated by category or as measured by NIHSS scale) | Not stated/unclear |
| Subgroup 3: Group compared to | Individual sessions |

| | |
|---|--|
| individual sessions of music therapy | |
| Subgroup 4: Delivered in hospital compared to delivered in the community | Hospital sessions |
| Subgroup 5: Type of intervention | Receptive interventions in which participants listen to music |
| Population subgroups | No additional information. |
| Comparator | <p>No treatment N=30</p> <p>Acupuncture needling only for the same timings.</p> <p>Concomitant therapy: No additional information.</p> <p>A third arm was included in the study:</p> <p>Control N=32</p> <p>50 mg sertraline hydrochloride tablets (Pfizer Pharmaceutical Co., Ltd.; approval No. H10980141) in the morning - This group was not included in the analysis as they were not comparable to the music intervention arm.</p> |
| Number of participants | 92 (60 in our analysis) |
| Duration of follow-up | 17 days (end of intervention) |

| | |
|----------------------------|---------------------------|
| Indirectness | No additional information |
| Additional comments | ITT (no dropouts) |

Study arms

Music intervention delivered by healthcare professionals (N = 30)

Acupuncture needling at Baihui (GV20) and acupoint injection at Yanglingquan (GB34) with a five phase music therapy intervention. Music was selected based on the principle of treating emotional disturbance with hyperaffectivity. Acupuncture was conducted during one of the daily music therapy times. Music therapy was administered twice daily, once in the morning and once in the afternoon for 20 minutes per session. All treatments were administered in a 5 day cycle, for three continuous cycles with an interval between two cycles of 1 day (17 days in total). Concomitant therapy: No additional information.

No treatment (N = 30)

Acupuncture needling only for the same timings. Concomitant therapy: No additional information.

Characteristics

Arm-level characteristics

| Characteristic | Music intervention delivered by healthcare professionals (N = 30) | No treatment (N = 30) |
|------------------------------|--|------------------------------|
| % Female | n = 13 ; % = 43.33 | n = 20 ; % = 66.66 |
| Sample size | | |
| Mean age (SD) (years) | 68.8 (11.5) | 72.9 (10.4) |
| Mean (SD) | | |

| Characteristic | Music intervention delivered by healthcare professionals (N = 30) | No treatment (N = 30) |
|----------------------------------|--|------------------------------|
| Ethnicity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Comorbidities | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Time after stroke (years) | NR (NR) | NR (NR) |
| Mean (SD) | | |
| Severity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Type of stroke | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |

Outcomes

Study timepoints

- Baseline
- 17 day (End of intervention. <6 months.)

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - continuous outcomes

| Outcome | Music intervention delivered by healthcare professionals, Baseline, N = 30 | Music intervention delivered by healthcare professionals, 17 day, N = 30 | No treatment, Baseline, N = 30 | No treatment, 17 day, N = 30 |
|---|--|--|--------------------------------|------------------------------|
| Activities of daily living (Activities of daily living score) Scale range: Unclear. Change scores. Mean (SD) | 36 (17) | 18 (5) | 33 (22) | 12 (5) |
| Psychological distress - Depression (Hamilton's Depression Scale-17) Scale range: 0-56. Change scores. Mean (SD) | 19.8 (3.6) | -6.2 (2.1) | 19.5 (4.1) | -4.9 (1.5) |

Activities of daily living (Activities of daily living score) - Polarity - Higher values are better

Psychological distress - Depression (Hamilton's Depression Scale-17) - Polarity - Lower values are better

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - dichotomous outcome

| Outcome | Music intervention delivered by healthcare professionals, Baseline, N = 30 | Music intervention delivered by healthcare professionals, 17 day, N = 30 | No treatment, Baseline, N = 30 | No treatment, 17 day, N = 30 |
|---|--|--|--------------------------------|------------------------------|
| Withdrawal due to adverse events No of events | n = NA ; % = NA | n = 0 ; % = 0 | n = NA ; % = NA | n = 0 ; % = 0 |

Withdrawal due to adverse events - Polarity - Lower values are better

Critical appraisal - Cochrane Risk of Bias tool (RoB 2.0) Normal RCT**Music intervention delivered by healthcare professionals compared to no treatment at <6 months - continuous outcomes - Activities of daily living (Activities of daily living score) - Mean SD - Music intervention delivered by healthcare professionals - No treatment - t17**

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - continuous outcomes - Psychological distress - Depression (Hamilton's Depression Scale-17) - Mean SD - Music intervention delivered by healthcare professionals - No treatment - t17

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - dichotomous outcome - Withdrawal due to adverse events - No Of Events - Music intervention delivered by healthcare professionals - No treatment - t17

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | Some concerns |
| Overall bias and Directness | Overall Directness | Directly applicable |

Luft, 2004**Bibliographic Reference**

Luft, A. R.; McCombe-Waller, S.; Whittall, J.; Forrester, L. W.; Macko, R.; Sorkin, J. D.; Schulz, J. B.; Goldberg, A. P.; Hanley, D. F.; Repetitive bilateral arm training and motor cortex activation in chronic stroke: a randomized controlled trial; JAMA; 2004; vol. 292 (no. 15); 1853-61

Study details

| | |
|---|--|
| Secondary publication of another included study- see primary study for details | No additional information |
| Other publications associated with this study included in review | No additional information |
| Trial name / registration number | No additional information |
| Study type | Randomised controlled trial (RCT) |
| Study location | United States of America |
| Study setting | Outpatient rehabilitation care in hospital |
| Study dates | 2001 to 2004. |
| Sources of funding | This study was funded by National Institutes of Health grants from the National Institute on Aging (P60AG 12583); University of Maryland Claude D. Pepper Older Americans Independence Center, National Institute on Disability and Rehabilitation Research (H133G010111); the Baltimore Department of Veterans Affairs Geriatrics Research, Education and Clinical Center (GRECC); National Institute of Neurological Disorders and Stroke 1RO1 NS 24282-08; the France-Merrick Foundation; the Johns Hopkins GCRC (NCRR MO1-00052); and the Eleanor Naylor Dana Charitable Trust, Deutsche Forschungsgemeinschaft (Lu 748/2, 748/3). |

| | |
|---|--|
| Inclusion criteria | Residual upper extremity spastic hemiparesis following a single cortical or subcortical ischaemic stroke; ability to move the affected limb (at least partial range antigravity movement); had completed 3-6 months of conventional rehabilitation therapy; adequate language and neurocognitive function to understand instructions |
| Exclusion criteria | People with multiple clinical strokes; a history of other neurological disease; chronic pain; emotional disorders. |
| Recruitment / selection of participants | Conducted as part of the University of Maryland School of Medicine, National Institute on Aging–Claude D. Pepper Older Americans Independence Center in collaboration with the Johns Hopkins University Division of Brain Injury Outcomes. This study took participants from a larger study and completed fMRI scans with them (the larger study could not be identified from this paper). |
| Intervention(s) | <p>Music intervention delivered by healthcare professionals N=11</p> <p>Bilateral arm training with rhythmic auditory cueing in hour long therapy sessions (four 5 minute movement periods with 10 minute rest periods between) 3 times per week for 6 weeks. Auditory cues were determined for the individual between 0.67 to 0.97 Hz.</p> <p>Concomitant therapy: No additional information.</p> |
| Subgroup 1: Time after stroke at the start of the trial | Chronic (>6 months) |
| Subgroup 2: Severity (as stated by category or as measured by NIHSS scale) | Not stated/unclear |
| Subgroup 3: Group compared to individual sessions of music therapy | Individual sessions |
| Subgroup 4: Delivered in | Hospital sessions |

| | |
|--|---|
| hospital compared to delivered in the community | |
| Subgroup 5: Type of intervention | Rhythmic auditory cueing |
| Population subgroups | No additional information. |
| Comparator | No treatment N=15 Dose matched therapeutic exercise (same timings). Concomitant therapy: No additional information. |
| Number of participants | 26. |
| Duration of follow-up | 6 weeks (end of intervention). |
| Indirectness | No additional information. |
| Additional comments | Not ITT (missing cases were excluded). |

Study arms

Music intervention delivered by healthcare professionals (N = 11)

Bilateral arm training with rhythmic auditory cueing in hour long therapy sessions (four 5 minute movement periods with 10 minute rest periods between) 3 times per week for 6 weeks. Auditory cues were determined for the individual between 0.67 to 0.97 Hz.

Concomitant therapy: No additional information.

No treatment (N = 15)

Dose matched therapeutic exercise (same timings). Concomitant therapy: No additional information.

Characteristics**Arm-level characteristics**

| Characteristic | Music intervention delivered by healthcare professionals (N = 11) | No treatment (N = 15) |
|-----------------------------------|--|------------------------------|
| % Female | n = 2 ; % = 22.2 | n = 7 ; % = 58.3 |
| Sample size | | |
| Mean age (SD) | 63.3 (15.3) | 59.6 (10.5) |
| Mean (SD) | | |
| Ethnicity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Comorbidities | n = NR ; % = NR | n = NR ; % = NR |
| No of events | | |
| Time after stroke (Months) | 75 (37.9 to 84.5) | 45.5 (22.6 to 66.3) |
| Median (IQR) | | |
| Severity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Type of stroke | n = NA ; % = NA | n = NA ; % = NA |
| Sample size | | |

| Characteristic | Music intervention delivered by healthcare professionals (N = 11) | No treatment (N = 15) |
|--------------------|---|-----------------------|
| Cortical | n = 6 ; % = 66.7 | n = 6 ; % = 50 |
| Sample size | | |
| Subcortical | n = 2 ; % = 22.2 | n = 4 ; % = 33.3 |
| Sample size | | |
| Brainstem | n = 1 ; % = 11.1 | n = 2 ; % = 16.7 |
| Sample size | | |

Outcomes

Study timepoints

- Baseline
- 6 week (End of intervention. <6 months.)

Music interventions delivered by healthcare professionals compared to no treatment at <6 months - continuous outcome

| Outcome | Music intervention delivered by healthcare professionals, Baseline, N = 9 | Music intervention delivered by healthcare professionals, 6 week, N = 9 | No treatment, Baseline, N = 10 | No treatment, 6 week, N = 10 |
|---|---|---|--------------------------------|------------------------------|
| Activities of daily living (University of Maryland Arm Questionnaire for Stroke) Scale range: 0-4 (5 point ordinal scale). Change scores. | NR (NR) | 2 (1.4) | NR (NR) | 1.7 (2.11) |

| Outcome | Music intervention delivered by healthcare professionals, Baseline, N = 9 | Music intervention delivered by healthcare professionals, 6 week, N = 9 | No treatment, Baseline, N = 10 | No treatment, 6 week, N = 10 |
|-----------|---|---|--------------------------------|------------------------------|
| Mean (SE) | | | | |

Activities of daily living (University of Maryland Arm Questionnaire for Stroke) - Polarity - Higher values are better

Critical appraisal - Cochrane Risk of Bias tool (RoB 2.0) Normal RCT

Music interventions delivered by healthcare professionals compared to no treatment at <6 months - continuous outcome - Activities of daily living (University of Maryland Arm Questionnaire for Stroke) - Mean SE - Music intervention delivered by healthcare professionals - No treatment - t6

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Nayak, 2000

Bibliographic Reference

Nayak, S.; Wheeler, B. L.; Shiflett, S. C.; Agostinelli, S.; Effect of music therapy on mood and social interaction among individuals with acute traumatic brain injury and stroke; Rehabilitation Psychology; 2000; vol. 45 (no. 3); 274-283

Study details

| | |
|---|---|
| Secondary publication of another included study- see primary study for details | No additional information. |
| Other publications associated with this study included in review | No additional information. |
| Trial name / registration number | No additional information. |
| Study type | Randomised controlled trial (RCT) |
| Study location | United States of America. |
| Study setting | Inpatient facility at the Kessler Institute for Rehabilitation. |
| Study dates | No additional information. |
| Sources of funding | Support for this research was provided by National Institutes of Health Grant U24-HD32994. |
| Inclusion criteria | Moderate to severe impairments at admission (defined as an average Functional Independence Measure score of 4.5 or lower and depression score of 4 or higher on the 7-point Faces Scale). If the person's self-reported rating of mood on the Faces Scale did not meet the inclusion cutoff, a family member was also asked to rate their mood and if that value was 4 or higher, the person could enter the study. |
| Exclusion criteria | People with nasogastric tubes or intravenous lines; uncontrolled agitated behaviour; any serious medical condition; those with sensory, perceptual or marked cognitive impairments that might have interfered with their ability to participate in the study. |
| Recruitment / selection of participants | No additional information. |

| | |
|---|---|
| Intervention(s) | <p>Music therapy delivered by trained music therapists N=10</p> <p>Music therapy 3 treatments per week for up to 10 treatments (treatment two or three times per week). A variety of procedures were used based on the needs of the person. Each session began with an opening song or activity. This was frequently some type of instrumental improvisation using simple percussion and melodic instruments. The therapist may structure the improvisation by asking the person to express how they were feeling and for everyone else to play in a manner reflecting their mood. Additional activities could include singing, composing, playing instruments, improvising, performing and listening.</p> <p>Concomitant therapy: Conventional rehabilitation was provided to both groups.</p> |
| Subgroup 1: Time after stroke at the start of the trial | Acute (72 hours - 7 days) |
| Subgroup 2: Severity (as stated by category or as measured by NIHSS scale) | <p>Moderate (or NIHSS 5-14)</p> <p>Moderate-severe</p> |
| Subgroup 3: Group compared to individual sessions of music therapy | Group sessions |
| Subgroup 4: Delivered in hospital compared to delivered in the community | Hospital sessions |
| Subgroup 5: Type of intervention | <p>Any combination of the above</p> <p>Combination of musical instruments, singing and songwriting</p> |

| | |
|-------------------------------|--|
| Population subgroups | No additional information |
| Comparator | No treatment N=8 Standard rehabilitation. Concomitant therapy: No additional information (same as the intervention arm). |
| Number of participants | 18 |
| Duration of follow-up | 3 weeks (end of intervention) |
| Indirectness | People were included who could have had traumatic brain injury or stroke. The number of participants with each is not stated. This study has been included but will be downgraded due to indirectness. |
| Additional comments | ITT (no dropouts). |

Study arms

Music therapy delivered by trained music therapists (N = 10)

Music therapy 3 treatments per week for up to 10 treatments (treatment two or three times per week). A variety of procedures were used based on the needs of the person. Each session began with an opening song or activity. This was frequently some type of instrumental improvisation using simple percussion and melodic instruments. The therapist may structure the improvisation by asking the person to express how they were feeling and for everyone else to play in a manner reflecting their mood. Additional activities could include singing, composing, playing instruments, improvising, performing and listening. Concomitant therapy: Conventional rehabilitation was provided to both groups.

No treatment (N = 8)

Standard rehabilitation. Concomitant therapy: No additional information (same as the intervention arm).

Characteristics**Study-level characteristics**

| Characteristic | Study (N = 18) |
|------------------------------|-----------------------|
| Mean age (SD) (years) | 59.89 (16.3) |
| Mean (SD) | |
| Ethnicity | n = NR ; % = NR |
| Sample size | |
| Comorbidities | n = NR ; % = NR |
| Sample size | |
| Time after stroke | NR (NR) |
| Mean (SD) | |
| Severity | n = NR ; % = NR |
| No of events | |
| Type of stroke | n = NR ; % = NR |
| No of events | |

Arm-level characteristics

| Characteristic | Music therapy delivered by trained music therapists (N = 10) | No treatment (N = 8) |
|----------------|--|----------------------|
| % Female | n = 6 ; % = 60 | n = 6 ; % = 75 |
| Sample size | | |

Outcomes**Study timepoints**

- Baseline
- 3 week (End of intervention. <6 months.)

Music therapy delivered by trained music therapists compared to no treatment at <6 months - continuous outcomes

| Outcome | Music therapy delivered by trained music therapists, Baseline, N = 10 | Music therapy delivered by trained music therapists, 3 week, N = 10 | No treatment, Baseline, N = 8 | No treatment, 3 week, N = 8 |
|---|---|---|-------------------------------|-----------------------------|
| Psychological distress - Depression (Faces Scale) Scale range: 0-7. Final values. | 4.6 (1.71) | 2.8 (1.32) | 5 (1.41) | 3.88 (1.36) |
| Mean (SD) | | | | |
| Participation in leisure activities/social groups (Sickness Impact Profile Social Interaction Subscale) Scale range: 0-102. Final values. | 37.6 (7.55) | 29.6 (4.5) | 44.5 (7.19) | 42.88 (8.34) |
| Mean (SD) | | | | |

Psychological distress - Depression (Faces Scale) - Polarity - Lower values are better

Participation in leisure activities/social groups (Sickness Impact Profile Social Interaction Subscale) - Polarity - Lower values are better

Music therapy delivered by trained music therapists compared to no treatment at <6 months - dichotomous outcome

| Outcome | Music therapy delivered by trained music therapists, Baseline, N = 10 | Music therapy delivered by trained music therapists, 3 week, N = 10 | No treatment, Baseline, N = 8 | No treatment, 3 week, N = 8 |
|---|---|---|-------------------------------|-----------------------------|
| Withdrawal due to adverse events | n = NA ; % = NA | n = 0 ; % = 0 | n = NA ; % = NA | n = 0 ; % = 0 |
| No of events | | | | |

Withdrawal due to adverse events - Polarity - Lower values are better

Critical appraisal - Cochrane Risk of Bias tool (RoB 2.0) Normal RCT

Music therapy delivered by trained music therapists compared to no treatment at <6 months - continuous outcomes - Psychological distress - Depression (Faces Scale) - Mean SD - Music therapy delivered by trained music therapists - No treatment - t3

| Section | Question | Answer |
|-----------------------------|------------------------|--|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Partially applicable (Due to population indirectness) |

Music therapy delivered by trained music therapists compared to no treatment at <6 months-continuous outcomes- Participation in leisure activities/social groups (Sickness Impact Profile Social Interaction Subscale)-Mean SD-Music therapy delivered by trained music therapists-No treatment-t3

| Section | Question | Answer |
|-----------------------------|------------------------|--|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Partially applicable (Due to population indirectness) |

Music therapy delivered by trained music therapists compared to no treatment at <6 months-dichotomous outcome- Withdrawal due to adverse events-No Of Events-Music therapy delivered by trained music therapists-No treatment-t3

| Section | Question | Answer |
|-----------------------------|------------------------|--|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Partially applicable (Due to population indirectness) |

Palumbo, 2022

Bibliographic Reference Palumbo, Anna; Aluru, Viswanath; Battaglia, Jessica; Geller, Daniel; Turry, Alan; Ross, Marc; Cristian, Adrian; Balagula, Caitlin; Ogedegbe, Gbenga; Khatri, Latika; Chao, Moses V; Froemke, Robert C; Urbanek, Jacek K; Raghavan, Preeti; Music Upper Limb Therapy-Integrated Provides a Feasible Enriched Environment and Reduces Post-stroke Depression: A Pilot Randomized Controlled Trial.; American journal of physical medicine & rehabilitation; 2022; vol. 101 (no. 10); 937-946

Study details

| | |
|---|---|
| Secondary publication of another included study- see primary study for details | No additional information. |
| Other publications associated with this study included in review | No additional information. |
| Trial name / registration number | No additional information. |
| Study type | Randomised controlled trial (RCT) |
| Study location | New York, US. |
| Study setting | Kingsbrook Jewish Medical Center, a full-service teaching hospital. |
| Study dates | No additional information. |
| Sources of funding | This research was supported in part by New York University Clinical Translational Science Award UL1TR000038 from the National Center for Advancing Translational Sciences (NCATS) and U54NS081765, National Institutes of Health, and in part by grants from the GRAMMY Foundation and the John and Jennifer Clay Foundation. |
| Inclusion criteria | Adults with chronic hemiparesis from a stroke at least 1 month before and the ability to open and close the hand partially with the affected side, suggesting the presence of adequate neural substrate for recovery. |
| Exclusion criteria | People with hearing deficits that might affect response to music using the Hearing Handicap Inventory for Adults, severe aphasia, cognitive or perceptual deficits including inability to follow directions and attend to task, and motor and ideational apraxia or neglect that would prevent participation in the intervention. |
| Recruitment / selection of participants | A convenience sample of 30 adults with unilateral hemiparesis post-stroke were recruited from Kingsbrook Jewish Medical Center, a full-service teaching hospital located in a metropolitan area. |

| | |
|---|---|
| Intervention(s) | Music Upper Limb Therapy-Integrated (MULT-I) intervention consists of groups of five participants each which were led collaboratively by one occupational therapist and one music therapist. MULT-I utilized the Nordoff-Robbins approach to music therapy, which incorporates live, interactive music-making. The 45-min intervention was provided twice a week for 6 wks. |
| Subgroup 1: Time after stroke at the start of the trial | Subacute (7 days - 6 months) |
| Subgroup 2: Severity (as stated by category or as measured by NIHSS scale) | Not stated/unclear |
| Subgroup 3: Group compared to individual sessions of music therapy | Group sessions |
| Subgroup 4: Delivered in hospital compared to delivered in the community | Hospital sessions |
| Subgroup 5: Type of intervention | Interventions where musical instruments are played (including clinical improvisation) |
| Population subgroups | No additional information. |
| Comparator | Home Exercise Programme (HEP) participants received a packet including photos and written instructions for a HEP consisting of bimanual passive and active assisted range of motion exercises for the upper limb with a cane/stick. The exercises targeted repetitive upper limb movements including shoulder flexion, abduction and adduction, elbow flexion and extension, forearm supination and pronation, and wrist flexion and extension. HEP participants were instructed to perform the exercises for 45 mins twice a week for 6 weeks, matching the dose of physical movements in the MULT-I intervention, and track their sessions on a tracking sheet. |

| | |
|-------------------------------|---|
| Number of participants | 15 |
| Duration of follow-up | 6 weeks. |
| Indirectness | Comparator indirectness (does not match protocol) and unclear if exercises in comparator group were the same as those used in the intervention group. |
| Additional comments | No additional comments. |

Study arms

Music therapy delivered by trained music therapists (Music upper limb therapy-integrated [MULT-I]) (N = 15)

The MULT-I intervention consists of groups of five participants each which were led collaboratively by one occupational therapist and one music therapist. MULT-I utilized the Nordoff-Robbins approach to music therapy, which incorporates live, interactive music-making. The 45-min intervention was provided twice a week for 6 wks.

No treatment (Home exercise programme [HEP]) (N = 15)

HEP participants received a packet including photos and written instructions for a HEP consisting of bimanual passive and active assisted range of motion exercises for the upper limb with a cane/stick. The exercises targeted repetitive upper limb movements including shoulder flexion, abduction and adduction, elbow flexion and extension, forearm supination and pronation, and wrist flexion and extension. HEP participants were instructed to perform the exercises for 45 mins twice a week for 6 weeks, matching the dose of physical movements in the MULT-I intervention, and track their sessions on a tracking sheet.

Characteristics***Arm-level characteristics***

| Characteristic | Music therapy delivered by trained music therapists (Music upper limb therapy-integrated [MULT-I]) (N = 15) | No treatment (Home exercise programme [HEP]) (N = 15) |
|--------------------------|--|--|
| % Female | n = NR ; % = 61.54 | n = NR ; % = 33.33 |
| Sample size | | |
| Age | 61.23 (9.13) | 61.75 (12.75) |
| Mean (SD) | | |
| Ethnic group | n = NR ; % = 92.31 | n = NR ; % = 83.33 |
| Sample size | | |
| Comorbidities | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Time after stroke | 12.85 (18.45) | 28.5 (31.92) |
| Mean (SD) | | |
| Severity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Type of stroke | n = NR ; % = NR | n = NR ; % = NR |
| No of events | | |

Outcomes

Study timepoints

- Baseline
- 6 week (< 6 months)

Music upper limb therapy-integrated versus home exercise programme at < 6 months - continuous outcomes

| Outcome | Music therapy delivered by trained music therapists (Music upper limb therapy-integrated [MULT-I]), Baseline, N = 13 | Music therapy delivered by trained music therapists (Music upper limb therapy-integrated [MULT-I]), 6 week, N = 13 | No treatment (Home exercise programme [HEP]), Baseline, N = 12 | No treatment (Home exercise programme [HEP]), 6 week, N = 9 |
|--|--|--|--|---|
| Psychological distress - depression (Patient Health Questionnaire (PHQ)-9) Scale range: 0-27. Final values. Mean (SD) | 8.31 (6.34) | 3.15 (3) | 4.25 (4.39) | 3.44 (5.7) |

Psychological distress - depression (Patient Health Questionnaire (PHQ)-9) - Polarity - Lower values are better

Music upper limb therapy-integrated versus home exercise programme at < 6 months - continuous outcomes

| Outcome | Music therapy delivered by trained music therapists (Music upper limb therapy-integrated [MULT-I]), Baseline, N = 13 | Music therapy delivered by trained music therapists (Music upper limb therapy-integrated [MULT-I]), 6 week, N = 13 | No treatment (Home exercise programme [HEP]), Baseline, N = 12 | No treatment (Home exercise programme [HEP]), 6 week, N = 12 |
|--|--|--|--|--|
| Stroke-specific Patient-Reported Outcome Measures (Stroke | NR (NR) | NR (NR) | NR (NR) | NR (NR) |

| Outcome | Music therapy delivered by trained music therapists (Music upper limb therapy-integrated [MULT-I]), Baseline, N = 13 | Music therapy delivered by trained music therapists (Music upper limb therapy-integrated [MULT-I]), 6 week, N = 13 | No treatment (Home exercise programme [HEP]), Baseline, N = 12 | No treatment (Home exercise programme [HEP]), 6 week, N = 12 |
|---|---|---|---|---|
| Impact Scale) Scale range: 0-100. Final values. | | | | |
| Mean (SD) | | | | |
| SIS Physical strength | 27.31 (16.41) | 38.46 (14.05) | 38.33 (17.62) | 49.17 (18.93) |
| Mean (SD) | | | | |
| SIS Activities | 36.92 (17.68) | 46.94 (17.67) | 47.5 (13.99) | 53.33 (17.44) |
| Mean (SD) | | | | |
| SIS Mobility | 34.36 (21.99) | 47.69 (22.66) | 55.37 (18.24) | 58.7 (15.9) |
| Mean (SD) | | | | |
| SIS Hand use | 8.62 (18.1) | 19.08 (25.04) | 16 (21.37) | 31 (30.04) |
| Mean (SD) | | | | |
| SIS Memory | 64.62 (16.56) | 70.11 (15.19) | 68.33 (19.66) | 71.19 (17.34) |
| Mean (SD) | | | | |
| SIS Emotions | 53.5 (11.17) | 60.51 (13.76) | 60.74 (14.47) | 58.89 (14.17) |
| Mean (SD) | | | | |
| SIS Communications | 68.35 (16.06) | 71.43 (15.56) | 73.1 (14.23) | 73.57 (14.16) |
| Mean (SD) | | | | |

| Outcome | Music therapy delivered by trained music therapists (Music upper limb therapy-integrated [MULT-I]), Baseline, N = 13 | Music therapy delivered by trained music therapists (Music upper limb therapy-integrated [MULT-I]), 6 week, N = 13 | No treatment (Home exercise programme [HEP]), Baseline, N = 12 | No treatment (Home exercise programme [HEP]), 6 week, N = 12 |
|--|---|---|---|---|
| SIS Participation | 25.28 (13.67) | 46.54 (17.52) | 45.27 (16.76) | 55 (19.45) |
| Mean (SD) | | | | |
| SIS Recovery | 48.46 (16.12) | 61.54 (12.65) | 57.25 (13.88) | 62.75 (18.48) |
| Mean (SD) | | | | |
| Wellbeing Scores (World Health Organisation (WHO-5), five item well-being index) (five item well-being index) Scale range: 0-25. Final values. | 17.77 (7.32) | 21.46 (4.31) | 18.83 (5.13) | 19.08 (5.96) |
| Mean (SD) | | | | |

Stroke-specific Patient-Reported Outcome Measures (Stroke Impact Scale) - Polarity - Higher values are better

Wellbeing Scores (World Health Organisation (WHO-5), five item well-being index) - Polarity - Higher values are better

Critical appraisal - Cochrane Risk of Bias tool (RoB 2.0) Normal RCT**Music upper limb therapy-integrated versus home exercise programme at <6 months-continuous outcomes-Psychological distress-depression-MeanSD-Music upper limb therapy-integrated (MULT-I)-Home exercise programme (HEP)-t6**

| Section | Question | Answer |
|-----------------------------|------------------------|---|
| Overall bias and Directness | Risk of bias judgement | High (No information on allocation concealment, baseline differences, and unexplained loss of participants in control group) |
| Overall bias and Directness | Overall Directness | Partially applicable (Comparator indirectness (does not match protocol) and unclear if exercises in comparator group were the same as those used in the intervention group.) |

Music upper limb therapy-integrated versus home exercise programme at <6 months-continuous outcomes-SIS Physical strength-MeanSD-Music upper limb therapy-integrated (MULT-I)-Home exercise programme (HEP)-t6

| Section | Question | Answer |
|-----------------------------|------------------------|---|
| Overall bias and Directness | Risk of bias judgement | High (No information on allocation concealment and baseline differences) |
| Overall bias and Directness | Overall Directness | Partially applicable (Comparator indirectness (does not match protocol) and unclear if exercises in comparator group were the same as those used in the intervention group.) |

Music upper limb therapy-integrated versus home exercise programme at <6 months-continuous outcomes-SIS Activities-MeanSD-Music upper limb therapy-integrated (MULT-I)-Home exercise programme (HEP)-t6

| Section | Question | Answer |
|-----------------------------|------------------------|---|
| Overall bias and Directness | Risk of bias judgement | High (No information on allocation concealment and baseline differences) |

| Section | Question | Answer |
|-----------------------------|--------------------|---|
| Overall bias and Directness | Overall Directness | Partially applicable (Comparator indirectness (does not match protocol) and unclear if exercises in comparator group were the same as those used in the intervention group.) |

Musicupperlimbtherapy-integratedversushomeexerciseprogrammeat<6months-continuousoutcomes-SISMobility-MeanSD-Music upper limb therapy-integrated (MULT-I)-Home exercise programme (HEP)-t6

| Section | Question | Answer |
|-----------------------------|------------------------|---|
| Overall bias and Directness | Risk of bias judgement | High (No information on allocation concealment and baseline differences) |
| Overall bias and Directness | Overall Directness | Partially applicable (Comparator indirectness (does not match protocol) and unclear if exercises in comparator group were the same as those used in the intervention group.) |

Musicupperlimbtherapy-integratedversushomeexerciseprogrammeat<6months-continuousoutcomes-SISHanduse-MeanSD-Music upper limb therapy-integrated (MULT-I)-Home exercise programme (HEP)-t6

| Section | Question | Answer |
|-----------------------------|------------------------|---|
| Overall bias and Directness | Risk of bias judgement | High (No information on allocation concealment and baseline differences) |
| Overall bias and Directness | Overall Directness | Partially applicable (Comparator indirectness (does not match protocol) and unclear if exercises in comparator group were the same as those used in the intervention group.) |

Music upper limb therapy-integrated versus home exercise programme at <6 months-continuous outcomes-SIS Memory-Mean SD-Music upper limb therapy-integrated (MULT-I)-Home exercise programme (HEP)-t6

| Section | Question | Answer |
|-----------------------------|------------------------|---|
| Overall bias and Directness | Risk of bias judgement | High (No information on allocation concealment and baseline differences) |
| Overall bias and Directness | Overall Directness | Partially applicable (Comparator indirectness (does not match protocol) and unclear if exercises in comparator group were the same as those used in the intervention group.) |

Music upper limb therapy-integrated versus home exercise programme at <6 months-continuous outcomes-SIS Emotions-Mean SD-Music upper limb therapy-integrated (MULT-I)-Home exercise programme (HEP)-t6

| Section | Question | Answer |
|-----------------------------|------------------------|---|
| Overall bias and Directness | Risk of bias judgement | High (No information on allocation concealment and baseline differences) |
| Overall bias and Directness | Overall Directness | Partially applicable (Comparator indirectness (does not match protocol) and unclear if exercises in comparator group were the same as those used in the intervention group.) |

Music upper limb therapy-integrated versus home exercise programme at <6 months-continuous outcomes-SIS Communications-Mean SD-Music upper limb therapy-integrated (MULT-I)-Home exercise programme (HEP)-t6

| Section | Question | Answer |
|-----------------------------|------------------------|---|
| Overall bias and Directness | Risk of bias judgement | High (No information on allocation concealment and baseline differences) |

| Section | Question | Answer |
|-----------------------------|--------------------|---|
| Overall bias and Directness | Overall Directness | Partially applicable (Comparator indirectness (does not match protocol) and unclear if exercises in comparator group were the same as those used in the intervention group.) |

Music upper limb therapy-integrated versus home exercise programme at <6 months-continuous outcomes-SIS Participation-Mean SD-Music upper limb therapy-integrated (MULT-I)-Home exercise programme (HEP)-t6

| Section | Question | Answer |
|-----------------------------|------------------------|---|
| Overall bias and Directness | Risk of bias judgement | High (No information on allocation concealment and baseline differences) |
| Overall bias and Directness | Overall Directness | Partially applicable (Comparator indirectness (does not match protocol) and unclear if exercises in comparator group were the same as those used in the intervention group.) |

Music upper limb therapy-integrated versus home exercise programme at <6 months-continuous outcomes-SIS Recovery-Mean SD-Music upper limb therapy-integrated (MULT-I)-Home exercise programme (HEP)-t6

| Section | Question | Answer |
|-----------------------------|------------------------|---|
| Overall bias and Directness | Risk of bias judgement | High (No information on allocation concealment and baseline differences) |
| Overall bias and Directness | Overall Directness | Partially applicable (Comparator indirectness (does not match protocol) and unclear if exercises in comparator group were the same as those used in the intervention group.) |

Music upper limb therapy-integrated versus home exercise programme at <6 months-continuous outcomes-Wellbeing Scores-Mean SD-Music upper limb therapy-integrated (MULT-I)-Home exercise programme (HEP)-t6

| Section | Question | Answer |
|-----------------------------|------------------------|---|
| Overall bias and Directness | Risk of bias judgement | High (No information on allocation concealment and baseline differences) |
| Overall bias and Directness | Overall Directness | Partially applicable (Comparator indirectness (does not match protocol) and unclear if exercises in comparator group were the same as those used in the intervention group.) |

Pocwierz-Marciniak, 2017

Bibliographic Reference

Pocwierz-Marciniak, Ilona; Bidzan, Mariola; The influence of music therapy on quality of life after a stroke; Health psychology report; 2017; vol. 5 (no. 2); 173-185

Study details

| | |
|---|----------------------------|
| Secondary publication of another included study- see primary study for details | No additional information. |
| Other publications associated with this study included in review | No additional information. |
| Trial name / registration number | No additional information. |

| | |
|--|---|
| Study type | Randomised controlled trial (RCT) |
| Study location | Gdynia (Northern Poland). |
| Study setting | Inpatient neurological rehabilitation in hospital. |
| Study dates | No additional information. |
| Sources of funding | No additional information. |
| Inclusion criteria | First episode of stroke (ischaemic or haemorrhagic); presence of a motor disability; absence of evident cognitive or executive disorders; a score of at least 27 points on the Mini-Mental State Examination; a score of at least 16 points on the Frontal Assessment battery. |
| Exclusion criteria | No additional information. |
| Recruitment / selection of participants | No additional information. |
| Intervention(s) | <p>Neurologic music therapy delivered by trained music therapists N=30</p> <p>One-to-one sessions using mainly a receptive approach based on cognitive music therapy and guided imagery and music. Classical and film music compositions were used. This took place twice a week and was identical for each patient, involving 10 meetings which followed the same pattern. Each session was divided into three parts: an introduction, which involved describing the patient's current mood and doing breathing exercises while listening to music or using their voice; the main part, including addressing the session's main theme while listening to music, conversation and psychoeducational activities as well as music therapy exercises; and the final part, which consists of relaxation while listening to music and a summing-up of the conversation. The themes addressed were grouped into three cycles: "Around the illness", "Around emotions" and "Around interpersonal relations".</p> <p>Concomitant therapy: All people were undergoing an inpatient neurological rehabilitation in hospital and receiving standard care, including physiotherapy, ergotherapy, psychological diagnosis and maintenance psychotherapy.</p> |
| Subgroup 1: Time after stroke at the start of the trial | <p>Acute (72 hours - 7 days)</p> <p>Assumed as inpatient rehabilitation</p> |

| | |
|---|--|
| Subgroup 2: Severity (as stated by category or as measured by NIHSS scale) | Not stated/unclear |
| Subgroup 3: Group compared to individual sessions of music therapy | Individual sessions |
| Subgroup 4: Delivered in hospital compared to delivered in the community | Hospital sessions |
| Subgroup 5: Type of intervention | Receptive interventions in which participants listen to music |
| Population subgroups | No additional information |
| Comparator | No treatment N=31 No listening materials. Concomitant therapy: All people were undergoing an inpatient neurological rehabilitation in hospital and receiving standard care, including physiotherapy, ergotherapy, psychological diagnosis and maintenance psychotherapy. |
| Number of participants | 61 |
| Duration of follow-up | 5 weeks (end of intervention) |
| Indirectness | No additional information |

| | |
|----------------------------|---|
| Additional comments | ITT (no dropouts) - does not explicitly mention this but all people appear to be accounted for and does not mention any participant withdrawal. This study reports multiple measures for quality of life (SF-36 and the Cantril Ladder). As per the protocol, we only extracted the values for SF-36 as this was a high priority measure. |
|----------------------------|---|

Study arms

Neurologic music therapy delivered by trained music therapists (N = 30)

One-to-one sessions using mainly a receptive approach based on cognitive music therapy and guided imagery and music. Classical and film music compositions were used. This took place twice a week and was identical for each patient, involving 10 meetings which followed the same pattern. Each session was divided into three parts: an introduction, which involved describing the patient's current mood and doing breathing exercises while listening to music or using their voice; the main part, including addressing the session's main theme while listening to music, conversation and psychoeducational activities as well as music therapy exercises; and the final part, which consists of relaxation while listening to music and a summing-up of the conversation. The themes addressed were grouped into three cycles: "Around the illness", "Around emotions" and "Around interpersonal relations". Concomitant therapy: All people were undergoing an inpatient neurological rehabilitation in hospital and receiving standard care, including physiotherapy, ergotherapy, psychological diagnosis and maintenance psychotherapy.

No treatment (N = 31)

No listening materials. Concomitant therapy: All people were undergoing an inpatient neurological rehabilitation in hospital and receiving standard care, including physiotherapy, ergotherapy, psychological diagnosis and maintenance psychotherapy.

Characteristics

Arm-level characteristics

| Characteristic | Neurologic music therapy delivered by trained music therapists (N = 30) | No treatment (N = 31) |
|-----------------------|--|------------------------------|
| % Female | n = 16 ; % = 53.3 | n = 15 ; % = 51.6 |

| Characteristic | Neurologic music therapy delivered by trained music therapists (N = 30) | No treatment (N = 31) |
|------------------------------|--|------------------------------|
| Sample size | | |
| Mean age (SD) (years) | 44 to 84 | 47 to 84 |
| Range | | |
| Mean age (SD) (years) | 65 (NR) | 63 (NR) |
| Mean (SD) | | |
| Ethnicity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Comorbidities | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Time after stroke | NR (NR) | NR (NR) |
| Mean (SD) | | |
| Severity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Type of stroke | n = NA ; % = NA | n = NA ; % = NA |
| Sample size | | |
| Ischaemic | n = 22 ; % = 73.3 | n = 27 ; % = 87.1 |
| Sample size | | |
| Haemorrhagic | n = 8 ; % = 26.7 | n = 4 ; % = 12.9 |
| Sample size | | |

Outcomes

Study timepoints

- Baseline
- 5 week (End of follow up. <6 months.)

Neurologic music therapy delivered by trained music therapists compared to no treatment at <6 months - continuous outcomes

| Outcome | Neurologic music therapy delivered by trained music therapists, Baseline, N = 30 | Neurologic music therapy delivered by trained music therapists, 5 week, N = 30 | No treatment, Baseline, N = 31 | No treatment, 5 week, N = 31 |
|---|--|--|--------------------------------|------------------------------|
| Person/participant generic health-related quality of life (SF-36) Scale range: 0-100. Final values. | NA (NA) | NA (NA) | NA (NA) | NA (NA) |
| Mean (SD) | | | | |
| Physical functioning | 12.93 (3.01) | 17.43 (4.73) | 13.19 (4.22) | 16 (5.38) |
| Mean (SD) | | | | |
| Physical limitations (role physical) | 4.23 (0.5) | 5 (1.05) | 4.32 (0.83) | 4.71 (0.94) |
| Mean (SD) | | | | |
| Bodily pain | 8.53 (2.1) | 8.63 (2.31) | 7.23 (2.91) | 8.16 (2.67) |
| Mean (SD) | | | | |
| General health perceptions | 14.33 (3.69) | 17.43 (3.1) | 15.32 (3.44) | 15.97 (3.7) |
| Mean (SD) | | | | |

| Outcome | Neurologic music therapy delivered by trained music therapists, Baseline, N = 30 | Neurologic music therapy delivered by trained music therapists, 5 week, N = 30 | No treatment, Baseline, N = 31 | No treatment, 5 week, N = 31 |
|---|---|---|---------------------------------------|-------------------------------------|
| Vitality | 14.03 (3.83) | 18.63 (3.19) | 12.61 (4.29) | 14.52 (3.84) |
| Mean (SD) | | | | |
| Social functioning | 5.73 (2.27) | 7.37 (1.87) | 5.97 (2.86) | 6.81 (2.47) |
| Mean (SD) | | | | |
| Emotional limitations (role emotional) | 5.03 (1.27) | 5.83 (0.53) | 4.52 (1.36) | 5 (1.29) |
| Mean (SD) | | | | |
| Mental health | 20.03 (5.33) | 24.23 (3.67) | 18.23 (6.68) | 20.48 (5.81) |
| Mean (SD) | | | | |
| Stroke-specific Patient Reported Outcome Measures (Stroke adjusted Sickness Impact Profile 30) Scale range: 0-68. Final values. | 20.8 (4.26) | 14.9 (4.62) | 19.48 (4.24) | 16.42 (4.63) |
| Mean (SD) | | | | |

Person/participant generic health-related quality of life (SF-36) - Polarity - Higher values are better

Stroke-specific Patient Reported Outcome Measures (Stroke adjusted Sickness Impact Profile 30) - Polarity - Lower values are better

Critical appraisal - Cochrane Risk of Bias tool (RoB 2.0) Normal RCT**Neurologic music therapy delivered by trained music therapists compared to no treatment at <6 months-continuous outcomes- Person/participant generic health-related quality of life (SF-36)-Physical functioning-Mean SD-Neurologic music therapy delivered by trained music therapists-No treatment-t5**

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Neurologic music therapy delivered by trained music therapists compared to no treatment at <6 months-continuous outcomes- Person/participant generic health-related quality of life (SF-36)-Physical limitations (role physical)-Mean SD-Neurologic music therapy delivered by trained music therapists-No treatment-t5

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Neurologic music therapy delivered by trained music therapists compared to no treatment at <6 months-continuous outcomes- Person/participant generic health-related quality of life (SF-36)-Bodily pain-Mean SD-Neurologic music therapy delivered by trained music therapists-No treatment-t5

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Neurologic music therapy delivered by trained music therapists compared to no treatment at <6 months-continuous outcomes- Person/participant generic health-related quality of life (SF-36)-General health perceptions-Mean SD-Neurologic music therapy delivered by trained music therapists-No treatment-t5

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Neurologic music therapy delivered by trained music therapists compared to no treatment at <6 months-continuous outcomes- Person/participant generic health-related quality of life (SF-36)-Vitality-Mean SD-Neurologic music therapy delivered by trained music therapists-No treatment-t5

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Neurologic music therapy delivered by trained music therapists compared to no treatment at <6 months-continuous outcomes- Person/participant generic health-related quality of life (SF-36)-Social functioning-Mean SD-Neurologic music therapy delivered by trained music therapists-No treatment-t5

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Neurologic music therapy delivered by trained music therapists compared to no treatment at <6 months-continuous outcomes- Person/participant generic health-related quality of life (SF-36)-Emotional limitations (role emotional)-Mean SD-Neurologic music therapy delivered by trained music therapists-No treatment-t5

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Neurologic music therapy delivered by trained music therapists compared to no treatment at <6 months-continuous outcomes- Person/participant generic health-related quality of life (SF-36)-Mental health-Mean SD-Neurologic music therapy delivered by trained music therapists-No treatment-t5

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Neurologic music therapy delivered by trained music therapists compared to no treatment at <6 months-continuous outcomes-Stroke-specific Patient Reported Outcome Measures (Stroke adjusted Sickness Impact Profile 30)-Mean SD-Neurologic music therapy delivered by trained music therapists-No treatment-t5

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Raglio, 2021**Bibliographic Reference**

Raglio, A.; Panigazzi, M.; Colombo, R.; Tramontano, M.; Iosa, M.; Mastrogiacomo, S.; Baiardi, P.; Molteni, D.; Baldissarro, E.; Imbriani, C.; Imarisio, C.; Eretti, L.; Hamedani, M.; Pistarini, C.; Imbriani, M.; Mancardi, G. L.; Caltagirone, C.; Hand rehabilitation with sonification techniques in the subacute stage of stroke; Scientific Reports; 2021; vol. 11 (no. 1); 7237

Study details

| | |
|---|--|
| Secondary publication of another included study- see primary study for details | No additional information. |
| Other publications associated with this study included in review | No additional information. |
| Trial name / registration number | NCT03306797. |
| Study type | Randomised controlled trial (RCT) |
| Study location | Italy |
| Study setting | Rehabilitation units (outpatient follow up). |
| Study dates | March 27th 2017 to April 15th 2019 |
| Sources of funding | This work was partially supported by the "Ricerca Corrente" funding provided by the Italian Ministry of Health. |
| Inclusion criteria | Age 40-85 years; ischaemic lesion in a single hemisphere (right or left hemiplegia/hemiparesis); clinically evaluable residual movement capacity of the paretic upper limb (ability to autonomously make postural adjustments during reaching tasks); Mini Mental State Examination >24; acute onset no more than 180 days prior to enrollment in the study. |

| | |
|---|---|
| Exclusion criteria | Presence of neglect (evaluated through a clinical and functional assessment); previous or concomitant diseases affecting upper limb functions (e.g. Parkinson's disease, multiple sclerosis, shoulder periarthritis, Dupuytren's disease ect.); rehabilitation treatments with music in the last year. |
| Recruitment / selection of participants | People were recruited in 5 Italian rehabilitative units located at Maugeri Scientific Clinical Institutes of Pavia, Montescano and Nervi, S. Lucia Foundation IRCCS (Rome) and Neurological Clinic of S. Martino Hospital (University of Genoa). |
| Intervention(s) | <p>Music interventions delivered by healthcare professionals N=33</p> <p>Sonofication - properly selected set of sonorous-music stimuli activated by patient's movements with the mediation of a sensor (the Leap Motion Controller). Synthesized sounds/musical texture and their parameters (mainly rhythm, pitch/melody, intensity/dynamics, harmony and timbre) are used to represent movements characteristics, especially from a temporal and spatial point of view. Delivered as 35 minute sessions (passive treatment without sonification for 15 minutes, and a second phase for 20 minutes in which motor exercises were supported by sonification techniques). Delivered by physiotherapists or occupational therapists 5 days a week for 4 weeks, for a total of 20 sessions.</p> <p>Concomitant therapy: All people received usual care for people with subacute strokes (such as occupational therapy, speech therapy, psychological support, lower extremity rehabilitation ect.). All people received standard motor exercises (with or without sonification). The first phase included passive treatment (15 minutes) while the second phase included active movements (20 minutes).</p> |
| Subgroup 1: Time after stroke at the start of the trial | Subacute (7 days - 6 months) |
| Subgroup 2: Severity (as stated by category or as measured by NIHSS scale) | Not stated/unclear |
| Subgroup 3: Group compared to | Individual sessions |

| | |
|---|---|
| individual sessions of music therapy | |
| Subgroup 4: Delivered in hospital compared to delivered in the community | Hospital sessions |
| Subgroup 5: Type of intervention | Other Sonofication - synthesizer connected to the movements of the individual |
| Population subgroups | No additional information. |
| Comparator | No treatment N=32 No music intervention. Concomitant therapy: All people received usual care for people with subacute strokes (such as occupational therapy, speech therapy, psychological support, lower extremity rehabilitation ect.). All people received standard motor exercises (with or without sonofication). The first phase included passive treatment (15 minutes) while the second phase included active movements (20 minutes). |
| Number of participants | 65 |
| Duration of follow-up | 4 weeks of intervention, 1 month of additional follow up (2 months in total). |
| Indirectness | No additional information. |
| Additional comments | ITT and per-protocol. ITT data does not report the 2 month follow up, so per-protocol analysis needs to be used. |

Study arms

Music interventions delivered by healthcare professionals (N = 33)

Sonofication - properly selected set of sonorous-music stimuli activated by patient's movements with the mediation of a sensor (the Leap Motion Controller). Synthesized sounds/musical texture and their parameters (mainly rhythm, pitch/melody, intensity/dynamics, harmony and timbre) are used to represent movements characteristics, especially from a temporal and spatial point of view. Delivered as 35 minute sessions (passive treatment without sonification for 15 minutes, and a second phase for 20 minutes in which motor exercises were supported by sonification techniques). Delivered by physiotherapists or occupational therapists 5 days a week for 4 weeks, for a total of 20 sessions. Concomitant therapy: All people received usual care for people with subacute strokes (such as occupational therapy, speech therapy, psychological support, lower extremity rehabilitation ect.). All people received standard motor exercises (with or without sonofication). The first phase included passive treatment (15 minutes) while the second phase included active movements (20 minutes).

No treatment (N = 32)

No music intervention. Concomitant therapy: All people received usual care for people with subacute strokes (such as occupational therapy, speech therapy, psychological support, lower extremity rehabilitation ect.). All people received standard motor exercises (with or without sonofication). The first phase included passive treatment (15 minutes) while the second phase included active movements (20 minutes).

Characteristics

Arm-level characteristics

| Characteristic | Music interventions delivered by healthcare professionals (N = 33) | No treatment (N = 32) |
|------------------------------|---|------------------------------|
| % Female | n = 14 ; % = 46.8 | n = 16 ; % = 48.5 |
| Sample size | | |
| Mean age (SD) (years) | 64.7 (16) | 62.4 (8.9) |
| Mean (SD) | | |

| Characteristic | Music interventions delivered by healthcare professionals (N = 33) | No treatment (N = 32) |
|---------------------------------|---|------------------------------|
| Ethnicity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Comorbidities | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Time after stroke (days) | 31.75 | 38.25 |
| Interquartile Range | | |
| Time after stroke (days) | 29.5 (13 to 180) | 34.5 (12 to 180) |
| Median Min-Max | | |
| Severity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Type of stroke | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |

Outcomes

Study timepoints

- Baseline
- 2 month (1 month after the intervention finished (reported as T3 in the supplementary information). <6 months.)

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - continuous outcome

| Outcome | Music interventions delivered by healthcare professionals, Baseline, N = 13 | Music interventions delivered by healthcare professionals, 2 month, N = 13 | No treatment, Baseline, N = 16 | No treatment, 2 month, N = 16 |
|--|---|--|--------------------------------|-------------------------------|
| Person/participant generic health-related quality of life (McGill Quality of Life) Scale range: 0-10. Final values. Mean (SD) | 6.29 (1.67) | 6.85 (1.81) | 6.81 (1.44) | 7.34 (1.24) |

Person/participant generic health-related quality of life (McGill Quality of Life) - Polarity - Higher values are better

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - dichotomous outcome

| Outcome | Music interventions delivered by healthcare professionals, Baseline, N = 33 | Music interventions delivered by healthcare professionals, 2 month, N = 33 | No treatment, Baseline, N = 32 | No treatment, 2 month, N = 32 |
|--|---|--|--------------------------------|-------------------------------|
| Withdrawal due to adverse events Intervention: 1 hospitalised, 3 worsening of clinical condition. Control: 3 worsening of clinical condition. No of events | n = NA ; % = NA | n = 4 ; % = 12.1 | n = NA ; % = NA | n = 3 ; % = 9.38 |

Withdrawal due to adverse events - Polarity - Lower values are better

Critical appraisal - Cochrane Risk of Bias tool (RoB 2.0) Normal RCT

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - continuous outcome - Person/participant generic health-related quality of life (McGill Quality of Life) - Mean SD - Music interventions delivered by healthcare professionals - No treatment - t2

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - dichotomous outcome - Withdrawal due to adverse events - No Of Events - Music interventions delivered by healthcare professionals - No treatment - t2

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | Low |
| Overall bias and Directness | Overall Directness | Directly applicable |

Raglio, 2017**Bibliographic Reference**

Raglio, A.; Zaliani, A.; Baiardi, P.; Bossi, D.; Sguazzin, C.; Capodaglio, E.; Imbriani, C.; Gontero, G.; Imbriani, M.; Active music therapy approach for stroke patients in the post-acute rehabilitation; Neurological Sciences; 2017; vol. 38 (no. 5); 893-897

Study details

| | |
|---|--|
| Secondary publication of another included study- see primary study for details | No additional information. |
| Other publications associated with this study included in review | No additional information. |
| Trial name / registration number | No additional information. |
| Study location | Italy. |
| Study setting | Inpatients. |
| Study dates | No additional information. |
| Sources of funding | The authors received no financial support for the research, authorship, and/or publication of this article. |
| Inclusion criteria | Mini Mental State examination at least 18; age at least 40 years; stable clinical condition; absence of other neurologic and psychiatric diseases; sufficient autonomy in motor functions of upper limbs to use musical instruments, including in the music therapy setting; cooperation during music therapy interaction. |
| Exclusion criteria | People with a total aphasia; amusia; previous neurological or psychiatric diseases; those who underwent previous music therapy treatments or musical training or practice |
| Recruitment / selection of participants | People were recruited immediately after the acute phase (during a rehabilitation period of 6-8 weeks). |
| Intervention(s) | Music therapy delivered by trained music therapists N=19 |

| | |
|---|--|
| | <p>Relational active music therapy approach using rhythmical-melodic instrumentation (i.e. xylophones, glockenspiels, drums, bongos, ethnic percussions ect.) in a non-verbal setting. The music therapist invites people to play an active role and to interact using musical instruments. 20 sessions lasting 30 minutes each, three weekly (duration = around 7 weeks).</p> <p>Concomitant therapy: The standard of care treatment consisted of daily sessions of physiotherapy (passive/assisted active mobilization and neurorehabilitative techniques of paretic upper limbs, coordination and balance exercises, and gait training) and occupational therapy (exercises improving fine motor skills and recovering activities of daily living).</p> |
| Subgroup 1: Time after stroke at the start of the trial | Subacute (7 days - 6 months) |
| Subgroup 2: Severity (as stated by category or as measured by NIHSS scale) | Mild (or NIHSS 1-5) |
| Subgroup 3: Group compared to individual sessions of music therapy | Individual sessions |
| Subgroup 4: Delivered in hospital compared to delivered in the community | Hospital sessions |
| Subgroup 5: Type of intervention | Interventions where musical instruments are played (including clinical improvisation) |
| Population subgroups | No additional information. |

| | |
|-------------------------------|---|
| Comparator | No treatment N=19 No music intervention. Concomitant therapy: The standard of care treatment consisted of daily sessions of physiotherapy (passive/assisted active mobilization and neurorehabilitative techniques of paretic upper limbs, coordination and balance exercises, and gait training) and occupational therapy (exercises improving fine motor skills and recovering activities of daily living). |
| Number of participants | 38 |
| Duration of follow-up | 7 weeks (end of intervention). |
| Indirectness | No additional information. |
| Additional comments | No additional information (no statement about dropout). |

Study arms

Music therapy delivered by trained music therapists (N = 19)

Relational active music therapy approach using rhythmical-melodic instrumentation (i.e. xylophones, glockenspiels, drums, bongos, ethnic percussions ect.) in a non-verbal setting. The music therapist invites people to play an active role and to interact using musical instruments. 20 sessions lasting 30 minutes each, three weekly (duration = around 7 weeks). Concomitant therapy: The standard of care treatment consisted of daily sessions of physiotherapy (passive/assisted active mobilization and neurorehabilitative techniques of paretic upper limbs, coordination and balance exercises, and gait training) and occupational therapy (exercises improving fine motor skills and recovering activities of daily living).

No treatment (N = 19)

No music intervention. Concomitant therapy: The standard of care treatment consisted of daily sessions of physiotherapy (passive/assisted active mobilization and neurorehabilitative techniques of paretic upper limbs, coordination and balance exercises, and gait training) and occupational therapy (exercises improving fine motor skills and recovering activities of daily living).

Characteristics**Arm-level characteristics**

| Characteristic | Music therapy delivered by trained music therapists (N = 19) | No treatment (N = 19) |
|------------------------------|---|------------------------------|
| % Female | n = 11 ; % = 58 | n = 11 ; % = 58 |
| Sample size | | |
| Mean age (SD) (years) | 70.4 (8.9) | 75.4 (7.6) |
| Mean (SD) | | |
| Ethnicity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Comorbidities | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Time after stroke | NR (NR) | NR (NR) |
| Mean (SD) | | |
| Severity | 4.63 (2.29) | 5.05 (2.27) |
| Mean (SD) | | |

| Characteristic | Music therapy delivered by trained music therapists (N = 19) | No treatment (N = 19) |
|-----------------------|---|------------------------------|
| Type of stroke | n = NA | n = NA ; % = NA |
| Sample size | | |
| Ischaemic | n = 17 ; % = 89.5 | n = 18 ; % = 94.7 |
| Sample size | | |
| Haemorrhagic | n = 2 ; % = 10.5 | n = 1 ; % = 5.3 |
| Sample size | | |
| Aphasia | n = 1 ; % = 5.3 | n = 1 ; % = 5.3 |
| Sample size | | |
| Dysarthria | n = 9 ; % = 47.4 | n = 8 ; % = 42.1 |
| Sample size | | |

Outcomes

Study timepoints

- Baseline
- 7 week (End of intervention. <6 months.)

Music therapy delivered by music therapists compared to no treatment at <6 months - continuous outcomes

| Outcome | Music therapy delivered by trained music therapists, Baseline, N = 19 | Music therapy delivered by trained music therapists, 7 week, N = 19 | No treatment, Baseline, N = 19 | No treatment, 7 week, N = 19 |
|--|--|--|---------------------------------------|-------------------------------------|
| Person/participant generic health-related quality of life (McGill Quality of Life) Scale range: 0-10. Final values. Mean (SD) | 6.8 (1.88) | 7.76 (1.34) | 7.27 (1.86) | 7.49 (1.68) |
| Psychological distress - Depression (HADS-D) Scale range: 0-42. Final values. Mean (SD) | 7.18 (4.89) | 4.47 (3.57) | 3.8 (3.6) | 4.33 (3.11) |
| Psychological distress - Anxiety (HADS-A) Scale range: 0-42. Final values. Mean (SD) | 6.17 (4.29) | 4.83 (3.5) | 5.8 (3.69) | 5.73 (3.97) |
| Activities of daily living (functional independence measure) Scale range: 18-126. Final values. Mean (SD) | 76.58 (20.35) | 110.47 (9.9) | 71.26 (19.33) | 106.89 (16.83) |

Person/participant generic health-related quality of life (McGill Quality of Life) - Polarity - Higher values are better

Psychological distress - Depression (HADS-D) - Polarity - Lower values are better

Psychological distress - Anxiety (HADS-A) - Polarity - Lower values are better

Activities of daily living (functional independence measure) - Polarity - Higher values are better

Critical appraisal - Cochrane Risk of Bias tool (RoB 2.0) Normal RCT

Music therapy delivered by music therapists compared to no treatment at <6 months-continuous outcomes-Person/participant generic health-related quality of life (McGill Quality of Life)-Mean SD-Music therapy delivered by trained music therapists-No treatment-t7

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music therapy delivered by music therapists compared to no treatment at <6 months-continuous outcomes-Psychological distress-Depression (HADS-D)-Mean SD-Music therapy delivered by trained music therapists-No treatment-t7

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music therapy delivered by music therapists compared to no treatment at <6 months-continuous outcomes-Psychological distress-Anxiety (HADS-A)-Mean SD-Music therapy delivered by trained music therapists-No treatment-t7

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music therapy delivered by music therapists compared to no treatment at <6 months-continuous outcomes-Activities of daily living (functional independence measure)-Mean SD-Music therapy delivered by trained music therapists-No treatment-t7

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Tarrant, 2018

Bibliographic Reference

Tarrant, M.; Carter, M.; Dean, S. G.; Taylor, R. S.; Warren, F. C.; Spencer, A.; Adamson, J.; Landa, P.; Code, C.; Calitri, R.; Singing for people with aphasia (SPA): a protocol for a pilot randomised controlled trial of a group singing intervention to improve well-being; BMJ Open; 2018; vol. 8 (no. 9); e025167

Study details

| | |
|---|---|
| Secondary publication of another included study- see primary study for details | Tarrant, M., Carter, M., Dean, S. G. et al. (2021) Singing for people with aphasia (SPA): results of a pilot feasibility randomised controlled trial of a group singing intervention investigating acceptability and feasibility. BMJ Open 11(1): e040544 |
| Other publications associated with this study included in review | No additional information |

Tarrant, 2021**Bibliographic Reference**

Tarrant, M.; Carter, M.; Dean, S. G.; Taylor, R.; Warren, F. C.; Spencer, A.; Adamson, J.; Landa, P.; Code, C.; Backhouse, A.; Lamont, R. A.; Calitri, R.; Singing for people with aphasia (SPA): results of a pilot feasibility randomised controlled trial of a group singing intervention investigating acceptability and feasibility; BMJ Open; 2021; vol. 11 (no. 1); e040544

Study details

| | |
|---|--|
| Secondary publication of another included study- see primary study for details | No additional information. |
| Other publications associated with this study included in review | Tarrant, M., Carter, M., Dean, S. G. et al. (2018) Singing for people with aphasia (SPA): a protocol for a pilot randomised controlled trial of a group singing intervention to improve well-being. BMJ Open 8(9): e025167 |
| Study type | Randomised controlled trial (RCT) Mixed methods including an RCT and qualitative data |
| Study location | United Kingdom |
| Study setting | Three community settings: a church hall, a community centre and a dedicated music venue. |
| Study dates | May 2017 and April 2018. |
| Sources of funding | The trial is funded by the Stroke Association (QQ12/TSA 2016/14). Excess treatment costs have been covered by South Devon and Torbay Clinical Commissioning Group, North East and West Devon Clinical Commissioning group and the University of Exeter Medical School. The report is independent research supported by the National Institute for Health Research Applied Research Collaboration South West Peninsula. |
| Inclusion criteria | Diagnosis of aphasia following a stroke (confirmed by general practitioners); willingness to be randomised to either intervention and attend the intervention venue; conversational English pre-stroke; capacity to consent to participate |

| | |
|--|---|
| Exclusion criteria | Under the age of 18 years old; currently attending a speech and language therapy; intended to relocate outside of the geographic region; participating in another group intervention study; currently attending an existing singing or music group |
| Recruitment / selection of participants | Several routes: the South West Peninsula Clinical Research Network (Stroke); through speech and language therapists who provided study information and patient information sheets to potential participants; advertising through local support groups and on relevant websites; via local stroke support networks identified through national organisations (eg, the Stroke Association and Different strokes) and word of mouth, study flyers, adverts and information sheets placed in community settings and on the host university website. |
| Intervention(s) | <p>Music intervention delivered by non-healthcare professionals N=20</p> <p>Singing for people with aphasia intervention consisting of 10 weekly sessions delivered in a community facility across three sites in the South-West of England, with each session lasting 90 minutes. Sessions comprised approximately 45 minutes of group singing, with 45 minutes allocated to settling in/warm-up, mid-session break and departure. Three separate venues were used: a church hall, a community centre and a dedicated music venue. All venues were arranged such that there were separate singing and social areas. The sessions were facilitated by one of two experienced community music leaders who also provided music accompaniment (keyboard or guitar) and were supported by a 'singing champion'. Both facilitator and champion ran all sessions for their assigned group/s which ranged from six to seven participants. Small auxiliary percussion instruments were available for participants to play, and supported the engagement of participants with limited singing ability. Session content was flexible and included a range of songs, mainly popular 'classics', suggested by both facilitator and participants. Each group worked toward a final activity, either the development of a 'playlist' or a performance for family/friends to be delivered in the last session. Carers were welcome to support participants and joint in with the singing programme. The intervention was manualised and facilitators were trained in communication with people with aphasia.</p> <p>Concomitant therapy: All people received a resource pack in aphasia-friendly format, constructed for the purpose of the study, which provided information on living with aphasia and the available local community services.</p> |
| Subgroup 1: Time after stroke at the start of the trial | Chronic (>6 months) |
| Subgroup 2: Severity (as stated by category or as | Mild (or NIHSS 1-5) Based on mean aphasia severity |

| | |
|---|---|
| measured by NIHSS scale) | |
| Subgroup 3: Group compared to individual sessions of music therapy | Group sessions |
| Subgroup 4: Delivered in hospital compared to delivered in the community | Community sessions |
| Subgroup 5: Type of intervention | Singing and music-based voice interventions |
| Population subgroups | No additional information |
| Comparator | No treatment N=21 No additional treatment. Concomitant therapy: All people received a resource pack in aphasia-friendly format, constructed for the purpose of the study, which provided information on living with aphasia and the available local community services. |
| Number of participants | 41 |
| Duration of follow-up | 3 months and 6 months |
| Indirectness | Outcome indirectness - the adverse events outcome include serious adverse events rather than withdrawal due to adverse events (as this was not clearly stated). |

| | |
|----------------------------|---|
| Additional comments | Intention to treat - whereby participants were analysed according to their allocated group irrespective of the treatment received. Observed data only were used for the analysis, with no use of methods to address missing data (such as multiple imputation). |
|----------------------------|---|

Study arms

Music intervention delivered by non-healthcare professionals (N = 20)

Singing for people with aphasia intervention consisting of 10 weekly sessions delivered in a community facility across three sites in the South-West of England, with each session lasting 90 minutes. Sessions comprised approximately 45 minutes of group singing, with 45 minutes allocated to settling in/warm-up, mid-session break and departure. Three separate venues were used: a church hall, a community centre and a dedicated music venue. All venues were arranged such that there were separate singing and social areas. The sessions were facilitated by one of two experienced community music leaders who also provided music accompaniment (keyboard or guitar) and were supported by a 'singing champion'. Both facilitator and champion ran all sessions for their assigned group/s which ranged from six to seven participants. Small auxiliary percussion instruments were available for participants to play, and supported the engagement of participants with limited singing ability. Session content was flexible and included a range of songs, mainly popular 'classics', suggested by both facilitator and participants. Each group worked toward a final activity, either the development of a 'playlist' or a performance for family/friends to be delivered in the last session. Carers were welcome to support participants and joint in with the singing programme. The intervention was manualised and facilitators were trained in communication with people with aphasia. Concomitant therapy: All people received a resource pack in aphasia-friendly format, constructed for the purpose of the study, which provided information on living with aphasia and the available local community services.

No treatment (N = 21)

No additional treatment. Concomitant therapy: All people received a resource pack in aphasia-friendly format, constructed for the purpose of the study, which provided information on living with aphasia and the available local community services.

Characteristics***Arm-level characteristics***

| Characteristic | Music intervention delivered by non-healthcare professionals (N = 20) | No treatment (N = 21) |
|----------------------------------|--|------------------------------|
| % Female | n = 8 ; % = 40 | n = 8 ; % = 38 |
| Sample size | | |
| Mean age (SD) (years) | 65.2 (12.2) | 67.7 (8.3) |
| Mean (SD) | | |
| Ethnicity | n = NA ; % = NA | n = NA ; % = NA |
| Sample size | | |
| White | n = 20 ; % = 100 | n = 19 ; % = 90 |
| Sample size | | |
| Black | n = 0 ; % = 0 | n = 1 ; % = 5 |
| Sample size | | |
| Asian | n = 0 ; % = 0 | n = 1 ; % = 1 |
| Sample size | | |
| Comorbidities | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Time after stroke (years) | 4.6 (3.8) | 5.6 (6.7) |
| Mean (SD) | | |

| Characteristic | Music intervention delivered by non-healthcare professionals (N = 20) | No treatment (N = 21) |
|-------------------------------------|--|------------------------------|
| Severity Aphasia severity | n = NA ; % = NA | n = NA ; % = NA |
| Sample size | | |
| Mild | n = 14 ; % = 70 | n = 13 ; % = 62 |
| Sample size | | |
| Moderate | n = 3 ; % = 15 | n = 4 ; % = 19 |
| Sample size | | |
| Severe | n = 3 ; % = 15 | n = 4 ; % = 19 |
| Sample size | | |
| Type of stroke | n = NR ; % = NR | n = NR ; % = NR |
| No of events | | |

Outcomes

Study timepoints

- Baseline
- 3 month (<6 months)
- 6 month (≥6 months)

Music intervention delivered by non-healthcare professionals compared to no treatment at <6 months and ≥6 months - continuous outcomes

| Outcome | Music intervention delivered by non-healthcare professionals, Baseline, N = 20 | Music intervention delivered by non-healthcare professionals, 3 month, N = 17 | Music intervention delivered by non-healthcare professionals, 6 month, N = 18 | No treatment, Baseline, N = 21 | No treatment, 3 month, N = 19 | No treatment, 6 month, N = 16 |
|---|---|--|--|---------------------------------------|--------------------------------------|--------------------------------------|
| Person/participant generic health-related quality of life (EQ-5D-5L) Scale range: -0.11-1. Final values. Mean (SD) | 0.71 (0.24) | 0.68 (0.27) | 0.75 (0.24) | 0.73 (0.16) | 0.73 (0.24) | 0.65 (0.24) |
| Carer generic health-related quality of life (CarerQoL-7D) Scale range: 0-14. Final values. Mean (SD) | 9.2 (2.3) | NR (NR) | 9 (2.7) | 9.6 (2.6) | NR (NR) | 9 (3.1) |
| Wellbeing scores (ICEpop CAPability measure for adults) Scale range: 0-1. Final values. Mean (SD) | 0.83 (0.12) | 0.82 (0.096) | 0.81 (0.1) | 0.76 (0.16) | 0.75 (0.82) | 0.78 (0.16) |
| Stroke-specific Patient-Reported Outcome Measures (Stroke and Aphasia Quality of Life Scale) | 3.6 (0.8) | 3.6 (0.8) | 3.7 (0.8) | 3.5 (0.6) | 3.6 (0.7) | 3.4 (0.7) |

| Outcome | Music intervention delivered by non-healthcare professionals, Baseline, N = 20 | Music intervention delivered by non-healthcare professionals, 3 month, N = 17 | Music intervention delivered by non-healthcare professionals, 6 month, N = 18 | No treatment, Baseline, N = 21 | No treatment, 3 month, N = 19 | No treatment, 6 month, N = 16 |
|--|---|--|--|---------------------------------------|--------------------------------------|--------------------------------------|
| Scale range: 1-5. Final values. | | | | | | |
| Mean (SD) | | | | | | |
| Participation in leisure activities/social groups scores (modified Reintegration to Normal Living Index) Scale range: 0-100. Final values. | 20.4 (7.1) | 23.1 (5.3) | 23.9 (6.2) | 20.1 (6.7) | 22.1 (8) | 22.1 (7.5) |
| Mean (SD) | | | | | | |

Person/participant generic health-related quality of life (EQ-5D-5L) - Polarity - Higher values are better

Carer generic health-related quality of life (CarerQoL-7D) - Polarity - Higher values are better

Wellbeing scores (ICEpop CAPability measure for adults) - Polarity - Higher values are better

Stroke-specific Patient-Reported Outcome Measures (Stroke and Aphasia Quality of Life Scale) - Polarity - Higher values are better

Participation in leisure activities/social groups scores (modified Reintegration to Normal Living Index) - Polarity - Higher values are better

Music intervention delivered by non-healthcare professionals compared to no treatment at <6 months and ≥6 months - dichotomous outcome

| Outcome | Music intervention delivered by non-healthcare professionals, Baseline, N = 20 | Music intervention delivered by non-healthcare professionals, 3 month, N = 20 | Music intervention delivered by non-healthcare professionals, 6 month, N = 20 | No treatment, Baseline, N = 21 | No treatment, 3 month, N = 21 | No treatment, 6 month, N = 21 |
|---|--|---|---|--------------------------------|-------------------------------|-------------------------------|
| <p>Serious adverse events The study does not report withdrawal due to adverse events. However, this will be included as indirect evidence. Events: Intervention arm: 1 person (fall and death), 1 person (lung infection and fall). Control: No events. Data reported overall but to avoid double counting has been counted at the 3 month period.</p> <p>No of events</p> | n = NA ; % = NA | n = 2 ; % = 10 | n = NA ; % = NA | n = NA ; % = NA | n = 0 ; % = 0 | n = NA ; % = NA |

Serious adverse events - Polarity - Lower values are better

Critical appraisal - Cochrane Risk of Bias tool (RoB 2.0) Normal RCT

Music intervention delivered by non-healthcare professionals compared to no treatment at <6 months and ≥6 months - continuous outcomes - Person/participant generic health-related quality of life (EQ-5D-5L) - Mean SD - Music intervention delivered by non-healthcare professionals - No treatment - t3

| Section | Question | Answer |
|-----------------------------|------------------------|---------------|
| Overall bias and Directness | Risk of bias judgement | Some concerns |

| Section | Question | Answer |
|-----------------------------|--------------------|---------------------|
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by non-healthcare professionals compared to no treatment at <6 months and ≥6 months-continuous outcomes- Person/participant generic health-related quality of life (EQ-5D-5L)-Mean SD-Music intervention delivered by non-healthcare professionals-No treatment-t6

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by non-healthcare professionals compared to no treatment at <6 months and ≥6 months-continuous outcomes- Carer generic health-related quality of life (Carer QoL-7D)-Mean SD-Music intervention delivered by non-healthcare professionals-No treatment-t6

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by non-healthcare professionals compared to no treatment at <6 months and ≥6 months-continuous outcomes- Wellbeing scores (ICEpopCAP ability measure for adults)-Mean SD-Music intervention delivered by non-healthcare professionals-No treatment-t3

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by non-healthcare professionals compared to no treatment at <6 months and ≥6 months-continuous outcomes- Wellbeing scores (ICEpopCAP ability measure for adults)-Mean SD-Music intervention delivered by non-healthcare professionals-No treatment-t6

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by non-healthcare professionals compared to no treatment at <6 months and ≥6 months-continuous outcomes- Stroke-specific Patient-Reported Outcome Measures (Stroke and Aphasia Quality of Life Scale)-Mean SD-Music intervention delivered by non-healthcare professionals-No treatment-t3

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | Some concerns |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by non-healthcare professionals compared to no treatment at <6 months and ≥6 months-continuous outcomes-Stroke-specific Patient-Reported Outcome Measures (Stroke and Aphasia Quality of Life Scale)-Mean SD-Music intervention delivered by non-healthcare professionals-No treatment-t6

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by non-healthcare professionals compared to no treatment at <6 months and ≥6 months-continuous outcomes-Participation in leisure activities/social group scores(modified Reintegration to Normal Living Index)-Mean SD-Music intervention delivered by non-healthcare professionals-No treatment-t3

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | Some concerns |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by non-healthcare professionals compared to no treatment at <6 months and ≥6 months-continuous outcomes-Participation in leisure activities/social group scores(modified Reintegration to Normal Living Index)-Mean SD-Music intervention delivered by non-healthcare professionals-No treatment-t6

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by non-healthcare professionals compared to no treatment at <6 months and ≥6 months - dichotomous outcome - Serious adverse events - No Of Events - Music intervention delivered by non-healthcare professionals - No treatment - t3

| Section | Question | Answer |
|-----------------------------|------------------------|---|
| Overall bias and Directness | Risk of bias judgement | Low |
| Overall bias and Directness | Overall Directness | Partially applicable (Due to outcome indirectness (serious adverse events rather than withdrawal due to adverse events)) |

Tian, 2020

Bibliographic Reference

Tian, R.; Zhang, B.; Zhu, Y.; Rhythmic Auditory Stimulation as an Adjuvant Therapy Improved Post-stroke Motor Functions of the Upper Extremity: A Randomized Controlled Pilot Study; Frontiers in Neuroscience; 2020; vol. 14; 649

Study details

| | |
|---|--|
| Secondary publication of another included study- see primary study for details | No additional information |
| Other publications associated with this study included in review | No additional information |
| Trial name / registration number | Chinese Clinical Trial Registry Number1900026665 |

| | |
|--|--|
| Study type | Randomised controlled trial (RCT) |
| Study location | China. |
| Study setting | Inpatient |
| Study dates | No additional information. |
| Sources of funding | Supported by the Project fund of Shanghai Science and technology commission. The project number was 18411962300 |
| Inclusion criteria | Confirmed diagnosis of stroke with evidence on MRI or CT; having motor impairments in the upper extremity with a Brunnstrom Stages IV-VI; first-time stroke with or without previous lacunar infarction which resulted in no functional consequences; 40-80 years old; vital signs stable; inpatient rehabilitation status |
| Exclusion criteria | Having Parkinson's Disease or other neurological conditions causing motor dysfunction; having cognitive (MMSE <24) or auditory (tuning-fork test) impairment; having cancer or severe cardiopulmonary diseases; participating in other research projects; unable to follow commands; having pacemaker placement. |
| Recruitment / selection of participants | No additional information. |
| Intervention(s) | <p>Music intervention delivered by healthcare professionals N=16</p> <p>Rhythmic auditory stimulation 30 minutes every day, 5 days per week for 4 weeks. Paced activities based the applicable tempo that was gradually increased during the study. No more than a 5% increase was allowed in a day.</p> <p>Concomitant therapy: Everyone received 30 minutes individualised physical therapy and 30 minutes individualised occupational therapy per day, 5 days per week for 4 weeks.</p> |
| Subgroup 1: Time after stroke at the start of the trial | Subacute (7 days - 6 months) |
| Subgroup 2: Severity (as stated by category or as | Not stated/unclear |

| | |
|---|--|
| measured by NIHSS scale) | |
| Subgroup 3: Group compared to individual sessions of music therapy | Individual sessions |
| Subgroup 4: Delivered in hospital compared to delivered in the community | Hospital sessions |
| Subgroup 5: Type of intervention | Rhythmic auditory cueing |
| Population subgroups | No additional information. |
| Comparator | No treatment N=16 An additional 15 minutes regular physical therapy and 15 minutes of regular occupational therapy per day. Concomitant therapy: Everyone received 30 minutes individualised physical therapy and 30 minutes individualised occupational therapy per day, 5 days per week for 4 weeks. |
| Number of participants | 32 |
| Duration of follow-up | 4 weeks (end of intervention) |
| Indirectness | No additional information |
| Additional comments | No additional information |

Study arms

Music intervention delivered by healthcare professionals (N = 16)

Rhythmic auditory stimulation 30 minutes every day, 5 days per week for 4 weeks. Paced activities based the applicable tempo that was gradually increased during the study. No more than a 5% increase was allowed in a day. Concomitant therapy: Everyone received 30 minutes individualised physical therapy and 30 minutes individualised occupational therapy per day, 5 days per week for 4 weeks.

No treatment (N = 16)

An additional 15 minutes regular physical therapy and 15 minutes of regular occupational therapy per day. Concomitant therapy: Everyone received 30 minutes individualised physical therapy and 30 minutes individualised occupational therapy per day, 5 days per week for 4 weeks.

Characteristics

Arm-level characteristics

| Characteristic | Music intervention delivered by healthcare professionals (N = 16) | No treatment (N = 16) |
|------------------------------|--|------------------------------|
| % Female | n = 2 ; % = 13.3 | n = 5 ; % = 33.3 |
| Sample size | | |
| Mean age (SD) (years) | 66.67 (13.59) | 64.4 (13.41) |
| Mean (SD) | | |
| Ethnicity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |

| Characteristic | Music intervention delivered by healthcare professionals (N = 16) | No treatment (N = 16) |
|-----------------------------------|--|------------------------------|
| Comorbidities | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Time after stroke (Months) | 5 (7.55) | 3.77 (9) |
| Mean (SD) | | |
| Severity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Type of stroke | n = NA ; % = NA | n = NA ; % = NA |
| Sample size | | |
| Frontal temporal lobe | n = 1 ; % = 7 | n = 0 ; % = 0 |
| Sample size | | |
| Frontal lobe | n = 0 ; % = 0 | n = 1 ; % = 7 |
| Sample size | | |
| Corona radiate | n = 1 ; % = 7 | n = 0 ; % = 0 |
| Sample size | | |
| Capsule externa | n = 1 ; % = 7 | n = 0 ; % = 0 |
| Sample size | | |
| Thalamus | n = 2 ; % = 13 | n = 1 ; % = 7 |
| Sample size | | |

| Characteristic | Music intervention delivered by healthcare professionals (N = 16) | No treatment (N = 16) |
|------------------------|---|-----------------------|
| Basal ganglia | n = 9 ; % = 59 | n = 8 ; % = 52 |
| Sample size | | |
| Brainstem | n = 1 ; % = 7 | n = 3 ; % = 20 |
| Sample size | | |
| Paraventricular | n = 0 ; % = 0 | n = 1 ; % = 7 |
| Sample size | | |
| Cerebellum | n = 0 ; % = 0 | n = 1 ; % = 7 |
| Sample size | | |

Outcomes

Study timepoints

- Baseline
- 4 week (End of intervention. <6 months.)

Music intervention delivered by healthcare professional compared to no treatment at <6 months - continuous outcome

| Outcome | Music intervention delivered by healthcare professionals, Baseline, N = 16 | Music intervention delivered by healthcare professionals, 4 week, N = 15 | No treatment, Baseline, N = 16 | No treatment, 4 week, N = 15 |
|---|--|--|--------------------------------|------------------------------|
| Activities of daily living (barthel index) | 60.67 (10.33) | 80.33 (8.96) | 60.33 (6.4) | 69.67 (7.19) |

| Outcome | Music intervention delivered by healthcare professionals, Baseline, N = 16 | Music intervention delivered by healthcare professionals, 4 week, N = 15 | No treatment, Baseline, N = 16 | No treatment, 4 week, N = 15 |
|--------------------------------------|--|--|--------------------------------|------------------------------|
| Scale range: 0-100. Final values. | | | | |
| Mean (SD) | | | | |

Activities of daily living (barthel index) - Polarity - Higher values are better

Music intervention delivered by healthcare professional compared to no treatment at <6 months - dichotomous outcome

| Outcome | Music intervention delivered by healthcare professionals, Baseline, N = 16 | Music intervention delivered by healthcare professionals, 4 week, N = 16 | No treatment, Baseline, N = 16 | No treatment, 4 week, N = 16 |
|---|--|--|--------------------------------|------------------------------|
| Withdrawal due to adverse events | n = NA ; % = NA | n = 0 ; % = 0 | n = NA ; % = NA | n = 0 ; % = 0 |
| No of events | | | | |

Withdrawal due to adverse events - Polarity - Lower values are better

Critical appraisal - Cochrane Risk of Bias tool (RoB 2.0) Normal RCT

Music intervention delivered by healthcare professional compared to no treatment at <6 months - continuous outcome - Activities of daily living (barthel index) - Mean SD - Music intervention delivered by healthcare professionals - No treatment - t4

| Section | Question | Answer |
|-----------------------------|------------------------|--------|
| Overall bias and Directness | Risk of bias judgement | High |

| Section | Question | Answer |
|-----------------------------|--------------------|---------------------|
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professional compared to no treatment at <6 months - dichotomous outcome - Withdrawal due to adverse events - No Of Events - Music intervention delivered by healthcare professionals - No treatment - t4

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | Some concerns |
| Overall bias and Directness | Overall Directness | Directly applicable |

van Delden, 2009

Bibliographic Reference

van Delden, A. L.; Peper, C. L.; Harlaar, J.; Daffertshofer, A.; Zijp, N. I.; Nienhuys, K.; Koppe, P.; Kwakkel, G.; Beek, P. J.; Comparing unilateral and bilateral upper limb training: the ULTRA-stroke program design; BMC Neurology; 2009; vol. 9; 57

Study details

| | |
|---|---|
| Secondary publication of another included study- see primary study for details | van Delden, A. L.; Peper, C. L.; Nienhuys, K. N.; Zijp, N. I.; Beek, P. J.; Kwakkel, G.; Unilateral versus bilateral upper limb training after stroke: the Upper Limb Training After Stroke clinical trial; Stroke; 2013; vol. 44 (no. 9); 2613-6 |
| Other publications associated with | No additional information |

**this study included
in review****van Delden, 2013****Bibliographic
Reference**

van Delden, A. L.; Peper, C. L.; Nienhuys, K. N.; Zijp, N. I.; Beek, P. J.; Kwakkel, G.; Unilateral versus bilateral upper limb training after stroke: the Upper Limb Training After Stroke clinical trial; Stroke; 2013; vol. 44 (no. 9); 2613-6

Study details

| | |
|---|---|
| Secondary publication of another included study- see primary study for details | No additional information |
| Other publications associated with this study included in review | van Delden, A. L.; Peper, C. L.; Harlaar, J.; Daffertshofer, A.; Zijp, N. I.; Nienhuys, K.; Koppe, P.; Kwakkel, G.; Beek, P. J.; Comparing unilateral and bilateral upper limb training: the ULTRA-stroke program design; BMC Neurology; 2009; vol. 9; 57 |
| Trial name / registration number | ULTRA-Stroke |
| Study type | Randomised controlled trial (RCT) |
| Study location | The Netherlands |
| Study setting | A rehabilitation centre (outpatient follow up) |
| Study dates | No additional information. |

| | |
|---|--|
| Sources of funding | This study was funded by the Dutch Scientific College of Physiotherapy of the Royal Dutch Society for Physical Therapy. |
| Inclusion criteria | A first-ever ischemic or hemorrhagic stroke in one of the hemispheres, as verified by CT and/or MRI scan; an upper limb deficit, however with minimal control of the paretic wrist and fingers (i.e., able to execute at least 10° of active wrist extension, at least 10° of thumb abduction/extension, and at least 10° extension in at least 2 additional digits); a score on the Action Research Arm Test (ARAT) of less than 53 points; between 18 and 80 years of age; written or oral informed consent; sufficient motivation to participate. |
| Exclusion criteria | Upper extremity orthopaedic limitations; not being able to communicate (i.e., < 4 points on the Utrecht Communication Observation, UCO [63]); a Mini Mental State Examination (MMSE) score of < 24 points |
| Recruitment / selection of participants | People admitted to the Rehabilitation Centre Amsterdam. |
| Intervention(s) | <p>Music intervention delivered by a healthcare professional N=19</p> <p>Modified bilateral arm training rhythmic auditory cueing group applied by physiotherapists and/or occupational therapists working at the rehabilitation centre. Treatment was given as 60 minute sessions, 3 days a week for 6 weeks. This included 3 minute movement periods interspersed with 5-minute rest periods (effectively 21 minutes of active movement). During rest periods and before the first exercise, people received visual and oral feedback on the previous exercise and instructions for the following exercise. The movements were paced by an auditory metronome. The tempo of the auditory cues depended on the severity of the upper limb deficit and was selected individually. The training was increased over the course of the training.</p> <p>Concomitant therapy: No additional information.</p> |
| Subgroup 1: Time after stroke at the start of the trial | Subacute (7 days - 6 months) |
| Subgroup 2: Severity (as stated by category or as measured by NIHSS scale) | Not stated/unclear |

| | |
|---|---|
| Subgroup 3: Group compared to individual sessions of music therapy | Individual sessions |
| Subgroup 4: Delivered in hospital compared to delivered in the community | Hospital sessions |
| Subgroup 5: Type of intervention | Rhythmic auditory cueing |
| Population subgroups | No additional information |
| Comparator | <p>No treatment N=19</p> <p>Dose matched control treatment using an exercise therapy based on existing guidelines for upper extremity treatment after stroke as presented by the Dutch Society of Occupational Therapy and Royal Dutch Society of Physical Therapy. Exercise was the same as in the music intervention group but without the rhythmic cues. This was provided for 60 minute sessions, 3 days a week for 6 weeks.</p> <p>Concomitant therapy: No additional information.</p> <p>A third group was reported (N=22). This group received constraint induced movement therapy which was deemed to be greater than no treatment and so was not included in the analysis.</p> |
| Number of participants | 60 |

| | |
|------------------------------|--|
| Duration of follow-up | 6 weeks (end of intervention) and 12 weeks |
| Indirectness | No additional information |
| Additional comments | No additional information |

Study arms

Music intervention delivered by a healthcare professional (N = 19)

Modified bilateral arm training rhythmic auditory cueing group applied by physiotherapists and/or occupational therapists working at the rehabilitation centre. Treatment was given as 60 minute sessions, 3 days a week for 6 weeks. This included 3 minute movement periods interspersed with 5-minute rest periods (effectively 21 minutes of active movement). During rest periods and before the first exercise, people received visual and oral feedback on the previous exercise and instructions for the following exercise. The movements were paced by an auditory metronome. The tempo of the auditory cues depended on the severity of the upper limb deficit and was selected individually. The training was increased over the course of the training. Concomitant therapy: No additional information.

No treatment (N = 19)

Dose matched control treatment using an exercise therapy based on existing guidelines for upper extremity treatment after stroke as presented by the Dutch Society of Occupational Therapy and Royal Dutch Society of Physical Therapy. Exercise was the same as in the music intervention group but without the rhythmic cues. This was provided for 60 minute sessions, 3 days a week for 6 weeks. Concomitant therapy: No additional information.

Characteristics***Arm-level characteristics***

| Characteristic | Music intervention delivered by a healthcare professional (N = 19) | No treatment (N = 19) |
|----------------------------------|---|------------------------------|
| % Female | n = 8 ; % = 42.1 | n = 3 ; % = 15.8 |
| Sample size | | |
| Mean age (SD) | 62.6 (9.8) | 56.9 (12.7) |
| Mean (SD) | | |
| Ethnicity | NR | NR |
| Nominal | | |
| Comorbidities | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Time after stroke (Weeks) | 7.8 (4.9) | 11.1 (6.8) |
| Mean (SD) | | |
| Severity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Type of stroke | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |

Outcomes

Study timepoints

- Baseline
- 12 week (<6 months)

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - continuous outcome

| Outcome | Music intervention delivered by a healthcare professional, Baseline, N = 19 | Music intervention delivered by a healthcare professional, 12 week, N = 17 | No treatment, Baseline, N = 19 | No treatment, 12 week, N = 15 |
|--|---|--|--------------------------------|-------------------------------|
| Stroke-specific Patient Reported Outcome Measures (Stroke Impact Scale) Scale range: 0-100. Change scores. | NR (NR) | NR (NR) | NR (NR) | NR (NR) |
| Mean (SD) | | | | |
| Strength | 49.7 (15.4) | -1.1 (13.1) | 52.5 (14.2) | 11.7 (13.1) |
| Mean (SD) | | | | |
| Memory | 85.9 (16.1) | 0.2 (9.6) | 83.8 (16.4) | -0.9 (8.2) |
| Mean (SD) | | | | |
| Emotion | 78.1 (16) | 5.8 (18.1) | 83 (14.3) | 3.9 (11.7) |
| Mean (SD) | | | | |
| Communication | 90.2 (13.6) | 2.3 (11.9) | 87.2 (14.2) | 3.1 (12.8) |
| Mean (SD) | | | | |

| Outcome | Music intervention delivered by a healthcare professional, Baseline, N = 19 | Music intervention delivered by a healthcare professional, 12 week, N = 17 | No treatment, Baseline, N = 19 | No treatment, 12 week, N = 15 |
|-----------------------------------|--|---|---------------------------------------|--------------------------------------|
| Activities of daily living | 56.4 (19.1) | 2.8 (13.5) | 61.4 (17.1) | 0 (12.7) |
| Mean (SD) | | | | |
| Mobility | 56 (28.9) | 1.8 (7.3) | 68.9 (22.1) | 3.1 (10.4) |
| Mean (SD) | | | | |
| Hand function | 27.4 (27.2) | 11.5 (22.8) | 33.7 (26.6) | 9.7 (22.6) |
| Mean (SD) | | | | |
| Social participation | 37.8 (22.6) | 6.8 (19.3) | 44.6 (17.5) | 3.1 (20.5) |
| Mean (SD) | | | | |

Stroke-specific Patient Reported Outcome Measures (Stroke Impact Scale) - Polarity - Higher values are better

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - dichotomous outcome

| Outcome | Music intervention delivered by a healthcare professional, Baseline, N = 19 | Music intervention delivered by a healthcare professional, 12 week, N = 19 | No treatment, Baseline, N = 19 | No treatment, 12 week, N = 19 |
|---|--|---|---------------------------------------|--------------------------------------|
| Withdrawal due to adverse events | n = NA ; % = NA | n = 0 ; % = 0 | n = NA ; % = NA | n = 0 ; % = 0 |
| No of events | | | | |

Withdrawal due to adverse events - Polarity - Lower values are better

Critical appraisal - Cochrane Risk of Bias tool (RoB 2.0) Normal RCT

Music intervention delivered by healthcare professionals compared to no treatment at <6 months-continuous outcome-Stroke-specific Patient Reported Outcome Measures (Stroke Impact Scale)-Strength-Mean SD-Music intervention delivered by a healthcare professional-No treatment-t12

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months-continuous outcome-Stroke-specific Patient Reported Outcome Measures (Stroke Impact Scale)-Memory-Mean SD-Music intervention delivered by a healthcare professional-No treatment-t12

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months-continuous outcome-Stroke-specific Patient Reported Outcome Measures (Stroke Impact Scale)-Emotion-Mean SD-Music intervention delivered by a healthcare professional-No treatment-t12

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months-continuous outcome-Stroke-specific Patient Reported Outcome Measures (Stroke Impact Scale)-Communication-Mean SD-Music intervention delivered by a healthcare professional-No treatment-t12

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months-continuous outcome-Stroke-specific Patient Reported Outcome Measures (Stroke Impact Scale)-Activities of daily living-Mean SD-Music intervention delivered by a healthcare professional-No treatment-t12

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months-continuous outcome-Stroke-specific Patient Reported Outcome Measures (Stroke Impact Scale)-Mobility-Mean SD-Music intervention delivered by a healthcare professional-No treatment-t12

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - continuous outcome - Stroke-specific Patient Reported Outcome Measures (Stroke Impact Scale) - Hand function - Mean SD - Music intervention delivered by a healthcare professional - No treatment - t12

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - continuous outcome - Stroke-specific Patient Reported Outcome Measures (Stroke Impact Scale) - Social participation - Mean SD - Music intervention delivered by a healthcare professional - No treatment - t12

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music intervention delivered by healthcare professionals compared to no treatment at <6 months - dichotomous outcome - Withdrawal due to adverse events - No Of Events - Music intervention delivered by a healthcare professional - No treatment - t12

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | Some concerns |
| Overall bias and Directness | Overall Directness | Directly applicable |

Whitall, 2011**Bibliographic Reference**

Whitall, J.; Waller, S. M.; Sorkin, J. D.; Forrester, L. W.; Macko, R. F.; Hanley, D. F.; Goldberg, A. P.; Luft, A.; Bilateral and unilateral arm training improve motor function through differing neuroplastic mechanisms: a single-blinded randomized controlled trial; *Neurorehabilitation & Neural Repair*; 2011; vol. 25 (no. 2); 118-29

Study details

| | |
|---|---|
| Secondary publication of another included study- see primary study for details | No additional information. |
| Other publications associated with this study included in review | No additional information. |
| Trial name / registration number | No additional information. |
| Study type | Randomised controlled trial (RCT) |
| Study location | United States of America |
| Study setting | Outpatient follow up |
| Study dates | No additional information. |
| Sources of funding | P60AG12583; PI AG, NIDDR H H133G010111, the Baltimore Veterans Administration Geriatrics Research, Education and Clinical Center (GRECC). Andreas Luft was supported by DFG SFB 550, C 12. |
| Inclusion criteria | People with unilateral stroke >6 months earlier; could follow simple instructions; had volitional control of the nonparetic arm; the ability to flex the paretic arm shoulder 3 inches from a neutral position. |

| | |
|---|--|
| Exclusion criteria | Symptomatic heart disease; uncontrolled hypertension (>180/100 mm Hg); significant orthopaedic or chronic pain conditions; untreated poststroke depression (Center for Epidemiological Studies Depression Scale; cutoff >16); active cancer; severe obstructive pulmonary disease; cognitive loss measured using the Folstein Mini Mental State Exam. |
| Recruitment / selection of participants | People were recruited from the Baltimore Veterans Affairs Medical Centre and involved referrals from the University of Maryland Medical System Hospital and regionwide advertisements. |
| Intervention(s) | <p>Music intervention delivered by healthcare professionals N=55</p> <p>Bilateral arm training with rhythmic auditory cueing for 3 times a week for 6 weeks (for a total of 18 sessions). Completed 5 minutes of training with arms moving simultaneously with auditory cuing, followed by 10 minutes at rest. Training continued for 5 minutes with arms moving alternately (antiphase) again with auditory cuing before 10 minutes of rest. This was then repeated once each achieving 20 minutes of active therapy in 1 hour sessions.</p> <p>Concomitant therapy: No additional information.</p> |
| Subgroup 1: Time after stroke at the start of the trial | Chronic (>6 months) |
| Subgroup 2: Severity (as stated by category or as measured by NIHSS scale) | Not stated/unclear |
| Subgroup 3: Group compared to individual sessions of music therapy | Individual sessions |
| Subgroup 4: Delivered in hospital compared | Hospital sessions |

| | |
|---|---|
| to delivered in the community | |
| Subgroup 5: Type of intervention | Rhythmic auditory cueing |
| Population subgroups | No additional information. |
| Comparator | No treatment N=56 Dose matched therapeutic exercises for the same time period. Based on neurodevelopmental principles, including thoracic spine mobilisation with weight shifting, scapular mobilization, weight bearing with the paretic arm (elbow fixed) and opening the hand with finger extension. This was completed for 4 cycles of active continuous 5 minute training followed by 10 minutes of rest. Concomitant therapy: No additional information |
| Number of participants | 111 |
| Duration of follow-up | 6 weeks (end of intervention) and 4 months. The 4 months data will be used where possible. |
| Indirectness | No additional information. |
| Additional comments | Intention-to-treat analysis including all participants at each time regardless of study completion |

Study arms

Music intervention delivered by healthcare professionals (N = 55)

Bilateral arm training with rhythmic auditory cueing for 3 times a week for 6 weeks (for a total of 18 sessions). Completed 5 minutes of training with arms moving simultaneously with auditory cueing, followed by 10 minutes at rest. Training continued for 5 minutes with

arms moving alternately (antiphase) again with auditory cuing before 10 minutes of rest. This was then repeated once each achieving 20 minutes of active therapy in 1 hour sessions. Concomitant therapy: No additional information.

No treatment (N = 56)

Dose matched therapeutic exercises for the same time period. Based on neurodevelopmental principles, including thoracic spine mobilisation with weight shifting, scapular mobilization, weight bearing with the paretic arm (elbow fixed) and opening the hand with finger extension. This was completed for 4 cycles of active continuous 5 minute training followed by 10 minutes of rest. Concomitant therapy: No additional information

Characteristics

Arm-level characteristics

| Characteristic | Music intervention delivered by healthcare professionals (N = 55) | No treatment (N = 56) |
|---------------------------|---|-----------------------|
| % Female | n = 16 ; % = 29 | n = 26 ; % = 46 |
| Sample size | | |
| Mean age (SD) (years) | 59.8 (9.9) | 57.7 (12.5) |
| Mean (SD) | | |
| Ethnicity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Comorbidities | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Time after stroke (years) | 4.5 (4.1) | 4.1 (5.2) |

| Characteristic | Music intervention delivered by healthcare professionals (N = 55) | No treatment (N = 56) |
|-----------------------|--|------------------------------|
| Mean (SD) | | |
| Severity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Type of stroke | n = NA ; % = NA | n = NA ; % = NA |
| Sample size | | |
| Brainstem | n = 3 ; % = 5.4 | n = 3 ; % = 5.4 |
| Sample size | | |
| Cerebellar | n = 0 ; % = 0 | n = 2 ; % = 3.6 |
| Sample size | | |
| Cortex | n = 19 ; % = 34.6 | n = 20 ; % = 35.7 |
| Sample size | | |
| Multiple | n = 3 ; % = 5.5 | n = 0 ; % = 0 |
| Sample size | | |
| Subcortical | n = 7 ; % = 16.7 | n = 12 ; % = 24 |
| Sample size | | |

Outcomes

Study timepoints

- Baseline

- 6 week (End of intervention. <6 months (4 month follow up data not reported).)
- 4 month (For withdrawal due to adverse events only. <6 months.)

Music interventions delivered by healthcare professionals compared to no treatment at <6 months - continuous outcome

| Outcome | Music intervention delivered by healthcare professionals, Baseline, N = 55 | Music intervention delivered by healthcare professionals, 6 week, N = 37 | Music intervention delivered by healthcare professionals, 4 month, N = 37 | No treatment, Baseline, N = 56 | No treatment, 6 week, N = 42 | No treatment, 4 month, N = 42 |
|--|---|---|--|---------------------------------------|-------------------------------------|--------------------------------------|
| Stroke-specific Patient Reported Outcome Measures (Stroke Impact Scale) Scale range: Unclear. Change scores. Mean (SD) | 549 (17) | 12 (6.1) | NR (NR) | 578 (15) | 26 (8.9) | NR (NR) |

Stroke-specific Patient Reported Outcome Measures (Stroke Impact Scale) - Polarity - Higher values are better

Music interventions delivered by healthcare professionals compared to no treatment at <6 months - dichotomous outcome

| Outcome | Music intervention delivered by healthcare professionals, Baseline, N = 55 | Music intervention delivered by healthcare professionals, 6 week, N = 55 | Music intervention delivered by healthcare professionals, 4 month, N = 55 | No treatment, Baseline, N = 56 | No treatment, 6 week, N = 56 | No treatment, 4 month, N = 56 |
|---|---|---|--|---------------------------------------|-------------------------------------|--------------------------------------|
| Withdrawal due to adverse events Intervention: Medical = 10, death | n = NA ; % = NA | n = NA ; % = NA | n = 11 | n = NA ; % = NA | n = NA ; % = NA | n = 11 ; % = 19.6 |

| Outcome | Music intervention delivered by healthcare professionals, Baseline, N = 55 | Music intervention delivered by healthcare professionals, 6 week, N = 55 | Music intervention delivered by healthcare professionals, 4 month, N = 55 | No treatment, Baseline, N = 56 | No treatment, 6 week, N = 56 | No treatment, 4 month, N = 56 |
|-----------------------------|--|--|---|--------------------------------|------------------------------|-------------------------------|
| = 1. Control: Medical = 11. | | | | | | |
| No of events | | | | | | |

Withdrawal due to adverse events - Polarity - Lower values are better

Critical appraisal - Cochrane Risk of Bias tool (RoB 2.0) Normal RCT

Music interventions delivered by healthcare professionals compared to no treatment at <6 months - continuous outcome - Stroke-specific Patient Reported Outcome Measures (Stroke Impact Scale) - Mean SD - Music intervention delivered by healthcare professionals - No treatment - t6

| Section | Question | Answer |
|-----------------------------|------------------------|---------------------|
| Overall bias and Directness | Risk of bias judgement | High |
| Overall bias and Directness | Overall Directness | Directly applicable |

Music interventions delivered by healthcare professionals compared to no treatment at <6 months - dichotomous outcome - Withdrawal due to adverse events - No Of Events - Music intervention delivered by healthcare professionals - No treatment - t4

| Section | Question | Answer |
|-----------------------------|------------------------|--------|
| Overall bias and Directness | Risk of bias judgement | High |

| Section | Question | Answer |
|-----------------------------|--------------------|---------------------|
| Overall bias and Directness | Overall Directness | Directly applicable |

Zhang, 2021

| | |
|--------------------------------|---|
| Bibliographic Reference | Zhang, Xiao-Ying; Yu, Wei-Yong; Teng, Wen-Jia; Lu, Meng-Yang; Wu, Xiao-Li; Yang, Yu-Qi; Chen, Chen; Liu, Li-Xu; Liu, Song-Huai; Li, Jian-Jun; Effectiveness of Melodic Intonation Therapy in Chinese Mandarin on Non-fluent Aphasia in Patients After Stroke: A Randomized Control Trial.; <i>Frontiers in neuroscience</i> ; 2021; vol. 15; 648724 |
|--------------------------------|---|

Study details

| | |
|---|---|
| Secondary publication of another included study- see primary study for details | No additional information. |
| Other publications associated with this study included in review | No additional information. |
| Trial name / registration number | Chinese Clinical Trial Registry (Registration No. ChiCTR2000037871) |
| Study type | Randomised controlled trial (RCT) |
| Study location | Beijing, China. |
| Study setting | China Rehabilitation Research Center (CRRRC) |
| Study dates | April 2020 to October 2020 |

| | |
|--|--|
| Sources of funding | This research was supported by the China Rehabilitation Research Center (CRRC). This research received a grant from the Scientific Research Project 2020CZ-10 of the Chinese Institute of Rehabilitation Science. It is a national non-profit foundation program and was approved by the Ministry of Finance of China. |
| Inclusion criteria | Diagnosed with fMRI or CT imaging, showing left ischemic stroke or hemorrhagic stroke; the ninth language score on the National Institutes of Health Stroke Scale (NIHSS) is 1— mild to moderate aphasia and 2—severe aphasia. Meets the diagnostic criteria for non-fluent aphasia: less active speech expression, lack of fluency in speaking, acceptable hearing ability, can give a sign of yes/no questions, willing to express, good cooperation, and emotional stability; Aphasia for more than 15 days after stroke, hospitalized patients; aged 18–70; tolerance to lying therapy for more than half an hour without postural hypotension. The medication and other brain metabolism enhancers are the same; physical therapy, occupational therapy, and routine care are the same. None of the participants had professional musical experience. |
| Exclusion criteria | Severe auditory dysfunction; having epilepsy, malignant arrhythmia, or other serious physical diseases; and patients with mental symptoms and obvious emotional agitation. |
| Recruitment / selection of participants | Recruited from China Rehabilitation Research Centre |
| Intervention(s) | Neurologic music therapy delivered by trained music therapists (melodic intonation therapy) N=20 All patients trained for 30 minutes per session, five sessions a week, for 8 weeks. The training process is carried out by music therapy professionals who have been trained in neurological musical therapy and have obtained a registered music therapist license to ensure the music professionalism of the intervention. The intervention steps of MIT strictly follow the operational steps of Chinese Mandarin MIT. According to the different three levels of speech rehabilitation, the music therapist trained the aphasia patients to intone and chant the targeted speech items and then fade slowly with tapping to let the patients speak out the targeted sentences in the first level. The music therapist leads the patients to sing and speak out in the same way in the second and third levels; the only difference is the length of the melodic target language (the second level is 5– 9-word sentences, and the third level is 10-word sentences and above). All the melodic phrases are noted according to the natural phonic pitches of targeted Mandarin sentences. The music therapist uses a keyboard or guitar to accompany while they are singing the melody with the patients. The effective behavioural performance of the intervention is that, when the therapist asks the target question, the patient can speak the target language at a natural speed without the melody and rhythm, and the behaviour performance can last for more than 3 weeks without regression. Concomitant therapy: All patients underwent routine treatment during the study period, including taking medication and other care and support. |

| | |
|---|---|
| Subgroup 1: Time after stroke at the start of the trial | Not stated/unclear |
| Subgroup 2: Severity (as stated by category or as measured by NIHSS scale) | Mixed |
| Subgroup 3: Group compared to individual sessions of music therapy | Individual sessions |
| Subgroup 4: Delivered in hospital compared to delivered in the community | Hospital sessions |
| Subgroup 5: Type of intervention | Singing and music-based voice interventions |
| Population subgroups | No additional information. |
| Comparator | No treatment (Control group - speech therapy) N=20 Treated by speech therapy that was conducted by speech therapist. The sessions were 30 min per day, five times a week, for a total of eight consecutive weeks. Concomitant therapy: All patients underwent routine treatment during the study period, including taking medication and other care and support. |
| Number of participants | 40 |
| Duration of follow-up | 8 weeks. |
| Indirectness | No indirectness. |

| | |
|----------------------------|----------------------------|
| Additional comments | No additional information. |
|----------------------------|----------------------------|

Study arms

Neurologic music therapy delivered by trained music therapists (melodic intonation therapy) (N = 20)

| | |
|------------------------|---|
| Intervention(s) | <p>Melodic intonation therapy (MIT) delivered by music therapy professionals N=20</p> <p>All patients trained for 30 minutes per session, five sessions a week, for 8 weeks. The training process is carried out by music therapy professionals who have been trained in neurological musical therapy and have obtained a registered music therapist license to ensure the music professionalism of the intervention. The intervention steps of MIT strictly follow the operational steps of Chinese Mandarin MIT. According to the different three levels of speech rehabilitation, the music therapist trained the aphasia patients to intone and chant the targeted speech items and then fade slowly with tapping to let the patients speak out the targeted sentences in the first level. The music therapist leads the patients to sing and speak out in the same way in the second and third levels; the only difference is the length of the melodic target language (the second level is 5– 9-word sentences, and the third level is 10-word sentences and above). All the melodic phrases are noted according to the natural phonic pitches of targeted Mandarin sentences. The music therapist uses a keyboard or guitar to accompany while they are singing the melody with the patients. The effective behavioural performance of the intervention is that, when the therapist asks the target question, the patient can speak the target language at a natural speed without the melody and rhythm, and the behaviour performance can last for more than 3 weeks without regression. Concomitant therapy: All patients underwent routine treatment during the study period, including taking medication and other care and support.</p> |
| Comparator | Control group - speech therapy from therapist. |

All patients trained for 30 minutes per session, five sessions a week, for 8 weeks. The training process is carried out by music therapy professionals who have been trained in neurological musical therapy and have obtained a registered music therapist license to ensure the music professionalism of the intervention. The intervention steps of MIT strictly follow the operational steps of Chinese Mandarin MIT. According to the different three levels of speech rehabilitation, the music therapist trained the aphasia patients to intone and chant the targeted speech items and then fade slowly with tapping to let the patients speak out the targeted sentences in the first level. The music therapist leads the patients to sing and speak out in the same way in the second and third levels; the only difference is the length of the melodic target language (the second level is 5– 9-word sentences, and the third level is 10-word sentences and above). All the melodic phrases are noted according to the natural phonic pitches of targeted Mandarin sentences. The music therapist uses a keyboard or guitar to accompany while they are singing the melody with the patients. The effective behavioural performance of the

intervention is that, when the therapist asks the target question, the patient can speak the target language at a natural speed without the melody and rhythm, and the behaviour performance can last for more than 3 weeks without regression. Concomitant therapy: All patients underwent routine treatment during the study period, including taking medication and other care and support.

No treatment (Control group - speech therapy) (N = 20)

| | |
|------------------------|---|
| Intervention(s) | <p>Melodic intonation therapy (MIT) delivered by music therapy professionals N=20</p> <p>All patients trained for 30 minutes per session, five sessions a week, for 8 weeks. The training process is carried out by music therapy professionals who have been trained in neurological musical therapy and have obtained a registered music therapist license to ensure the music professionalism of the intervention. The intervention steps of MIT strictly follow the operational steps of Chinese Mandarin MIT. According to the different three levels of speech rehabilitation, the music therapist trained the aphasia patients to intone and chant the targeted speech items and then fade slowly with tapping to let the patients speak out the targeted sentences in the first level. The music therapist leads the patients to sing and speak out in the same way in the second and third levels; the only difference is the length of the melodic target language (the second level is 5– 9-word sentences, and the third level is 10-word sentences and above). All the melodic phrases are noted according to the natural phonic pitches of targeted Mandarin sentences. The music therapist uses a keyboard or guitar to accompany while they are singing the melody with the patients. The effective behavioural performance of the intervention is that, when the therapist asks the target question, the patient can speak the target language at a natural speed without the melody and rhythm, and the behaviour performance can last for more than 3 weeks without regression. Concomitant therapy: All patients underwent routine treatment during the study period, including taking medication and other care and support.</p> |
| Comparator | Control group - speech therapy from therapist. |

Treated by speech therapy that was conducted by speech therapist. The sessions were 30 min per day, five times a week, for a total of eight consecutive weeks. Concomitant therapy: All patients underwent routine treatment during the study period, including taking medication and other care and support.

Characteristics**Arm-level characteristics**

| Characteristic | Neurologic music therapy delivered by trained music therapists (melodic intonation therapy) (N = 20) | No treatment (Control group - speech therapy) (N = 20) |
|--------------------------------|---|---|
| % Female (Sample size) | n = 4 ; % = 20 | n = 5 ; % = 25 |
| Sample size | | |
| Mean age (SD) | 52.9 (9.08) | 54.05 (10.81) |
| Mean (SD) | | |
| Ethnicity | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Comorbidities | n = NR ; % = NR | n = NR ; % = NR |
| Sample size | | |
| Time after stroke | 2.57 (1.74) | 1.96 (1.38) |
| Standardised Mean (SD) | | |
| Severity | NR | NR |
| Nominal | | |
| Left cortical ischaemic | 10 | 14 |
| Nominal | | |

| Characteristic | Neurologic music therapy delivered by trained music therapists (melodic intonation therapy) (N = 20) | No treatment (Control group - speech therapy) (N = 20) |
|----------------------------------|--|--|
| Left cortical hemorrhagic | 10 | 6 |
| Nominal | | |

Outcomes

Study timepoints

- Baseline
- 8 week (< 6 months)

Melodic intonation therapy delivered by music therapy professionals compared to speech therapy at < 6 months - continuous

| Outcome | Baseline, Neurologic music therapy delivered by trained music therapists (melodic intonation therapy), N = 20 | Baseline, No treatment (Control group - speech therapy), N = 20 | 8 week, Neurologic music therapy delivered by trained music therapists (melodic intonation therapy), N = 20 | 8 week, No treatment (Control group - speech therapy), N = 20 |
|--|---|---|---|---|
| Psychological distress - anxiety (Hamilton Anxiety Rating Scale) Scale range: 0-56. Final values. | 12.15 (3.16) | 12.75 (2.47) | 8.6 (2.68) | 9.65 (1.8) |
| Mean (SD) | | | | |
| Psychological distress - depression (Hamilton Depression Scale) (17 item score battery using a five-level scoring method of 0-4 points. Score ranges from | 16.15 (2.52) | 16.45 (2.25) | 8.95 (1.97) | 10.9 (1.64) |

| Outcome | Baseline, Neurologic music therapy delivered by trained music therapists (melodic intonation therapy), N = 20 | Baseline, No treatment (Control group - speech therapy), N = 20 | 8 week, Neurologic music therapy delivered by trained music therapists (melodic intonation therapy), N = 20 | 8 week, No treatment (Control group - speech therapy), N = 20 |
|---|---|---|---|---|
| asymptomatic to extremely severe) Scale range: 0-56. Final values. | | | | |
| Mean (SD) | | | | |
| Psychological distress - anxiety (Hamilton Anxiety Rating Scale) - Polarity - Lower values are better | | | | |
| Psychological distress - depression (Hamilton Depression Scale) - Polarity - Lower values are better | | | | |

Critical appraisal - Cochrane Risk of Bias tool (RoB 2.0) Normal RCT

Melodic intonation therapy delivered by music therapy professionals compared to speech therapy at <6 months - continuous - Psychological distress - Mean SD - Melodic intonation therapy (MIT) delivered by music therapy professionals - Control group - speech therapy - t8

| Section | Question | Answer |
|-----------------------------|------------------------|---|
| Overall bias and Directness | Risk of bias judgement | Some concerns (Allocation concealment not mentioned) |
| Overall bias and Directness | Overall Directness | Directly applicable |

Melodic intonation therapy delivered by music therapy professionals compared to speech therapy at <6 months-continuous-Hamilton Depression Scale-Mean SD-Melodic intonation therapy (MIT) delivered by music therapy professionals-Control group - speech therapy-t8

| Section | Question | Answer |
|-----------------------------|------------------------|---|
| Overall bias and Directness | Risk of bias judgement | Some concerns (Allocation concealment not mentioned) |
| Overall bias and Directness | Overall Directness | Directly applicable |

Zhao, 2022

Bibliographic Reference

Zhao, Li; Lyu, Xiaokang; Jiang, He; Gao, Xinhai; Musicokinetic and exercise therapies decrease the depression level of elderly patients undergoing post-stroke rehabilitation: The moderating effect of health regulatory focus.; *Frontiers in psychology*; 2022; vol. 13; 889510

Study details

| | |
|--|----------------------------|
| Secondary publication of another included study- see primary study for details | No additional information. |
| Other publications associated with this study included in review | No additional information. |
| Trial name / registration number | No additional information. |

| | |
|--|--|
| Study type | Randomised controlled trial (RCT) |
| Study location | Shanghai, China |
| Study setting | Hospital in Shanghai, China. |
| Study dates | December 2019 to March 2020. |
| Sources of funding | This work was supported partially by the Tianjin Social Science Foundation of China (TJJX21-011) and the Developmental Program of Liberal and Social Sciences of Nankai University (ZB22BZ0109). |
| Inclusion criteria | Patients who (1) were at least 60 years old; (2) met the diagnostic criteria in the 2018 edition of the Chinese Guidelines for the Management of Acute Ischemic Stroke or the 2019 edition of the Chinese Guidelines for the Management of Cerebral Hemorrhage; (3) had approximately two weeks after the stroke; (4) with neurological sequelae requiring rehabilitative exercise therapy; (5) received at least eight points on the Chinese version of Hamilton Depression Rating Scale with 24 items (HDRS-24); and (6) received at least 27 points on the Mini-mental State Examination (MMSE). |
| Exclusion criteria | Patients (1) with other serious organ diseases, such as a malignant tumor, myocardial infarction, and comatose; (2) who were unable to take exercise therapy after the stroke attack; (3) or their relatives who refuse to take the exercise or musicokinetic therapy; (4) who received antidepressant medication in the past month; and (5) who withdrew from treatments due to transfer and personal reasons. |
| Recruitment / selection of participants | A convenience sampling method was used, and data were collected from elderly patients undergoing post-stroke rehabilitation treatments in one hospital located in Shanghai, China. |
| Intervention(s) | <p>Musicokinetic therapy group: Exercise treatment and music component of background music in the rehabilitation room. The Edifier S2000MKIII speaker in the rehabilitation room was used as the music-playing equipment. Participants listened to the same type of ambient pure music from Bandari. Bandari produces music with slower and softer rhythm and many people in China choose to listen to Bandar music to relax. The volume of the music was set to 50 decibels. The musicokinetic therapy lasted for 30 minutes each time and was performed twice a day. The whole musicokinetic therapy lasted 8 weeks.</p> <p>Exercise treatment: participants focused on active exercises, including isometric muscle, joint function, speech and swallowing function, balance function, and gait training. The length of each exercise was 30 minutes and performed twice a day. The rehabilitation room in this study was about 80 m² and exercise therapy lasted 8 weeks.</p> |

| | |
|---|---|
| | Concomitant treatment: All participants ingested one tablet of Sertraline Hydrochloride (50mg) which is a type of antidepressant drug, every morning during the experimental period. |
| Subgroup 1: Time after stroke at the start of the trial | Subacute (7 days - 6 months) |
| Subgroup 2: Severity (as stated by category or as measured by NIHSS scale) | Not stated/unclear |
| Subgroup 3: Group compared to individual sessions of music therapy | Individual sessions |
| Subgroup 4: Delivered in hospital compared to delivered in the community | Hospital sessions |
| Subgroup 5: Type of intervention | Receptive interventions in which participants listen to music |
| Population subgroups | No additional information. |
| Comparator | <p>Exercise therapy group: participants focused on active exercises, including isometric muscle, joint function, speech and swallowing function, balance function, and gait training. The length of each exercise was 30 minutes and performed twice a day. The rehabilitation room in this study was about 80 m² and exercise therapy lasted 8 weeks.</p> <p>Concomitant treatment: All participants ingested one tablet of Sertraline Hydrochloride (50mg) which is a type of antidepressant drug, every morning during the experimental period.</p> |

| | |
|-------------------------------|----------------------------|
| Number of participants | 65 |
| Duration of follow-up | 8 weeks. |
| Indirectness | No indirectness. |
| Additional comments | No additional information. |

Study arms

Music intervention delivered by healthcare professionals (Musicokinetic therapy group) (N = 32)

Exercise treatment and music component of background music in the rehabilitation room. The Edifier S2000MKIII speaker in the rehabilitation room was used as the music-playing equipment. Participants listened to the same type of ambient pure music from Bandari. Bandari produces music with slower and softer rhythm and many people in China choose to listen to Bandar music to relax. The volume of the music was set to 50 decibels. The musicokinetic therapy lasted for 30 minutes each time and was performed twice a day. The whole musicokinetic therapy lasted 8 weeks.

No treatment (Exercise therapy group) (N = 33)

Participants focused on active exercises, including isometric muscle, joint function, speech and swallowing function, balance function, and gait training. The length of each exercise was 30 minutes and performed twice a day. The rehabilitation room in this study was about 80 m² and exercise therapy lasted 8 weeks.

Characteristics***Study-level characteristics***

| Characteristic | Study (N = 65) |
|--------------------------|-----------------------|
| % Female | n = 37 ; % = 56.9 |
| No of events | |
| Mean age (SD) | 81.14 (8.33) |
| Mean (SD) | |
| Ethnicity | NR |
| Nominal | |
| Comorbidities | NR |
| Nominal | |
| Time after stroke | NR |
| Nominal | |
| Severity | NR |
| Nominal | |
| Type of stroke | NR |
| Nominal | |

Arm-level characteristics

| Characteristic | Music intervention delivered by healthcare professionals (Musicokinetic therapy group) (N = 32) | No treatment (Exercise therapy group) (N = 33) |
|--------------------------|--|---|
| % Female | n = NR ; % = NR | n = NR ; % = NR |
| No of events | | |
| Mean age (SD) | NR (NR) | NR (NR) |
| Mean (SD) | | |
| Ethnicity | NR | NR |
| Nominal | | |
| Comorbidities | NR | NR |
| Nominal | | |
| Time after stroke | NR | NR |
| Nominal | | |
| Severity | NR | NR |
| Nominal | | |
| Type of stroke | NR | NR |
| Nominal | | |

Outcomes

Study timepoints

- Baseline
- 8 week (<6 months)

Hamilton Depression Rating Scale-24

| Outcome | Baseline, Music intervention delivered by healthcare professionals (Musicokinetic therapy group), N = 32 | Baseline, No treatment (Exercise therapy group), N = 33 | 8 week, Music intervention delivered by healthcare professionals (Musicokinetic therapy group), N = 32 | 8 week, No treatment (Exercise therapy group), N = 33 |
|--|--|---|--|---|
| <p>Psychological distress - depression (Hamilton Depression Rating Scale-24) (In the Chinese version of HDRS-24, 14 items were scored on a 5-point Likert scale from 0 to 4, and 10 items were scored on a 3-point scale from 0 to 2, representing from absent to severe, respectively. If the final score was less than 8, the patient could be considered to be normal/non-depressed, and if the final score was 8 or higher, the patient could be considered to have varying degrees of depressive symptoms) Scale range: 0-42. Final values.</p> <p>Mean (SD)</p> | 23.97 (11.33) | 32.3 (17.9) | 16.38 (9.92) | 29.7 (19.43) |

Psychological distress - depression (Hamilton Depression Rating Scale-24) - Polarity - Lower values are better

Critical appraisal - Cochrane Risk of Bias tool (RoB 2.0) Normal RCT**HamiltonDepressionRatingScale-24-HamiltonDepressionRatingScale-24-MeanSD-Musicokinetic therapy group-Exercise therapy group-t8**

| Section | Question | Answer |
|-----------------------------|------------------------|--|
| Overall bias and Directness | Risk of bias judgement | High <i>(No mention of allocation concealment, no mention of blinding and subjective outcome and baseline measures of outcome were different)</i> |
| Overall bias and Directness | Overall Directness | Directly applicable |

Appendix E – Forest plots

E.1 Neurologic music therapy delivered by trained music therapists compared to no treatment

Figure 2: Person/participant generic health-related quality of life (SF-36 physical functioning, 0-100, higher values are better, final values) at <6 months

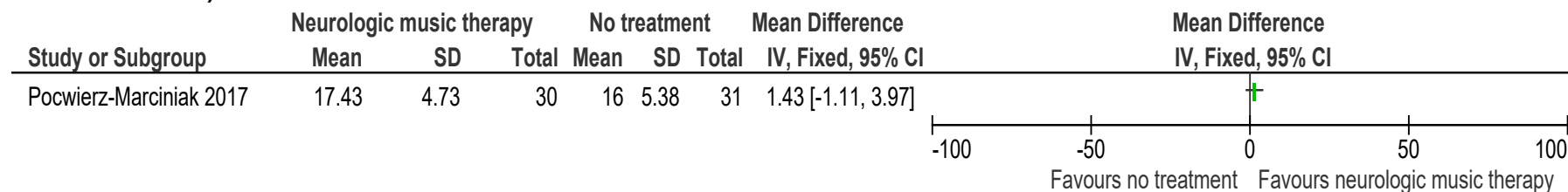


Figure 3: Person/participant generic health-related quality of life (SF-36 bodily pain, 0-100, higher values are better, final values) at <6 months

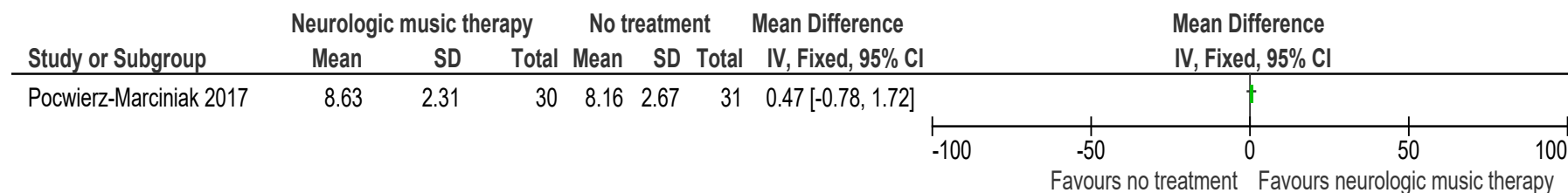


Figure 4: Person/participant generic health-related quality of life (SF-36 role physical, 0-100, higher values are better, final values) at <6 months

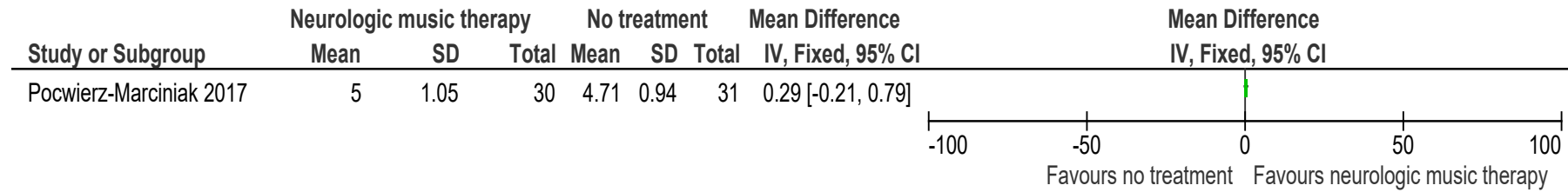


Figure 5: Person/participant generic health-related quality of life (SF-36 vitality, 0-100, higher values are better, final values) at <6 months

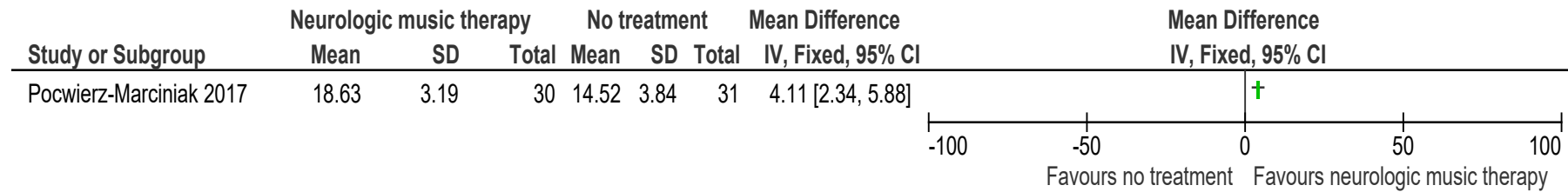


Figure 6: Person/participant generic health-related quality of life (SF-36 general health, 0-100, higher values are better, final values) at <6 months

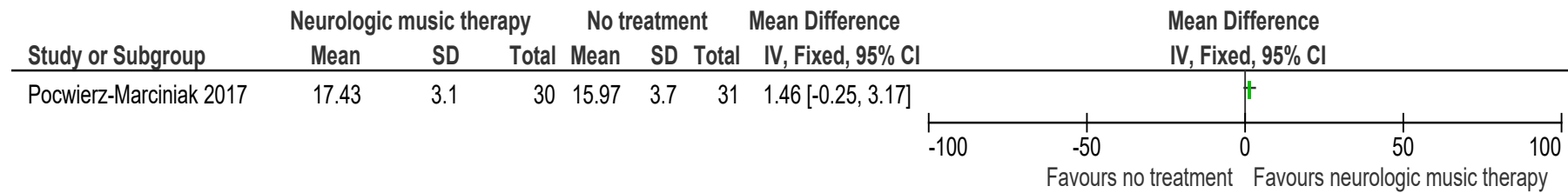


Figure 7: Person/participant generic health-related quality of life (SF-36 role emotional, 0-100, higher values are better, final values) at <6 months

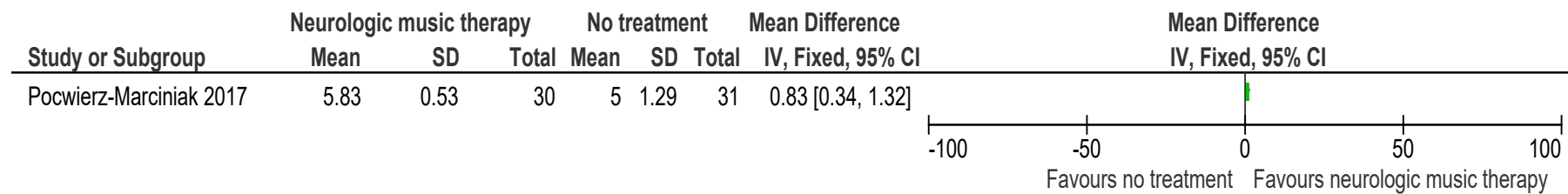


Figure 8: Person/participant generic health-related quality of life (SF-36 mental health, 0-100, higher values are better, final values) at <6 months

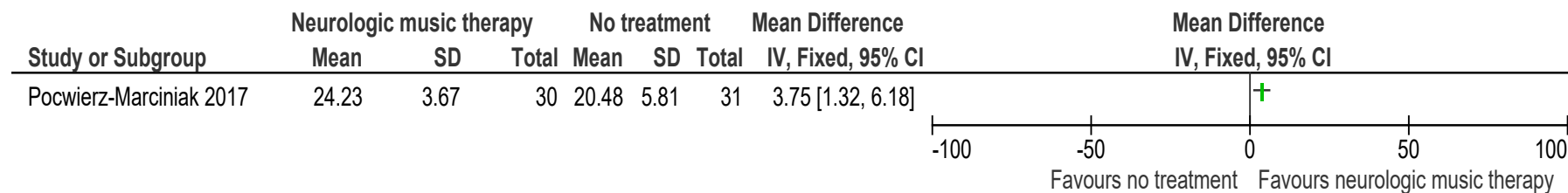


Figure 9: Person/participant generic health-related quality of life (SF-36 social functioning, 0-100, higher values are better, final values) at <6 months

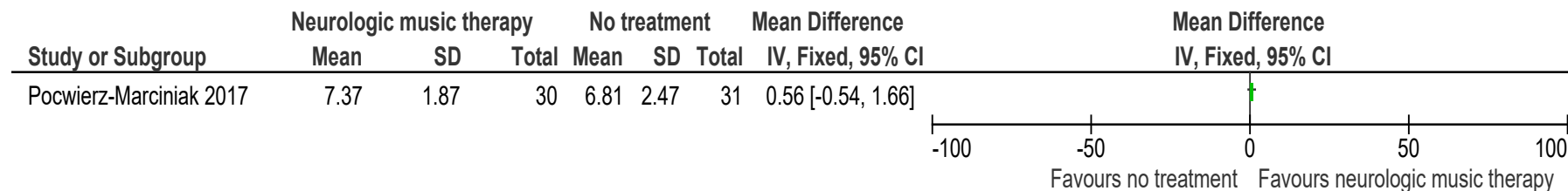


Figure 10: Psychological distress - depression (Hamilton Depression Scale, 0-56, lower values are better, final value) at <6 months

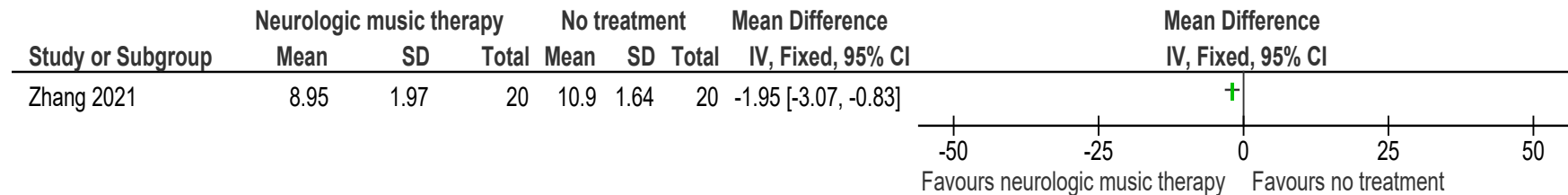


Figure 11: Psychological distress - anxiety (Hamilton Anxiety Rating Scale, 0-56, lower values are better, final value) at <6 months

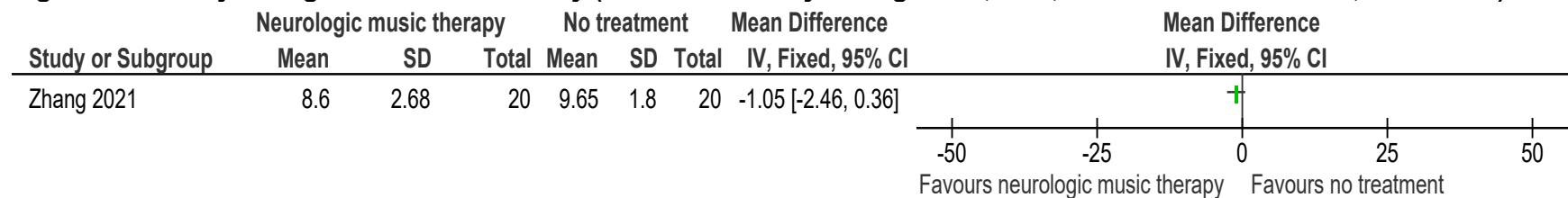
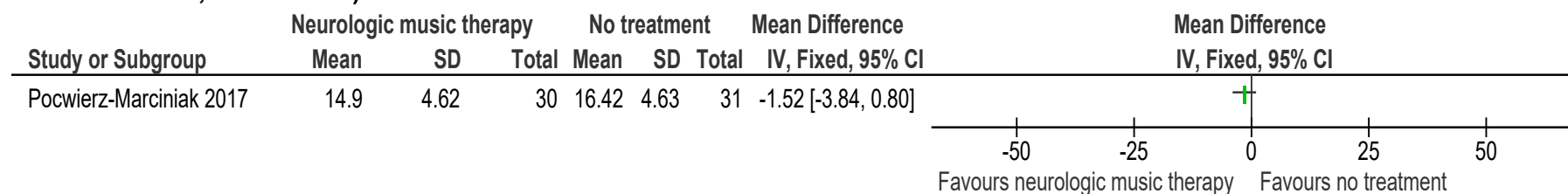


Figure 12: Stroke-specific Patient-Reported Outcome Measures (Stroke adjusted Sickness Impact Profile 30, 0-68, lower values are better, final values) at <6 months



E.2 Music therapy delivered by trained music therapists compared to no treatment

Figure 13: Patient/participant generic health-related quality of life (McGill Quality of Life, 0-10, higher values are better, final value) at <6 months

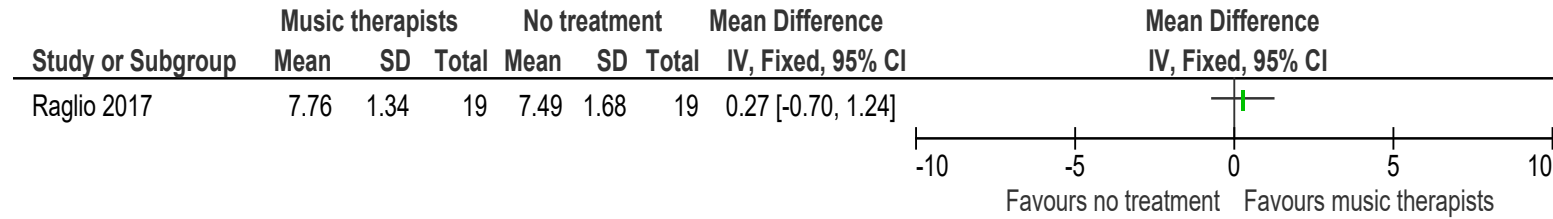


Figure 14: Activities of daily living (Korean-Modified Barthel Index, 0-100, higher values are better, change score) at <6 months

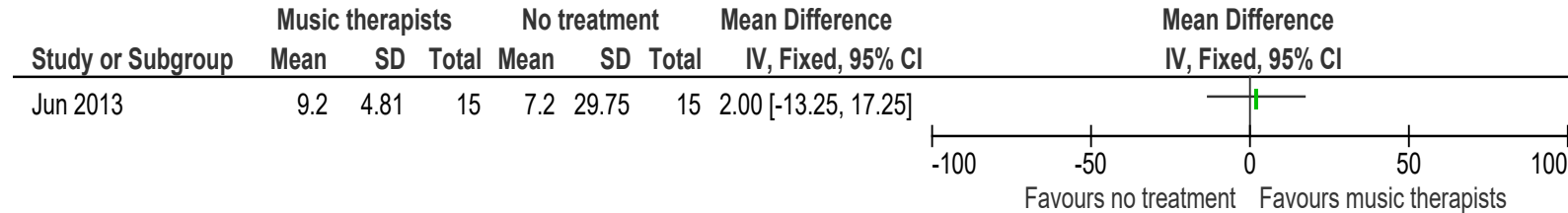


Figure 15: Activities of daily living (Functional independence measure, 18-126, higher values are better, final value) at <6 months

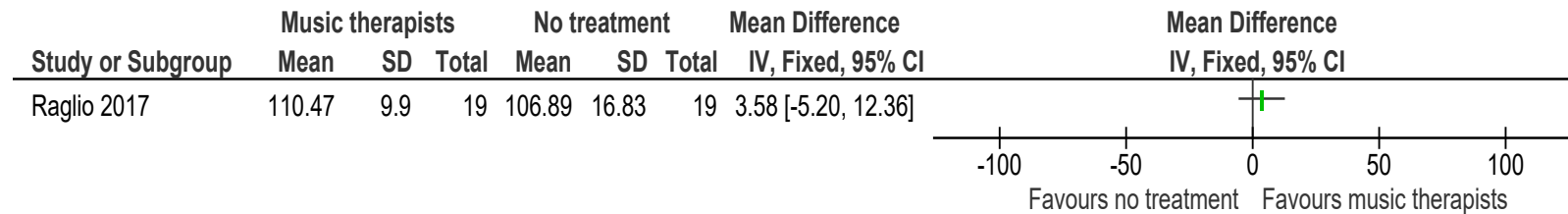


Figure 16: Psychological distress - Depression (HADS-D, BDI, Faces scale, PANAS - negative affect [different scale ranges], lower values are better, final values) at <6 months

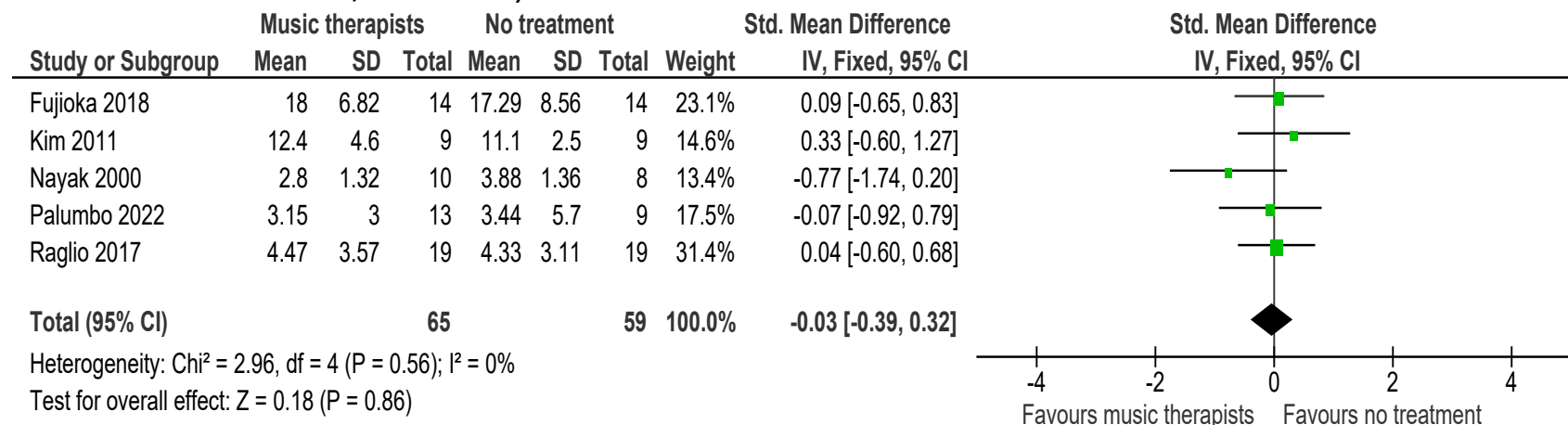


Figure 17: Psychological distress - Depression (Center for Epidemiologic Studies Depression Scale, 0-60, lower values are better, change score) at <6 months

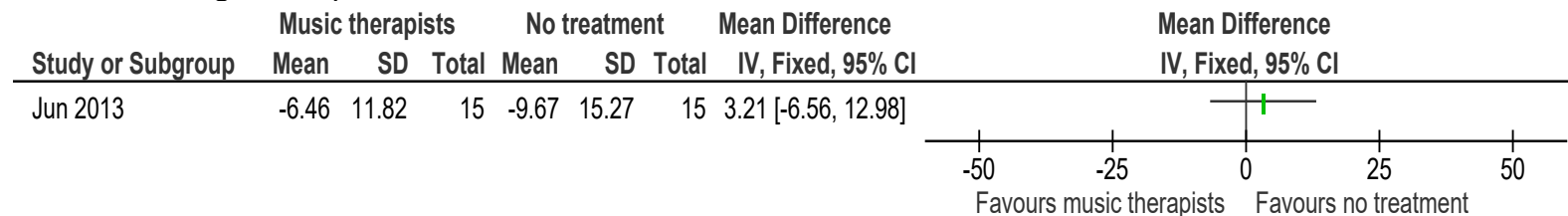


Figure 18: Psychological distress - Anxiety (HADS-A, BAI [different scale ranges], lower values are better, final values) at <6 months

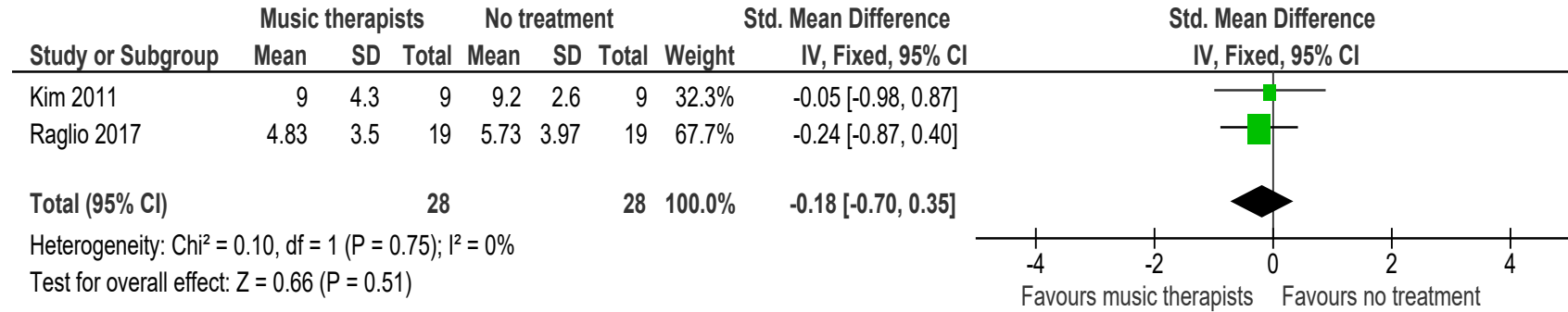


Figure 19: Psychological distress (PANAS - positive affect, 10-50, higher values are better, final value) at <6 months

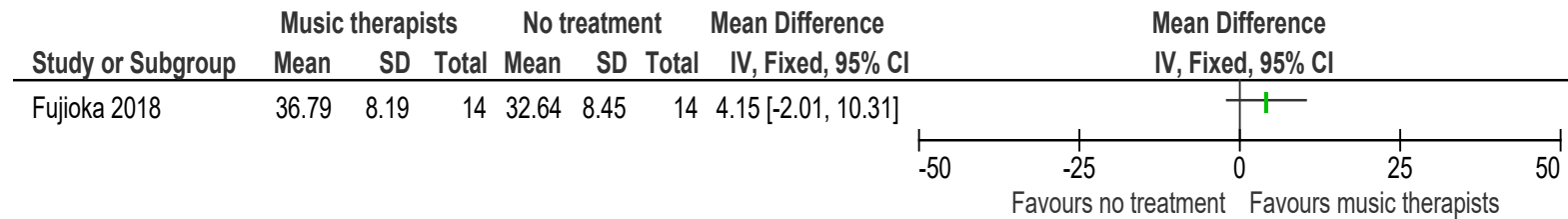


Figure 20: Stroke-specific Patient-Reported Outcome Measures (Stroke specific quality of life, 49-245, higher values are better, final value) at <6 months

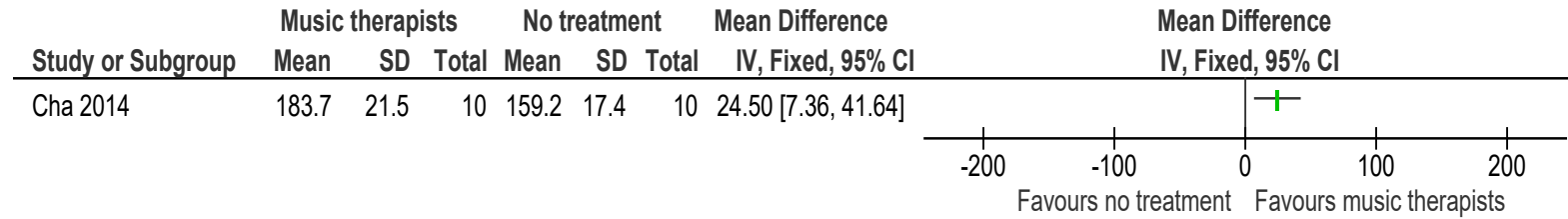


Figure 21: Stroke-specific Patient-Reported Outcome Measures (SIS physical strength, 0-100, higher values are better, final value) at <6 months

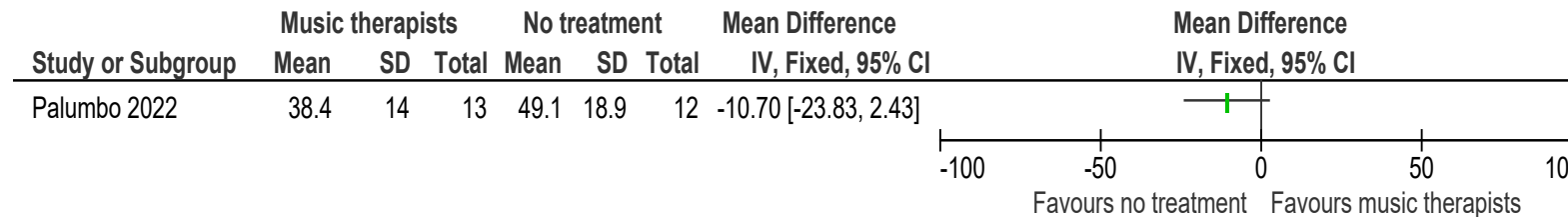


Figure 22: Stroke-specific Patient-Reported Outcome Measures (SIS activities, 0-100, higher values are better, final value) at <6 months

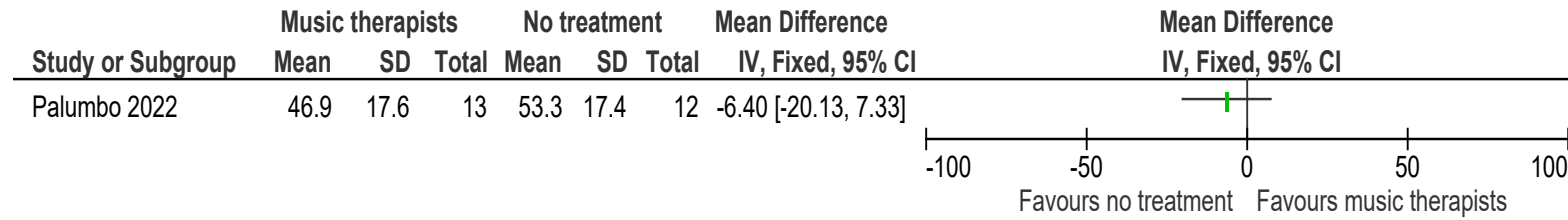


Figure 23: Stroke-specific Patient-Reported Outcome Measures (SIS hand use, 0-100, higher values are better, final value) at <6 months

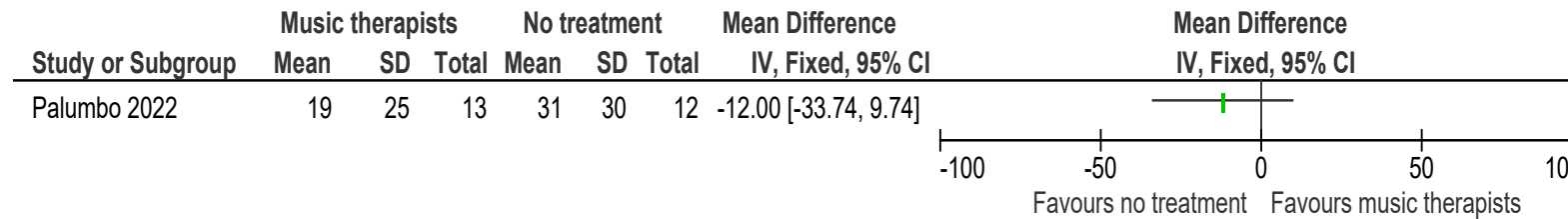


Figure 24: Stroke-specific Patient-Reported Outcome Measures (SIS mobility, 0-100, higher values are better, final value) at <6 months

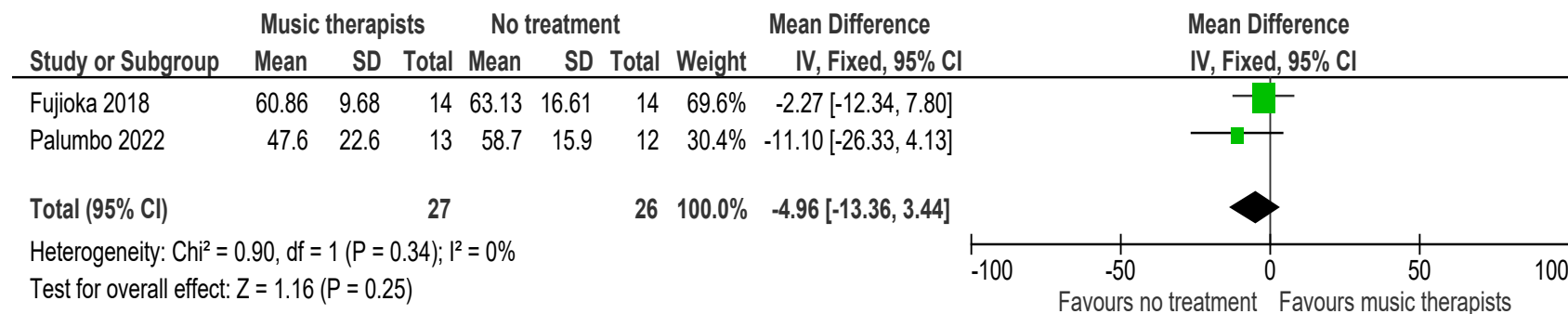


Figure 25: Stroke-specific Patient-Reported Outcome Measures (SIS communication, 0-100, higher values are better, final value) at <6 months

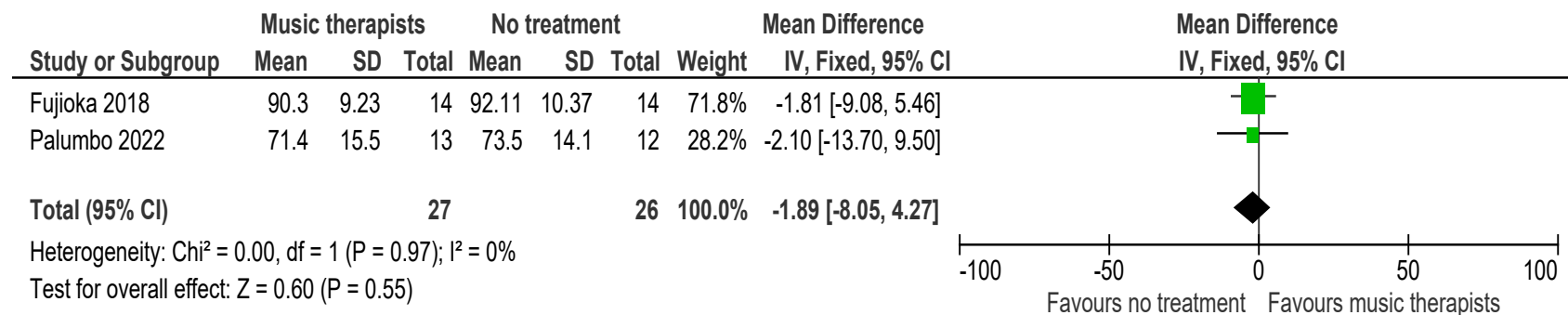


Figure 26: Stroke-specific Patient-Reported Outcome Measures (SIS memory, 0-100, higher values are better, final value) at <6 months

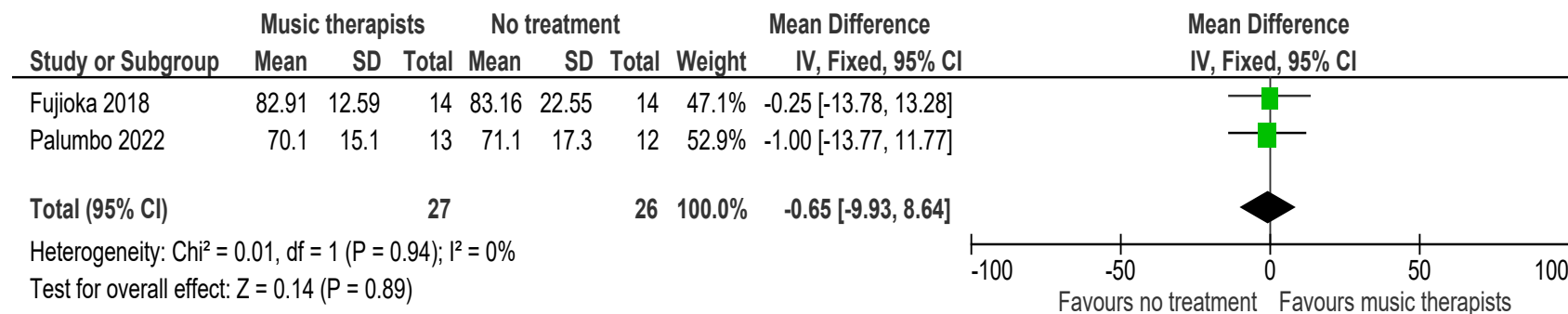


Figure 27: Stroke-specific Patient-Reported Outcome Measures (SIS emotion, 0-100, higher values are better, final value) at <6 months

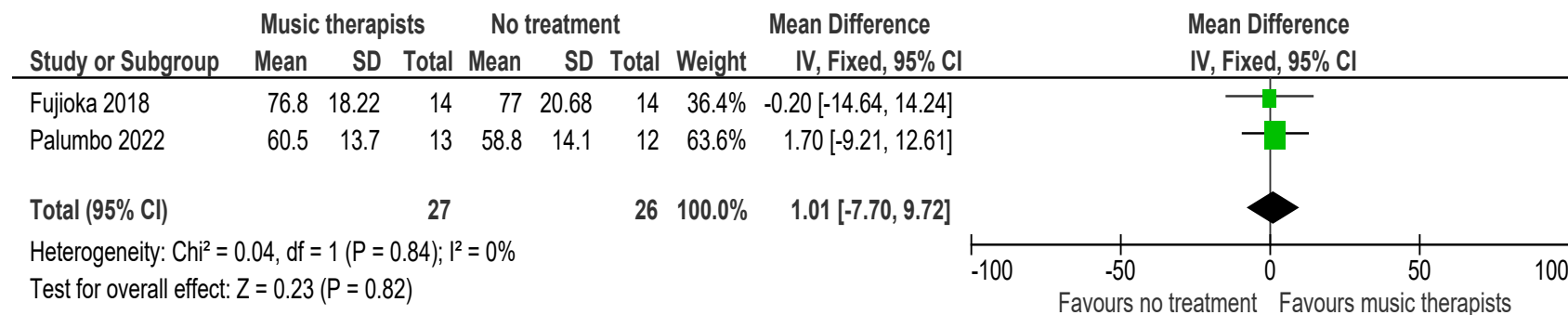


Figure 28: Stroke-specific Patient-Reported Outcome Measures (SIS social/participation, 0-100, higher values are better, final value) at <6 months

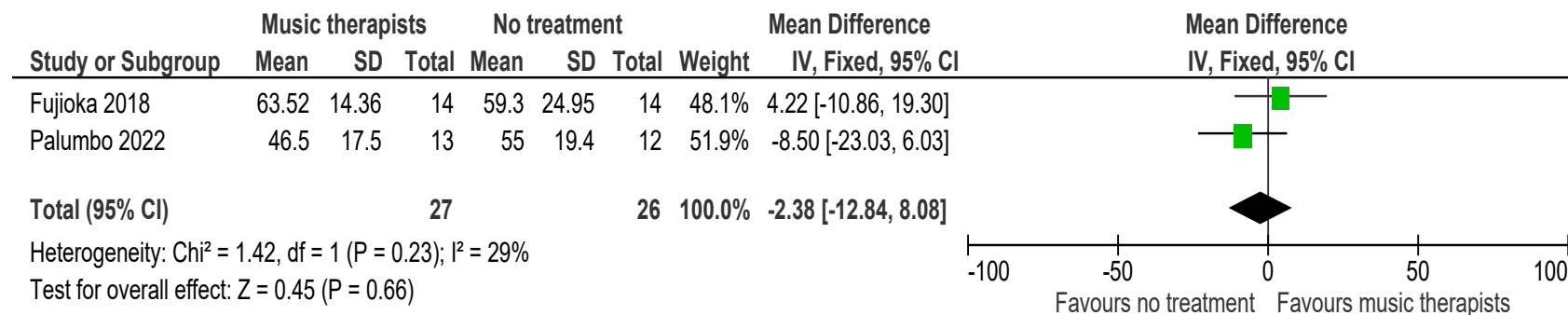


Figure 29: Stroke-specific Patient-Reported Outcome Measures (SIS recovery, 0-100, higher values are better, final value) at <6 months

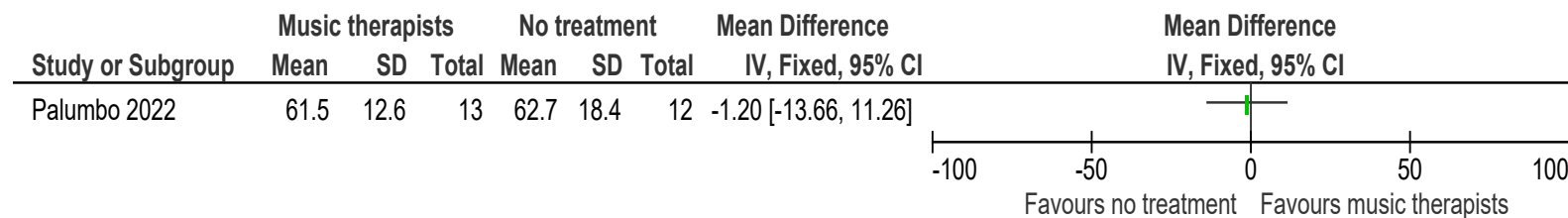


Figure 30: Wellbeing scores (WHO five item well-being index, 0-25, higher values are better, final value) at <6 months

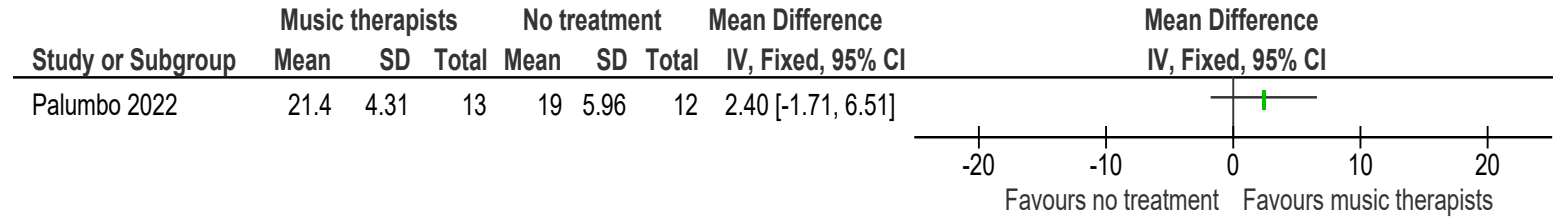
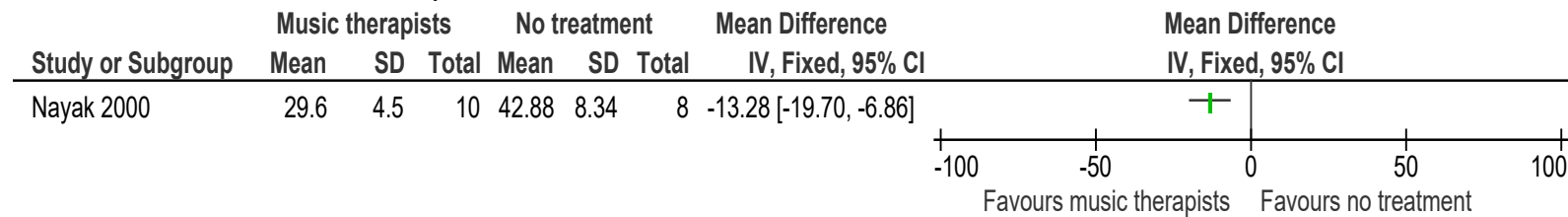
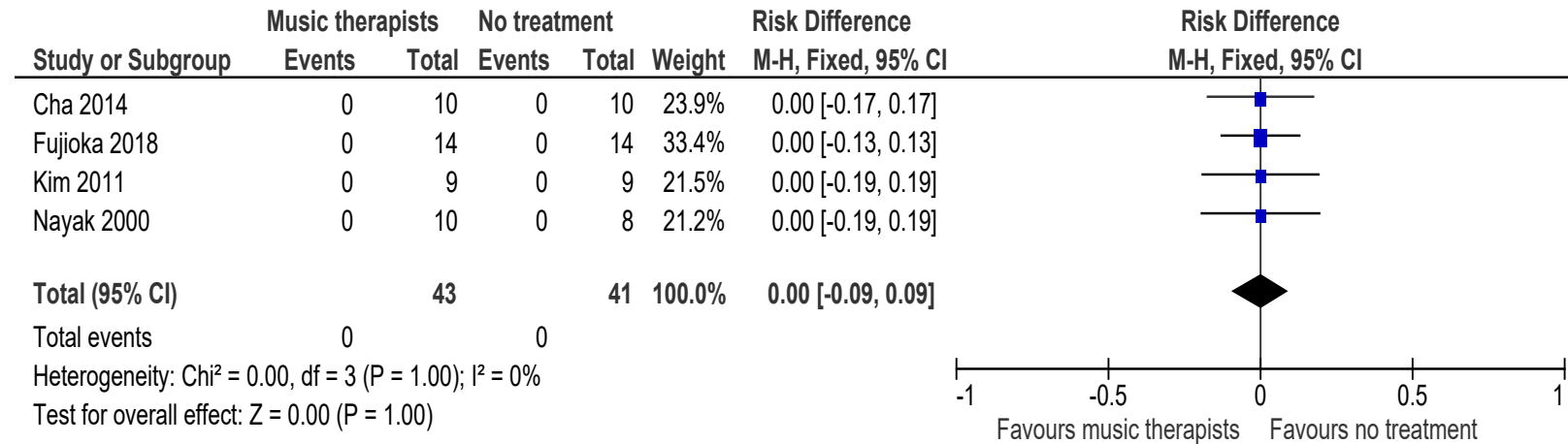


Figure 31: Participation in leisure activities/social groups (Sickness Impact Profile Social Interaction subscale, 0-102, lower values are better, final value) at <6 months



Note: Baseline music therapists: 37.6 (7.55). Baseline no treatment: 44.5 (7.19).

Figure 32: Withdrawal due to adverse events at <6 months



E.3 Music intervention delivered by healthcare professionals compared to passive music listening

Figure 33: Withdrawal due to adverse events at <6 months

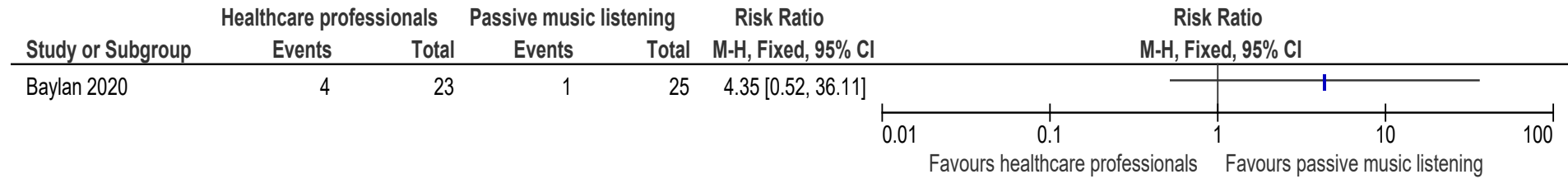
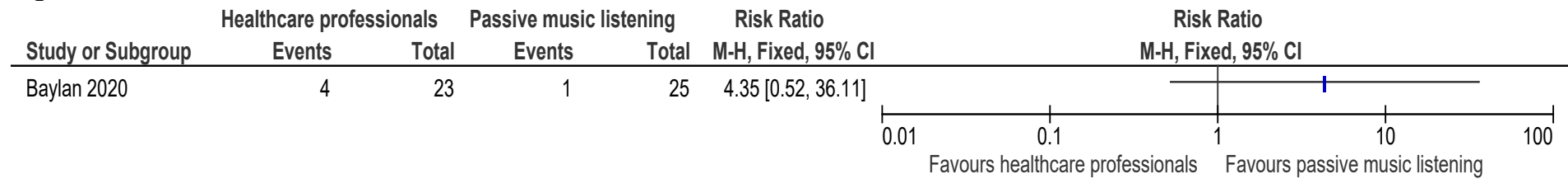


Figure 34: Withdrawal due to adverse events at ≥6 months



E.4 Music intervention delivered by healthcare professionals compared to placebo music therapy

Figure 35: Psychological distress - Depression (HADS-D, 0-42, higher values are better, mean difference) at <6 months

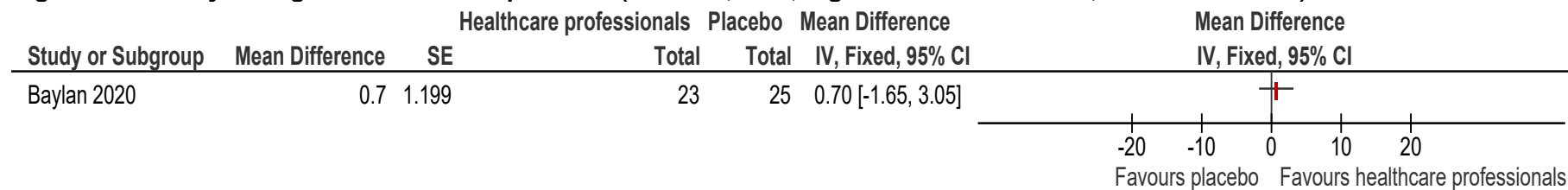


Figure 36: Psychological distress - Depression (HADS-D, 0-42, higher values are better, mean difference) at ≥6 months

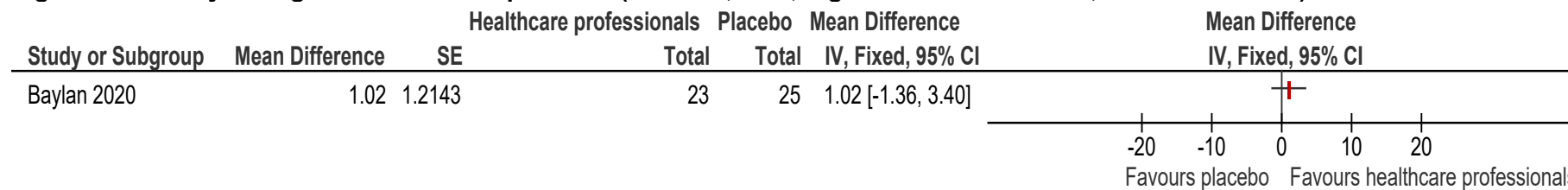


Figure 37: Psychological distress - Anxiety (HADS-A, 0-42, higher values are better, mean difference) at <6 months

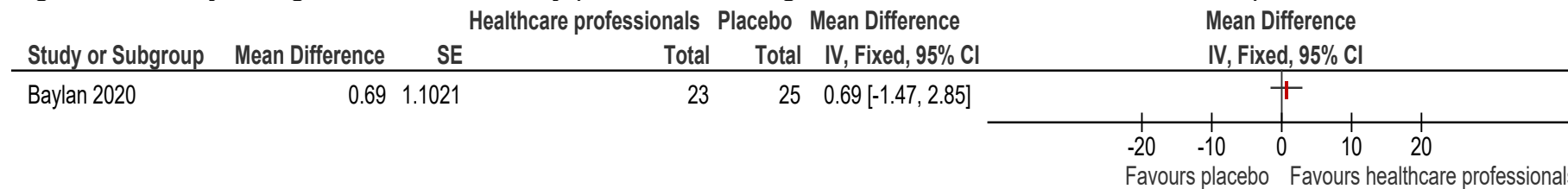


Figure 38: Psychological distress - Anxiety (HADS-A, 0-42, higher values are better, mean difference) at ≥6 months

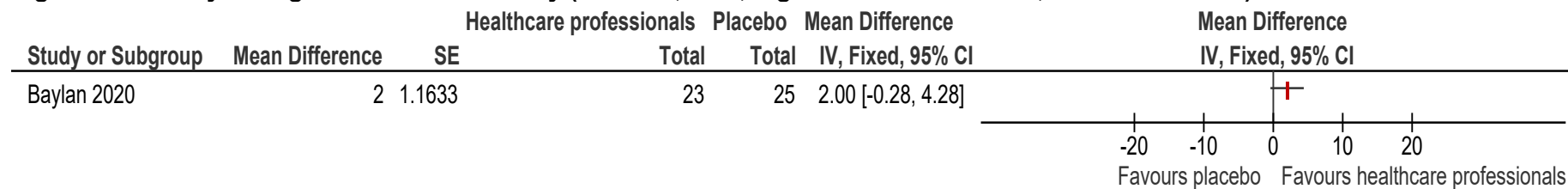


Figure 39: Participation in leisure activities/social group scores (Mayo-Portland Adaptability Inventory 4 participation, 0-30, higher values are better, mean difference) at <6 months

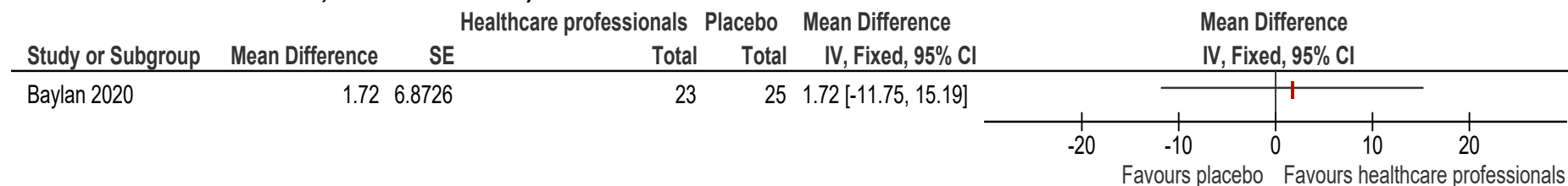


Figure 40: Withdrawal due to adverse events at <6 months

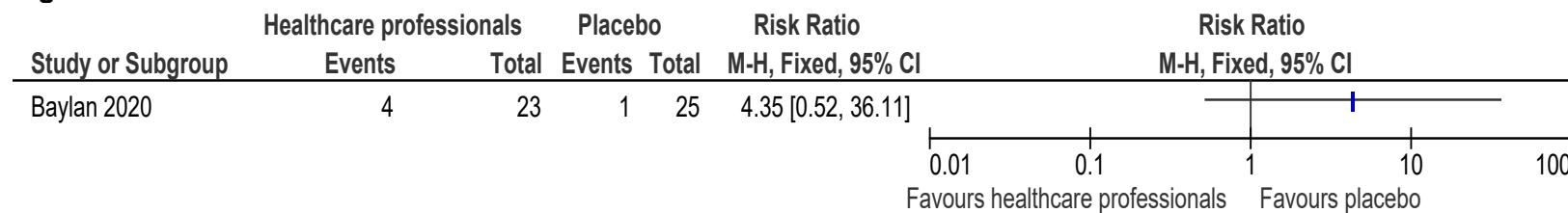
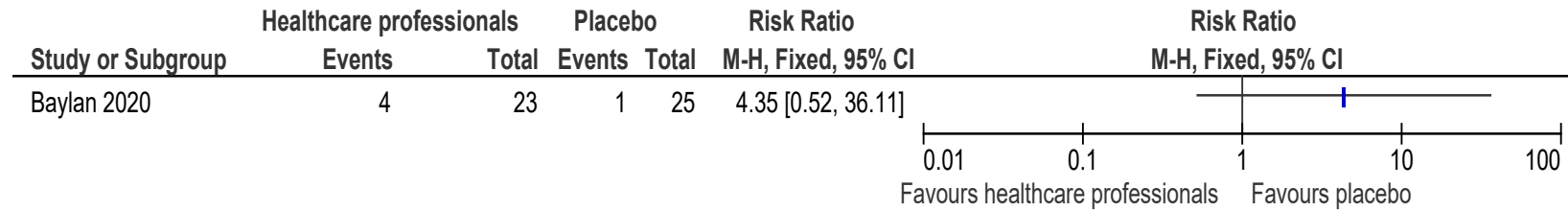


Figure 41: Withdrawal due to adverse events at ≥6 months



E.5 Music intervention delivered by healthcare professionals compared to no treatment

Figure 42: Person/participant generic health-related quality of life (SF-36 physical function, 0-100, higher values are better, final value) at <6 months

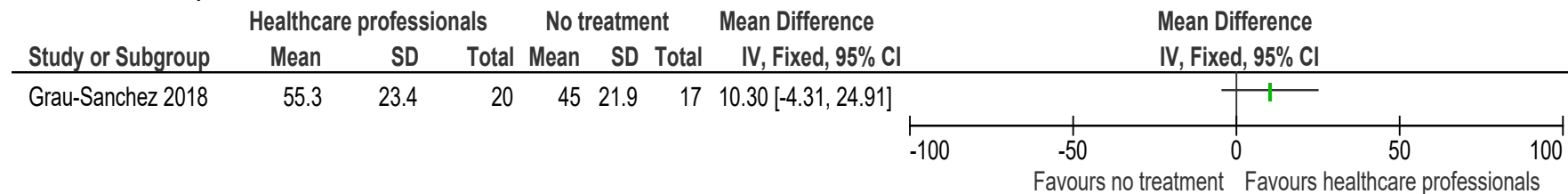


Figure 43: Person/participant generic health-related quality of life (SF-36 bodily pain, 0-100, higher values are better, final value) at <6 months

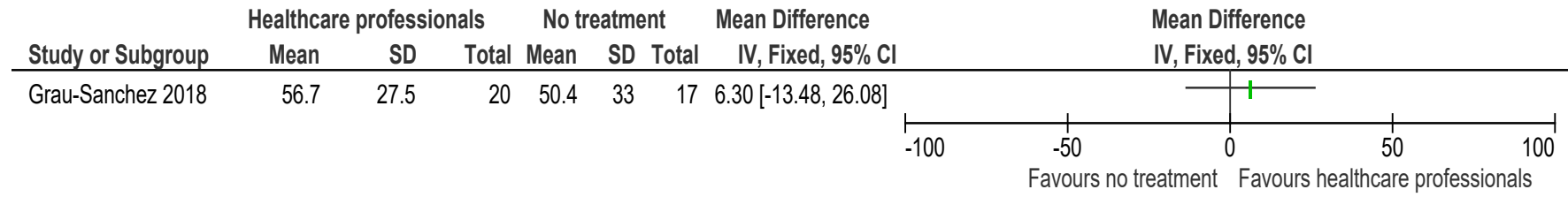


Figure 44: Person/participant generic health-related quality of life (SF-36 role physical, 0-100, higher values are better, final value) at <6 months

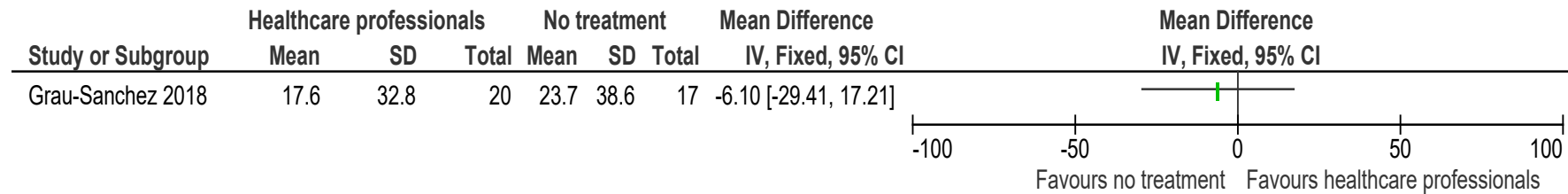


Figure 45: Person/participant generic health-related quality of life (SF-36 vitality, 0-100, higher values are better, final value) at <6 months

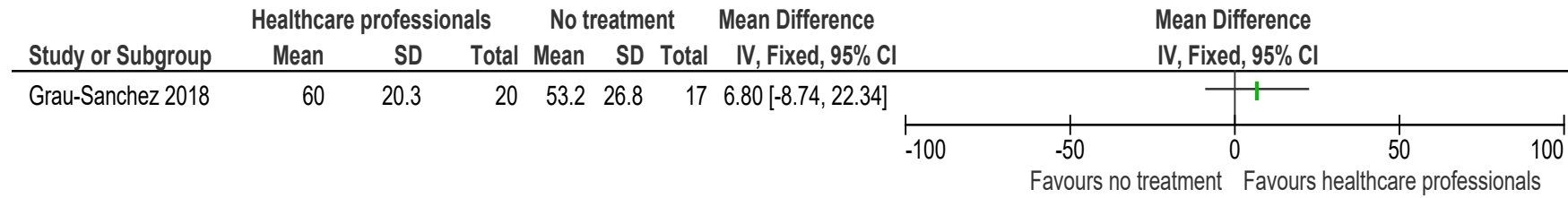


Figure 46: Person/participant generic health-related quality of life (SF-36 general health, 0-100, higher values are better, final value) at <6 months

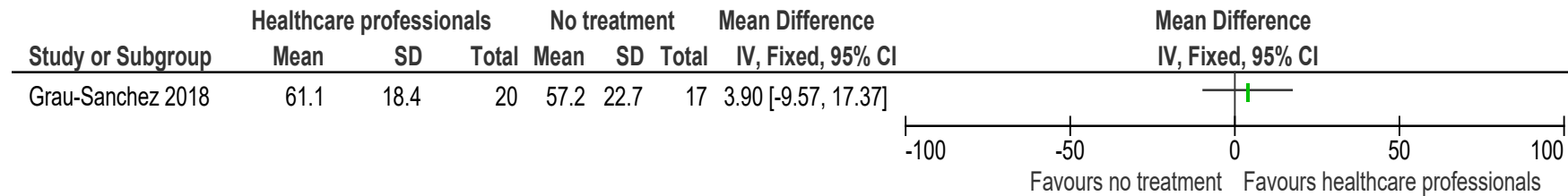


Figure 47: Person/participant generic health-related quality of life (SF-36 role emotional, 0-100, higher values are better, final value) at <6 months

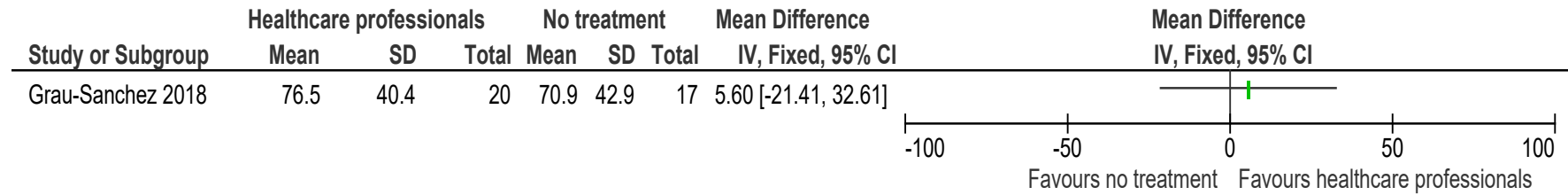


Figure 48: Person/participant generic health-related quality of life (SF-36 mental health, 0-100, higher values are better, final value) at <6 months

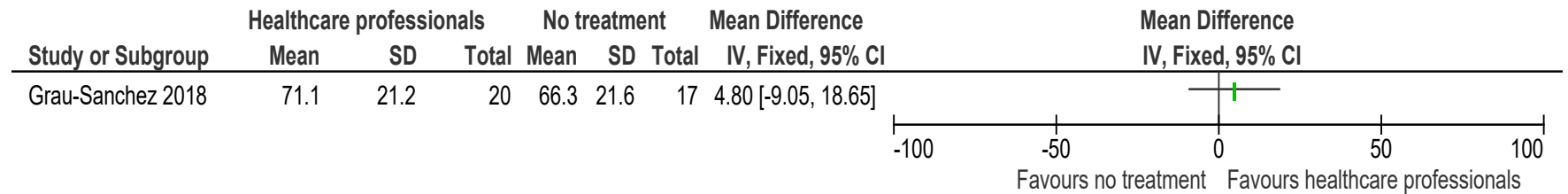


Figure 49: Person/participant generic health-related quality of life (SF-36 social function, 0-100, higher values are better, final value) at <6 months

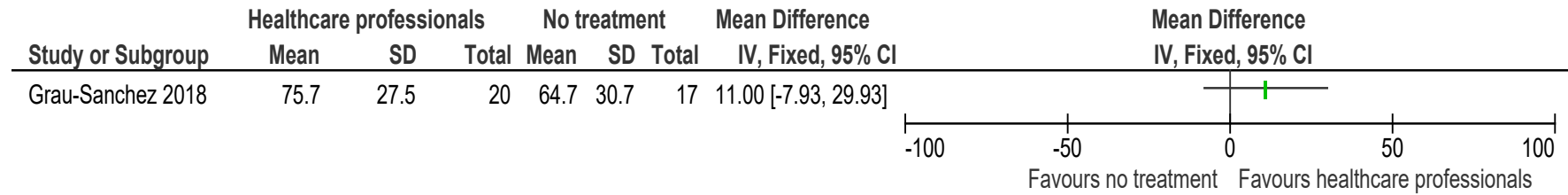


Figure 50: Person/participant health-related quality of life (McGill Quality of Life, 0-10, higher values are better, final values) at <6 months

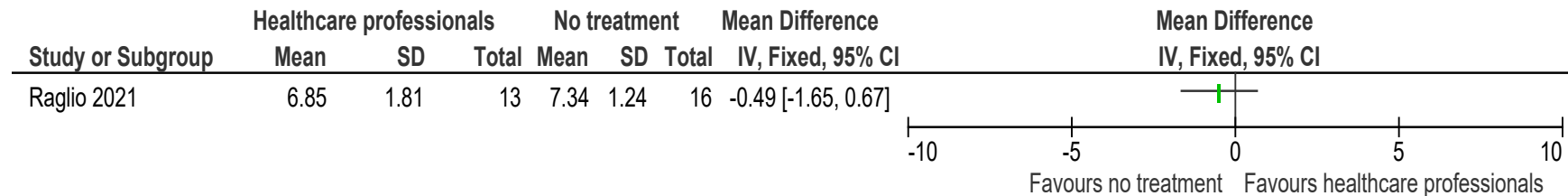


Figure 51: Activities of daily living (Canadian Occupational Performance Measure, University of Maryland Arm Questionnaire for Stroke, Activities of daily living score [different scale ranges], higher values are better, change scores) at <6 months

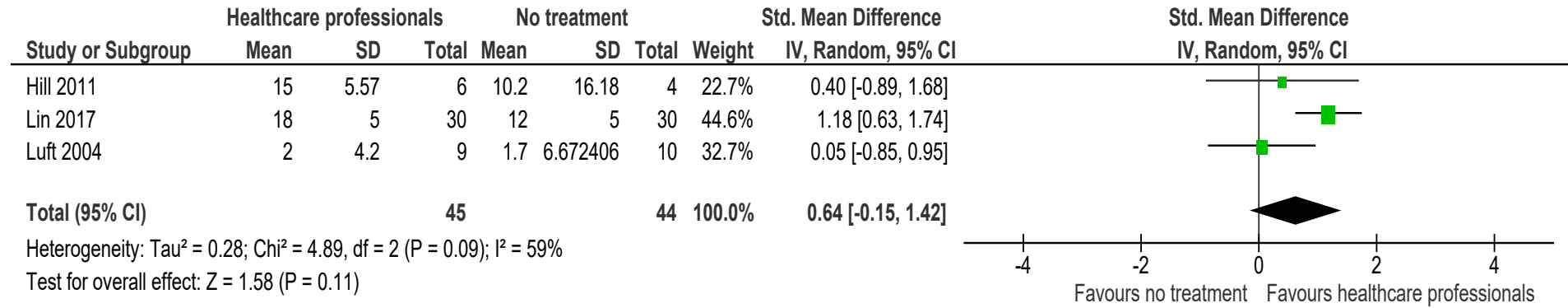


Figure 52: Activities of daily living (Barthel index, 0-100, higher values are better, final value) at <6 months

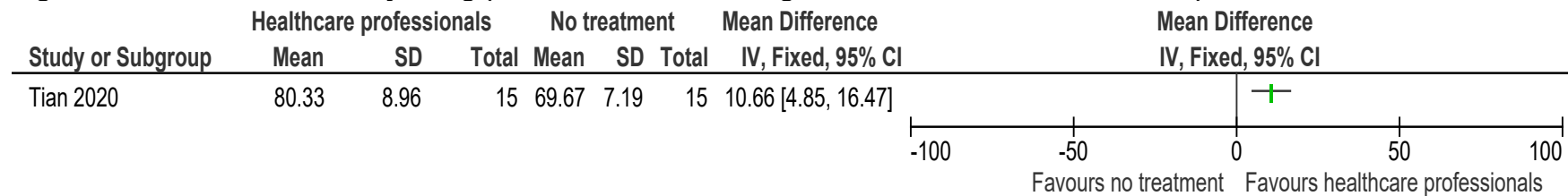


Figure 53: Psychological distress - Depression (Hamilton's Depression Scale-17, 0-56, lower values are better, change score) at <6 months

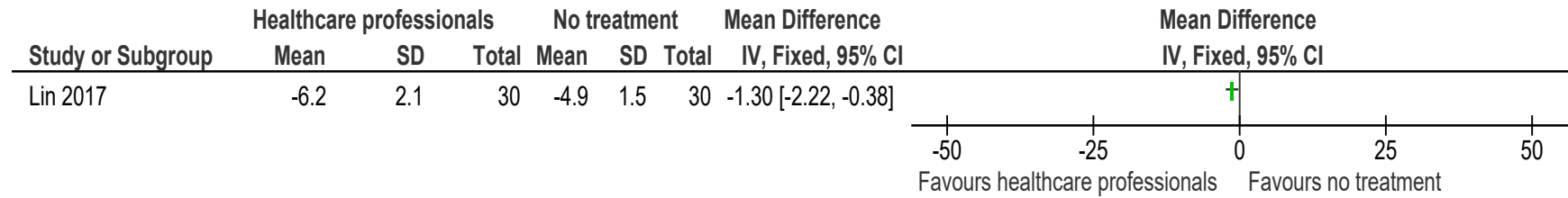


Figure 54: Psychological distress - Depression (BDI, Hamilton Depression Rating Scale-24, profile of mood states [different scale ranges], lower values are better, final values) at <6 months

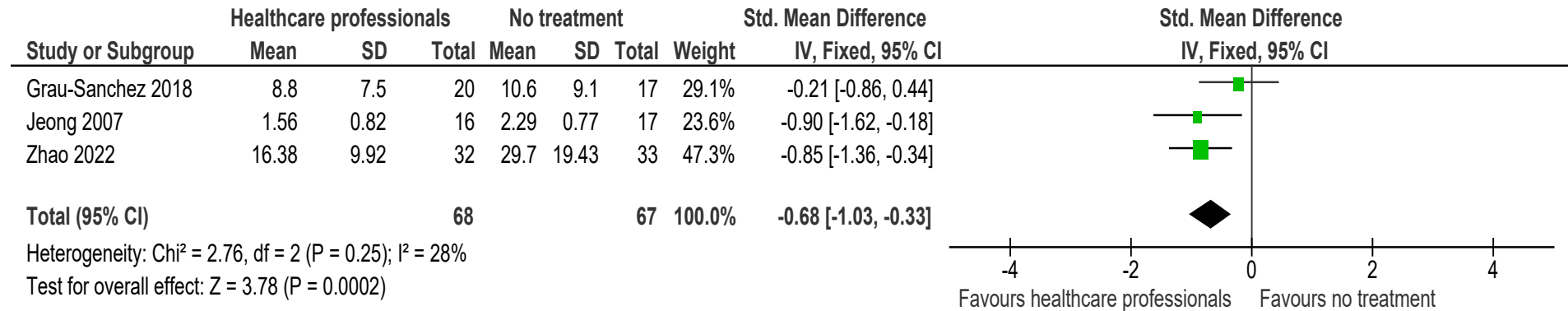


Figure 55: Psychological distress (PANAS positive affect, 10-50, higher values are better, change score) at <6 months

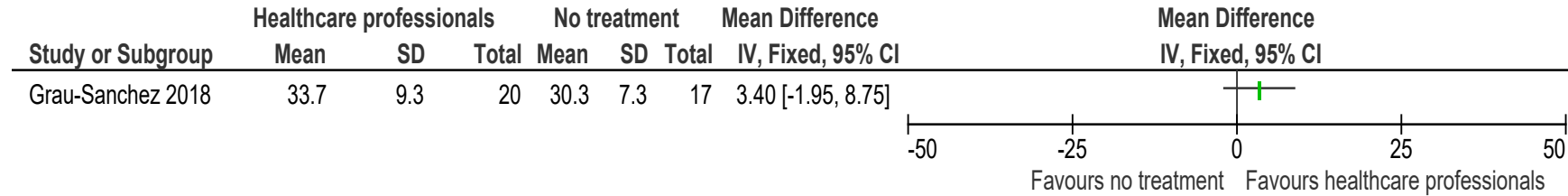


Figure 56: Stroke-specific Patient-Reported Outcome Measures (SIS [different scale ranges], higher values are better, change scores) at <6 months

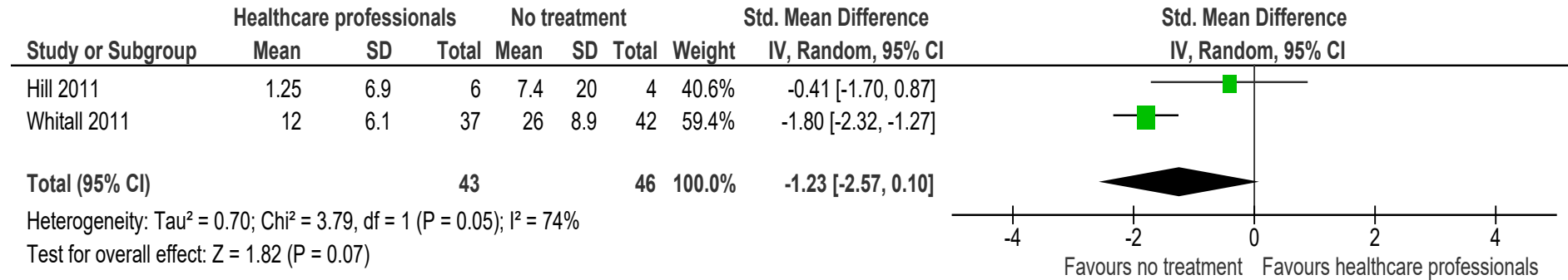


Figure 57: Stroke-specific Patient-Reported Outcome Measures (SIS strength subscale, 0-100, higher values are better, change score) at <6 months

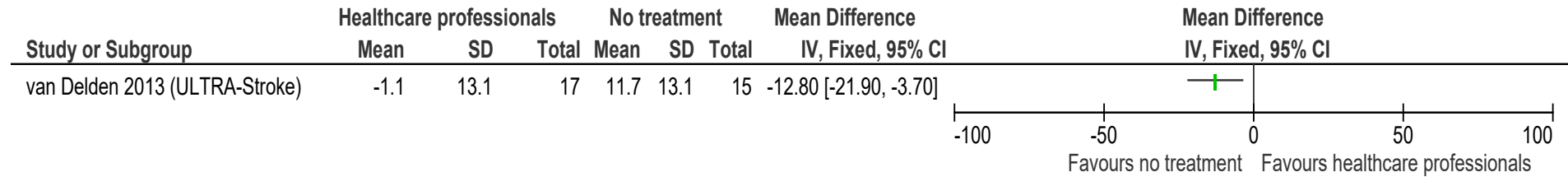


Figure 58: Stroke-specific Patient-Reported Outcome Measures (SIS memory subscale, 0-100, higher values are better, change score) at <6 months

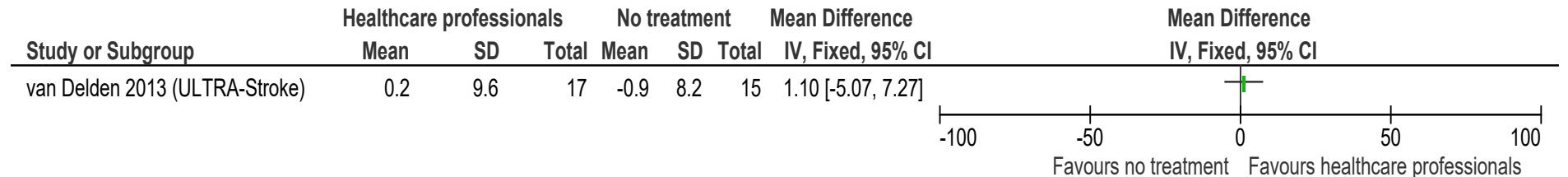


Figure 59: Stroke-specific Patient-Reported Outcome Measures (SIS emotion subscale, 0-100, higher values are better, change score) at <6 months

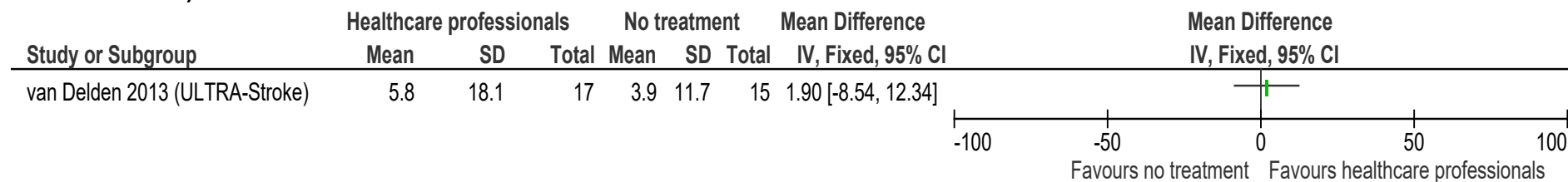


Figure 60: Stroke-specific Patient-Reported Outcome Measures (SIS communication subscale, 0-100, higher values are better, change score) at <6 months

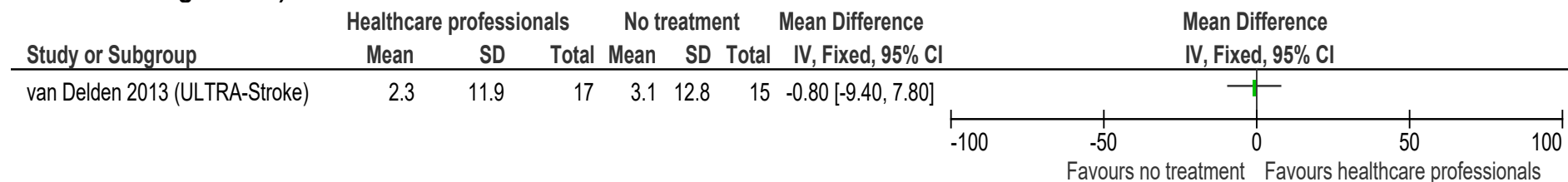


Figure 61: Stroke-specific Patient-Reported Outcome Measures (SIS ADL subscale, 0-100, higher values are better, change score) at <6 months

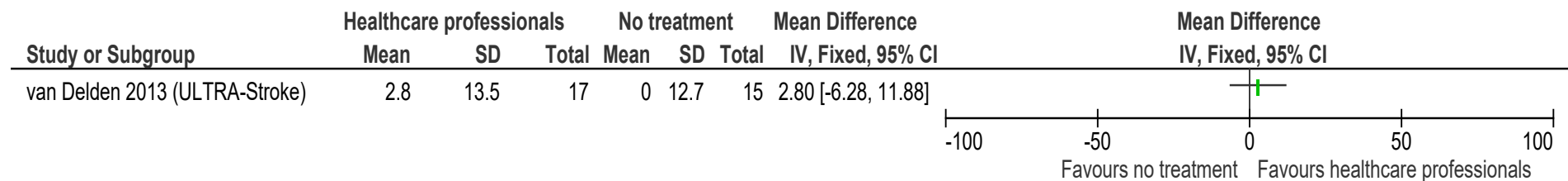


Figure 62: Stroke-specific Patient-Reported Outcome Measures (SIS mobility subscale, 0-100, higher values are better, change score) at <6 months

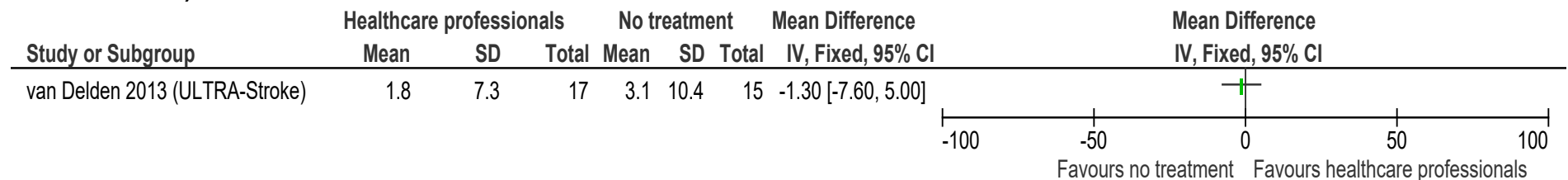


Figure 63: Stroke-specific Patient-Reported Outcome Measures (SIS hand function subscale, 0-100, higher values are better, change score) at <6 months

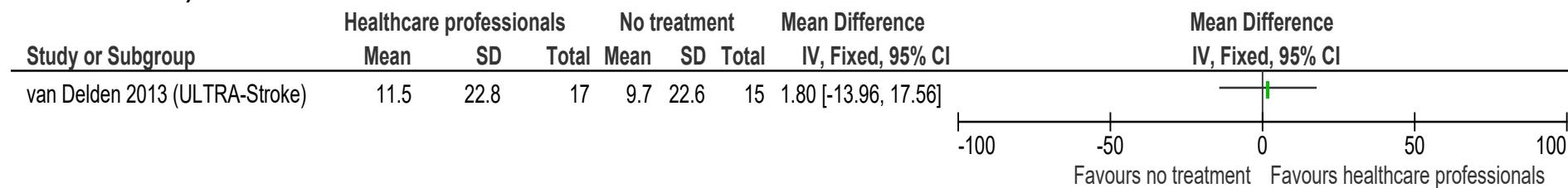


Figure 64: Stroke-specific Patient-Reported Outcome Measures (SIS social participation subscale, 0-100, higher values are better, change score) at <6 months

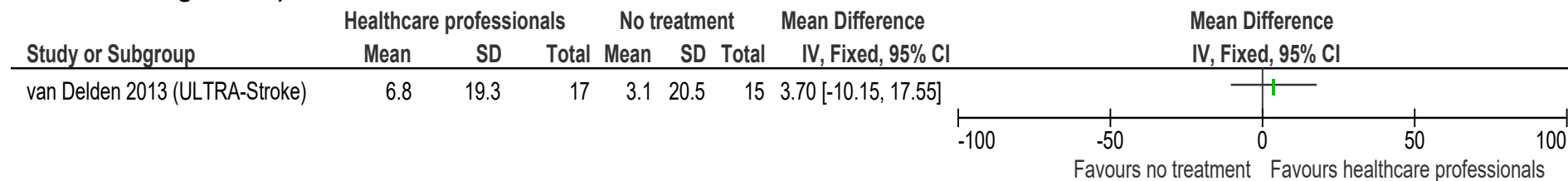


Figure 65: Stroke-specific Patient-Reported Outcome Measures (Stroke Specific Quality of Life [different scale ranges], higher values are better, final values) at <6 months

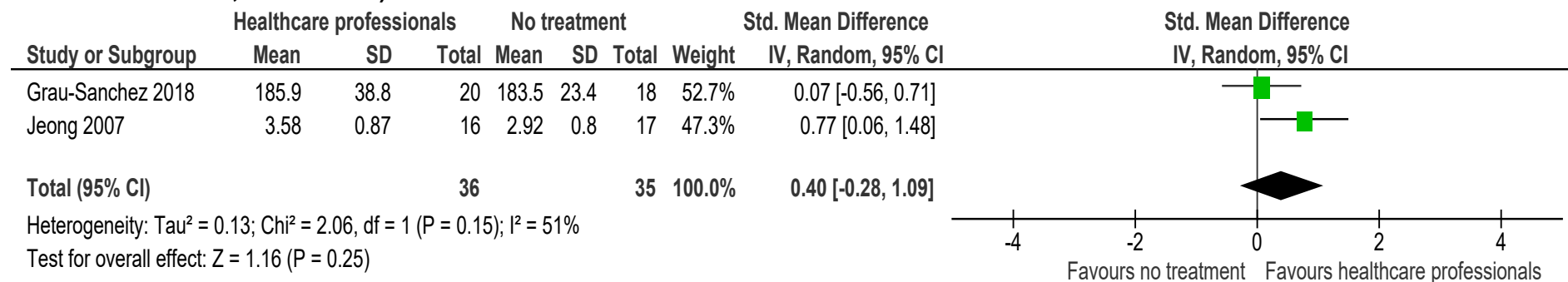
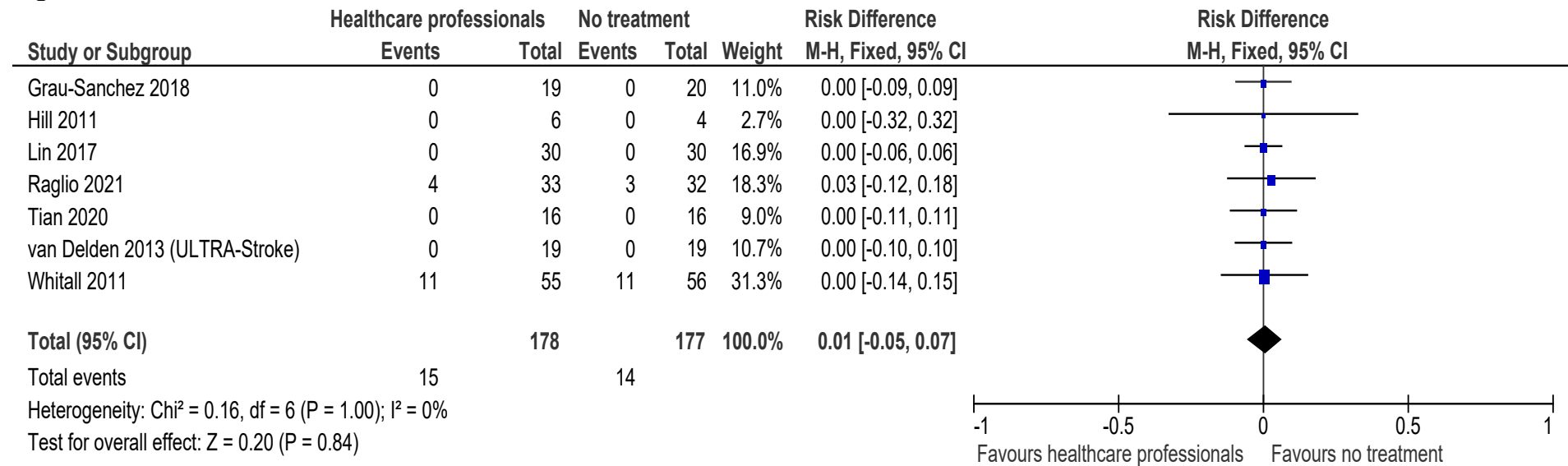


Figure 66: Withdrawal due to adverse events at <6 months



E.6 Music intervention delivered by non-healthcare professionals compared to no treatment

Figure 67: Person/participant generic health-related quality of life (EQ-5D 5L, -0.11-1, higher values are better, final value) at <6 months

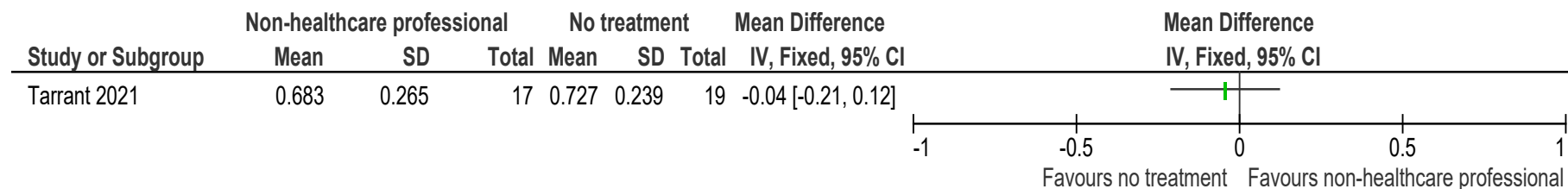


Figure 68: Person/participant generic health-related quality of life (EQ-5D 5L, -0.11-1, higher values are better, final value) at ≥6 months

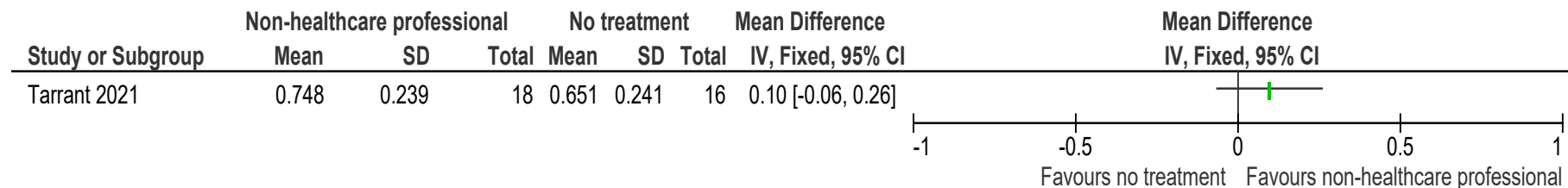


Figure 69: Carer generic health-related quality of life (CarerQoL-7D, 0-14, higher values are better, final value) at ≥6 months

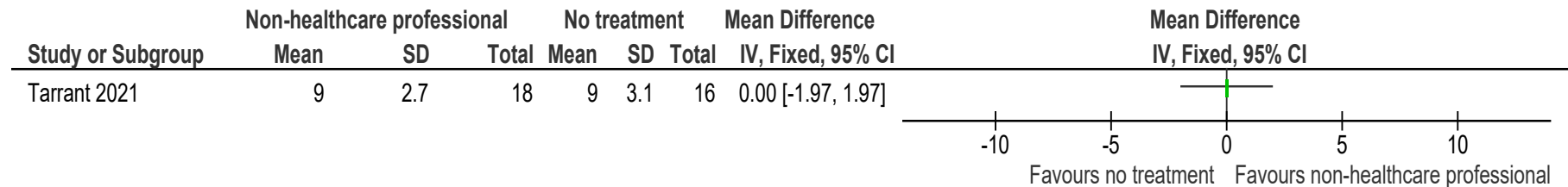


Figure 70: Stroke-specific Patient-Reported Outcome Measures (Stroke and Aphasia Quality of Life, 1-5, higher values are better, final value) at <6 months

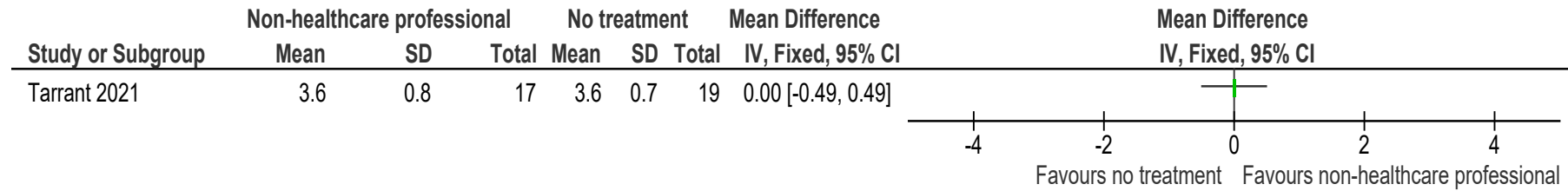


Figure 71: Stroke-specific Patient-Reported Outcome Measures (Stroke and Aphasia Quality of Life, 1-5, higher values are better, final value) at ≥6 months

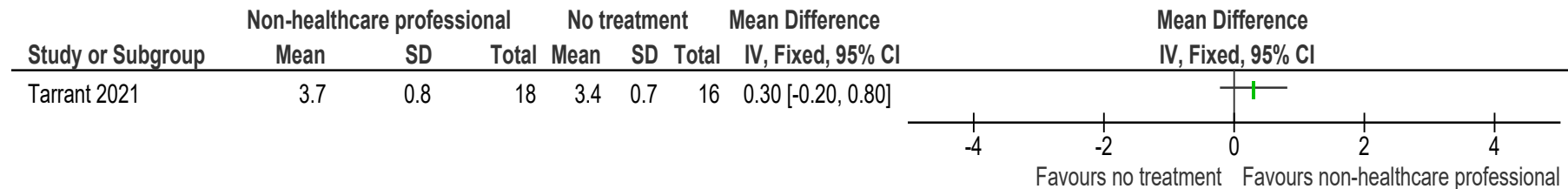


Figure 72: Wellbeing scores (ICEpop CAPability measure for adults, 0-1, higher values are better, final value) at <6 months

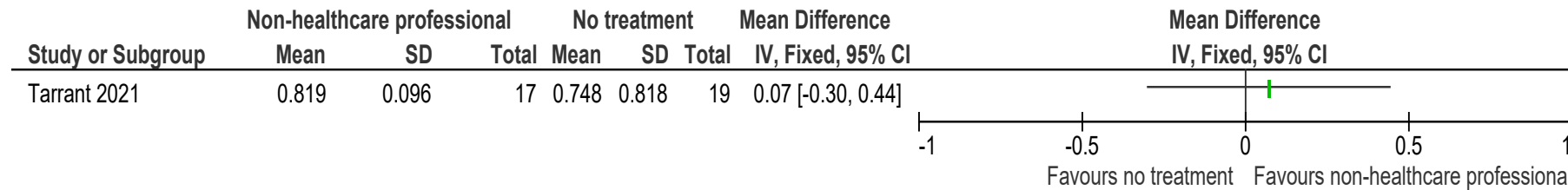


Figure 73: Participation in leisure activities/social groups scores (modified Reintegration to Normal Living Index, 0-100, higher values are better, final value) at <6 months

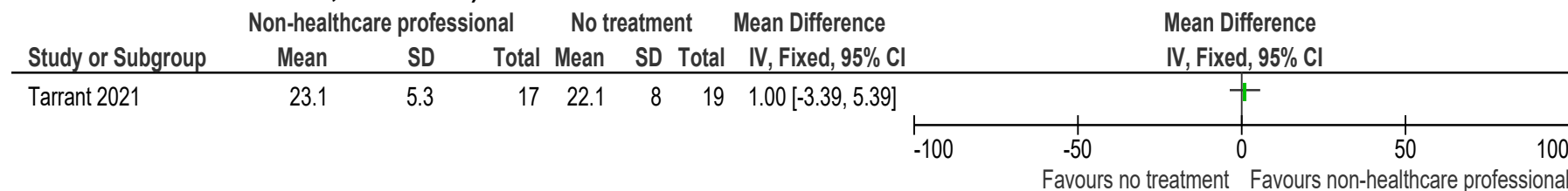


Figure 74: Wellbeing scores (ICEpop CAPability measure for adults, 0-1, higher values are better, final value) at ≥6 months

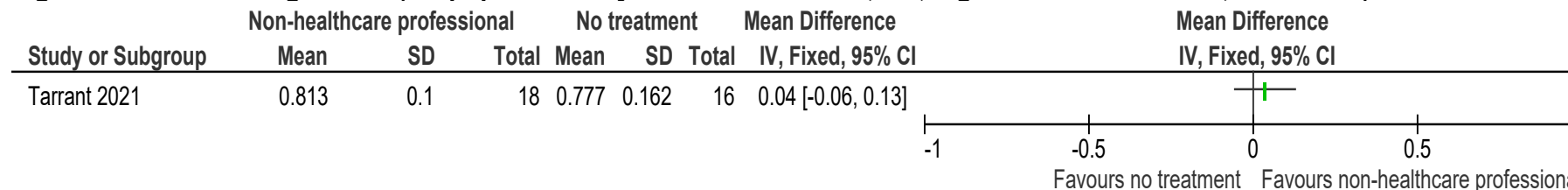


Figure 75: Participation in leisure activities/social groups scores (modified Reintegration to Normal Living Index, 0-100, higher values are better, final value) at ≥6 months

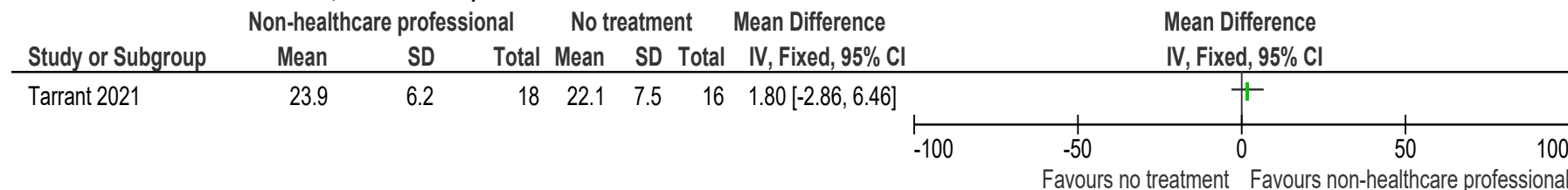
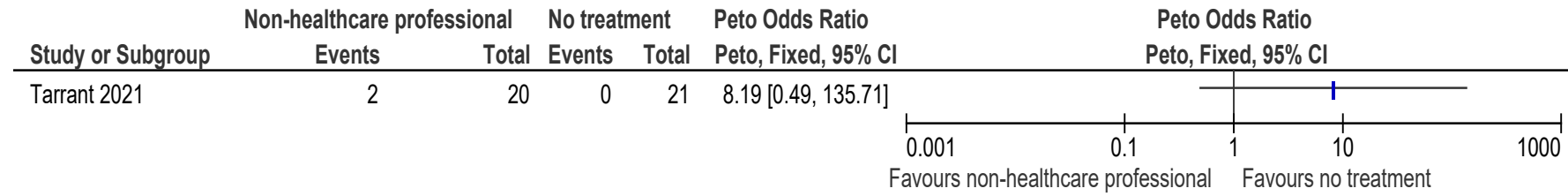


Figure 76: Withdrawal due to adverse events at <6 months

Appendix F – GRADE tables

F.1 Neurologic music therapy delivered by trained music therapists compared to no treatment

Table 16: Clinical evidence profile: neurologic music therapy delivered by trained music therapists compared to no treatment

| Certainty assessment | | | | | | | No of patients | | Effect | | Certainty | Importance |
|----------------------|--------------|--------------|---------------|--------------|-------------|----------------------|--|--------------|-------------------|-------------------|-----------|------------|
| No of studies | Study design | Risk of bias | Inconsistency | Indirectness | Imprecision | Other considerations | neurologic music therapy delivered by trained music therapists | no treatment | Relative (95% CI) | Absolute (95% CI) | | |

Person/participant generic health-related quality of life (SF-36 physical functioning, 0-100, higher values are better, final values) at <6 months (follow-up: 5 weeks; assessed with: SF-36 physical functioning; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|--|------------------|----------|
| 1 | randomised trials | very serious ^a | not serious | not serious | serious ^b | none | 30 | 31 | - | MD 1.43 higher (1.11 lower to 3.97 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|--|------------------|----------|

Person/participant generic health-related quality of life (SF-36 bodily pain, 0-100, higher values are better, final values) at <6 months (follow-up: 5 weeks; assessed with: SF-36 bodily pain; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|-------------|------|----|----|---|--|-------------|----------|
| 1 | randomised trials | very serious ^a | not serious | not serious | not serious | none | 30 | 31 | - | MD 0.47 higher (0.78 lower to 1.72 higher) | ⊕⊕○○ Low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|-------------|------|----|----|---|--|-------------|----------|

Person/participant generic health-related quality of life (SF-36 role physical, 0-100, higher values are better, final values) at <6 months (follow-up: 5 weeks; assessed with: SF-36 role physical; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|-------------|------|----|----|---|--|-------------|----------|
| 1 | randomised trials | very serious ^a | not serious | not serious | not serious | none | 30 | 31 | - | MD 0.29 higher (0.21 lower to 0.79 higher) | ⊕⊕○○ Low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|-------------|------|----|----|---|--|-------------|----------|

Person/participant generic health-related quality of life (SF-36 vitality, 0-100, higher values are better, final values) at <6 months (follow-up: 5 weeks; assessed with: SF-36 vitality; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|-------------|------|----|----|---|---|-------------|----------|
| 1 | randomised trials | very serious ^a | not serious | not serious | not serious | none | 30 | 31 | - | MD 4.11 higher (2.34 higher to 5.88 higher) | ⊕⊕○○ Low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|-------------|------|----|----|---|---|-------------|----------|

| Certainty assessment | | | | | | | No of patients | | Effect | | Certainty | Importance |
|----------------------|--------------|--------------|---------------|--------------|-------------|----------------------|--|--------------|-------------------|-------------------|-----------|------------|
| No of studies | Study design | Risk of bias | Inconsistency | Indirectness | Imprecision | Other considerations | neurologic music therapy delivered by trained music therapists | no treatment | Relative (95% CI) | Absolute (95% CI) | | |

Person/participant generic health-related quality of life (SF-36 general health, 0-100, higher values are better, final values) at <6 months (follow-up: 5 weeks; assessed with: SF-36 general health; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|--|------------------|----------|
| 1 | randomised trials | very serious ^a | not serious | not serious | serious ^b | none | 30 | 31 | - | MD 1.46 higher (0.25 lower to 3.17 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|--|------------------|----------|

Person/participant generic health-related quality of life (SF-36 role emotional, 0-100, higher values are better, final values) at <6 months (follow-up: 5 weeks; assessed with: SF-36 role emotional; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|-------------|------|----|----|---|---|-------------|----------|
| 1 | randomised trials | very serious ^a | not serious | not serious | not serious | none | 30 | 31 | - | MD 0.83 higher (0.34 higher to 1.32 higher) | ⊕⊕○○ Low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|-------------|------|----|----|---|---|-------------|----------|

Person/participant generic health-related quality of life (SF-36 mental health, 0-100, higher values are better, final values) at <6 months (follow-up: 5 weeks; assessed with: SF-36 mental health; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|---|------------------|----------|
| 1 | randomised trials | very serious ^a | not serious | not serious | serious ^b | none | 30 | 31 | - | MD 3.75 higher (1.32 higher to 6.18 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|---|------------------|----------|

Person/participant generic health-related quality of life (SF-36 social functioning, 0-100, higher values are better, final values) at <6 months (follow-up: 5 weeks; assessed with: SF-36 social functioning; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|-------------|------|----|----|---|--|-------------|----------|
| 1 | randomised trials | very serious ^a | not serious | not serious | not serious | none | 30 | 31 | - | MD 0.56 higher (0.54 lower to 1.66 higher) | ⊕⊕○○ Low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|-------------|------|----|----|---|--|-------------|----------|

Psychological distress - depression (Hamilton Depression Scale, 0-56, lower values are better, final value) at <6 months (follow-up: 8 weeks; assessed with: Hamilton Depression Scale; Scale from: 0 to 56)

| | | | | | | | | | | | | |
|---|-------------------|----------------------|-------------|-------------|----------------------|------|----|----|---|--|-------------|----------|
| 1 | randomised trials | serious ^c | not serious | not serious | serious ^b | none | 20 | 20 | - | MD 1.95 lower (3.07 lower to 0.83 lower) | ⊕⊕○○ Low | CRITICAL |
|---|-------------------|----------------------|-------------|-------------|----------------------|------|----|----|---|--|-------------|----------|

Psychological distress - anxiety (Hamilton Anxiety Rating Scale, 0-56, lower values are better, final value) at <6 months (follow-up: 8 weeks; assessed with: Hamilton Anxiety Rating Scale; Scale from: 0 to 56)

| Certainty assessment | | | | | | | № of patients | | Effect | | Certainty | Importance |
|----------------------|-------------------|----------------------|---------------|--------------|----------------------|----------------------|--|--------------|-------------------|---|-------------|------------|
| № of studies | Study design | Risk of bias | Inconsistency | Indirectness | Imprecision | Other considerations | neurologic music therapy delivered by trained music therapists | no treatment | Relative (95% CI) | Absolute (95% CI) | | |
| 1 | randomised trials | serious ^c | not serious | not serious | serious ^b | none | 20 | 20 | - | MD 1.05 lower (2.46 lower to 0.36 higher) | ⊕⊕○○ Low | CRITICAL |

Stroke-specific Patient-Reported Outcome Measures (Stroke adjusted Sickness Impact Profile 30, 0-68, lower values are better, final values) at <6 months (follow-up: 5 weeks; assessed with: Stroke adjusted Sickness Impact Profile 30; Scale from: 0 to 68)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|--|------------------|----------|
| 1 | randomised trials | very serious ^a | not serious | not serious | serious ^b | none | 30 | 31 | - | MD 1.52 lower (3.84 lower to 0.8 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|--|------------------|----------|

CI: confidence interval; MD: mean difference

Explanations

- a. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias arising from the randomisation process, bias due to deviations from the intended interventions and bias in measurement of the outcome)
- b. Downgraded by 1 increment if the confidence interval crossed one MID or by 2 increments if the confidence interval crossed both MIDs
- c. Downgraded by 1 increment as the majority of the evidence was of high risk of bias (due to bias arising from the randomisation process)

F.2 Music therapy delivered by trained music therapists compared to no treatment

Table 17: Clinical evidence profile: music therapy delivered by trained music therapists compared to no treatment

| Certainty assessment | | | | | | | № of patients | | Effect | | Certainty | Importance |
|----------------------|--------------|--------------|---------------|--------------|-------------|----------------------|---|--------------|-------------------|-------------------|-----------|------------|
| № of studies | Study design | Risk of bias | Inconsistency | Indirectness | Imprecision | Other considerations | music therapy delivered by trained music therapists | no treatment | Relative (95% CI) | Absolute (95% CI) | | |

Patient/participant generic health-related quality of life (McGill Quality of Life, 0-10, higher values are better, final value) at <6 months (follow-up: 2 months; assessed with: McGill Quality of Life; Scale from: 0 to 10)

| Certainty assessment | | | | | | | No of patients | | Effect | | Certainty | Importance |
|--|-------------------|---------------------------|---------------|--------------|---------------------------|----------------------|---|--------------|-------------------|--|------------------|------------|
| No of studies | Study design | Risk of bias | Inconsistency | Indirectness | Imprecision | Other considerations | music therapy delivered by trained music therapists | no treatment | Relative (95% CI) | Absolute (95% CI) | | |
| 1 | randomised trials | very serious ^a | not serious | not serious | serious ^b | none | 19 | 19 | - | MD 0.27 higher (0.7 lower to 1.24 higher) | ⊕○○○ Very low | CRITICAL |
| Activities of daily living (Korean-Modified Barthel Index, 0-100, higher values are better, change score) at <6 months (follow-up: 8 weeks; assessed with: Korean-Modified Barthel Index; Scale from: 0 to 100) | | | | | | | | | | | | |
| 1 | randomised trials | very serious ^c | not serious | not serious | very serious ^b | none | 15 | 15 | - | MD 2 higher (13.25 lower to 17.25 higher) | ⊕○○○ Very low | CRITICAL |
| Activities of daily living (Functional independence measure, 18-126, higher values are better, final value) at <6 months (follow-up: 2 months; assessed with: Functional independence measure; Scale from: 18 to 126) | | | | | | | | | | | | |
| 1 | randomised trials | very serious ^a | not serious | not serious | not serious | none | 19 | 19 | - | MD 3.58 higher (5.2 lower to 12.36 higher) | ⊕⊕○○ Low | CRITICAL |
| Psychological distress - Depression (HADS-D, BDI, Faces scale, PANAS - negative affect [different scale ranges], lower values are better, final values) at <6 months (follow-up: mean 8 weeks; assessed with: HADS-D, BDI, Faces scale, PANAS - negative affect) | | | | | | | | | | | | |
| 5 | randomised trials | very serious ^d | not serious | not serious | not serious | none | 65 | 59 | - | SMD 0.03 SD lower (0.39 lower to 0.32 higher) | ⊕⊕○○ Low | CRITICAL |
| Psychological distress - Depression (Center for Epidemiologic Studies Depression Scale, 0-60, lower values are better, change score) at <6 months (follow-up: 8 weeks; assessed with: Center for Epidemiologic Studies Depression Scale; Scale from: 0 to 60) | | | | | | | | | | | | |
| 1 | randomised trials | very serious ^c | not serious | not serious | very serious ^b | none | 15 | 15 | - | MD 3.21 higher (6.56 lower to 12.98 higher) | ⊕○○○ Very low | CRITICAL |
| Psychological distress - Anxiety (HADS-A, BAI [different scale ranges], lower values are better, final values) at <6 months (follow-up: mean 6 weeks; assessed with: HADS-A, BAI) | | | | | | | | | | | | |
| 2 | randomised trials | very serious ^a | not serious | not serious | serious ^b | none | 28 | 28 | - | SMD 0.18 SD lower (0.7 lower to 0.35 higher) | ⊕○○○ Very low | CRITICAL |

| Certainty assessment | | | | | | | No of patients | | Effect | | Certainty | Importance |
|----------------------|--------------|--------------|---------------|--------------|-------------|----------------------|---|--------------|-------------------|-------------------|-----------|------------|
| No of studies | Study design | Risk of bias | Inconsistency | Indirectness | Imprecision | Other considerations | music therapy delivered by trained music therapists | no treatment | Relative (95% CI) | Absolute (95% CI) | | |

Psychological distress (PANAS - positive affect, 10-50, higher values are better, final value) at <6 months (follow-up: 4 months; assessed with: PANAS - positive affect; Scale from: 10 to 50)

| | | | | | | | | | | | | |
|---|-------------------|-------------|-------------|-------------|----------------------|------|----|----|---|--|------------------|----------|
| 1 | randomised trials | not serious | not serious | not serious | serious ^b | none | 14 | 14 | - | MD 4.15 higher (2.01 lower to 10.31 higher) | ⊕⊕⊕○ Moderate | CRITICAL |
|---|-------------------|-------------|-------------|-------------|----------------------|------|----|----|---|--|------------------|----------|

Stroke-specific Patient-Reported Outcome Measures (Stroke specific quality of life, 49-245, higher values are better, final value) at <6 months (follow-up: 6 weeks; assessed with: Stroke specific quality of life; Scale from: 49 to 245)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|---|------------------|----------|
| 1 | randomised trials | very serious ^a | not serious | not serious | serious ^b | none | 10 | 10 | - | MD 24.5 higher (7.36 higher to 41.64 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|---|------------------|----------|

Stroke-specific Patient-Reported Outcome Measures (SIS physical strength, 0-100, higher values are better, final value) at <6 months (follow-up: 6 weeks; assessed with: SIS physical strength; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|----------------------|----------------------|------|----|----|---|---|------------------|----------|
| 1 | randomised trials | very serious ^f | not serious | serious ^a | serious ^b | none | 13 | 12 | - | MD 10.7 lower (23.83 lower to 2.43 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|----------------------|----------------------|------|----|----|---|---|------------------|----------|

Stroke-specific Patient-Reported Outcome Measures (SIS activities, 0-100, higher values are better, final value) at <6 months (follow-up: 6 weeks; assessed with: SIS activities; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|----------------------|----------------------|------|----|----|---|--|------------------|----------|
| 1 | randomised trials | very serious ^f | not serious | serious ^a | serious ^b | none | 13 | 12 | - | MD 6.4 lower (20.13 lower to 7.33 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|----------------------|----------------------|------|----|----|---|--|------------------|----------|

Stroke-specific Patient-Reported Outcome Measures (SIS hand use, 0-100, higher values are better, final value) at <6 months (follow-up: 6 weeks; assessed with: SIS hand use; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|----------------------|----------------------|------|----|----|---|---|------------------|----------|
| 1 | randomised trials | very serious ^f | not serious | serious ^a | serious ^b | none | 13 | 12 | - | MD 12 lower (33.74 lower to 9.74 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|----------------------|----------------------|------|----|----|---|---|------------------|----------|

Stroke-specific Patient-Reported Outcome Measures (SIS mobility, 0-100, higher values are better, final value) at <6 months (follow-up: mean 11 weeks; assessed with: SIS mobility; Scale from: 0 to 100)

| Certainty assessment | | | | | | | No of patients | | Effect | | Certainty | Importance |
|----------------------|-------------------|--------------|---------------|--------------|----------------------|----------------------|---|--------------|-------------------|--|------------------|------------|
| No of studies | Study design | Risk of bias | Inconsistency | Indirectness | Imprecision | Other considerations | music therapy delivered by trained music therapists | no treatment | Relative (95% CI) | Absolute (95% CI) | | |
| 2 | randomised trials | not serious | not serious | not serious | serious ^b | none | 27 | 26 | - | MD 4.96 lower (13.36 lower to 3.44 higher) | ⊕⊕⊕○ Moderate | CRITICAL |

Stroke-specific Patient-Reported Outcome Measures (SIS communication, 0-100, higher values are better, final value) at <6 months (follow-up: mean 11 weeks; assessed with: SIS communication; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|-------------|-------------|-------------|----------------------|------|----|----|---|---|------------------|----------|
| 1 | randomised trials | not serious | not serious | not serious | serious ^b | none | 27 | 26 | - | MD 1.89 lower (8.05 lower to 4.27 higher) | ⊕⊕⊕○ Moderate | CRITICAL |
|---|-------------------|-------------|-------------|-------------|----------------------|------|----|----|---|---|------------------|----------|

Stroke-specific Patient-Reported Outcome Measures (SIS memory, 0-100, higher values are better, final value) at <6 months (follow-up: mean 11 weeks; assessed with: SIS memory; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|----------------------|----------------------|------|----|----|---|---|------------------|----------|
| 1 | randomised trials | very serious ^f | not serious | serious ^a | serious ^b | none | 27 | 26 | - | MD 0.65 lower (9.93 lower to 8.64 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|----------------------|----------------------|------|----|----|---|---|------------------|----------|

Stroke-specific Patient-Reported Outcome Measures (SIS emotion, 0-100, higher values are better, final value) at <6 months (follow-up: mean 11 weeks; assessed with: SIS emotion; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|-------------|-------------|----------------------|----------------------|------|----|----|---|---|-------------|----------|
| 1 | randomised trials | not serious | not serious | serious ^a | serious ^b | none | 27 | 26 | - | MD 1.01 higher (7.7 lower to 9.72 higher) | ⊕⊕○○ Low | CRITICAL |
|---|-------------------|-------------|-------------|----------------------|----------------------|------|----|----|---|---|-------------|----------|

Stroke-specific Patient-Reported Outcome Measures (SIS social/participation, 0-100, higher values are better, final value) at <6 months (follow-up: mean 11 weeks; assessed with: SIS social; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|-------------|-------------|----------------------|----------------------|------|----|----|---|--|-------------|----------|
| 1 | randomised trials | not serious | not serious | serious ^a | serious ^b | none | 27 | 26 | - | MD 2.38 lower (12.84 lower to 8.08 higher) | ⊕⊕○○ Low | CRITICAL |
|---|-------------------|-------------|-------------|----------------------|----------------------|------|----|----|---|--|-------------|----------|

Stroke-specific Patient-Reported Outcome Measures (SIS recovery, 0-100, higher values are better, final value) at <6 months (follow-up: 6 weeks; assessed with: SIS recovery; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|----------------------|---------------------------|------|----|----|---|--|------------------|----------|
| 1 | randomised trials | very serious ^f | not serious | serious ^a | very serious ^b | none | 13 | 12 | - | MD 1.2 lower (13.66 lower to 11.26 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|----------------------|---------------------------|------|----|----|---|--|------------------|----------|

Wellbeing scores (WHO five item well-being index, 0-25, higher values are better, final value) at <6 months (follow-up: 6 weeks; assessed with: WHO five item well-being index; Scale from: 0 to 25)

| Certainty assessment | | | | | | | No of patients | | Effect | | Certainty | Importance |
|----------------------|-------------------|---------------------------|---------------|----------------------|----------------------|----------------------|---|--------------|-------------------|---|------------------|------------|
| No of studies | Study design | Risk of bias | Inconsistency | Indirectness | Imprecision | Other considerations | music therapy delivered by trained music therapists | no treatment | Relative (95% CI) | Absolute (95% CI) | | |
| 1 | randomised trials | very serious ^f | not serious | serious ^a | serious ^b | none | 13 | 12 | - | MD 2.4 higher (1.71 lower to 6.51 higher) | ⊕○○○ Very low | CRITICAL |

Participation in leisure activities/social groups (Sickness Impact Profile Social Interaction subscale, 0-102, lower values are better, final value) at <6 months (follow-up: 4 months; assessed with: Sickness Impact Profile Social Interaction subscale,; Scale from: 0 to 102)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|----------------------|-------------|------|----|---|---|---|------------------|----------|
| 1 | randomised trials | very serious ^a | not serious | serious ^b | not serious | none | 10 | 8 | - | MD 13.28 lower (19.7 lower to 6.86 lower) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|----------------------|-------------|------|----|---|---|---|------------------|----------|

Withdrawal due to adverse events at <6 months (follow-up: 7 weeks)

| | | | | | | | | | | | | |
|---|-------------------|----------------------|-------------|-------------|----------------------|------|-------------|-------------|-------------------------|---|-------------|----------|
| 4 | randomised trials | serious ⁱ | not serious | not serious | serious ⁱ | none | 0/43 (0.0%) | 0/41 (0.0%) | RD 0.00 (-0.09 to 0.09) | 0 fewer per 1,000 (from 90 fewer to 90 more) ^k | ⊕⊕○○ Low | CRITICAL |
|---|-------------------|----------------------|-------------|-------------|----------------------|------|-------------|-------------|-------------------------|---|-------------|----------|

CI: confidence interval; MD: mean difference; SMD: standardised mean difference

Explanations

- a. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias arising from the randomisation process and bias due to deviations from the intended interventions)
- b. Downgraded by 1 increment if the confidence interval crossed one MID or by 2 increments if the confidence interval crossed both MIDs
- c. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias arising from the randomisation process, bias due to deviations from the intended interventions, bias due to missing outcome data and bias in measurement of the outcome)
- d. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to a mixture of bias arising from the randomisation process, bias due to deviations from the intended interventions, bias due to missing outcome data and bias in measurement of the outcome)
- e. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to a mixture of bias arising from the randomisation process, bias due to deviations from the intended interventions and bias in measurement of the outcome)
- f. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias arising from the randomisation process and bias due to missing outcome data)
- g. Downgraded by 1 increment due to comparator indirectness (due to the comparator group in 1 study receiving an exercise intervention that may have been more intense than that received than the intervention group, however it was unclear as to whether this was the case from the information provided in the study)
- h. Downgraded by 1 increments due to population indirectness (10-20% of people in a study having had a traumatic brain injury rather than a stroke)

- i. Downgraded by 1 increment as the majority of the evidence was of high risk of bias (due to bias arising from the randomisation process)
- j. Downgraded by 1 to 2 increments for imprecision due to zero events and small sample size
- k. Absolute effect calculated by risk difference due to zero events in at least one arm of one study

F.3 Music interventions delivered by healthcare professionals compared to passive music listening

Table 18: Clinical evidence profile: music interventions delivered by healthcare professionals compared to passive music listening

| Certainty assessment | | | | | | | No of patients | | Effect | | Certainty | Importance |
|----------------------|--------------|--------------|---------------|--------------|-------------|----------------------|--|-------------------------|-------------------|-------------------|-----------|------------|
| No of studies | Study design | Risk of bias | Inconsistency | Indirectness | Imprecision | Other considerations | music intervention delivered by healthcare professionals | passive music listening | Relative (95% CI) | Absolute (95% CI) | | |

Withdrawal due to adverse events at <6 months (follow-up: 3 months)

| | | | | | | | | | | | | |
|---|-------------------|----------------------|-------------|-------------|---------------------------|------|--------------|-------------|----------------------------|---|------------------|----------|
| 1 | randomised trials | serious ^a | not serious | not serious | very serious ^b | none | 4/23 (17.4%) | 1/25 (4.0%) | RR 4.35 (0.52 to 36.11) | 134 more per 1,000 (from 19 fewer to 1,000 more) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|----------------------|-------------|-------------|---------------------------|------|--------------|-------------|----------------------------|---|------------------|----------|

Withdrawal due to adverse events at ≥6 months (follow-up: 6 months)

| | | | | | | | | | | | | |
|---|-------------------|----------------------|-------------|-------------|---------------------------|------|--------------|-------------|----------------------------|---|------------------|----------|
| 1 | randomised trials | serious ^a | not serious | not serious | very serious ^b | none | 4/23 (17.4%) | 1/25 (4.0%) | RR 4.35 (0.52 to 36.11) | 134 more per 1,000 (from 19 fewer to 1,000 more) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|----------------------|-------------|-------------|---------------------------|------|--------------|-------------|----------------------------|---|------------------|----------|

CI: confidence interval; RR: risk ratio

Explanations

- a. Downgraded by 1 increment as the majority of the evidence was of high risk of bias (due to bias due to missing outcome data)
- b. Downgraded by 1 increment if the confidence interval crossed one MID or by 2 increments if the confidence interval crossed both MIDs


F.4 Music interventions delivered by healthcare professionals compared to placebo music therapy

Table 19: Clinical evidence profile: music interventions delivered by healthcare professionals compared to placebo music therapy


| Certainty assessment | | | | | | | No of patients | | Effect | | Certainty | Importance |
|--|-------------------|---------------------------|---------------|--------------|----------------------|----------------------|--|-----------------------|-------------------|---|------------------|------------|
| No of studies | Study design | Risk of bias | Inconsistency | Indirectness | Imprecision | Other considerations | music intervention delivered by healthcare professionals | placebo music therapy | Relative (95% CI) | Absolute (95% CI) | | |
| Psychological distress - Depression (HADS-D, 0-42, higher values are better, mean difference) at <6 months (follow-up: 3 months; assessed with: HADS-D; Scale from: 0 to 42) | | | | | | | | | | | | |
| 1 | randomised trials | very serious ^a | not serious | not serious | serious ^b | none | 23 | 25 | - | MD 0.7 higher (1.65 lower to 3.05 higher) | ⊕○○○ Very low | CRITICAL |
| Psychological distress - Depression (HADS-D, 0-42, higher values are better, mean difference) at ≥6 months (follow-up: 6 months; assessed with: HADS-D; Scale from: 0 to 42) | | | | | | | | | | | | |
| 1 | randomised trials | very serious ^a | not serious | not serious | serious ^b | none | 23 | 25 | - | MD 1.02 higher (1.36 lower to 3.4 higher) | ⊕○○○ Very low | CRITICAL |
| Psychological distress - Anxiety (HADS-A, 0-42, higher values are better, mean difference) at <6 months (follow-up: 3 months; assessed with: HADS-A; Scale from: 0 to 42) | | | | | | | | | | | | |
| 1 | randomised trials | very serious ^a | not serious | not serious | serious ^b | none | 23 | 25 | - | MD 0.69 higher (1.47 lower to 2.85 higher) | ⊕○○○ Very low | CRITICAL |
| Psychological distress - Anxiety (HADS-A, 0-42, higher values are better, mean difference) at ≥6 months (follow-up: 6 months; assessed with: HADS-A; Scale from: 0 to 42) | | | | | | | | | | | | |
| 1 | randomised trials | very serious ^a | not serious | not serious | serious ^b | none | 23 | 25 | - | MD 2 higher (0.28 lower to 4.28 higher) | ⊕○○○ Very low | CRITICAL |

| Certainty assessment | | | | | | | № of patients | | Effect | | Certainty | Importance |
|----------------------|--------------|--------------|---------------|--------------|-------------|----------------------|--|-----------------------|-------------------|-------------------|-----------|------------|
| № of studies | Study design | Risk of bias | Inconsistency | Indirectness | Imprecision | Other considerations | music intervention delivered by healthcare professionals | placebo music therapy | Relative (95% CI) | Absolute (95% CI) | | |


Participation in leisure activities/social group scores (Mayo-Portland Adaptability Inventory 4 participation, 0-30, higher values are better, mean difference) at <6 months (follow-up: 3 months; assessed with: Mayo-Portland Adaptability Inventory 4 participation; Scale from: 0 to 30)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|-------------|------|----|----|---|---|--|----------|
| 1 | randomised trials | very serious ^a | not serious | not serious | not serious | none | 23 | 25 | - | MD 1.72 higher (11.75 lower to 15.19 higher) |  Low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|-------------|------|----|----|---|---|--|----------|

Withdrawal due to adverse events at <6 months (follow-up: 3 months)

| | | | | | | | | | | | | |
|---|-------------------|----------------------|-------------|-------------|---------------------------|------|--------------|-------------|----------------------------|---|---|----------|
| 1 | randomised trials | serious ^c | not serious | not serious | very serious ^b | none | 4/23 (17.4%) | 1/25 (4.0%) | RR 4.35 (0.52 to 36.11) | 134 more per 1,000 (from 19 fewer to 1,000 more) |  Very low | CRITICAL |
|---|-------------------|----------------------|-------------|-------------|---------------------------|------|--------------|-------------|----------------------------|---|---|----------|

Withdrawal due to adverse events at ≥6 months (follow-up: 6 months)

| | | | | | | | | | | | | |
|---|-------------------|----------------------|-------------|-------------|---------------------------|------|--------------|-------------|----------------------------|---|---|----------|
| 1 | randomised trials | serious ^c | not serious | not serious | very serious ^b | none | 4/23 (17.4%) | 1/25 (4.0%) | RR 4.35 (0.52 to 36.11) | 134 more per 1,000 (from 19 fewer to 1,000 more) |  Very low | CRITICAL |
|---|-------------------|----------------------|-------------|-------------|---------------------------|------|--------------|-------------|----------------------------|---|---|----------|

CI: confidence interval; MD: mean difference; RR: risk ratio

Explanations

- a. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias due to deviations from the intended interventions and bias due to missing outcome data)
- b. Downgraded by 1 increment if the confidence interval crossed one MID or by 2 increments if the confidence interval crossed both MIDs
- c. Downgraded by 1 increment as the majority of the evidence was of high risk of bias (due to bias due to missing outcome data)

F.5 Music interventions delivered by healthcare professionals compared to no treatment

Table 20: Clinical evidence profile: music interventions delivered by healthcare professionals compared to no treatment

| Certainty assessment | | | | | | | No of patients | | Effect | | Certainty | Importance |
|---|-------------------|---------------------------|---------------|--------------|---------------------------|----------------------|--|--------------|-------------------|---|------------------|------------|
| No of studies | Study design | Risk of bias | Inconsistency | Indirectness | Imprecision | Other considerations | music intervention delivered by healthcare professionals | no treatment | Relative (95% CI) | Absolute (95% CI) | | |
| Person/participant generic health-related quality of life (SF-36 physical function, 0-100, higher values are better, final value) at <6 months (follow-up: 4 weeks; assessed with: SF-36 physical function; Scale from: 0 to 100) | | | | | | | | | | | | |
| 1 | randomised trials | very serious ^a | not serious | not serious | very serious ^b | none | 20 | 17 | - | MD 10.3 higher (4.31 lower to 24.91 higher) | ⊕○○○ Very low | CRITICAL |
| Person/participant generic health-related quality of life (SF-36 bodily pain, 0-100, higher values are better, final value) at <6 months (follow-up: 4 weeks; assessed with: SF-36 bodily pain; Scale from: 0 to 100) | | | | | | | | | | | | |
| 1 | randomised trials | very serious ^a | not serious | not serious | very serious ^b | none | 20 | 17 | - | MD 6.3 higher (13.48 lower to 26.08 higher) | ⊕○○○ Very low | CRITICAL |
| Person/participant generic health-related quality of life (SF-36 role physical, 0-100, higher values are better, final value) at <6 months (follow-up: 4 weeks; assessed with: SF-36 role physical; Scale from: 0 to 100) | | | | | | | | | | | | |
| 1 | randomised trials | very serious ^a | not serious | not serious | very serious ^b | none | 20 | 17 | - | MD 6.1 lower (29.41 lower to 17.21 higher) | ⊕○○○ Very low | CRITICAL |
| Person/participant generic health-related quality of life (SF-36 vitality, 0-100, higher values are better, final value) at <6 months (follow-up: 4 weeks; assessed with: SF-36 vitality; Scale from: 0 to 100) | | | | | | | | | | | | |
| 1 | randomised trials | very serious ^a | not serious | not serious | very serious ^b | none | 20 | 17 | - | MD 6.8 higher (8.74 lower to 22.34 higher) | ⊕○○○ Very low | CRITICAL |
| Person/participant generic health-related quality of life (SF-36 general health, 0-100, higher values are better, final value) at <6 months (follow-up: 4 weeks; assessed with: SF-36 general health; Scale from: 0 to 100) | | | | | | | | | | | | |
| 1 | randomised trials | very serious ^a | not serious | not serious | very serious ^b | none | 20 | 17 | - | MD 3.9 higher (9.57 lower to 17.37 higher) | ⊕○○○ Very low | CRITICAL |

| Certainty assessment | | | | | | | № of patients | | Effect | | Certainty | Importance |
|----------------------|--------------|--------------|---------------|--------------|-------------|----------------------|--|--------------|-------------------|-------------------|-----------|------------|
| № of studies | Study design | Risk of bias | Inconsistency | Indirectness | Imprecision | Other considerations | music intervention delivered by healthcare professionals | no treatment | Relative (95% CI) | Absolute (95% CI) | | |

Person/participant generic health-related quality of life (SF-36 role emotional, 0-100, higher values are better, final value) at <6 months (follow-up: 4 weeks; assessed with: SF-36 role emotional; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|---------------------------|------|----|----|---|---|------------------|----------|
| 1 | randomised trials | very serious ^a | not serious | not serious | very serious ^b | none | 20 | 17 | - | MD 5.6 higher (21.41 lower to 32.61 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|---------------------------|------|----|----|---|---|------------------|----------|

Person/participant generic health-related quality of life (SF-36 mental health, 0-100, higher values are better, final value) at <6 months (follow-up: 4 weeks; assessed with: SF-36 mental health; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|---------------------------|------|----|----|---|--|------------------|----------|
| 1 | randomised trials | very serious ^a | not serious | not serious | very serious ^b | none | 20 | 17 | - | MD 4.8 higher (9.05 lower to 18.65 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|---------------------------|------|----|----|---|--|------------------|----------|

Person/participant generic health-related quality of life (SF-36 social function, 0-100, higher values are better, final value) at <6 months (follow-up: 4 weeks; assessed with: SF-36 social function; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|---------------------------|------|----|----|---|---|------------------|----------|
| 1 | randomised trials | very serious ^c | not serious | not serious | very serious ^b | none | 20 | 17 | - | MD 11 higher (7.93 lower to 29.93 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|---------------------------|------|----|----|---|---|------------------|----------|

Person/participant health-related quality of life (McGill Quality of Life, 0-10, higher values are better, final value) at <6 months (follow-up: 2 months; assessed with: McGill Quality of Life; Scale from: 0 to 10)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|-------------|------|----|----|---|---|-------------|----------|
| 1 | randomised trials | very serious ^d | not serious | not serious | not serious | none | 13 | 16 | - | MD 0.49 lower (1.65 lower to 0.67 higher) | ⊕⊕○○ Low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|-------------|------|----|----|---|---|-------------|----------|

Activities of daily living (Canadian Occupational Performance Measure, University of Maryland Arm Questionnaire for Stroke, Activities of daily living score [different scale ranges], higher values are better, change scores) at <6 months (follow-up: 6 weeks; assessed with: Canadian Occupational Performance Measure, University of Maryland Arm Questionnaire for Stroke, Activities of daily living score)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|----------------------|-------------|----------------------|------|----|----|---|--|------------------|----------|
| 3 | randomised trials | very serious ^a | serious ^f | not serious | serious ^b | none | 45 | 44 | - | SMD 0.64 SD higher (0.15 lower to 1.42 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|----------------------|-------------|----------------------|------|----|----|---|--|------------------|----------|

Activities of daily living (Barthel index, 0-100, higher values are better, final value) at <6 months (follow-up: 4 weeks; Scale from: 0 to 100)

| Certainty assessment | | | | | | | No of patients | | Effect | | Certainty | Importance |
|----------------------|-------------------|---------------------------|---------------|--------------|-------------|----------------------|--|--------------|-------------------|---|-------------|------------|
| No of studies | Study design | Risk of bias | Inconsistency | Indirectness | Imprecision | Other considerations | music intervention delivered by healthcare professionals | no treatment | Relative (95% CI) | Absolute (95% CI) | | |
| 1 | randomised trials | very serious ^a | not serious | not serious | not serious | none | 15 | 15 | - | MD 10.66 higher (4.85 higher to 16.47 higher) | ⊕⊕○○ Low | CRITICAL |

Psychological distress - Depression (Hamilton's Depression Scale-17, 0-56, lower values are better, change score) at <6 months (follow-up: 17 days; assessed with: Hamilton's Depression Scale-17; Scale from: 0 to 56)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|---|------------------|----------|
| 1 | randomised trials | very serious ^a | not serious | not serious | serious ^b | none | 30 | 30 | - | MD 1.3 lower (2.22 lower to 0.38 lower) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|---|------------------|----------|

Psychological distress - Depression (BDI, Hamilton Depression Rating Scale-24, profile of mood states [different scale ranges], lower values are better, final values) at <6 months (follow-up: mean 6 weeks; assessed with: BDI, Hamilton Depression Rating Scale-24, profile of mood states)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|--|------------------|----------|
| 3 | randomised trials | very serious ^b | not serious | not serious | serious ^b | none | 68 | 67 | - | SMD 0.68 SD lower (1.03 lower to 0.33 lower) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|--|------------------|----------|

Psychological distress (PANAS positive affect, 10-50, higher values are better, change score) at <6 months (follow-up: 4 weeks; assessed with: PANAS positive affect; Scale from: 10 to 50)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|---|------------------|----------|
| 1 | randomised trials | very serious ^a | not serious | not serious | serious ^b | none | 20 | 17 | - | MD 3.4 higher (1.95 lower to 8.75 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|---|------------------|----------|

Stroke-specific Patient-Reported Outcome Measures (SIS [different scale ranges], higher values are better, change scores) at <6 months (follow-up: 8 weeks)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|----------------------|-------------|----------------------|------|----|----|---|--|------------------|----------|
| 2 | randomised trials | very serious ^b | serious ^f | not serious | serious ^b | none | 43 | 46 | - | SMD 1.23 SD lower (2.57 lower to 0.1 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|----------------------|-------------|----------------------|------|----|----|---|--|------------------|----------|

Stroke-specific Patient-Reported Outcome Measures (SIS strength subscale, 0-100, higher values are better, change score) at <6 months (follow-up: 12 weeks; assessed with: SIS strength subscale; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|---|------------------|----------|
| 1 | randomised trials | very serious ^f | not serious | not serious | serious ^b | none | 17 | 15 | - | MD 12.8 lower (21.9 lower to 3.7 lower) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|---|------------------|----------|

| Certainty assessment | | | | | | | No of patients | | Effect | | Certainty | Importance |
|----------------------|--------------|--------------|---------------|--------------|-------------|----------------------|--|--------------|-------------------|-------------------|-----------|------------|
| No of studies | Study design | Risk of bias | Inconsistency | Indirectness | Imprecision | Other considerations | music intervention delivered by healthcare professionals | no treatment | Relative (95% CI) | Absolute (95% CI) | | |

Stroke-specific Patient-Reported Outcome Measures (SIS memory subscale, 0-100, higher values are better, change score) at <6 months (follow-up: 12 weeks; assessed with: SIS memory subscale; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|---------------------------|------|----|----|---|---|------------------|----------|
| 1 | randomised trials | very serious ⁱ | not serious | not serious | very serious ^p | none | 17 | 15 | - | MD 1.1 higher (5.07 lower to 7.27 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|---------------------------|------|----|----|---|---|------------------|----------|

Stroke-specific Patient-Reported Outcome Measures (SIS emotion subscale, 0-100, higher values are better, change score) at <6 months (follow-up: 12 weeks; assessed with: SIS emotion subscale; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|---------------------------|------|----|----|---|--|------------------|----------|
| 1 | randomised trials | very serious ⁱ | not serious | not serious | very serious ^p | none | 17 | 15 | - | MD 1.9 higher (8.54 lower to 12.34 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|---------------------------|------|----|----|---|--|------------------|----------|

Stroke-specific Patient-Reported Outcome Measures (SIS communication subscale, 0-100, higher values are better, change score) at <6 months (follow-up: 12 weeks; assessed with: SIS communication subscale; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|---------------------------|------|----|----|---|--|------------------|----------|
| 1 | randomised trials | very serious ⁱ | not serious | not serious | very serious ^p | none | 17 | 15 | - | MD 0.8 lower (9.4 lower to 7.8 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|---------------------------|------|----|----|---|--|------------------|----------|

Stroke-specific Patient-Reported Outcome Measures (SIS ADL subscale, 0-100, higher values are better, change score) at <6 months (follow-up: 12 weeks; assessed with: SIS ADL subscale; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|--|------------------|----------|
| 1 | randomised trials | very serious ⁱ | not serious | not serious | serious ^p | none | 17 | 15 | - | MD 2.8 higher (6.28 lower to 11.88 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|--|------------------|----------|

Stroke-specific Patient-Reported Outcome Measures (SIS mobility subscale, 0-100, higher values are better, change score) at <6 months (follow-up: 12 weeks; assessed with: SIS mobility subscale; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|---------------------------|------|----|----|---|--------------------------------------|------------------|----------|
| 1 | randomised trials | very serious ⁱ | not serious | not serious | very serious ^p | none | 17 | 15 | - | MD 1.3 lower (7.6 lower to 5 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|---------------------------|------|----|----|---|--------------------------------------|------------------|----------|

Stroke-specific Patient-Reported Outcome Measures (SIS hand function subscale, 0-100, higher values are better, change score) at <6 months (follow-up: 12 weeks; assessed with: SIS hand function subscale; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|---------------------------|------|----|----|---|---|------------------|----------|
| 1 | randomised trials | very serious ⁱ | not serious | not serious | very serious ^p | none | 17 | 15 | - | MD 1.8 higher (13.96 lower to 17.56 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|---------------------------|------|----|----|---|---|------------------|----------|

| Certainty assessment | | | | | | | No of patients | | Effect | | Certainty | Importance |
|----------------------|--------------|--------------|---------------|--------------|-------------|----------------------|--|--------------|-------------------|-------------------|-----------|------------|
| No of studies | Study design | Risk of bias | Inconsistency | Indirectness | Imprecision | Other considerations | music intervention delivered by healthcare professionals | no treatment | Relative (95% CI) | Absolute (95% CI) | | |

Stroke-specific Patient-Reported Outcome Measures (SIS social participation subscale, 0-100, higher values are better, change score) at <6 months (follow-up: 12 weeks; assessed with: SIS social participation subscale; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|---------------------------|------|----|----|---|---|------------------|----------|
| 1 | randomised trials | very serious ⁱ | not serious | not serious | very serious ^o | none | 17 | 15 | - | MD 3.7 higher (10.15 lower to 17.55 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|---------------------------|------|----|----|---|---|------------------|----------|

Stroke-specific Patient-Reported Outcome Measures (Stroke Specific Quality of Life [different scale ranges], higher values are better, final values) at <6 months (follow-up: mean 6 weeks; assessed with: Stroke Specific Quality of Life)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|----------------------|-------------|----------------------|------|----|----|---|---|------------------|----------|
| 2 | randomised trials | very serious ^h | serious ^f | not serious | serious ^o | none | 36 | 35 | - | SMD 0.4 SD higher (0.28 lower to 1.09 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|----------------------|-------------|----------------------|------|----|----|---|---|------------------|----------|

Withdrawal due to adverse events at <6 months (follow-up: mean 7 weeks)

| | | | | | | | | | | | | |
|---|-------------------|----------------------|----------------------|-------------|---------------------------|------|---------------|---------------|-------------------------|--|------------------|----------|
| 7 | randomised trials | serious ⁱ | serious ^k | not serious | very serious ^l | none | 15/178 (8.4%) | 14/177 (7.9%) | RD 0.01 (-0.05 to 0.07) | 10 fewer per 1,000 (from 70 fewer to 50 more) ^m | ⊕○○○ Very low | CRITICAL |
|---|-------------------|----------------------|----------------------|-------------|---------------------------|------|---------------|---------------|-------------------------|--|------------------|----------|

CI: confidence interval; MD: mean difference; SMD: standardised mean difference

Explanations

- a. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias arising from the randomisation process, bias due to deviations from the intended interventions and bias in selection of the reported result)
- b. Downgraded by 1 increment if the confidence interval crossed one MID or by 2 increments if the confidence interval crossed both MIDs
- c. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias due to deviations from the intended interventions and bias in selection of the reported result)
- d. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias due to deviations from the intended interventions and bias due to missing outcome data)
- e. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to a mixture of bias arising from the randomisation process, bias due to deviations from the intended interventions and bias in measurement of the outcome)
- f. Downgraded by 1 or 2 increments because heterogeneity, unexplained by subgroup analysis
- g. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias arising from the randomisation process and bias due to deviations from the intended interventions)

- h. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to a mixture of bias arising from the randomisation process, bias due to deviations from the intended interventions, bias due to missing outcome data, bias in measurement of the outcome and bias in selection of the reported result)
- i. Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias due to deviations from the intended interventions and bias due to missing outcome data)
- j. Downgraded by 1 increment as the majority of the evidence was of high risk of bias (due to a mixture of bias arising from the randomisation process, bias due to missing outcome data, bias in measurement of the outcome and bias in selection of the reported result)
- k. Downgraded for heterogeneity due to conflicting number of events in different studies (zero events in one or more studies)
- l. Downgraded by 1 to 2 increments for imprecision due to zero events and small sample size
- m. Absolute effect calculated by risk difference due to zero events in at least one arm of one study

F.6 Music interventions delivered by non-healthcare professionals compared to no treatment

Table 21: Clinical evidence profile: music interventions delivered by non-healthcare professionals compared to no treatment

| Certainty assessment | | | | | | | No of patients | | Effect | | Certainty | Importance |
|--|-------------------|---------------------------|---------------|--------------|---------------------------|----------------------|--|--------------|-------------------|--|------------------|------------|
| No of studies | Study design | Risk of bias | Inconsistency | Indirectness | Imprecision | Other considerations | music intervention delivered by non-healthcare professionals | no treatment | Relative (95% CI) | Absolute (95% CI) | | |
| Person/participant generic health-related quality of life (EQ-5D 5L, -0.11-1, higher values are better, final value) at <6 months (follow-up: 3 months; assessed with: EQ-5D 5L; Scale from: -0.11 to 1) | | | | | | | | | | | | |
| 1 | randomised trials | serious ^a | not serious | not serious | very serious ^b | none | 17 | 19 | - | MD 0.04 lower (0.21 lower to 0.12 higher) | ⊕○○○ Very low | CRITICAL |
| Person/participant generic health-related quality of life (EQ-5D 5L, -0.11-1, higher values are better, final value) at ≥6 months (follow-up: 6 months; assessed with: EQ-5D 5L; Scale from: -0.11 to 1) | | | | | | | | | | | | |
| 1 | randomised trials | very serious ^c | not serious | not serious | very serious ^b | none | 18 | 16 | - | MD 0.1 higher (0.06 lower to 0.26 higher) | ⊕○○○ Very low | CRITICAL |
| Carer generic health-related quality of life (CarerQoL-7D, 0-14, higher values are better, final value) at ≥6 months (follow-up: 6 months; assessed with: CarerQoL-7D; Scale from: 0 to 14) | | | | | | | | | | | | |
| 1 | randomised trials | very serious ^c | not serious | not serious | very serious ^b | none | 18 | 16 | - | MD 0 (1.97 lower to 1.97 higher) | ⊕○○○ Very low | CRITICAL |

| Certainty assessment | | | | | | | № of patients | | Effect | | Certainty | Importance |
|----------------------|--------------|--------------|---------------|--------------|-------------|----------------------|--|--------------|-------------------|-------------------|-----------|------------|
| № of studies | Study design | Risk of bias | Inconsistency | Indirectness | Imprecision | Other considerations | music intervention delivered by non-healthcare professionals | no treatment | Relative (95% CI) | Absolute (95% CI) | | |

Stroke-specific Patient-Reported Outcome Measures (Stroke and Aphasia Quality of Life, 1-5, higher values are better, final value) at <6 months (follow-up: 3 months; assessed with: Stroke and Aphasia Quality of Life; Scale from: 1 to 5)

| | | | | | | | | | | | | |
|---|-------------------|----------------------|-------------|-------------|---------------------------|------|----|----|---|-------------------------------------|------------------|----------|
| 1 | randomised trials | serious ^a | not serious | not serious | very serious ^b | none | 17 | 19 | - | MD 0 (0.49 lower to 0.49 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|----------------------|-------------|-------------|---------------------------|------|----|----|---|-------------------------------------|------------------|----------|

Stroke-specific Patient-Reported Outcome Measures (Stroke and Aphasia Quality of Life, 1-5, higher values are better, final value) at ≥6 months (follow-up: 6 months; assessed with: Stroke and Aphasia Quality of Life; Scale from: 1 to 5)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|--|------------------|----------|
| 1 | randomised trials | very serious ^c | not serious | not serious | serious ^b | none | 18 | 16 | - | MD 0.3 higher (0.2 lower to 0.8 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|--|------------------|----------|

Wellbeing scores (ICEpop CAPability measure for adults, 0-1, higher values are better, final value) at <6 months (follow-up: 3 months; assessed with: ICEpop CAPability measure for adults; Scale from: 0 to 1)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|---------------------------|------|----|----|---|--|------------------|----------|
| 1 | randomised trials | very serious ^d | not serious | not serious | very serious ^b | none | 17 | 19 | - | MD 0.07 higher (0.3 lower to 0.44 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|---------------------------|------|----|----|---|--|------------------|----------|


Wellbeing scores (ICEpop CAPability measure for adults, 0-1, higher values are better, final value) at ≥6 months (follow-up: 6 months; assessed with: ICEpop CAPability measure for adults; Scale from: 0 to 1)

| | | | | | | | | | | | | |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|---|------------------|----------|
| 1 | randomised trials | very serious ^d | not serious | not serious | serious ^b | none | 18 | 16 | - | MD 0.04 higher (0.06 lower to 0.13 higher) | ⊕○○○ Very low | CRITICAL |
|---|-------------------|---------------------------|-------------|-------------|----------------------|------|----|----|---|---|------------------|----------|


Participation in leisure activities/social groups scores (modified Reintegration to Normal Living Index, 0-100, higher values are better, final value) at <6 months (follow-up: 3 months; assessed with: modified Reintegration to Normal Living Index; Scale from: 0 to 100)

| | | | | | | | | | | | | |
|---|-------------------|----------------------|-------------|-------------|----------------------|------|----|----|---|--|-------------|----------|
| 1 | randomised trials | serious ^a | not serious | not serious | serious ^b | none | 17 | 19 | - | MD 1 higher (3.39 lower to 5.39 higher) | ⊕⊕○○ Low | CRITICAL |
|---|-------------------|----------------------|-------------|-------------|----------------------|------|----|----|---|--|-------------|----------|

Participation in leisure activities/social groups scores (modified Reintegration to Normal Living Index, 0-100, higher values are better, final value) at ≥6 months (follow-up: 6 months; assessed with: modified Reintegration to Normal Living Index; Scale from: 0 to 100)

| Certainty assessment | | | | | | | No of patients | | Effect | | Certainty | Importance |
|----------------------|-------------------|---------------------------|---------------|--------------|----------------------|----------------------|--|--------------|-------------------|---|---|------------|
| No of studies | Study design | Risk of bias | Inconsistency | Indirectness | Imprecision | Other considerations | music intervention delivered by non-healthcare professionals | no treatment | Relative (95% CI) | Absolute (95% CI) | | |
| 1 | randomised trials | very serious ^c | not serious | not serious | serious ^b | none | 18 | 16 | - | MD 1.8 higher (2.86 lower to 6.46 higher) |  Very low | CRITICAL |

Withdrawal due to adverse events at <6 months (follow-up: 3 months)

| | | | | | | | | | | | | |
|---|-------------------|-------------|-------------|-------------|---------------------------|------|--------------|-------------|--------------------------|---|--|----------|
| 1 | randomised trials | not serious | not serious | not serious | very serious ^b | none | 2/20 (10.0%) | 0/21 (0.0%) | OR 8.19 (0.49 to 135.71) | 100 more per 1,000 (from 50 fewer to 250 more) ^e |  Low | CRITICAL |
|---|-------------------|-------------|-------------|-------------|---------------------------|------|--------------|-------------|--------------------------|---|--|----------|

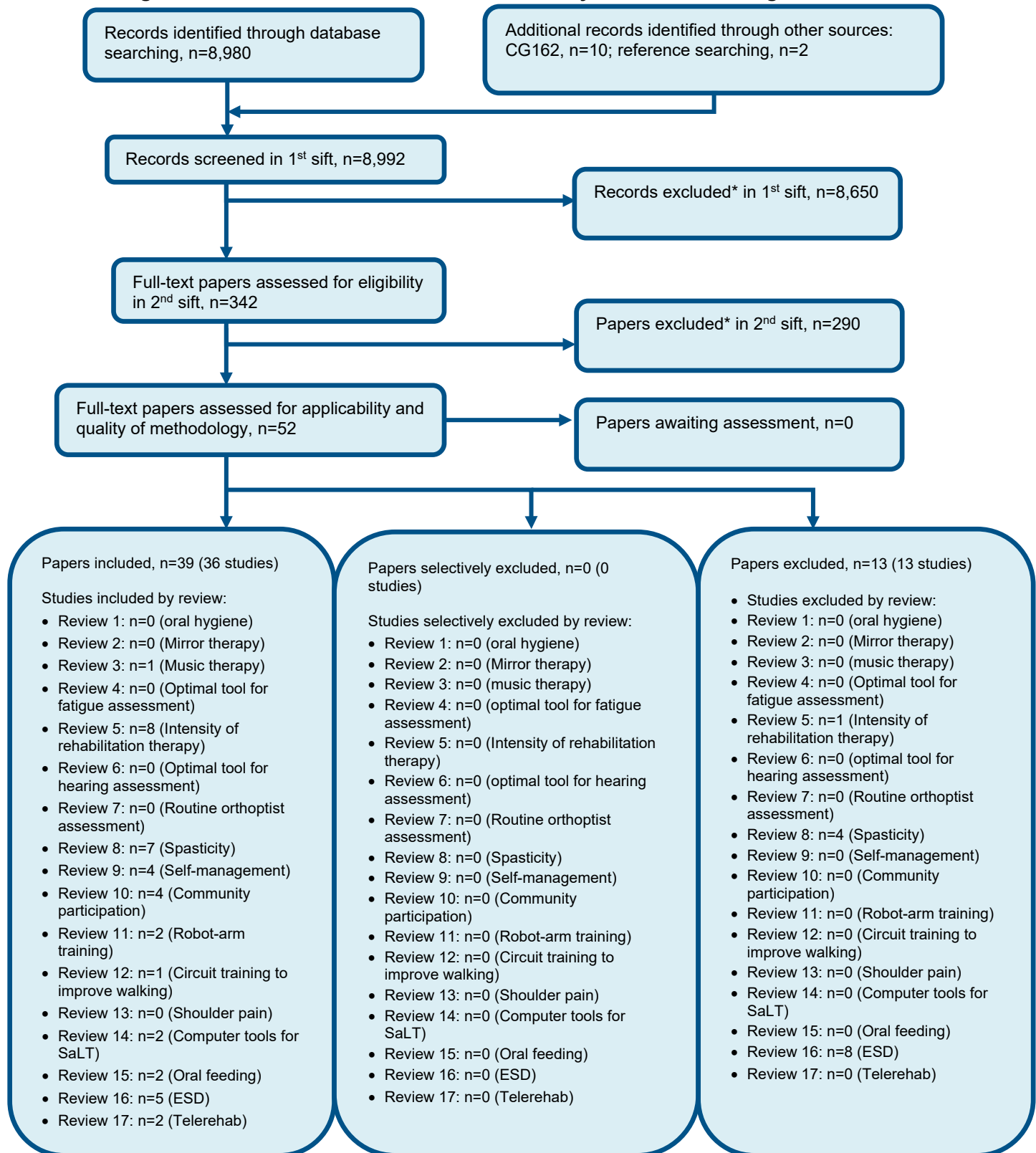
CI: confidence interval; MD: mean difference; OR: odds ratio

Explanations

- Downgraded by 1 increment as the majority of the evidence was of high risk of bias (due to bias due to deviations from the intended interventions)
- Downgraded by 1 increment if the confidence interval crossed one MID or by 2 increments if the confidence interval crossed both MIDs
- Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias due to deviations from the intended interventions and bias due to missing outcome data)
- Downgraded by 2 increments as the majority of the evidence was of very high risk of bias (due to bias arising from the randomisation process, bias due to deviations from the intended interventions and bias due to missing outcome data)
- Absolute effect calculated by risk difference due to zero events in at least one arm of one study

Appendix G – Economic evidence study selection

Figure 22: Flow chart of health economic study selection for the guideline



* Non-relevant population, intervention, comparison, design or setting; non-English language

Appendix H – Economic evidence tables

| Study | Tarrant et al. 2021 ²⁰ | | | |
|---|---|--|---|--|
| Study details | Population & interventions | Costs | Health outcomes | Cost effectiveness |
| <p>Economic analysis: CUA (health outcome: QALYs)</p> <p>Study design: Within trial analysis (RCT – same paper) (note the study aims to investigate acceptability and feasibility of a definitive RCT that would include full cost effectiveness assessment.)</p> <p>Approach to analysis: A micro-costing approach was adopted to estimate the intervention costs associated with SPA. Other healthcare resource use was collected but not included in cost analysis.</p> <p>Perspective: UK. It is not stated that an NHS and PSS perspective is taken however, the costs included are all considered relevant if</p> | <p>Population: Adults with post-stroke aphasia</p> <p>Patient characteristics N=41 Mean age: 66.5 years (SD:10.3) Male: 61%</p> <p>Intervention 1: Control group (no treatment)</p> <p>Intervention 2: Singing groups for people with aphasia (SPA), sessions held for 90 minutes, once a week for 10 weeks led by a music facilitator and assisted by an individual with poststroke aphasia.</p> | <p>Total costs (mean per patient): Intervention 1: £0 Intervention 2: £399 Incremental (2–1): £399^(a) (95% CI: NR; p=NR)</p> <p>Currency & cost year: 2019 UK pounds</p> <p>Cost components incorporated: Training costs, staff time during training and delivery of the intervention, travel costs for facilitators and singing champions, course materials (song books, percussion instruments, badges and flip charts), venue costs and refreshments.</p> <p>Note that other healthcare resource use was collected but not included in cost analysis (authors note the objective of the study was to assess feasibility of collection).</p> | <p>From clinical review (2 vs 1) – same paper:</p> <p>EQ-5D-5L scores ^(b) 3-month follow-up: -0.04 (95% CI: -0.21, 0.12) 6-month follow-up: 0.10 (95% CI: -0.06, 0.26)</p> <p>QALY gained (2 vs 1): 0.05</p> <p>Carer generated health-related quality of life (CarerQoL-7D) ^(b) 3-month follow-up: NR 6-month follow-up: 0.00 (95% CI: -1.97, 1.97)</p> <p>Other outcomes were reported and can be seen in clinical evidence table.</p> | <p>ICER (Intervention 2 versus Intervention 1): £7,980 per QALY gained^(c)</p> <p>Probability Intervention 2 cost effective (£20K/30K threshold): NR</p> <p>Analysis of uncertainty: None.</p> |

| | | | |
|---|---|--|--|
| <p>the intervention is funded by the NHS.</p> <p>Follow-up: 6 months Discounting: Costs: n/a; Outcomes: n/a</p> | <p>Health resource use included primary care (GP and nurse), secondary care (hospital admissions and accident and emergency) and social care.</p> <p>The study did not present differences in other resource use as costs but these were higher for the intervention group so inclusion of such costs could impact results.</p> | | |
|---|---|--|--|

Data sources

Health outcomes: Within-RCT analysis (same paper, included in the clinical review) – UK setting. **Quality-of-life weights:** QALYs were not reported but these were calculated for this review using EQ-5D-5L scores collected in the study (UK population valuation tariff). **Cost sources:** Within-trial analysis – micro-costing from 3 centres involved in trial. Total costs associated with providing therapy were calculated and then divided by number of participants to give cost per participant. Costs were inflated to 2019 using the Adult Personal Social Services pay and prices index inflation index reported in PSSRU unit costs report 2019. Other health and social care resource use was collected using self-reported data but not included in cost analysis (authors note the objective of the study was to assess feasibility of collection).

Comments

Source of funding: Stroke Association. **Limitations:** Mean EQ-5D-5L scores (UK tariff) at 6-months were used to calculate the cost per QALY gained for this review: the NICE reference case currently prefers EQ5D-3L. It is not stated that an NHS and PSS perspective is taken however, if the costs included are all considered relevant if the intervention is funded by the NHS. 6-month time horizon may not capture full health benefits of the intervention if these persist. Pilot feasibility RCT (n=41) that was not powered to test the effectiveness of the intervention or differences in healthcare resource use; the aim was to inform a future study where effectiveness and cost-effectiveness could be assessed. Within-trial analysis only reflects health outcomes and resource use from single trial; however, as this was the only trial included in the clinical review for music interventions delivered by non-healthcare professionals and so reflects the best currently available evidence. It is unclear if all relevant costs are included; interventions costs were included but other healthcare resource use was collected but not included; other healthcare resource used was numerically higher with the interventions but authors note the study was not powered to detect differences and did not present these as costs. Sensitivity analysis was not performed. **Other:** none.

Overall applicability: Partially applicable^(d) **Overall quality:** Potentially serious limitations^(e)

Abbreviations: 95% CI= 95% confidence interval; CUA= cost-utility analysis; ICER= incremental cost-effectiveness ratio; NA = not applicable; NR= not reported; QALYs= quality-adjusted life years.

(a) See Table 21 for cost breakdown.

(b) Mean difference taken from figures 60, 65 and 66 of guideline clinical review.

(c) Cost per QALY gained not reported but was estimated using 6-month EQ-5D-5L scores collected within the study and assuming no difference in mortality.

(d) Directly applicable / Partially applicable / Not applicable

(e) Minor limitations / Potentially serious limitations / Very serious limitations

Table 22: Course running costs based on three community-based cohorts in the South West of England²⁰

| Item | Total cost (£) |
|---|------------------|
| Salary | |
| Music facilitators | £2,725.00 |
| Music facilitator training | £743.75 |
| Singing champion* | £675.00 |
| Trainer | £310.08 |
| Administrator cohort set-up (42 hours) | £1,008.00 |
| Senior coordinator of sessions (15.25 hours) | £367.20 |
| Junior coordinator of sessions (20 hours) | £223.51 |
| Sub-total (training costs) | £1,095.29 |
| Sub-total (salary) | £6,662.35 |
| Course | |
| Facilitator travel | £0.00 |
| Singing champion travel | £425.45 |
| Course materials | £91.91 |
| Hospitality (venue costs) | £718.02 |
| Refreshments | £88.82 |
| Sub-total (course) | £1,324.20 |
| Total cost of intervention for (salary + course) | £7,986.55 |
| Cost per participant (including training) | £399.33 |
| Cost per participant (excluding training) | £344.56 |
| Cost per participant (excluding training and assistance during singing sessions which may be provided by another carer) | £325.62 |

**unit cost based on experience of within trial payment of £25 in 2017 prices and adjusted to 2019 prices*

Appendix I – Health economic model

New cost-effectiveness analysis was not conducted in this area.

Appendix J – Excluded studies

Clinical studies

Table 22: Studies excluded from the clinical review

| Study | Code [Reason] |
|---|---|
| Acalha, T., Mg, Suzuki, S. O. et al. (2010) Effects of the task oriented and auditory cues for chronic stroke patients. <i>Revista terapia manual</i> 8(39): 441-447 | - Full text paper not available |
| Ala-Ruona, E. (2009) Active music therapy for post-stroke recovery. | - Full text paper not available |
| Altenmuller, E., Marco-Pallares, J., Munte, T. F. et al. (2009) Neural reorganization underlies improvement in stroke-induced motor dysfunction by music-supported therapy. <i>Annals of the New York Academy of Sciences</i> 1169: 395-405 | - Study reported outcomes that were not included in the protocol |
| Anonymous (2008) Music in stroke rehabilitation. <i>Lancet</i> 371(9614): 698 | - Commentary only |
| Aravantinou-Fatorou, K. and Fotakopoulos, G. (2021) Efficacy of exercise rehabilitation program accompanied by experiential music for recovery of aphasia in single cerebrovascular accidents: a randomized controlled trial. <i>Irish Journal of Medical Science</i> 190(2): 771-778 | - Data not reported in an extractable format or a format that can be analysed |
| Argstatter, H., Hillecke, Th, Thaut, M. et al. (2007) Music therapy in motor rehabilitation. Evaluation of a musico-medical gait training program for hemiparetic stroke patients. <i>Neurologie und rehabilitation</i> 13(3): 159-165 | - Study not reported in English |
| Bamiou, D. E. (2016) Auditory rehabilitation in stroke patients with auditory processing disorders. | - Full text paper not available |
| Barnes, C. L., Smith, M. B., Harriet, E. et al. (2006) A pilot study of bilateral arm training with repetitive auditory cueing in subjects with low functioning upper limb hemiparesis as a result of chronic stroke. <i>Journal of neurologic physical therapy</i> 4: 221 | - Conference abstract |

| Study | Code [Reason] |
|--|--|
| Baylan, S., Quinn, T., Cullen, B. et al. (2016) The effects of music listening on mood and cognition post-stroke. International journal of stroke 11(suppl4): 29 | - Study not reported in English |
| Baylan, S., Swann-Price, R., Peryer, G. et al. (2016) The effects of music listening interventions on cognition and mood post-stroke: a systematic review. Expert Review of Neurotherapeutics 16(11): 1241-1249 | - Systematic review used as source of primary studies |
| Bittman, B., Poornima, I., Smith, M. A. et al. (2020) Gospel Music: A Catalyst for Retention, Engagement, and Positive Health Outcomes for African Americans in a Cardiovascular Prevention and Treatment Program. Advances in Mind-Body Medicine 34(1): 8-16 | - Population not relevant to this review protocol |
| Blythe LaGasse, A. and Knight, A. (2011) Rhythm and music in rehabilitation: A critical review of current research. Critical Reviews in Physical and Rehabilitation Medicine 23(14): 49-67 | - Review article but not a systematic review |
| Breitenfeld, T., Jergovi, K., Vargek Solter, V. et al. (2005) Music therapy in aphatic stroke patients - a pilot study. European journal of neurology 12(suppl2): 55p1060 | - Conference abstract |
| Breitenfeld, T., Vargek Solter, V., Breitenfeld, D. et al. (2005) Is there a benefit for aphasic stroke patients treated with music therapy? - preliminary results. Cerebrovascular diseases (basel, switzerland) 19 (Suppl 2): 92-93 | - Full text paper not available |
| Bunketorp Kall, L., Lundgren-Nilsson, A., Blomstrand, C. et al. (2012) The effects of a rhythm and music-based therapy program and therapeutic riding in late recovery phase following stroke: a study protocol for a three-armed randomized controlled trial. BMC Neurology 12: 141 | - Study reported outcomes that were not included in the protocol |
| Bunketorp-Kall, L., Lundgren-Nilsson, A., Nilsson, M. et al. (2018) Multimodal rehabilitation in the late phase after stroke enhances the life situation of informal caregivers. Topics in Stroke Rehabilitation 25(3): 161-167 | - Study reported outcomes that were not included in the protocol |

| Study | Code [Reason] |
|---|---|
| <p>Bunketorp-Kall, L., Lundgren-Nilsson, A., Samuelsson, H. et al. (2017) Long-Term Improvements After Multimodal Rehabilitation in Late Phase After Stroke: A Randomized Controlled Trial. Stroke 48(7): 1916-1924</p> | <p>- Study reported outcomes that were not included in the protocol</p> |
| <p>Bunketorp-Kall, L., Pekna, M., Pekny, M. et al. (2019) Effects of horse-riding therapy and rhythm and music-based therapy on functional mobility in late phase after stroke. Neurorehabilitation 45(4): 483-492</p> | <p>- Study reported outcomes that were not included in the protocol</p> |
| <p>Cha, Y. J., Kim, J. D., Choi, Y. R. et al. (2018) Effects of gait training with auditory feedback on walking and balancing ability in adults after hemiplegic stroke: a preliminary, randomized, controlled study. International Journal of Rehabilitation Research 41(3): 239-243</p> | <p>- Study reported outcomes that were not included in the protocol</p> |
| <p>Cha, Y.; Kim, Y.; Chung, Y. (2014) Immediate effects of rhythmic auditory stimulation with tempo changes on gait in stroke patients. Journal of physical therapy science 26: 479-482</p> | <p>- Study reported outcomes that were not included in the protocol</p> |
| <p>Chen, F. J., Li, L., Sun, J. L. et al. (2015) The research into the functions of TCM five elements music therapy on anxious with anxiety due to stroke. Henan traditional chinese medicine [he nan zhong yi] 35(6): 1279-1280</p> | <p>- Study not reported in English</p> |
| <p>Choi, W.; Lee, G.; Lee, S. (2015) Effect of the cognitive-motor dual-task using auditory cue on balance of survivors with chronic stroke: a pilot study. Clinical Rehabilitation 29(8): 763-70</p> | <p>- Study reported outcomes that were not included in the protocol</p> |
| <p>Chouhan, S. and Kumar, S. (2012) Comparing the effects of rhythmic auditory cueing and visual cueing in acute hemiparetic stroke. International journal of therapy and rehabilitation 19(6): 344-351</p> | <p>- Study reported outcomes that were not included in the protocol</p> |
| <p>Chouhan, S., Kumar, S., Walker, S. et al. (2012) Comparative study of the effects of rhythmic auditory cueing and visual cueing in acute hemiparetic stroke. International journal of therapy & rehabilitation 19(5): 1-8</p> | <p>- Duplicate reference</p> |
| <p>Cofrancesco, E. M. (1985) The effect of music therapy on hand grasp strength and functional</p> | <p>- Study design not relevant to this review protocol</p> |

| Study | Code [Reason] |
|--|--|
| task performance in stroke patients. Journal of music therapy 22(3): 129-145 | |
| Conklyn, D., Novak, E., Boissy, A. et al. (2012) The effects of modified melodic intonation therapy on nonfluent aphasia: a pilot study. Journal of Speech Language & Hearing Research 55(5): 1463-71 | - Study reported outcomes that were not included in the protocol |
| Copland, D. and Roxbury, T. (2019) To examine whether daily music listening in addition to usual care will result in superior aphasia recovery compared to usual care only, as measured by standard clinical language and communication assessments at 2-4 weeks, 3 months and 6 months post stroke-onset. | - Study not reported in English |
| Crosby, LD, Wong, JS, Chen, JL et al. (2020) An Initial Investigation of the Responsiveness of Temporal Gait Asymmetry to Rhythmic Auditory Stimulation and the Relationship to Rhythm Ability Following Stroke. Frontiers in neurology 11 | - Study design not relevant to this review protocol <i>Single arm trial</i> |
| Dispa, D.; Lejeune, T.; Thonnard, J. L. (2013) The effect of repetitive rhythmic precision grip task-oriented rehabilitation in chronic stroke patients: a pilot study. International Journal of Rehabilitation Research 36(1): 81-7 | - Comparator in study does not match that specified in this review protocol |
| Do, A. (2016) To determine the therapeutic effect of the music glove and conventional hand exercises to subacute stroke patients. | - Full text paper not available |
| Douglass-Kirk, Pedro, Grierson, Mick, Ward, Nick S et al. (2022) Real-time auditory feedback may reduce abnormal movements in patients with chronic stroke. Disability and rehabilitation: 1-7 | - Study reported outcomes that were not included in the protocol - Study does not contain an intervention relevant to this review protocol <i>Intervention lasted for 1 session only</i> |
| Elsner, B. (2018) Auditory stimulation for improving mobility after stroke. | - Full text paper not available |
| Elsner, B., Scholer, A., Kon, T. et al. (2020) Walking with rhythmic auditory stimulation in chronic patients after stroke: A pilot randomized | - Study reported outcomes that were not included in the protocol |

| Study | Code [Reason] |
|--|--|
| controlled trial . <i>Physiotherapy Research International</i> 25(1): e1800 | |
| Fachner, J. C. (2014) Music Therapy for The Rehabilitation of Upper Limb With Stroke Patients (TIMPStro). | - Full text paper not available |
| Ford, M.; Wagenaar, R.; Newell, K. (2007) The effects of auditory rhythms and instruction on walking patterns in individuals post stroke. <i>Gait & posture</i> 26: 150-155 | - Study design not relevant to this review protocol |
| Fotakopoulos, G. and Kotlia, P. (2018) The Value of Exercise Rehabilitation Program Accompanied by Experiential Music for Recovery of Cognitive and Motor Skills in Stroke Patients. <i>Journal of Stroke & Cerebrovascular Diseases</i> 27(11): 2932-2939 | - Data not reported in an extractable format or a format that can be analysed <i>Reports outcomes for all groups combined</i> |
| Fouad, M. A. (2016) Effect of rhythmic auditory stimulation on gait in patients with stroke. <i>International journal of medical and health sciences</i> 3(6) | - Conference abstract |
| Friedman, N., Chan, V., Reinkensmeyer, A. N. et al. (2014) Retraining and assessing hand movement after stroke using the MusicGlove: comparison with conventional hand therapy and isometric grip training. <i>Journal of Neuroengineering & Rehabilitation</i> 11: 76 | - Study reported outcomes that were not included in the protocol |
| Garcia-Casares, Natalia; Barros-Cano, Amanda; Garcia-Arnes, Juan A (2022) Melodic Intonation Therapy in Post-Stroke Non-Fluent Aphasia and Its Effects on Brain Plasticity. <i>Journal of clinical medicine</i> 11(12) | - Systematic review used as source of primary studies |
| Ghai, S. (2018) Effects of Real-Time (Sonification) and Rhythmic Auditory Stimuli on Recovering Arm Function Post Stroke: A Systematic Review and Meta-Analysis. <i>Frontiers in neurology [electronic resource]</i> . 9: 488 | - Systematic review used as source of primary studies |
| Ghai, S. and Ghai, I. (2019) Effects of (music-based) rhythmic auditory cueing training on gait and posture post-stroke: A systematic review & dose-response meta-analysis. <i>Scientific Reports</i> 9(1): 2183 | - Systematic review used as source of primary studies |

| Study | Code [Reason] |
|--|--|
| Goh, M. (2001) The role of music therapy in the rehabilitation of people who have had strokes, specifically focusing on depression. National research register. Issue 1 | - Full text paper not available |
| Grau-Sanchez, J., Segura, E., Sanchez-Pinsach, D. et al. (2021) Enriched Music-supported Therapy for chronic stroke patients: a study protocol of a randomised controlled trial. BMC Neurology 21(1): 19 | - Protocol only |
| Haire, C. (2017) Therapeutic instrumental music performance with sensory-enhanced motor imagery in chronic post-stroke rehabilitation. | - Comparator in study does not match that specified in this review protocol |
| Haire, Catherine M, Tremblay, Luc, Vuong, Veronica et al. (2021) Therapeutic Instrumental Music Training and Motor Imagery in Post-Stroke Upper-Extremity Rehabilitation: A Randomized-Controlled Pilot Study. Archives of rehabilitation research and clinical translation 3(4): 100162 | - Comparator in study does not match that specified in this review protocol <i>Compares therapeutic instrumental music training with the same intervention with motor imagery in addition (with and without metronome cueing) therefore comparing different types of music therapy, which is not a comparison listed in the protocol.</i> |
| Hankinson, Katherine, Shaykevich, Alex, Vallenge, Ann-Maree et al. (2022) A Tailored Music-Motor Therapy and Real-Time Biofeedback Mobile Phone App ('GotRhythm') to Promote Rehabilitation Following Stroke: A Pilot Study. Neuroscience insights 17: 26331055221100587 | - Data not reported in an extractable format or a format that can be analysed <i>Outcomes reported as median and interquartile range values</i> |
| Haro-Martinez, A. M., Lubrini, G., Madero-Jarabo, R. et al. (2019) Melodic intonation therapy in post-stroke nonfluent aphasia: a randomized pilot trial. Clinical Rehabilitation 33(1): 44-53 | - Study reported outcomes that were not included in the protocol |
| Haro-Martinez, Ana, Perez-Araujo, Carmen M, Sanchez-Caro, Juan M et al. (2021) Melodic Intonation Therapy for Post-stroke Non-fluent Aphasia: Systematic Review and Meta-Analysis. Frontiers in neurology 12: 700115 | - Systematic review used as source of primary studies |
| Hewitt, L. and Sanctuary, C. (2015) Stroke sounds: music listening in stroke rehabilitation. International journal of stroke 10(suppl3): 64 | - Conference abstract |

| Study | Code [Reason] |
|---|---|
| <p>Huang, Wen-Hao, Dou, Zu-Lin, Jin, Hui-Min et al. (2021) The Effectiveness of Music Therapy on Hand Function in Patients With Stroke: A Systematic Review of Randomized Controlled Trials. <i>Frontiers in neurology</i> 12: 641023</p> | <p>- Systematic review used as source of primary studies</p> |
| <p>Jarvinen-Lepisto, P.; Burger, B.; Ala-Ruona, E. (2014) Motor performance in post-stroke recovery using active music therapy. 13th international conference for music perception and cognition / 5th conference of asia-pacific society</p> | <p>- Crossover trial (unit of randomisation = participant)</p> |
| <p>Jia, C., Zhang, H., Ni, G. et al. (2017) Spasmodic hemiplegia after stroke treated with scalp acupuncture, music therapy and rehabilitation: a randomized controlled trial. <i>Zhongguo zhen jiu [Chinese acupuncture & moxibustion]</i> 37(12): 1271-1275</p> | <p>- Study not reported in English</p> |
| <p>Jiang, Y., Yang, Y., Xiang, R. et al. (2015) Clinical study of post-stroke speech apraxia treated with scalp electric acupuncture under anatomic orientation and rehabilitation training. <i>Zhongguo zhen jiu [Chinese acupuncture & moxibustion]</i> 35(7): 661-664</p> | <p>- Study not reported in English</p> |
| <p>John, S.; Khanna, G. L.; Kotwal, P. (2010) Effect of music therapy and meditation along with conventional physiotherapy management in sub-acute stroke patients. <i>British journal of sports medicine</i> 44(suppl1): i14</p> | <p>- Conference abstract</p> |
| <p>Kang, T. W. (2015) Robot-assisted walking training for patients with subacute stroke: randomized controlled pilot trial of rhythmic arm swing versus arm fixation during training.</p> | <p>- Full text paper not available</p> |
| <p>Kang, T. W., Oh, D. W., Lee, J. H. et al. (2018) Effects of integrating rhythmic arm swing into robot-assisted walking in patients with subacute stroke: a randomized controlled pilot study. <i>International Journal of Rehabilitation Research</i> 41(1): 57-62</p> | <p>- Study does not contain an intervention relevant to this review protocol</p> <p><i>Rhythmic arm swing without a music component built into the intervention</i></p> |
| <p>Keller, I. and Lefin-Rank, G. (2010) Improvement of visual search after audiovisual exploration training in hemianopic patients. <i>Neurorehabilitation and Neural Repair</i> 24(7): 666-673</p> | <p>- Study does not contain an intervention relevant to this review protocol</p> |

| Study | Code [Reason] |
|---|--|
| <p>Kim, J. H., Park, S. G., Lim, H. J. et al. (2012) Effects of the combination of rhythmic auditory stimulation and task-oriented training on functional recovery of subacute stroke patients. Journal of physical therapy science 24(12): 1307-1313</p> | <p>- Study reported outcomes that were not included in the protocol</p> |
| <p>Kim, J. S. and Oh, D. W. (2012) Home-based auditory stimulation training for gait rehabilitation of chronic stroke patients. Journal of physical therapy science 24: 775-777</p> | <p>- Study reported outcomes that were not included in the protocol</p> |
| <p>Kiper, P. (2017) Proprioceptive Stimulation With Manual Bilateral Rhythmic Exercise in Post-stroke Patients (BAT).</p> | <p>- Full text paper not available</p> |
| <p>Klinke, M. E., Hafsteinsdottir, T. B., Hjaltason, H. et al. (2015) Ward-based interventions for patients with hemispatial neglect in stroke rehabilitation: a systematic literature review. International Journal of Nursing Studies 52(8): 1375-403</p> | <p>- Study does not contain an intervention relevant to this review protocol</p> |
| <p>Kumari, N. (2017) Effects of rhythmic auditory cueing along with task oriented activities on upper limb functions in stroke patients.</p> | <p>- Full text paper not available</p> |
| <p>Le Danseur, M., Crow, A. D., Stutzman, S. E. et al. (2019) Music as a Therapy to Alleviate Anxiety During Inpatient Rehabilitation for Stroke. Rehabilitation Nursing Journal 44(1): 29-34</p> | <p>- Comparator in study does not match that specified in this review protocol</p> |
| <p>Lee, S. H.; Lee, K. J.; Song, C. H. (2012) Effects of rhythmic auditory stimulation (RAS) on gait ability and symmetry after stroke. Journal of physical therapy science 24(4): 311-314</p> | <p>- Study design not relevant to this review protocol</p> |
| <p>Lee, S.; Lee, K.; Song, C. (2018) Gait Training with Bilateral Rhythmic Auditory Stimulation in Stroke Patients: A Randomized Controlled Trial. Brain Sciences 8(9): 31</p> | <p>- Study reported outcomes that were not included in the protocol</p> |
| <p>Lin, S. I. (2007) Effect of rhythmic auditory cues on gait of stroke patients. Cerebrovascular diseases (basel, switzerland) 23(suppl2): 128</p> | <p>- Full text paper not available</p> |

| Study | Code [Reason] |
|---|--|
| <p>Magee, WI, Clark, I, Tamplin, J et al. (2017) Music interventions for acquired brain injury. Cochrane Database of Systematic Reviews</p> | <p>- Population not relevant to this review protocol</p> |
| <p>Mainka, S., Wissel, J., Voller, H. et al. (2018) The Use of Rhythmic Auditory Stimulation to Optimize Treadmill Training for Stroke Patients: A Randomized Controlled Trial. Frontiers in neurology [electronic resource]. 9: 755</p> | <p>- Study reported outcomes that were not included in the protocol</p> |
| <p>McCombe Waller, S.; Liu, W.; Whittall, J. (2008) Temporal and spatial control following bilateral versus unilateral training. Human Movement Science 27(5): 749-58</p> | <p>- Study reported outcomes that were not included in the protocol</p> |
| <p>McCue, P., Del Din, S., Hunter, H. et al. (2020) Auditory rhythmical cueing to improve gait and physical activity in community-dwelling stroke survivors (ACTIVATE): study protocol for a pilot randomised controlled trial. Pilot & Feasibility Studies 6: 68</p> | <p>- Protocol only</p> |
| <p>McIntosh, G. C., Rice, R. R., Prassas, S. G. et al. (1993) Rhythmic auditory-motor entrainment as gait rehabilitation technique with stroke patients. International congress on stroke rehabilitation: 43</p> | <p>- Duplicate reference</p> |
| <p>McIntosh, G. C., Thaut, M. H., Rice, R. R. et al. (1993) Auditory rhythmic cuing in gait rehabilitation with stroke patients. Canadian journal of neurological sciences 20(suppl4): 168</p> | <p>- Full text paper not available</p> |
| <p>Moon, S. Y. (2008) The effects of piano-playing music therapy on motor coordination of stroke patients using midi-based computer analysis. Neurorehabilitation and neural repair 22(5): 593</p> | <p>- Full text paper not available</p> |
| <p>Moumdjian, L., Sarkamo, T., Leone, C. et al. (2017) Effectiveness of music-based interventions on motricity or cognitive functioning in neurological populations: a systematic review. European journal of physical & rehabilitation medicine. 53(3): 466-482</p> | <p>- Population not relevant to this review protocol</p> |
| <p>Nikmaram, N., Scholz, D. S., Grosbach, M. et al. (2019) Musical Sonification of Arm Movements in Stroke Rehabilitation Yields Limited Benefits. Frontiers in Neuroscience 13: 1378</p> | <p>- Data not reported in an extractable format or a format that can be analysed</p> |

| Study | Code [Reason] |
|---|---|
| Oiga, L. (2014) The effect of music and rhythmic auditory stimulation on upper motor strength rehabilitation of hemiparetic stroke patients in a tertiary hospital: a randomized controlled study. International journal of stroke 9suppl3: 237 | - Conference abstract |
| Olson, D. M., Perera, A., Atem, F. et al. (2019) Music in mechanically ventilated stroke patients. British journal of neuroscience nursing 15: 8 | - Comparator in study does not match that specified in this review protocol |
| Park, I. M., Oh, D. W., Kim, S. Y. et al. (2010) Clinical feasibility of integrating fast-tempo auditory stimulation with self-adopted walking training for improving walking function in post-stroke patients: a randomized, controlled pilot trial. Journal of physical therapy science 22: 295-300 | - Study reported outcomes that were not included in the protocol |
| Park, M. O. and Lee, S. H. (2018) Effects of cognitive-motor dual-task training combined with auditory motor synchronization training on cognitive functioning in individuals with chronic stroke: A pilot randomized controlled trial. Medicine 97(22): e10910 | - Study reported outcomes that were not included in the protocol |
| Prassas, S. G., Thaut, M. H., McIntosh, G. C. et al. (1997) Effect of auditory rhythmic cuing on gait kinematic parameters in hemiparetic stroke patients. Gait & posture 6: 218-223 | - Study design not relevant to this review protocol |
| Purdie, H.; Hamilton, S.; Baldwin, S. (1997) Music therapy: facilitating behavioural and psychological change in people with stroke--a pilot study. International Journal of Rehabilitation Research 20(3): 325-7 | - Data not reported in an extractable format or a format that can be analysed <i>Does not report the number of participants in each study arm and so unable to interpret results</i> |
| Raglio, A. (2017) Music therapy for rehabilitation in stroke patients (SONICHAND). | - Full text paper not available |
| Raglio, A., Attardo, L., Gontero, G. et al. (2015) Effects of music and music therapy on mood in neurological patients. World Journal of Psychiatry 5(1): 68-78 | - Population not relevant to this review protocol |
| Raglio, A., Oasi, O., Gianotti, M. et al. (2016) Improvement of spontaneous language in stroke patients with chronic aphasia treated with music therapy: a randomized controlled trial. | - Data not reported in an extractable format or a format that can be analysed <i>Reports median and interquartile range values for outcomes only</i> |

| Study | Code [Reason] |
|---|---|
| International Journal of Neuroscience 126(3): 235-42 | |
| Reagon, C., Gale, N., Enright, S. et al. (2016) A mixed-method systematic review to investigate the effect of group singing on health related quality of life. Complementary Therapies in Medicine 27: 1-11 | - Population not relevant to this review protocol |
| Renna, L., Frkovic, N., Spear, M. et al. (2012) Stroke sounds: music listening in stroke rehabilitation. International journal of stroke 7(suppl1): 58 | - Conference abstract |
| Richards, L. G., Senesac, C. R., Davis, S. B. et al. (2008) Bilateral arm training with rhythmic auditory cueing in chronic stroke: not always efficacious. Neurorehabilitation and neural repair 22: 180-184 | - Study design not relevant to this review protocol |
| Rodriguez-Fornells, A. (2014) Music Therapy to Restore Motor Deficits After Stroke (NEUROMUSIC). | - Full text paper not available |
| Rosenberg, K. (2017) Multimodal Interventions Improve Stroke Recovery. American Journal of Nursing 117(10): 61 | - Commentary only |
| Sarkamo, T., Leo T. Sihvonen A. Ripolles P. Rodri and guez-Fornells, A. Tervaniemi M. (2016) Cognitive, emotional and neural benefits of music on stroke recovery. European stroke journal 1(suppl1): 730-731 | - Conference abstract |
| Sarkamo, T., Pihko, E., Laitinen, S. et al. (2010) Music and speech listening enhance the recovery of early sensory processing after stroke. Journal of Cognitive Neuroscience 22(12): 2716-27 | - Data not reported in an extractable format or a format that can be analysed |
| Sarkamo, T., Ripolles, P., Vepsalainen, H. et al. (2014) Structural changes induced by daily music listening in the recovering brain after middle cerebral artery stroke: a voxel-based morphometry study. Frontiers in Human Neuroscience 8: 245 | - Study reported outcomes that were not included in the protocol |
| Sarkamo, T., Tervaniemi, M., Laitinen, S. et al. (2008) Music listening enhances cognitive | - Data not reported in an extractable format or a format that can be analysed |

| Study | Code [Reason] |
|--|---|
| recovery and mood after middle cerebral artery stroke . Brain 131(pt3): 866-76 | <i>Reports as graph data only</i> |
| Schauer, M. and Mauritz, K. H. (2003) Musical motor feedback (MMF) in walking hemiparetic stroke patients: randomized trials of gait improvement . Clinical Rehabilitation 17(7): 713-22 | - Study reported outcomes that were not included in the protocol |
| Schneider, S., Schonle, P. W., Altenmuller, E. et al. (2007) Using musical instruments to improve motor skill recovery following a stroke . Journal of Neurology 254(10): 1339-46 | - Study reported outcomes that were not included in the protocol |
| Scholz, D. S., Rohde, S., Nikmaram, N. et al. (2016) Sonification of Arm Movements in Stroke Rehabilitation - A Novel Approach in Neurologic Music Therapy . Frontiers in neurology [electronic resource]. 7: 106 | - Data not reported in an extractable format or a format that can be analysed |
| Shaw, Lisa, McCue, Patricia, Brown, Philip et al. (2022) Auditory rhythmical cueing to improve gait in community-dwelling stroke survivors (ACTIVATE): a pilot randomised controlled trial . Pilot and feasibility studies 8(1): 239 | - Study reported outcomes that were not included in the protocol |
| Shen, J., Shen, X., Chang, G. F. et al. (1994) Effect of music electrotherapy on cerebral infarction. Chinese journal of physical therapy 17(3): 162-164 | - Full text paper not available |
| Shen, J., Shen, X., Chang, G. et al. (1994) Effect of music electrotherapy treatment on cerebral infarction. Chinese journal of physical therapy 17(3): 162-164 | - Full text paper not available |
| Shin, J. and Chung, Y. (2017) Influence of visual feedback and rhythmic auditory cue on walking of chronic stroke patient induced by treadmill walking in real-time basis . Neurorehabilitation 41(2): 445-452 | - Study reported outcomes that were not included in the protocol |
| Shin, Jin and Chung, Yijung (2022) The effects of treadmill training with visual feedback and rhythmic auditory cue on gait and balance in chronic stroke patients: A randomized controlled trial . NeuroRehabilitation | - Study reported outcomes that were not included in the protocol |

| Study | Code [Reason] |
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| <p>Sihvonen, A. J., Leo, V., Ripolles, P. et al. (2020) Vocal music enhances memory and language recovery after stroke: pooled results from two RCTs. <i>Annals of Clinical & Translational Neurology</i> 7(11): 2272-2287</p> | <p>- Pooled analysis of a published and an unpublished trial with inappropriate methodology for this review</p> |
| <p>Silveira, T. M. (2018) Examining the effect of FES+iPad-based music therapy on upper limb function and wellbeing outcomes for stroke survivors.</p> | <p>- Full text paper not available</p> |
| <p>Soinila, S. (2012) Music Listening and Stroke Recovery (MUKU2).</p> | <p>- Full text paper not available</p> |
| <p>Stewart, C., Subbarayan, S., Paton, P. et al. (2019) Non-pharmacological interventions for the improvement of post-stroke quality of life amongst older stroke survivors: a systematic review of systematic reviews (The SENATOR ONTOP series). <i>European Geriatric Medicine</i> 10(3): 359-386</p> | <p>- Study design not relevant to this review protocol</p> |
| <p>Stinear, J. W. and Byblow, W. D. (2004) Rhythmic bilateral movement training modulates corticomotor excitability and enhances upper limb motricity poststroke: a pilot study. <i>Journal of clinical neurophysiology</i> 21(2): 124-131</p> | <p>- Study design not relevant to this review protocol</p> |
| <p>Street, A. J., Magee, W. L., Bateman, A. et al. (2018) Home-based neurologic music therapy for arm hemiparesis following stroke: results from a pilot, feasibility randomized controlled trial. <i>Clinical Rehabilitation</i> 32(1): 18-28</p> | <p>- Crossover trial (unit of randomisation = participant)</p> |
| <p>Street, A. J., Magee, W. L., Odell-Miller, H. et al. (2015) Home-based neurologic music therapy for upper limb rehabilitation with stroke patients at community rehabilitation stage-a feasibility study protocol. <i>Frontiers in Human Neuroscience</i> 9: 480</p> | <p>- Protocol only</p> |
| <p>Studebaker, S. (2007) The effect of a music therapy protocol on the attentional abilities of stroke patients. Unpublished masters thesis. University of Kansas</p> | <p>- Full text paper not available</p> |
| <p>Suh, J. H., Han, S. J., Jeon, S. Y. et al. (2014) Effect of rhythmic auditory stimulation on gait and balance in hemiplegic stroke patients. <i>Neurorehabilitation</i> 34(1): 193-9</p> | <p>- Study reported outcomes that were not included in the protocol</p> |

| Study | Code [Reason] |
|---|--|
| Sukumaran, S., Sivadasan, S., Sakunthala, P. T. et al. (2019) Effect of combined visual-auditory-sensory stimulation in hemineglect syndrome following right hemispheric ischemic strokes: a randomized control trial. | <ul style="list-style-type: none"> - Thesis paper - Study reported outcomes that were not included in the protocol |
| Syros, Apostolis; Kotlia, Polikceni; Fotakopoulos, George (2022) Preliminary findings from an acupuncture and experiential/traditional music therapy during the standard care of rehabilitation exercise program for recovery on post-stroke upper limb dysfunction. The International journal of neuroscience 132(11): 1110-1117 | <ul style="list-style-type: none"> - Data not reported in an extractable format or a format that can be analysed <p><i>Does not report the mean and standard deviations for the relevant outcomes instead reporting alternative analyses that are not as relevant</i></p> |
| Thaut, M. H., Hoemberg, B., Hurt, C. P. et al. (1998) Rhythmic entrainment of paretic arm movements in stroke patients. Proceedings of the society for neuroscience 24: 1663 | <ul style="list-style-type: none"> - Full text paper not available |
| Thaut, M. H., Hoemberg, V., Hurt, C. P. et al. (1998) Rhythmic entrainment of hemiparetic arm movements in stroke patients. Society for neuroscience abstracts 24: 1663 | <ul style="list-style-type: none"> - Full text paper not available |
| Thaut, M. H., McIntosh, C. G., Rice, R. et al. (1993) Effect of rhythmic cuing on temporal stride parameters and EMG patterns in hemiparetic gait of stroke patients. Journal of neurological rehabilitation 7: 9-16 | <ul style="list-style-type: none"> - Study design not relevant to this review protocol |
| Thaut, M. H.; McIntosh, G. C.; Rice, R. R. (1997) Rhythmic facilitation of gait training in hemiparetic stroke rehabilitation. Journal of the Neurological Sciences 151(2): 207-12 | <ul style="list-style-type: none"> - Study reported outcomes that were not included in the protocol |
| Thaut, M. H., McIntosh, G. C., Rice, R. R. et al. (1995) Rhythmic auditory motor training in gait rehabilitation of stroke patients. Journal of stroke and cerebrovascular diseases 5(2): 100 | <ul style="list-style-type: none"> - Full text paper not available |
| Tinga, A. M., Visser-Meily, J. M., van der Smagt, M. J. et al. (2016) Multisensory Stimulation to Improve Low- and Higher-Level Sensory Deficits after Stroke: A Systematic Review. Neuropsychology Review 26(1): 73-91 | <ul style="list-style-type: none"> - Study does not contain an intervention relevant to this review protocol |
| Tong, Y., Forreider, B., Sun, X. et al. (2015) Music-supported therapy (MST) in improving | <ul style="list-style-type: none"> - Study reported outcomes that were not included in the protocol |

| Study | Code [Reason] |
|--|--|
| <p>post-stroke patients' upper-limb motor function: a randomised controlled pilot study. Neurological Research 37(5): 434-40</p> | |
| <p>Van Criekinge, T., D'Aout, K., O'Brien, J. et al. (2019) The Influence of Sound-Based Interventions on Motor Behavior After Stroke: A Systematic Review. Frontiers in neurology [electronic resource]. 10: 1141</p> | <p>- Systematic review used as source of primary studies</p> |
| <p>Van Der Meulen, I., Van De Sandt-Koenderman, M. W., Heijnenbrok, M. H. et al. (2016) Melodic Intonation Therapy in Chronic Aphasia: Evidence from a Pilot Randomized Controlled Trial. Frontiers in Human Neuroscience 10: 533</p> | <p>- Study reported outcomes that were not included in the protocol</p> |
| <p>van der Meulen, I., van de Sandt-Koenderman, W. M., Heijnenbrok-Kal, M. H. et al. (2014) The Efficacy and Timing of Melodic Intonation Therapy in Subacute Aphasia. Neurorehabilitation & Neural Repair 28(6): 536-44</p> | <p>- Study reported outcomes that were not included in the protocol</p> |
| <p>van Vugt, F. T., Kafczyk, T., Kuhn, W. et al. (2016) The role of auditory feedback in music-supported stroke rehabilitation: A single-blinded randomised controlled intervention. Restorative Neurology & Neuroscience 34(2): 297-311</p> | <p>- Comparator in study does not match that specified in this review protocol</p> |
| <p>Volpi, J. J. (2018) Stroke recovery and music or no music.</p> | <p>- Study not reported in English</p> |
| <p>Wang, Y., Pan, W. Y., Li, F. et al. (2021) Effect of Rhythm of Music Therapy on Gait in Patients with Stroke. Journal of Stroke and Cerebrovascular Diseases 30 (3)</p> | <p>- Study reported outcomes that were not included in the protocol</p> |
| <p>Wheeler, B. L.; Shiflett, S. C.; Nayak, S. (2003) Effects of number of sessions and group or individual music therapy on the mood and behavior of people who have had strokes or traumatic brain injuries. Nordic journal of music therapy 12(2): 139-151</p> | <p>- Population not relevant to this review protocol <i>Less than 80% of participants had a stroke</i></p> |
| <p>Whitall, J., McCombe Waller, S., Silver, K. H. et al. (2000) Repetitive bilateral arm training with rhythmic auditory cueing improves motor function in chronic hemiparetic stroke. Stroke; a journal of cerebral circulation 31(10): 2390-2395</p> | <p>- Study design not relevant to this review protocol</p> |

| Study | Code [Reason] |
|--|---|
| Whitall, J., McCombe-Waller, S., Gordes, K. et al. (1999) Locomotor training with and without rhythmic auditory stimulation in patients with chronic stroke. <i>Neurology report</i> 23(5): 190 | - Conference abstract |
| Wright, R. L., Brownless, S. B., Pratt, D. et al. (2017) Stepping to the Beat: Feasibility and Potential Efficacy of a Home-Based Auditory-Cued Step Training Program in Chronic Stroke. <i>Frontiers in neurology</i> [electronic resource]. 8: 412 | - Comparator in study does not match that specified in this review protocol |
| Yakupov, E. Z., Nalbat, A. V., Semenova, M. V. et al. (2019) Efficacy of music therapy in the rehabilitation of stroke patients. <i>Neuroscience and behavioral physiology</i> 49(1): 121-128 | - Study reported outcomes that were not included in the protocol |
| Yakupov, E. Z., Nalbat, A. V., Semenova, M. V. et al. (2017) Music therapy as an effective method of neurorehabilitation. <i>Zhurnal neurologii i psikiatrii imeni S.S. Korsakova</i> 117(5): 14-21 | - Study not reported in English |
| Yoo, G. E. and Kim, S. J. (2016) Rhythmic Auditory Cueing in Motor Rehabilitation for Stroke Patients: Systematic Review and Meta-Analysis. <i>Journal of Music Therapy</i> 53(2): 149-77 | - Systematic review used as source of primary studies |
| Yoon, S. K. and Kang, S. H. (2016) Effects of inclined treadmill walking training with rhythmic auditory stimulation on balance and gait in stroke patients. <i>Journal of Physical Therapy Science</i> 28(12): 3367-3370 | - Study reported outcomes that were not included in the protocol |
| Young, Hui-Ju, Mehta, Tapan, Herman, Cassandra et al. (2021) The Effects of a Movement-to-Music (M2M) Intervention on Physical and Psychosocial Outcomes in People Poststroke: A Randomized Controlled Trial. <i>Archives of rehabilitation research and clinical translation</i> 3(4): 100160 | - Study reported outcomes that were not included in the protocol |
| Zhang, J. and Chen, C. (2016) Effect of audio training on executive dysfunction in patients with stroke. <i>Chinese journal of cerebrovascular diseases</i> 13(7): 356-359 | - Full text paper not available |
| Zhang, Xiaoying; Li, Jianjun; Du, Yi (2021) Melodic Intonation Therapy on Non-fluent Aphasia After Stroke: A Systematic Review and | - Systematic review used as source of primary studies |

| Study | Code [Reason] |
|---|--|
| Analysis on Clinical Trials . Frontiers in neuroscience 15: 753356 | |
| Zhang, Y., Cai, J., Zhang, Y. et al. (2016) Improvement in Stroke-induced Motor Dysfunction by Music-supported Therapy: A Systematic Review and Meta-analysis . Scientific Reports 6: 38521 | - Systematic review used as source of primary studies |
| Zhang, Y.; Yao, Y.; Lu, X. (2015) Therapeutic effect of music therapy and speech language therapy on post-stroke patients with non-fluent aphasia. Chinese journal of neurology 48(4): 274-278 | - Study not reported in English |
| Zondervan, D. K., Friedman, N., Chang, E. et al. (2016) Home-based hand rehabilitation after chronic stroke: Randomized, controlled single-blind trial comparing the MusicGlove with a conventional exercise program . Journal of Rehabilitation Research & Development 53(4): 457-72 | - Study reported outcomes that were not included in the protocol |

Health Economic studies

Published health economic studies that met the inclusion criteria (relevant population, comparators, economic study design, published 2006 or later and not from non-OECD country or USA) but that were excluded following appraisal of applicability and methodological quality are listed below. See the health economic protocol for more details.

Table 23: Studies excluded from the health economic review

| Reference | Reason for exclusion |
|-----------|----------------------|
| None | |

Appendix K – Research recommendations – full details

K.1 Research recommendation

What is the clinical and cost-effectiveness of music therapy for people after a first stroke or recurrent strokes?

K.1.1 Why this is important

Music therapy is an evidence based clinical intervention, delivered by trained music therapists with the aim to help people achieve their therapeutic goals. It is becoming increasingly used to help people after a stroke to support people's emotional, cognitive, physical and communication needs. This review identified studies that in general reported positive outcomes of music interventions. However, the majority of the evidence was for music interventions not delivered by trained music therapists. Furthermore, the evidence base was limited due to small sample sizes and a lack of cost effectiveness data. Therefore, it was not possible to make a recommendation for use in the NHS at this time. High quality randomised controlled trials, with a larger number of participants that include cost effectiveness data and compare music therapy with a time matched appropriate comparator are needed. Research should also include outcomes important to people who have had a stroke such as stroke-specific Patient-reported Outcome Measures and activities of daily living to fully explore the possible benefits of this therapy. The committee highlighted that additional therapies may be present that incorporate sound and could therefore be beneficial for people after stroke (for example: sound therapy). While this was not investigated during this guideline update, this was highlighted as a potential area that could benefit from further investigation.

K.1.2 Rationale for research recommendation

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| Importance to 'patients' or the population | Music therapy is becoming increasingly popular amongst stroke survivors. Lay members of the committee shared their own personal experiences of music therapy and stated that it significantly improved both their quality of life and that of their family members. It is currently not widely used in an NHS setting and therefore further research is important to be able to recommend its use in the NHS. |
| Relevance to NICE guidance | There is a growing body of evidence into the use of music therapy for a variety of conditions covered by NICE guidance. This review showed benefits of music therapy for a number of reported outcome measures included health-related quality of life and psychological distress. However, due to the limited evidence base it was not possible to make a recommendation at this stage. Further evidence would help to answer the original review question and to inform future guidance. |
| Relevance to the NHS | Music therapy is an emerging intervention with a growing evidence base. It may be an effective therapy for a wide range of conditions so has the scope to help a large number of patients. Currently the evidence base is too small to be |

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| | able make a positive recommendation for its use. As music therapy is not widely used in the NHS there would likely be a large resource impact so further robust randomised controlled trials with cost effectiveness data are required. |
| National priorities | None identified. |
| Current evidence base | This review identified studies relating to different types of music interventions which included a number of benefits including health-related quality of life and psychological distress. However, the evidence base was limited due to small sample sizes, lack of cost effectiveness data and no time matched comparator therapy. Large, high-quality randomised controlled trials that include cost effectiveness data and compare music therapy with a time matched social activity as a comparator are therefore required. The committee highlighted that additional therapies may be present that incorporate sound and could therefore be beneficial for people after stroke (for example: sound therapy). While this was not investigated during this guideline update, this was highlighted as a potential area that could benefit from further investigation. |
| Equality considerations | No specific equality considerations were identified. The committee noted that in general throughout the guideline, people with communication and cognitive difficulties, older people and people who have had a previous stroke or transient ischaemic attack were excluded from trials but are people that the guideline is for. Therefore, research should aim to include these people where possible. |

K.1.3 Modified PICO table

| | |
|--------------|---|
| Population | <p>Inclusion:</p> <ul style="list-style-type: none"> Adults (age ≥ 16 years) who have had a first stroke or recurrent stroke (including people after a subarachnoid haemorrhage) <p>Exclusion:</p> <ul style="list-style-type: none"> Children (age < 16 years) People who have had a transient ischaemic attack |
| Intervention | <ul style="list-style-type: none"> Music therapy delivered by trained music therapists <p>(Music therapy can include any type of music therapy or sound based therapy deemed appropriate by the therapist)</p> |

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| Comparator | <ul style="list-style-type: none"> • Time matched intervention of equivalent interaction with a healthcare professional that does not involve music • No additional treatment (usual care comparison) |
| Outcome | <p>At time period</p> <ul style="list-style-type: none"> • <6 months • ≥6 months <ul style="list-style-type: none"> • Person/participant generic health-related quality of life • Carer generic health-related quality of life • Activities of daily living • Psychological distress • Stroke-specific Patient-Reported Outcome Measures • Wellbeing scores • Participation in leisure activities/social groups scores • Withdrawal due to adverse events |
| Study design | Randomised controlled trial |
| Timeframe | 6 months |
| Additional information | <p>Subgroup analyses:</p> <ul style="list-style-type: none"> • Severity of stroke (NIHSS: mild, moderate, severe, very severe) • Time after stroke at the start of the trial (hyperacute, acute, subacute, chronic) |