Inadvertent overdosing of neonates as a result of the dead space of the syringe hub and needle

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Transcutaneous temperature taken at a thermally insulated site (in this instance the baby’s back, insulated by a non-conducting mattress) represents core body temperature once the insulated skin warms to body temperature.1 Our observations suggested that this equilibration took around seven minutes (fig 1). We took the temperature at seven minutes as representative of the baseline and attributed all changes in temperature that occurred after seven minutes to the postnatal thermal care. Fetal temperature is higher than maternal temperature.2 We defined the upper limit of normal temperature at birth as 37.5°C. Hypothermia was defined as <36°C.

Twenty seven infants (14 boys and 13 girls) were studied. Mean (range) birth weight and gestation were 916 g (490–1470) and 26+4 weeks (24–28+5). Mean (range) temperature at 15 minutes was 37.3°C (36.3–38.1). No infant became hypothermic. Sixteen infants had temperatures that never exceeded 37.5°C. Three infants with normal baseline temperature were warmed to temperatures above 37.5°C during stabilisation. The increases in temperature were 0.2°C, 0.6°C, and 0.6°C with the maximum temperature reached being 38.1°C. Eight (30%) infants had baseline temperatures above 37.5°C of these, five gradually cooled towards 37°C during stabilisation and the remaining three warmed by 0.2°C, 0.3°C, and 0.4°C with a maximum temperature of 38°C. Figure 2 categorises temperature progression of the infants during the 15 minute study period.

These data confirm that early neonatal hypothermia can be eliminated by the use of polythene bags during initial stabilisation of small preterm infants. A significant proportion (30%) of infants are probably hyperthermic at birth. Iatrogenic temperature increases occurred in a minority of the infants in this study and were small. The significance of transient small temperature increases is uncertain.

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References

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Chappell and Newman1 have asked for urgent initiatives to ensure the manufacture of neonatal target products to reduce the risks associated with intravenous drug administration. We endorse their view and report a little recognised problem with the use of adult formulations in neonatal nurseries.

This investigation was conducted after we noticed symptoms of digoxin overdose (bradyarrhythmia) in a neonate. Retrospective
The nursing drug dose manual *Pediatric drugs and nursing implications* gives the maintenance dose of intravenous digoxin as 2.5 μg/kg/dose. The nursing instruction states that the dose must be diluted, with at least four times the volume, using normal saline or 5% dextrose, and the drug must be given over five minutes. In practice, for a 2 kg neonate who is to receive a 5 μg digoxin intravenous injection (250 μg/ml), the drug volume required is 0.02 ml. This is drawn up in a 1 ml syringe up to the 0.02 ml mark. The nursing practice in our nursery was to draw up normal saline to dilute this four times and give it intravenously, slowly, over five minutes. Cognizance is not taken of the dead space in the needle hub and syringe.

We estimated this dead space by drawing up saline in a 1 ml syringe and flushing out the syringe (Syringe and Precision Glide Needle; 1 ml; 26 G; 0.5 inch; Becton- Dickinson, Singapore). The syringe piston was then withdrawn again, and the volume of saline in the dead space was drawn into the syringe barrel. This volume was noted. The dead space volume is 0.07 ml in this syringe. When the diluent is drawn up, the drug in the dead space is also drawn up resulting in toxicity. In the case of digoxin, the baby received 0.09 ml digoxin instead of 0.02 ml.

Ordinarily if the digoxin is drawn up to the 0.02 ml mark and injected directly, the drug in the dead space is retained in the syringe, and there is no overdose delivered. However, when the diluent is drawn up into the syringe for dilution, the drug in the dead space is also drawn up, and this results in the overdosing.

We looked at the magnitude of error induced by the dead space in some of the drugs routinely used in the nursery. Table 1 shows the standard volume of drug required for 2 kg neonate and the magnitude of error introduced by the dead space of a 1 ml syringe. Its is assumed that these drugs were first drawn in a 1 ml syringe and then diluted in the same syringe. The neonate will get 4.5 times the recommended dose if 250 μg/ml digoxin is used, and 2.4 times the recommended dose if 100 μg/ml digoxin is used.

### Table 1: Inadvertent overdose for different drugs

<table>
<thead>
<tr>
<th>Drug</th>
<th>Drug concentration of drug in vial/ampoule</th>
<th>Amount of drug for a neonate weighing 2 kg</th>
<th>Drug volume taken in a 1 ml syringe</th>
<th>Volume of drug in dead space</th>
<th>Amount of drug in dead space</th>
<th>Total amount of drug given if in normal saline or 5% dextrose</th>
<th>Inadvertent overdose factor</th>
</tr>
</thead>
<tbody>
<tr>
<td>Digoxin</td>
<td>(1 ml = 0.25 mg)</td>
<td>0.025</td>
<td>0.005</td>
<td>0.02</td>
<td>0.07</td>
<td>0.0175</td>
<td>0.005 + 0.0175 = 0.0225</td>
</tr>
<tr>
<td>Digoxin</td>
<td>(1 ml = 0.1 mg)</td>
<td>0.025</td>
<td>0.005</td>
<td>0.05</td>
<td>0.07</td>
<td>0.007</td>
<td>0.005 + 0.007 = 0.012</td>
</tr>
<tr>
<td>Adrenaline</td>
<td>(1 ml = 1 mg; 1:1000)</td>
<td>0.03</td>
<td>0.06</td>
<td>0.06</td>
<td>0.07</td>
<td>0.07</td>
<td>0.06 + 0.07 = 0.13</td>
</tr>
<tr>
<td>Furosemide</td>
<td>(1 ml = 10 mg)</td>
<td>1</td>
<td>2</td>
<td>0.2</td>
<td>0.07</td>
<td>0.7</td>
<td>2 + 0.7 = 2.7</td>
</tr>
<tr>
<td>Dexamethasone</td>
<td>(1 ml = 4 mg)</td>
<td>0.1</td>
<td>0.2</td>
<td>0.05</td>
<td>0.07</td>
<td>0.28</td>
<td>0.2 + 0.28 = 0.48</td>
</tr>
<tr>
<td>Midazolam</td>
<td>(1 ml = 5 mg)</td>
<td>0.1</td>
<td>0.2</td>
<td>0.04</td>
<td>0.07</td>
<td>0.35</td>
<td>0.2 + 0.35 = 0.55</td>
</tr>
</tbody>
</table>

Assuming the drug for a 2 kg neonate is drawn up in a 1 ml syringe and diluted by drawing up the diluent into the same syringe. Doses of drugs taken from Pediatric drugs and nursing implications. * Digoxin is available in two formulations: 250 μg/ml and 100 μg/ml.

### References


**BOOK REVIEWS**

### Managing newborn problems: a guide for doctors, nurses and midwives


It is a sobering thought that world wide there are more than four million neonatal deaths per year, the majority occurring in the first week of life with an equal number of stillbirths. Most of these deaths are preventable and the result of a combination of poor maternal health and nutrition and inadequate perinatal care. This manual is the result of a collaboration between the WHO, UNFPA, UNICEF, and the World Bank and represents a renewed effort to address this problem. It forms one of the WHO IMPAC (Integrated Management of Pregnancy and Childbirth) series. Its remit is to complement the Integrated Management of Childhood Illness (IMCI) guidelines and expand these into a comprehensive evidence based perinatal care manual with the emphasis on illnesses in the first week of life. It is targeted at health professionals (doctors, nurses, and midwives) in under-resourced settings where the vast majority of these deaths occur.

The guide has four distinct sections. The first, “Assessment, findings, and management”, is in the IMCI emergency management style. It guides the recognition and diagnosis of sick neonates and the institution of appropriate management of a range of common clinical scenarios such as fast breathing, maternal fever, poor feeding. In section 2 (“Principles of newborn care”) the basics of safe general care from temperature management to immunisation are reviewed. Section 3 describes frequently used procedures including resuscitation, and section 4, the appendices, deals with medical record keeping. Many management pathways will begin in section 1, the scenario/problem, and will be followed through by cross referencing the later sections. There are also additional training packages to accompany the IMPAC guides.
The manual is both detailed and thoughtful: detailed in its cover of most common neonatal illnesses in a clear, well thought through style, and thoughtful in the emphasis afforded to often neglected areas. Emotional support to the family, note taking, and coping with a death are good examples of the way in which the manual looks beyond the immediate clinical realm. It is also gratifying that in the main) appropriate levels of care are addressed; for example, IPPV is not discussed in the management of respiratory distress but instead suitable, more appropriate, and (for most) equally effective alternatives of delivery such as nasal and ambient oxygen. The book might sometimes miss the mark in trying to be “all things to all men”. There will inevitably be differences in training and experience between the professionals for whom it is intended, and, for some, the amount of detail and need for cross referencing may be “too much information”. This could confuse and risk that less experienced less educated caregivers revert to the “tried and tested” treatment format. Greater use of short, flow chart style algorithms within the guidelines (as opposed to in a separate manual) would aid interpretation. Some medical workers may find a more pictorial format easier. I assume the manual has been translated into a number of languages but could find no details about which (English being far from universal).

One other (small) criticism is the emphasis placed on facilities for investigation, particularly lab service. This is a “luxury” that is placed on facilities for investigation, particularly lab service. This is a “luxury” that is far from universal).

One final point is that, although the manual comprehensively covers care of the neonate, it does not mention either prevention or recurrence. It is debatable as to whose responsibility this is, but ensuring access to good antenatal care including maternal antitetanus vaccination, nutrition, and family planning before the next baby is surely at least partly the remit of the paediatric team.

These points aside, this is a commendable and impressive guide, which comes as close as any I’ve seen to combining the factual information needed in a reference with practical management in the field. One can only hope that it impacts on neonatal morbidity and mortality in the way it deserves to, as these have been static for far too long in the settings for which the book is intended.

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Neonatal respiratory disorders, 2nd edn
Edited by A Greenough, A D Milner. Published by Arnold, 2003, £115.00 (hardback), pp 533. ISBN 0340808136

The title of this second edition should be borne in mind by prospective readers. Those looking for a manual of respiratory care in the wee small hours should look elsewhere. Instead, Greenough and Milner have set themselves the more ambitious task of recording current expert opinion on the causes, management, and outcomes of respiratory disorders affecting the newborn. To achieve this, the editors have drawn on the expertise of those researchers around the world who are recognised leaders in this field and have allowed them a refreshing degree of freedom in expressing their opinions. Herein lies the strength of this text.

The 15 contributors from the first edition have expanded to more than 40. There is far greater international representation and presumably anticipation of an international readership (traditionalists among us will mourn the demise of caesarean and aetiology in favour of cesarean and etiology). Despite the increased number of authors, this second edition is only marginally longer than the first. No space has been allowed for excess verbiage; fact is backed by reference, leaving opinion to stand alone.

Although the chapters dealing with basic science have been retained, change in authorship means that many have been rewritten and all have been updated. New chapters include lung liquid, immunology, and microbiology. The clinical management chapters have been rewritten to reflect advances in the use of high frequency oscillation, ECMO, and nitric oxide and to emphasise the importance of feeding. Emphasis is given to long term outcomes in individual conditions, reflecting the increased data available. Additional chapters include respiratory presentation of cardiac disease, apnoea, and bradycardia of prematurity and respiratory problems of infants with neurological disease. Certainly, those who bought the first edition will find enough here to justify investing in the second edition.

The enthusiasm of contributors for their own areas of research has been allowed to shine through, yet a balance of views has been maintained—Robertson and Johansson’s new chapter on surfactant springs to mind. This leads, inevitably, to differences in style and some idiosyncrasies—how else to explain the fact that CNEP is given almost as many column inches as CPAP? Nonetheless, this book achieves the hallmark of successful medical texts, namely that each chapter can stand alone as a thesis with reasoned argument as to why a particular approach is advocated.

There are rewards too for those who seek answers to clinical questions. Greenough and Davenport’s revised chapter on abnormalities of the diaphragm complements Nicolaides’ chapter on antenatal imaging and therapy and provides a concise and up to date review of the causes, associations, management, and outcomes of congenital diaphragmatic hernia. The answers to almost all questions raised during antenatal consultation for this condition can be found here.

Who should buy it? The first edition had a readership wish list of students, physiotherapists, and obstetricians, as well as doctors and nurses working in neonatal intensive care. This edition is aimed at the neonatal clinician and in my view hits the target, thus earning its place in the unit library. In addition, those wishing to explore the literature underpinning current understanding of newborn respiratory disease would be well rewarded by spending some time with this text.

This then is an eclectic collection of opinions on neonatal respiratory disease built on a sound foundation of basic science. It is a stimulating, rewarding, and enjoyable read—words that don’t often spring to this reader’s mind when contemplating medical texts—and fills the gap between traditional textbook and latest research admirably.

K McCormick