Advances in Imaging Chest Tuberculosis: Blurring of Differences Between Children and Adults

Savvas Andronikou, MBBCh, FCRad, FRCR, PhD\textsuperscript{a,b,*}, Filip M. Vanhoenacker, MD, PhD\textsuperscript{c,d}, Adelard I. De Backer, MD, PhD\textsuperscript{e}

ROLE OF IMAGING

If there was a reliable, cheap, and fast clinical test to diagnose tuberculosis (TB), then imaging would probably be relegated to looking for complications and providing alternative diagnoses in nonresponders. As things stand however, current clinical signs and tests for diagnosing TB do not do the job well enough, cheaply enough, or quickly enough and imaging continues to play a role in the diagnosis and management of TB. Sputum microscopy (and culture) is specific for diagnosis and may be widely available, however, a large proportion of patients, and children in particular, are found to be smear-negative. Imaging remains useful for diagnosis, detection of complications, monitoring response to therapy, and for evaluating outcome.

Diagnosis using imaging is difficult for several reasons: changing patterns of disease; effects of human immunodeficiency virus (HIV) coinfection and AIDS\textsuperscript{1}; inability to identify drug resistance; nonspecific radiographic signs\textsuperscript{1}; subjective interpretation with inter- and intraobserver variability of readers\textsuperscript{1–3}; possibility of a normal radiograph\textsuperscript{1}; problems distinguishing active from inactive disease and infection from disease; imaging is also expensive and often unavailable; radiography is subject to variable quality in technique.

HAS OUR THINKING CHANGED?

The traditional classification of TB into primary and postprimary (reactivation TB) should be avoided\textsuperscript{4} as the pathologic differences between these and the corresponding classic imaging patterns characterizing disease in adults and children have blurred. The age-related distinction has changed because primary infection can occur at any age (especially in countries with low TB incidence)\textsuperscript{5}; because of exogenous reinfection in endemic areas\textsuperscript{4,8–10}; cavitation occurring within 6 months of initial infection (reducing its status as indicator of reactivation),\textsuperscript{4} and because HIV infection results in atypical patterns of disease. A radiological classification of disease is more appropriate.

\textsuperscript{a} Diagnostic Imaging Working Group, Medecins Sans Frontieres, Plantage middenlaan 14, Amsterdam, The Netherlands
\textsuperscript{b} Department of Radiology, University of Cape Town, Anzio Road, Observatory, Cape Town, South Africa
\textsuperscript{c} Department of Radiology, Sint-Maarten Hospital, Duffel-Mechelen, Belgium
\textsuperscript{d} University Hospital Antwerp, UZA, University of Antwerp, Wilrijkstraat, 10, B-2650, Edegem, Belgium
\textsuperscript{e} Department of Radiology, Sint-Lucas Hospital, Groenebriel, 1, B-9000, Ghent, Belgium
\textsuperscript{*} Corresponding author. Department of Radiology, University of Cape Town, Anzio Road, Observatory, Cape Town, South Africa.

E-mail address: docsav@mweb.co.za (S. Andronikou).

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RADIOLOGICAL CLASSIFICATION
- Lymph node TB (gangliopulmonary TB)
- Air-space parenchymal TB (consolidation)
- Tuberculoma
- Miliary TB
- Cavities
- Pleural TB
- Fibrosis and destruction

ADDITIONAL FACTORS
Various situations require different information from the image reader and in turn require more information to be supplied to the reader for an insightful and meaningful interpretation: drug resistance requires the reader to give information on the presence and location of cavities and progression or stability of radiographic findings; HIV coinfection requires a different level of suspicion and a specific differential diagnosis depending on the combination of radiographic findings and clinical information (e.g., CD4 count); complications of TB should be looked for, depending on the clinical presentation and previous imaging findings; treatment centers require comment on activity and whether findings indicate infection or disease, which affects management.

NATIONAL OR PROGRAM POLICY
The role, influence, and level of imaging depend on various factors within each country, project, and setting, and may even vary according to current universal attitude or personal experience of individuals. These variations are influenced by the incidence of TB in a community or the world at large, geography, socioeconomic factors, the age of patients, HIV coinfection, drug resistance, and philosophy of the program managers. There are currently active programs that use imaging because: it is mandatory to screen the general population; smear microscopy and culture are unavailable; of the predominant number of smear-negative patients suspected of having TB, HIV-infected patients require exclusion of active TB before initiation of highly active antiretroviral therapy (HAART); radiographs are useful to guide a change, continuation, or termination of treatment in patients with drug-resistant TB; only the tuberculin skin test is positive in a patient without symptoms; at the end of treatment it is useful to predict relapse of disease. More advanced programs use: computed tomography (CT) when radiographs are normal or equivocal but there are symptoms of TB; multidetector CT with multiplanar reconstruction in an attempt to replace bronchoscopy in complicated lymphobronchial TB; ultrasound to diagnose TB lymphadenopathy in children; magnetic resonance imaging (MRI) to detect and differentiate TB lymphadenopathy from other causes of mediastinal widening; positron emission tomography (PET) to differentiate solitary nodules of TB (tuberculoma) from other causes such as malignancy. Conversely, there is no or little use of radiography when services are unavailable, expensive, require referral elsewhere, are poorly performed or interpreted, or when other more specific tests are proving successful.

DIFFERENCES IN IMAGING CHILDREN AND ADULTS
Children are different in size, anatomy, and physiology from adults. The thymus for example confounds interpretation of the mediastinal width on radiographs. Children are also imaged with a different technique to adults (anteroposterior (AP) instead of posteroanterior (PA) with different settings), provide opportunities for alternative imaging (imaging the mediastinum using ultrasound), and require significant considerations with regard to radiation dose.

IMAGING FINDINGS
Lymph Node TB (Gangliopulmonary TB)
TB lymph nodes in the mediastinum and hilar regions drain a primary parenchymal focus of Fig. 1. Calcified Ghon (Ranke) complex. Plain radiograph (detailed view of the left hilum and left lower lobe). Note the presence of multiple calcified parenchymal foci in the lingula and calcified lymph nodes at the left hilum (arrowhead) and aortopulmonary window (black arrow).
infection. Together the parenchymal focus and lymphadenopathy are known as the “Ghon complex” (Fig. 1).12

Lymphadenopathy was not a major feature of what was previously termed “postprimary TB” in adults (only 5% of cases),13 but recently, especially with HIV coinfection, this tendency has reversed. Enlarged TB lymph nodes may cause complications involving the airways and other surrounding structures (see later discussion on complications of lymphobronchial TB).

Chest radiographs (CXR)
Parenchymal abnormality may be small, peripheral, and difficult to identify.11 In children and immune-suppressed adults, the focal abnormality may not be contained and may present as an air-space process (see later discussion on air-space disease).14,15 Lymphadenopathy in children is only obvious when it projects beyond the cardiac margins and is less often seen on the left (Fig. 2A). Lateral radiographs are useful for detecting lymphadenopathy posterior and inferior to the bronchus intermedius (Fig. 2B). Calcification of lymphadenopathy (and the pulmonary focus) represents a healed lesion but is rare in childhood (Fig. 3).

CT
Characteristic TB lymphadenopathy shows the “rim sign” on contrast-enhanced studies, with a low density center and an enhancing rim (Fig. 4A, B).16,17 There are other causes for the rim sign including atypical mycobacteria,18 lymphoma, and carcinoma.12 More delicate and bizarre enhancement patterns particularly in matted nodes in children have been described as ghostlike (Fig. 5).14 Calcification is easily detected on CT particularly in adults but is rare in children (Fig. 6).14

MRI
Lymphadenopathy may have a characteristic low signal on T2-weighted short Tau inversion recovery (STIR) imaging, probably related to free radicals, which are paramagnetic and associated with caseous necrosis (Fig. 7A, B). Some TB

![Fig. 2. Hilar lymphadenopathy (gangliopulmonary TB) in a child. (A) On the AP radiograph there is a multilobulated mass of lymph nodes projecting beyond the right cardiac margin (short arrows). There is resultant compression of the bronchus intermedius (long arrow). (B) On the lateral radiograph there is an oval dense mass often referred to as a “doughnut” representing a mass of hilar lymphadenopathy (arrowheads).](image)

![Fig. 3. Calcification on a lateral radiograph in a child. Small calcified foci (arrows) representing calcification within TB lymphadenopathy are an unusual finding in children.](image)
lymphadenopathy shows solid nodular enhancement, whereas necrotic nodes show rim enhancement (Fig. 8A, B).

**Air-space Parenchymal Disease (Consolidation)**

This pattern is seen with primary infection (especially in children) as a complication of bronchial erosion and resultant bronchogenic spread of disease or as a complication of bronchial compression with distal parenchymal disease including volume changes (collapse or hyperexpansion). During primary infection the small peripheral focus forms a granuloma, which limits initial replication and spread. In children predominantly, the infection is not well controlled with increasing numbers of mycobacteria and dissemination via lymphatics and the blood stream. Air-space disease is seen in approximately 25% of children with TB.

**CXR**

Confluent areas of opacity often affecting 1 lobe (Fig. 9A) and showing air bronchograms and a positive “silhouette sign” (obliterating the crisp cardiac, mediastinal, or diaphragmatic margins) (Fig. 9B) are the main features. The edge of an air-space process may be ill defined or may be well defined by a fissure that may bulge when there is volume gain (Fig. 9C). In children these may occur in any part of the lung parenchyma.

**CT**

Lung becomes isodense to muscle and shows air bronchograms. Viable (non-necrotic) lung tissue enhances with contrast and shows enhancing vessels branching within it (Fig. 10), whereas necrosis shows lower density and does not enhance, often losing the vascular detail.

**MRI**

The signal of air-space disease is isointense to muscle on T1-weighted MR images and hyperintense on T2-weighted imaging (see Fig. 7B). The signal enhances with intravenous gadolinium when there is no necrosis (Fig. 11). The type of necrosis that may take place can be distinguished using T2-weighted imaging. Liquefactive necrosis

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**Fig. 4.** Lymph node TB (gangliopulmonary tuberculosis). (A) Contrast-enhanced CT image at the level of the aortic arch in an adult showing huge lymph nodes with a necrotic center and peripheral rim enhancement. Note also some smaller solid enhancing lymph nodes anterior to the carina (Courtesy of Dr. A. Snoeckx, Antwerp, Belgium). (B) CT reveals multiple rim-enhancing TB lymph nodes in the mediastinum of a child, clearly demarcated from each other and from the thymus, which is displaced to the left. Note that the trachea is compressed from either side by lymphadenopathy. (From Andronikou S, Wieselthaler N. Imaging for tuberculosis in children. In: Schaaf HS, Zumla A, editors. Tuberculosis: a comprehensive clinical reference. Philadelphia: Saunders Elsevier; 2009. p. 266; with permission.)

**Fig. 5.** CT in a child with primary infection shows characteristic ghost like enhancement. The lymph nodes have various shapes with a suggested translucency created by faintly enhancing margins and low density necrotic centers. (From Andronikou S, Wieselthaler N. Imaging for tuberculosis in children. In: Schaaf HS, Zumla A, editors. Tuberculosis: a comprehensive clinical reference. Philadelphia: Saunders Elsevier; 2009. p. 266; with permission.)
has a high T2 signal and caseous necrosis a low T2 signal within the lung, which already has a high T2 signal caused by the air-space process.

**Tuberculoma/Parenchymal Nodules**

Round or oval lesions involving the lung parenchyma in TB are usually granulomas. By definition, granulomas greater than 1 cm are termed tuberculomas. Tuberculomas may show central necrosis and cavitate. Only calcified lesions should be considered inactive.

**CXR**

Lesions are round or oval and show smooth, sharply defined margins (Fig. 12A). Size varies from 0.4 to 5 cm and the majority are stable in size over time (Fig. 12B, C). Calcification is present in 20% to 30% of tuberculomas in adults. In 80% of tuberculomas there are characteristic satellite lesions in the immediate vicinity of the main lesion (see Fig. 12B, C).

**CT**

Lesions are more easily identified (Fig. 13) and CT is more sensitive for showing calcification. Some lesions may show central low density in keeping with necrosis.

**MRI**

Tuberculomas are isointense to muscle on T1-weighted images but on T2-weighted images signal intensity varies according to the stage/type of necrosis (liquefactive necrosis being of high signal and caseous necrosis of low signal intensity (Fig. 14).

**Miliary Pattern**

Miliary nodules result from and indicate hematogenous dissemination of TB bacilli into the lungs and other organs, where innumerable...
granulomas develop. This is classically seen in children in endemic areas (up to one-third of cases)\(^\text{14}\) but there is an increasing incidence in adults.

**CXR, CT, MRI**

Innumerable nodules of similar size (1–4 mm) are scattered randomly and diffusely throughout both lungs (Fig. 15A). By definition the nodules, being

![Fig. 8. Axial gadolinium-enhanced MRI reveals nodular (short arrowhead) and rim-enhancing (long arrowhead) lymphadenopathy in (A), and distinctly rim-enhancing lymphadenopathy in (B). Note also the enhancing lung parenchyma where there is air-space disease and an area of signal void where there is an air-filled cavity.](image)

![Fig. 9. Air-space disease. (A) Plain radiograph shows a confluent area of peripheral density in the left upper lobe of an adult with primary infection. (Courtesy of Steve Beningfield, University of Cape Town, South Africa). (B) Typical primary infection in a child showing left upper lobe confluent density of an air-space process. In addition there is right hilar lymphadenopathy causing compression of the bronchus intermedius (arrows). (C) The right-sided air-space process in this child has a bulging inferior margin in keeping with an exudative process. In this case it is caused by compression of the right main bronchus by tuberculous lymphadenopathy (arrow). (From Andronikou S, Wieselthaler N. Imaging for tuberculosis in children. In: Schaaf HS, Zumla A, editors. Tuberculosis: a comprehensive clinical reference. Philadelphia: Saunders Elsevier; 2009. p. 268; with permission.)](image)
interstitial, do not coalesce and remain discreetly marginated (Fig. 15B). Even though the appearance is easily recognized on CXR, high-resolution CT (HRCT) is ideally suited to showing these and often also shows interlobular septal thickening (Fig. 15C).14,19 A tip for distinguishing nodules from normal vessels for inexperienced observers is to look in the costophrenic angles and the peripheral 1 cm of the lungs where few vessels are expected.

Cavities

These are formed by liquefaction of caseous necrosis and subsequent fibrosis with lung destruction. They are reported to occur in 40% to 50% of adults with a new diagnosis of TB. They also occur in children and should not be considered as an indication of reinfection. Risk of relapse after anti-TB treatment is significantly higher in patients with cavities whether these are present early on during treatment or at the end of treatment.9 It is also important to recognize cavities and report their position for surgical management when treatment options become limited. There are 4 mechanisms described for cavity formation:

- primary cavitation as early as 6 months after primary infection (Fig. 16A)
- reactivation of previous hematogenous spread with confluence of nodular lesions
- bronchial cavities caused by bronchiectasis or bronchial perforation and distal parenchymal cavitation
- Exogenous reinfection4,6,7

CXR and CT

Air-filled oval areas within an opacity (Fig. 16B) or a nodule (Fig. 16C) represent cavitation. “True” cavities have thick walls (3 mm) and may be nodular or smooth, whereas bullae or pneumatoceles have thin walls and little surrounding opacity (Fig. 16D). Air-fluid levels occur within 10% of cavities (Fig. 16E) (often caused by super-infection). Bronchiectasis is more difficult to confirm on CXR by identification of “ring shadows” with thick walls and parallel tubular markings (“tram-track sign”). HRCT shows the characteristic “signet-ring sign” of the ectatic thick-walled bronchus (representing the ring portion) adjacent to the smaller blood vessel (representing the “gem” of the ring). Traction bronchiectasis exists within distorted lung parenchyma with elevated fissures and hila and pleural adhesions (Fig. 16F).12

Pleural TB

Previously, exudative pleuritis was seen mainly in older children and adolescents. It is currently also seen in adults (in countries with a low incidence of TB infection) and younger children. It occurs 3 to 6 months after primary infection and is often unilateral (and asymptomatic). Pleural fluid only yields culture of the organism in 20% to 40% of samples in patients with TB.20
CXR

Effusions are usually accompanied by parenchymal and nodal disease. They may, however, be the only radiographic sign in a minority of primary TB infections. Blunting of the costophrenic angle alone in adults is not considered significant for active disease by the Centers for Disease Control and Prevention (CDC) as a CXR screening criterion. Only larger amounts of pleural fluid are taken into consideration. Pleural apical capping is also not considered to be suggestive of active TB disease and may represent fatty proliferation (Fig. 17).

Ultrasound

Ultrasound (US) is a useful and rapid way of detecting pleural fluid and guiding a pleural tap.

CT

Contrast-enhanced scans show thickening of the visceral and parietal pleura ("split-pleura sign").

Fibrosis, Scarring or Destruction

Complete or partial destruction of the lung is not uncommon in the end stage of parenchymal and nodal disease.
airway involvement. Fibrosis may be stable, progress or regress, but once the lung is destroyed, activity is difficult to assess.

CXR/CT
Cicatrization atelectasis is common after cavitary disease and involves atelectasis of the upper lobe, retraction of the hilum, compensatory hyper-inflation of the lower lobe, and mediastinal shift toward the fibrotic lung. Distortion of lung parenchyma with volume loss also results in pleural adhesions (Fig. 18) and formation of traction bronchiectasis. Apical pleural thickening is another association of fibrosis and may be caused by proliferation of extrapleural fat and peripheral atelectasis.12

IMPORTANT ADDITIONAL CONSIDERATIONS

Drug Resistance (Multidrug-Resistant or Extensive Drug-Resistant TB)

Multidrug-resistant (MDR) TB is resistance to isoniazid and rifampicin. Extensive drug-resistant (XDR) TB is MDR plus resistance to the fluoroquinolones and at least 1 of the second-line injectable

Fig. 14. Tuberculoma. MRI (STIR sequence) shows a peripheral tuberculoma in the right lung (arrowhead) with characteristic low signal intensity in keeping with caseous necrosis.

Fig. 15. Miliary tuberculosis. (A, B) Plain chest radiographs showing diffuse 2- to 3-mm widespread nodules throughout the lungs. (From Andronikou S, Wieselthaler N. Modern imaging of tuberculosis in children: thoracic, central nervous system and abdominal tuberculosis. Pediatr Radiol 2004;34(11):85; with kind permission of Springer Science + Business Media.) (C) High-resolution CT showing multiple small nodules in a random distribution. Note subtle subpleural and subfissural nodules.
drugs. Imaging plays a role in identifying lesions contributing to drug resistance such as large cavities, which harbor mycobacteria within an area where there is limited drug penetration.

**CXR and CT**

Two imaging patterns have emerged. Patients with new drug resistance (ie, no previous TB treatment or treatment for ≤1 month) usually show noncavitating disease and pleural effusion. Patients with MDR and a history of previous anti-TB treatment longer than 1 month show cavitating disease. Overall, patients with drug-resistant TB show multiple cavities and bronchiectasis more commonly (Fig. 19A, B). Identification and localization of cavities on CT serves as a road map for planning surgery.17

**AIDS, HIV Coinfection and Immune Reconstitution Inflammatory Syndrome**

HIV infection enhances the susceptibility to TB, hastens its progression, and makes it more likely for a patient to be exposed to a case of TB.15 Patients infected with HIV can have massive hematogenous dissemination after initial infection, resulting in a higher risk for a fulminant course. TB is a major cause of death in patients with HIV. CXR for TB in patients with HIV may be confusing.

**Fig. 16.** Cavities. (A) Coronal reformatted CT in a child with primary infection showing multiple cavities in the right lung and necrotic mediastinal lymphadenopathy. (B) Plain radiograph in an adult showing multiple cavities developing bilaterally within large areas of air-space disease. There are air-fluid levels within some of the cavities (arrowhead). (C) Plain radiograph of a thick-walled cavity that has developed within an apical tuberculoma (arrow). (D) Multiple thin-walled cavities within the right upper zone. (E) CT image showing an air-fluid level (arrowhead) in a cavity within the apex of the right lower lobe. (F) Cavities occurring in association with lung distortion and associated endobronchial TB shown on coronal CT scan. There are multiple thick-walled cavities within the left upper lobe, ill defined apicohilar opacities, extensive distortion of the lung parenchyma and hila, and thickening of the apical pleura. Note also traction bronchiectasis and endobronchial spread (“tree-in-bud” sign).

**Fig. 17.** Pleural calcifications in an elderly patient with a past history of tuberculous pleuritis. Plain radiograph shows extensive calcification of the pleura.
because TB and HIV share some imaging features, TB features may be more severe in patients with HIV, there are multiple possible pathologic conditions that may occur simultaneously (Kaposi sarcoma, lymphocytic interstitial pneumonitis, bacterial pneumonia), and because CXR in patients with TB and HIV coinfection may be normal.15

CXR and CT
Appearances depend on the level of immunosuppression.23

- Early stage (immunocompetent) CD4 200–500/mm³: appearances are those of postprimary disease24
- Late stage (immunosuppression) CD4 <200/mm³: up to 20% of CXR may be normal12,25; findings are usually those of lymph node TB regardless of previous exposure24,26
- Severe and advanced (immunosuppression) CD4 <100/mm³: nonspecific findings with diffuse coarse pattern24
- All stages and degrees of immunosuppression: miliary23,27
- Post-HAART initiation (restored immunity): development of immune reconstitution inflammatory syndrome (IRIS) represents a paradoxic clinical and radiological reaction to antigens from TB infection that provokes an inflammatory response. Worsening or new lymphadenopathy, pulmonary disease, and/or effusion are the major chest features.28

Fig. 18. Fibrosis and scarring. There are long linear densities in the left upper zone with distortion of lung architecture, in addition to multiple cavities. The right and left hila are displaced superiorly by traction from the fibrotic scarring.

Fig. 19. Multidrug-resistant (MDR) TB with (A) initial, and (B) 5-month follow-up radiographs. (A) The initial radiograph shows that in addition to the bilateral upper zone ring shadows representing traction bronchiectasis and thin-walled cavities, there is a confluent process in the right upper zone containing thick-walled cavities suggestive of active disease. The active cavitating disease indicates a higher risk for relapse of disease. (B) The routine follow-up radiograph (for management decisions in MDR TB) at 5 months shows that the right upper zone process has extended rather than resolved, with further cavitation and fluid levels in the right mid-zone. There was no culture conversion of sputum in this patient requiring revision of therapy and further drug susceptibility testing.
**Infection Versus Disease, Active Versus Inactive Disease**

Distinguishing TB infection from disease is important as treatment differs (from 1 drug to 3–4 drugs) and because laboratory tests are often unable to make this distinction. This is difficult in children because of the rapid progression from infection to disease. Parenchymal air-space or miliary patterns, lymphadenopathy, and pleural effusion are considered to represent disease. A single non-cavitating, calcified granuloma (Ghon focus) is considered to represent previous infection.

The distinction between active and inactive disease has numerous implications including therapeutic and social (immigration, employment, and health care benefit). Once the lung is destroyed, activity is difficult to assess with radiology studies because fibrosis is known to regress or remain stable. Air-space disease and cavities, with or without lymphadenopathy, are considered active disease. High-resolution CT shows large nodules, “tree-in-bud” densities, lobular consolidation, or mass lesions, and is considered the most accurate imaging method for determining activity.29 Nodules and scars may contain slowly multiplying TB bacilli and have the potential to progress to active disease.21 Patients infected with HIV are a greater challenge as even a normal CXR may be associated with active disease.21 Labeling CXR features as inactive TB should be done with caution. Pulmonary nodules without fibrotic scars and volume loss are considered to represent old healed TB, but it is better to only consider these findings as inactive when smear or culture is negative and the CXR findings are stable over time.12 Calcified nodular lesions have a low risk of active disease.21

**COMPLICATIONS**

**Progressive Primary TB**

Children with progressive primary disease tend to have extensive cavitations and are very ill. They often require admission and ventilation. There is a high mortality rate.

**CXR and CT**

Multiple cavities are seen throughout both lungs with surrounding air-space disease (Fig. 20).30

**Lymphobronchial TB**

Lymph node TB may be complicated by perforation of adenopathy into a bronchus with obstructive pneumonia/atelectasis (epituberculosis).

**CXR**

The lung parenchyma may show air-space disease (consolidation) or obstructive hyperinflation.19 This usually affects the right main bronchus or bronchus intermedius (Fig. 9).12,31,32

**CT**

A progression from postobstructive air-space disease to bronchial filling by fluid (wet lung), progressive necrosis with loss of vascular markings, lack of lung parenchymal enhancement, bulging fissures, and eventual cavitation is well shown (Fig. 21).30,33

**MRI**

Lung necrosis shows nonenhancement within an area of consolidated enhancing lung parenchyma and a range of signal intensity possibilities on T2-weighted imaging. Caseous necrosis is of low signal intensity on T2-weighted images, whereas liquefactive necrosis, which is a precursor to cavitation, is of high signal intensity, and eventual cavitation with air shows signal void (Fig. 22).

**Bronchogenic Spread and Tracheobronchial TB**

Bronchogenic spread complicates 2% to 4% of pulmonary TB disease.34 This originates when there is perforation of lymphadenopathy into a bronchus.

**CXR**

Usually the radiograph is normal but parenchymal opacities, seen as ill defined micronodules that tend to coalesce in some parts, are noted.
Fig. 21. Lymphobronchial TB. Progressive stages of the effects on the lung shown on contrast-enhanced CT (in different patients) from earliest (A) to latest (E). (A) Lymphadenopathy is seen involving the right hilum (arrowhead) and, unlike on the left, the right bronchus is not visible (it is compressed). (B) Distal to a right-sided compression, there is air-space disease. Posteriorly in the parenchyma there are air bronchograms and visible vessels. Anteriorly (the right middle lobe) the bronchi are filled with fluid (wet lung) (arrowhead). The lung enhances throughout. (C) Distal to the bronchus intermedius compression (by hilar and subcarinal lymphadenopathy) the right middle lobe shows nonenhancement, lack of visible bronchi, and a bulging margin (arrowheads) indicating a necrotic lung lobe that is expansile. (D) The right middle lobe necrosis shows early breakdown with air pockets (arrowhead). In contrast, the left lower lobe shows air-space disease with air bronchograms and enhancing parenchyma (in keeping with viability). (E) Advanced cavitation (air-filled areas) in multiple locations within the right lung, which also shows some areas of necrosis (low density, nonenhancing) and some areas of viable consolidated lung (enhancing with air bronchograms). (Parts (C), (D) and (E) from Andronikou S, Wieselthaler N. Imaging for tuberculosis in children. In: Schaff HS, Zumla A, editors. Tuberculosis: a comprehensive clinical reference. Philadelphia: Saunders Elsevier; 2009. p. 270; with permission.)
Fig. 22. MRI of lung necrosis. (A) Postgadolinium T1-weighted MR image shows right-sided necrosis as geographic areas of nonenhancing low signal (arrowheads) compared with the enhancing “solid” air-space disease containing air bronchograms. (B) STIR/T2 imaging shows that some areas of necrosis have characteristic low signal (caseous necrosis) (short arrowhead), whereas other areas show very high signal (liquefactive necrosis) (long arrowhead) easily distinguished from air-space disease, which is generally of high signal with air bronchograms.

Fig. 23. Bronchogenic spread and tracheobronchial TB. (A) Plain radiograph shows widespread, fluffy nodules of various sizes with ill defined edges tending to confluence into areas of air-space disease (left upper zone) representing bronchogenic spread of TB. (B) Sagittal reformatted CT of the left lung showing multiple stenoses along the course of the left upper bronchi (arrowheads), thickening of the bronchial wall, ill defined opacity within the upper lobe, and linear branching opacities within the lower lobe (“tree-in-bud” pattern). Note associated cavitation in the apicoposterior segment of the left upper lobe. (C) Axial image in the same patient as in (B) showing “tree-in-bud” pattern in the left lower lung.

Fig. 24. Extensive pleural calcifications. (A) Plain radiographs and (B) CT showing extensive pleural calcifications on the left side.
This is distinguished from miliary nodules, which have sharp edges and are discreet (ie, separable from each other).

HRCT
Acute tracheobronchial TB is seen as circumferential bronchial narrowing with mural thickening.\(^{35,36}\)
In the late phase the effects on the bronchus are cicatricial bronchiectasis (Fig. 23B).\(^{36}\) Bronchogenic spread of TB is seen earlier and more often on HRCT than plain radiographs and presents as nodular and sharply branching opacities ("tree-in-bud" sign) (Fig. 23C).\(^{37}\)

Empyema, Bronchopleural Fistula, and Fibrothorax
TB pleurisy may become localized causing an empyema. This can also break through the parietal pleura to form a subcutaneous abscess\(^{38}\) or communicate with the bronchial tree as a bronchopleural fistula.

CXR and CT
Empyema is shown as a focal fluid collection with pleural thickening (with or without calcification) (Fig. 24). Fibrothorax is shown as diffuse pleural thickening without pleural effusion and suggests inactivity.\(^{36}\) Bronchopleural fistula shows air in the pleural space that has a changing air-fluid level on sequential CXR. On CT a direct communication may be seen between the pleural space and bronchial tree or lung parenchyma.\(^{39}\)

Aspergilloma
A residual TB cavity can be colonized by \textit{Aspergillus} and presents as an aspergilloma with...
hemoptysis in 60% of cases (and results in death in 5%).

CXR and CT
Early on there is thickening of the walls of a TB cavity and later a spherical nodule is seen with a crescent area of air separating the nodule from the adjacent cavity wall (Fig. 25).\(^{40}\) The nodule is mobile on supine or prone positioning.

TB Pericarditis and Pericardial Effusion
TB pericarditis complicates 1% of TB cases and especially affects immunosuppressed individuals. In sub-Saharan Africa TB is the most common cause of pericardial effusion in adults\(^ {41}\) and effusion is seen in 14% of all TB cases (as high as 22% in those coinfected with HIV).\(^ {42}\) In children with TB, pericardial effusion may be the result of severe malnutrition.\(^ {41}\) Constrictive pericarditis is a late complication affecting 10% of patients with TB pericarditis.\(^ {36}\)

Ultrasound and CT
Pericardial thickening with or without effusion are evident. There may be associated distension of the inferior vena cava, pleural effusions, and deformation of the interventricular septum.\(^ {43}\) Constrictive pericarditis is shown as thickening of the pericardium greater than 3 mm with or without calcification (Fig. 26).

Myocardial TB
This is a rare complication of TB usually associated with other foci of TB.

Fig. 25. Esophageal TB and trachea-esophageal fistula. (A) Traction diverticulae of the esophagus on a barium study, as a result of longstanding TB. (B) Contrast swallow in a child shows contrast entering the trachea and main bronchi in the absence of aspiration indicating a trachea-esophageal fistula. This was repaired using an esophageal stent.

Fig. 26. Chest wall tuberculosis. Coronal fatsuppressed T1-weighted image after gadolinium contrast showing marked enhancement of a large soft tissue lesion at the right thoracic wall. The areas of central nonenhancement reflect caseation and/or liquefaction necrosis.
Fig. 29. Airway compression. (A) Cropped and contrast manipulated view of the airway in a child showing significant compression of the right main bronchus and bronchus intermedius (arrowheads) and less obvious compression of the left main bronchus by TB lymphadenopathy. (B) Coronal reformat of a contrast-enhanced CT of the chest in a child showing the significant airway compression of the right main bronchus and bronchus intermedius by massive rim-enhancing TB lymphadenopathy. (C) Chest radiograph shows complete obliteration of the right main bronchus and bronchus intermedius with resultant right upper lobe and right middle lobe parenchymal lung disease in addition to partial compression of the left main bronchus (arrowhead). (D) Corresponding CT of the patient in (C) shows the offending lymphadenopathy and the severe compression of the bronchus intermedius (arrowhead). (E) Follow-up radiograph of the patient in (C) after surgical enucleation of the right hilar and subcarinal lymphadenopathy shows reconstitution of the caliber of right- and left-sided airways and resolution of the parenchymal disease. (Parts (A), (C) and (D) from Andronikou S, Wieselthaler N. Imaging for tuberculosis in children. In: Schaaf HS, Zumla A, editors. Tuberculosis: a comprehensive clinical reference. Philadelphia: Saunders Elsevier; 2009. p. 272–3; with permission.)
CT and MRI
Single or multiple cardiac tuberculomas may be shown as sharply demarcated lesions especially involving the right heart chambers. CT shows calcification more successfully with or without thrombus formation, which may extend into the vena cava.

Acute Respiratory Distress Syndrome
Acute respiratory distress syndrome may result as an evolution of miliary TB.

CXR
A “snow-storm” appearance is characteristic. After recovery, multiple cystic lesions (pneumatoceles or bullae) may remain.

Broncholithiasis
Broncholithiasis is an uncommon complication occurring after rupture of a calcified peribronchial lymph node into a bronchus.

CT
CT shows the calcified lymph node associated with bronchial obstruction, atelectasis, obstructive pneumonitis, branching calcification, focal hyperinflation, or bronchiectasis.

Fibrosing Mediastinitis
This uncommon complication arises after coalescence of multiple mediastinal lymph nodes and multiple tuberculomas creating an acute inflammatory reaction and reactive fibrosis.
**CT**
A mediastinal and hilar mass with calcification, tracheobronchial narrowing, and vascular encasement (and in some cases superior vena cava syndrome) are evident.\(^{47}\)

**Pneumothorax**
Pneumothorax may complicate severe cavitary disease and heralds the onset of a bronchopleural fistula.\(^{12}\)

**CXR and CT**
A crescent of air (with no discernable lung markings) is seen in the nondependent portion of the chest (erect, apicolateral; supine, inferiorly and medially). There may be mass effect phenomena on the mediastinum when the pneumothorax is large.

**Esophageal TB and Tracheo-esophageal Fistula**
Esophageal TB is more common in patients infected with HIV.\(^{48}\) It results from ingestion of

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*Fig. 32. Lateral radiographs. (A) A normal lateral radiograph shows no oval densities behind the trachea/bronchus intermedium or below its apparent termination. The density of the main pulmonary arteries and posterior aspect of the aortic arch are visible. (B) Obvious lymphadenopathy on a lateral radiograph is shown as oval densities posterior and inferior to the trachea/bronchus intermedium (arrowheads). Associated with the normal dense structures more superiorly, this fits the description known as the "doughnut sign". There is also lower lobe consolidation projected over the distal visible vertebral bodies. (C) Oval density representing subcarinal lymphadenopathy is noted posterior and inferior to the trachea/bronchus intermedium (arrowheads). (D) Another example where oval density representing subcarinal lymphadenopathy is noted posterior and inferior to the trachea/bronchus intermedium (arrowheads). (E) Multiplanar CT reconstruction with cross-hair cross-referencing used to show the positional projection of subcarinal and hilar lymphadenopathy on a lateral radiograph, by inference from the sagittal reconstruction (top right corner). (F) Sagittal STIR/T2 MRI shows the hilar mass of low signal TB lymph nodes (large arrowhead) which would be shown as a doughnut on a lateral image. On MRI the triad of normal vessels that form the superior aspect of the doughnut show flow void (small arrowheads).*
infected sputum (intrinsic), spread from an adjacent TB lymph node or pulmonary TB (extrinsic) or may be the result of hematogenous or lymphatic spread.49

**Barium study**
Barium studies show narrowing and displacement, often at the subcarinal level caused by proximity of the esophagus to the lymphadenopathy in this region. Advanced disease results in esophageal ulceration, fistula formation, stricture, and traction diverticula (Fig. 27A).48,49 Tracheo-esophageal fistula may develop and is shown with contrast in the airways (in the absence of aspiration) (Fig. 27B).50

**CT**
The extent of disease and the extrinsic causes may be evident.

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**Chylous Effusion**
This is an extremely rare complication described in children caused by infiltration or compression of the thoracic duct by right paratracheal lymphadenopathy. The chylous nature of the effusion is best determined with an aspirate.

**US and CT**
Large effusions are shown either uni- or bilaterally (10–20 HU). The enlarged lymph nodes in the right paratracheal and other locations may be a clue to TB as the cause.51

**Phrenic Nerve Palsy**
This is a rare complication reported in children. Large lymphadenopathy in the AP window causes compression and infiltration of the phrenic nerve causing diaphragmatic paralysis.52
CXR and CT
The elevated hemidiaphragm is always on the left. Usually there is accompanying left-sided consolidation with volume loss. On CT this is shown to be caused by compression of the left main bronchus. This is not only caused by lymphadenopathy but also because the mediastinum is displaced to the right allowing the displaced aorta to contribute to the compression of the bronchus.52

Chest Wall TB
The ribs are a frequent location for involvement of tuberculous osteomyelitis. TB granulomatous inflammation and abscesses of the chest wall are most commonly found at the parasternal region, costovertebral junction, and along the shafts of the ribs. It is thought that lymph node enlargement and subsequent caseation necrosis may burrow through the chest wall.53

CXR and CT
CT is more accurate to show subtle rib involvement than CXR. There is usually associated TB pleuritis and intrathoracic lymph node enlargement in contiguity with these chest wall collections. The internal mammary nodes are most commonly involved.

MRI
MRI is the preferred imaging method to assess isolated pyomyositis of the chest wall. The lesion

Fig. 33. Techniques for showing the airway. (A) High kilovolt radiographs used to better show airway stenosis of the left main bronchus (arrowhead). (From Andronikou S, Wieselthaler N. Imaging for tuberculosis in children. In: Schaaf HS, Zumla A, editors. Tuberculosis: a comprehensive clinical reference. Philadelphia: Saunders Elsevier; 2009. p. 272; with permission.) (B) Thick slab minimum intensity projection CT reconstruction shows a normal airway caliber and outline. (C) Thick slab minimum intensity projection CT reconstruction shows how the technique is used to identify stenoses caused by TB and show their length (an advantage over bronchoscopy when the scope cannot pass a compressed area). (D) Three-dimensional volume rendered CT reconstruction technique with transparent setting. The bronchus intermedius and right lower lobe bronchus narrowing are well shown. (From du Plessis J, Goussard P, Andronikou S, Gie R, George R. Comparing three-dimensional volume-rendered CT images with fibreoptic tracheobronchoscopy in the evaluation of airway compression caused by tuberculous lymphadenopathy in children. Pediatr Radiol 2009;39(7):696; with kind permission of Springer Science + Business Media.)
is of low signal intensity on T1-weighted images and of high signal intensity on T2-weighted images. There is marked contrast enhancement after intravenous administration of gadolinium (Fig. 28). Abscess formation is the rule in all cases of advanced pyomyositis. The peripheral wall of the abscess shows a subtle hyperintensity on T1-weighted images and hypointensity on T2-weighted images. On gadolinium-enhanced images, a peripheral enhancing rim corresponding to the abscess wall is observed.

**Vascular**

Pulmonary arteries and veins in an area of active TB may show vasculitis and thrombosis. Bronchial arteries may be enlarged in TB bronchiectasis or parenchymal TB. Rasmussen aneurysms, which are pseudoaneurysms of the pulmonary arteries occurring adjacent to TB cavities, occur in 5% of patients. Hemothysis is the usual presentation and it ranges from being minimal to massive and life-threatening.

**CXR and CT**

Nodular and tubular structures are seen adjacent to areas of bronchiectasis.

**CT angiography**

CT angiography is useful for confirming that tubular structures on CXR are ectatic vessels.

**Angiographic arterial embolization**

Angiographic arterial embolization achieves control of massive hemothysis while awaiting definitive therapy.

**IMAGING TIPS AND NOVEL IDEAS**

**Tips for Reading Chest Radiographs**

**AP CXR in children**

- The most important feature to detect is airway displacement or compression. This may be an indirect sign of lymphadenopathy but is the most objective radiological sign as the airway is the only discernable structure within the density of the mediastinum (Fig. 29A). Airways in normal children are compressible and the normal trachea should be positioned to the right of the midline. Normal bronchi should taper distally and should also be of similar size at an equal distance from the carina on both sides. Lymphadenopathy causes compression of the major airways and may displace the trachea to the left (Fig. 29B). Offending lymphadenopathy can be enucleated to relieve the obstruction and allow re-expansion of the lung (Fig. 29C–E).

- The lobulated soft tissue masses of lymphadenopathy at the paratracheal region and hilar regions are often hidden behind the thymic shadow or the heart shadow, which is relatively large in children. Noting only the outwardly convex lobulated masses that extend beyond the cardiac margin ensures that only definite lymphadenopathy is recorded (Fig. 30).

**PA CXR in adults**

Lymphadenopathy is represented by a polycyclic, curvilinear or lobulated outline to the mediastinum. Enlarged paratracheal or pretracheal lymph nodes result in widening or obliteration of the right paratracheal stripe within a wide mediastinum. A normal hilar point should show a V on its side, convex outward (Fig 31A). Filling of this point, failure to identify it, or an outwardly convex or lobulated margin may represent a mass, including lymphadenopathy (Fig. 31B).

**Lateral CXR in children**

Subcarinal and retrocarinal lymphadenopathy is represented as a lobulated density inferior and posterior to the bronchus intermedius. Lymphadenopathy in this region completes the lower half of a doughnut-shaped density (the upper half of which is made up of the right and left main pulmonary arteries and aortic arch) (Fig. 32).

**Advanced Airway Assessment (High Kilovolt, Three-dimensional Reconstruction, Virtual Bronchoscopy)**

Use of high kilovolt radiographs using a filter technique is not recommended for routine use but can assist in showing the airway, which if compressed or displaced strongly suggests lymphadenopathy (Fig. 33A). In more advanced departments, reconstruction of multidetector CT (MDCT) is
possible in many ways including minimum intensity projections (Fig. 33B, C), volume rendered three-dimensional (Fig. 33D) and virtual bronchoscopy. The three-dimensional volume rendered technique has been assessed against bronchoscopy for detection and characterization of the airway and the causative extrinsic mass. The CT technique was shown to not only match bronchoscopy but also to assess the degree of compression more accurately, and managed to show more distal obstructions of bronchi where the bronchoscope could not navigate because of proximal obstruction.10 Bronchoscopy can be reserved for when biopsy may be necessary.

Routine Availability of HRCT with MDCT (Combi-scanning)

Current MDCT scanning produces slices thin enough to be useful as high-resolution imaging (after reconstruction with appropriate filters). This means that every time an MDCT is performed with contrast for visualization of mediastinal nodes, effusions, air-space disease, and other complications of TB, the data set is available for reconstruction and review as an HRCT study. This is useful for detecting miliary and other interstitial nodules (and lines), bronchiectasis, tree-in-bud features of bronchogenic spread, and air-filled cysts. This is known to add to the diagnostic confidence in TB assessment and the determination of outcome, and can affect management (eg, detection of cavities and bronchiectasis).57 The converse is also possible; that is, when clinicians request HRCT for TB, the study should be performed as a postcontrast, thin-slice mediastinal study (combi-scan) and the HRCT component can be produced in addition to the mediastinal window with a comment on the presence of lymphadenopathy and other information.

Ultrasound for Diagnosis

Ultrasound has been used successfully to detect mediastinal lymphadenopathy via the suprasternal and parasternal approaches in children.58 Follow-up

Fig. 35. Thymus. (A) Plain radiograph shows how the normal thymus can cause the mediastinum to appear wide and have outwardly convex margins. There is no displacement or compression of mediastinal structures by normal thymic tissue. (B) When there is doubt about a mediastinal width, suggesting lymphadenopathy or a mass, ultrasound is useful to show the homogenous texture of the normal thymus seen here (as opposed to the nodular heterogenous appearance of a mass or lymphadenopathy).

Fig. 36. Cropped plain radiograph of the right chest to show how effusion, termed laminar effusions, in children spares the costophrenic angle and tracks superiorly along the lateral chest wall. A tip for detecting these is to check that the lung reaches the rib margins. In normal children there are no significant companion shadows and when a linear density separates the ribs from the lung an effusion is suspected.
Fig. 37. A tick sheet designed for use in children investigated for TB to record information in a retrievable manner and to ensure that the correct aspects of diagnosis are reviewed by the readers in order of priority. (Courtesy of S. Moyo, M. Hatherill, and the South African Tuberculosis Vaccine Initiative, Cape Town, South Africa; with permission.)
ultrasound using the same method has successfully showed response to therapy with decrease in size or disappearance of lymphadenopathy. Using ultrasound to diagnose pulmonary TB by identifying associated abdominal lymphadenopathy has been suggested but shown to be positive in only 25% of patients. The increasing incidence of extrapulmonary TB may make this more important over time especially when used to additionally detect axillary lymphadenopathy (shown to have a moderate correlation with the presence of intrathoracic lymphadenopathy using CT) and cervical lymphadenopathy. Fine-needle aspiration biopsy can be guided by ultrasound in various locations including the mediastinum.

CONSIDERATIONS IN CHILDREN

Thymus

The mediastinum on CXR in children less than 5 years of age shows a prominent thymus that can be as wide as the thoracic cavity. Even though it may have a variable size, margin, and appearance, the thymus is a soft organ and never displaces or compresses the airways (unless infiltrated and abnormal or replaced by lymphadenopathy) (Fig. 35A). Suprasternal ultrasound, CT or MRI can identify normal thymic tissue (Fig. 35B) and distinguish it from lymphadenopathy.

Heart

The heart occupies a larger proportion of the chest in a child than in an adult. This makes it difficult to evaluate the hilar points for lymphadenopathy as described in adults. Instead of considering this a disadvantage, identification of lobulated shadows projecting beyond the cardiac margins should add confidence to the certainty of the presence of lymphadenopathy.

Trachea and Bronchi

Children have soft and compressible airways, which results in the complication of lymphobronchial TB (with compression and distal parenchymal disease) more often than in adults. However, the compressibility of the airways can be used to advantage to infer the presence of lymphadenopathy when noted on CXR.

Effusions

In children, effusions seen on CXR tend to spare the costophrenic angle and extend cranially along the lateral chest wall as they increase in amount. They are termed lamellar effusions and should be checked by looking for a linear density separating the air-filled edge of lung from the lateral chest wall (Fig. 36). Children lack the soft tissue companion shadows noted in adults that would be difficult to distinguish from this imaging pattern.

Radiation

Children are more vulnerable to radiation from medical investigations because of the immature developing organs, the close proximity of sensitive areas (shorter bodies), and the longer life ahead in which to develop malignancies. The lateral CXR is not only an additional view but it imparts approximately twice the radiation dose of the AP CXR. However, the dose is relatively low compared with traditional CT, which is equivalent to approximately 100 CXR. Modern departments have reduced CT doses dramatically by decreasing the tube current setting.

Lymphadenopathy: What Size is Abnormal?

This is a relevant question that has been raised regarding chest CT imaging for TB in children but has also been addressed with regard to abdominal lymphadenopathy in children. Because normal children have been shown to have lymphadenopathy up to 5 mm, the authors recommend noting only lymphadenopathy greater than 1 cm unless there is clear central necrosis on CT, calcification, or low signal intensity on T2-weighted MRI images.

RECORDING SYSTEMS AND TICK SHEETS

Many recording systems have been devised to assist recording of features of TB (Fig 37), particularly for the purposes of research, screening, and data collection. These systems not only ensure that a systematic approach is followed but also systematic recording of the findings using a common language. This makes reporting applicable for follow-up especially if this is not always by the same physician or radiologist. These systems are useful for training residents and non-radiologists in identifying and recording the pertinent features of TB in an organized fashion. Testing of 1 such tick sheet has suggested that CXR may be more useful as a screening tool for TB than previously recognized.

SUMMARY

This article reviews the ongoing role of imaging in the diagnosis of TB and its complications. The blurring of differences in the presentation patterns must be absorbed into daily practice. Communication between radiologists and clinicians and improved interpretation skills by clinicians is increasingly important in the field for making
management decisions, especially where there are limited resources. Recognition of special imaging, anatomic, and vulnerability differences between children and adults is more important than trying to define patterns of disease exclusive to children. Most importantly, radiologists must continue to push ahead with research into identifying more accurate and cheaper methods of diagnosing TB. Even though some aspects of imaging are being discredited, they persist in practice because laboratory and clinical tests have been disappointing or unaffordable. High-end imaging should be used to confirm the most confident and reliable aspects of diagnosis on cheaper modalities such as plain radiographs. These can then be used with greater confidence in the less fortunate parts of the world that suffer the most from this disease.

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