LETTERS TO THE EDITOR

Individualised pulse oximetry limits in neonatal intensive care

Editor,—Gupta et al have successfully demonstrated that they were unable to accurately predict PaO₂ from saturation monitoring even after standardising from a previous measurement. The rest of the conclusions presented in their discussion are however based upon interpretation of other research findings which is not further supported by their own study.

They correctly point out that the poor relation between SpO₂ and PaO₂ is related to differing proportions of fetal haemoglobin, Pco₂, and acid-base balance. This begs a question, which their discussion fails to address, on whether the PaO₂ or the SpO₂ is the most useful index of oxygenation. It is certainly the case that normal in utero PO₂ is within a range which they would describe as “hypoxic”. On the other hand, in the presence of 100% fetal haemoglobin, a saturations monitor should, in these circumstances, correctly indicate adequate saturation.

The authors also remark that transcutaneous oxygen monitoring is “a better way of non-invasively assessing PaO₂”. They provide no evidence for this remark. It is certainly a common experience to find a saturation monitoring alarm high when a transcutaneous monitor is apparently recording a normal or even low PaO₂, because of an undetected poor contact. It is also not the case that transcutaneous oxygen monitoring, particularly on the extremely premature infant, is entirely non-invasive.

In the long run, the purpose of oxygen monitoring is to detect degrees of hypoxia which are likely to cause acidosis or tissue damage and levels of hyperoxia which may risk retinopathy of prematurity. To date there would appear to be no study comparing different measurement methods with respect to these outcomes. However, the authors’ own discussion of the reasons for the poor correlation between SpO₂ and PaO₂ provides an excellent theoretical argument in favour of the former over the latter!

ROLLO CLIFFORD Consultant Paediatrician, Dorset County Hospital, Williams Avenue, Dorchester, Dorset DT1 2UY, UK

Dr Yoxall and Dr Shaw respond: Dr Clifford has raised the important question of whether PaO₂ or SpO₂ is the best index of arterial oxygenation. The answer to this question is, of course, unknown and the aim of our study was not to attempt to provide an answer. Most of the work defining hypoxia and harmful hyperoxia in neonates was performed in the era prior to pulse oximetry and therefore defines these situations in terms of partial pressure rather than oxyhaemoglobin saturation. The guidelines for good practice in the management of neonatal respiratory distress syndrome published by BAPM and RCP state that arterial blood sampling is the “gold standard” for assessing arterial oxygenation. In the absence of evidence to the contrary, we would agree with this.

Pulse oximetry is very widely used during neonatal intensive care. Our study has shown that it is not possible to accurately predict PaO₂ from SpO₂ even after standardising from a previous measurement. The information provided by measurements of PaO₂ and SpO₂ is different and should not be interpreted as interchangeable. As SpO₂ monitoring is non-invasive, non-inconsequential, and has a rapid response time, it useful for monitoring trends and particularly in detecting episodes of sudden deoxygenation. Monitoring of PaO₂ is possible using transcutaneous monitoring and this is a better way of continuously monitoring PaO₂ than trying to extrapolate from the SpO₂ signal. The ease with which pulse oximetry can be applied has led to the virtual abandonment of transcutaneous monitoring by many units. There are technical difficulties with transcutaneous monitoring as pointed out by Dr Clifford, but these can be overcome if the staff caring for the babies are familiar with the technique and use it on a routine basis. The anecdotal experience of those units that continue to use transcutaneous monitoring as part of their clinical routine is that it remains an extremely useful technique.

The purpose of maintaining adequate arterial oxygenation is to provide oxygen to meet the metabolic demands of the baby, prevent pulmonary hypertension, and avoid oxygen toxicity. We do not have a satisfactory method of defining what constitutes adequate oxygenation during intensive care at present and it is necessary to take into account other variables, such as haemoglobin concentration or tissue perfusion, which also determine oxygen delivery when we assess our patients.


Dr Slack and Dr Schapira respond:

We thank Dr Stalker for his comments about our report. He is quite correct that neither we nor any of those authors who have previously described these events have provided any evidence that it is either the pertussis component of the diphtheria/tetanus/whole cell pertussis vaccine or its endotoxin content that is responsible for these reactions. That is why we described it as “the presumed cause”. It is possible that any of the vaccine components or adjuvants are responsible. It may simply be a response to a pain, although this seems unlikely. However, given the evidence of reduced local and systemic reactions with acellular pertussis vaccines in term infants compared with the UK whole cell vaccine, it seems plausible that the pertussis component may be involved.

He is partially correct in stating that the way to answer this would be to conduct a three way randomised trial between placebo, acellular (DTPα) and whole cell pertussis (DTP) vaccines. In fact, it would require multiple randomisations between Hib, DT, DTPα, one, two, three, and five component DTPα vaccines, and placebo. Even if the use of a placebo were ethically acceptable, which we believe would not be the case, the low incidence of these events would make the number of infants required for such a study impossible large.

We are, however, conducting a multicentre study to determine the incidence of apnoeas following immunisation using a three component acellular vaccine. Finally, we do not accept that we have joined the ranks of those who have shaken confidence in the vaccination programme.

The question is not whether preterm infants should be immunised against pertussis but when and with what vaccine.


Randomised controlled trial of cisapride in preterm infants

EDITOR—I read with great interest the study by McClure et al1 reporting delayed gastric emptying and a non-significant increase in whole gut transit time in premature infants treated with cisapride.2 In contrast, placebo controlled studies evaluating cisapride in premature infants have reported improvement in symptoms of gastro-oesophageal reflux disease or improvement in oesophageal pH results and/or manometry.3 As acknowledged by the authors, other studies in preterm infants have reported a reduction in gastric residue and improved feed tolerance. In addition, we have reported a reduction in gastro-oesophageal reflux in premature infants.4

In this latest trial the authors chose hydroxypropylmethyl cellulose as a placebo, given at the “same volume” as cisapride; however, the actual quantity administered is not clear. Cellulose derivatives are commonly employed as laxatives, where it is proposed that they act by absorbing water, so softening the faeces and increasing stool volume. This in turn stimulates faecal propulsion.5 Clearly, depending on the dose of hydroxypropylmethyl cellulose used, the choice of this compound as a placebo is nonsensical as there could be a marked impact on gastrointestinal transit time. From the publication it is also not clear why gastrointestinal motility was measured after three days dosing; it is likely that any cathartic effect from a laxative would be less marked after more prolonged treatment time.

The manufacturer had recommended against the use of cisapride in premature infants because of concern that the metabolic pathway of cisapride may not be fully developed. However, in most European countries and the United States, cisapride is commonly used in premature (gestational age > 34 weeks) infants, and, despite this widespread use, the incidence of clinically important cardiovascular effects is very low. Such events are often associated with high doses (> 0.8 mg/kg/day) or concomitant administration of drugs known to inhibit cytochrome P450 that act on cisapride, are known to prolong the QT interval.

In my opinion, in view of the clinical experience with cisapride in severe gastro-oesophageal reflux disease and feed intolerance in premature infants, and the few clinically relevant cardiovascular events, treatment with cisapride with appropriate monitoring (electrocardiogram before and after two or three days of treatment) should not be withheld from these infants if clinically indicated.

YVAN VANDENPLAS
Academic Children’s Hospital Free University of Brussels Laarbeeklaan 101 1090 Brussels, Belgium


Mechanism of blood pressure increase induced by dopamine in hypotensive preterm neonates

EDITOR—The paper of Zhang et al on the effects of dopamine relies on the assumption that left ventricular output (LVO) is a measure of systemic blood flow (SBF). This is not true in the presence of any shunt through the ductus arteriosus where LVO becomes the sum of SBF and volume of blood shunting back into the lungs through the duct, and so overestimates SBF. While Zhang et al define ducts in their study babies as insignificant, we would question the validity of the criteria used to make this definition. We have never seen shunting only in diastole, chamber enlargement is an inconsistent sign of ductal shunting that develops after day one, and shunt velocity has little relation to shunt size during week one. In fact, the left to right velocity often increases as the duct constricts and the shunt diminishes in size.2 Bidirectional shunting is also usually predominantly left to right. There is much in the data presented by Zhang et al to suggest that these ducts were highly significant; firstly, they selected a population at high risk for a significant duct,


F75
and secondly they measured a mean diameter of 2.9 mm. We showed that this measure is the most accurate predictor of early haemodynamic significance and that preterm ducts over 2 mm in diameter, with very few exceptions, have a haemodynamic impact. In this haemodynamic milieu, LVO is actually a measure of pulmonary blood flow, and right ventricular output becomes the better measure of SBF, but even this can be confounded by the common finding of left to right atrial shunting. That this haemodynamic impact is often present from the very early postnatal period is emphasised in fig 1 which is of a 5 hour old infant of 26 weeks gestation, in whom we measured an LVO of 350 ml/min per kg but a right ventricular output (RVO) of 90 ml/min per kg. This baby had an unrestricted duct (2.6 mm) with a low velocity (0.7 m/s) but high volume left to right shunt. In other words LVO was overestimating SBF by a factor of over 300%. This is not an unusual finding.

This means that the changes in LVO in response to dopamine documented by Zhang et al may have occurred in the systemic circulation but, equally, could have occurred in the pulmonary circulation. The same uncertainty applies to changes in calculated vascular resistance. In other words, LVO may have increased or decreased solely because dopamine has changed the volume of the ductal shunt back into the lungs.

Early preterm ductal shunting should not be assumed to be inconsequential, and interpretation of early postnatal measures of either right or left ventricular output in preterm infants should be approached with caution.

Dr Zhang et al respond:
Dr Evans and his colleagues have raised two main issues about our paper. The first relates to the haemodynamic significance of shunting through the ductus in our subjects. We readily acknowledge that shunting was present and that the measured left ventricular output was therefore an approximation of systemic tissue blood flow. This was unavoidable, as most babies were studied on the first day after birth, when the ductus is proceeding to close. However, on the basis of clinical assessment as well as other data not presented—for example, average left ventricular shortening fraction was 29%—we considered that myocardial dysfunction rather than ductal shunting was the most likely cause of the systemic hypotension shown.

Our study was confined to a select subgroup of babies with hypotension who failed to respond to volume loading and were thus started on inotropes. In contrast, the studies cited by Evans and colleagues were performed in groups of infants who may or may not have received volume loading and/or inotropic treatment and in whom the level of myocardial function ranged from depressed to normal. Furthermore, most of our subjects were studied during the initial 24 hours after birth, whereas the studies cited were performed over the course of the first postnatal week. Given the substantial physiological changes in the cardiovascular system, findings obtained in the middle or end of the initial week may not necessarily be applicable to the first day after birth.

Evans and colleagues state that “left ventricular output may have increased or decreased solely because dopamine changed the volume of the ductal shunt back to the lungs”. The physiological changes we observed suggest that this was unlikely and that dopamine exerted a vasoactive effect within the systemic circulation. Thus the premise that the increase in left ventricular output was related only to greater left to right ductal shunting is not in accord with the concomitant rise in systemic blood pressure induced by dopamine. The contraction that fails in left ventricular output were merely due to a reduction in ductal shunting is inconsistent with the observed fall in superior mesenteric artery velocity and rise in superior mesenteric vascular resistance.

A possibility deserving of consideration is that dopamine changed the degree of ductal shunting and that this comprised one component of the alteration in left ventricular output. We are not aware of any published data in newborn infants that support direct relaxation or constriction of the ductus by dopamine or of a differential effect of dopamine on the systemic and pulmonary vascular beds. Indeed, in studies performed on newborn lambs, we observed that, at infusion rates of up to 15 µg/min per kg, the systemic and pulmonary vascular effects of dopamine were proportionally similar in magnitude (J J Smolich, H Park, and D J Penny, unpublished observations). In this view therefore it was reasonable to assume that, while ductal shunting was present in our babies, the degree was not altered to a substantial extent by dopamine. However, the notion that inotropic treatment could alter ductal shunting in preterm infants has potentially important clinical ramifications and so is worth investigating.

Finally, although we agree that the illustrative example of the degree of dissociation which may occur between the level of left and right ventricular outputs is striking, examination of group data from the same laboratory suggests that, on average, left ventricular output was only 7–15% higher than right ventricular output in ventilated preterm infants. Moreover, while we did not include the data in our paper, the pulmonary trunk velocity-time integral in our infants was significantly different from the ascending aortic velocity-time integral (p < 0.9), suggesting that the marked degree of dissociation between ventricular outputs evident in the illustrative example was not a feature of the babies in our study.

J J SMOLICH
J ZHANG
D J PENNY
V H YU
Centre for Heart and Chest Research
Department of Medicine
Monash University
Department of Paediatrics
Monash University


Figure 1. Doppler measures taken from a 26 week infant at 5 hours of age. (A) High velocity flow (about 1.0 m/s) in the ascending aorta representing a left ventricular output of 350 ml/min per kg. (B) Low velocity flow (about 0.2 m/s) in the pulmonary artery representing a right ventricular output of 90 ml/min per kg. (C) Low velocity (about 0.7 m/s) high volume laminar left to right flow through an unsteconstricted duct.

Letters, Book reviews
Retained umbilical artery catheter presenting as an umbilical abscess

**Editor,—** We report the delayed recognition of an umbilical artery catheter in an 18 month infant who presented with recurrent umbilical discharge.

A male infant was delivered by caesarian section which was complicated by a hypoglycaemic fit. Resuscitation included the attempted intubation of an umbilical venous catheter. The line was found to be arterial and thus removed. A second attempt also entered the umbilical artery, and, as the infant was now stable, no further attempt was made.

At 15 months, the infant presented with an umbilical abscess. This was investigated by drainage to exclude a urachal remnant. An ultrasonic and abdominal radiograph showed a foreign body within the common iliac artery, and, using an infraumbilical approach, part of an umbilical catheter was removed. The remnant was 10 cm in length.

During the neonatal period, the umbilicus is a useful route of vascular access. The umbilical artery can be used for blood pressure and gas tension measurement but all catheters should be removed early to avoid central infection. Other complications such as thrombosis and necrotising enterocolitis have been reported. Migration of umbilical vein catheters into the left atrium and pulmonary veins may occur. Retrieval of fractured umbilical catheters during the neonatal period has been reported.

In this case, the retained catheter was not recognised until the infant presented late with recurrent umbilical infections and illustrates a need addition to our list of differential diagnoses. The history also re-emphasises that it is essential to check central lines for completeness on removal.

R KOTNIS
A SALLOUM
R HITCHCOCK
Department of Paediatric Surgery, John Radcliffe Hospital, Oxford

Table 1 Results of survey of premedication practice for tracheal intubation in term and preterm neonates in Australia and the United Kingdom

<table>
<thead>
<tr>
<th></th>
<th>United Kingdom</th>
<th>Australia</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Term</td>
<td>Preterm</td>
</tr>
<tr>
<td>Routine premedication (%)</td>
<td>22 (42)</td>
<td>18 (34)</td>
</tr>
<tr>
<td>Opiate</td>
<td>BDZ</td>
<td>BDZ + Opiate</td>
</tr>
<tr>
<td>Opiate + BDZ</td>
<td>1</td>
<td>0</td>
</tr>
<tr>
<td>BDZ + muscle relaxant + atropine</td>
<td>1</td>
<td>0</td>
</tr>
<tr>
<td>BDZ, benzodiapine</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

Mat was a semi-structured telephone interview of the nurse in charge of the shift when the call was made. All interviews were conducted by one of two of the authors (S H and J B) in September 1999. There was a 100% response rate, and the results are given in table 1.

Several different combinations of premedication drugs were in routine use in Australia compared with 14 different combinations in the United Kingdom.

In Australian units, the routine administration of premedication for non-emergency tracheal intubation of term and preterm neonates is common practice, and there is some uniformity in the combinations of drugs used. In contrast, this practice is less common in the United Kingdom and there is more diversity of prescribing. In both countries, premedication was more commonly used for term neonates. This difference in practice may reflect the fact that larger babies are more likely to struggle when intubated, making the procedure more technically demanding.

We agree with Whyte et al that there is a strong evidence-based argument for premedication for tracheal intubation in neonates to be routine. Our work brings added clarity to the existing picture and confirms that there is little consensus as to the best combination of drugs to use. Further work to define best practice is urgently required.

STEPHEN HANCOCK
Department of Anaesthesia
Bradford Royal Infirmary

SIMON NEWELL
St James’s University Hospital
Leeds

JOE BRIERLEY
ANDREW BERRY
New South Wales Newborn and Paediatric Emergency Transport Service
 Wentworthville, NSW, Australia


Neonatal pain relief

**Editor,—** The study of Jain and Rutter indicates that, after an hour of application, topical amethocaine gel exerts a demonstrable anaesthetic effect 54.8% of the time, indicating low potential for practical use. In the authors’ own words “a successful . . . anaesthetic should be quick acting, effective and safe . . .”. It is therefore disappointing to note that in the discussion no mention was made of oral sucrose as an analgesic agent for use in neonates undergoing painful procedures. This is despite a large body of evidence suggesting that it reduces the indicators of pain from various sources. Unfortunately there is a widespread reluctance of clinicians to use oral sucrose before performing painful procedures on neonates. We showed this in a recent study.

Questionnaires were sent to the medical directors of the 18 neonatal units in New Zealand in order to determine the knowledge, attitudes, and practice with regard to commonly performed painful neonatal procedures such as blood taking or line placement. In the 15 replies, there was a high degree of awareness that the procedures caused pain (100%), that the physiological stress of pain was more hazardous than the risks of analgesia and analgesia (67%), and that oral sucrose was a safe and effective analgesic (87%) and yet only five units (33%) ever used any type of analgesia for these procedures and usually for less than 10% of the time. Only one unit (our own) used oral sucrose routinely. We find it to

Physiopaedia, pnuemomediastinum, pneumothorax), air can easily reach the cerebral vasculature, especially when there is persistent pulmonary hypertension and right to left intracardiac shunts.

In the light of our observations, as air microemboli could not be detected by ultrasound, I suspect that the perinatal cortical infarction may have been due to cerebral arterial air embolisation in some patients in the study by Govaert et al.

PETER TEMESVARI
Department of Paediatrics
University Teaching Hospital Koczenyi, Hungary
email: temesvari@hotmail.com


Arch Dis Child Fetal Neonatal Ed: first published as 10.1136/fn.83.1.F74e on 1 July 2000. Downloaded from http://fn.bmj.com/ on April 4, 2022 by guest. Protected by copyright.
be highly effective, simple to use, and rapid in action. It is also exceedingly cheap. We do not know why oral sucrose has failed to find much favour in neonatal units. Clearly neither lack of understanding of neonatal pain perception nor lack of knowledge about the analgesic properties of sucrose are significant factors. Perhaps there is an unrecognised prejudice against anything that is low tech, cheap, and not promoted by pharmaceutical companies.

PAUL HEATON
DAVID HERD
Child and Adolescent Community Centre
Taranaki Health
Private Bag 2016
New Plymouth 4620, New Zealand
email: paed.cacc@thcl.co.nz


BOOK REVIEWS


After fifty years, my only real memory of the paediatrics teaching of Professor A V Neale in Bristol is the benefit to children of sunlight and the aphorism “Preconceptual and Prenatal care is crippled by a large proportion of the disadvantaged population who further disadvantage themselves by their own actions”. Regrettably this text from the USA shows us that little has changed. Over these decades the rate of prematurity, intrauterine growth retardation (IUGR) and perinatal mortality has changed very little in spite of all our collective clinical efforts.

The male partner is hardly mentioned in this book so it seems that, to American eyes, the sole responsibility for producing healthy or unhealthy babies lies with the mother. All of us now know that many varied vaginal infections are associated with premature/prelabour rupture of the membranes and prematurity itself and that antibiotic treatment may have a reducing effect on these problems. Yet with a close reading of the chapter on vaginal infections, which is very well referenced, there appear to be no papers indicating that with all these infections possibly being sexually transmitted, no trials have included investigation and treatment of the male partner. For example, although Bacterial Vaginosis, Candida and Haemolytic Streptococcus are not usually recognised as sexually transmitted diseases (STD), all gynaecologists have witnessed cases where they have been the cause of recurrent vaginal infections.

Many chapters emphasise the adverse effects of alcohol, smoking, recreation drugs and poverty on the unborn fetus. I have always had a healthily sceptical interest in Jongbloet’s theory that alcohol consumption just before and at the time of conception may be responsible for a large number of chromosomal aberrations but this attractive theory is nowhere to be found in the book.

Meanwhile we are beset by new threats. A recent study, not mentioned in this book, shows ecstasy exposure being implicated in some congenital anomalies.

This is a well researched book with excellent references. Throughout, areas of research yet to be carried out are mentioned. For that reason alone it could serve as a good “ideas” source book for junior paediatricians and obstetricians to involve themselves in research projects which may one day help to reduce the continuing high incidence of potentially-damaged babies.

JOHN MCGARRY
Consultant Obstetrician and Gynaecologist in Independent Practice North Devon