Usefulness of lateral radiographs for detecting tuberculous lymphadenopathy in children – confirmation using sagittal CT reconstruction with multiplanar cross-referencing

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Background. Diagnosis of pulmonary tuberculosis (PTB) in children remains difficult. Lateral chest radiographs are frequently used to facilitate diagnosis, but interpretation is variable. In this study, lateral chest radiographs (CXRs) are evaluated against sagittal CT reconstructions for the detection of mediastinal lymphadenopathy.

Aim. To correlate suspected lymphadenopathy on lateral CXR with sagittal CT reconstructions and determine which anatomical group of lymph nodes contributes to each lateral CXR location.

Methods and materials. Thirty TB-positive children’s lateral CXRs were retrospectively reviewed for presence of mediastinal lymphadenopathy in 3 pre-determined locations in relation to the carina: retrocarinal, subcarinal and precarinal. Findings of the CT sagittal reconstructions were then correlated with the CXRs for the presence of lymphadenopathy in the same 3 pre-determined areas across the width of the mediastinum. Axial and coronal CT cross-referencing confirmed the position of the lymphadenopathy.

Results. The most frequent locations for lymphadenopathy were the subcarinal (28) and right hilar (25). Sensitivity and specificity values of the CXRs were moderate, with the precarinal region having the best sensitivity and specificity for presence of lymphadenopathy. Contribution to each zonal group on lateral CXR were from multiple anatomical lymph node sites.

Conclusion. The precarinal zone on CXR had the best specificity and sensitivity, and represented mainly subcarinal and right hilar lymph node groups. Attention should be paid to this area on lateral CXRs for detecting lymphadenopathy in children with suspected PTB.

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Diagnosis of pulmonary tuberculosis (PTB) in children remains a difficult problem,1,2 and radiography is frequently used for elucidation.1-3 Interpretation of chest radiographs (CXRs), however, is variable when measured against computed tomography (CT), which is considered the gold standard.4,5 In the present study, the lateral CXR – a readily available and inexpensive diagnostic tool – is evaluated against sagittal CT reconstructions for the detection of mediastinal and hilar lymphadenopathy. Densities suspected of representing lymphadenopathy on lateral CXR have not been confirmed using CT previously, and the contribution of different lymph node groups to the lateral CXR locations remain unproven. Sagittal CT reconstructions were used for comparison with lateral radiographs for determining the position of suspected TB lymphadenopathy. Multiplanar cross-referencing was used to confirm the presence and location of lymphadenopathy. The sensitivity and specificity of lateral CXR locations for the presence of lymphadenopathy was determined against the CT gold standard.

Aim
We set out to compare detection of tuberculous lymphadenopathy on lateral CXRs with sagittal CT reconstructions and determine which anatomical group of lymph nodes contributes to locations on lateral CXR by using multiplanar cross-referencing.

Methods and materials
A retrospective descriptive study was undertaken. Children <13 years of age were included when a diagnosis of TB was made by positive culture of bronchoscopic washings or gastric aspirates and when CXRs (PA and lateral) as well as a chest CT were available (CT scans were performed for the indication of airway compression, as not all children with TB undergo CT routinely). Thirty children meeting the
criteria were included, working retrospectively and consecutively from the start date. Patients were excluded when there was severe airspace disease (i.e. whiteout) and very poor-quality radiographs. Where a case failed to meet the study criteria, the next sequential case was selected.

The chest CTs were retrieved from our electronic database, and multiplanar reconstructions (MPRs) were performed retrospectively using Voxar 3D 4.2 (Barco, Belgium). Lateral CXRs were read from hard copy. Reading of plain radiographs and CT was performed by a paediatric radiologist (SA) with a special interest in TB. Plain radiographs were read first; CTs were read at a different sitting with the reader blinded to patient details, history and previous radiological findings.

Lateral CXRs were read for presence of lymphadenopathy in 3 predetermined locations as ‘negative’ or ‘positive’. Location positions were determined by extending lines from the anterior and posterior walls of the trachea beyond the carina and drawing another line horizontally (perpendicularly) to these below the level of the right upper lobe bronchus. The horizontal line was drawn at this level to exclude densities representing the major vascular structures around the upper regions of the hila (right and left main pulmonary arteries and aortic arch which are indistinguishable from lymph node soft tissue densities). Three areas were thus viewed in relation to the carina: retrocarinal (posterior to the lines drawn), subcarinal (between the lines drawn) and precarinal (anterior to the lines drawn) to represent actual lymph node locations (Fig. 1). Lymphadenopathy in the 3 zones was reported when conglomerate oval densities were noted. Densities were reported to be vascular if these were linear and diverging from the central point. Airspace disease was reported when confluent densities containing air bronchograms were present.

Sagittal MPRs were then read for the presence of lymphadenopathy. The same 3 lines that were drawn on the lateral CXR were digitally superimposed on the mid- and para-sagittal CT reconstructions. The sagittal MPR scan was then read for the presence of lymphadenopathy in the same 3 predetermined areas as a surrogate for lateral CXR, in the midline and every para-sagittal position across the mediastinum. Position of adenopathy was confirmed on axial and coronal plane CT using the cross-referencing cross-hair (Fig. 2). No size dimension for pathological mediastinal nodes was used and the presence of any lymph nodes was recorded as abnormal.

Zonal findings on lateral CXR and sagittal MPR were then correlated. Sensitivity, specificity, positive predictive value and negative predictive value were determined overall and for each of the predetermined positions using the CT images as the gold standard. Ethical approval for this study was obtained from the local ethics board; the study was conducted in conformance with the guiding principles of the Declaration of Helsinki.

Results

The 30 children in this study (15 male and 15 female) ranged from 0 - 7 years, with a mean age of 2 years. Table 1 shows how lymphadenopathy was distributed as seen in the predetermined zones on lateral CXR and sagittal MPR. Anatomical locations for all lymph nodes seen on (axial and MPR) CT were in descending order of frequency: subcarinal (28), right hilar (25), right para-tracheal and azygo-oesophageal (22), precarinal (21) and left hilar (15).

Mismatches in diagnosis of the presence or absence of lymph nodes are summarised in Table 2. Sensitivity and specificity values of CXR (taking CT as the gold standard) were moderate, with the precardinal region having the best sensitivity and specificity for the presence of lymphadenopathy. Positive predictive values were high at all zones, as was the incidence of lymphadenopathy in the chosen population; Table...
3 provides a summary.

Contributions of the different lymph node groups to each of the zonal groups on the lateral CXR (taking only true positives into account) show a predominant contribution to the retrocarinal zone from azygo-oesophageal nodes, subcarinal zone from subcarinal nodes, and the precarinal zone from right hilar and subcarinal nodes. A more detailed summary of this correlation is in Table 4.

**Discussion**

TB remains a significant cause of mortality and morbidity in South Africa and worldwide. The Western Cape in South Africa had an incidence of 468 cases per 100 000 people in 1998, which is among the highest in the world.6 HIV has further increased the incidence of TB. Diagnosing the disease in children is a challenge, as 56 - 65% of children are asymptomatic at the time of diagnosis. The Mantoux test reaction lags behind radiological findings.1-3,7 Sputum is frequently unavailable, and gastric washings are positive in only 30 - 40% of cases.3 Because of all these factors, *Mycobacterium tuberculosis* is isolated in less than 50% of paediatric cases. The diagnosis of PTB therefore often relies on imaging findings.

Children from birth to 3 years have a high prevalence of lymphadenopathy (92 - 96%) and a lower prevalence of parenchymal abnormalities, and this is the radiological hallmark of primary TB.2,3,6 In almost half of early childhood cases, lymphadenopathy is the sole radiological manifestation.3 Because of these factors, strong reliance is placed on CXR in aiding the diagnosis through detection of mediastinal adenopathy. As the WHO’s proposed criteria for PTB include a suggestive appearance on CXR,8 children diagnosed with PTB through radiography are treated with a full course of anti-tuberculous drugs.9

The lateral CXR is still frequently used in South Africa, as it is thought to assist with the detection of hilar and mediastinal lymph nodes.10 It demonstrates lymphadenopathy as lobulated densities appearing in the lower half of a doughnut configuration around the carina. The upper half of the doughnut is made up of the aortic arch.
and the main pulmonary arteries,\textsuperscript{11, 12} which is the reason why we have defined the region below the right upper lobe bronchus for evaluation in our methodology.

Worldwide use of the lateral CXR is still a contentious issue. Some feel that it adds value to frontal projections, which are frequently confusing owing to overlapping anatomical structures.\textsuperscript{13} In the presence of other pathology (e.g. infiltrates and miliary TB), the lateral projection was found to be more accurate in detecting hilar lymphadenopathy by Smuts et al.\textsuperscript{14} Their study confirmed the value of the lateral CXR in paediatric PTB but found difficulty in differentiating left and right hilar lymphadenopathy on frontal CXR, as the heart obscures the left hilum.\textsuperscript{15}

Swingler et al.\textsuperscript{9} found low accuracy among expert observers in detecting adenopathy on plain film. Their study found that the lateral CXR adds little extra value, but that diagnostic accuracy could be improved by refining radiological criteria for lymphadenopathy.\textsuperscript{4} Such criteria have not yet been determined.

TB lymphadenopathy occurs predominantly in the subcarinal and right hilar areas.\textsuperscript{4, 15} Hila are affected more often than the paratracheal
region since lymph drainage occurs from the hilum to paratracheal nodes.14 Glazer et al. found the largest mediastinal lymph nodes in normal individuals also occur in the subcarinal and right tracheobronchial areas.15 These features are useful when trying to determine diagnostic criteria on radiographs.

CT is considered the gold standard for the detection of lymphadenopathy, while CXR has been shown to be insensitive for lymphadenopathy detection.16 However, in many areas of South Africa and the developing world, CT is not readily available. Cost constraints and the high radiation dose of CT limit routine use of this modality in diagnosing PTB. It is therefore best to use CT to define more sensitive and specific CXR criteria.

The size of pathological lymph nodes in PTB on CT is also not well determined.17 Some studies adopt the notion that any visible mediastinal and hilar lymph nodes on CT are abnormal in children. Newer scanners, however, detect smaller nodes, which creates high sensitivity but lower specificity as some visualised nodes may only reflect reactive changes.6 Diagnosing nodes of TB specifically may be more accurate on CT owing to the typical post-contrast enhancement patterns.18,19 Calcification helps in lymph node detection (Fig. 3) but is present in only 15 - 21% of children with PTB, and not before 6 months of age.6

A previous study done Hammersley et al. showed high correlation between subcarinal density on PA films and subcarinal lymphadenopathy on CT.20 No studies to date correlating findings of lymphadenopathy on lateral CXR with those on CT could be found.

In our study, we chose a group with a (pre-selection bias) high incidence of lymphadenopathy, as the goal was not to demonstrate an incidence of lymphadenopathy but rather to confirm the true site of origin of suspected lymphadenopathy on lateral CXR. Numerous locations of nodes are a common finding in thin TB, and multiple groups contributed to each positive zone on lateral CXR. Predominant groups in certain zones follow the anatomical distribution of nodes. Examples are the axygo-oesophageal nodes in the retrocardiac zone. However logical this may seem, it has not been proven to date for lateral CXR imaging. Even though there was a high incidence of nodes, there was still significant mismatch between lateral CXR and MPR. Sensitivity in all zones was only moderately high, considering the high incidence and multiple sites of lymph nodes involved. Specificity for most zones was poor. Overall, the most frequent sites for lymphadenopathy in this study were subcarinal and right hilar and these two lymph node groups presented predominantly in the subcarinal and precarinal zones of the lateral CXR. The precarinal location on lateral CXR (Fig. 4) was the only group with high sensitivity and specificity for detecting lymph nodes when correlated with MPR.

Limitations

The population chosen had a very high incidence of lymph nodes. No normal control cases were used. Therefore, PPV and NPV might not reflect the situation in a regular population group. However, ethical considerations relating to high radiation exposure of chest CT in children prevent us from generating a control group. Use of CT scans for pathology other than TB as controls is also unsuited in our population, as TB is endemic with the highest per capita incidence in the world.

Conclusion

This study proves only moderate correlation between findings of lymphadenopathy on lateral CXR and CT. The precarinal zone on CXR had the best specificity and sensitivity. In addition, this zone was contributed to by all the lymph node groups, and in particular by the subcarinal and right hilar groups, which are most frequently involved in primary TB. Most attention should be paid to the area anterior to a line extended from the anterior tracheal wall and below the right upper lobe bronchus when assessing the CXR for lymphadenopathy in children with suspected PTB.

The lateral CXR is still used in South Africa and many other parts of the world in assessing lymphadenopathy in PTB in children, and research should aim to assist clinicians to focus on the areas of lateral CXR that have higher yield for the presence of lymphadenopathy.