Comparison of this simplified fluid volume calculation with the standard recommended method shows the close agreement as regards total volumes administered per 24 hours and also complies with the World Health Organisation policy on oral rehydration.

We do not claim that clinical assessment is not required in dehydrating diarrhoea. We do, however, suggest that a careful assessment of the state of the circulation and of the need for resuscitation are the important issues in diarrhoeal dehydration. Once these have been taken care of, there is no real further urgent need to achieve rehydration within a set time frame, provided the patient receives more fluids than he is actually losing. Oral fluid therapy has been shown to achieve rehydration at least as rapidly as the intravenous route, even though the prescription of how much fluid is to be given or how fast is generally much less specific than in the case of intravenous therapy. Expressing the degree of dehydration in terms of a percentage of body weight for the purpose of calculating fluid volumes has little further management relevance in a Third-World context and can safely be relegated to the files of medical history.

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TB or not TB?

An evaluation of children with an incorrect initial diagnosis of pulmonary tuberculosis

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Object. The aim of the study was to identify diagnoses that are confused with pulmonary tuberculosis in children.

Design. Prospective, investigative clinical study.

Setting. Tertiary care teaching hospital and an urban tuberculosis clinic in an area with a very high incidence of pulmonary tuberculosis (> 800 new cases/100 000/year).

Patients. Children suspected of having tuberculosis, children followed up for pulmonary infiltrates with eosinophilia and children with congenital pulmonary anomalies were investigated.

Intervention(s). None.

Outcome measure. Pulmonary tuberculosis was diagnosed using modified World Health Organisation criteria and the diagnoses of those children not suffering from pulmonary tuberculosis were analysed.

Results. Of the 354 children initially suspected of suffering from tuberculosis 71 (20%) were found to be suffering from other pulmonary disease, viz. pneumonia or bronchopneumonia (29%), bronchopneumonia with wheezing (18%), and asthma with lobar or segmental collapse (12%). Of 14 children suffering from pulmonary infiltrates with peripheral eosinophilia 6 (43%) were initially incorrectly diagnosed and treated for tuberculosis. Of 54 children suffering from pulmonary tuberculosis were analysed.

Conclusions. The criteria for diagnosing tuberculosis in children is complicated in areas with a high incidence of tuberculosis and poor socio-economic circumstances where many children presenting with conditions other than tuberculosis will be in contact with an adult case of pulmonary tuberculosis. The commonest conditions confused with tuberculosis are pneumonia, bronchopneumonia and asthma. Pulmonary infiltrates with
Peripheral eosinophilia and congenital lung abnormalities should be considered especially if the children have an atypical clinical picture or do not respond to tuberculosis treatment.

The diagnosis of pulmonary tuberculosis (PTB) in childhood is fraught with imprecision and is usually made by drawing upon a constellation of physical signs, a history of contact with an adult case of PTB, the result of a Mantoux test and a chest radiograph. The symptoms and signs of PTB in childhood are often nonspecific and radiology is often not diagnostic. Tuberculin skin testing will frequently be negative in children with tuberculosis and the skin test reading is influenced by a variety of factors, including technique, BCG administration after birth, nutritional status, intermittent viral infection and age of the child. Gastric aspiration for culture of Mycobacterium tuberculosis will seldom be undertaken outside of referral hospitals and when done usually yields positive results in less than 30% of cases. In areas with a high tuberculosis incidence children from disadvantaged socio-economic backgrounds are often exposed to a variety of infections, parasitic diseases and allergens which, when causing disease, will often present with the same nonspecific symptoms and signs as tuberculosis. These children will also frequently be in contact with adults with PTB, so that the diagnosis of childhood tuberculosis will frequently be considered and this may lead to inappropriate anti-tuberculosis treatment.

In this report we describe firstly our experience with a group of children initially suspected but later shown not to have tuberculosis, and secondly the diagnostic confusion caused by cases of pulmonary infiltrates with eosinophilia and congenital pulmonary anomalies which were initially diagnosed as PTB and treated as such.

Patients and methods

The study was undertaken at Tygerberg Hospital, a tertiary care hospital situated in an area with a tuberculosis incidence of more than 800 new cases per 100 000 population per year.

During the study period 354 children were initially suspected of having active PTB by the attending physicians on the basis of history, physical examination or chest radiography. The children were drawn from three groups. The first group were part of a prospective study which over a period of 16 months evaluated 340 children in the overnight ward who were thought to have PTB.

The second group of 14 children were admitted to the paediatric pulmonology unit with a diagnosis of pulmonary infiltrates with eosinophilia. This diagnosis was made if the child had an absolute eosinophil count greater than 2.5 x 10^9/L and a chest radiograph showed widespread alveolar opacification. All these children responded to antimicrobial drugs, bronchodilators and steroids. In addition, in the majority of cases the diagnosis was confirmed by bronchoalveolar lavage where more than 10% of the cells recovered were eosinophils. Of the 14 children seen over a 6-month period, 6 (43%) had previously been treated for PTB.

The third group of 54 children were referred for evaluation of congenital pulmonary anomalies. All these children were extensively investigated and the anomalies diagnosed by bronchoscopy, bronchography and computed tomography (CT), and confirmed by excision biopsy. Of the 54 children, 8 (15%) had received treatment for PTB.

All 354 children suspected of having tuberculosis were evaluated for tuberculosis. Evaluation included taking a history of close contact with an adult suffering from active PTB, confirmed weight loss, a Mantoux skin test using 5 IU of PPD and a chest radiograph. Owing to the fact that more than 80% of South African children receive BCG in the neonatal period, a Mantoux test was regarded as significantly positive in the presence of ≥ 15 mm induration. Early morning gastric aspirate specimens from each child were submitted for M. tuberculosis culture using a radiometric assay (Bactec). Other investigations, done as clinically indicated, included a full blood count, C-reactive protein concentrations, blood culture, pleural fluid aspiration for microscopy and culture and serology for detection of causative organisms.

Subsequently, these results, as well as the child's response to therapy, were taken into account and the children were reclassified using the criteria proposed by the World Health Organisation and modified by other workers as having probable tuberculosis, confirmed tuberculosis (if M. tuberculosis was cultured from the gastric aspirate), or as not having tuberculosis.

In an attempt to ensure that the children classified as not having tuberculosis were correctly diagnosed their hospital folders were re-evaluated after 12 - 24 months to assess if the diagnosis was correct and whether the children subsequently developed tuberculosis.

Results

Of the 354 children suspected of having PTB, 86 (24%) (median age 24.5 months) had probable and 191 (54%) (median age 19 months) confirmed tuberculosis, while 71 (22%) (median age 19 months) were subsequently diagnosed as not having tuberculosis. The diagnoses of the children not having tuberculosis are shown in Table I.

Table I. Final diagnosis in 71 children initially suspected but subsequently found not to have tuberculosis

<table>
<thead>
<tr>
<th>Diagnosis</th>
<th>Number</th>
</tr>
</thead>
<tbody>
<tr>
<td>Pneumonia/bronchopneumonia</td>
<td>22</td>
</tr>
<tr>
<td>Bronchopneumonia with bronchospasm</td>
<td>15</td>
</tr>
<tr>
<td>Asthma</td>
<td>9</td>
</tr>
<tr>
<td>Pneumonia with cavity formation</td>
<td>3</td>
</tr>
<tr>
<td>Paraffin inhalation</td>
<td>3</td>
</tr>
<tr>
<td>Persistent collapse of a lobe</td>
<td>2</td>
</tr>
<tr>
<td>Foreign body</td>
<td>1</td>
</tr>
<tr>
<td>Bronchiectasis</td>
<td>1</td>
</tr>
<tr>
<td>Morexella vincenti effusion</td>
<td>1</td>
</tr>
<tr>
<td>AV canal defect with azygos vein</td>
<td>1</td>
</tr>
<tr>
<td>Liver abscess</td>
<td>1</td>
</tr>
<tr>
<td>Pulmonary hypertension due to upper airway obstruction</td>
<td>1</td>
</tr>
<tr>
<td>Aspiration pneumonia due to cleft palate</td>
<td>1</td>
</tr>
<tr>
<td>Retropharyngeal abscess</td>
<td>1</td>
</tr>
<tr>
<td>Chronic cough of unknown origin</td>
<td>1</td>
</tr>
<tr>
<td>Congenital pulmonary anomalies</td>
<td>8</td>
</tr>
<tr>
<td>Pulmonary infiltrates with eosinophilia</td>
<td>6</td>
</tr>
</tbody>
</table>
Not surprisingly, pneumonia/bronchopneumonia (29%), bronchopneumonia with bronchospasm (18%) and asthma (12%), especially if associated with gastro-enteritis and malnutrition, were the commonest cause of confusion. The most common clinical findings of the children suspected of having tuberculosis were wheezing (37%), crepitations (17%) and weight below the third percentile for age (37%). The children had been in contact with an adult with active PTB in 15 (21%) of the 71 cases where the information was available.

Twenty children (26%) with repeated respiratory symptoms and signs were, without further evidence, erroneously treated for tuberculosis. This was particularly common in the children with asthma (4) and pneumonia with eosinophilia (6) and pulmonary congenital anomalies (8). Poorly controlled asthma can lead to mucus plugging of the bronchi, causing segmental or lobar collapse which on chest radiography is confused with PTB (Fig. 1).

Six (43%) of the children with pulmonary infiltrates with eosinophilia (mean age 39 months, range 15 - 94 months) were initially diagnosed as having and treated for tuberculosis. These children had a mean duration of symptoms of 450 days (range 90 - 840 days), only 1 had a significantly positive Mantoux test, none had positive cultures of M. tuberculosis and none responded to anti-tuberculosis therapy. The chest radiographs of these children revealed perihilar opacifications (interpreted as hilar adenopathy) in 4 and lobar or segmental collapse (interpreted as endobronchial tuberculosis) in 2 (Fig. 2). In addition, 1 child also had a pleural effusion. The single most important diagnostic finding in these children was a mean absolute peripheral eosinophilia count of 9.1 x 10^9/l (range 2.6 - 28.9 x 10^9/l). The cause of this severe peripheral eosinophilia was most probably Ascaris lumbricoides infestation as all the children had Ascaris eggs in their stools. All the children responded to anthelmintics, bronchodilators and a short course of steroids. None of the children followed up developed tuberculosis.
Fig. 4. Chest radiograph (above) and CT scan (below) of a boy with unilateral pulmonary hypoplasia (arrow).

In the 8 children (mean age 34 months) with congenital pulmonary anomalies the most significant finding was that they had experienced symptoms for a mean of 481 days and all 8 children had been fully treated (some more than once) for tuberculosis even though none had a positive skin test or gastric aspirate culture for M. tuberculosis. None of the children had responded to anti-tuberculosis therapy. The chest radiographs of these 8 children showed large masses originally considered as hilar or mediastinal glands in 6 children and dense opacifications regarded as endobronchial tuberculosis in 2. The final diagnoses confirmed on surgery and histology were that of bronchogenic cyst in 4 children (Fig. 3), unilateral lung hypoplasia in 2 (Fig. 4) and a tracheal bronchus supplying a portion of the right upper lobe in 2.

Only 1 of the 71 children initially diagnosed as not having tuberculosis subsequently developed tuberculosis. The child who initially presented with vague symptoms and an interstitial pneumonitis on chest radiography and whom we initially thought did not have tuberculosis, developed hilar adenopathy on subsequent radiography and justifiably was started on anti-tuberculosis therapy.

Discussion

The decision to institute anti-tuberculosis therapy in a child is influenced on the one hand by the degree of diagnostic certainty and on the other hand by the possible consequences of failure to start treatment timely. One of the factors which influences mortality and morbidity is the age of the child; in particular, those less than 1 year of age are known to be at greatly increased risk of morbidity and mortality as a result of progressive and disseminated forms of tuberculosis. It is therefore not unexpected that children suspected of having tuberculosis but subsequently thought not to have tuberculosis should be younger (median age 19 months) although not significantly younger than those with probable (median age 24.5 months) or confirmed tuberculosis (median age 19 months).

Wheezeing as a result of airway narrowing caused by tuberculosis adenopathy is known to be a relatively common presenting symptom in young children with tuberculosis and in a series of bacteriologically proven cases of childhood tuberculosis presenting to our hospital wheezeing was the presenting symptom in 17%. When asthma is associated with pulmonary opacification or collapse of a lobe or segment due to mucus plugging, especially if the child is in contact with an adult tuberculosis patient, it is not surprising that some confusion should ensue. During the follow-up of the group of 71 children initially suspected of having tuberculosis, 20 were subsequently erroneously treated for tuberculosis. Of these 20 children, 4 had asthma and were repeatedly admitted, sometimes with mucus plugging and lobar or segmental collapse. The same mechanisms are probably the cause of lobar and segmental collapse in children with pneumonia and eosinophilia.

In developing communities failure to thrive or loss of weight are all too common and a significant proportion of young children in the Western Cape experience a fall in growth velocity during the 2nd year of life. While wheezeing as a result of airway narrowing caused by tuberculosis adenopathy is known to be a relatively common presenting symptom in young children with tuberculosis and in a series of bacteriologically proven cases of childhood tuberculosis presenting to our hospital wheezeing was the presenting symptom in 17%. When asthma is associated with pulmonary opacification or collapse of a lobe or segment due to mucus plugging, especially if the child is in contact with an adult tuberculosis patient, it is not surprising that some confusion should ensue. During the follow-up of the group of 71 children initially suspected of having tuberculosis, 20 were subsequently erroneously treated for tuberculosis. Of these 20 children, 4 had asthma and were repeatedly admitted, sometimes with mucus plugging and lobar or segmental collapse. The same mechanisms are probably the cause of lobar and segmental collapse in children with pneumonia and eosinophilia.

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have a chronic lung problem associated with malnutrition. Careful consideration of the child’s previous history, review of previous chest radiographs, a full blood count to exclude peripheral eosinophilia and a Mantoux skin test will minimise the errors. Critical evaluation of all available chest radiographs for perihilar opacification, segmental or lobar collapse and congenital pulmonary anomalies will further decrease incorrect diagnoses.

In conclusion, our experience draws attention to some of the diagnostic pitfalls in the diagnosis of tuberculosis in childhood, HIV infection and AIDS in children is not yet a serious factor in our community, but seems likely in due course to complicate the already difficult task of diagnosing primary tuberculosis and its complications in childhood even further.

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Subclinical pertussis in incompletely vaccinated and unvaccinated infants

A. Ramkisson, H. M. Coovadia, W. E. K. Loening

Incidental to a phase II study of acellular and whole-cell pertussis vaccines involving 342 infants who were clinically observed from birth until the age of 9 months, subclinical pertussis was retrospectively diagnosed in 10 infants on the basis of serological evidence. IgG and IgA to filamentous haemagglutinin (FHA), pertussis toxin (PT) and agglutinogens 2 and 3 (AGG2,3) were assayed by enzyme-linked immunosorbent assay (ELISA) in serum obtained at birth and at 2, 4, 6 and 9 months of age. All 10 infants had > 4-fold rises in at least two different pertussis IgG antibodies. Nine of the 10 infants had > 4-fold increases in all three IgG antibodies measured. One infant had > 4-fold increases in IgG-FHA and IgG-AGG2,3 but not IgG-PT. Seven infants had raised IgA antibodies to PT and FHA and 4 infants had raised IgA antibodies to AGG2,3. Subclinical infection provoked differing degrees of antibody production in response to multiple antigens.

Subclinical infection was detected in both unvaccinated infants (4) and in infants who had been vaccinated from 2 months of age with either acellular (4) or whole-cell vaccines (2). Subjects were 8 months of age or younger and only 1 had completed primary vaccination. Other infections of infancy were commonly detected; 4 infants had upper respiratory disease about the time of subclinical pertussis. None had a household member with symptomatic pertussis. Likelihood of subclinical infection was related to significantly lower levels of maternally acquired pertussis IgG-AGG2,3 antibodies but not associated with infants’ nutritional status. Subclinical pertussis is described in very young babies at an age when the disease is most severe, and therefore has implications for infant morbidity and mortality; it is also relevant to disease surveillance and vaccine efficacy studies.


Subclinical pertussis is important for a number of reasons: it may cause silent spread of infection in the community and among household contacts; it may be a major cause of unrecognised infant morbidity and mortality; and it distorts surveillance data and influences vaccine efficacy studies (although the latter are conventionally defined in terms of protection against clinical disease). Furthermore, this muted expression of disease consolidates our information on the