



Clinical Evaluation of Streptococcus pneumoniae Polymerase Chain Reaction in Children with Suspected Septicemia

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Clinical evaluation of *Streptococcus pneumoniae* PCR in children with suspected septicaemia.

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Pneumococcal PCR in children with suspected sepsis.

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Manuscript

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5 Invasive infection with *Streptococcus pneumoniae* is the second commonest cause of
6 meningitis and septicaemia in childhood, resulting in 10 to 15% of cases in the UK ^{1,2}.
7
8 Isolation of *Streptococcus pneumoniae* from a normally sterile site remains the gold
9 standard for diagnosis. Blood culture sensitivity is low (around 45%) and decreases
10 further when antibiotics have been administered (around 20%)³. Despite its accuracy
11 and potential to detect pneumococcus in culture negative patients, the use of
12 molecular testing for *Streptococcus pneumoniae* is not widespread and not yet
13 recommended as part of routine investigations of suspected septicaemia^{1,2}.
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23 We retrospectively tested 117 blood specimens for *Streptococcus pneumoniae lytA*
24 using previously published PCR assays⁴. These specimens were collected from
25 children with suspected meningitis or septicaemia as part of an ethically approved
26 study on the diagnostic accuracy of *Neisseria meningitidis* Loop Mediated Isothermal
27 Assay⁵.
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34 Four of the 117 children tested had pneumococcal *lytA* detected by PCR. Only two of
35 these had invasive pneumococcal disease confirmed by blood culture. The two
36 children who were culture negative had clinical signs of septicaemia: A 10 month old
37 boy with fever vomiting and listlessness who required fluid resuscitation and
38 admission to intensive care; A two year old girl with fever, cough, poor feeding, poor
39 perfusion and focal chest signs.
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48 Evaluating *lytA* PCR as a confirmatory test for pneumococcus is complicated because
49 it appears to perform better than the gold standard traditional culture. The children
50 included in this study were defined as 'those whom the attending clinician suspected
51 as having meningitis or septicaemia'. Review of the clinical details of the children
52 in this study in combination with a positive *lytA* PCR make it likely that these children
53 had invasive pneumococcal disease which conventional culture methods failed to
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diagnose. Although our study was small and pneumococcal disease was rare these data suggest that addition of *lytA* PCR to the routine investigations of children presenting with signs of septicaemia and meningitis is likely to greatly improve laboratory diagnosis of this serious infection. This is clinically a very desirable outcome as treatment for invasive pneumococcal disease requires a longer course of treatment than meningococcal disease². We believe that reviewed National Institute for Health and Care Excellence and other guidelines should consider recommending this test to the panel of investigations of children with suspected meningitis or septicaemia.

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