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1 Recommendations

Parents and carers have the right to be involved in discussions and make informed decisions about their baby's care, as described in [your care](#).

[Making decisions using NICE guidelines](#) explains how we use words to show the strength (or certainty) of our recommendations, and has information about prescribing medicines (including off-label use), professional guidelines, standards and laws (including on consent and mental capacity), and safeguarding.

2 1.1 ***Risk factors for bronchopulmonary dysplasia***

3 1.1.1 Be aware that the risk factors for bronchopulmonary dysplasia (BPD)
4 include those shown in table 1:

5 **Table 1 Identified risk factors for bronchopulmonary dysplasia^a**

In babies born before 32 weeks	<ul style="list-style-type: none"> • lower gestational age • lower birthweight • small for gestational age • male sex • core body temperature of less than 35°C on admission to neonatal unit • clinical sepsis with or without positive blood cultures • feeding with formula milk (exclusively or in addition to breast milk) • treated with surfactant^b • treated for a patent ductus arteriosus^b
In babies born before 30 weeks	<ul style="list-style-type: none"> • cardiopulmonary resuscitation performed at birth
In babies born before 28 weeks	<ul style="list-style-type: none"> • invasive ventilation begun within 24 hours of birth, especially if it lasts longer than 48 hours
<p>^a These risk factors have been identified, but other gestational ages and other risk factors not listed here might also be associated with increased risk of bronchopulmonary dysplasia.</p> <p>^b Be aware that 'treated with surfactant' and 'treated for a patent ductus arteriosus (PDA)' may reflect the severity of the baby's condition rather than being a causal link. Surfactant should be used, and a PDA should be treated, where clinically appropriate.</p>	

6

To find out why the committee made the recommendation on risk factors for BPD and how it might affect services, see [rationale and impact](#).

1 **1.2 Respiratory support for preterm babies**

2 **Respiratory support before admission to the neonatal unit**

3 1.2.1 When stabilising preterm babies who need respiratory support, use
4 continuous positive airways pressure (CPAP) where clinically appropriate,
5 rather than invasive ventilation.

To find out why the committee made the recommendation on respiratory support before admission to the neonatal unit and how it might affect services, see [rationale and impact](#).

6 **Surfactant**

7 1.2.2 Give surfactant to preterm babies who need invasive ventilation for
8 stabilisation.

9 1.2.3 When giving surfactant¹ to a preterm baby who does not need invasive
10 ventilation, use a minimally invasive administration technique. If this is not
11 feasible, use endotracheal intubation to give surfactant, with early
12 extubation afterwards.

To find out why the committee made the recommendations on giving surfactant and how they might affect services, see [rationale and impact](#).

13 **Oxygen**

14 1.2.4 Choose between nasal cannula and incubator oxygen for preterm babies
15 who need supplemental oxygen, depending on the age of the baby and
16 their clinical stability.

¹ At the time of consultation (October 2018), some brands of surfactant did not have a UK marketing authorisation for minimally invasive administration. The prescriber should follow relevant professional guidance, taking full responsibility for the decision. Informed consent should be obtained and documented. See the General Medical Council's [Prescribing guidance: prescribing unlicensed medicines](#) for further information.

To find out why the committee made the recommendation on oxygen and how it might affect services, see [rationale and impact](#).

1 **Non-invasive ventilation techniques in the neonatal unit**

2 1.2.5 For preterm babies who need non-invasive ventilation, consider nasal
3 CPAP or nasal high-flow therapy as the primary mode of respiratory
4 support. Base the decision on the age of the baby and their prematurity.

5 **Invasive ventilation techniques in the neonatal unit**

6 1.2.6 For preterm babies who need invasive ventilation, use volume-targeted
7 ventilation (VTV) as the primary mode of respiratory support. If VTV is not
8 effective, consider high-frequency oscillatory ventilation (HFOV).

9 1.2.7 Do not use synchronised pressure-limited ventilation such as assist
10 control (AC), synchronised intermittent positive pressure ventilation
11 (SIPPV), patient-triggered ventilation (PTV), pressure support ventilation
12 (PSV) or synchronised time-cycled pressure-limited ventilation (STCPLV).

To find out why the committee made the recommendations on ventilation techniques and how they might affect services, see [rationale and impact](#).

13 **Nitric oxide**

14 1.2.8 Do not use inhaled nitric oxide for preterm babies who need respiratory
15 support for respiratory distress syndrome (RDS).

16 1.2.9 Consider inhaled nitric oxide² for preterm babies with pulmonary
17 hypoplasia.

To find out why the committee made the recommendations on nitric oxide and how they might affect services, see [rationale and impact](#).

² At the time of consultation (October 2018), inhaled nitric oxide did not have a UK marketing authorisation for this indication. The prescriber should follow relevant professional guidance, taking full responsibility for the decision. Informed consent should be obtained and documented. See the General Medical Council's [Prescribing guidance: prescribing unlicensed medicines](#) for further information.

1 **1.3** *Managing respiratory disorders*

2 **Corticosteroids**

3 1.3.1 For preterm babies who are 8 days or older and still receiving invasive
4 ventilation, consider dexamethasone³ to reduce the risk of BPD. Take into
5 account the risk factors for BPD in [table 1](#) when deciding whether to use
6 dexamethasone.

7 1.3.2 Before starting treatment with dexamethasone, discuss with parents and
8 carers the possible benefits and harms. Topics to discuss include those in
9 table 2.

10 1.3.3 For preterm babies who are less than 8 days old, be aware that
11 dexamethasone increases the risk of gastrointestinal perforation.

12 1.3.4 Do not use dexamethasone with non-steroidal anti-inflammatory drugs
13 (NSAIDs).

14 1.3.5 Monitor the blood pressure of babies who receive dexamethasone,
15 because of the risk of hypertension.

³ Although this use is common in UK clinical practice, at the time of consultation (October 2018), dexamethasone did not have a UK marketing authorisation for this indication. The prescriber should follow relevant professional guidance, taking full responsibility for the decision. Informed consent should be obtained and documented. See the General Medical Council's [Prescribing guidance: prescribing unlicensed medicines](#) for further information.

1 **Table 2 Benefits and harms of dexamethasone in preterm babies 8 days or**
 2 **older**

Mortality before discharge	There is no difference in mortality before discharge in babies who receive dexamethasone compared with babies who do not receive dexamethasone.
BPD at 36 weeks' postmenstrual age	Babies who receive dexamethasone are less likely to develop BPD compared with babies who do not receive dexamethasone. On average, if 100 preterm babies are given dexamethasone, 16 fewer babies will develop BPD compared with 100 preterm babies who do not receive dexamethasone.
Cerebral palsy	There is no difference in the incidence of cerebral palsy in babies who receive dexamethasone compared with babies who do not receive dexamethasone. However, this is uncertain because there is not much good evidence, so the possibility of cerebral palsy occurring should not be excluded.
Other neurodevelopmental outcomes (neurodevelopmental delay and neurosensory impairment)	There is no difference in other neurodevelopmental outcomes in babies who receive dexamethasone compared with babies who do not receive dexamethasone.
Days on invasive ventilation	Babies who receive dexamethasone are likely to have fewer days on invasive ventilation compared with babies who do not receive dexamethasone.
Gastrointestinal perforation	There is no evidence about gastrointestinal perforation in babies who receive dexamethasone compared with babies who do not receive dexamethasone.
Hypertension	Babies who receive dexamethasone are more likely to develop hypertension compared with babies who do not receive dexamethasone. On average, if 100 preterm babies are given dexamethasone, 8 more babies will develop hypertension compared with 100 babies who do not receive dexamethasone.
Abbreviation: BPD, bronchopulmonary dysplasia.	

3

To find out why the committee made the recommendations on dexamethasone and how they might affect services, see [rationale and impact](#).

4

1 Diuretics

To find out why the committee did not make any recommendations on diuretics, see [rationale and impact](#).

2 Caffeine

3 1.3.6 Use caffeine citrate routinely in preterm babies born at or before
4 30 weeks, starting it as early as possible and ideally before 3 days of age.

5 1.3.7 Consider stopping caffeine citrate at 33–35 weeks' corrected gestational
6 age if the baby is clinically stable.

7 1.3.8 Consider caffeine citrate for any preterm baby with apnoea.

8 1.3.9 Give a loading dose of 20 mg/kg of caffeine citrate, followed 24 hours later
9 by a maintenance dosage of 5 mg/kg once daily, increasing up to
10 20 mg/kg daily⁴ if apnoeas persist.

11 1.3.10 Consider a maintenance dosage higher than 20 mg/kg daily⁴ if therapeutic
12 efficacy is not achieved, while ensuring that safe plasma levels are
13 maintained.

To find out why the committee made the recommendations on caffeine and how they might affect services, see [rationale and impact](#).

14 Patent ductus arteriosus

15 1.3.11 Do not treat a patent ductus arteriosus (PDA) in a preterm baby unless it
16 causes a significant clinical problem, for example, difficulty weaning the
17 baby from a ventilator.

To find out why the committee made the recommendation on patent ductus arteriosus and how it might affect services, see [rationale and impact](#).

⁴ At the time of consultation (October 2018), caffeine citrate did not have a marketing authorisation for use in children and young people at this dosage. The prescriber should follow relevant professional guidance, taking full responsibility for the decision. Informed consent should be obtained and documented. See the General Medical Council's [Prescribing guidance: prescribing unlicensed medicines](#) for further information.

1

2 **1.4 Monitoring**

3 **Oxygen**

4 1.4.1 Use continuous pulse oximetry to measure oxygen saturation in preterm
5 babies, supplemented by arterial sampling if clinically indicated.

6 1.4.2 Aim for an oxygen saturation of 91–95% in preterm babies.

7 1.4.3 For preterm babies on invasive ventilation who are clinically unstable,
8 consider transcutaneous oxygen monitoring.

To find out why the committee made the recommendations on oxygen monitoring and how they might affect services, see [rationale and impact](#).

9 **Carbon dioxide**

10 1.4.4 For preterm babies on invasive ventilation, aim for a carbon dioxide partial
11 pressure (pCO₂) of:

- 12 • 4.5–8.5 kPa on days 1–3 **and**
- 13 • 4.5–10 kPa from day 4 onwards.

14 1.4.5 Reduce minute ventilation without delay in preterm babies with low pCO₂,
15 and check the pCO₂ within an hour of the low measurement being
16 identified.

To find out why the committee made the recommendations on carbon dioxide monitoring and how they might affect services, see [rationale and impact](#).

17 **Blood pressure**

18 1.4.6 Do not treat preterm babies for hypotension based solely on specific blood
19 pressure thresholds, but take into account other factors, such as evidence
20 of poor tissue perfusion. The aim of treatment should be to improve
21 perfusion.

To find out why the committee made the recommendation on blood pressure and how it might affect services, see [rationale and impact](#).

1

2 **1.5 Sedation and analgesia**

3 **Morphine**

4 1.5.1 Do not routinely use morphine for preterm babies on respiratory support.

5 1.5.2 Consider morphine⁵ if the baby is in pain, using a validated pain score.

6 1.5.3 Reassess babies on morphine regularly to ensure that morphine is
7 stopped as soon as possible.

To find out why the committee made the recommendations on morphine and how they might affect services, see [rationale and impact](#).

8 **Premedication before intubation**

9 1.5.4 Consider premedication before elective non-urgent intubation in preterm
10 babies.

11 1.5.5 If giving premedication, consider either:

- 12 • an opioid analgesic (for example, morphine⁵ or fentanyl⁶), combined
- 13 with a neuromuscular blocking agent (for example, suxamethonium) **or**
- 14 • propofol⁷ alone.

⁵ Although this is common in UK clinical practice, at the time of consultation (October 2018), morphine did not have a UK marketing authorisation for children under 12 years (intravenous administration) or under 1 year (oral administration). The prescriber should follow relevant professional guidance, taking full responsibility for the decision. Informed consent should be obtained and documented. See the General Medical Council's [Prescribing guidance: prescribing unlicensed medicines](#) for further information.

⁶ Although this is common in UK clinical practice, at the time of consultation (October 2018), fentanyl did not have a UK marketing authorisation for children under 2 years. The prescriber should follow relevant professional guidance, taking full responsibility for the decision. Informed consent should be obtained and documented. See the General Medical Council's [Prescribing guidance: prescribing unlicensed medicines](#) for further information.

⁷ Although this is common in UK clinical practice, at the time of consultation (October 2018), propofol did not have a UK marketing authorisation for children under 1 month. The prescriber should follow relevant professional guidance, taking full responsibility for the decision. Informed consent should be

To find out why the committee made the recommendations on premedication for intubation and how they might affect services, see [rationale and impact](#).

1

2 **1.6 *Involving, supporting and informing parents and carers***

3 **Involving parents and carers while their preterm baby is on respiratory support**

4 1.6.1 Explain to the parents and carers of preterm babies on respiratory support
5 that non-nutritive sucking (using a dummy):

- 6 • is beneficial during nasogastric tube feeds if the baby is awake,
7 because it can reduce the length of the baby's hospital stay **and**
- 8 • can help soothe the baby between feeds.

9 1.6.2 Consider providing the Newborn individualized developmental care and
10 assessment program (NIDCAP®) to improve cognitive development in
11 babies born at less than 27 weeks.

12 1.6.3 Tell parents and carers about the benefits of using touch to communicate
13 with their baby, for example, through skin-to-skin contact.

To find out why the committee made the recommendations on involving parents and carers and how they might affect services, see [rationale and impact](#).

14 **Supporting parents and carers while their preterm baby is on respiratory** 15 **support**

16 1.6.4 Recognise parents and carers as partners in their baby's care, and
17 support them in this role.

18 1.6.5 Encourage and support parents and carers to:

- 19 • be involved in planning and providing their baby's day-to-day care, for
20 example, feeding and nappy changing

obtained and documented. See the General Medical Council's [Prescribing guidance: prescribing unlicensed medicines](#) for further information.

- 1 • participate in discussions and decisions about their baby during ward
2 rounds, providing input into planning and providing care.

3 1.6.6 Provide regular opportunities and time for parents and carers to discuss
4 their baby's care, ask questions about the information they have been
5 given, and discuss concerns.

6 1.6.7 Give parents and carers the time, support and encouragement they need
7 to become confident in caring effectively for their baby.

8 1.6.8 Offer parents and carers psychological support from a professional who is
9 trained to deliver this type of help and advice.

10 **Providing information to parents and carers while their preterm baby is on**
11 **respiratory support**

12 1.6.9 Ask parents and carers about how and when they would like to receive
13 information about their baby's treatment and progress.

14 1.6.10 Support discussions with parents and carers using written information.
15 Ensure that information is up to date, relevant, appropriate to the parents'
16 and carers' needs and preferences, and consistent between healthcare
17 professionals. For more guidance on communication (including different
18 formats and languages), providing information, and shared decision-
19 making, see the NICE guideline on [patient experience in adult NHS](#)
20 [services](#).

21 1.6.11 Ensure that information for parents and carers is delivered by an
22 appropriate healthcare professional, and information for hospitalised
23 mothers who cannot visit their baby is delivered by a senior healthcare
24 professional, for example, a neonatologist or specialist registrar, face-to-
25 face whenever possible.

26 1.6.12 Be sensitive about the timing of discussions with parents and carers. In
27 particular, discuss significant perinatal events without delay, providing the
28 mother has sufficiently recovered from the birth.

29 1.6.13 Provide information for parents and carers that includes:

- 1 • explanations and regular updates about their baby's condition and
- 2 treatment, especially if there are any changes
- 3 • what happens in the neonatal unit, and the equipment being used to
- 4 support their baby
- 5 • what respiratory support is being provided for their baby
- 6 • how to get involved in their baby's day-to-day care, interact with their
- 7 baby and interpret the baby's neurobehavioural cues
- 8 • the roles and responsibilities of different members of their baby's
- 9 healthcare team, and key contacts
- 10 • information about caring for a premature baby to share with family and
- 11 friends, and practical suggestions about how to get help and support
- 12 from family and friends
- 13 • opportunities for peer support from neonatal unit graduate parents or
- 14 parent buddies
- 15 • details of local support groups, online forums and national charities,
- 16 and how to get in touch with them.

To find out why the committee made the recommendations on supporting and informing parents and carers and how they might affect services, see [rationale and impact](#).

17 **Neonatal services for preterm babies on respiratory support**

18 1.6.14 Those responsible for planning and delivering neonatal services should
19 ensure that neonatal units:

- 20 • are welcoming and friendly
- 21 • foster positive and supportive relationships by providing parents and
- 22 carers with 24-hour access to their baby
- 23 • provide privacy for skin-to-skin contact and feeding
- 24 • have private areas for difficult conversations
- 25 • have comfortable furniture and provide a relaxing environment for
- 26 families.

- 1 1.6.15 Ensure that healthcare professionals in neonatal units can support
2 parents and carers by being competent in:
- 3 • communicating complex and sensitive information clearly
 - 4 • tailoring information and support to the person's individual needs and
5 circumstances.

To find out why the committee made the recommendations on neonatal services and how they might affect services, see [rationale and impact](#).

6

7 **1.7 Discharge planning**

8 **Planning safe discharge from the neonatal unit for preterm babies on** 9 **respiratory support**

- 10 1.7.1 Neonatal units should consider appointing a member of staff as a
11 designated neonatal discharge coordinator to discuss the following with
12 parents and carers:

- 13 • ongoing support and follow-up after discharge (also see the NICE
14 guideline on [developmental follow-up of children and young people
15 born preterm](#))
- 16 • how to care for their baby at home
- 17 • how to use specialist equipment safely
- 18 • how to travel with their baby and specialist equipment.

- 19 1.7.2 When planning to discharge a preterm baby on respiratory support from
20 the neonatal unit:

- 21 • follow the principles in the NICE guideline on [postnatal care](#)
- 22 • consider early referral to, and regular contact with, community and
23 continuing healthcare teams
- 24 • consider an interim discharge placement to, for example, a hospice,
25 alternative family member's home, step-down unit, transitional care
26 unit, or alternative suitable accommodation, where appropriate.

To find out why the committee made the recommendations on planning safe discharge and how they might affect services, see [rationale and impact](#).

1 **Supporting and informing parents and carers of preterm babies on respiratory** 2 **support – preparing for discharge**

3 1.7.3 Recognise parents and carers as partners in the discharge planning
4 process. Answer their questions and concerns as they arise, and support
5 them in making joint decisions with the discharge team.

6 1.7.4 Throughout the baby's neonatal admission, provide support and guidance
7 for parents and carers with constructive and supportive feedback about
8 how to care for their baby and how to use specialist equipment. Use a
9 formal competency-based assessment tool to evaluate the safe use of
10 specialist equipment.

11 1.7.5 Discuss any modifications that parents and carers might need to make to
12 their home as soon as possible.

13 1.7.6 Educate parents and carers about possible emergencies that may arise,
14 how to deal with them and who to contact for help and advice. This should
15 include how to carry out cardiopulmonary resuscitation, and what to do if
16 there are problems with any specialist equipment.

17 1.7.7 Provide parents and carers with opportunities to care for their baby
18 overnight.

19 1.7.8 Provide information for parents and carers to help them care for their baby
20 safely and confidently after discharge. Follow the principles on
21 communication and information-giving in section 1.6 of this guideline, and
22 also see the NICE guideline on [postnatal care](#). Information should include:

- 23 • how to recognise signs of deterioration in their baby, and what to do
- 24 • how to adapt routines such as feeding and sleeping after discharge,
25 and information about safe sleep guidance
- 26 • how to make follow-up appointments and timing of immunisations

- 1 • who to contact after discharge, as well as a list of useful medical
2 contacts.

3 1.7.9 Tell parents and carers about sources of support after discharge, for
4 example:

- 5 • opportunities for peer support
6 • help and support for their own needs, for example, postnatal
7 depression (also see the NICE guideline on [antenatal and postnatal](#)
8 [mental health](#)).

To find out why the committee made the recommendations on supporting and informing parents as part of discharge planning, and how they might affect services, see [rationale and impact](#).

9

10 ***Terms used in this guideline***

11 **Automated oxygen titration**

12 A control system that measures the oxygen saturation and automatically adjusts the
13 oxygen flow to maintain the oxygen saturation within a predefined target range.

14 **Invasive ventilation**

15 Administration of respiratory support via an endotracheal tube or tracheostomy,
16 using a mechanical ventilator.

17 **Minimally invasive administration technique**

18 Administration of surfactant through a small endotracheal catheter without insertion
19 of an endotracheal tube or invasive ventilation.

20 **Minute ventilation**

21 The tidal volume of each breath in millilitres (ml) multiplied by the number of breaths
22 per minute gives the minute ventilation in ml/min (usually expressed as ml/kg/min,
23 which is achieved by dividing by the baby's weight in kg).

1 **Neurobehavioural cues**

2 Sounds, characteristics of movements including facial expressions and physiological
3 parameters such as heart rate, breathing patterns and skin tone that reflect the
4 baby's current level of sensitivity or wellbeing, and reveal their current developmental
5 stage.

6 **Neurodevelopmental outcomes**

7 In this guideline, neurodevelopmental outcomes at 18 months or older have been
8 defined as:

- 9 • cerebral palsy (reported as presence or absence of condition, not severity)
- 10 • neurodevelopmental delay (reported as dichotomous outcomes, not continuous
11 outcomes such as mean change in score):
 - 12 – severe (score of more than 2 standard deviation [SD] below normal on
13 validated assessment scales, or a score of less than 70 on the Bayley scale of
14 infant development mental developmental index [MDI] or psychomotor
15 developmental index [PDI], or complete inability to assign score because of
16 cerebral palsy or severe cognitive delay)
 - 17 – moderate (score of 1–2 SD below normal on validated assessment scales, or a
18 score of 70–84 on the Bayley scale of infant development MDI or PDI)
- 19 • neurosensory impairment (reported as presence or absence of condition, not
20 severity):
 - 21 – severe hearing impairment (for example, deaf)
 - 22 – severe visual impairment (for example, blind).

23 **Non-invasive ventilation**

24 Administration of respiratory support using a ventilator or flow driver, but not via an
25 endotracheal tube or tracheostomy.

26 **Perinatal**

27 In this guideline, the perinatal period is defined as the period of time from 48 hours
28 before birth up until 7 completed days after birth.

1 **Preterm**

2 A baby born before 37 weeks. This can be subdivided further:

- 3 • extremely preterm: babies born at less than 28 weeks
4 • very preterm: babies born between 28 and 31⁺⁶ weeks
5 • moderate to late preterm: babies born between 32 and 36⁺⁶ weeks.

6 **Skin-to-skin contact**

7 Holding a baby on the skin of a parent or carer, usually on the chest.

8 **Stabilisation**

9 Facilitating and supporting a smooth transition from fetal to neonatal life. The
10 process involves careful assessment of heart rate, colour (oxygenation) and
11 breathing, with provision of appropriate interventions where indicated.

12 **Recommendations for research**

13 ***Key recommendations for research***

14 **1 Non-invasive ventilation techniques**

15 What is the effectiveness of high-pressure non-invasive positive pressure ventilation
16 (NIPPV) compared with continuous positive airways pressure (CPAP) flow driver as
17 the primary mode of ventilation?

18 To find out why the committee made the research recommendation on non-invasive
19 ventilation techniques, see [rationale and impact](#).

20 **2 Surfactant**

21 What is the best technique for delivering surfactant in a minimally invasive manner?

22 To find out why the committee made the research recommendation on surfactant,
23 see [rationale and impact](#).

24 **3 Diuretics**

25 What is the effectiveness of diuretics compared with placebo in preventing
26 bronchopulmonary dysplasia (BPD) in preterm babies on respiratory support?

1 To find out why the committee made the research recommendation on diuretics, see
2 [rationale and impact](#).

3 **4 Oxygen monitoring**

4 Does targeting higher oxygen saturations of 92–97% in preterm babies lead to
5 improved survival without significant complications?

6 To find out why the committee made the research recommendation on oxygen
7 monitoring, see [rationale and impact](#).

8 **5 Premedication before intubation**

9 What is the most effective combination of an analgesic with a neuromuscular
10 blocker, or an analgesic with an anaesthetic agent, for premedication in preterm
11 babies requiring elective or semi-elective intubation?

12 To find out why the committee made the research recommendation on
13 premedication before intubation, see [rationale and impact](#).

14 ***Other recommendations for research***

15 **Respiratory support before admission to the neonatal unit**

16 Does CPAP plus prophylactic surfactant, administered by a non-invasive technique
17 in the delivery room, improve outcomes compared with CPAP alone in preterm
18 babies?

19 **Surfactant**

20 What is the optimal dosing regimen of surfactant when delivered in a minimally
21 invasive manner?

22 **Oxygen administration**

23 What is the effectiveness of humidified and non-humidified supplemental low-flow
24 oxygen in preterm babies?

25 What should be the target oxygen saturation range for preterm babies when using an
26 automated oxygen titration system that creates a normal frequency saturation curve?

1 **Invasive ventilation techniques**

2 Are there differences in the long-term neurodevelopmental outcomes for preterm
3 babies receiving volume-targeted ventilation (VTV) compared with high-frequency
4 oscillatory ventilation (HFOV) as their primary mode of ventilation?

5 **Corticosteroids**

6 What is the comparative efficacy of hydrocortisone compared with dexamethasone
7 for preventing BPD in preterm babies requiring respiratory support?

8 Is nebulised budesonide effective compared to placebo in preventing BPD in preterm
9 babies requiring respiratory support?

10 **Diuretics**

11 What is the effectiveness of diuretics compared with placebo in the treatment of BPD
12 in preterm babies on respiratory support?

13 **Caffeine**

14 What is the optimal maintenance dose of caffeine citrate in order to optimise
15 neurodevelopmental outcomes in preterm babies?

16 **Patent ductus arteriosus**

17 Are any echocardiographic parameters able to improve the predictive course of
18 patent ductus arteriosus (PDA) and therefore suggest a group of babies who would
19 benefit from PDA treatment?

20 **Oxygen monitoring**

21 What is the accuracy of pulse oximetry and transcutaneous measurement of partial
22 pressure of oxygen compared with arterial oxygen levels for detecting hyperoxia and
23 hypoxia in preterm babies?

24 **Carbon dioxide monitoring**

25 What is the optimal carbon dioxide target range in preterm babies on non-invasive
26 ventilation at different gestational ages?

1 **Blood pressure**

2 What is the optimal method and frequency of measuring blood pressure for preterm
3 babies requiring respiratory support?

4 What is the optimal target blood pressure range for preterm babies requiring
5 respiratory support?

6 **Morphine**

7 What is the effectiveness of morphine compared with containment holding for
8 preterm babies receiving respiratory support?

9 **Involving parents and carers**

10 What is the impact of parental involvement as part of Family integrated care (FIC) or
11 the Newborn individualised developmental care and assessment programme
12 (NIDCAP®) on the incidence of bronchopulmonary dysplasia and length of hospital
13 stay in preterm babies?

14 **Discharge planning**

15 What is best practice around discharge planning for preterm babies on respiratory
16 support?

17

18 **Rationale and impact**

19 These sections briefly explain why the committee made the recommendations and
20 how they might affect services. They link to details of the evidence and a full
21 description of the committee's discussion.

22 ***Risk factors for bronchopulmonary dysplasia***

23 Recommendation [1.1.1](#)

24 **Why the committee made the recommendation**

25 There was evidence that lower gestational age, lower birth weight, being small for
26 gestational age, male sex, lower body temperature, sepsis, any formula feeding,
27 surfactant use, treatment for a patent ductus arteriosus (PDA), cardiopulmonary

1 resuscitation and mechanical ventilation, are all independent risk factors for
2 bronchopulmonary dysplasia (BPD) in preterm babies.

3 There was no evidence of a link between antenatal steroids, chorioamnionitis,
4 intrauterine growth restriction, ethnicity or race, or postnatal steroid use, and BPD.
5 However, the committee did not prioritise these areas for further research.

6 The committee was concerned that including surfactant use and treatment for PDA
7 as risk factors for BPD could lead to a reduction in surfactant use and PDA
8 treatment. They agreed that there was unlikely to be a causal link – rather, the
9 increased risk of BPD associated with these factors is more likely to reflect the
10 severity of the baby’s condition, and that surfactant should be used, and a PDA
11 should be treated, where clinically appropriate.

12 The committee noted that there was an absence of evidence for certain risk factors
13 for BPD; some evidence was for specific gestational ages at birth from which the
14 committee was unable to extrapolate to other gestational ages, and for some risk
15 factors, the evidence was underpowered to detect an effect. The committee
16 therefore concluded that other gestational ages and other risk factors not listed here
17 might also be associated with increased risk of BPD.

18 No evidence was found for some of the potential risk factors that had been
19 suggested by the committee (such as necrotising enterocolitis and supplementary
20 oxygen), but these were not prioritised by the committee for further research.

21 **How the recommendation might affect services**

22 Knowledge of BPD risk factors means healthcare professionals can identify preterm
23 babies who are more likely to develop BPD, and prioritise treatment regimens
24 accordingly. This may reduce the incidence of BPD, which will lead to long-term
25 savings for the NHS.

26 Full details of the evidence and the committee’s discussion are in [evidence review A:
27 diagnosing respiratory disorders](#).

28 [Return to recommendations](#)

29

1 ***Respiratory support before admission to the neonatal unit***

2 Recommendation [1.2.1](#)

3 **Why the committee made the recommendation**

4 The evidence did not show a clear difference between continuous positive airways
5 pressure (CPAP) alone and invasive ventilation with surfactant when used in preterm
6 babies in the delivery room, for any of the outcomes that the committee had
7 prioritised (mortality, BPD and neurodevelopmental outcomes). However, the
8 evidence showed a possible reduction in mortality before discharge, and a possible
9 reduction in the incidence of BPD at 36 weeks' postmenstrual age (PMA) with CPAP.

10 One large study found that just over half of those who received CPAP instead of
11 intubation did need to be intubated at some point during their hospitalisation.
12 However, the committee agreed that this was a very positive result, as around half of
13 babies avoided all the risks of invasive intervention.

14 However, the committee agreed that it is preferable to avoid invasive ventilation
15 wherever possible, so agreed that when stabilising a preterm baby in the delivery
16 room, the non-invasive ventilation technique of CPAP should be used rather than
17 invasive ventilation with surfactant, unless clinically inappropriate (for example, the
18 baby is not breathing and requires invasive ventilation). The committee agreed that
19 this approach would not be suitable for preterm babies born very early, for example
20 at less than 25 weeks, because these babies may not have the necessary
21 respiratory drive, and because the failure rate of non-invasive ventilation is high in
22 babies of this age. The committee agreed that for these very young preterm babies,
23 it may be more practical to use invasive ventilation with surfactant in the delivery
24 room, but as this would be a clinical decision it was not appropriate to set a particular
25 age cut-off.

26 Because there was not enough evidence to make recommendations on the use of
27 CPAP with surfactant compared to CPAP without surfactant in the delivery room, the
28 committee recommended that further research be done in this area.

1 **How the recommendation might affect services**

2 Current practice in most units is to routinely intubate preterm babies (below a certain
3 gestation, often 27–28 weeks, but specific cut-offs will vary) and give surfactant, so
4 this will be a change in practice for these units. Because CPAP is associated with
5 lower costs than invasive ventilation, this change is likely to lead to cost savings.

6 Full details of the evidence and the committee’s discussion are in [evidence review B:
7 respiratory support](#).

8 [Return to recommendations](#)

9

10 **Surfactant**

11 Recommendations [1.2.2 and 1.2.3](#)

12 **Why the committee made the recommendations**

13 It is established clinical practice in the UK to give surfactant to preterm babies
14 needing invasive ventilation, based on good evidence and extensive clinical
15 experience, so the committee agreed to make a recommendation that reinforces this.

16 In preterm babies who do not require invasive ventilation, there was evidence that
17 minimally invasive surfactant administration techniques reduce the incidence of BPD,
18 the number of days on invasive ventilation, and the incidence of pneumothorax,
19 compared with endotracheal administration.

20 However, not all neonatal units have the facilities to carry out minimally invasive
21 surfactant administration techniques, and not all healthcare professionals have been
22 trained to use them. The committee agreed that in these circumstances,
23 endotracheal surfactant administration followed by early extubation should be used,
24 because there was evidence that it reduces the incidence of BPD compared with
25 conventional administration of surfactant with continued ventilation.

26 Because there was not enough good evidence to make recommendations on which
27 minimally invasive administration technique leads to the best outcomes, or on

1 different surfactant dosing regimens, the committee recommended that further
2 research be done in these areas.

3 **How the recommendations might affect services**

4 Current practice for giving surfactant to preterm babies varies among neonatal units
5 because of differences in available facilities and training. The recommendations may
6 increase the trend towards using less invasive techniques of surfactant
7 administration. Neonatal units that currently use conventional endotracheal
8 administration of surfactant may therefore change practice to use minimally invasive
9 techniques or to extubate earlier.

10 Full details of the evidence and the committee's discussion are in [evidence review B:
11 respiratory support](#).

12 [Return to recommendations](#)

13

14 **Oxygen**

15 Recommendation [1.2.4](#)

16 **Why the committee made the recommendation**

17 There was no evidence to suggest any difference in the effectiveness or safety of
18 oxygen delivered by nasal cannula compared with oxygen delivered in the incubator.
19 The committee agreed that the decision about whether to deliver oxygen by nasal
20 cannula or in the incubator would depend on factors such as the age of the baby at
21 birth and how clinically stable they are.

22 There was evidence that automated oxygen titration reduces the number of days on
23 oxygen, reduces the number of manual adjustments for titration, and increases the
24 time that preterm babies spend in the optimal target oxygen saturation range.
25 However, the committee were concerned, based on their clinical knowledge, that the
26 cumulative frequency oxygen curves for oxygen saturation achieved by automated
27 titration may lead to the mean saturation level achieved by babies being reduced
28 (due to the normal distribution of the frequency-saturation curve) compared to
29 manual adjustments (where the frequency-saturation curve is skewed to the higher

1 end of the target saturation range). The committee therefore made a research
2 recommendation to determine the optimal target oxygen saturation range for use in
3 conjunction with an automated oxygen titration system.

4 There was no evidence comparing humidified to non-humidified oxygen, so the
5 committee made a research recommendation.

6 **How the recommendation might affect services**

7 The recommendation to use nasal cannula or incubator oxygen reflects current
8 clinical practice.

9 Full details of the evidence and the committee's discussion are in [evidence review B:
10 respiratory support](#).

11 [Return to recommendations](#)

12

13 ***Ventilation techniques***

14 Recommendations [1.2.5 to 1.2.7](#)

15 **Why the committee made the recommendations**

16 ***Non-invasive ventilation techniques***

17 The available evidence made it difficult to differentiate between the non-invasive
18 ventilation techniques. The evidence showed that nasal high-flow therapy had the
19 highest probability of being the best technique for reducing mortality before
20 discharge, compared with other non-invasive ventilation techniques. However, the
21 committee agreed that babies born extremely preterm are less likely to manage
22 successfully on nasal high-flow therapy as the primary mode of ventilation when
23 compared to babies born less preterm.

24 The evidence showed a reduction in the failure of non-invasive ventilation with CPAP
25 compared with nasal high-flow therapy. Using their clinical experience, the
26 committee agreed that CPAP would be a more suitable option for use in babies born
27 more preterm.

1 Because of the lack of good evidence, the committee agreed that CPAP or nasal
2 high-flow therapy should be used as a primary mode of ventilation in preterm babies
3 who need non-invasive ventilation, with the decision on which option to use being
4 made for individual babies, and depending on their age.

5 There was evidence that nasal intermittent positive pressure ventilation (NIPPV) had
6 lower rates of failed non-invasive ventilation and fewer days on invasive ventilation
7 than CPAP, but the delivery of NIPPV in the studies was significantly different to
8 routine clinical practice in the UK, so the committee recommended that further
9 research should be carried out comparing NIPPV and CPAP.

10 ***Invasive ventilation techniques***

11 There was evidence from the network meta-analysis that volume-targeted ventilation
12 (VTV) has the highest probability of being the best technique, both for mortality
13 before discharge and BPD at 36 weeks.

14 The committee agreed that VTV may not be appropriate for all preterm babies, for
15 example, if there is an air leak. There was evidence that if VTV is not effective, high-
16 frequency oscillatory ventilation (HFOV) should be considered as an alternative.

17 The committee agreed that synchronised pressure-limited ventilation should be
18 avoided because the evidence showed an increase in the incidence of mortality
19 before discharge, compared with non-synchronised pressure-limited ventilation,
20 HFOV and VTV. The evidence also showed an increase in days on invasive
21 ventilation and pneumothorax, compared with VTV.

22 The evidence from the pair-wise analysis showed no significant difference between
23 HFOV and VTV, and there was no evidence on neurodevelopmental outcomes at
24 18 months or older, so the committee recommended that further research should be
25 carried out.

26 **How the recommendations might affect services**

27 The recommendations should reinforce current clinical practice and lead to greater
28 consistency.

1 Full details of the evidence and the committee’s discussion are in [evidence review B:](#)
2 [respiratory support](#).

3 [Return to recommendations](#)

4

5 ***Nitric oxide***

6 Recommendations [1.2.8 and 1.2.9](#)

7 **Why the committee made the recommendations**

8 There was no evidence of benefit for inhaled nitric oxide in preterm babies who need
9 respiratory support for respiratory distress syndrome (RDS). There was some
10 evidence of adverse effects, and the treatment is unlikely to be cost effective. The
11 exception is for preterm babies with pulmonary hypoplasia in whom there may be
12 some survival benefits.

13 No research recommendations were made because the committee agreed that this
14 area is not a priority area for further research.

15 **How the recommendations might affect services**

16 The recommendations will reduce the use of inhaled nitric oxide for preterm babies
17 who need respiratory support, which may lead to cost savings to the NHS given the
18 high acquisition cost of inhaled nitric oxide.

19 Full details of the evidence and the committee’s discussion are in [evidence review B:](#)
20 [respiratory support](#).

21 [Return to recommendations](#)

22

23 ***Corticosteroids***

24 Recommendations [1.3.1 to 1.3.5](#)

1 **Why the committee made the recommendations on dexamethasone**

2 There was evidence that in babies 8 days or older, dexamethasone reduces the
3 incidence of BPD, but dexamethasone is associated with an increased risk of
4 hypertension. There was some evidence suggesting that dexamethasone reduces
5 the number of days on invasive ventilation.

6 In babies younger than 8 days, there was evidence that dexamethasone reduces the
7 incidence of BPD but is associated with an increased risk of gastrointestinal
8 perforation.

9 In babies 8 days or older, there was no evidence that dexamethasone is associated
10 with an increased risk of cerebral palsy or gastrointestinal perforation. However, the
11 committee emphasised that this lack of evidence should not be considered an
12 absence of effect.

13 There were no clinically important differences in mortality before discharge, or other
14 neurodevelopmental outcomes between babies who received dexamethasone and
15 those who did not.

16 The committee recommended that dexamethasone be considered for babies 8 days
17 or older, after taking into account risk factors for BPD. This is in line with current
18 practice, which is to use corticosteroids to assist weaning from ventilatory support
19 when a baby is 8 days or older, rather than using corticosteroids as 'prophylaxis' for
20 babies less than 8 days old.

21 The committee agreed the importance of discussing the risks of gastrointestinal
22 perforation, hypertension and cerebral palsy with parents and carers before starting
23 dexamethasone therapy, because there may be lifelong implications for the baby
24 and their family.

25 Although the combination of dexamethasone and non-steroidal anti-inflammatory
26 drugs (NSAIDs) was not reviewed, the committee confirmed that they should not be
27 used together because this increases the risk of gastrointestinal bleeding and
28 perforation. The committee agreed that although this risk is widely recognised, it
29 should be reinforced in the guideline to ensure that dexamethasone and NSAIDs are
30 not used together in clinical practice.

1 Because of the increased risk of hypertension with dexamethasone, the committee
2 recommended that babies' blood pressure should be monitored. There was no
3 evidence about when or for how long to monitor blood pressure, so the committee
4 agreed that this should be decided by the neonatologist responsible for the baby's
5 care.

6 The evidence did not show any differences between different dosing strategies, and
7 so the committee did not make any specific dosing recommendations.

8 **Why the committee didn't make any recommendations on hydrocortisone and** 9 **nebulised budesonide**

10 Evidence comparing hydrocortisone and placebo was inconclusive so the committee
11 did not make any recommendations. The committee was aware there is an ongoing,
12 large multicentre randomised controlled trial investigating hydrocortisone compared
13 with placebo in preterm babies who need respiratory support, so did not make a
14 research recommendation that would replicate this study. However, they agreed that
15 a comparison of dexamethasone and hydrocortisone could provide useful guidance
16 and so made a research recommendation for this comparison.

17 There was very little evidence for the use of nebulised budesonide and therefore the
18 committee made a research recommendation.

19 **How the recommendations might affect services**

20 Current practice is to use corticosteroids in preterm babies to assist weaning or
21 removal from ventilatory support, but they are not routinely used to prevent BPD in
22 all preterm babies. The choice of dexamethasone or hydrocortisone varies among
23 neonatal units. These recommendations are unlikely to affect how often
24 corticosteroids are used, but they might prompt units who currently use
25 hydrocortisone to consider dexamethasone as an alternative.

26 Full details of the evidence and the committee's discussion are in [evidence review C:
27 managing respiratory disorders](#).

28 [Return to recommendations](#)

29

1 ***Diuretics***

2 **Why the committee did not make any recommendations**

3 The evidence on the use of diuretics in preterm babies on respiratory support was
4 very limited. None of the studies identified assessed critical outcomes such as
5 mortality before discharge, BPD or neurodevelopmental outcomes. Although the
6 studies looked at short-term adverse effects associated with diuretics, it was not
7 clear whether there was an increased risk of adverse effects because of the small
8 sample size of the studies.

9 Because of the limited evidence and lack of clinical consensus, the committee could
10 not make any recommendations for or against diuretic use in preterm babies on
11 respiratory support. Instead, the committee recommended that further research be
12 done in this area.

13 **How the recommendations might affect services**

14 Although they did not make any recommendations, some of the committee members
15 thought that the lack of evidence identified may lead to healthcare professionals
16 reviewing their use of diuretics. This may lead to a reduction in the use of diuretics in
17 preterm babies on respiratory support, at least until further evidence is available.

18 Full details of the evidence and the committee's discussion are in [evidence review C:
19 managing respiratory disorders](#).

20 [Return to recommendations](#)

21

22 ***Caffeine***

23 Recommendations [1.3.6 to 1.3.10](#)

24 **Why the committee made the recommendations**

25 There was evidence that in preterm babies born before 31 weeks, caffeine reduces
26 the incidence of BPD, cerebral palsy (at 18–21 months' follow-up) and blindness (at
27 11-year follow-up) compared to placebo. Based on their clinical experience, the

1 committee agreed that administering caffeine would also reduce apnoea in older
2 preterm babies.

3 There was evidence that, compared with lower doses, higher doses of caffeine
4 reduce the incidence of BPD, continued apnoea and extubation failure.

5 Evidence showed that the treatment with caffeine before 3 days of age may lead to a
6 reduction in BPD. There was also evidence that treatment with caffeine for 15–
7 30 days reduces the incidence of BPD compared to a shorter duration, and that
8 treatment for greater than 30 days reduces the incidence of necrotising enterocolitis
9 compared with treatment for less than 15 days.

10 To determine when caffeine should be stopped, the committee referred back to the
11 studies and identified the age at which caffeine was started, the duration of caffeine,
12 and hence the age at which it had been stopped. The committee noted that caffeine
13 had been stopped in the studies between 33 and 35 weeks. This reflected the clinical
14 experience of the committee as the age at which preterm babies were no longer
15 expected to suffer from apnoea, and so this figure was used by the committee to
16 develop their recommendations.

17 The committee made their dosing recommendations based on evidence that a higher
18 dose is more effective than a lower dose, and on currently recommended doses
19 used in clinical practice. However, the variation in loading and maintenance doses
20 used across different clinical trials made selecting an optimal dose difficult, and
21 although higher doses appeared to improve early outcomes, there were few data on
22 long-term outcomes. For this reason, the committee recommended further research
23 to identify the maintenance dose of caffeine citrate needed to optimise
24 neurodevelopmental outcomes. The committee also discussed whether monitoring
25 caffeine levels was necessary and noted that the Evelina London Paediatric
26 Formulary advises that babies can receive 10 mg/kg of caffeine twice daily without
27 monitoring blood plasma levels (Evelina London 2015). The committee noted that
28 there are units that do not currently monitor blood levels, and increasing doses to
29 higher than 20 mg/kg daily may be a concern if units did not test blood levels at
30 these higher doses. Therefore, the committee made an additional recommendation

1 that if apnoea persists and a baby receives more than 20 mg/kg daily, caffeine levels
2 should be tested.

3 **How the recommendations might affect services**

4 The recommendations will have a minimal impact on current practice. The committee
5 noted that there is some variation in dosage regimens across the NHS, so these
6 recommendations should lead to greater consistency in the choice of dosage
7 regimens. In addition, there may be a small increase in the number of blood tests
8 performed to assess caffeine levels if higher doses are used.

9 Full details of the evidence and the committee's discussion are in [evidence review C:
10 managing respiratory disorders](#).

11 [Return to recommendations](#)

12

13 ***Patent ductus arteriosus***

14 Recommendation [1.3.11](#)

15 **Why the committee made the recommendation**

16 There was no evidence of benefit from treating a PDA, and there was evidence for
17 potential harms from treating it, with either medicines or surgery. However, the
18 committee agreed that for some babies, treatment might be appropriate, for
19 example, if there is difficulty weaning the baby from a ventilator. The committee
20 agreed that further research was needed to identify which groups of babies would
21 benefit most from PDA closure, and so made a research recommendation.

22 **How the recommendation might affect services**

23 The recommendation will reduce the unnecessary treatment of PDA and the number
24 of babies exposed to potential harms from its treatment. The recommendations may
25 result in cost savings because fewer procedures will be carried out.

26 Full details of the evidence and the committee's discussion are in [evidence review C:
27 managing respiratory disorders](#).

28 [Return to recommendations](#)

1

2 ***Oxygen monitoring***

3 Recommendations [1.4.1 to 1.4.3](#)

4 **Why the committee made the recommendations**

5 The evidence on the best method for measuring oxygen levels in diagnosing
6 hyperoxia or hypoxia in preterm babies was very limited. There were no studies
7 assessing the diagnostic accuracy of SpO₂ (peripheral capillary oxygen saturation)
8 compared with the standard PaO₂ (partial pressure of arterial oxygen) that met the
9 review's inclusion criteria. The committee agreed, based on clinical consensus and
10 their experience of clinical practice, that SpO₂ should remain the first-line method for
11 continuous monitoring of oxygen saturation levels in preterm babies because of its
12 widespread availability and non-invasive nature. The committee agreed that arterial
13 sampling of partial pressure of oxygen remained the 'gold standard', but is not
14 always possible and can never be continuous.

15 The only evidence on tcPO₂ (transcutaneous oxygen) was 1 study from the 1970s,
16 and the way this procedure is performed has changed substantially since then.
17 However, tcPO₂ is currently used in clinical practice, and in the committee's
18 experience it can provide useful information. This is particularly the case for preterm
19 babies on invasive ventilation who are clinically unstable and need continuous
20 monitoring to guide management, and in whom SpO₂ may not give the most
21 accurate picture.

22 Because of the lack of good evidence, the committee agreed that further research
23 needs to be conducted looking at the diagnostic accuracy of tcPO₂ and SpO₂ against
24 the gold standard arterial oxygen saturation in diagnosing hyperoxia and hypoxia in a
25 preterm baby population.

26 There was evidence that higher target oxygen saturation levels reduce mortality.
27 Although a higher target is associated with an increase in retinopathy of prematurity
28 and an increased risk of BPD, the evidence suggested no increase in severe visual
29 impairment at 18 months, and the reduction in mortality was considered to offset the
30 increased risk of BPD. The committee were aware that target oxygen levels (up to

1 97%) may be more beneficial but there was no evidence to support this, so they
2 made a research recommendation.

3 **How the recommendation might affect services**

4 The recommendations reflect current practice, where SpO₂ is generally used as
5 routine continuous oxygen monitoring in preterm babies, and tcPO₂ is reserved for
6 the more clinically unstable preterm babies as a continuous monitoring tool.

7 Many units already use 91 to 95% as their target saturation level for preterm babies,
8 but for those that do not, this will be a change in practice. This will reduce the
9 variation in clinical practice.

10 Full details of the evidence and the committee's discussion are in [evidence review D:
11 monitoring](#).

12 [Return to recommendations](#)

13

14 ***Carbon dioxide monitoring***

15 Recommendations [1.4.4 and 1.4.5](#)

16 **Why the committee made the recommendations**

17 The evidence showed no differences in the outcomes measured between higher and
18 lower target ranges for the partial pressure of carbon dioxide in preterm babies on
19 invasive ventilation. The committee recognised that the higher target ranges
20 specified in the studies were in line with the definition of permissive hypercapnia and
21 would probably not have any detrimental effects on clinical outcomes and long-term
22 neurodevelopmental outcomes. In view of this, the committee agreed that when
23 healthcare professionals are monitoring carbon dioxide levels in preterm babies on
24 invasive ventilation, a higher target range would be acceptable. This avoids the need
25 for frequent adjustment of the ventilators to reach an extremely tight target range.

26 There was variation in the target ranges of carbon dioxide used by different studies,
27 and the range of days at which at different permissive hypercapnia levels were
28 tolerated. The committee agreed to make a recommendation in line with the largest

1 and most recent study that looked at clinical and long-term neurodevelopmental
2 outcomes, but simplified the three-stage ranges (days 1–3, days 4–6 and day 7
3 onwards) used in this study to a two-stage range based on their clinical experience
4 that the difference in upper limits tolerated would be negligible and would have
5 minimal detrimental effects on a preterm baby on invasive ventilation.

6 There was no evidence on the action to be taken when a low carbon dioxide level
7 was detected, but the committee were aware that this was a dangerous situation, so
8 agreed the action to be taken based on their clinical knowledge and experience.

9 All the evidence for the optimal target range of carbon dioxide was in preterm babies
10 on invasive ventilation. The committee recognised the lack of evidence in preterm
11 babies on non-invasive ventilation, so they recommended further research in this
12 area.

13 **How the recommendations might affect services**

14 The recommendations reflect current practice, both where permissive hypercapnia is
15 accepted in the monitoring of carbon dioxide levels in preterm babies on invasive
16 ventilation, and for the action to be taken if hypocapnia is detected.

17 Full details of the evidence and the committee's discussion are in [evidence review D:
18 monitoring](#).

19 [Return to recommendations](#)

20

21 ***Blood pressure***

22 Recommendation [1.4.6](#)

23 **Why the committee made the recommendation**

24 There was no evidence to define what blood pressure is normal in preterm babies,
25 what blood pressure is abnormal, or how blood pressure should be measured. The
26 committee wanted to make healthcare professionals aware of this lack of evidence –
27 to prevent unnecessary blood pressure monitoring and treatment, and reduce the
28 risks of adverse effects from monitoring and treatment. The committee advised,

1 based on their clinical experience, that inadequate perfusion should be treated with
2 the aim of increasing perfusion, and not to aim for a particular blood pressure target.

3 Because there was no evidence, the committee made research recommendations to
4 determine both the optimal blood pressure target and method of measuring blood
5 pressure in preterm babies.

6 **How the recommendation might affect services**

7 For units that routinely monitor blood pressure in preterm babies and treat when
8 blood pressure falls outside certain limits, this may be a change in practice. The
9 recommendation will lead to less unnecessary monitoring and treatment of blood
10 pressure.

11 Full details of the evidence and the committee's discussion are in [evidence review D:
12 monitoring](#).

13 [Return to recommendations](#)

14

15 ***Morphine***

16 Recommendations [1.5.1 to 1.5.3](#)

17 **Why the committee made the recommendations**

18 The evidence showed that there was no difference in mortality prior to discharge in
19 babies who received morphine compared to placebo. Babies receiving morphine
20 took longer to achieve full enteral feeding, and babies born at 27–29 weeks'
21 gestation had an increased risk of severe intraventricular haemorrhage (IVH). There
22 was some evidence that, when compared with placebo, morphine improves sedation
23 and pain scores in preterm babies who need invasive respiratory support during
24 infusion. However, moderate quality evidence from a larger study showed no
25 difference in pain scores during endotracheal suctioning between babies who
26 received morphine compared to placebo.

27 The only evidence available comparing morphine to fentanyl showed no clinically
28 significant difference in rates of severe IVH.

1 There was some evidence that when compared with midazolam, babies receiving
2 morphine may have decreased rates of severe IVH.

3 Babies receiving morphine experienced less pain during infusion, but less sedation
4 after infusion.

5 Because of the mixed evidence regarding the effectiveness of morphine and taking
6 into account the risks, the committee agreed that morphine should not be used
7 routinely, but may be considered when it is clear the baby is in pain (using a
8 validated pain score).

9 The committee discussed other concerns about using morphine, such as suppressed
10 respiratory drive and opioid dependency. They agreed that regular reassessments
11 are important to ensure that morphine is stopped as soon as appropriate.

12 The committee did not make any recommendations for paracetamol or non-
13 pharmacological interventions because there was no evidence available. Instead, the
14 committee recommended that further research be done to compare morphine with
15 containment holding during respiratory support, because the committee agreed that
16 containment holding may improve outcomes in preterm babies, with a reduced risk of
17 adverse events compared to pharmacological therapy.

18 **How the recommendations might affect services**

19 Use of sedation and analgesia currently varies among units. The recommendations
20 will have little impact in units that do not routinely use morphine, but other units may
21 need to change practice and this may lead to a reduction in the use of morphine. The
22 recommendations will make practice more consistent across the NHS.

23 Full details of the evidence and the committee's discussion are in [evidence review E:
24 sedation and analgesia](#).

25 [Return to recommendations](#)

26

27 ***Premedication before intubation***

28 Recommendations [1.5.4 and 1.5.5](#)

1 **Why the committee made the recommendations**

2 There was some evidence from small, single studies that using an analgesic with a
3 neuromuscular blocker, or an anaesthetic such as propofol used alone, is an
4 effective regimen to achieve successful intubation in preterm babies, while avoiding
5 adverse effects.

6 However, there was a lack of evidence to show exactly which medicines or classes
7 of medicines form the best combination, so the committee recommended that
8 healthcare professionals should consider premedication before elective intubation
9 and recommended that further research be done in this area.

10 **How the recommendations might affect services**

11 Current practice of using premedication for elective intubation in preterm babies
12 varies among units. Units that currently use single medicines (such as morphine or
13 fentanyl) may need to change practice to follow the recommendation. The
14 recommendation will make practice more consistent across the NHS.

15 Full details of the evidence and the committee's discussion are in [evidence review E:
16 sedation and analgesia](#).

17 [Return to recommendations](#)

18

19 ***Involving parents and carers while their preterm baby is on
20 respiratory support***

21 Recommendation [1.6.1 to 1.6.3](#)

22 **Why the committee made the recommendations**

23 There was good evidence that using a dummy (non-nutritive sucking) during
24 nasogastric feeds reduces the length of the baby's hospital stay. In addition, there
25 was some evidence that the Newborn individualized developmental care and
26 assessment program (NIDCAP®) improved neurodevelopmental outcomes relating to
27 cognitive development and was a cost-effective intervention in babies born at less
28 than 27 weeks. Although the evidence for skin-to-skin contact did not show any

1 benefit, there was no evidence of harm. There was no evidence that Family
2 integrated care (FIC) provided any additional benefits compared to standard care.

3 Based on their experience and the clinical evidence, the committee recommended
4 explaining to parents and carers about the potential benefits of interacting with their
5 baby because early social development and relationship-forming are key to
6 successful emotional and behavioural development.

7 Because of the limited evidence available on FIC and NIDCAP[®], the committee
8 made it a priority to recommend that further research be done to investigate the
9 potential impact of NIDCAP[®] and FIC on length of stay and BPD.

10 **How the recommendations might affect services**

11 The committee agreed that the recommendations on non-nutritive sucking and using
12 positive touch (such as containment holding or skin-to-skin contact) would not result
13 in a major change in practice, but will help improve consistency in best practice.

14 Although there are cost implications for units to train professionals in NIDCAP[®], the
15 recommendation to consider NIDCAP[®] would lead to a more consistent approach
16 across neonatal care networks to practice linked with neurodevelopmental care. It
17 would also improve parent access to this neurodevelopmental care.

18 Full details of the evidence and the committee's discussion are in [evidence review F:
19 involving and supporting parents and carers](#).

20 [Return to recommendations](#)

21

22 ***Supporting and informing parents and carers while their preterm 23 baby is on respiratory support***

24 Recommendations [1.6.4 to 1.6.13](#)

1 **Why the committee made the recommendations**

2 ***Support***

3 There was good evidence that parents value emotional, psychological and practical
4 support from staff, friends and family, peers (such as other parents of preterm
5 babies) and employers when caring for a preterm baby receiving respiratory support.
6 Parents also value professional support and counselling.

7 There was also evidence that parents value being partners in their baby's care, want
8 to be supported by staff in caring for their baby, and need to be able to develop good
9 communication and relationships with the staff caring for their baby.

10 There was evidence that parents value a comfortable, homely environment on the
11 neonatal unit that is conducive to being involved in planning and providing care for
12 their baby. Parents also value having 24-hour access to the neonatal unit, with
13 private areas and privacy when needed.

14 ***Information***

15 There was good evidence that parents and carers value high-quality, relevant,
16 consistent information about their baby's health and care, including regular updates
17 on their baby's progress. Parents and carers value information that is appropriate for
18 their needs and explained clearly to them, and value the opportunity to ask
19 questions. There was evidence that the appropriate timing of information is important
20 to parents. The evidence also showed that parents and carers prefer information to
21 be provided by an appropriate healthcare professional, and for it to be backed up by
22 written information.

23 Parents value information on a range of topics, including how to interpret their baby's
24 neurobehavioural cues, breastfeeding, skin-to-skin contact, the medical equipment
25 used, who to contact, and other sources of information they could access
26 themselves.

27 **How the recommendations might affect services**

28 The committee agreed that the recommendations would not result in a major change
29 in practice, but will help improve consistency in best practice.

1 Full details of the evidence and the committee’s discussion are in [evidence review F:](#)
2 [involving and supporting parents and carers.](#)

3 [Return to recommendations](#)

4

5 ***Neonatal unit services***

6 Recommendations [1.6.14 and 1.6.15](#)

7 **Why the committee made the recommendations**

8 There was evidence that parents and carers value having 24-hour access to the
9 neonatal unit, which should be a homely environment with comfortable furniture and
10 private areas. In a number of the support and information themes, parents and
11 carers agreed that healthcare professionals who provide information and support
12 should be trained and competent in this, so the committee made an overarching
13 recommendation.

14 **How the recommendations might affect services**

15 The committee agreed that the recommendations would not result in a major change
16 in practice, but will help improve consistency in best practice.

17 Full details of the evidence and the committee’s discussion are in [evidence review F:](#)
18 [involving and supporting parents and carers.](#)

19 [Return to recommendations](#)

20

21 ***Discharge planning – planning safe discharge***

22 Recommendations [1.7.1 and 1.7.2](#)

23 **Why the committee made the recommendations**

24 There was evidence about the importance of good communication with parents
25 about their baby's discharge. The committee agreed that a designated neonatal
26 discharge coordinator, as a single point-of-contact, would facilitate the
27 communication of key information with parents and carers. The committee also

1 agreed that early referral to community and continuing healthcare teams would also
2 help parents prepare for their baby's discharge. Having the option to discharge to an
3 alternative location, such as to another relative's home or a hospice, would enable
4 parents and carers whose homes are not suitable for their preterm baby to be able to
5 care for their baby outside the hospital

6 The committee also recognised that some of the advice in the NICE guideline on
7 postnatal care was also relevant to babies born preterm and so made a cross
8 reference to this guideline.

9 However, because there were only 2 studies, and no evidence for a number of
10 themes identified by the committee, the committee agreed that more research could
11 better define best practice, and so made a research recommendation.

12 **How the recommendations might affect services**

13 The committee agreed that the recommendations would not result in a major change
14 in practice, but will help improve consistency in delivering best practice.

15 Full details of the evidence and the committee's discussion are in [evidence review G:
16 discharge planning](#).

17 [Return to recommendations](#)

18

19 ***Discharge planning – preparing for discharge***

20 Recommendations [1.7.3 to 1.7.9](#)

21 **Why the committee made the recommendations**

22 There was evidence that parents and carers value having support and information
23 about their baby's routine care, being involved in preparing for the baby's discharge,
24 and having information on equipment, identifying illness in their baby, and dealing
25 with emergencies. Parents and carers also value information about future care, such
26 as contact details, follow-up appointments and immunisations, ongoing peer support
27 and self-care for problems such as postnatal depression.

1 **How the recommendations might affect services**

2 The committee agreed that the recommendations would not result in a major change
3 in practice, but will help improve consistency in delivering best practice.

4 Full details of the evidence and the committee's discussion are in [evidence review G:
5 discharge planning](#).

6 [Return to recommendations](#)

7

8 **Context**

9 In 2016, a [national neonatal audit](#) found that approximately 13% of babies in the UK
10 need specialist neonatal care, either because they are born preterm (at less than
11 37 weeks) or because of an illness or condition.

12 A comparison of the [EPICure studies](#) published in 2012 found that, between 1995
13 and 2006, the number of babies born at less than 26 weeks and admitted to neonatal
14 units increased by 30% in England. Over the same period, survival rates for babies
15 born at 22–25 weeks and admitted for intensive care increased by 13%. In addition,
16 a higher proportion of these babies survived without disability (particularly babies
17 born at 24–25 weeks). [International comparisons](#) show that the neonatal mortality
18 rate varies significantly by country.

19 Preterm babies are at risk of respiratory disorders, including respiratory distress
20 syndrome and bronchopulmonary dysplasia (BPD). High-quality respiratory care can
21 reduce the length of hospital stay and risk of long-term disability. BPD is particularly
22 common in preterm babies who require assisted ventilation. Babies with BPD need
23 prolonged specialist care and respiratory support.

24 Respiratory support is used in different ways in different units, and it is unclear what
25 the best method is for providing ventilation and preventing BPD. There are many
26 other areas of uncertainty and variation in how respiratory support is provided. There
27 is also variation in other areas of respiratory management, including how
28 corticosteroids are used to prevent and manage BPD.

1 Since 2013, neonatal critical care services have been managed within Operational
2 Delivery Networks. For healthy babies and babies with minor problems, most care is
3 provided by the hospital they are born in. Neonatal intensive care units are
4 responsible for babies who have more complex problems. Neonatal intensive care,
5 and the [service specifications](#) for Neonatal Critical Care and Neonatal Intensive Care
6 Transport, are within the scope of the neonatal critical care Clinical Reference
7 Group.

8 This guideline is for:

- 9 • healthcare professionals in primary, secondary and tertiary care
- 10 • parents and carers of babies born preterm who need respiratory support
- 11 • commissioners and providers of specialist neonatal care services.

12 **Groups that are covered**

- 13 • Babies born preterm who need respiratory support (for example, oxygen
14 supplementation or assisted ventilation) in hospital, beginning in the neonatal
15 period.

16 **Groups that are not covered**

- 17 • Babies born at term.
- 18 • Babies who need respiratory support because of congenital disorders, for
19 example, congenital diaphragmatic hernia.

20 **Finding more information and resources**

21 To find out what NICE has said on topics related to this guideline, see our web page
22 on [postnatal care](#).