determining if one method was biased compared with the other. However, this does not answer the more important question, "Does the TMS-derived gestational age agree sufficiently well with the LMP-derived age for individual babies?" To answer this question it is necessary to consider the distribution of the within-person differences between the ages derived by each method (see, for example, Wu et al.\textsuperscript{3}).

A separate criticism of this paper concerns the division of LBW babies into term and preterm groups. While it may be of interest to see if the results are different for the two groups, the only relevant analysis is that for all LBW babies together. The whole point of the exercise is to see if the TMS method can accurately assess gestational age, and distinguishing between LBW babies of different gestations is an important aspect which Cater’s approach cannot fully consider. Even if the statistical approach were valid, a conclusion that the method works for term babies is of limited value if you do not know exactly which babies these are.

I recently made similar remarks\textsuperscript{3} about a paper by Serfontein and Jaroszewicz\textsuperscript{4} in which two other methods of assessing gestational age (Dubowitz’s and Robinson’s) were compared. Serfontein and Jaroszewicz\textsuperscript{4} had used correlation—a method which Dr Cater rightly rejects, but for the wrong reason. Their reply to my previous letter dismissed my criticisms as minor. They were however, fundamental, as are my criticisms of Dr Cater’s paper. It is not statistical quibbling to raise these points. Each type of analysis tests a different hypothesis; if you choose an incorrect analysis you will test an inappropriate hypothesis.

As in the earlier paper, Dr Cater’s conclusion that two different methods are comparable is totally unjustified. In both cases the authors have collected the correct data to enable them to answer their question, but have used incorrect analyses, and in both the data they present suggest that agreement may not be particularly good.

A minimum requirement for the comparison of methods is a scatter diagram relating the methods and indicating the line of equality, together with an analysis of the within-person differences between methods.

References
\begin{itemize}
  \item Cater J I. Confirmation of gestational age by external physical characteristics (total maturity score). \textit{Arch Dis Child} 1979; 54: 794–5.
\end{itemize}

\textbf{D G Altman}
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Dr Cater comments:

I am grateful to Mr Altman for his interest and comments. The infants studied were a sample from a larger survey of LBW babies and their controls were matched for paternal occupation, ordinal position in the family, maternal height, smoking, and sex of the baby. Those of certain gestation were selected as controls and were therefore not strictly matched.

Mr Altman is right in saying that the paired t test will not prove the equivalent of TMS and LMP methods. It is however a necessary if not sufficient condition if the two methods are equivalent for the means to be the same. It is precisely for this reason that the further analysis of subgroups is relevant since the equivalent of means for these subgroups should also be demonstrated.

I cannot agree that the only relevant analysis is to consider all the LBW babies together. Detection of bias in the TMS for particular subgroups may suggest possible recalibration to improve the scoring system.

As clinical methods for corroborating maternal menstrual data in neonatal work become more widespread, it is important to be aware of both the clinical and the statistical limitations of the available scoring methods. Further study of this is proceeding.

\textbf{John I Cater}

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Angus

\textbf{Exchange transfusion in newborn infants}

\textit{SIR},

We were interested to read the report by Pelet\textsuperscript{1} of the effect of exchange transfusion on opsonisation and granulocyte function. While he clearly shows that exchange transfusion increases the capacity of serum to promote phagocytosis of heat-killed brewers’ yeast, this may not be the same phenomenon as correction of the isolated defect of yeast opsonisation by plasma as described by Sothill and Harvey.\textsuperscript{2} Newborn infants have a number of immunodeficiencies and one in particular, namely deficiency of factor B of the alternative pathway of complement,\textsuperscript{3} is likely to be important in the pathogenesis of defective yeast opsonisation. We have shown that children with fulminant hepatic failure (FHF) not only have defective yeast opsonisation, but also profound defects of functional activity of both C3 and factor B (V F Larcher, R J Wyke, A P Mowat, R Williams, in preparation). The addition of subopsonising amounts of normal sera to FHF sera restores yeast opsonisation, as it does in children with primary yeast opsonisation deficiency, but addition of factor B-depleted sera, even in equal volumes, did not provide more than 50% restoration of opsonising activity. On the other hand, the opsonising capacity of sera from patients with primary isolated yeast opsonisation deficiency can be fully restored by the addition of factor B-depleted sera.
These results suggest that normal yeast opsonisation is also, to some extent, dependent on adequate functional levels of factor B. Therefore, unless activity of other complement components is measured in parallel, care is necessary in the interpretation of the yeast opsonisation test.

Finally, the deficiency of yeast opsonisation found in two of Pelet's donors may not have been due to incorrect storage of blood but to primary yeast opsonisation deficiency, which occurs in 5% of the normal population. The use of plasma from such individuals was thought to have been the explanation for the poor opsonic response to plasma infusion in one of our patients with protracted diarrhoea and defective yeast opsonisation.6

We believe that these observations stress the need to screen potential donors of blood for exchange transfusion for white cell and opsonic function if this procedure is to be used rationally in the treatment of septicaemia. The opsonising capacity of sera from normal healthy individuals does not seem to alter markedly with time (personal observations) so that there is a case for the prospective estimation of this parameter at least in potential donors.

References


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Dr Pelet comments:

I appreciate Drs Larcher and Mowat's comments concerning a possible defect in the 'functional activity' of both C3 and factor B. There is increasing evidence that in vitro factor B and C3 activation is defective in newborn sera.1-4 I found no correlation between the level of factor B or C3 and the grade of opsonisation deficiency as indicated by the opsonisation index. Furthermore, studying synthesis of C3 and analysing the degradation products both of factor B and C3 in normal and neonatal septicaemic newborn sera before, during, and after exchange transfusion, I came to the conclusion that in vivo as well as in vitro, such an activation deficiency should be postulated.5 Larcher and Mowat's observation on FHF seems to be another clinical situation where a similar defect is observed. In cases of hepatic failure there could be an additional and different mechanism, since liver is one of the sites for the synthesis of C3.

I fully agree that potential donors of blood for rational septicaemia treatment should be screened for white cells and sera functions.

References


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A suggested child-health clinic form

Sir,

Professor Illingworth1 invited comment on intervention programmes for children who speak late and have been properly tested for hearing defect, and of the effectiveness of such programmes. Cooper et al.2 set out to find and prove an effective way of helping language development in children with early language handicaps. Their 5-year study3 of 119 children in the age range 2 to 4½ years, together with field trials at clinics, showed that most 'programme' children made accelerated progress in all language-related areas of development and that this improved rate of progress was maintained. Sonksen4 showed that the accelerated progress occurred in children with all degrees of handicap and there was no evidence that it was related to degree, nor was there a relationship to the paediatric categories of 'causal' or 'developmental'. The Wolfson intervention programmes5 introduced last year by our speech therapists at the Newcomen Centre give encouraging results.

Professor Illingworth also questions the need for routine vision tests for children under age 5 years, relying instead on nystagmus, opacity, or persistent squint to reveal treatable visual acuity problems. The Stycar distant vision tests in daily use at the Newcomen Centre give convincing evidence of reliability in picking up visual impairment, and lead to early referral to an ophthalmologist for refraction. Near vision tests are more difficult to interpret. The 6-month-old infant's interest in a 1 mm sweet is taken as an indication of adequate near vision. Ophthalmologists are becoming increasingly concerned by the late discovery of children with squint and amblyopia, which have escaped detection until visual acuity is tested. Ingram,6 in a review of all cases referred to hospital and school eye clinics in his district in one calendar year, found that the majority (69%) of amblyopes presented after 5 years of age. Little more than half the children with esotropia had a cosmetically noticeable squint. He pointed out that no improvement can be expected for either straight-eyed amblyopia, or for the