

Appendix K-Forest Plots

Diagnosis

Diagnosis and accuracy of diagnostic test

Figure 6.1: forest plot of DSM-III-R diagnostic test with DSM-IV as a reference standard in a hospital setting

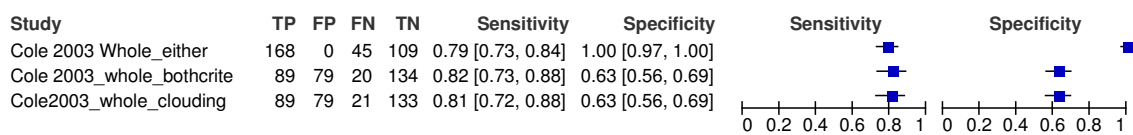


Figure 6.2: forest plot of DSM-III diagnostic test with DSM-III-R as a reference standard in a hospital setting

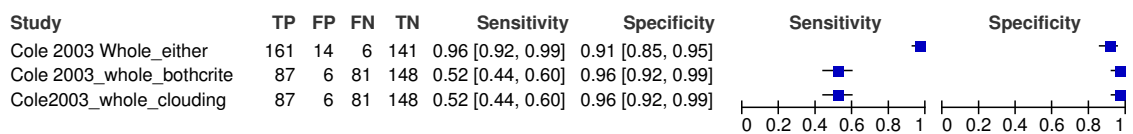


Figure 6.3: forest plot of ICD-10 diagnostic test with DSM-III-R as a reference standard in a hospital setting

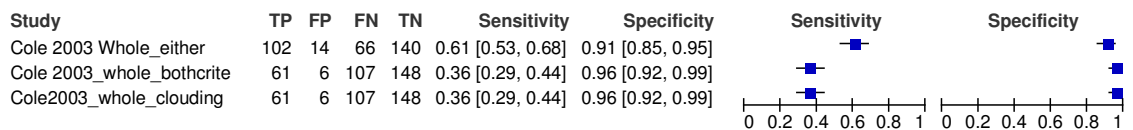


Figure 6.4: forest plot of DSM-III-R compared with DSM-IV in a hospital setting-subgroup analyses

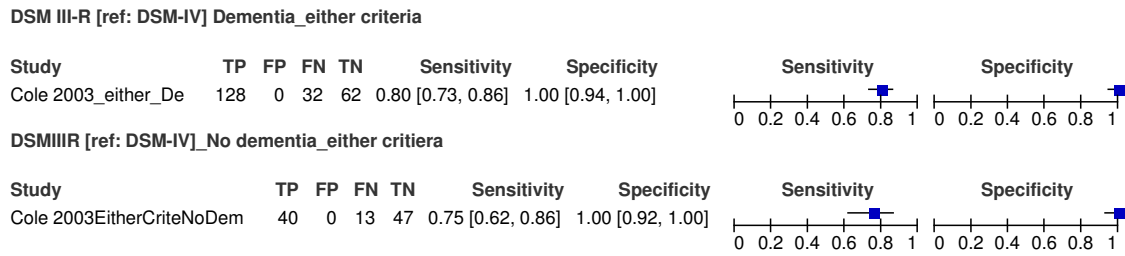


Figure 6.5: forest plot of DSM-III-R compared with DSM-III-R in a hospital setting - subgroup analyses

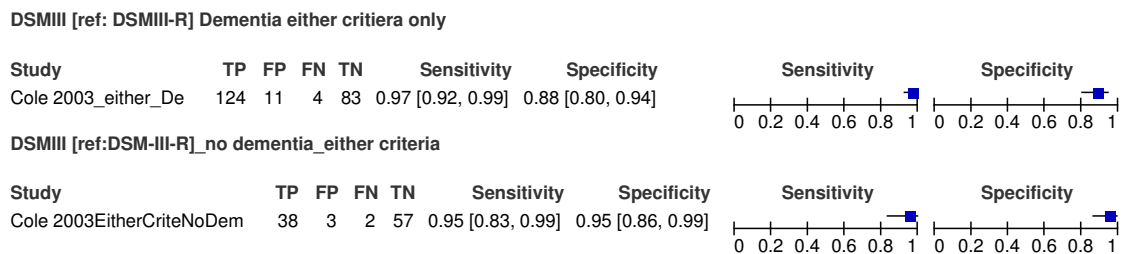


Figure 6.6: forest plot of ICD-10 compared with DSM-III-R in a hospital setting-subgroup analyses

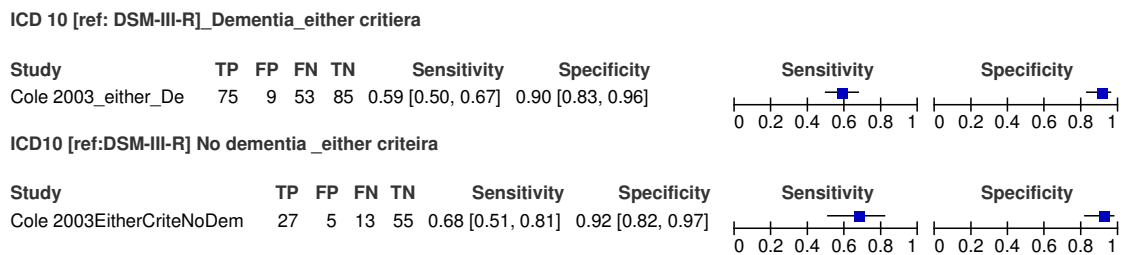
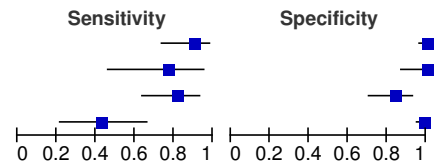


Figure 6.7: forest plot of index tests compared with DSM-IV in a hospital setting

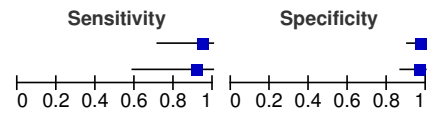
CAM short version [ref: DSMIV]

Study	TP	FP	FN	TN	Sensitivity	Specificity
Gonzalez 2004	27	0	3	93	0.90 [0.73, 0.98]	1.00 [0.96, 1.00]
Hestermann 2009_Rater 2	10	0	3	26	0.77 [0.46, 0.95]	1.00 [0.87, 1.00]
Laurila 2002	26	8	6	41	0.81 [0.64, 0.93]	0.84 [0.70, 0.93]
Radtke 2008	9	2	12	131	0.43 [0.22, 0.66]	0.98 [0.95, 1.00]



CAM long version [ref: DSMIV]

Study	TP	FP	FN	TN	Sensitivity	Specificity
Fabbri 2001	16	3	1	80	0.94 [0.71, 1.00]	0.96 [0.90, 0.99]
Yates 2009	10	2	1	49	0.91 [0.59, 1.00]	0.96 [0.87, 1.00]



DRS-R-98 [Ref: DSM-IV]

Study	TP	FP	FN	TN	Sensitivity	Specificity
Andrew 2009	31	16	24	74	0.56 [0.42, 0.70]	0.82 [0.73, 0.89]

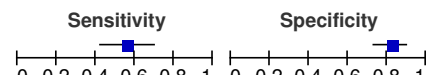


Figure 6.8: forest plot of DRS-R-98 compared with DSM-IV in a hospital setting- subgroup analysis

Study	TP	FP	FN	TN	Sensitivity	Specificity
Andrew 2009	13	12	9	24	0.59 [0.36, 0.79]	0.67 [0.49, 0.81]

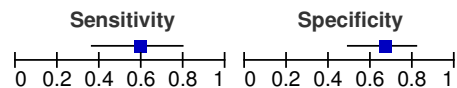


Figure 6.9: forest plot of CAM compared with ICD-10 in a hospital setting- subgroup analysis

Study	TP	FP	FN	TN	Sensitivity	Specificity
Laurila 2002	8	26	2	45	0.80 [0.44, 0.97]	0.63 [0.51, 0.75]

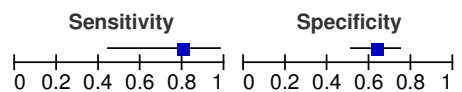
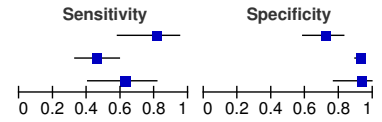


Figure 6.10: forest plot of index test compared with DSM-III-R

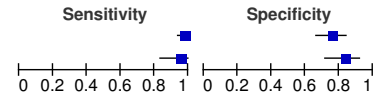
CAM short version [ref: DSM III-R]

Study	TP	FP	FN	TN	Sensitivity	Specificity
Laurila 2002	17	17	4	43	0.81 [0.58, 0.95]	0.72 [0.59, 0.83]
Pompei 1995	28	28	33	338	0.46 [0.33, 0.59]	0.92 [0.89, 0.95]
Rockwood 1994	15	2	9	26	0.63 [0.41, 0.81]	0.93 [0.76, 0.99]



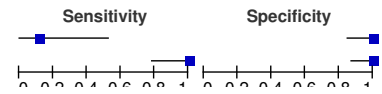
CAM long version [ref: DSM-III-R]

Study	TP	FP	FN	TN	Sensitivity	Specificity
Cole 2003 Dementia more than 6 smptoms	125	24	3	75	0.98 [0.93, 1.00]	0.76 [0.66, 0.84]
Cole 2003 No dementia; more than 6 symptoms	38	10	2	50	0.95 [0.83, 0.99]	0.83 [0.71, 0.92]



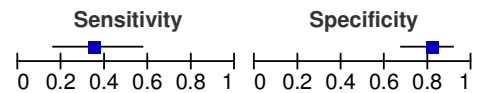
CAM [type of version unclear] [ref: DSM-III-R]

Study	TP	FP	FN	TN	Sensitivity	Specificity
Rolfson 1999b_nurseassess	1	0	7	22	0.13 [0.00, 0.53]	1.00 [0.85, 1.00]
Rolfson 1999b_physician	15	0	0	26	1.00 [0.78, 1.00]	1.00 [0.87, 1.00]



MMSE [ref: DSM-III-R]

Study	TP	FP	FN	TN	Sensitivity	Specificity
Rolfson 1999b	8	9	15	39	0.35 [0.16, 0.57]	0.81 [0.67, 0.91]



Clock drawing test [ref: DSM-III-R]

Study	TP	FP	FN	TN	Sensitivity	Specificity
Rolfson 1999b	2	1	21	47	0.09 [0.01, 0.28]	0.98 [0.89, 1.00]

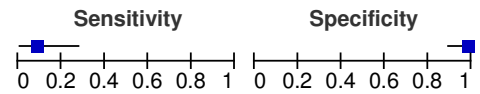
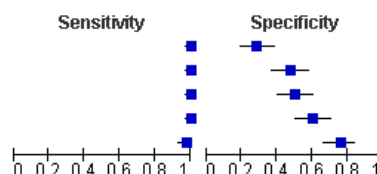


Figure 6.11: forest plot of number of symptoms in index test compared with DSMIII-R as the reference standard in a hospital setting

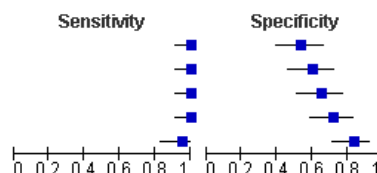
CAM [number of symptoms] [ref: DSMIII-R] patients with dementia

Study	TP	FP	FN	TN	Sensitivity	Specificity
Cole 2003 Dementia >2	128	67	0	27	1.00 [0.97, 1.00]	0.29 [0.20, 0.39]
Cole 2003 Dementia >3	128	52	0	47	1.00 [0.97, 1.00]	0.47 [0.37, 0.58]
Cole 2003 Dementia >4	128	49	0	50	1.00 [0.97, 1.00]	0.51 [0.40, 0.61]
Cole 2003 Dementia >5	128	39	0	60	1.00 [0.97, 1.00]	0.61 [0.50, 0.70]
Cole 2003 Dementia more than 6 smptoms	125	24	3	75	0.98 [0.93, 1.00]	0.76 [0.66, 0.84]



CAM [number of symptoms] [ref: DSMIII-R] no dementia

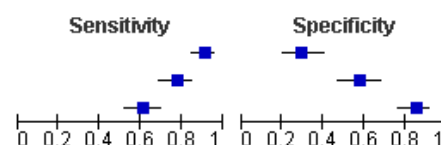
Study	TP	FP	FN	TN	Sensitivity	Specificity
Cole 2003 No dementia >2	40	28	0	32	1.00 [0.91, 1.00]	0.53 [0.40, 0.66]
Cole 2003 No dementia >3	40	24	0	36	1.00 [0.91, 1.00]	0.60 [0.47, 0.72]
Cole 2003 No dementia >4	40	21	0	39	1.00 [0.91, 1.00]	0.65 [0.52, 0.77]
Cole 2003 No dementia >5	40	17	0	43	1.00 [0.91, 1.00]	0.72 [0.59, 0.83]
Cole 2003 No dementia >6sy	38	10	2	50	0.95 [0.83, 0.99]	0.83 [0.71, 0.92]



NB: > indicates 'more than'; '>2' would indicate 'more than 2 symptoms'

DI- dementia

Study	TP	FP	FN	TN	Sensitivity	Specificity
Cole 2003 Dementia >2	116	66	12	28	0.91 [0.84, 0.95]	0.30 [0.21, 0.40]
Cole 2003 Dementia >3	99	40	29	54	0.77 [0.69, 0.84]	0.57 [0.47, 0.68]
Cole 2003 Dementia >4	78	14	50	80	0.61 [0.52, 0.69]	0.85 [0.76, 0.92]



DI- no dementia

Study	TP	FP	FN	TN	Sensitivity	Specificity
Cole 2003 No dementia >2	33	22	7	38	0.82 [0.67, 0.93]	0.63 [0.50, 0.75]
Cole 2003 No dementia >3	24	11	16	49	0.60 [0.43, 0.75]	0.82 [0.70, 0.90]
Cole 2003 No dementia >4	17	5	23	55	0.42 [0.27, 0.59]	0.92 [0.82, 0.97]

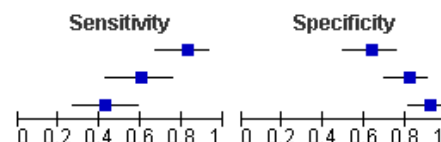


Figure 6.12: ROC plot of effects of varying threshold for CAM and DI compared with DSM-III-R

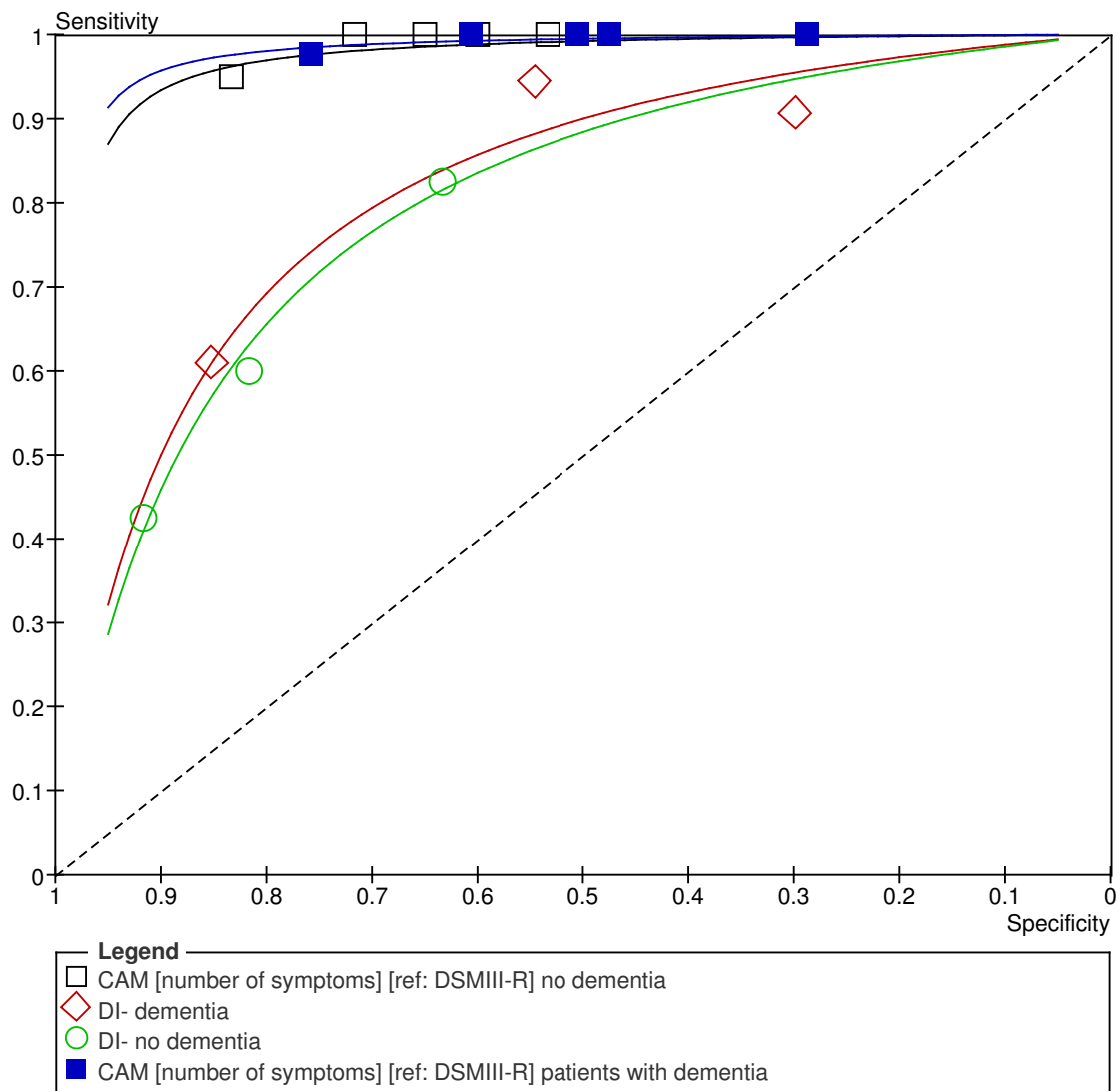
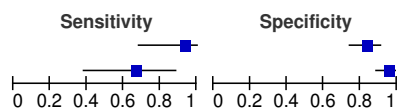


Figure 6.13: forest plot of index tests with DSM-III as the reference standard in a hospital setting

AMT [ref: DSMIII]- decline in score 2 points & 3 points

Study	TP	FP	FN	TN	Sensitivity	Specificity
Ni Chonchubhair 1995-2 pt	14	14	1	71	0.93 [0.68, 1.00]	0.84 [0.74, 0.91]
Ni Chonchubhair 1995-3pt	10	4	5	81	0.67 [0.38, 0.88]	0.95 [0.88, 0.99]



CAM [ref: DSM III]

Study	TP	FP	FN	TN	Sensitivity	Specificity
Laurila 2002	17	17	3	44	0.85 [0.62, 0.97]	0.72 [0.59, 0.83]

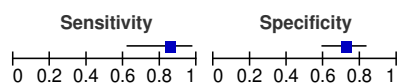


Figure 6.14: forest plot of index test compared with consensus diagnosis as the reference standard in a hospital setting

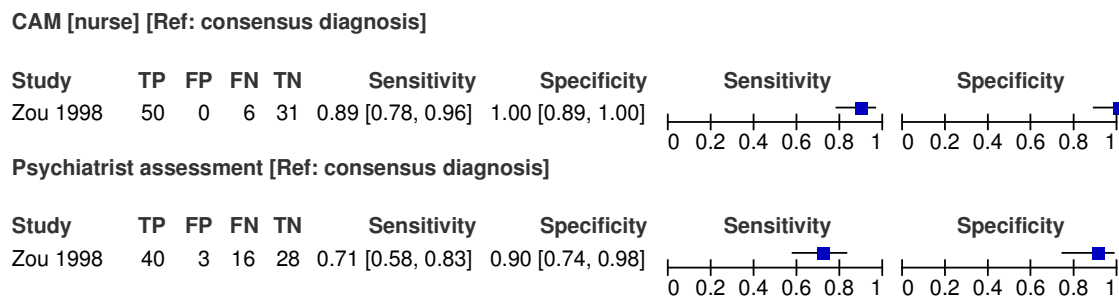


Figure 6.15: forest plot of index test compared with CAM (short version) and clinical interview as the reference standard

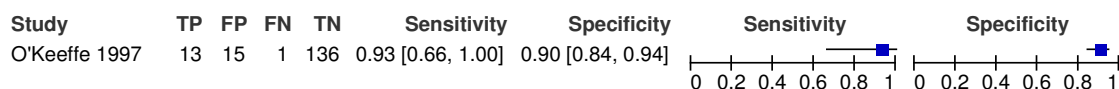


Figure 6.16: forest plot CAM (lay person) compared with CAM (geriatrician) - subgroup analyses

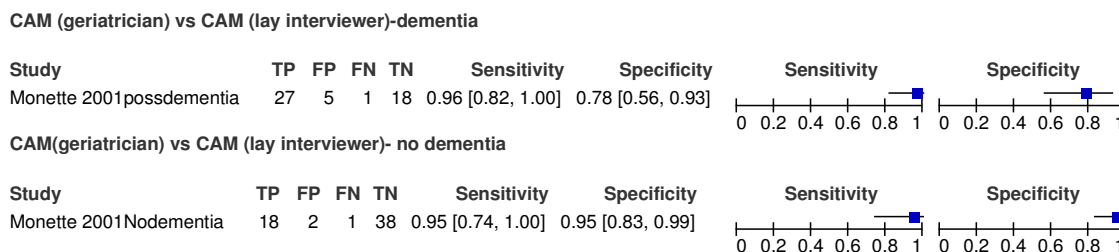


Figure 6.17: forest plot of CAM-ICU index test with DSM-IV as reference standard in an ICU setting

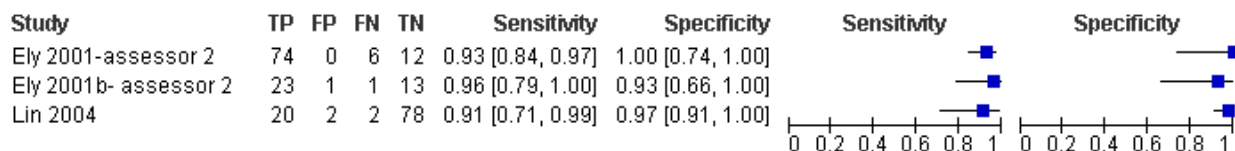
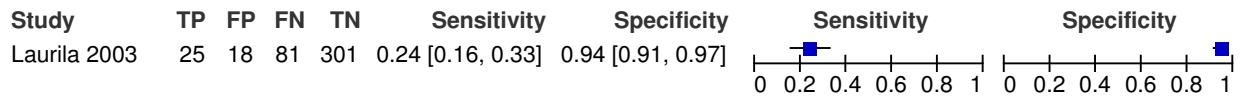
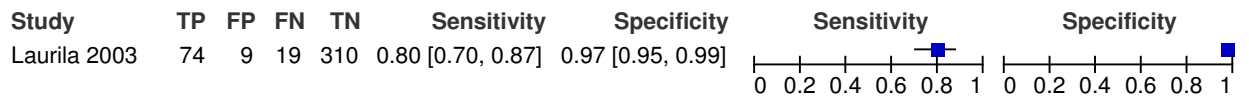


Figure 6.18: forest plot of ICD-10, DSM-III-R and DSM-III compared with DSM-IV; mixed setting (hospital and long-term care)

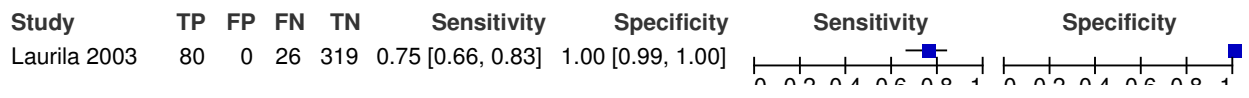
ICD10 [ref:DSM-IV]



DSM-III-R [ref: DSMIV]



DSMIII [ref: DSMIV]



Risk factors for delirium: non-pharmacological

Figure 7.1: hospital unit as a risk factor for an increased severity of delirium

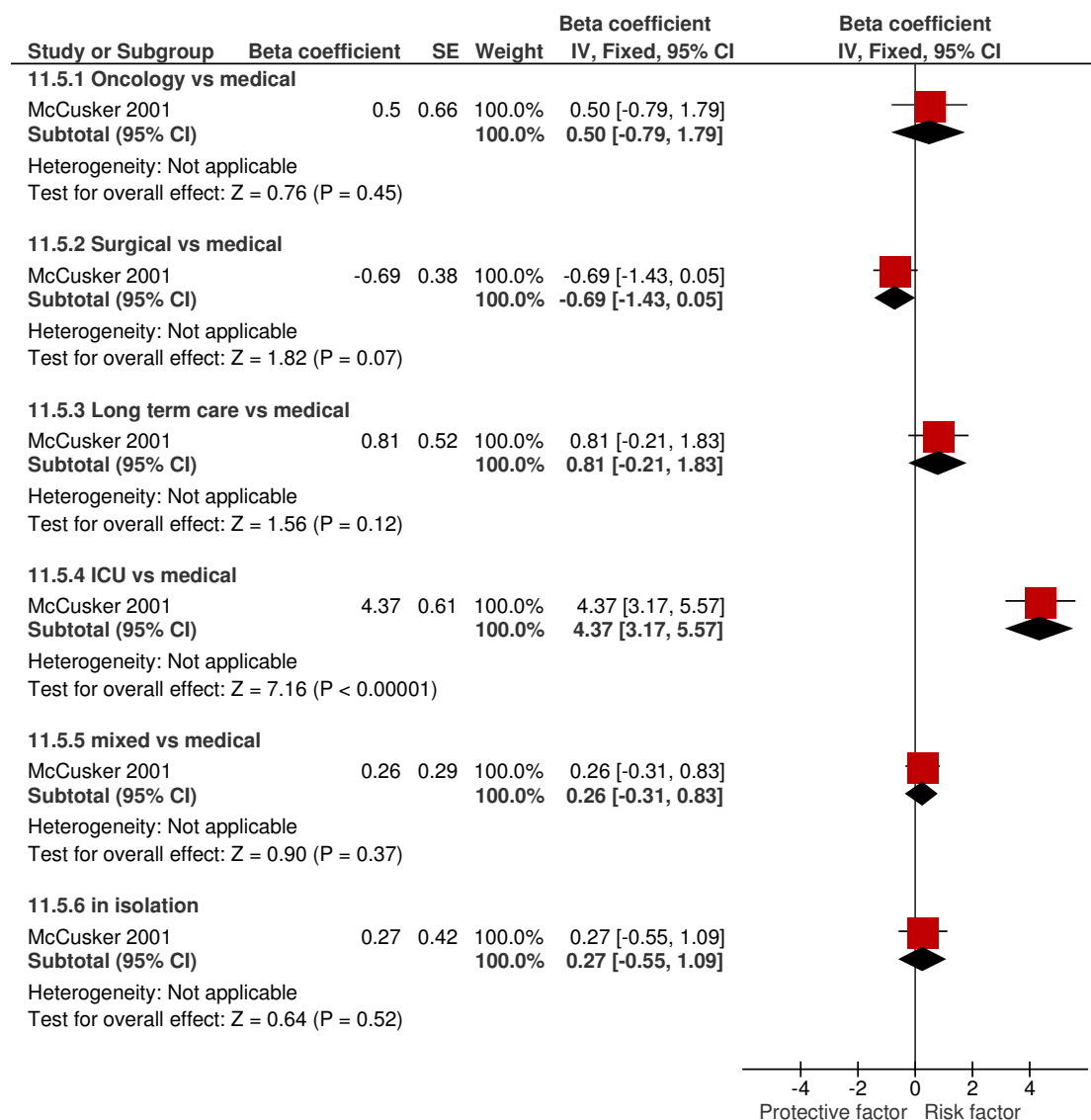


Figure 7.2: age as a risk factor: incidence of delirium

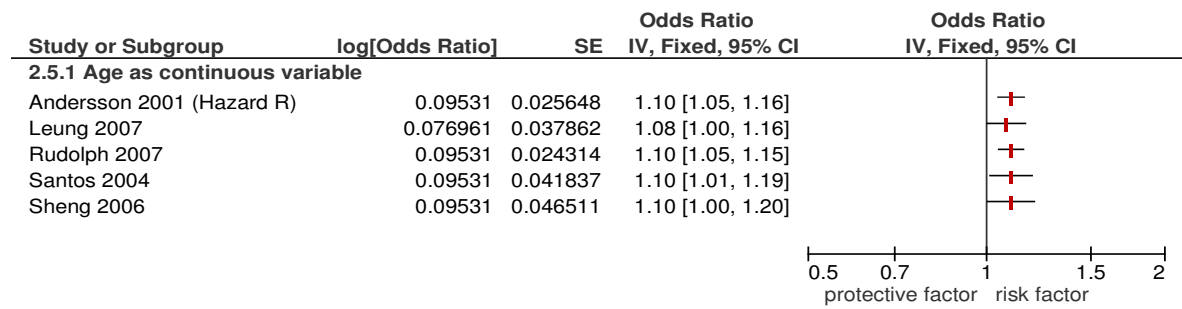


Figure 7.3a: age as a risk factor: incidence of delirium

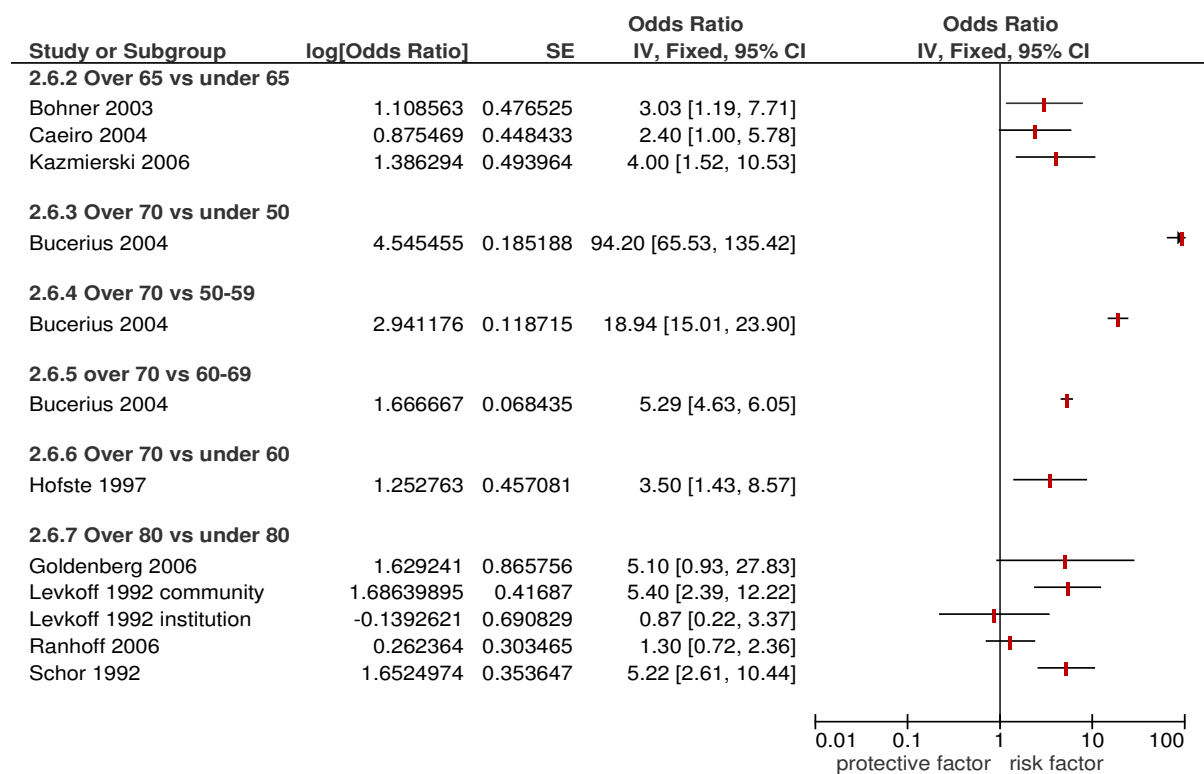


Figure 7.3b: age: incidence of delirium excluding studies with a low quality rating

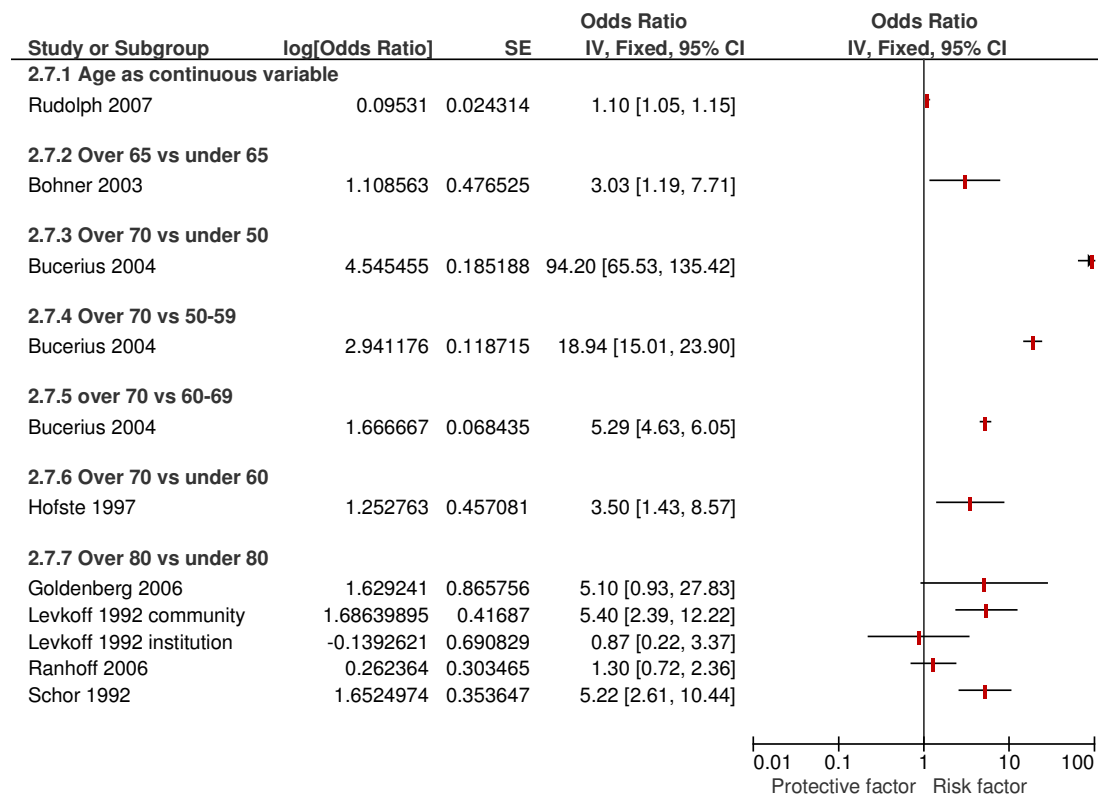


Figure 7.4: age as a risk factor: duration of delirium

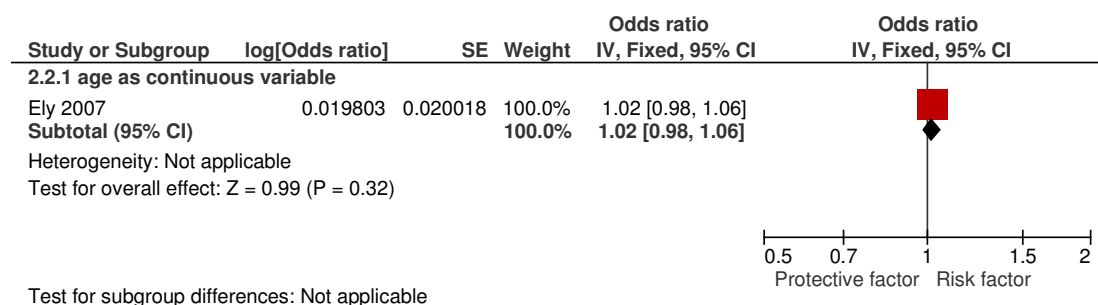


Figure 7.5: patient characteristics as risk factors: severity of delirium

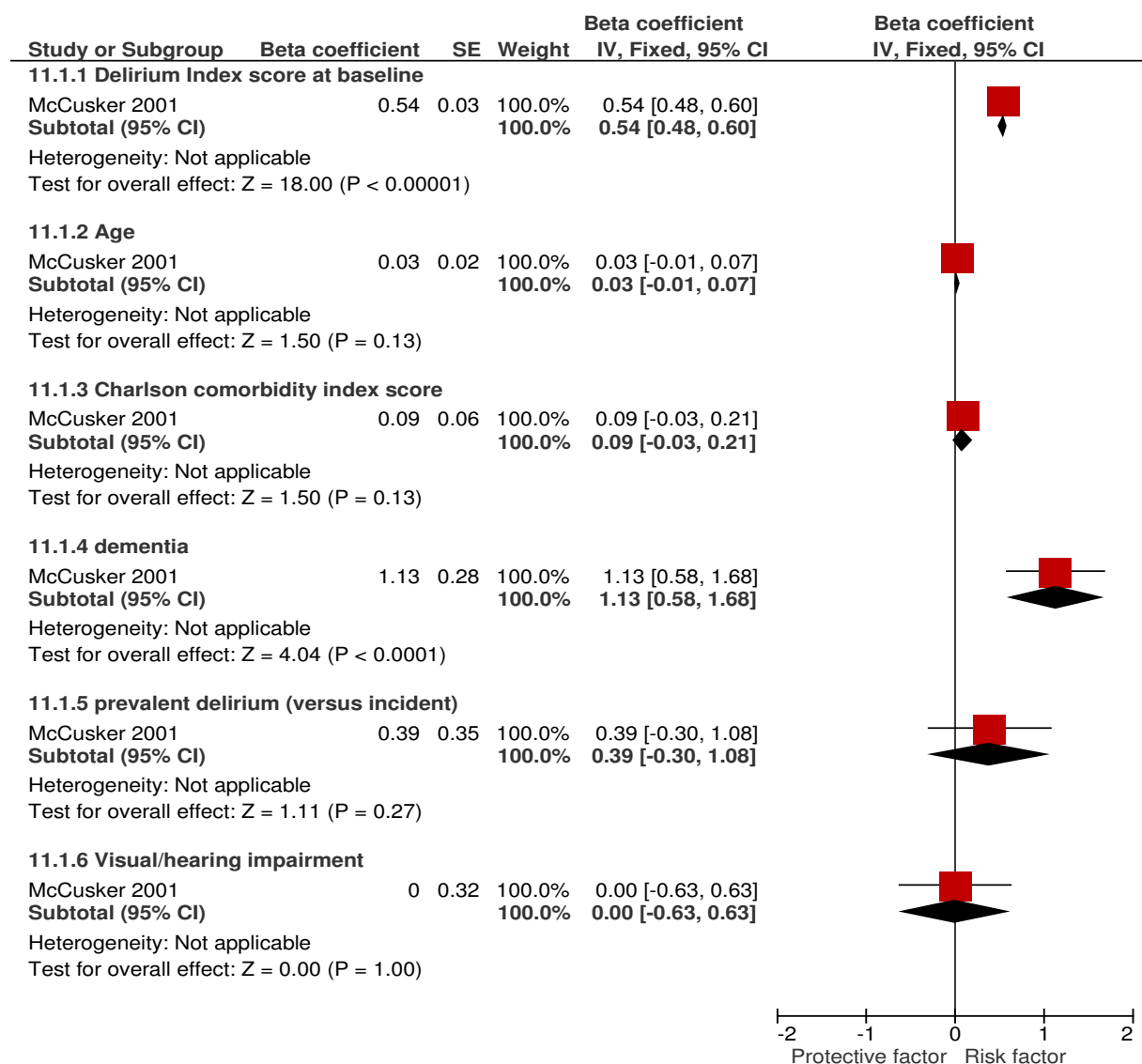


Figure 7.6a: cognitive impairment and/or dementia as a risk factor: incidence of delirium

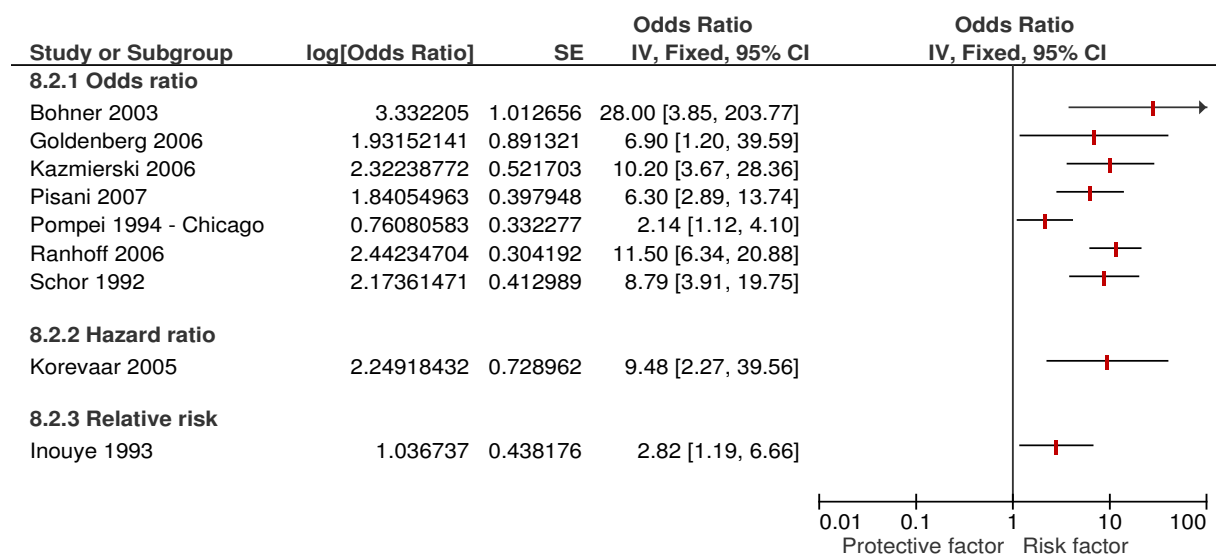


Figure 7.6b: cognitive impairment and/or dementia: incidence of delirium excluding studies with a low quality rating, and also Ranhoff (2006)

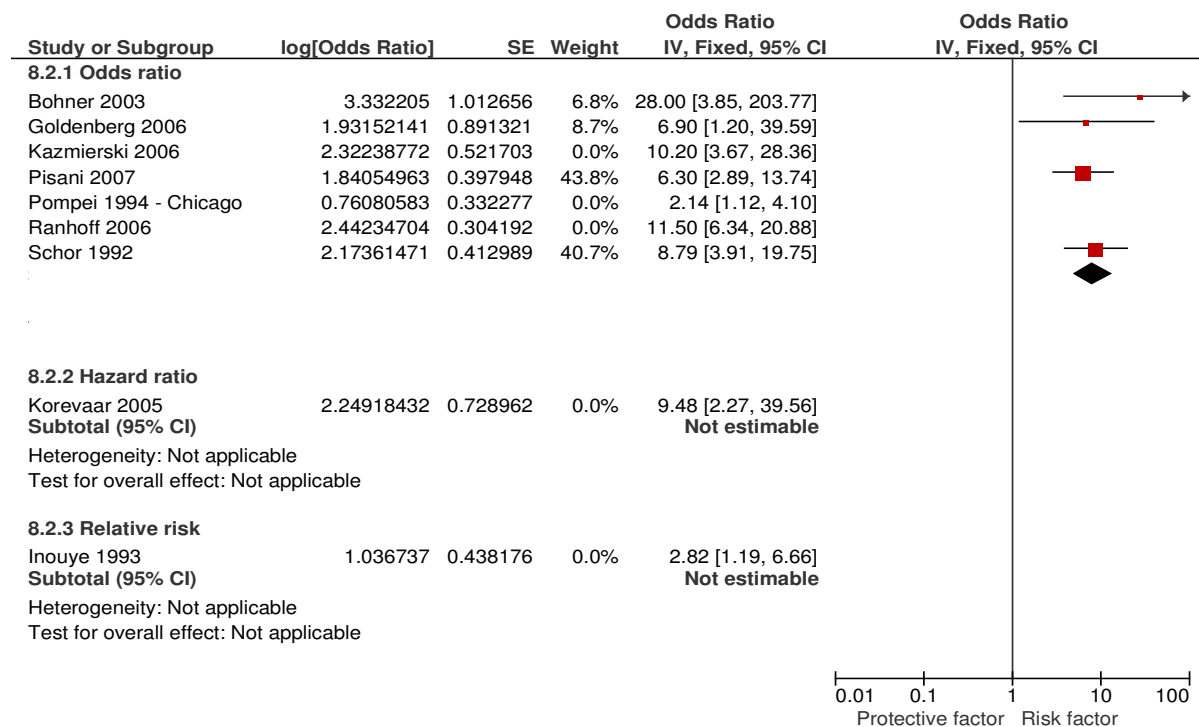


Figure 7.7: cognitive impairment and/or dementia as a risk factor: persistent delirium

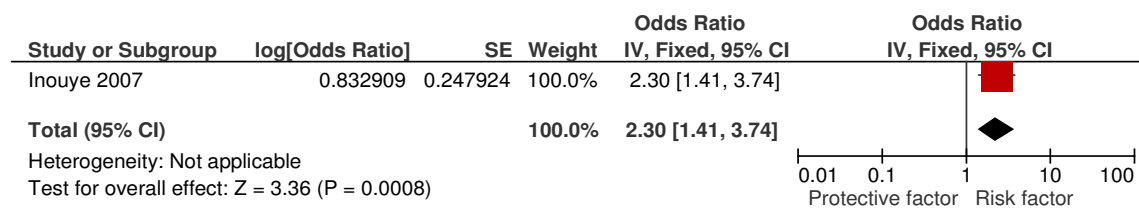


Figure 7.8: impaired vision as a risk factor: incidence of delirium

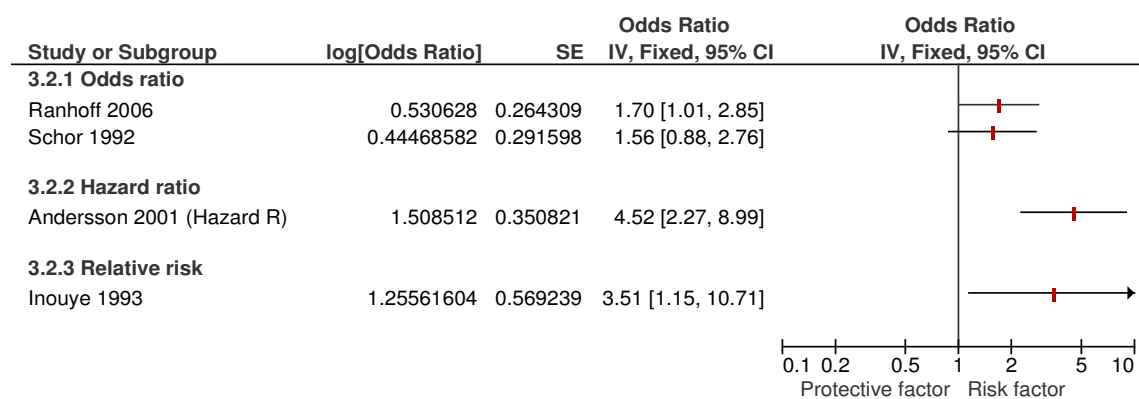


Figure 7.9: impaired vision as a risk factor: persistent delirium

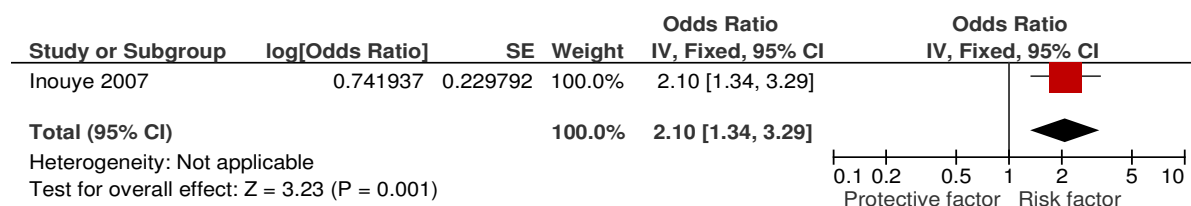


Figure 7.10: polypharmacy: incidence of delirium

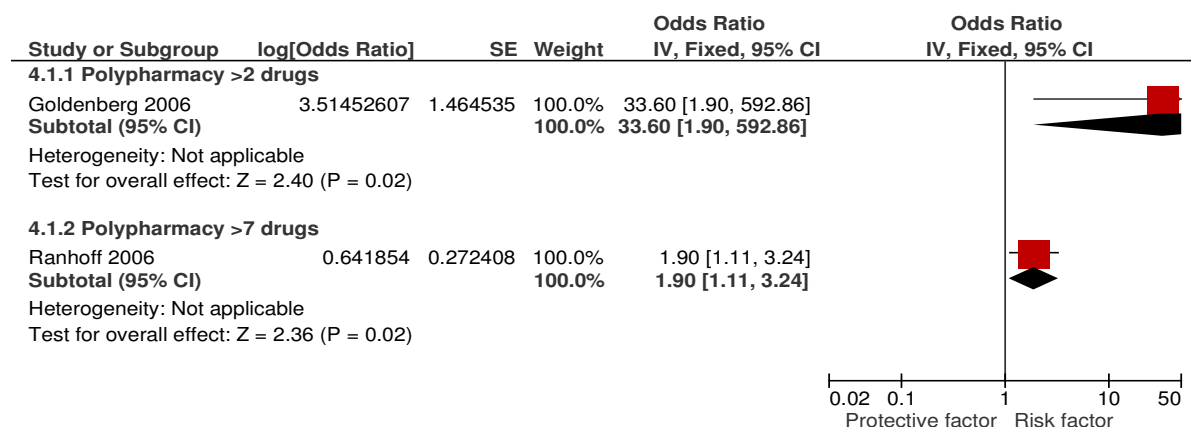


Figure 7.11: dehydration as a risk factor: incidence of delirium

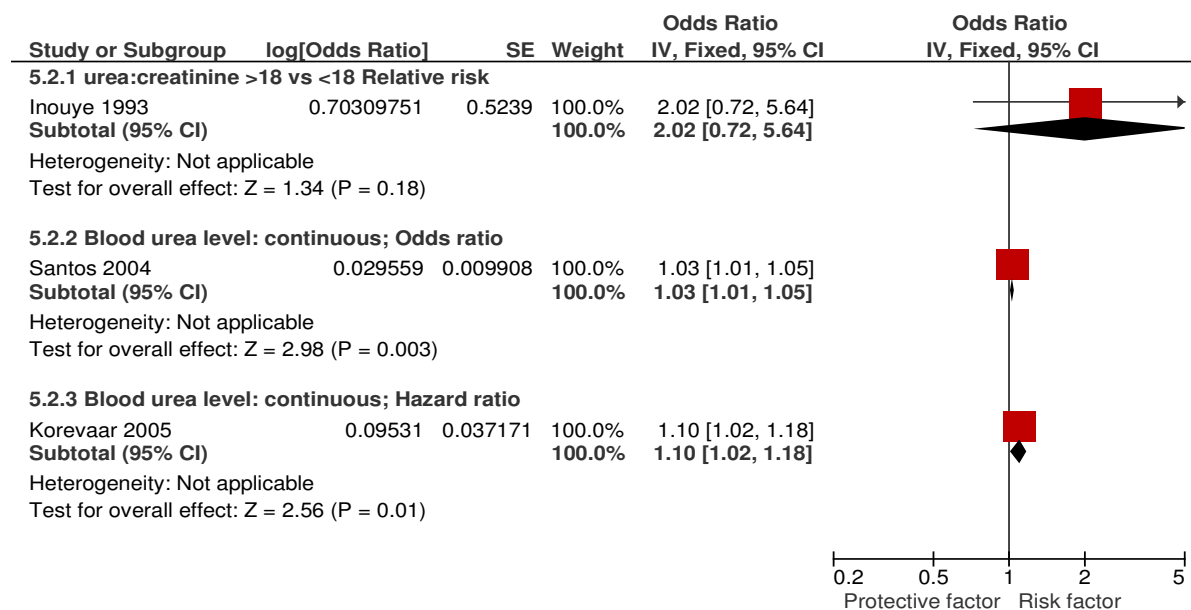


Figure 7.12: illness severity as a risk factor: incidence of delirium

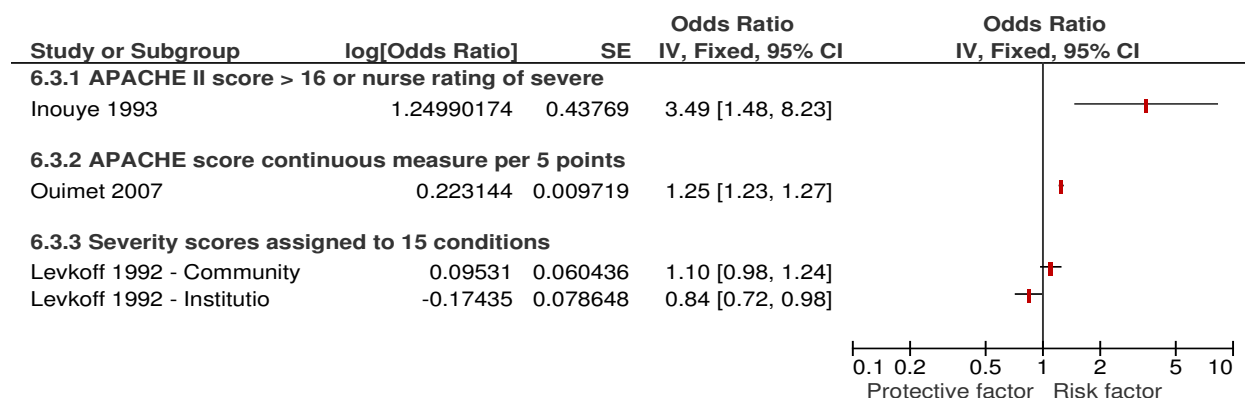


Figure 7.13: illness severity as a risk factor: duration of delirium

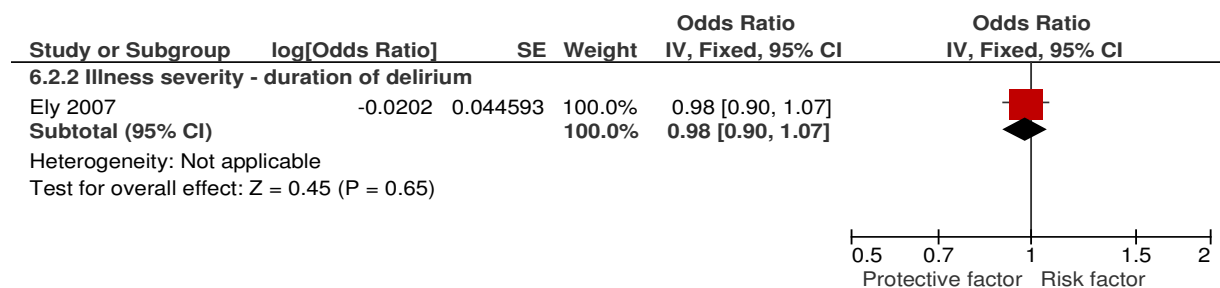


Figure 7.14: comorbidity as a risk factor: incidence of delirium

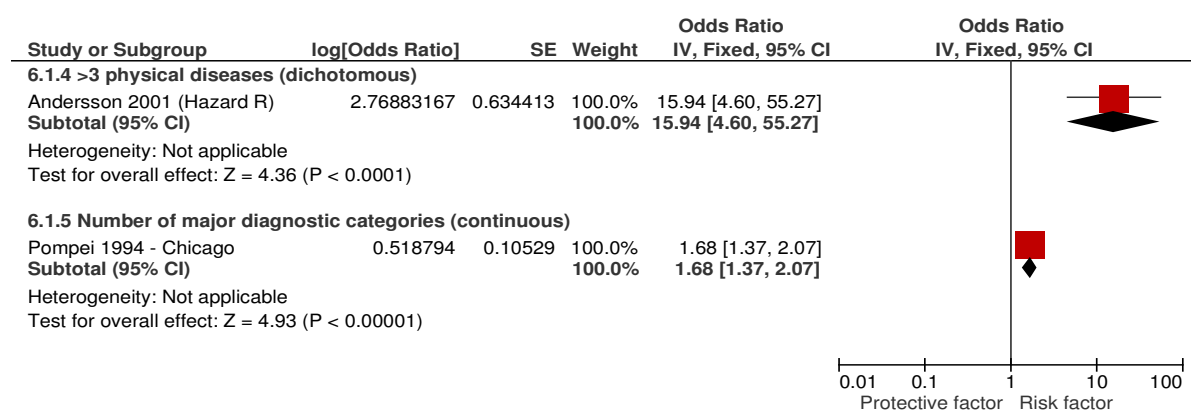


Figure 7.15: comorbidity as a risk factor: persistent delirium

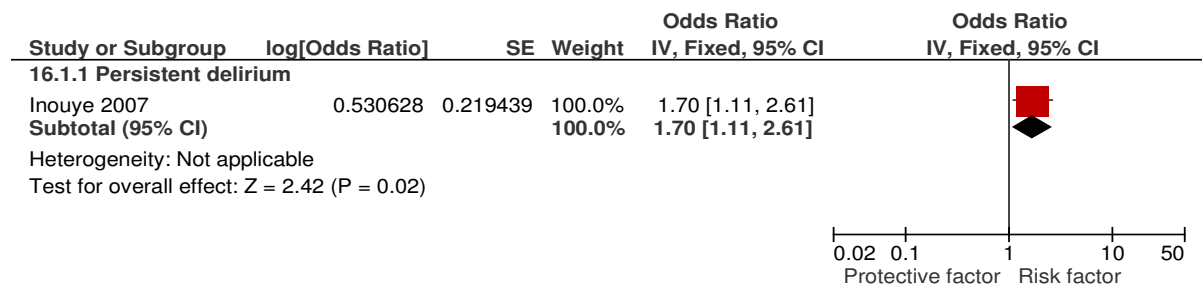


Figure 7.16: sex (male) as a risk factor: incidence of delirium

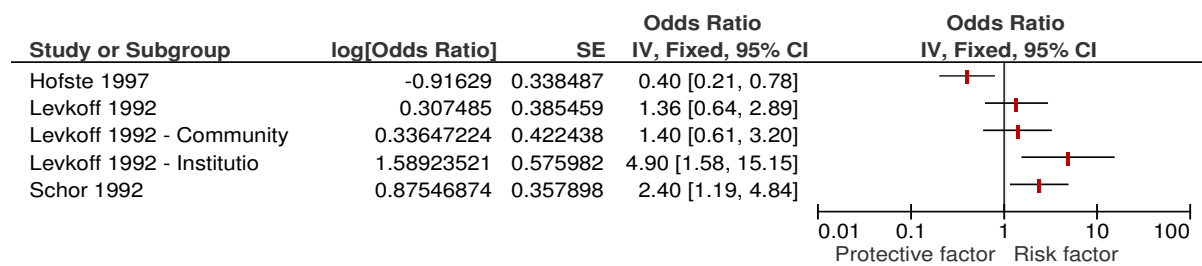


Figure 7.17: electrolyte disturbance as a risk factor: incidence of delirium

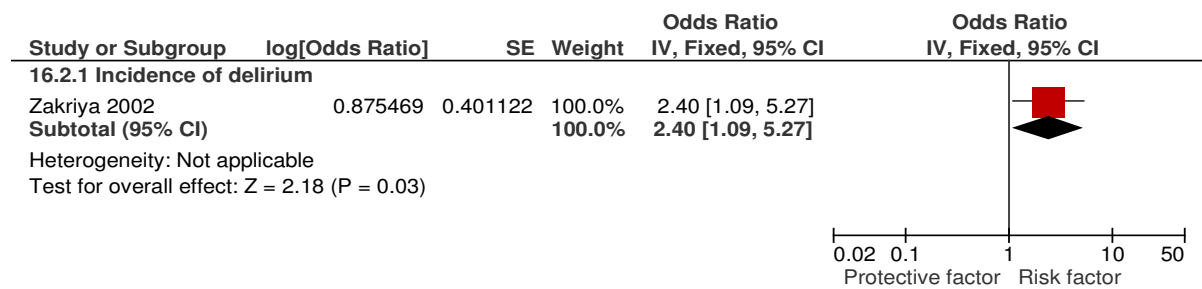


Figure 7.18: depression as a risk factor: incidence of delirium

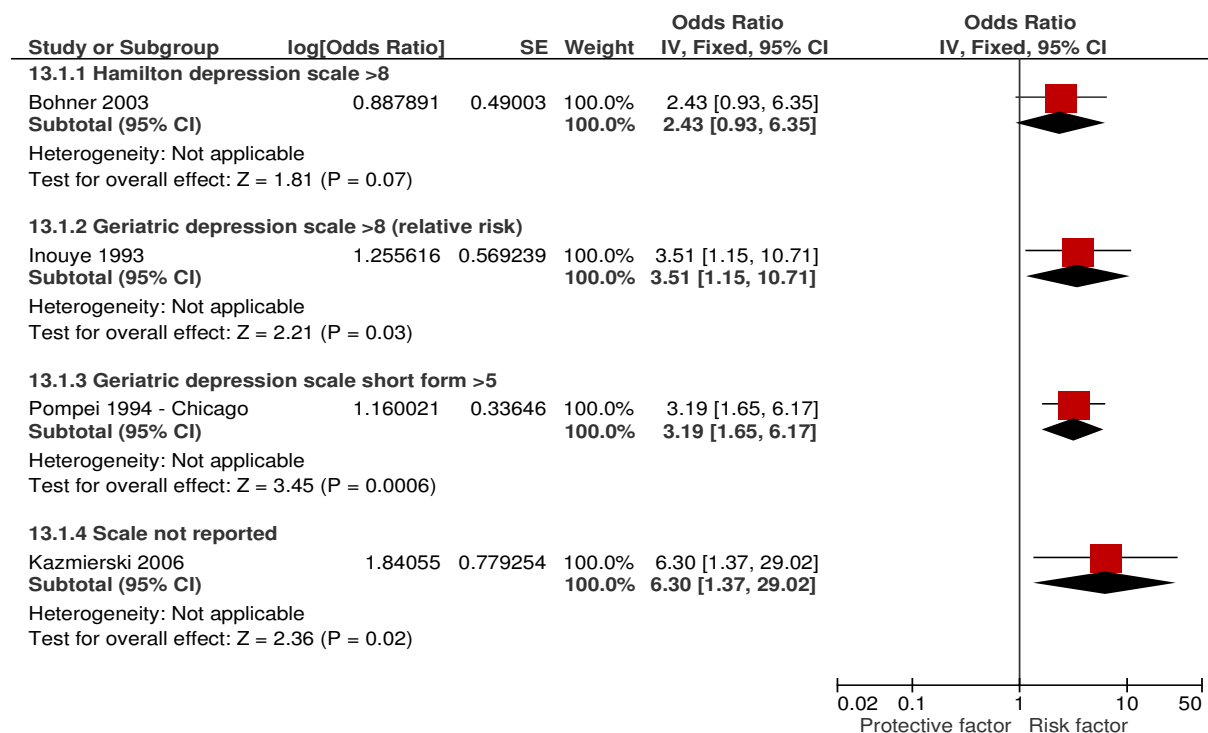


Figure 7.19: infection as a risk factor: incidence of delirium

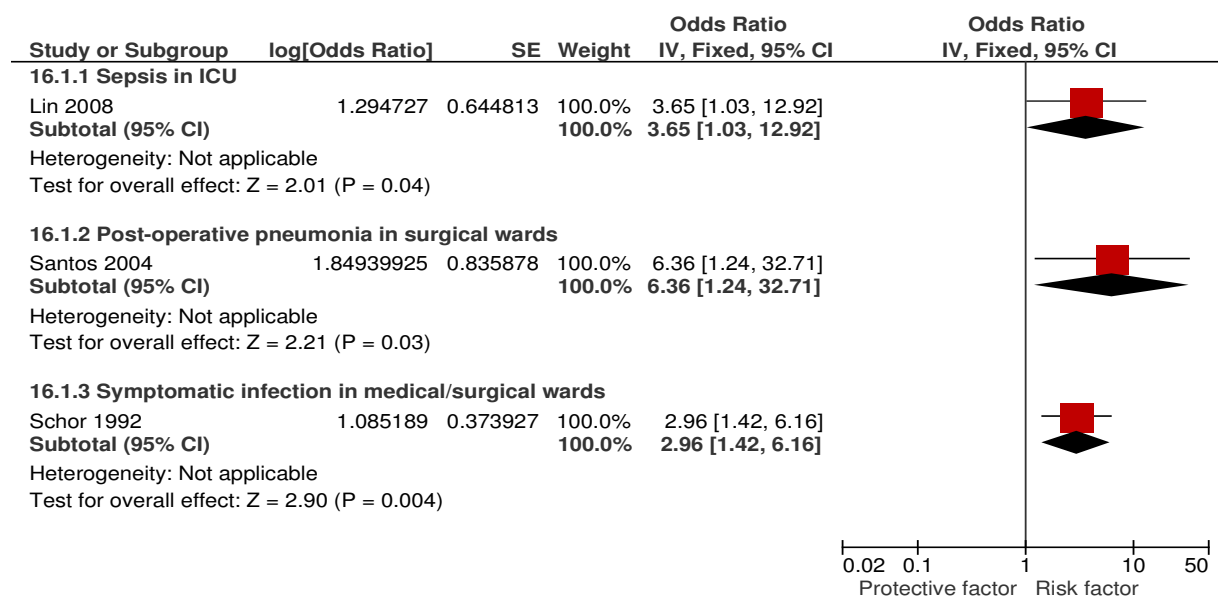


Figure 7.20: infection as a risk factor: duration of delirium

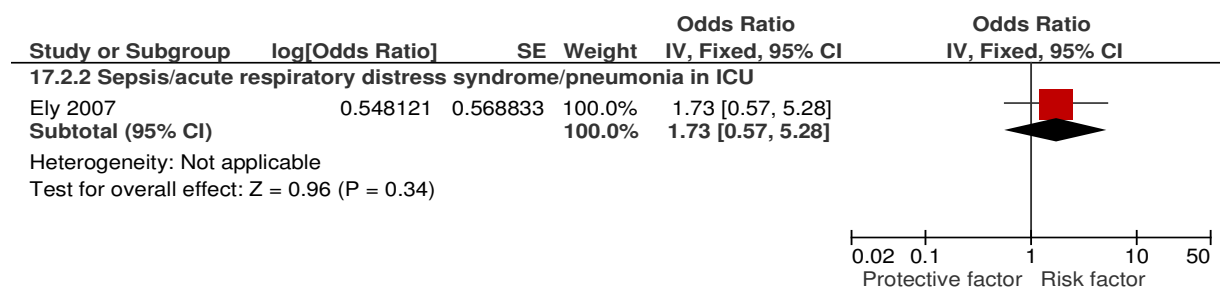


Figure 7.21: fracture on admission as a risk factor

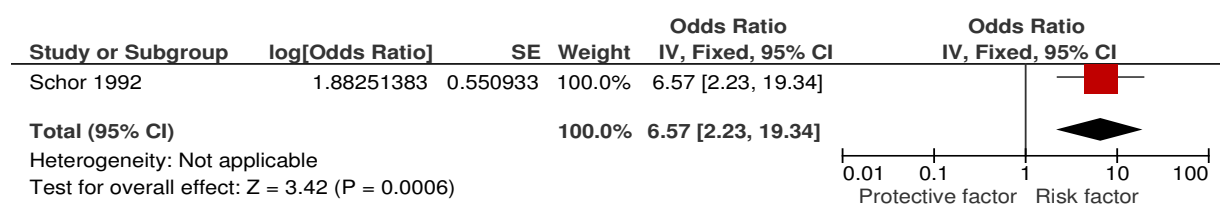


Figure 7.22: environmental risk factors: severity of delirium

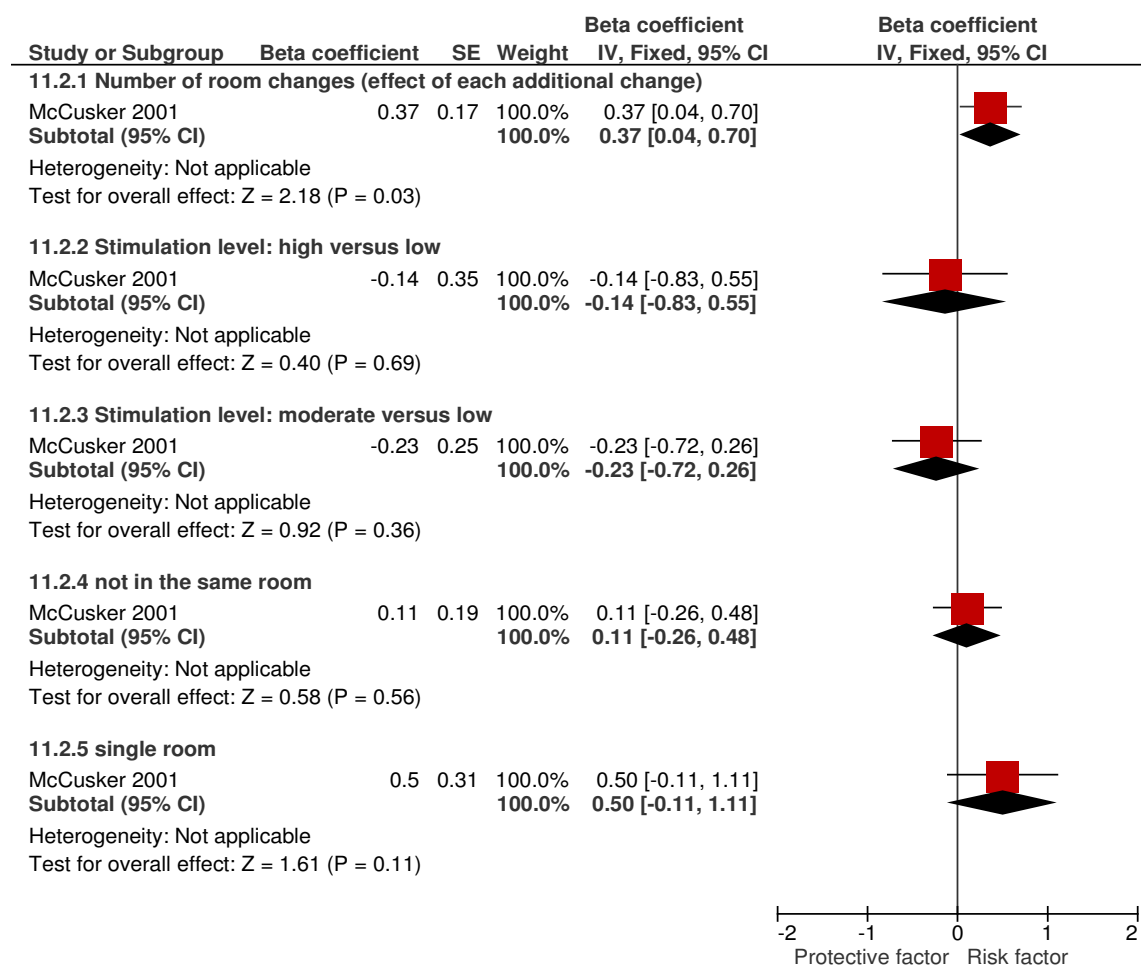
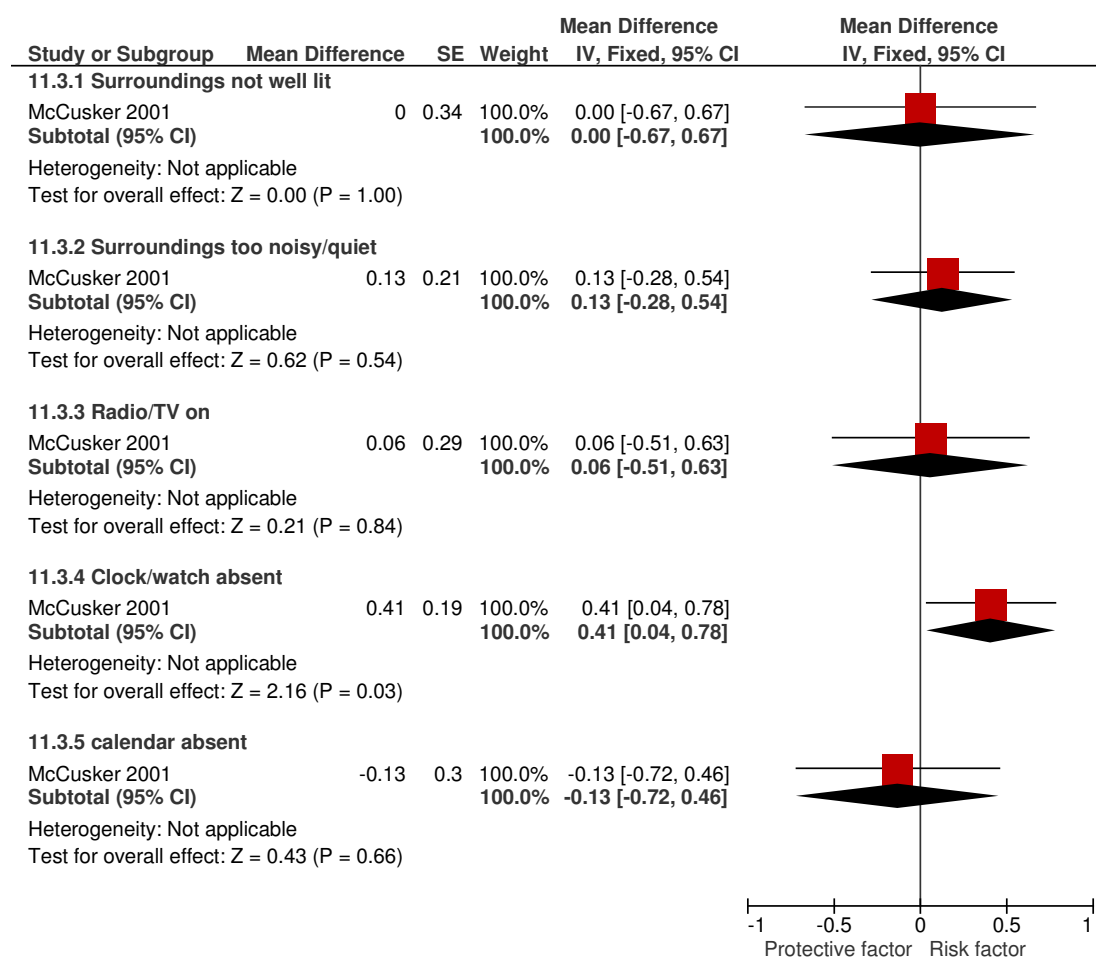


Figure 7.23: environmental risk factors: severity of delirium



NB scale -1 to +1

Figure 7.24: environmental risk factors: severity of delirium

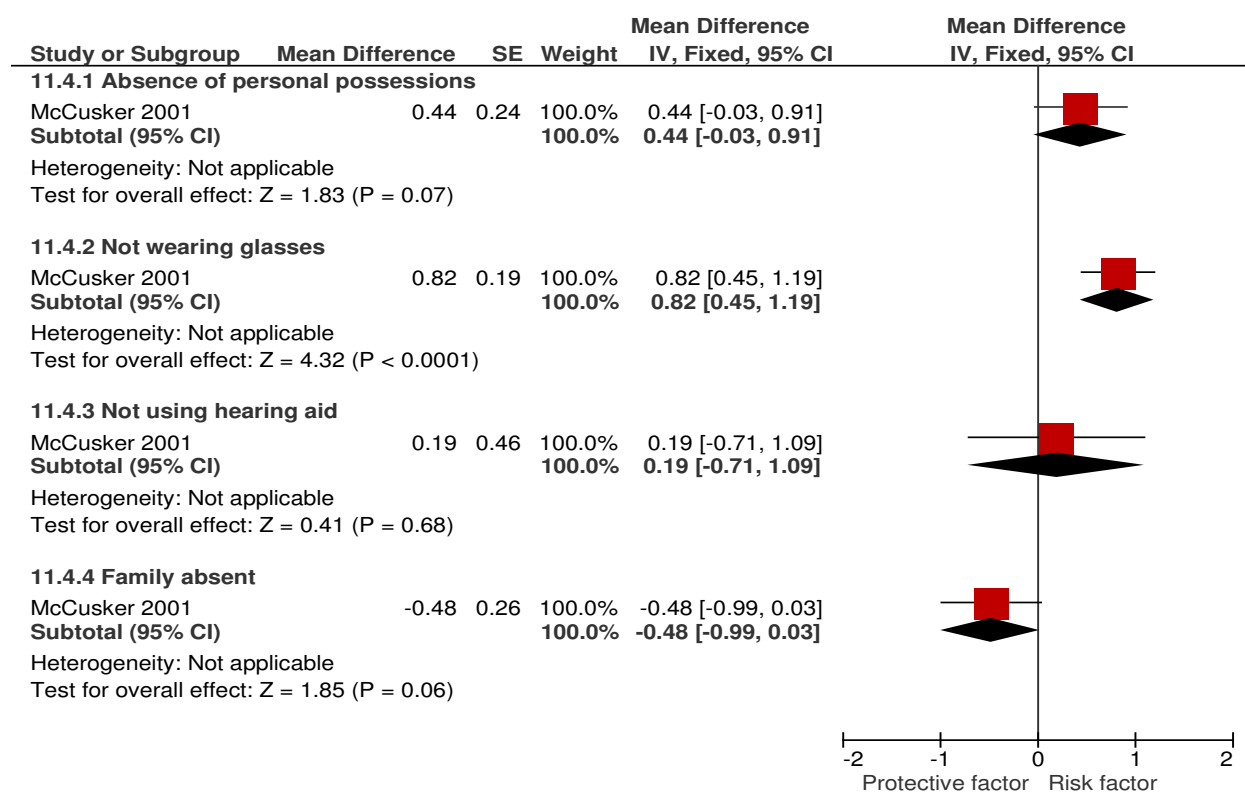


Figure 7.25: cardiac surgery risk factors: incidence of delirium

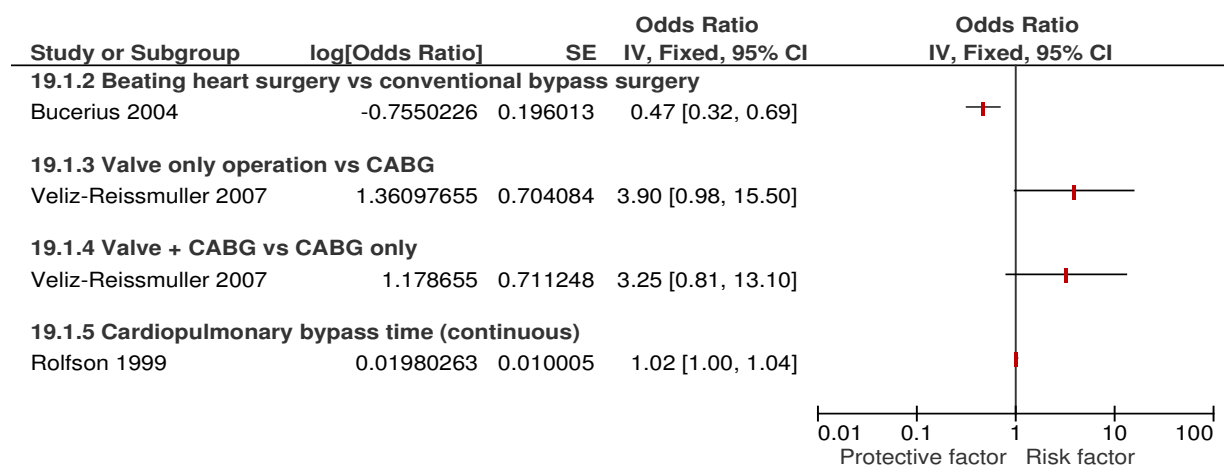


Figure 7.26: type of surgery a risk factor: incidence of delirium

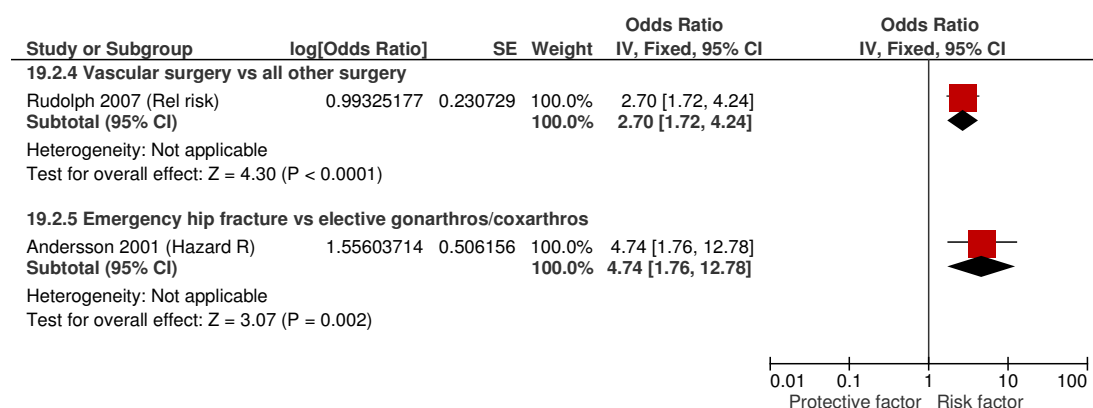


Figure 7.27: iatrogenic intervention as a risk factor: incidence of delirium

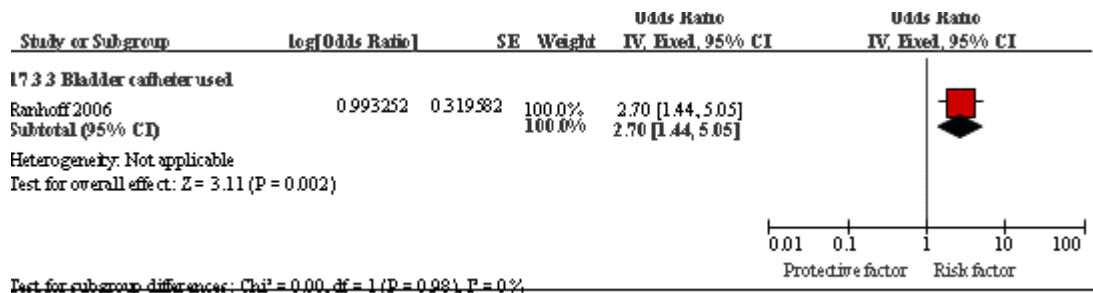


Figure 7.28: physical restraint during delirium: persistent delirium

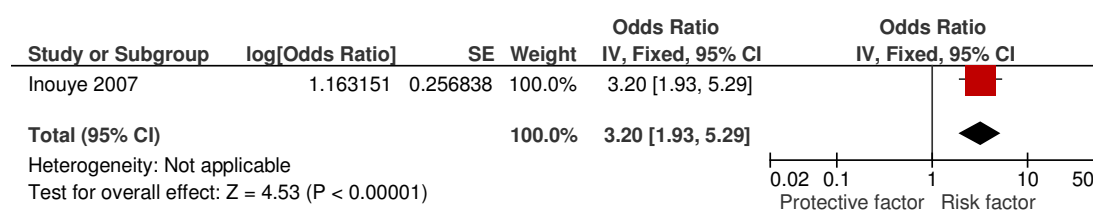
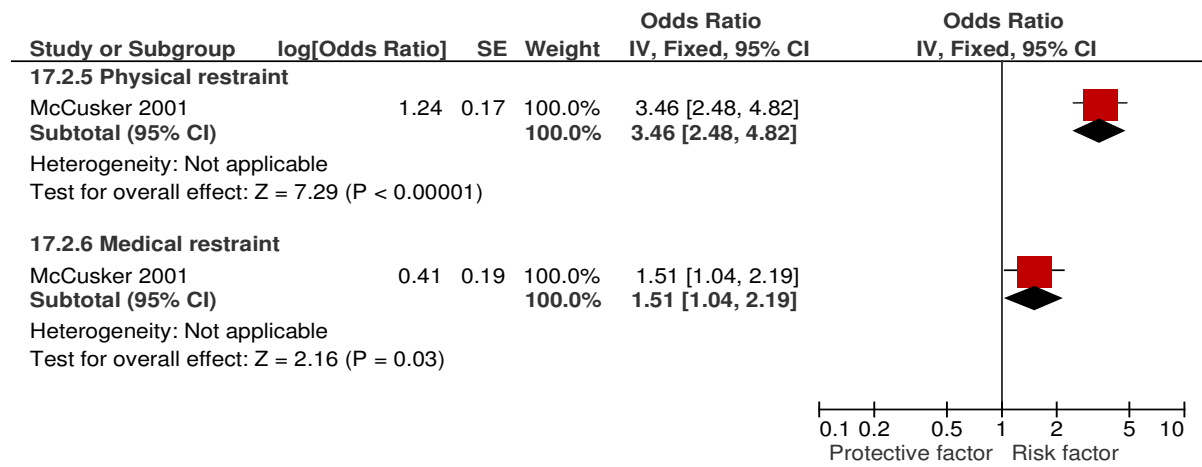


Figure 7.29: physical and medical restraint as a risk factor for the severity of delirium



Risk factors for delirium: pharmacological agents

Figure 8.1: Midazolam as a risk factor for development of delirium

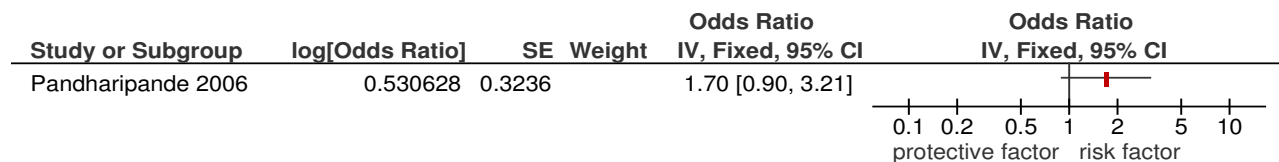


Figure 8.2: lorazepam as a risk factor for development of delirium

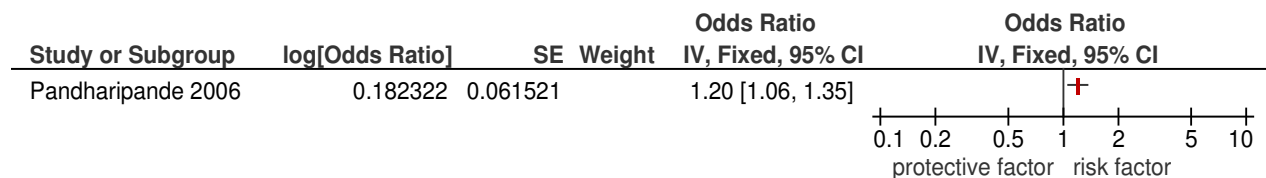


Figure 8.3: benzodiazepines as a risk factor for delirium

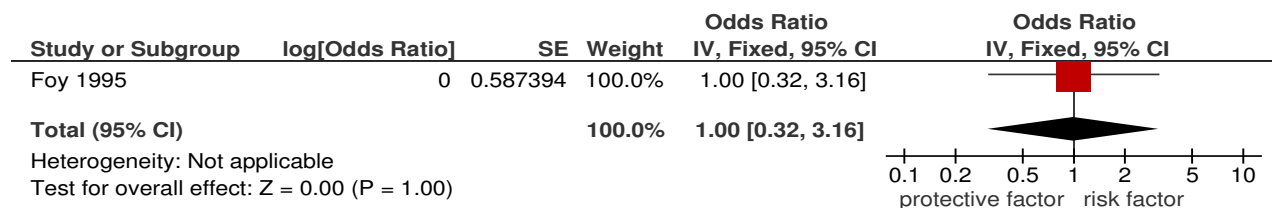
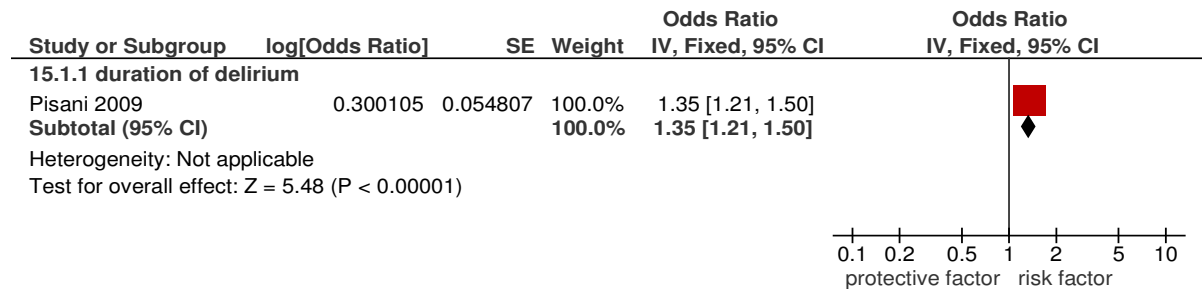


Figure 8.4: Haloperidol as a risk factor for duration of delirium



NB: Scale 0.1 to 10

Figure 8.5: antihistamines with anticholinergic activity

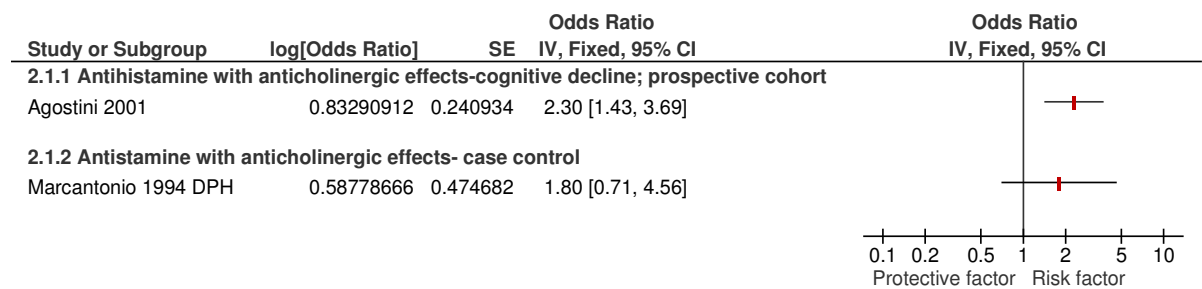


Figure 8.6: exposure to H2 blockers on the incidence of delirium

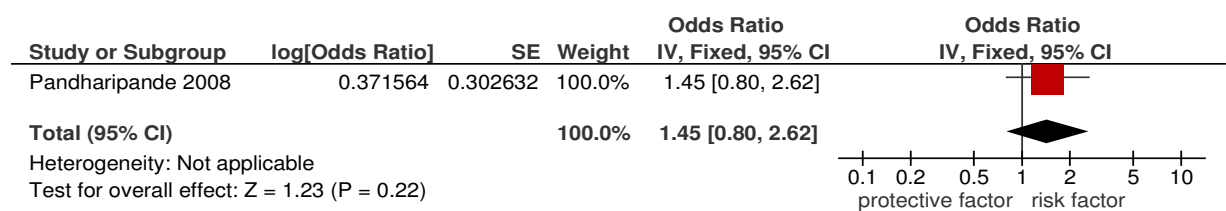
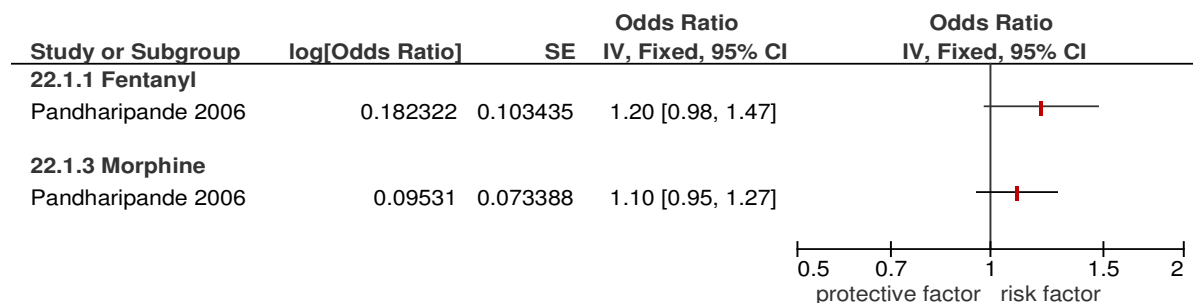


Figure 8.7a: effect of individual opioids on delirium



NB: Scale 0.5 to 2

Figure 8.7b: effect of individual opioids on delirium

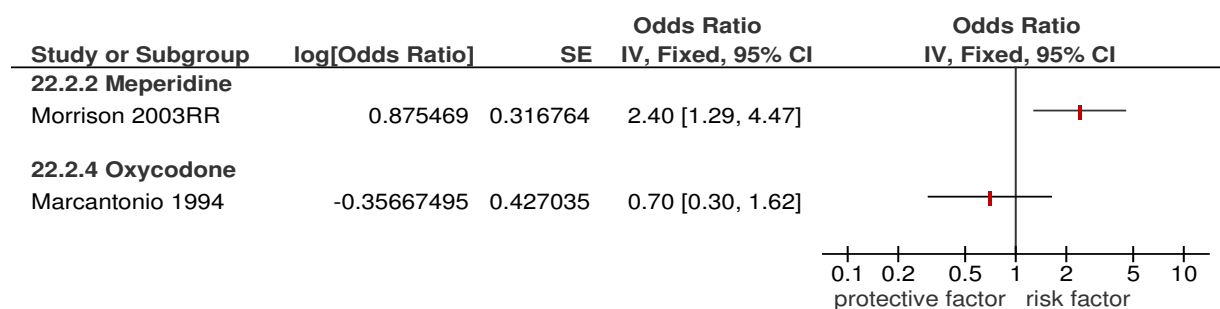


Figure 8.8: effect of opioids on the incidence of delirium

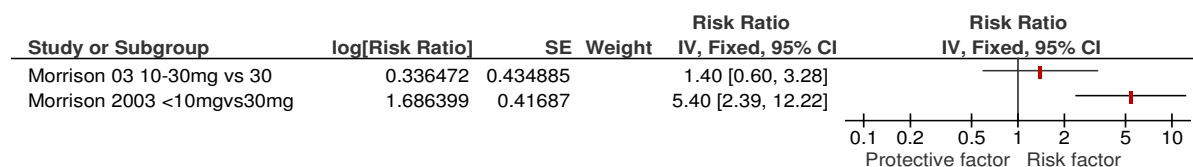


Figure 8.9: effect of intrathecal morphine + PCA morphine versus placebo + PCA morphine

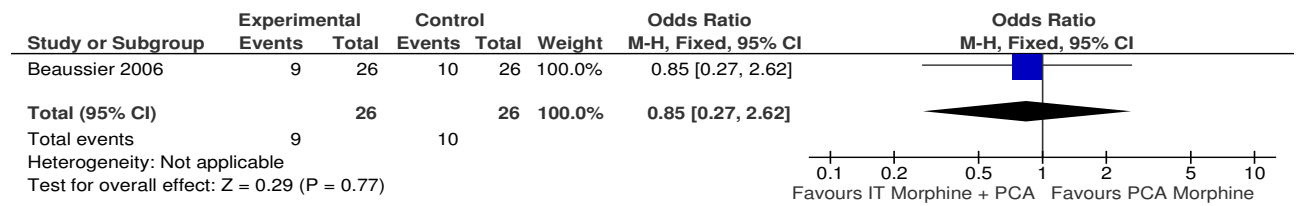


Figure 8.10: effect of PCA opioid analgesics versus oral opioids

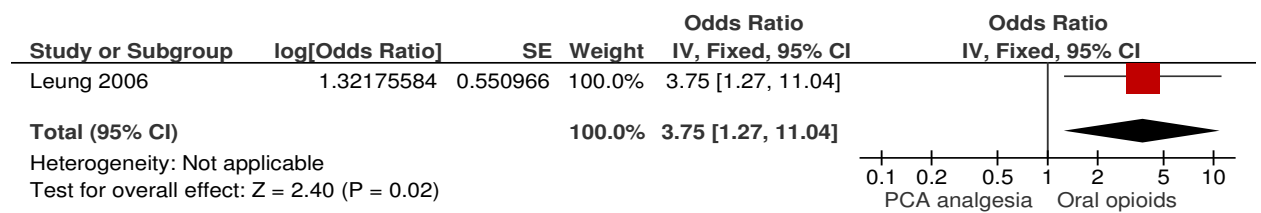


Figure 8.11: effect of general anaesthesia versus regional anaesthesia on delirium

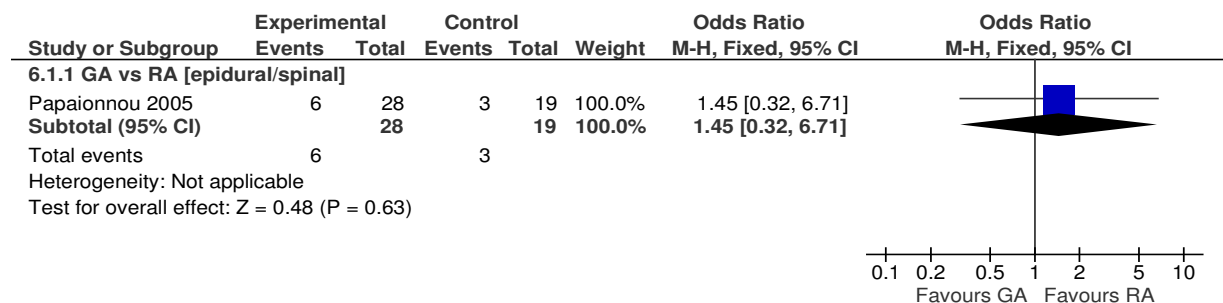


Figure 8.12: effect of N₂O plus O₂ versus O₂ on delirium

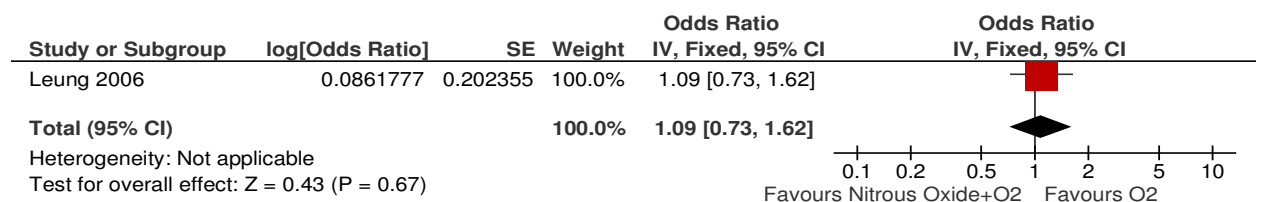
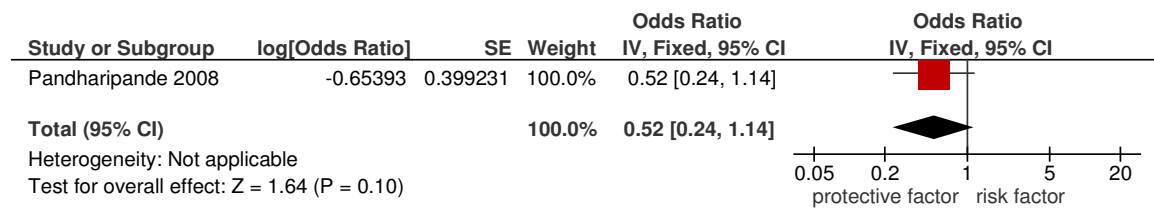
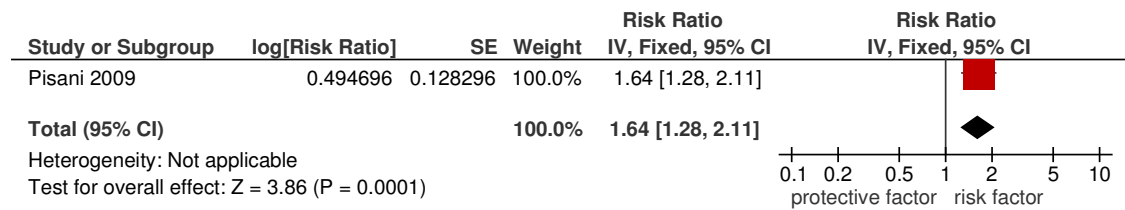


Figure 8.13: effect of anaesthetics on delirium



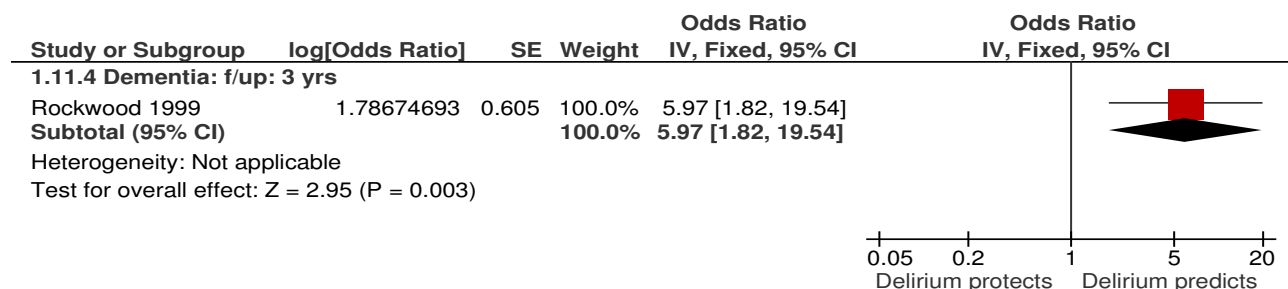
NB: Scale 0.05 to 20

Figure 8.14: effect of benzodiazepines or opioids on the duration of delirium



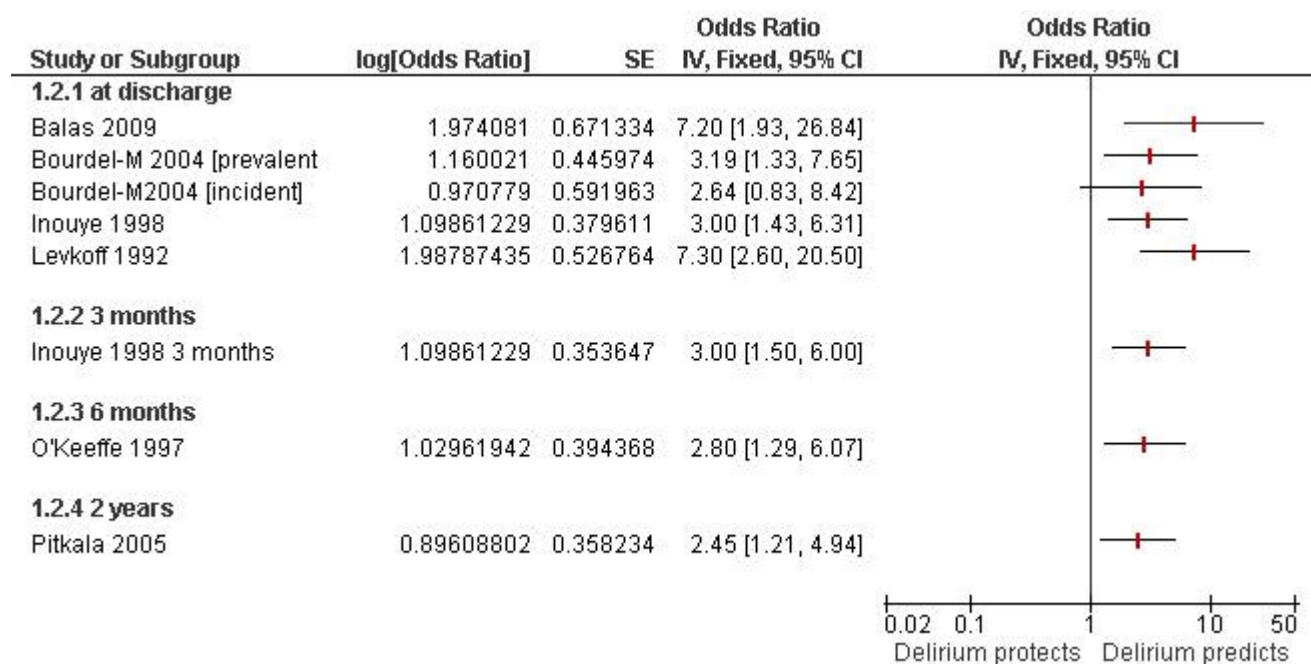
Consequences of delirium

Figure 9.1: dementia as a consequence of delirium



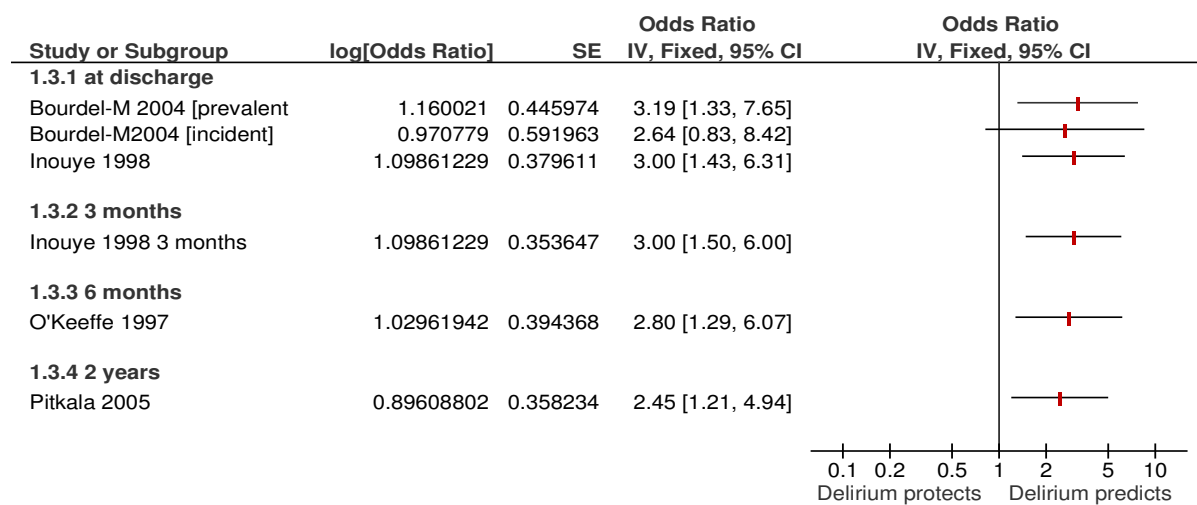
NB: Scale 0.05 to 20

Figure 9.2a: new admission to institution as a consequence of delirium



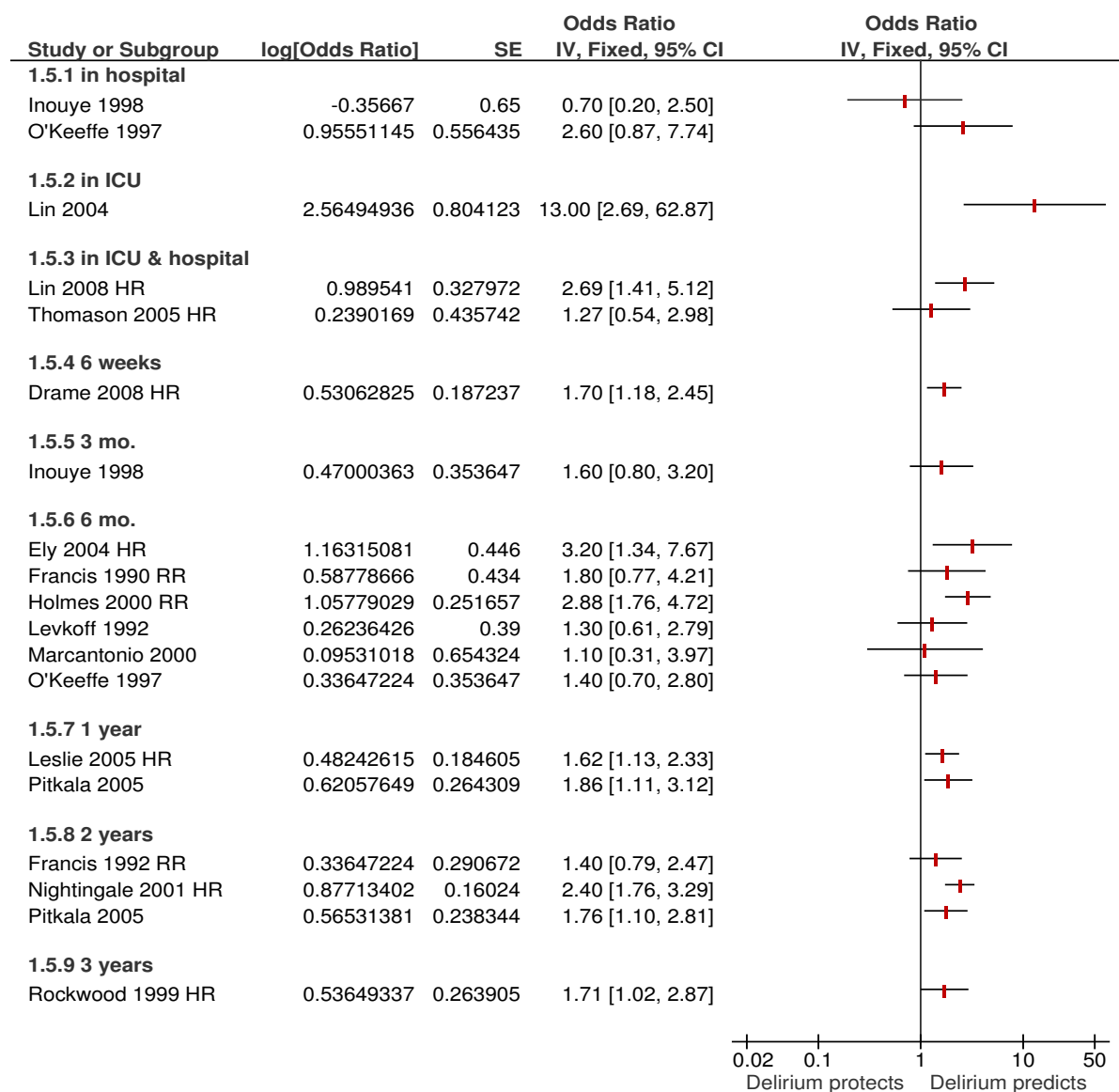
NB: Scale 0.05 to 20

Figure 9.2b: new admission to institution [moderate quality studies]



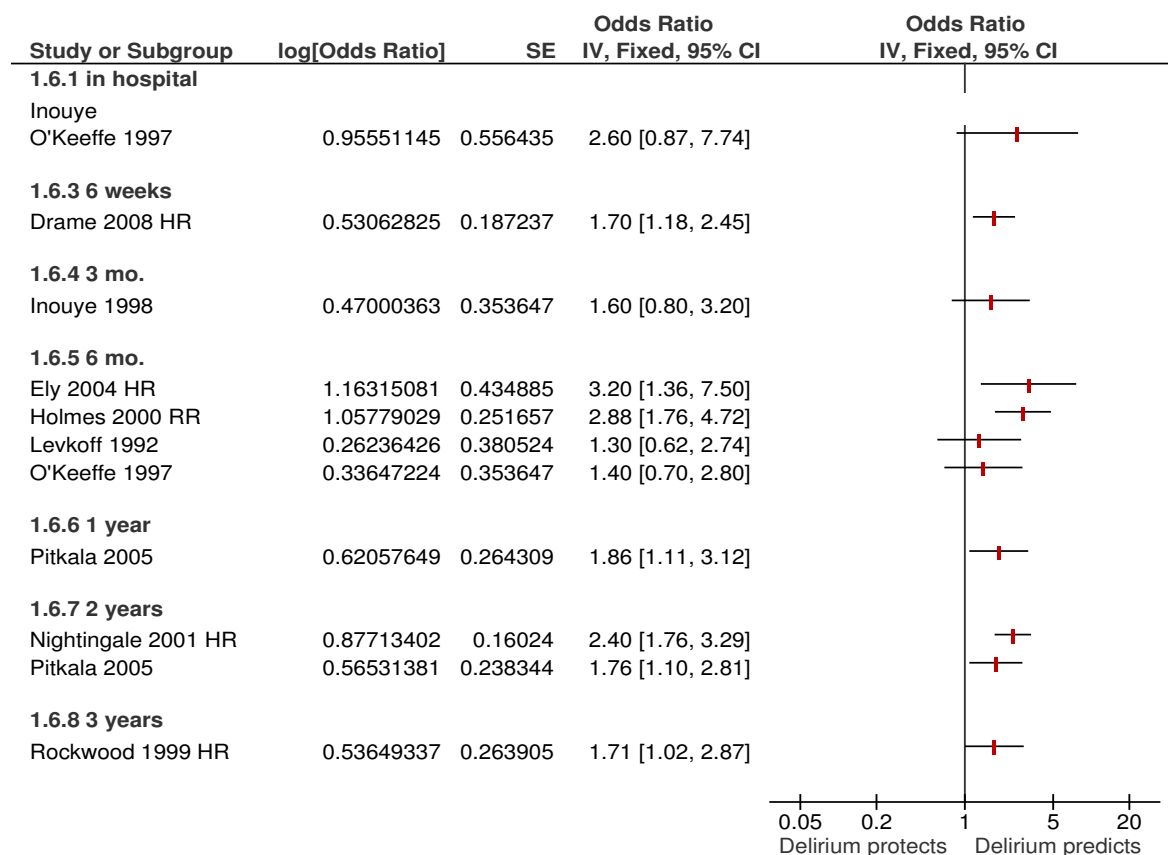
NB: Scale 0.1 to 10

Figure 9.3a: mortality as a consequence of delirium



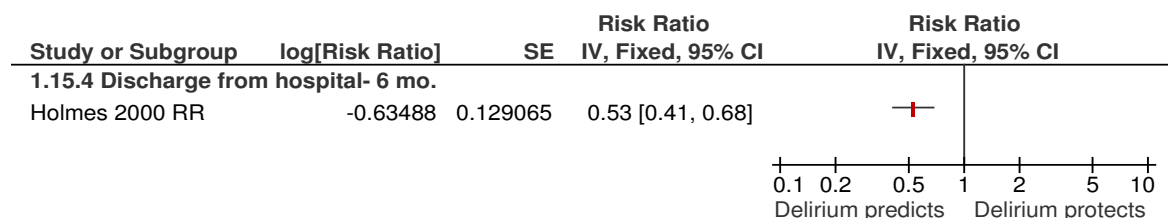
NB: Scale 0.02 to 50

Figure 9.3b: mortality as a consequence of delirium; high and moderate quality studies and restricting to the UK hospital study



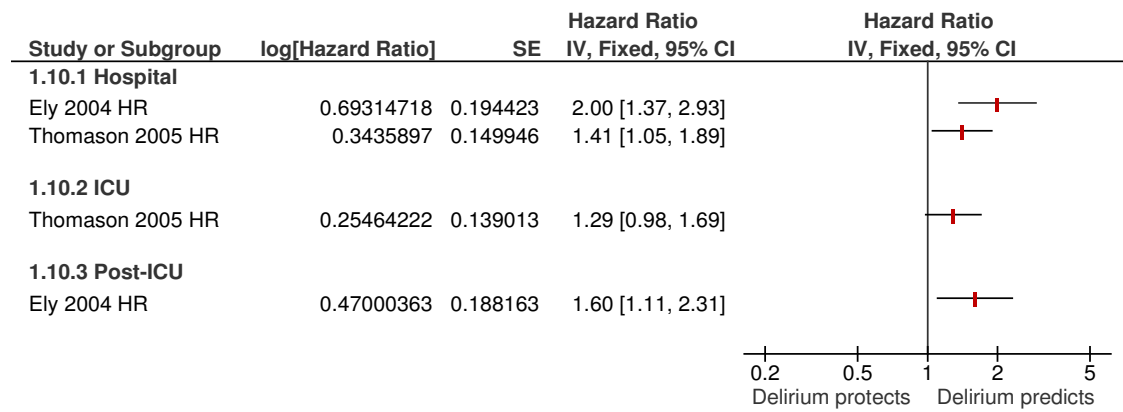
NB: Scale 0.05 to 20

Figure 9.4a: length of stay (discharge from hospital) as a consequence of delirium



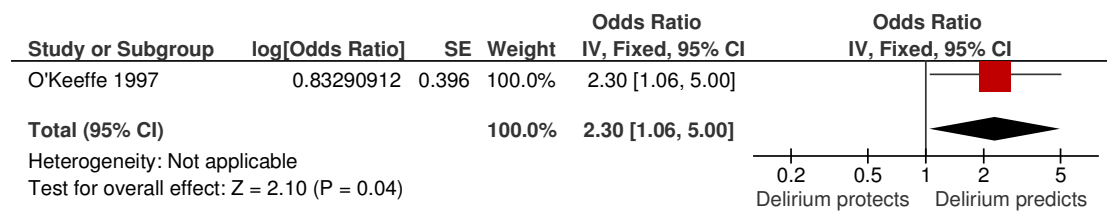
NB: Scale 0.1 to 10

Figure 9.4b: length of stay as a consequence of delirium



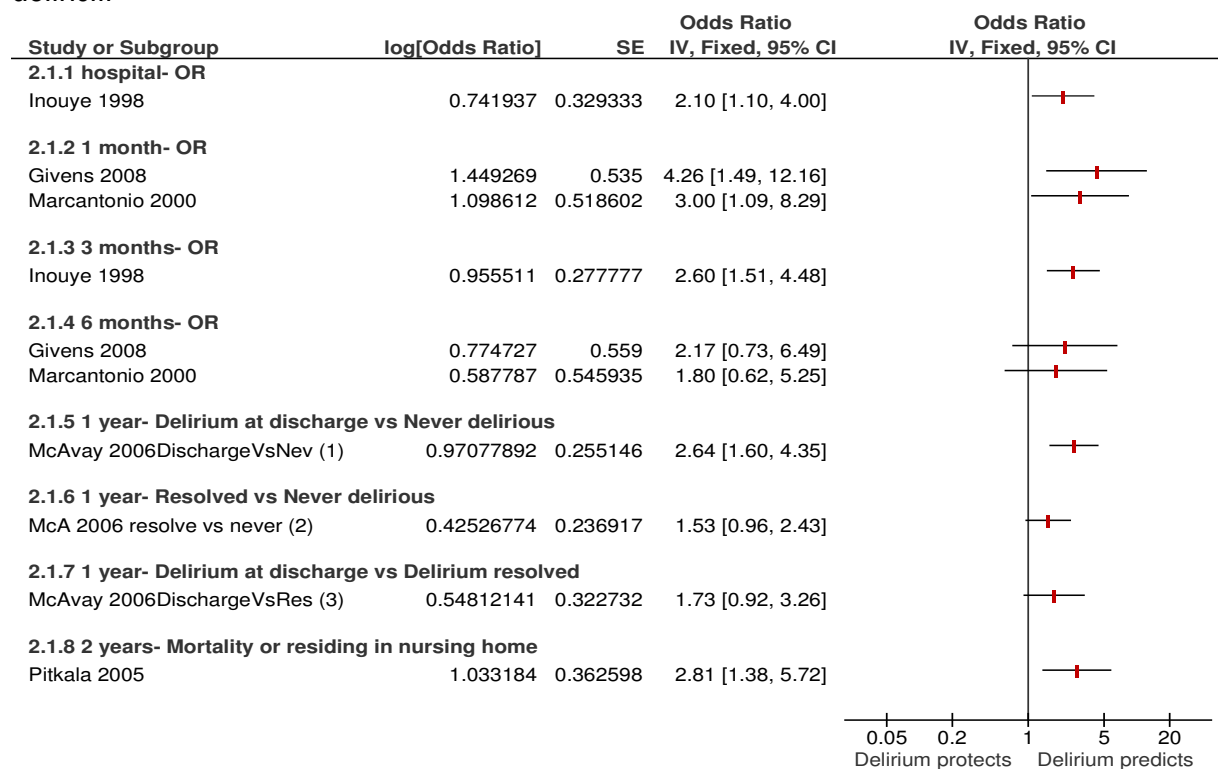
NB: Scale 0.2 to 5

Figure 9.5: hospital acquired complications as a consequence of delirium



NB: Scale 0.2 to 5

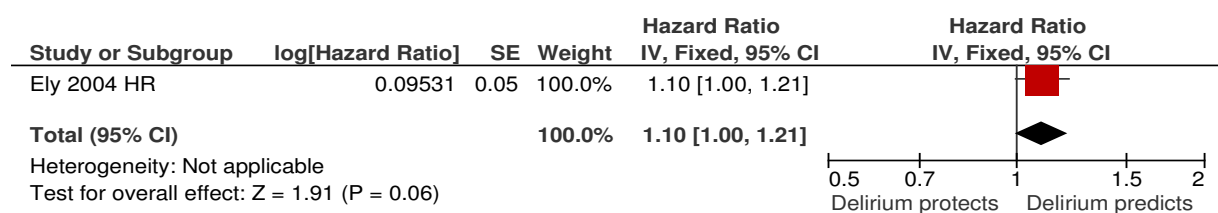
Figure 9.6: mortality or new admission to institution as a consequence of delirium



- (1) HR
- (2) HR
- (3) HR

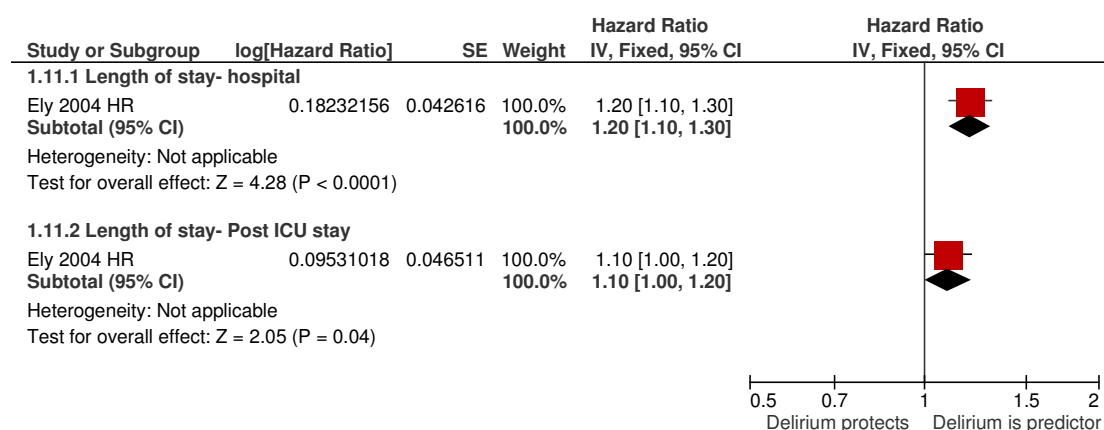
NB: Scale 0.05 to 20

Figure 9.7: mortality as a consequence of increased duration of delirium



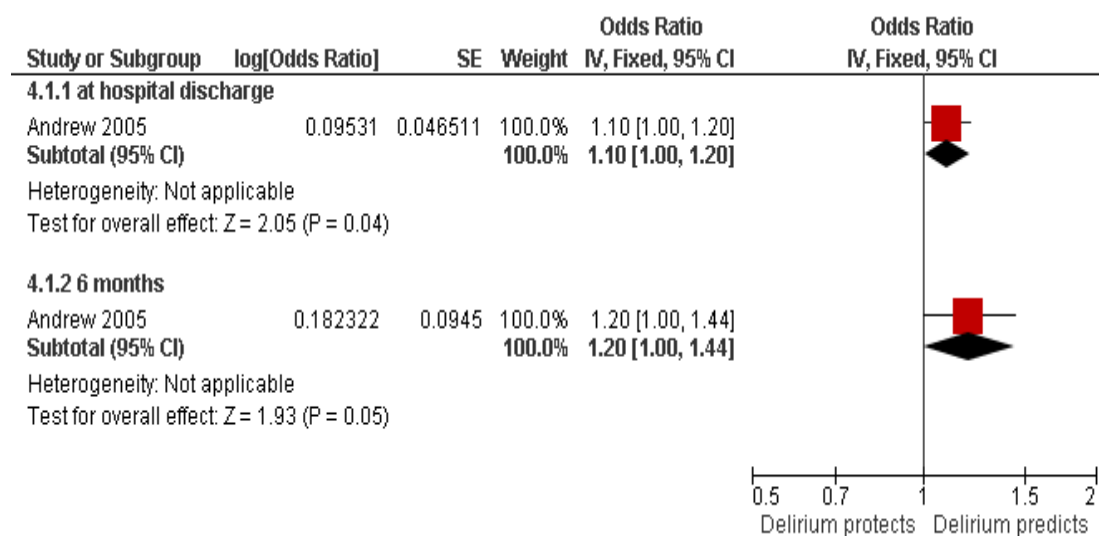
NB: Scale 0.5 to 2

Figure 9.8: length of stay as a consequence of increased duration of delirium



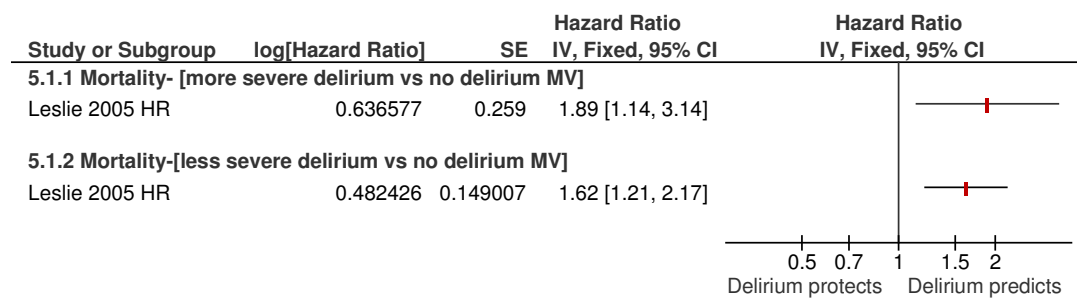
NB: Scale 0.5 to 2

Figure 9.9: mortality or functional decline as a consequence of increased duration of delirium



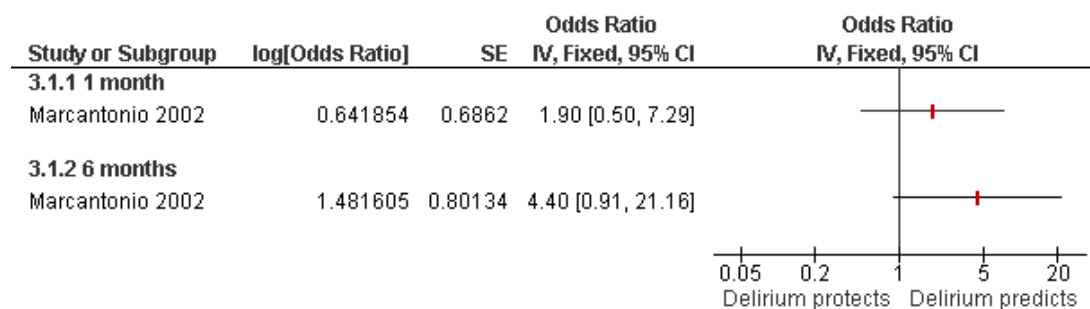
NB: Scale 0.5 to 2

Figure 9.10: mortality (at 1 year) as a consequence of delirium (severity)



NB: Scale 0.5 to 2

Figure 9.11: mortality or new admission to institution (at 1 month and 6 months) as a consequence of delirium severity



NB: Scale 0.05 to 20

Prevention of delirium: non-pharmacological

HYDRATION FOR THE PREVENTION OF DELIRIUM (LONG-TERM CARE SETTING)

Figure 10.1: acute confusion.

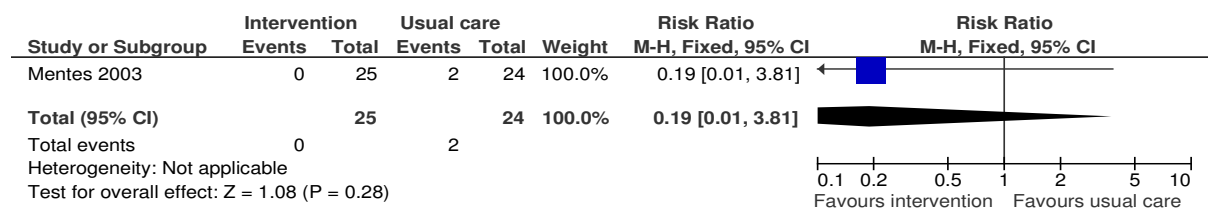
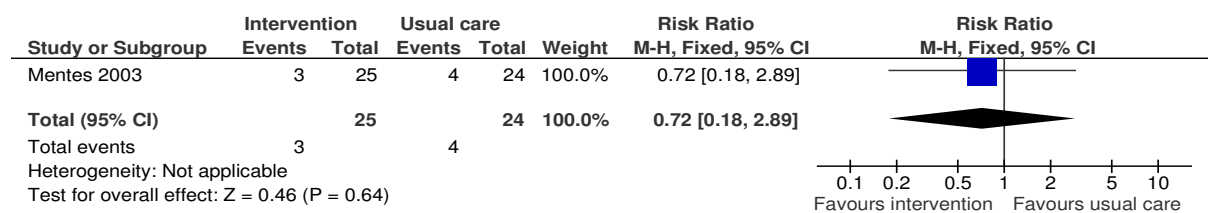
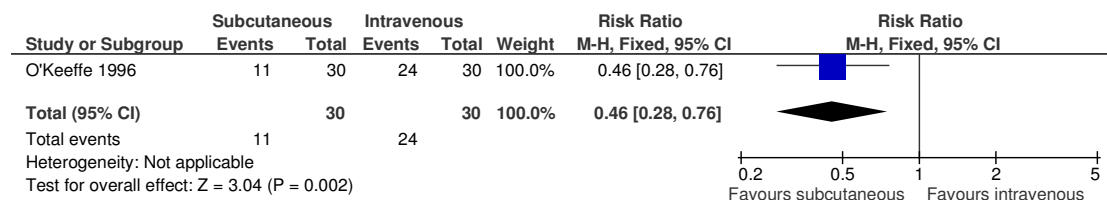


Figure 10.2: hydration-linked events.



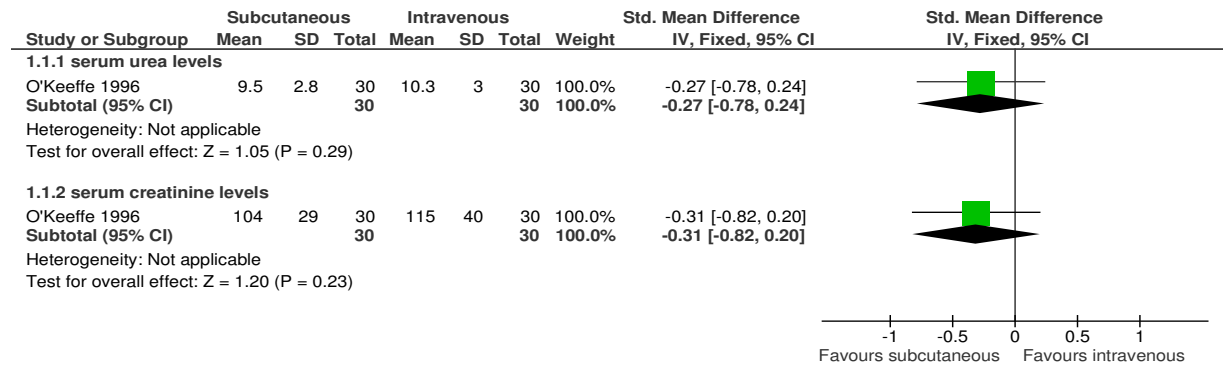
HYDRATION FOR THE PREVENTION OF DELIRIUM (HOSPITAL SETTING)

Figure 10.3: agitation



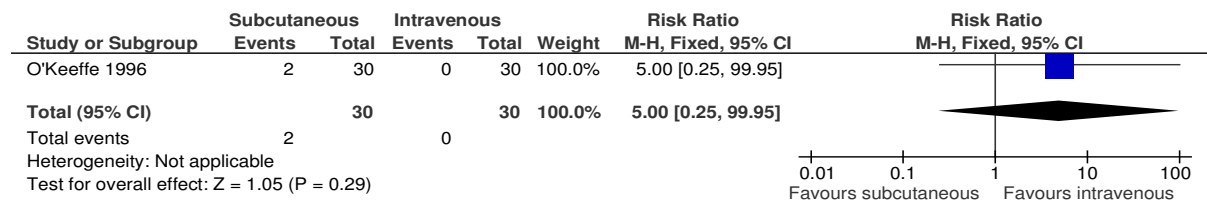
NB: Scale 0.2 to 5

Figure 10.4: serum levels



NB: Scale -1 to 1

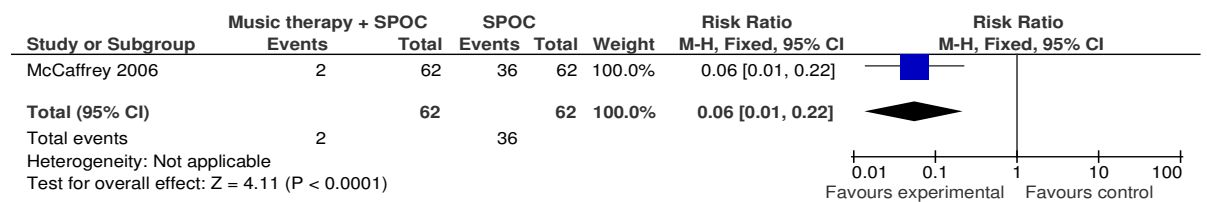
Figure 10.5: local oedema



NB: Scale 0.01 to 100

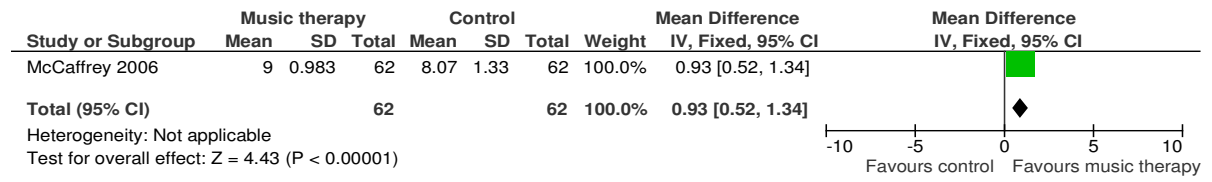
MUSIC THERAPY FOR THE PREVENTION OF DELIRIUM (HOSPITAL SETTING)

Figure 10.6: number of patients with delirium



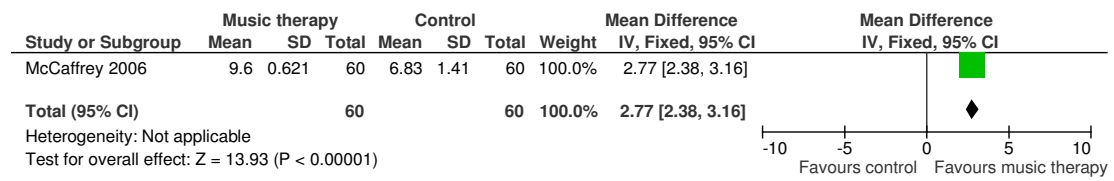
NB: forest plot scale 0.01 to 100

Figure 10.7: patient's readiness to ambulate after undergoing surgery



NB: Scale -10 to 10

Figure 10.8: patient satisfaction



NB: Scale -10 to 10

Multicomponent prevention

Figure 10.9: number of patients with delirium in hospital

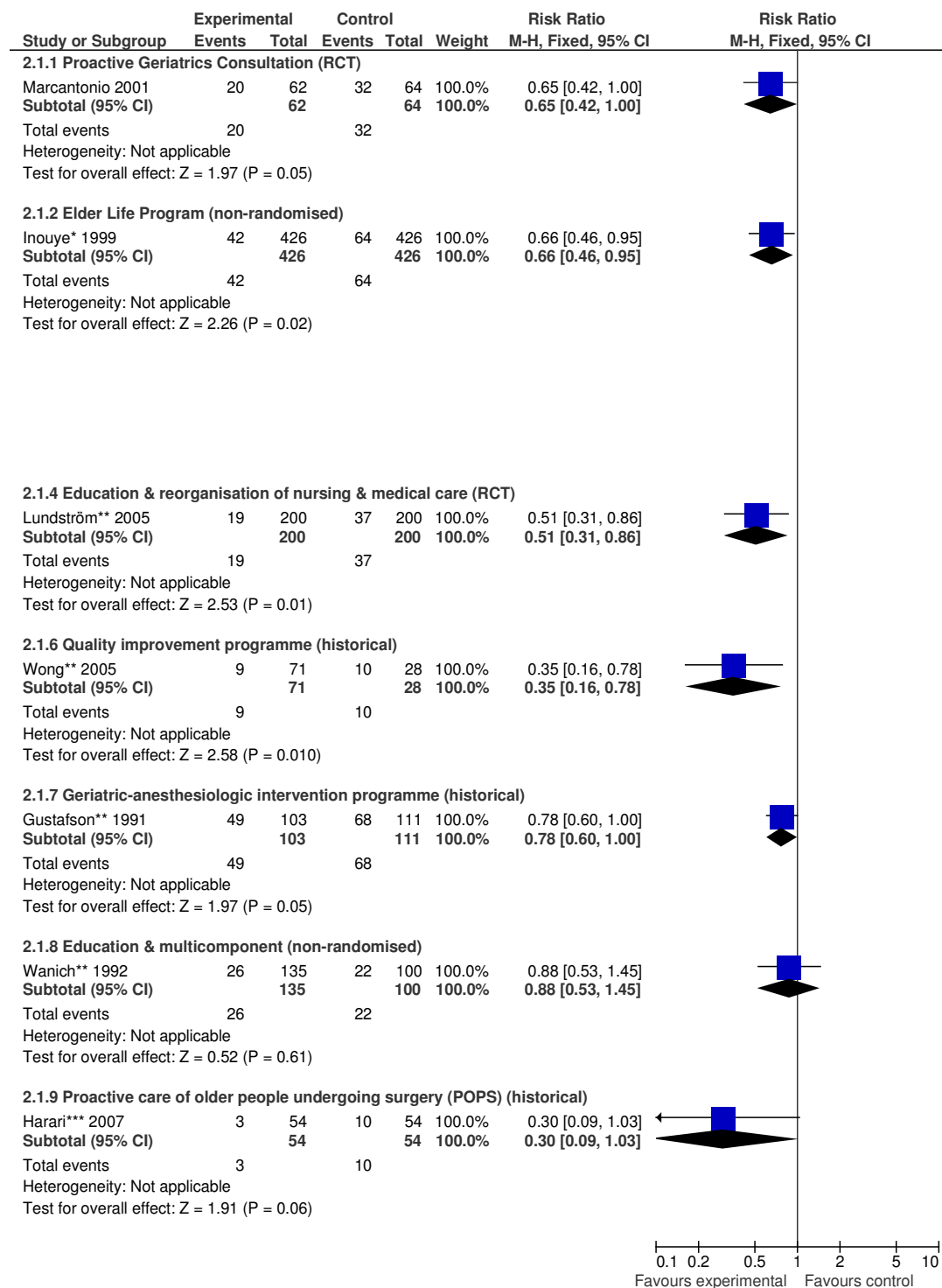


Figure 10.10: number of patients with delirium at 6 months follow-up

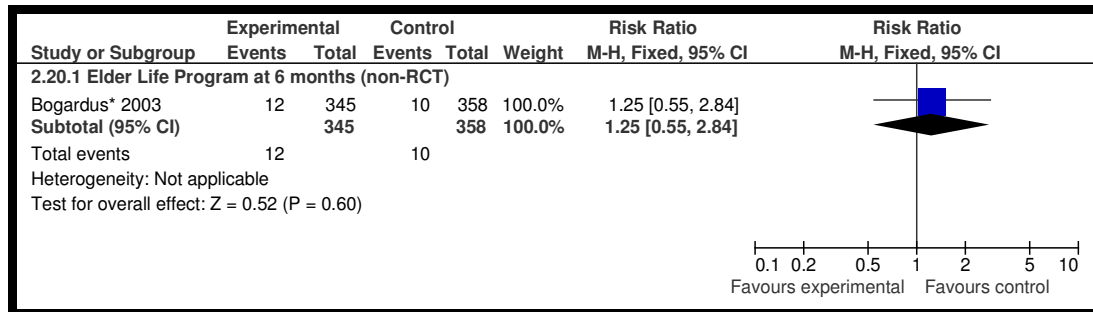


Figure 10.11: mean duration of delirium

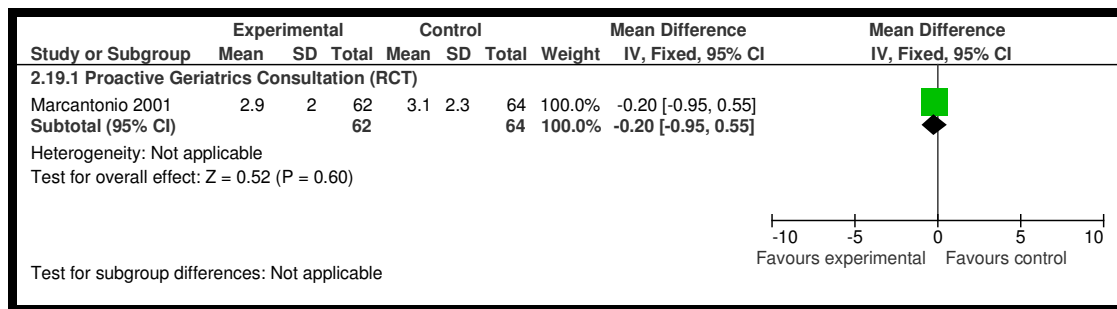


Figure 10.12: number of patients with delirium at 7 or more days

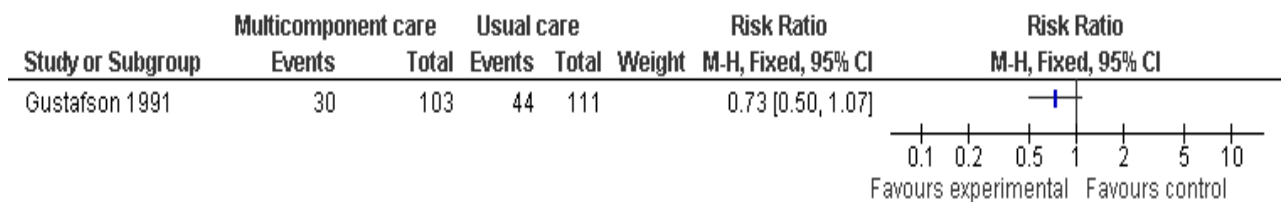


Figure 10.13: severity scores

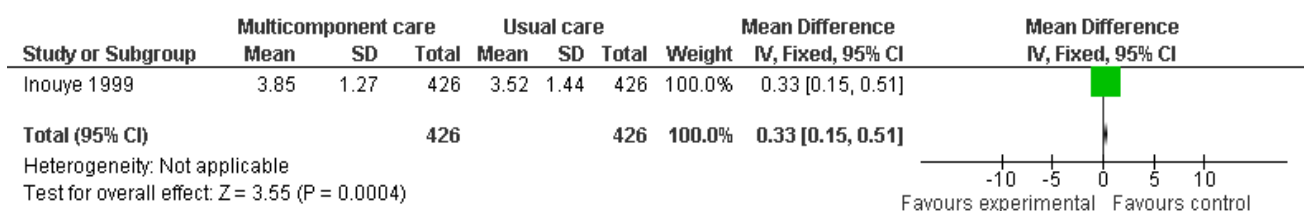


Figure 10.14: length of hospital stay

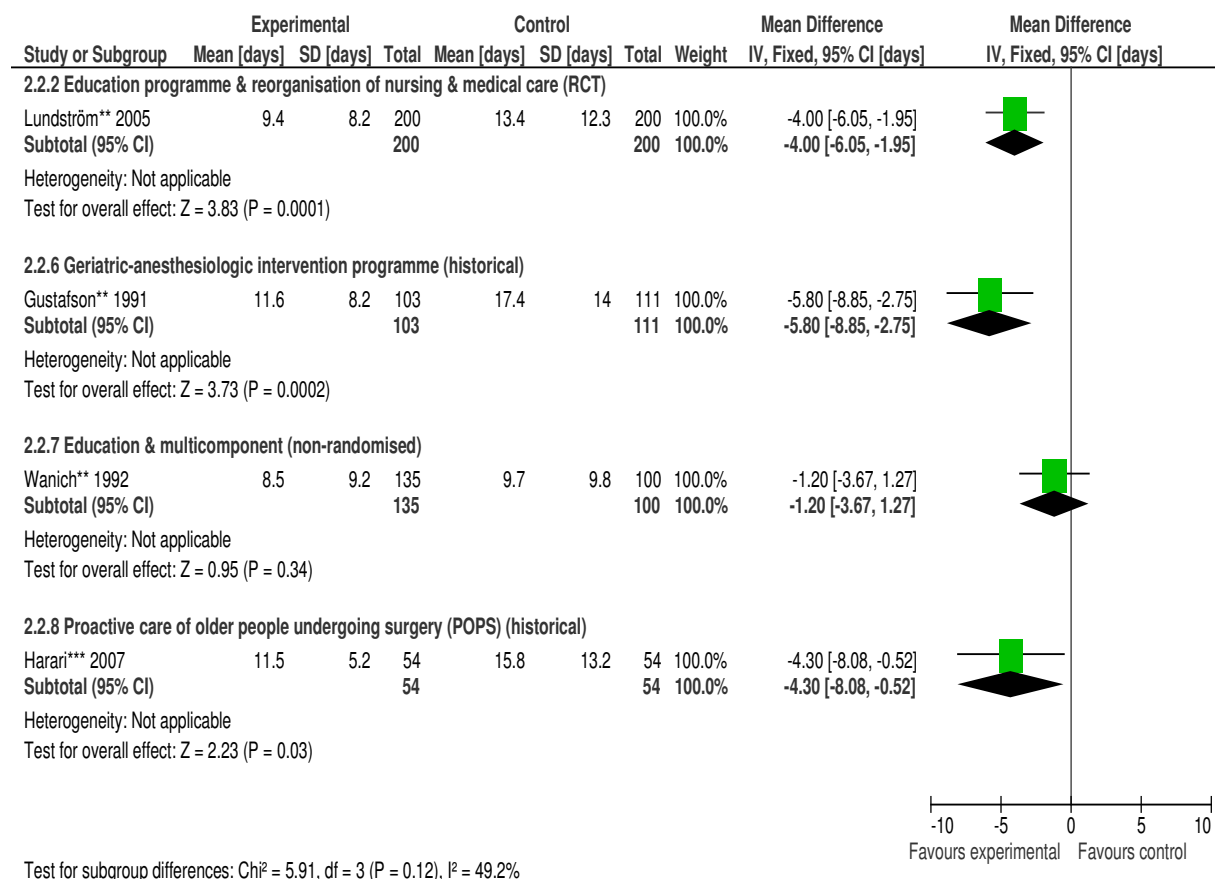


Figure 10.15: improvement in cognitive impairment at 5 days or discharge

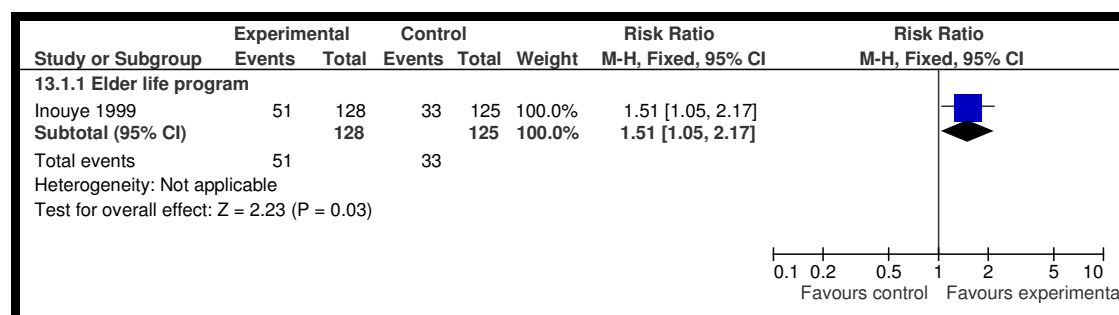


Figure 10.16: number of patients discharged to a new institutional setting

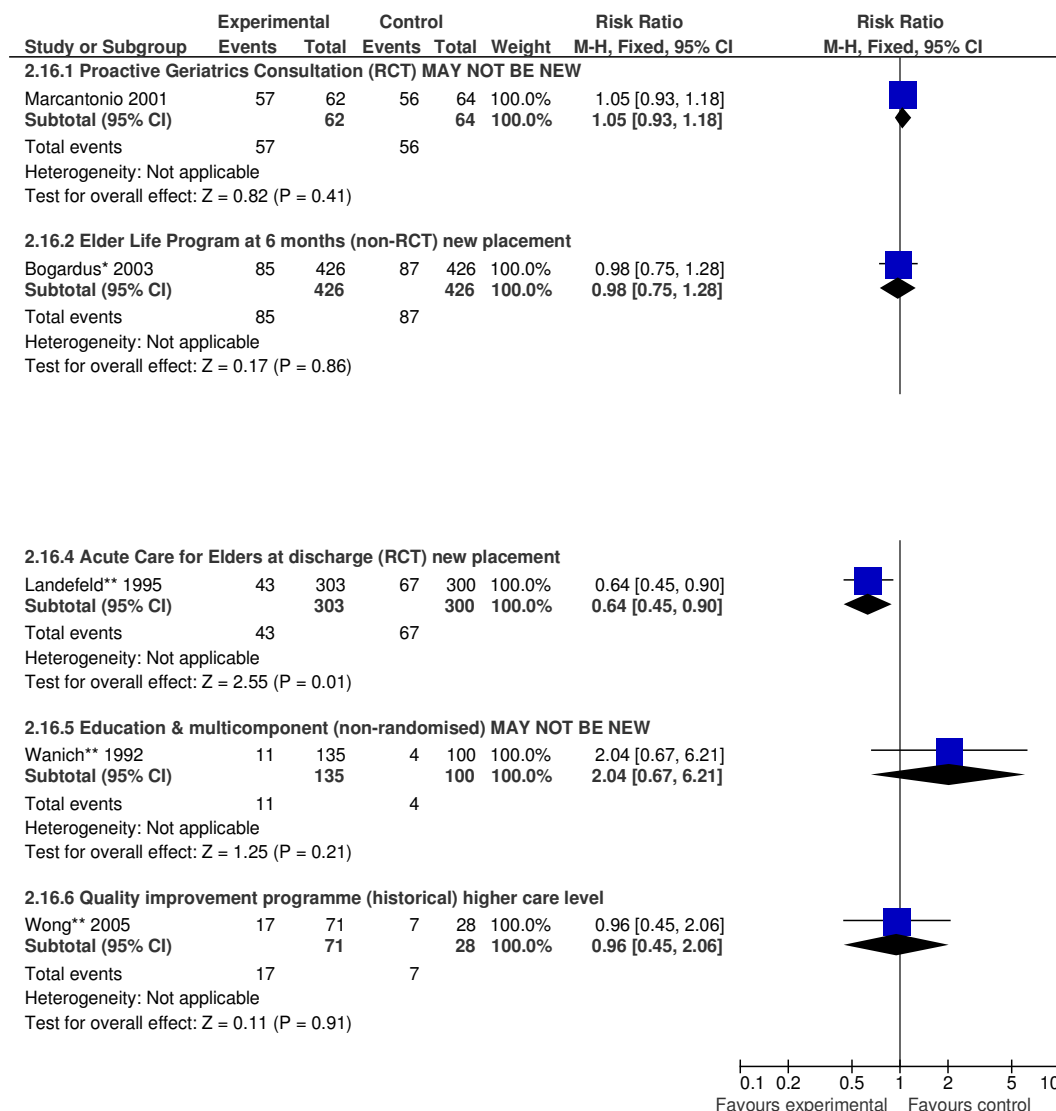


Figure 10.17: mortality in hospital

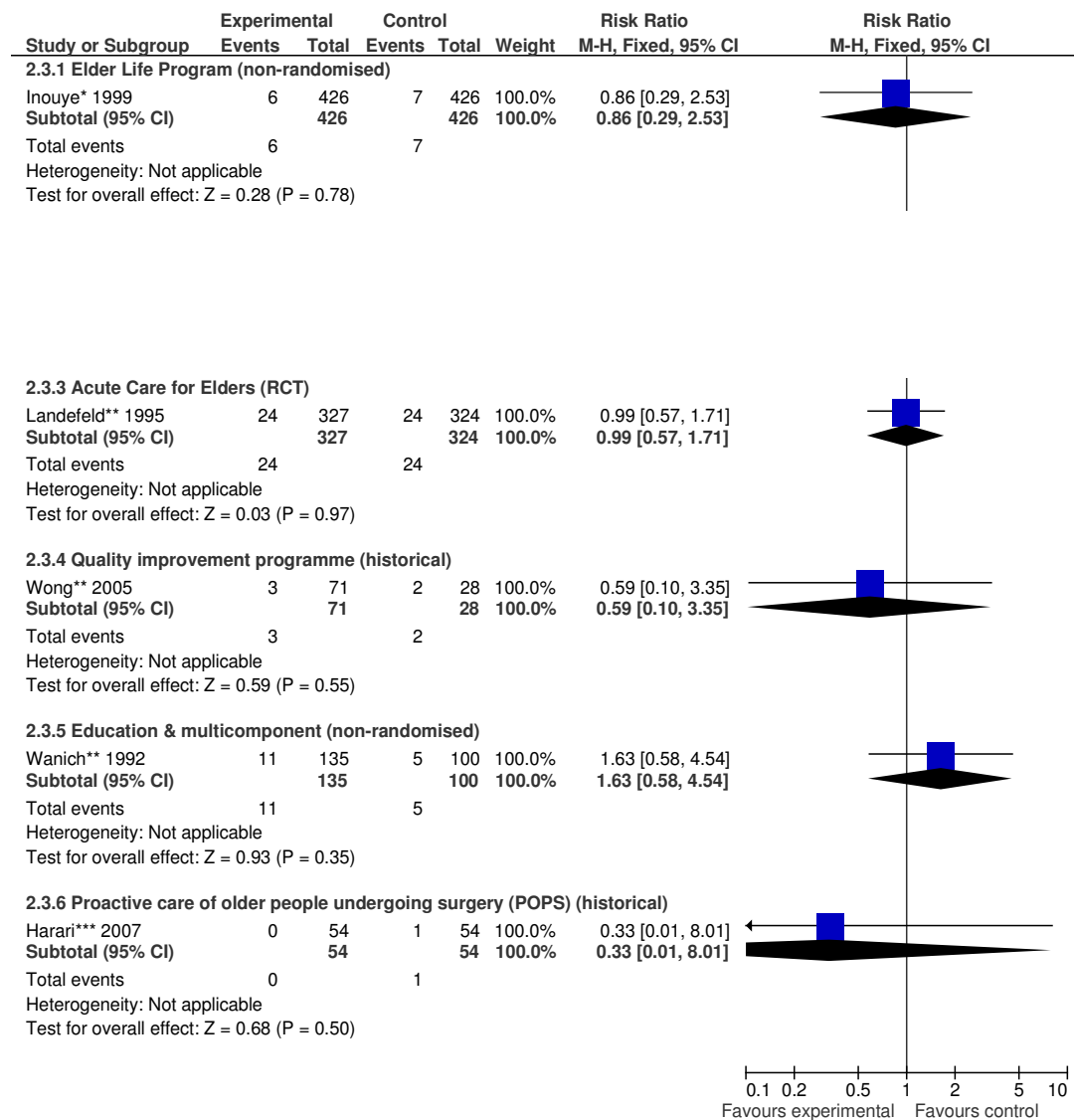


Figure 10.18: mortality at up to 6 months follow up

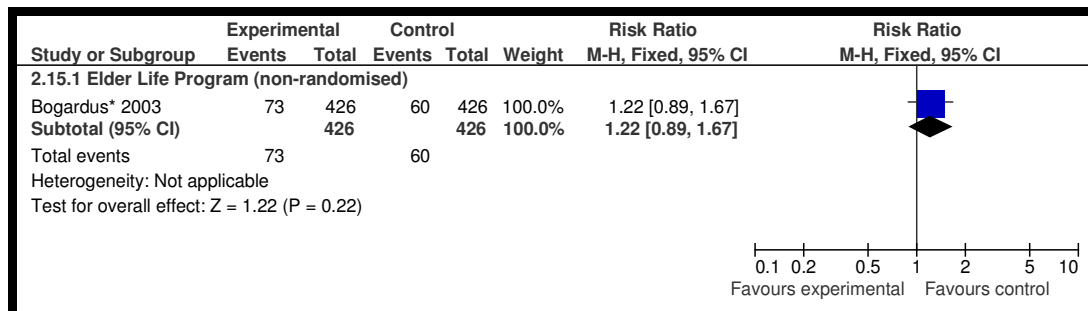


Figure 10.19: number of patients with an improvement in ADL

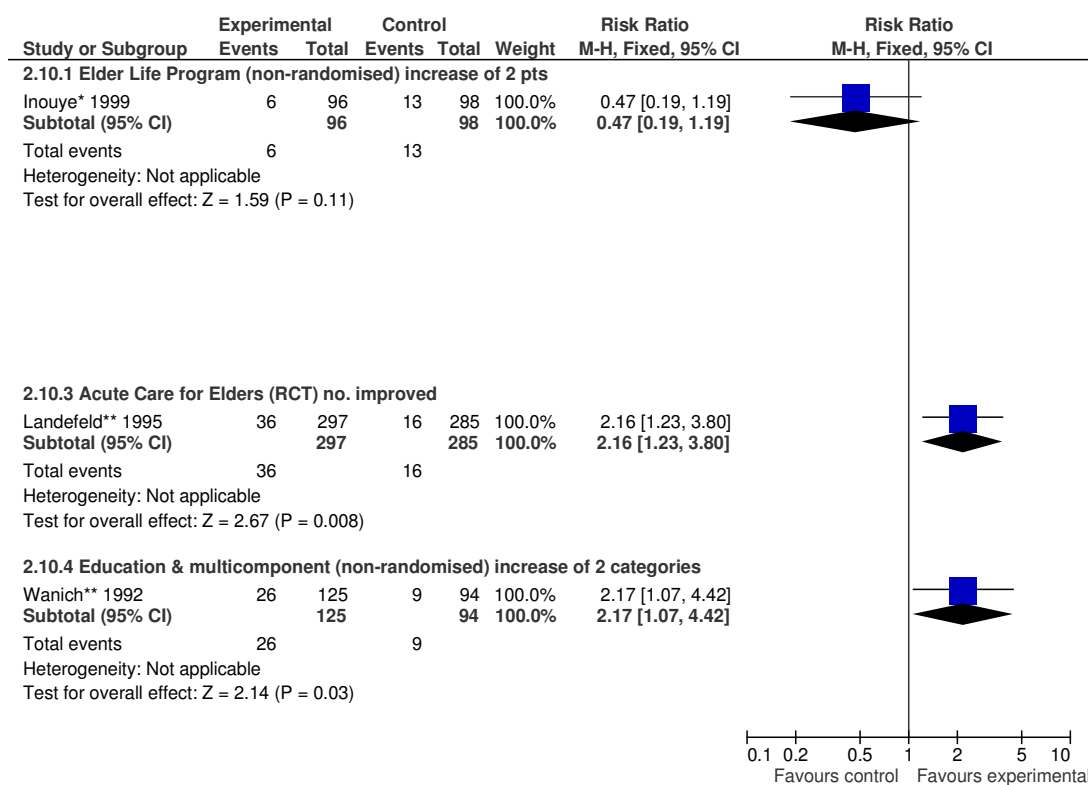


Figure 10.20: adjusted ADL score

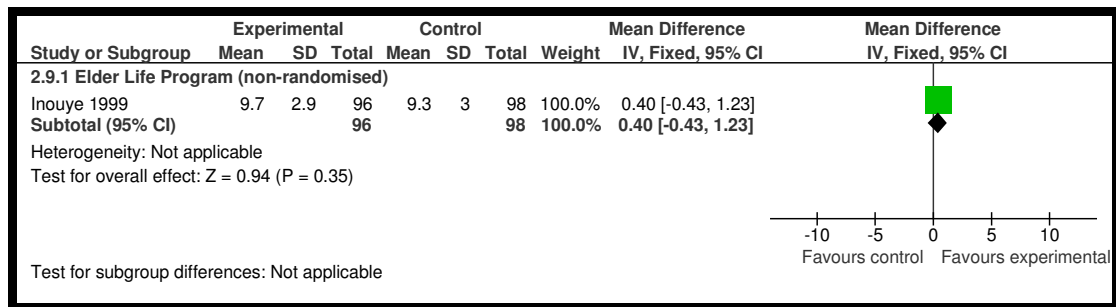


Figure 10.21: number of patients with severe falls

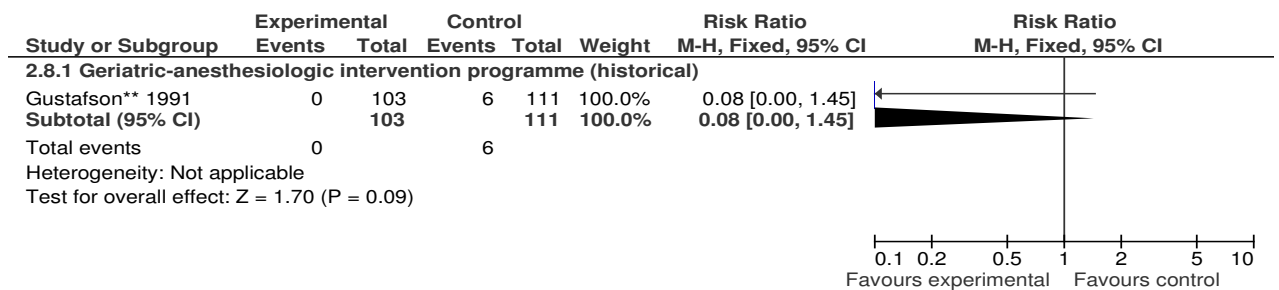


Figure 10.22: urinary tract infections

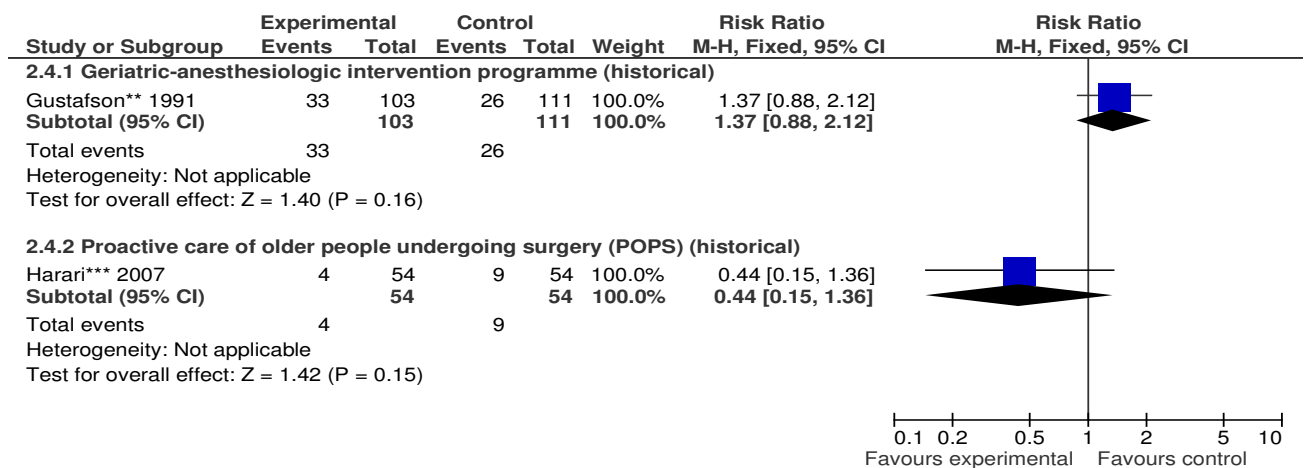
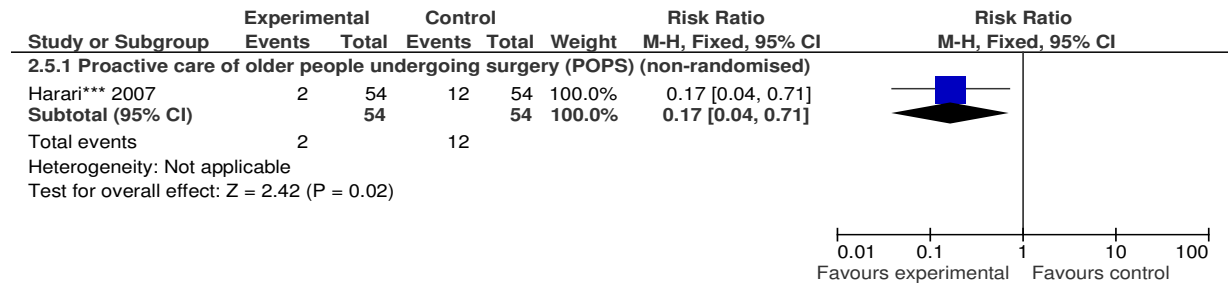


Figure 10.23: wound infections



NB scale 0.01 to 100

Figure 10.24: pressure ulcers

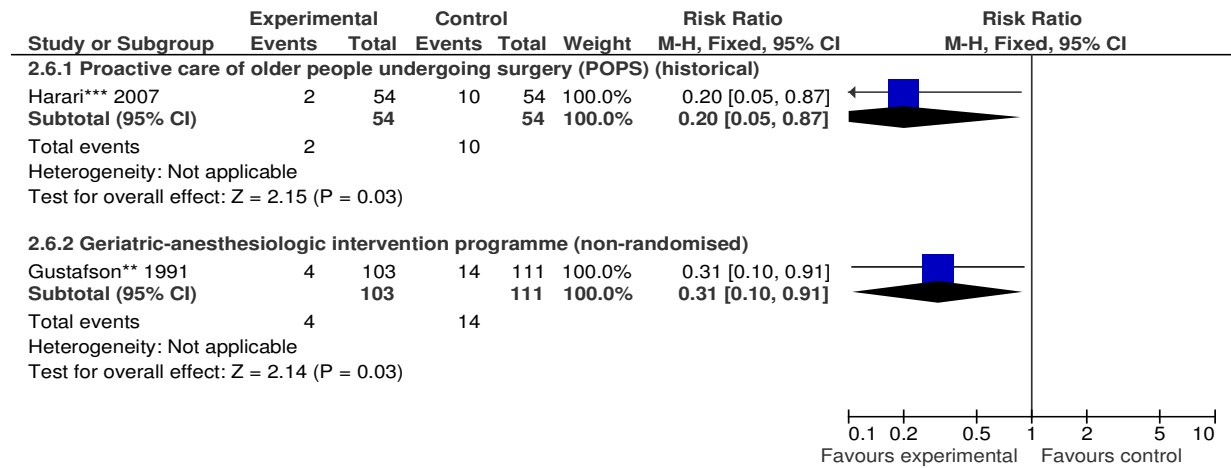


Figure 10.25: early vision correction at reassessment (day 5 or at discharge if earlier)

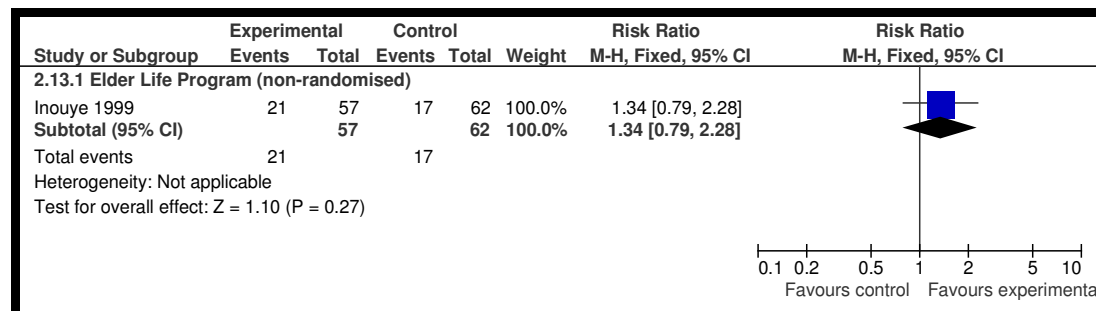


Figure 10.26: whisper test

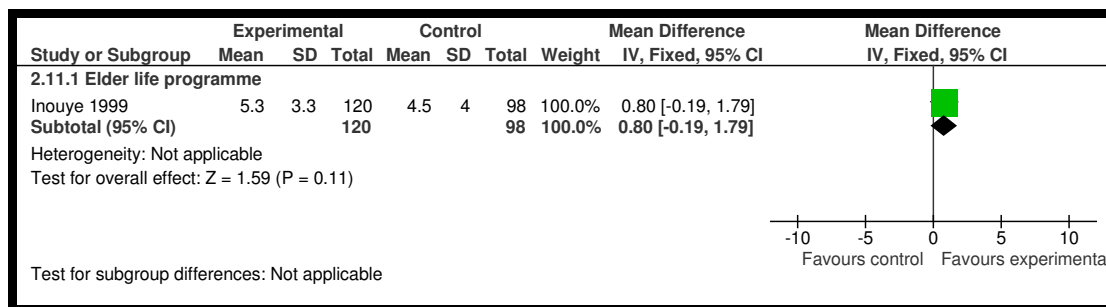


Figure 10.27: whisper test – number of patients with improvement by one point

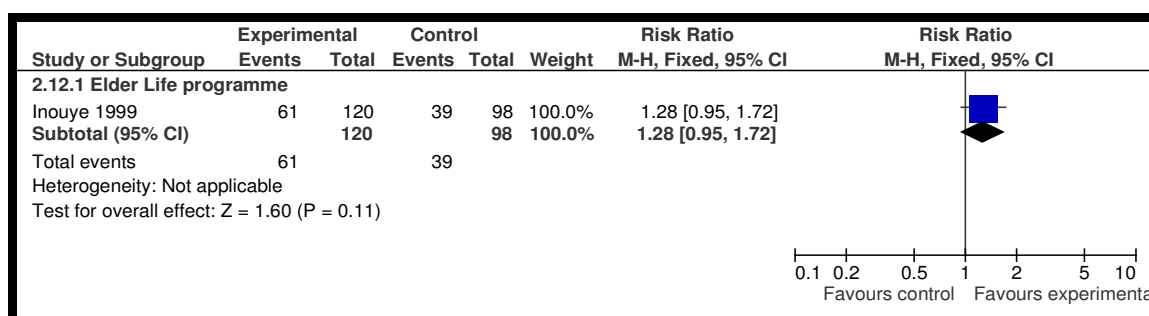


Figure 10.28: number of patients with improvement in dehydration

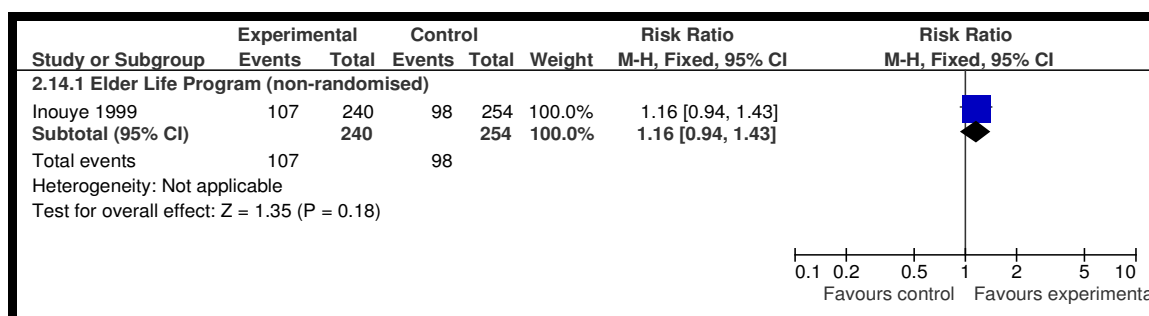


Figure 10.29: number of patients with dehydration

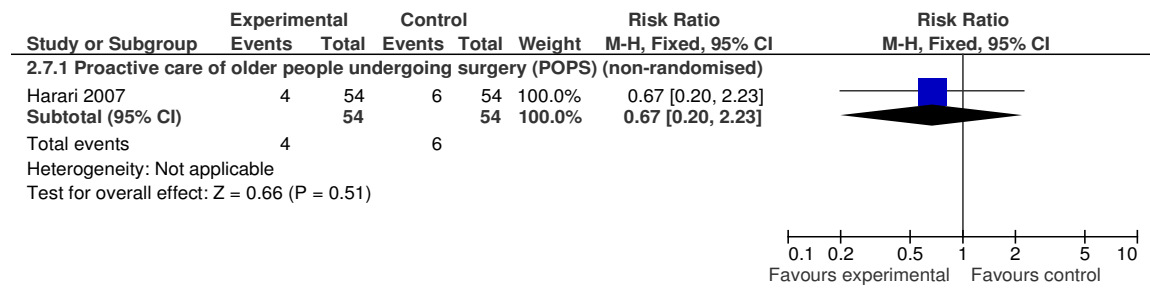
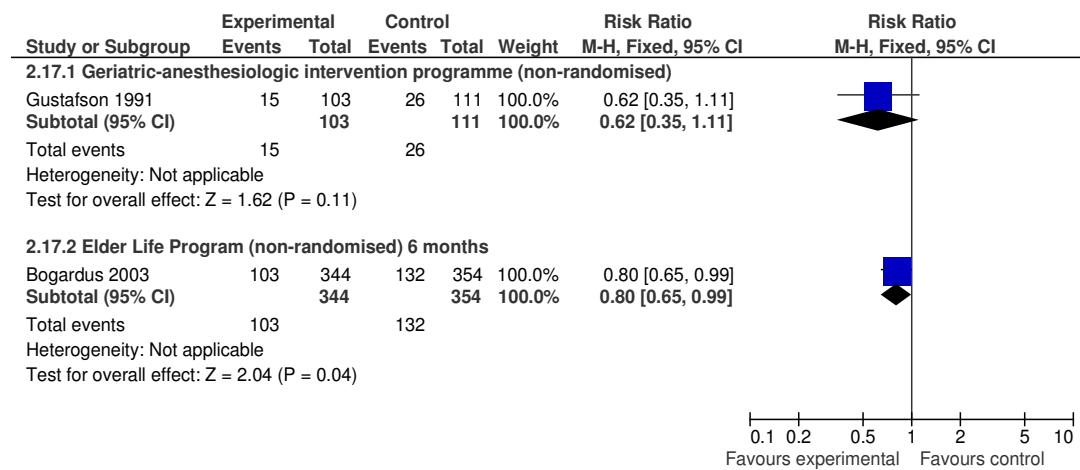


Figure 10.30: urinary incontinence



Prevention of delirium: pharmacological

Acetylcholinesterase inhibitor versus placebo

Figure 11.1: number of patients with delirium

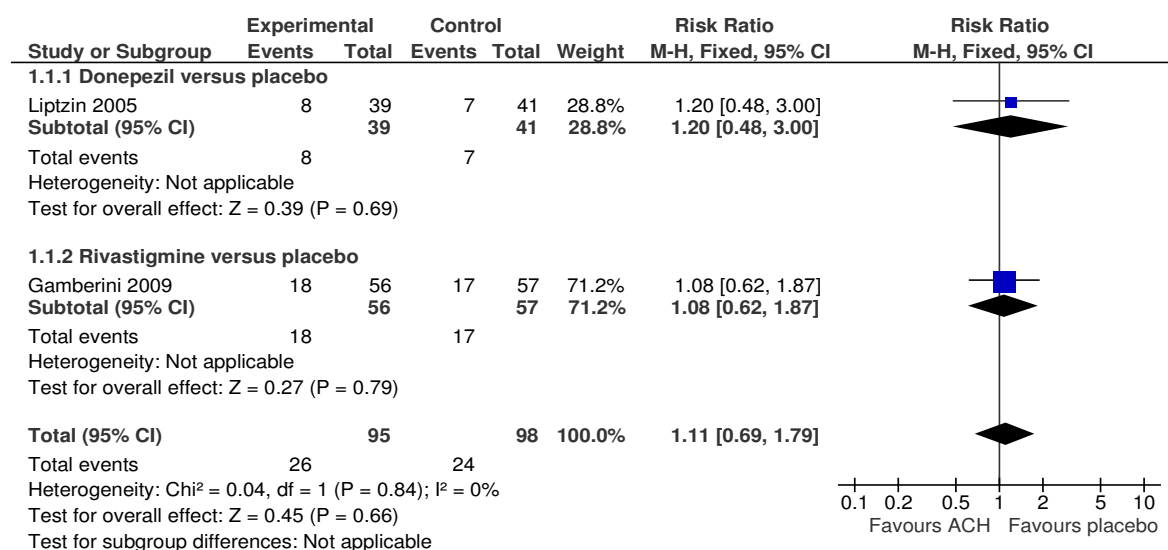


Figure 11.2: duration of delirium

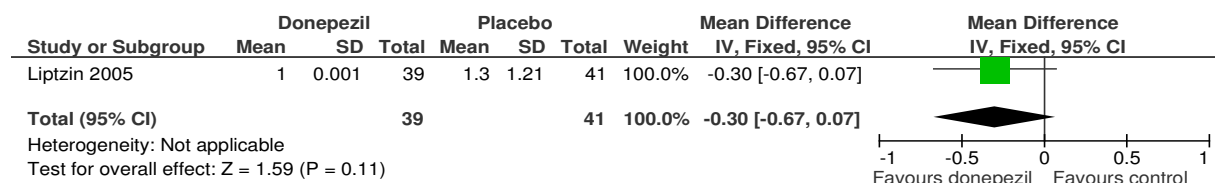


Figure 11.3: length of hospital stay

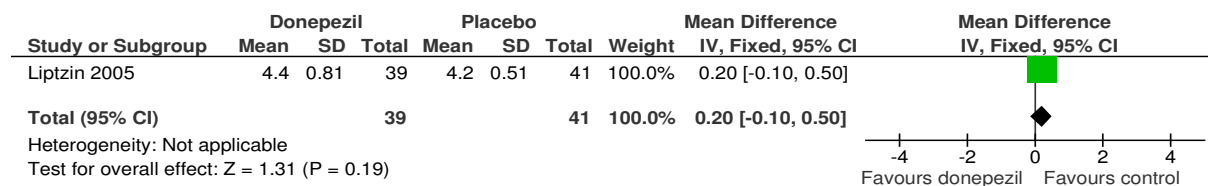
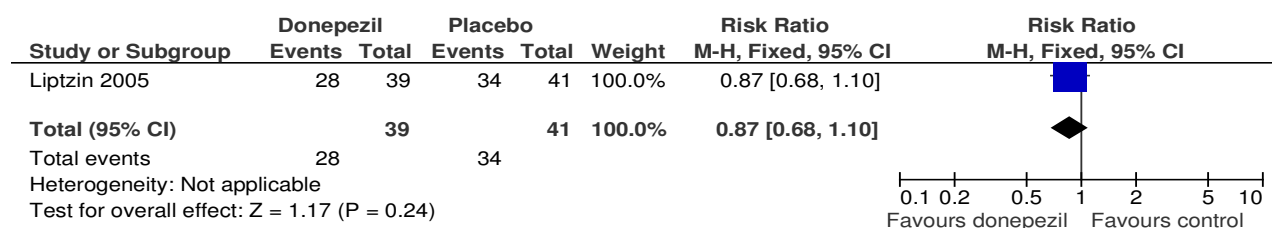


Figure 11.4: discharge to rehabilitation facility



Typical antipsychotics versus placebo

Figure 11.5: number of patients with postoperative delirium

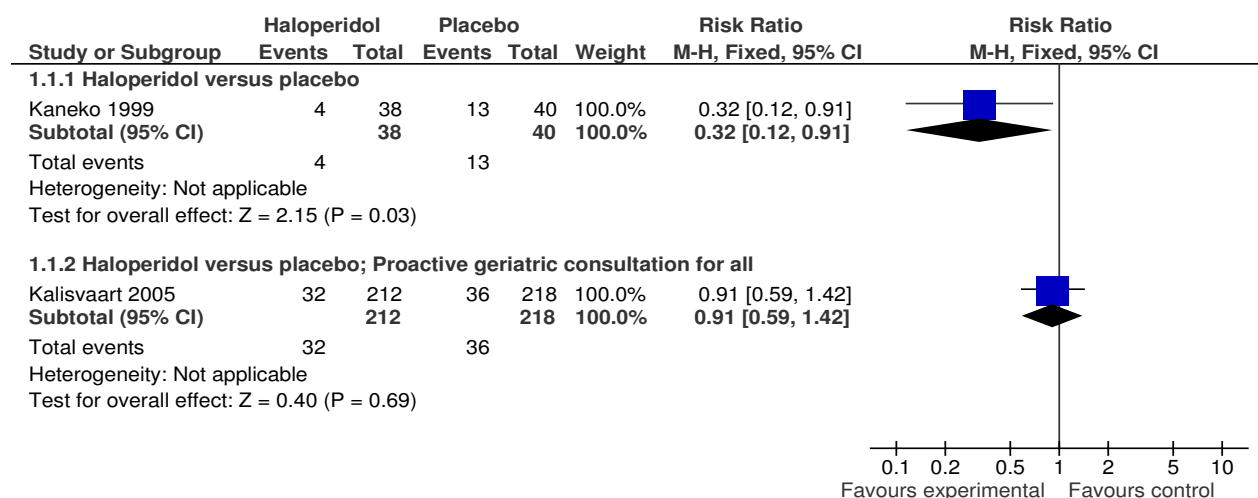


Figure 11.6: severity of delirium scores

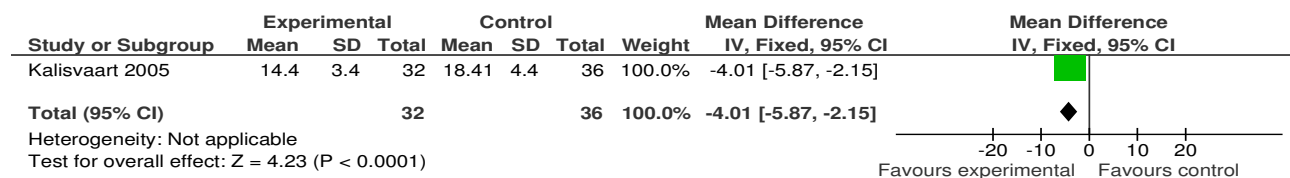


Figure 11.7: duration of delirium

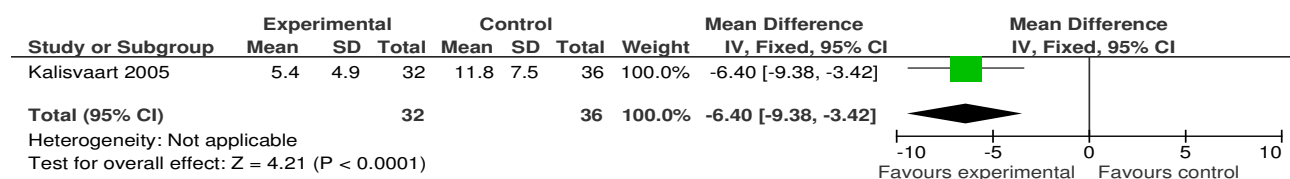
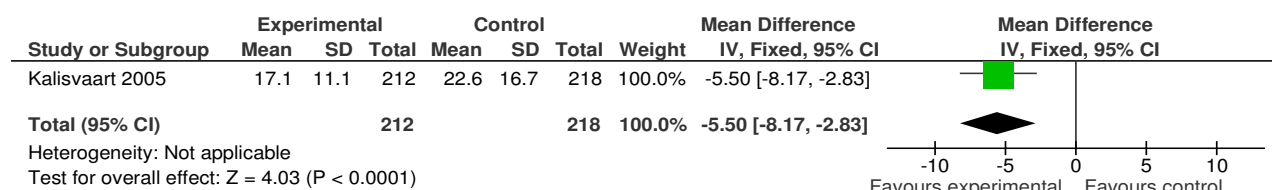
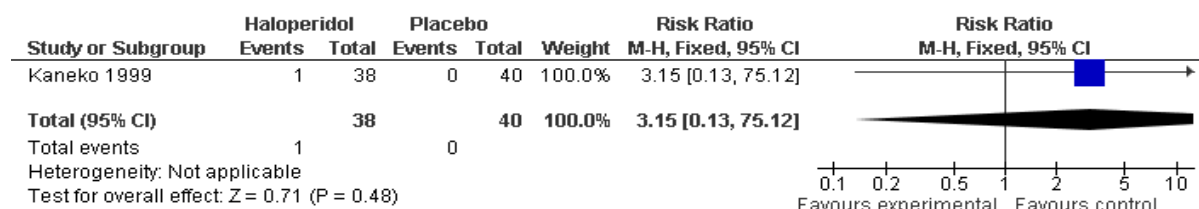


Figure 11.8: length of hospital stay



NB: Scale -10 to 10

Figure 11.9: number of patients with adverse events



Atypical antipsychotics versus placebo

Figure 11.10: number of patients with delirium

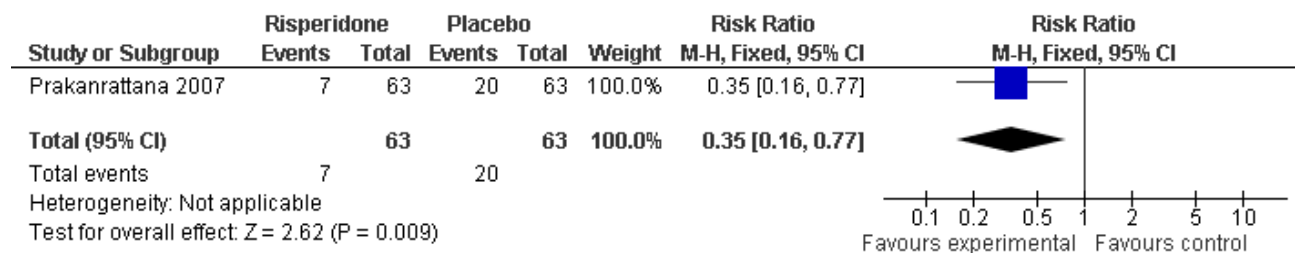


Figure 11.11: length of ICU stay

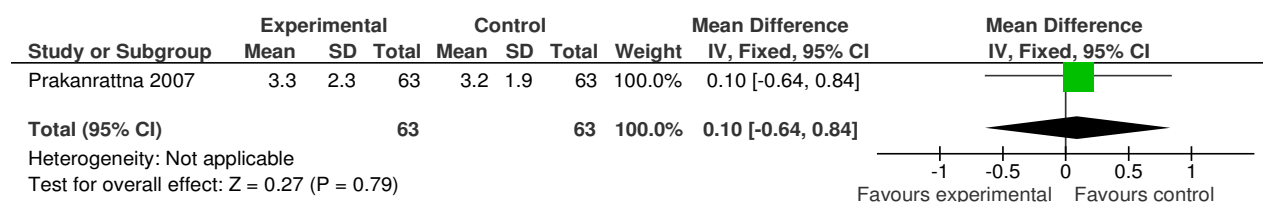
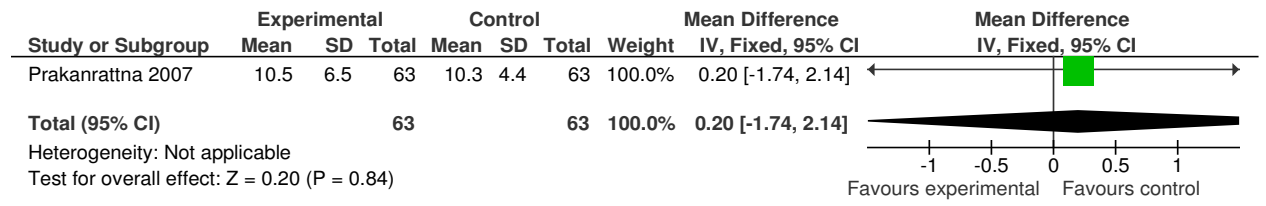


Figure 11.12: length of hospital stay



Pharmacological Prevention in long-term care

Acetylcholinesterase inhibitor versus placebo

Figure 11.13: incidence of delirium

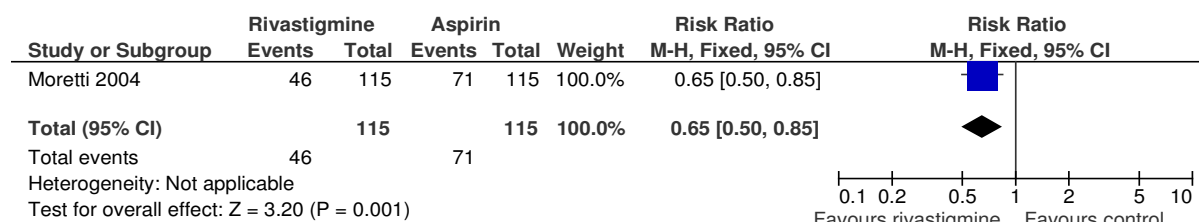


Figure 11.14a: duration of delirium (all patients)

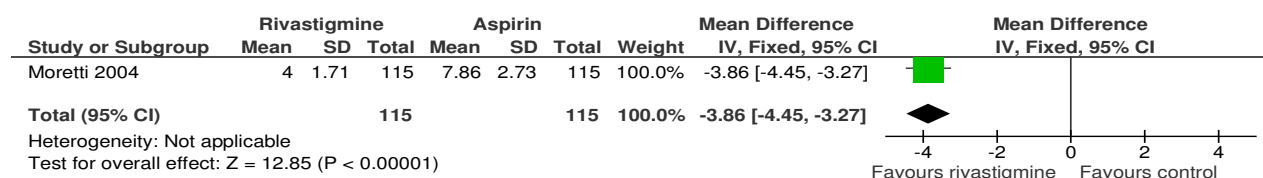


Figure 11.14b: duration of delirium assuming mean is across those with delirium

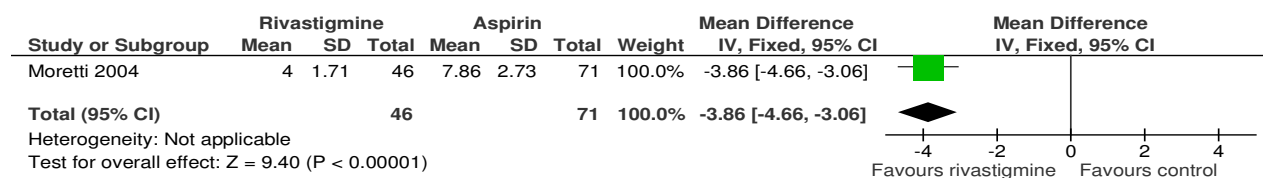


Figure 11.15: cognitive impairment (Clinical Dementia Rating change scores)

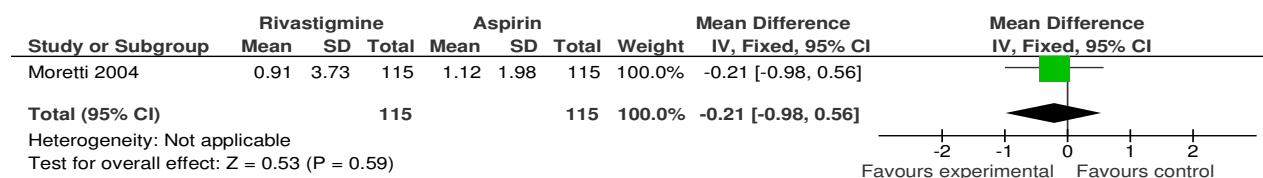


Figure 11.16a: BEHAVE-AD scale change scores

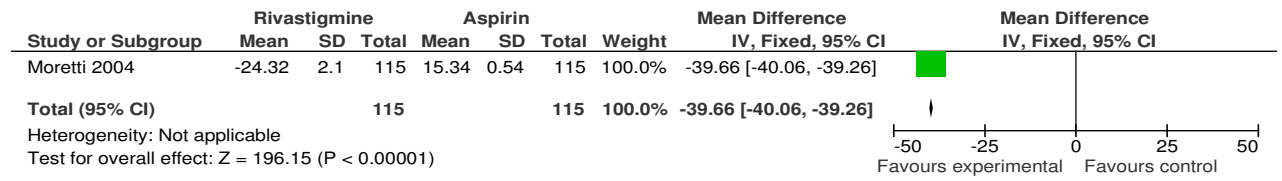
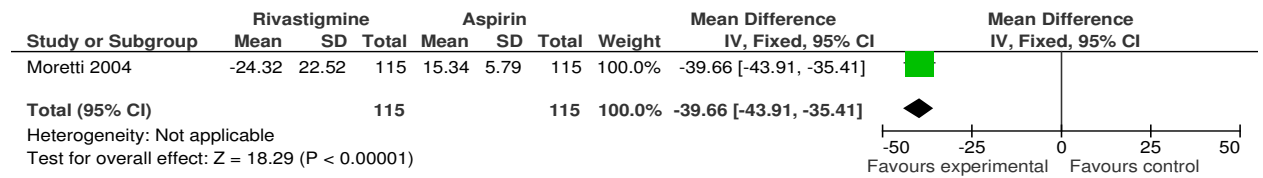


Figure 11.16b: BEHAVE-AD overall change scores



Treatment of delirium: non-pharmacological (hospital setting)

Multicomponent intervention

Figure 12.1: number of patients with complete response.

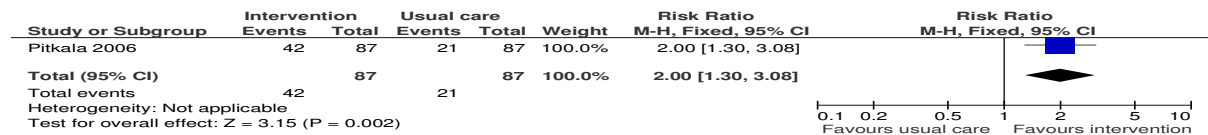


Figure 12.2: cognitive impairment

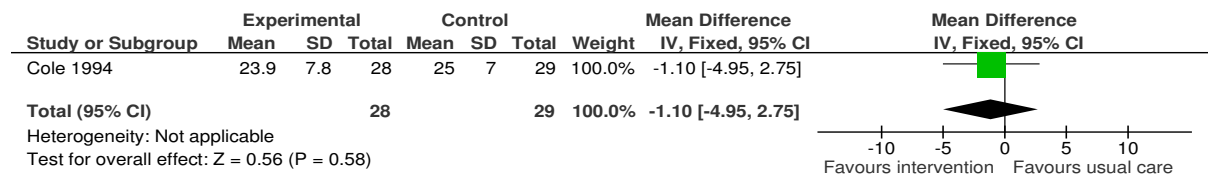


Figure 12.3: length of stay

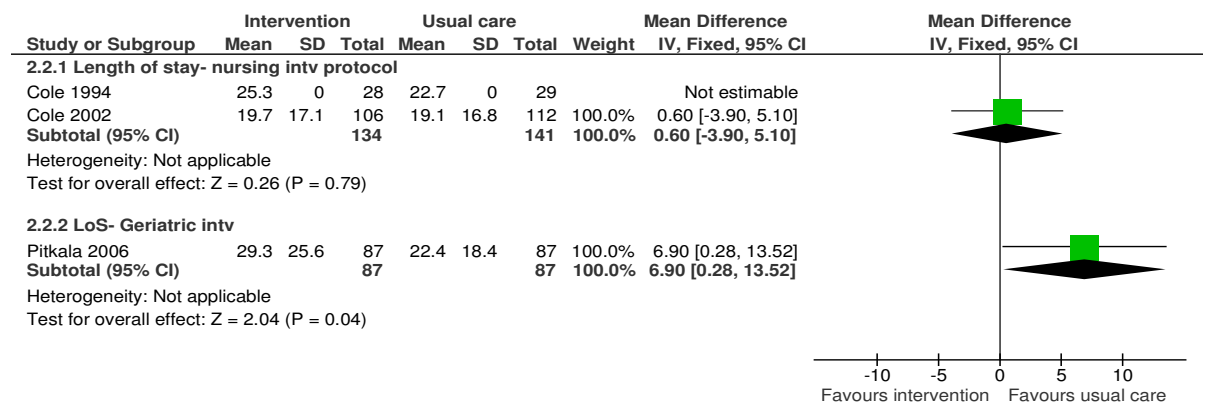


Figure 12.4: discharge to higher dependency or to new long-term care (RCTs)

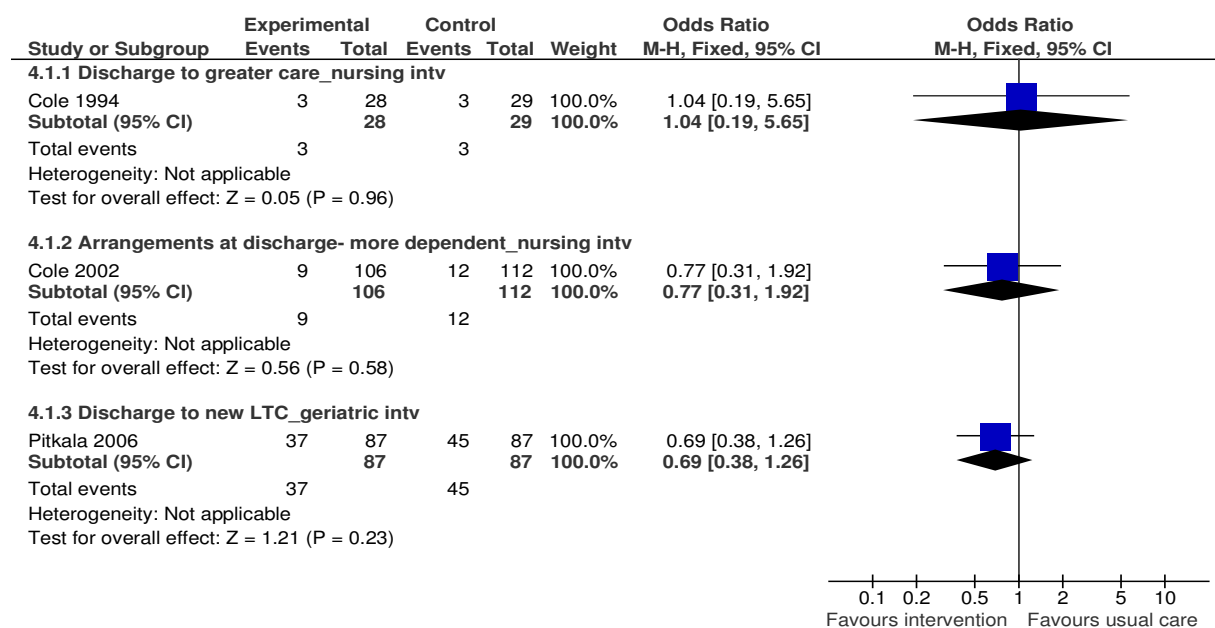
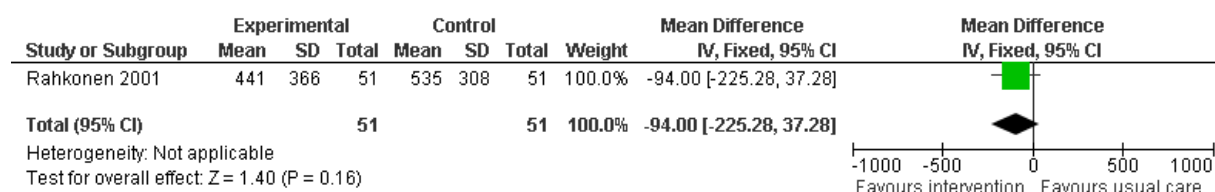
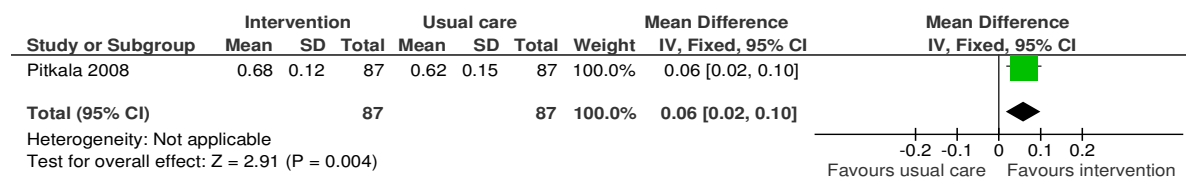


Figure 12.5: number of days in new long-term care (non RCT)



NB: Scale -1000 to 1000

Figure 12.6: improvement in HRQoL



NB: Scale -0.2 to 0.2

Figure 12.7: mortality (RCTs only)

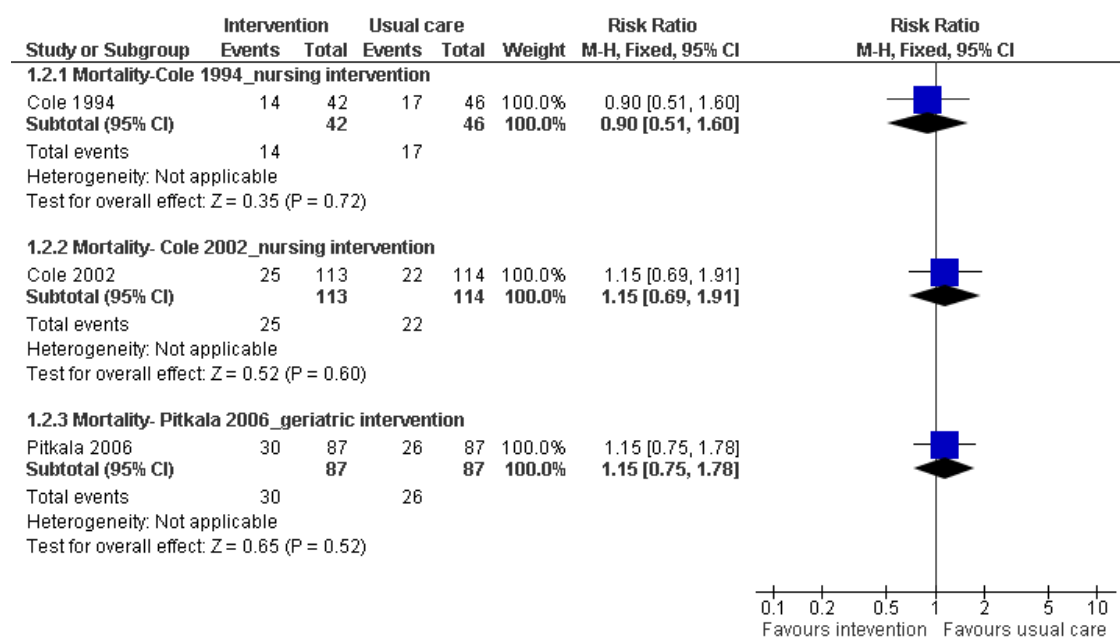
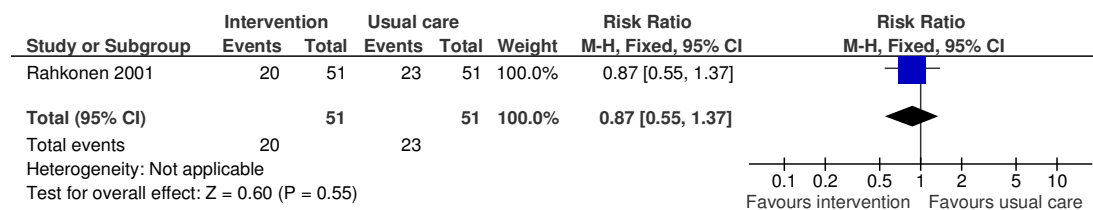


Figure 12.8: mortality (non-RCT)



Treatment of delirium: pharmacological

Typical antipsychotics versus placebo

Figure 13.1: complete response

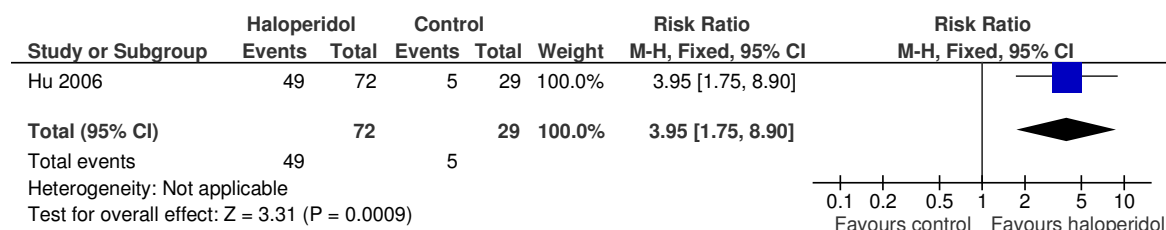
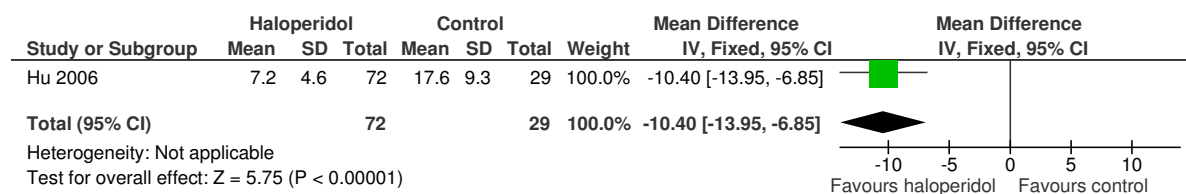


Figure 13.2: severity of delirium



Atypical antipsychotics versus placebo

Figure 13.3: complete response

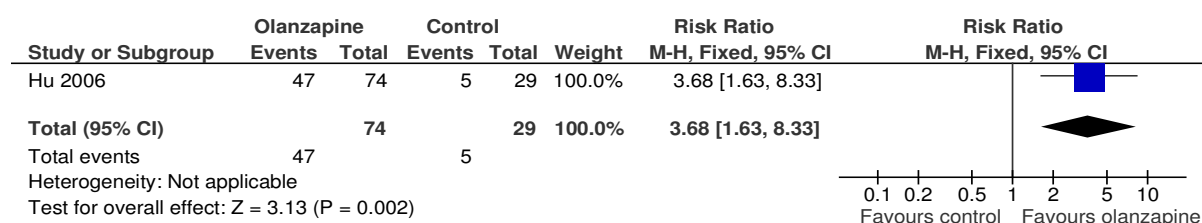
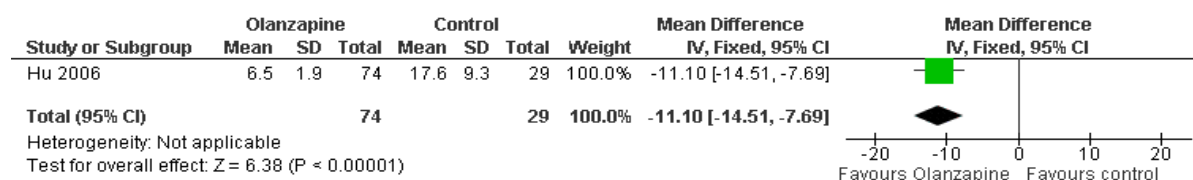


Figure 13.4: severity of delirium



NB: Scale -20 to 20

Atypical antipsychotics 1 versus atypical antipsychotics 2

Figure 13.5: duration of delirium

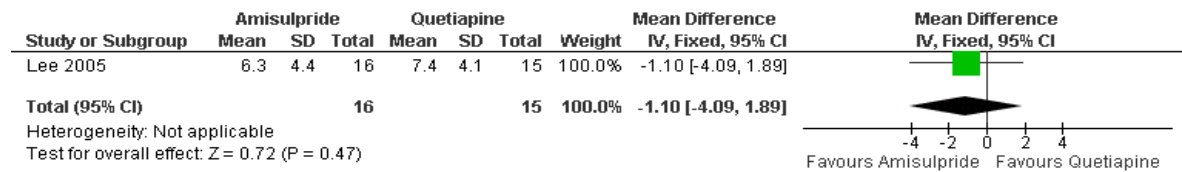


Figure 13.6: severity of delirium

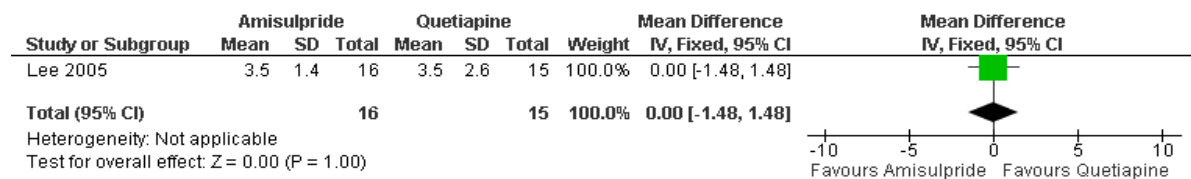


Figure 13.7: complete response

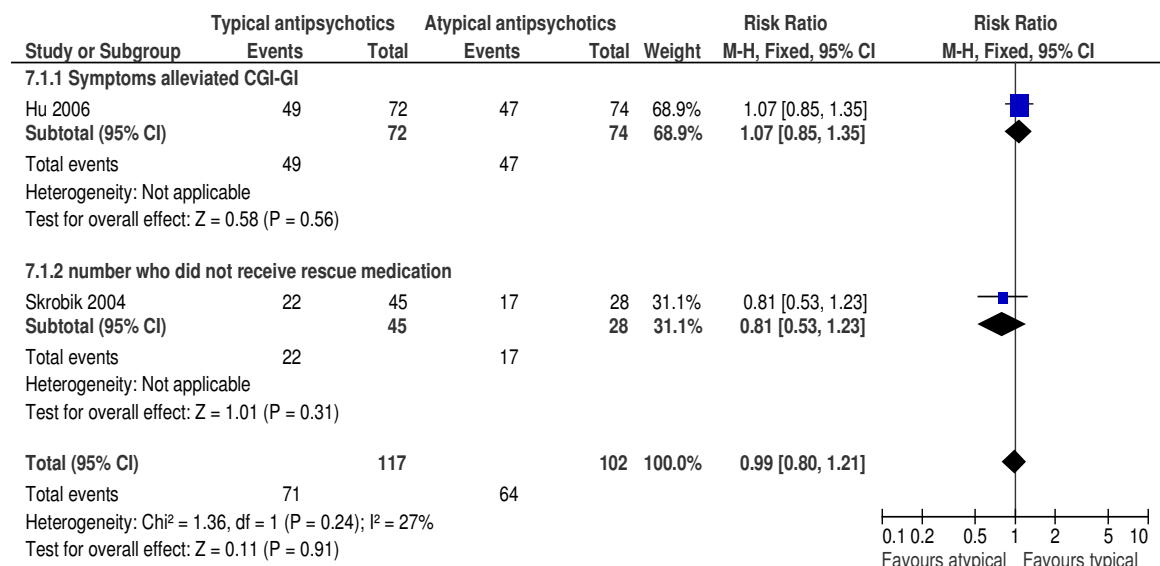
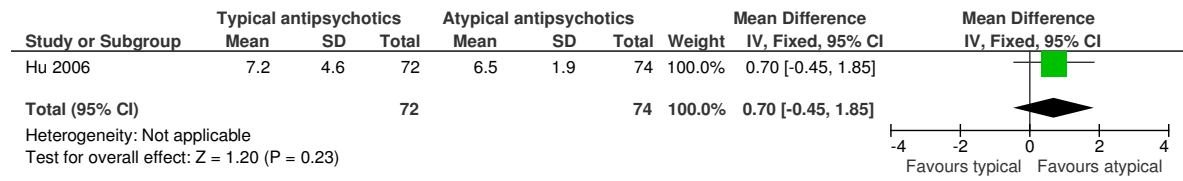
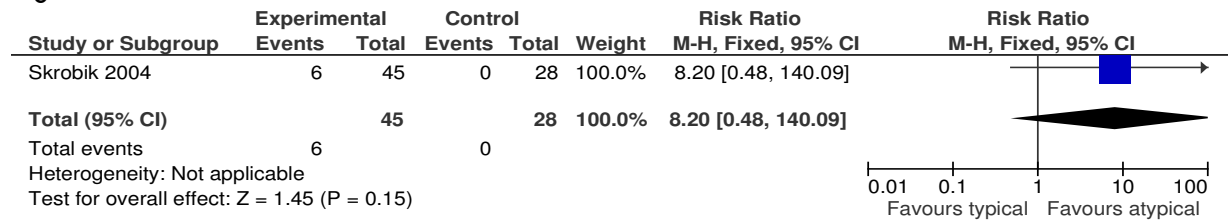


Figure 13.8: severity of delirium



NB: Scale -4 to 4

Figure 13.9 Adverse events



Adverse effects

Figure 14.1: antipsychotics as a risk factor for stroke

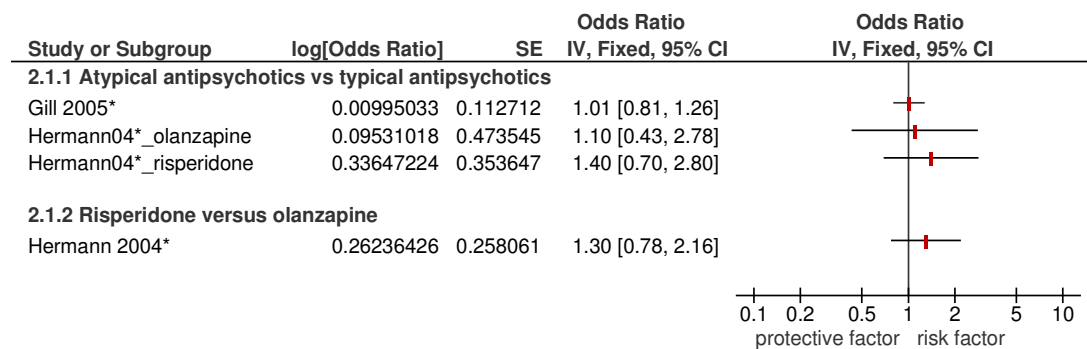


Figure 14.2: antipsychotics as a risk factor for stroke

