

Pediatric Abdominal Tuberculosis: A disease with many faces

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ABSTRACT:

Abdominal tuberculosis has been reported in 0.3-4% of all cases of childhood tuberculosis. It has been reported mainly from developing countries and is rarer in children as compared to adults. The commonest age group affected is 9-14 years. Abdominal pain, fever and weight loss are the most frequent symptoms on presentation and diagnosis is often delayed by 4–6 months.

There is a variability in the distribution of the disease within the abdomen reported from different centres and multiple abdominal sites are frequently involved. Overall, the spectrum of disease in children is different from adults, peritoneal and lymph nodal involvement being more common than gastrointestinal disease.

Chest x-ray shows pulmonary involvement in up to 25% even in the absence of symptoms and 1/3rd have history of contact with an infected adult. Abdominal imaging is an important preliminary investigation and helps in guiding further evaluation. It's important to try and establish bacteriological and/or histopathological confirmation by obtaining appropriate samples (ascitic fluid, endoscopic biopsies, imaging-guided aspiration from lymph nodes, omentum etc. depending on involved site), however it's possible in only 23–47% cases and one may often have to resort to a therapeutic trial. Standard anti-tuberculous drugs are generally effective and are given for 9-12 months. A proper follow-up is important and the improvement should be assessed both subjective improvement and by objective parameters (such as endoscopic healing and resolution of imaging features). Surgery is reserved for patients who have developed a perforation or obstruction not responding to medical management.

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KEYWORDS: Pediatric abdominal tuberculosis, Acid fast bacilli

ABBREVIATION LIST:

TB –Tuberculosis
 ATB –Abdominal Tuberculosis
 ITB –Intestinal Tuberculosis
 CD –Crohns disease
 GI –gastro-intestinal
 ATT –anti-tubercular therapy
 CBNAAT –cartridge based nucleic acid amplification test
 FNAC - Fine Needle Aspiration Cytology
 AFB –Acid fast bacilli
 SAAG - serum ascitic albumin gradient
 ADA –Adenosine Deaminase
 FDC - Fixed drug combinations

INTRODUCTION

Children have been neglected in the fight against tuberculosis TB for years. Of the 10 million cases of tuberculosis recorded world-wide in the year 2018, an estimated 1.1 million were children, of which an estimated 205,000 children died.(1)An alarming 96% of these deaths occurred among children who

were untreated which highlights the fact that tuberculosis in children is frequently unrecognized.(2) Apart from this, children also represent a significant but underappreciated proportion of the multi-drug resistant -TB burden with an estimated 30,000 children each year.(3) India has by far the highest burden of tuberculosis in children, which is not surprising considering its large size, demographic composition, and moderate tuberculosis prevalence.(4)

Pulmonary tuberculosis is overall the commonest site of tubercular involvement in children and abdominal tuberculosis has been reported to comprise of 0.3 - 4% of all cases of tuberculosis. (5-7) This is likely an underestimate. An autopsy study of children who died from TB showed that 15.7% children had abdominal involvement. (8) Abdominal tuberculosis is relatively rare in children as compared to adults. It has been reported mainly from developing countries, especially the Indian sub-continent and is rare in developed nations. Delisle et al. in a review spanning 70 years (1946 –2014) found only a total of 45 cases reported in literature from Europe, North

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America, New Zealand and Australia. Even amongst these all patients except one were from high risk populations, including aboriginals, foreign born and those with a history of recent travel to endemic countries. (8)

Abdominal TB occurs in four forms: tuberculosis of the abdominal lymph nodes, peritoneal tuberculosis, gastrointestinal tuberculosis and visceral tuberculosis and may occur as a result of haematogenous spread from a primary complex elsewhere, by the ingestion of the tubercle bacilli or by contiguous extension from other adjacent organs.

The protean clinical manifestations of abdominal tuberculosis make it a challenge for the physicians to establish the diagnosis and and this is even more so in the paediatric age group where a child's inability to define their problems accurately contribute to this challenge. It is important to remember that abdominal TB in a child is not like TB in a miniature adult. There are many important differences in the disease presentation, site and type of the disease, diagnostic evaluation and treatment, which have been highlighted in the next few sections.

Clinical features

Abdominal TB usually presents in childhood in the age –group of 9 –14 years and does not have a predilection to any gender. (7, 9-14) (**Table 1**) It is often initially confused with other conditions, and the diagnosis is usually delayed. A median delay of 4 –6 months in diagnosis has been reported in literature. A testimony of the deceptive nature of the illness is the fact that

this delay is also seen in endemic areas where there is a general awareness of this condition.

The clinical spectrum depends on the site of involvement. It ranges from nonspecific symptoms to those that may mimic Crohn's disease. Overall, abdominal pain (61.5 –81%), fever (31-90%), and weight loss (40 –74%) are the most frequent findings on presentation. (7, 9-14) In patients of the pediatric age group, loss of body weight is defined as >5% weight loss in the preceding 3 months. In a child presenting with these triad of symptoms one should suspect abdominal TB.

Table 1 includes the clinical features noted in the major series of patients with pediatric abdominal tuberculosis.

In 33-37% patients extra –abdominal symptoms including respiratory (cough, breathlessness etc.), neurological (headache, seizures), genitourinary symptoms, cervical/ axillary lymphadenopathy and dermatological manifestations (erythema nodosum) may be an associated presenting symptom with respiratory symptoms being most common amongst these.

Disease Distribution

There is considerable variability in the distribution of the disease within the abdomen reported from different centres. The reason for this variability includes –differences in clinical setting (secondary care center vs tertiary referral center) and the department (pediatrics vs pediatric gastroenterology vs pediatric surgeon) who publish their data. Overall, the spectrum of disease in children is different from adults, in whom

Table 1: Clinical features of children with abdominal tuberculosis

	Turkey (n = 35)	Taiwan (n = 10)	India Lucknow (n = 38)	India Chandigarh (n = 218)	India Ajmer (n = 125)	Tunisia (n = 13)	Europe, North America, NZ & Australia (n = 45)
Mean age	9.7 ± 4.3 years	14.7 years	11 (4–16) years	10 (0.25–12) years	9.5 (2-14) years	9.8 (7–14) years	5-9 years –3 10-13 years - 5 >14 years - 11
Gender (M)	54.3%	40%	45%	54%	40%	23%	64%
Duration of symptoms	109 days (10days–3years)	-	4.5 mths (15 days - 5 years)	4 mths (0.5–36) months	54 day (7 days –9 months)	4.2 mths (4 days –1 year)	-
Fever	31.4%	90%	66%	76%	80%	31%	71%
Weight loss	40%	72%	71%	74%	58%	54%	68%
Anorexia	-	-	76%	66%	54%	54%	27%
Abdominal Pain	62.9%	80%	66%	81%	80%	61.5%	76%
Abdominal Mass	-	-	11%	6%	32%	15.4%	32%
Diarrhea	-	0%	18%	21%	14%	-	29%
Intestinal obstruction	-	0%	21%	16%	36%	7.7%	-
Abdominal distension	45.7	-	63%	43%	38%	61.5%	-
Doughy abdomen	-	-	-	27%	12%	-	-
Hepatomegaly	16.1%	-	42%	32%	6%	7.7%	-
Splenomegaly	16.1%	-	21%	15%	4%	16%	-
Ascites	74.3%	40%	47%	26%	44%	69%	68%
Extra –gastrointestinal symptoms	31.4%	100%	21%	21%	16%	16%	37%
Mortality	2.8%	10%	8%	-	0%	0%	9%

Table 2: Distribution of the disease within the abdominal cavity

	Turkey (n = 35)	India Lucknow (n = 38)	India Chandigarh (n = 218)	India Ajmer (n = 125)	Tunisia (n = 13)	Europe, North America, NZ & Australia (n = 45)
Peritonitis	83%	24% (39.4%)	10.1% (35.3%)*	44%	38.4%	42%
Gastro-intestinal	14.3%	16% (39.4%)	16% (54.5%)	20%	-	49%
Lymph nodal	-	16% (39.4%)	17.9% (68.8%)	36%	7.7%	-
Visceral	-	5%	2% (8.2%)	-	-	-
Multiple sites	-	32%	54%	-	54%	-

(*) when those with involvement in multiple sites are also included

peritoneal and lymph nodal involvement is more common than gastrointestinal disease. (9) In the largest series of cases of pediatric abdominal TB reported from Chandigarh, lymph nodal involvement was overall the commonest (12) (Table 2)

Involvement of multiple abdominal sites is common. In the series from Chandigarh, a substantial number (54%) had involvement of multiple sites with a combination of intestinal and lymph nodal involvement being the commonest. In an autopsy series of 24 children with ATB, it was found that, in 82%(14/17) cases of intestinal TB, the intestinal lesion extended to the peritoneum, while 2/7 children with presumed isolated peritoneal TB had evidence of disease arising from other abdominal sites, reiterating the fact that coexistence of TB at multiple sites of the abdomen is common in children.(8)

Lymph nodal involvement varies from 7.7 –68% of children with abdominal TB. The most commonly involved lymph nodes are the mesenteric nodes, peri pancreatic, retroperitoneal, omental nodes and those along the celiac axis. However, it should be remembered that the presence of enlarged mesenteric lymph nodes alone does not mean that the child has abdominal TB as it is a common, non –specific finding in children. The diagnosis of TB should only be considered in the appropriate clinical context. The sonographic finding of oval and elongated lymph nodes with a short-axis diameter up to 10 mm in children, should be considered a normal finding and should not be misdiagnosed as an early manifestation of tuberculosis.(15)

Among children with Intestinal TB (ITB), ileocolonic involvement is the commonest and is seen in upto 3/4th of all children with ITB. Increased physiological stasis with a maximal period of contact, abundant lymphoid tissue and minimal digestive activity in this region are reasons for this occurrence. The left –sided colon may be involved in ~40%. Clinical symptoms specific to ITB include –abdominal pain (60%), diarrhea (40%), blood in stools (10%) and intestinal obstruction (20%).(16) Growth failure is seen in upto 60%. The differentiation between ITB and Crohns Disease is often a challenge as microbiologic confirmation of ITB is possible only in around 40% cases. In a study by Singh et al. 20 children with ITB were compared with 23 children with CD. Features of subacute intestinal obstruction (20% vs 0%), ascites s (30% vs 0%) and isolated ileocecal involvement (40% vs 8.7%) favored ITB. (16) The presence of blood in stool and left-sided colonic involvement were independent predictors of CD. On colonoscopy the presence of deep, longitudinal ulcers

with involvement of multiple segments is more suggestive of CD than TB. Perianal disease in children is seen exclusively in children with CD. This is contrast to adults, where even though perianal disease is more common in CD it has been reported even in ITB. (16)

At times even after taking all the clinical, endoscopic and biochemical parameters into account it is not possible to conclusively differentiate between the two diseases. A therapeutic trial of anti-tubercular therapy (ATT) is safer than starting steroid in children in whom there is a diagnostic dilemma and upto 1/3rd of children with CD receive ATT before a diagnosis of CD is made.

Overall, peritoneal involvement is reported in 35.3 –83% of children with abdominal tuberculosis. Similar to adults, it includes the wet type (which is characterized by ascites formation), a dry fibrotic type (associated with a thickened peritoneum, adhesions and omental thickening with little or no ascites), a mixed type (which is a combination of both), and abdominal cocoon (characterized by presence of a membranous sac around the intestinal loops). Children with the wet type of peritoneal TB generally present with progressive diffuse or localized abdominal distension and pain in association with constitutional symptoms like fever, anorexia and weight loss. Children with the dry fibrotic type or abdominal cocoon often present with features of abdominal pain and distension, vomiting and constipation suggesting intestinal obstruction. In a series of 110 children with abdominal TB presenting to a pediatric surgical department with features of intestinal obstruction, an abdominal lump or entero-umbilical fistula, 91% (n = 100) were found to have adhesive (dry fibrotic) peritonitis. Out of these 100 patients, 23 also had mesenteric lymph nodal involvement.(17) Abdominal cocoon is also known as subacute encapsulating peritonitis is a known cause of intestinal obstruction although it is rare in children. In a small series of 17 children all presented with features of small bowel obstruction.(18)

Visceral (hepatic, splenic) tuberculosis is usually associated with an active pulmonary disease or miliary tuberculosis and isolated involvement (<2%) is rare. Fever, weight loss, right or left hypochondriac pain and hepatosplenomegaly are the most frequently observed clinical findings. Jaundice is a very rare manifestation of tuberculous liver involvement and may be caused by extra or intrahepatic obstruction.

Tubercular involvement of the stomach, duodenum and esophagus in children is hitherto rare (12)

Congenital TB

A rare form of tuberculosis in children is congenital TB. By 2005 only ~ 400 cases had been reported in literature.(19) TB during pregnancy may lead to infection of the placenta or the genital tract, which may then be transmitted to the fetus either haematogenously from the placenta to the umbilical vein or by aspiration or ingestion of contaminated amniotic fluid. In infants in whom the mode of acquisition is transplacental the primary complex develops in the liver, while in those in whom it occurs by the ingestion of infected material the primary is in the gastro-intestinal (GI) tract. As it primarily involves the liver and GI tract i.e. has abdominal involvement it has been included in this chapter.

The Cantwell criteria is used for the diagnosis and is as follows (20) –

Proven tuberculosis lesions in the infant plus one of the following:

- i. Lesions occurring in the first week of life,
- ii. A primary hepatic complex
- iii. Maternal genital tract or placental tuberculosis,
- iv. Exclusion of postnatal transmission by thorough investigation of contacts

The average age at the onset of congenital tuberculosis is 24 days (range, 1–84 days). The symptoms are often non-specific and include fever, irritability, poor feeding, hepatosplenomegaly and respiratory symptoms. It is particularly hard to diagnose because it is seldom distinguishable from other neonatal and congenital infections seen in this age group.

Upto 60–70% of mothers of patients have no symptoms of TB, which makes this condition even difficult to recognise. Most mothers are diagnosed with tuberculosis only after the child has been diagnosed with tuberculosis. (20).

A high index of suspicion is needed for the diagnosis of congenital TB. Tuberculin testing is generally normal. Abdominal imaging (ultrasonography, CT) is a useful tool in diagnosing lesions in the liver and spleen. The sensitivity of liver biopsy for the diagnosis of congenital tuberculosis is 100% however it is an invasive test. (21) One may first try isolating the bacilli from other samples such as gastric aspirates, ascitic fluid (if present), pleural fluid (if present) and/or cerebrospinal fluid as the disease is often disseminated. A proportion (~ 50%) of patients may have an abnormal chest radiograph which may suggest the diagnosis.

Historically, the prognosis of congenital TB has been poor with up to 50% mortality seen. A delay in diagnosis being the most important reason for it.(22) However, with timely diagnosis and initiation of treatment, outcomes have improved.

Diagnosis

Establishing the diagnosis of abdominal tuberculosis can be challenging in children. The paucibacillary nature of the disease in children, difficulties in obtaining adequate samples for analysis and the technical problems in obtaining proper imaging are some of the things that contribute to this difficulty.

Diagnostic tests can be divided into two categories –

1. Test for definitive diagnosis

a. Demonstration of acid fast bacilli on smear or culture

It is the most definitive method to diagnose tuberculosis. Samples are obtained depending on the site of involvement.

- Ultrasound guided Fine Needle Aspiration Cytology (FNAC) of enlarged lymph nodes, focal lesions in the liver and spleen, omentum and thickened bowel may have a yield of 45 –58% (23, 24)
- Endoscopic biopsies from the lesions has a AFB detection rate of 36.1%. Ileocolonoscopy in younger children can be challenging and can be done only in tertiary centres where the appropriate expertise and equipment (pediatric colonoscopies) are available.
- Demonstration of AFB in Ascitic fluid is seen in only ~ 10% children. (25) Ascites with a low serum ascitic albumin gradient (SAAG) and lymphocytic predominance is characteristic of tuberculosis.

The rates of bacteriological confirmation have remained at ~50 % from 1976 to 2019. It appears that the advances in the diagnostic modalities have not translated into higher microbiological yields. (11)

b. Histopathology showing tubercular granuloma

- Tissue obtained during endoscopy may demonstrate the presence of caseating granulomas on histology in ~ 60% of children with intestinal TB
- Laparoscopy is an invaluable tool when the diagnosis is unclear, with a pick –up rate of 85%. Thickened peritoneum with miliary yellowish white tubercles with or without adhesions may be seen. However, it is invasive and is generally done only when the imaging features raise doubts about the possibility of a malignancy, since a lymphoma in a child can mimic TB in every way or there continue to be progressive symptoms even after initiation of anti–tubercular therapy i.e. failure of a therapeutic trial.
- In children with hepatic involvement, a liver biopsy may help in confirming the diagnosis.

c. Cartridge Based Nucleic Acid Assay (CBNAAT)

In all children with suspected abdominal TB the appropriate specimen from the involved site should be collected and subjected to CBNAAT. It is a rapid and fully automated and is based on polymerase chain reaction that detects deoxyribonucleic acid directly from the clinical specimens. It can also detect rpoB gene mutations that confer rifampicin-resistance. In adults it has been found to be positive in upto 67.9% with intestinal TB. (26) Pediatric data is lacking.

2. Investigations that support the diagnosis

a. Radiology

An abnormal imaging plays a pivotal role in the diagnosis of abdominal TB and has the highest yield. However, caution should be exercised in the interpretation of the radiological findings and a diagnosis of TB should only be considered in the appropriate clinical context. This is especially true in children with isolated mesenteric lymph node enlargement where unwarranted treatment is rife.

Chest x –ray –It is a useful adjunct in establishing the diagnosis of tuberculosis. A third of patients may not have pulmonary symptoms and are detected to have pulmonary involvement incidentally on a chest radiograph. Routine chest X- ray is a part of the evaluation in children with suspected ATB.

Ultrasound abdomen/Computed Tomography/
Magnetic resonance imaging

The most common findings in children with abdominal TB are enlarged intra-abdominal lymph nodes.(27). The upper para-aortic and mesenteric groups are more often involved in TB than in lymphoma. Presence of large and / or conglomerate lymph nodes with central hypodensity is suggestive of TB. However, they may be non-enhancing or show areas of calcification. Calcification does not imply inactivity. Ascites (septate ascites); bowel wall thickening; omental thickening “caking”; clumped bowel loops and solid organ involvement (tubercular abscess, calcifications) are other features seen. CT is superior to US because of the ability to identify all the features in a single study. This is especially so in the detection of caseating lymph nodes, high density ascites and bowel-wall thickening. Bowel thickening is considered less common in children than in adults.

b. Demonstration of AFB from extra –gastrointestinal sites.

In children with concomitant pulmonary involvement, gastric lavage samples have a detection rate of 40-92%.(28) The specimen is collected after 4-6 hours of fasting. Induced sputum by 3% nebulized saline is another alternate. Whatever method one chooses to use, one needs to collect at least two, preferably three, samples.

In children with concomitant peripheral lymphadenopathy, Fine needle aspiration cytology (FNAC) specimen may demonstrate AFB in 20-70% cases. (29)

c. Adenosine deaminase

ADA activity in the peritoneal fluid has been proved to be a simple and reliable method for early diagnosis of tuberculous peritonitis. Sensitivity and specificity levels over 90 percent have been reported. (30, 31) Similar observations about the utility of ADA (cut of 36- 40 IU/L) in the diagnosis of tubercular peritonitis have also been made,

d. Mantoux test

Tuberculin skin tests (Mantoux test) are examined 48–72 h after the intradermal injection of five tuberculin units of apurified protein derivative. Positive results have been seen in 17-90% children with abdominal TB.

e. Contact

A unique aspect about childhood tuberculosis is the hunt for an index case i.e. history of contact with an infected person. It has been reported in upto 21-65% children with abdominal TB. (11, 32) Risk of acquiring TB in the child is directly proportional to the number of bacilli to which he/she is exposed. Contagiousness is generally limited to subjects with pulmonary disease, and is greater among the patients with with positive sputum microscopy test results. Subjects with cavitating TB and cough with expectoration are more bacilliferous and contagious.

f. Response to a therapeutic trial

At times when the diagnosis cannot be established even after exhausting all investigative modalities one may have to give anti –tubercular drugs and assess the response. The follow-up of these patients is of extreme importance and demonstration of an objective response to therapy secures the diagnosis.

In children with a diagnostic confusion between TB and CD and who were initiated on ATT, the demonstration of endoscopic healing of the ulcers at end of ATT helps in confirming the diagnosis. A mere subjective response i.e. the resolution of symptoms is not enough as some patients with CD may also have symptomatic improvement.

In children with peritoneal, visceral or lymph nodal TB a repeat imaging at the end of therapy showing an improvement/resolution of the imaging features is warranted for an objective assessment.

It is also important to follow –up these patients for ~ 2 years after completion of therapy to look for a relapse of symptoms.

Based on these investigations the diagnosis of tuberculosis can be

- **Definitive** [Demonstration of AFB (in a tissue smear, histopathology or culture), positive CBNAAT or caseous granulomas on histopathology and/or unequivocal response to ATT]

- **Probable** [compatible clinical features + radiology features/positive mantoux/history of contact/ raised ADA/ demonstration of AFB from extra –gastrointestinal sites]

Table 3: Yield of various investigations for the diagnosis of pediatric abdominal tuberculosis

	Taiwan	India - Lucknow	India- Chandigarh n = 218				Tunisia	Developed
	n = 10	n = 35	GI	L	P	V	n = 13	n = 45
Bacteriological	40%	47%	36.1%	29.3%	29.8%	44.4%	23%	73%
Histopathology	50%	19% ±	54.7%	45.3%	72.2%	72%	46%	18%
Radiology	100% (60%)§	94.5%	81.5%	89.3%	92.2%	100%	100%	-
ADA	-	82%			53.8%		-	-
Abnormal chest x-ray	90%	16%			25.2%		15.3%	24%
TB contact	60%	21%			32.5%		7%	
Mantoux test	17%	45%	64.7%	68.7%	67.5%	72%	61%	90%

± in these patients histopathology alone helped in the diagnosis

¶ AFB on smear, culture or polymerase chain reaction, CBNAAT

§- Ultrasound only

GI –gastrointestinal, L –lymph node, P –Peritoneal, V –Visceral

Table 4: Dosage for Anti –tubercular drugs in children

	Range (mg/kg/d)	Average (mg/kg/d)	Maximum dose (mg)
Rifampicin	10 –20	15	600 mg
Isoniazid	7–15	10	300 mg
Pyrazinamide	30–40	35	2000 mg
Ethambutol	15–25	20	1500 mg

Treatment

In recent years the pharmacokinetics of all first line TB drugs have been re-visited and there has been an upward revision of the dosages needed for children.(33) This is because the pharmacokinetic data suggest higher dosages for maximising the area under curve above the minimum inhibitory concentration. Also, pediatric patients show a rapid metabolism of isoniazid and require a higher mg/kg body weight dose as compared to adults. The current dosages have been tabulated in Table 4

Fixed drug combinations (FDC) that incorporate multi-drug therapy are preferred due to safe and simplified treatment and to do away with the possibility of missing one or more of the combination drugs. Dispersible tablets are available for children.

For a newly diagnosed child with abdominal TB, Intensive phase consists of 8 weeks of Isoniazid, Rifampicin, Pyrazinamide and Ethambutol. This is followed by 16 –40 weeks of three drugs Isoniazid, Rifampicin and Ethambutol as a continuation phase.

The optimal duration of treatment for pediatric ATB is unclear. A shorter duration of treatment increases compliance and decrease the risk of toxicity of the drugs. However, it may also pose a risk for relapses. A review of three randomised controlled trials comprising 328 adult participants found a 6-month regimen (2-month intensive phase, 4-month continuation phase) to be efficacious, but whether it can be extrapolated to children or not is debatable.(34) Apart from including only adults, two of the three studies included only those who had intestinal tuberculosis, which may not be applicable to children where the majority have involvement of multiple abdominal sites. Moreover, the authors of the systemic review have conceded that the quality of evidence regarding the relapse estimate is very low, which is a cause of concern. Hence more data is required before recommendations regarding the optimal treatment duration can be made. Till then treatment duration should be tailored according to the disease extent, treatment response and treating physician's experience in managing such patients. Most centres treat for a total duration of 9-12 months.

Anti tubercular therapy induced hepatotoxicity occurs less frequently in children than adults, it is by no means uncommon. It contributes to 4-8% and 8.7% paediatric cases of drug induced liver injury in the west and India, respectively. It is important to keep it in the back of one's mind when evaluating a child on follow-up. (35)

Surgery is absolutely indicated when there is intestinal perforation. It constitutes ~ 15 % of all children who present with perforating peritonitis to a tertiary centre. (36) Partial intestinal obstruction, adhesive peritonitis and enterocutaneous fistulas are relative indications. In such children it is

prudent to first give a trial of ATT as a proportion of children may respond to it alone. In children with a tubercular enteric stricture, endoscopic dilatation may be attempted.

There has been a paradigm shift in the management, and the frequency of children who require surgery has gone down from 85 –100% in the 1990s to 4 % now.

Outcome

Most children respond well to therapy. Emergence of drug –resistant abdominal TB has recently been reported and should be considered in children who do not show an optimal response to anti –tubercular therapy. In a study from Mumbai, 12.5% children with Abdominal TB had drug –resistant TB.(37) With a timely diagnosis mortality is rare, however in children in whom the diagnosis is considerably delayed upto 10% mortality has been reported.

An algorithm for approaching a child with suspected abdominal TB has been given in **Fig. 1**

References

1. WHO Global TB Report. 2019 October 17 (2019)
2. Dodd PJ, Yuen CM, Sismanidis C, et al. The global burden of tuberculosis mortality in children: a mathematical modelling study. *Lancet Glob Health*. 2017;5: e898–e906
3. Dodd PJ, Sismanidis C, Seddon JA. Global burden of drug-resistant tuberculosis in children: a mathematical modelling study. *Lancet Infect Dis*. 2016; 16:1193–1201.
4. Dodd PJ, Gardiner E, Coghlan R, et al. Burden of childhood tuberculosis in 22 high-burden countries: a mathematical modelling study. *Lancet Glob Health*. 2014;2:e453–e459.
5. Pekcan S, Tana Aslan A, Kiper N, et al. Multicentric analysis of childhood tuberculosis in Turkey. *Turk J Pediatr* 2013; 55:121-9
6. Somu N, Vijayasekaran D, Ravikumar T, et al. Tuberculous disease in a pediatric referral centre: 16 years' experience. *Indian Pediatr*. 1994; 31:1245–1249.
7. Delisle M, Seguin J, Zeilinski D, et al. Paediatric abdominal tuberculosis in developed countries: case series and literature review [published correction appears in *Arch Dis Child*. 2016 Apr;101(4):411]. *Arch Dis Child*. 2016; 101:253–258
8. Ridaura-Sanz C, López-Corella E, Lopez-Ridaura R. Intestinal/Peritoneal tuberculosis in children: an analysis of autopsy cases. *Tuberc. Res. Treat*. 2012; 2012: 230814.
9. Kılıç Ö, Somer A, Hançerli Törün S, et al. Assessment of 35 children with abdominal tuberculosis. *Turk J Gastroenterol*. 2015; 26:128–132.
10. Lin YS, Huang YC, Lin TY. Abdominal tuberculosis in children: a diagnostic challenge. *J Microbiol Immunol Infect*. 2010; 43:188–193.
11. Malik R, Srivastava A, Yachha SK, et al. Childhood abdominal tuberculosis: Disease patterns, diagnosis, and drug resistance. *Indian J Gastroenterol*. 2015; 34:418–425.
12. Lal SB, Bolia R, Menon JV, et al. Abdominal tuberculosis in children: A real-world experience of 218 cases from an endemic region. *JGH Open*. 2019; 4:215–220.
13. Talwar BS, Talwar R, Chowdhary B, et al. Abdominal tuberculosis in children: an Indian experience. *J Trop Pediatr*. 2000; 46:368–370
14. Tinsa F, Essaddam L, Fitouri Z, et al. Abdominal tuberculosis in children. *J Pediatr Gastroenterol Nutr*. 2010; 50:634–638.
15. Karmazyn B, Werner EA, Rejaie B, et al. Mesenteric lymph nodes in children: what is normal? *Pediatr Radiol*. 2005; 35:774–777.
16. Singh SK, Srivastava A, Kumari N, et al. Differentiation Between Crohn Disease and Intestinal Tuberculosis in Children. *J Pediatr Gastroenterol Nutr*. 2018;66(1): e6–e11.
17. Sharma AK, Agarwal LD, Sharma CS, et al. Abdominal tuberculosis in children: experience over a decade. *Indian Pediatr*. 1993; 30:1149.
18. Singal R, Satyashree B, Mittal A et al. Tubercular abdominal cocoon in children - a single centre study in remote area of northern India. *Clujul Med*. 2017; 90: 179–84.

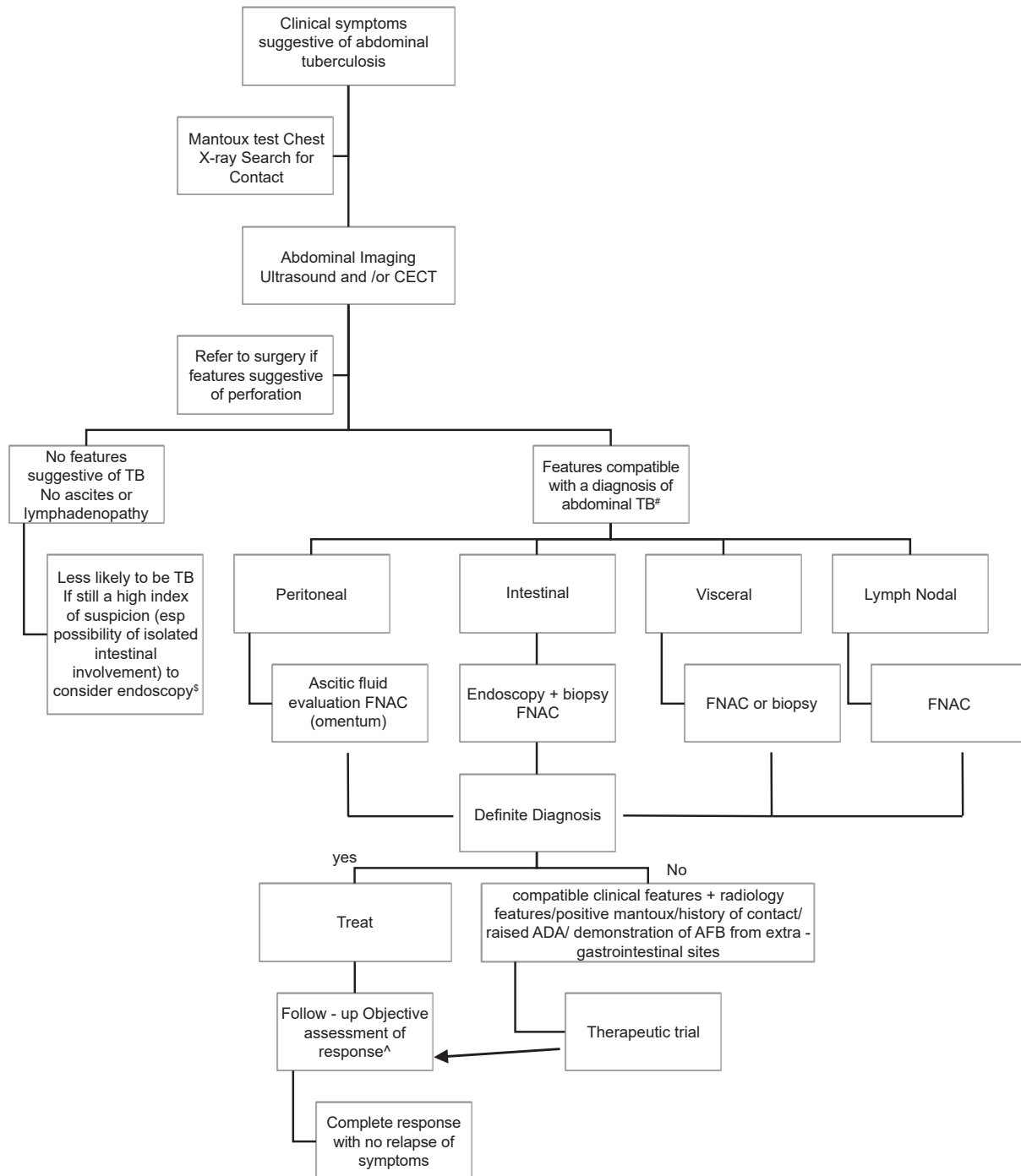


Figure 1: Algorithmic approach to Pediatric Abdominal Tuberculosis

- ascites (free or loculated). high density (on CT) with or without multiple, thin, complete and incomplete septae;

- lymphadenopathy (mesenteric, peri pancreatic, periportal, and para-aortic groups of lymph nodes) seen as conglomerate masses and/or as scattered enlarged nodes with hypochoic or an echoic centres (on USG)/peripheral rim enhancement, non-homogenous enhancement (on CT);
- bowel wall thickening, peritoneal thickening and nodularity, adhesions, mesenteric thickening, and irregular soft tissue densities in the omental area; and
- tiny, low-density foci or multiple low-attenuation, 1-3 cm round lesions scattered in the liver and or spleen.

\$ - Isolated intestinal involvement has the lowest diagnostic yield on imaging

^ - If the child continues to have symptoms then the diagnosis needs to be re-visited. If previous tissue diagnosis has been inconclusive one may consider re-obtaining/repeating it At this point one may consider a laparoscope to obtain better tissue samples for diagnostic evaluation. In children in whom the diagnosis is definite, the possibility of drug-resistance has to be considered. Some children with tubercular intestinal strictures may continue to have pain even after ATT in spite of healing of lesions. This needs to be tackled endoscopically / surgically

CECT –Contrast enhanced computed tomography, FNAC –Fine needle aspiration cytology, ADA –Adenosine deaminase

19. Li C, Liu L, Tao Y. Diagnosis and treatment of congenital tuberculosis: a systematic review of 92 cases. *Orphanet J Rare Dis*. 2019; 14:131. Published 2019 Jun 10.
20. Cantwell MF, Sehab ZM, Costello AM, et al. Congenital tuberculosis. *N Engl J Med*. 1994; 330:1051–4.
21. Dewan P, Gomber S, Das S. Congenital tuberculosis: a rare manifestation of a common disease. *Paediatr Int Child Health*. 2014; 34:60–2.
22. Saramba MI, Zhao D. A Perspective of the Diagnosis and Management of Congenital Tuberculosis. *J Pathog*. 2016; 2016:8623825.
23. Suri R, Gupta S, Gupta SK, Singh K, Suri S. Ultrasound guided fine needle aspiration cytology in abdominal tuberculosis. *Br J Radiol*. 1998; 71:723–727.
24. Radhika S, Rajwanshi A, Kochhar R, et al. Abdominal tuberculosis. Diagnosis by fine needle aspiration cytology. *Acta Cytol*. 1993; 37:673–678.
25. Dinler G, Sensoy G, Helek D, et al. Tuberculous peritonitis in children: report of nine patients and review of the literature. *World J Gastroenterol*. 2008; 14:7235–7239.
26. Makharia GK, Srivastava S, Das P, et al. Clinical, Endoscopic, and Histological Differentiations Between Crohn's Disease and Intestinal Tuberculosis. *Am J Gastroenterol* 2010; 105:642–651
27. Andronikou S, Welman CJ, Kader E. The CT features of abdominal tuberculosis in children. *Pediatr Radiol*. 2002; 32:75–81.
28. Singh, S., Singh, A., Prajapati, S. et al. Xpert MTB/RIF assay can be used on archived gastric aspirate and induced sputum samples for sensitive diagnosis of paediatric tuberculosis. *BMC Microbiol*. 2015; 15:191.
29. Working Group on Tuberculosis, Indian Academy of Pediatrics (IAP). Consensus statement on childhood tuberculosis. *Indian Pediatr*. 2010; 47:41–55.
30. Riquelme A, Calvo M, Salech F, et al. Value of adenosine deaminase (ADA) in ascitic fluid for the diagnosis of tuberculous peritonitis: a meta-analysis. *J Clin Gastroenterol* 2006; 40:705–10.
31. Tao L, Ning HJ, Nie HM, Guo XY, et al. Diagnostic value of adenosine deaminase in ascites for tuberculosis ascites: a meta-analysis. *Diagn Microbiol Infect Dis*. 2014; 79:102–107
32. Basu S, Ganguly S, Chandra PK, et al. Clinical profile and outcome of abdominal tuberculosis in Indian children. *Singapore Med J*. 2007; 48:900–905.
33. Singh V. Pediatric TB Management under RNTCP: What and Why? *Indian J Pediatr*. 2019; 86:707–713.
34. Jullien S, Jain S, Ryan H, et al. Six-month therapy for abdominal tuberculosis. *Cochrane Database Syst. Rev*. 2016; 11: CD012163.
35. Shah I, Uppuluri R. Clinical profile of abdominal tuberculosis in children. *Indian J Med Sci*. 2010; 64:204–209.
36. Devarbhavi H, Karanth D, Prasanna K, et al. Drug- Induced liver injury with hypersensitivity features has a better outcome: A single center experience of 39 children and adolescents. *Hepatology* 2011; 54:1344–50.
37. Dhar A, Bagga D, Taneja SB. Perforated tubercular enteritis of childhood: a ten year study. *Indian J. Pediatr*. 1990; 57: 713–6.