# Abdominal tuberculosis in children Report of 10 cases

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#### Abstract

Background: Abdominal tuberculosis is a rare location of this infection with a lot of diagnostic difficulties. The aim of this study was to review our experience of pediatric abdominal tuberculosis with that of the literature data.

Patients and methods: A retrospective study was conducted over a 7- year period; 10 cases of abdominal tuberculosis in children were reviewed (6 girls and 4 boys, mean age: 11 years, extremes 4-14). Eight patients enrolled in this study satisfied the following criteria: a positive culture for mycobacterium tuberculosis on samples of ascites fluid, sputum, urine, abscess puncture and/or caseating granulomas on histologic examination of biopsies obtained by endoscopy, percutaneous aspiration or needle biopsy or exploratory laparotomy. Two other patients (patients  $N^{\circ}$  1 & 5) had a favorable response within 10 weeks of antituberculous therapy.

Results: We observed 8 cases of peritoneal tuberculosis and 2 cases of intestinal tuberculosis. Extra-abdominal tuberculosis was found in 4 of those patients. Two patients had a contact with a tuberculosis positive individual. Nine children had BCG immunization.

The main clinical features were abdominal distension (6 cases) and abdominal pain (6 cases). Abdominal ultrasonography visualized mesenteric lymphadenopathies (6 cases), an abdominal mass (5 cases), free (1 case) and localized ascites (1 case). Barium enema and small bowell series showed small bowel stenosis (1 case) and ileal ulcerations (1 case).

Exploratory laparotomy, performed in 3 patients, showed whitish granulations and peritoneal abscesses with caseous necrosis on histology. Quadruple therapy with tuberculostatic agents was prescribed in all patients, associating isoniazid, rifampicin, pyrazinamide combined at the start of the treatment with streptomycin (7 cases) and ethambutol (3 cases). Short term outcome was favorable with no deaths. The long term outcome was complicated by persistent ascites (1 case) and the development of portal hypertension (1 case).

Conclusion: Abdominal tuberculosis remains a rare localization with a lot of diagnostic difficulties. (Acta gastroenterol. belg., 2004, 67, 245-249).

Key words: abdominal tuberculosis, mycobacterium, child.

In 1975, tuberculosis incidence in Tunisia was 49 cases per 100 000 people. Currently, it is 23 cases per 100 000 people. Almost 98.9 per cent of children are vaccinated by BCG vaccine. Tunisian national health situation has improved by the application of vaccination, pasteurization of milk, increased standard of living, and finally provision of health care. In developed countries, a surge in the number of patients presenting with gastrointestinal tuberculosis is reported (1). It is, partly, due to the epidemic of acquired immunodeficiency syndrome (AIDS) and the influx of patients from endemic areas of tuberculosis (2).

Pulmonary tuberculosis is by far the most frequent localization in Tunisia: 71.1% in 1992 (3). Abdominal tuberculosis is a rare disease characterized by its polymorphic clinico-pathological features and diagnostic difficulties.

The aim of this study was to review our experience of pediatric abdominal tuberculosis with that of the paediatric literature data.

#### Patients and methods

We conducted this retrospective study over a 7-year period, between 1989 and 1996 in our unit which is a part of a general pediatric department specialized in pediatric gastroenterology.

Ten cases of abdominal tuberculosis in children were reviewed (6 girls and 4 boys, mean age: 11 yrs, extremes 4 ½-14). Eight patients enrolled in this study satisfied the following criteria: a positive culture for mycobacterium tuberculosis on samples of ascites fluid, sputum, urine, abscess puncture and/or caseating granulomas on histologic examination of biopsies obtained by endoscopy, percutaneous aspiration or needle biopsy or exploratory laparotomy. Two other patients (patients N° 1 & 5) had a favorable response within 10 weeks of antituberculous therapy.

The Mantoux tuberculin skin test uses 5 tuberculin units of purified protein derivative (PPD) which are injected intradermally. The test is considered positive if an induration > 10 mm is obtained in children without prior BCG vaccination or > 15 mm in children with prior BCG vaccination.

Clinical characteristics, laboratory, radiological data, treatment and outcomes for the 10 children with abdominal tuberculosis are summarized in table I. Table II summaries the diagnostic criteria for the 10 enrolled children with abdominal tuberculosis.

# **Results**

During the 7 years of the study, 34357 children were hospitalized in our department. The incidence of

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Table I. — Clinical characteristics, laboratory and radiological data, treatment and outcomes for the 10 children with abdominal tuberculosis

	10	e Female	6 1/2		46 months 9 months	Insidious Insidious	Abdominal Ascites, weight loss, abdominal swelling, abdominal pain	ESR: 50/80 mm ESR: 28 mm Hb: 13 g/dL Hb: 11.6 g/dL	M. tuberculosis	Hepatomegaly, Localized ascites splenomegaly, multiple lymph nodes, portal hypertension				Mesenteric lymph Peritoneal node uberculosis uberculosis	tive negative	
	8	Male Male	4 1/2		10 days 46 m	acute Insid	Fever, weight loss, Abdd anorexia, pain Swelling of the right flank, abdominal pain	ESR: 60/105 mm ESR Hb: 9g/dL Hb:		Abdominal Hepa and pelvic mass splen multi node				Peritoneal and Mese mesenteric lymph node node tuberculosis	positive positive	
	7	Female	8	Grand father	2 months	Insidious	fever, pallor, peri umbilical mass, choreiform movements, clavicular lymphadenopathy	ESR: 100/125 mm Hb: 6.5 g/dL		Hypoechogenic mass, multiple nodes				Tuberculoma- Mesenteric lymph node and peritoneal tuberculosis	positive	
cicom	9	Female	13		9 months	Insidious	headache, fever, abdominal mass	ESR: 107 mm, Hb: 8.2 g/dL		Multiple nodes compressing inferior vena cava				Mesenteric lymph node and peritoneal tuberculosis	positive	
	5	Female	12		1 month	Insidious	Weight loss, fever, ascites, abdominal pain	ESR: 10/35 mm Hb: 11.5 g/dL		Hepatomegaly, splenomegaly, ascires; multiple mesenteric nodes	Internal capsula nodule		trepassing on left thalamus	Peritoneal tuberculosis	positive	
WIEII A	4	Male	13		2 1/2 years	Insidious	Chronic diarrhea, pallor, weight: -3SD, height: -3SD	ESR: 55/90 mm Hb: 8.4 g/dL	M. tuberculosis		Irregular narrowing of the distal portion of ileum.	Stenosing tumoral process, bulging in cecum, polyp of the left colon covered with congestive mucosa		Intestinal tuberculosis	negative	
	3	Female	111		15 months	Insidious	Right iliac osteitis, tuberculosis adenitis treated during 9 months, arthralgia, abdominal pain	ESR: 128/135 mm, Hb: 7.8 g/dL	M. tuberculosis	Splenomegaly, multiple mesenteric nodes, bilateral hydronephrosis	Ileum stenosis	Stenosis and fistula of the 3 <sup>st</sup> portion of duodenum		Intestinal, peritoneal, mesenteric lymph node tuberculosis, Miliary tuberculosis	negative	
	2	Female	14		1 month	Insidious	Weight loss, fever, mass of the hypogastrium, abdominal pain	ESR: 95/120 mm	M. tuberculosis	Multiple lymph nodes				Peritoneal and mesenteric lymph node tuberculosis-Miliary tuberculosis	negative	
	1	Female	10	Grand mother	1 year	Insidious	Cervical lymphadenopathy splenomegaly	ESR: 60/160 mm, Hb: 11 g/dL		Splenomegaly multiple lymph nodes				Mesenteric lymph node tuberculosis	positive	
	Case	Sex	Age (years)	Origin of contamination	Disease duration	Type of onset	Clinical data	Biology	Bacteriology	- Abdominal US:	- Barium enema / small bowel series :	- Fibroscopy/ coloscopy :	- CT scan:	Diagnosis	PPD skin test	

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Table II. — Diagnostic criteria for the 10 enrolled children with abdominal tuberculosis

	Observations
A positive culture for mycobacterium tuberculosis from:  - ascites liquid - sputum samples - urine - abscess puncture	N° 3, 10 N° 3, 4 N° 3 N° 3
Histologic examination of biopsy material obtained by:  - needle biopsy of lymph node - exploratory laparotomy  Favorable response to a trial of antituberculous therapy	N° 3, 6, 7 N° 2, 8, 9 N° 1, 5

abdominal tuberculosis was 2.91 cases per 10 000 hospitalizations. Abdominal tuberculosis represented 20% of all tuberculosis cases collected during this period.

The other tuberculosis infections were pulmonary tuberculosis and lymph node tuberculosis.

Six patients were country people. Data on the type of milk consumed were missing in the children's files. Two patients had a contact with a tuberculosis positive individual (grand parents). Nine children had BCG immunization. No one had AIDS.

Abdominal tuberculosis was primary in six children. Extra-abdominal tuberculosis was present in four patients: pulmonary tuberculosis in two patients, tuberculosis of the urinary tract in one case and tuberculous adenitis in another case.

Among our patients, there were 8 cases of peritoneal tuberculosis and two cases of small bowel tuberculosis (patients 3 and 4).

The main clinical features were abdominal distension (6 cases) and abdominal pain (6 cases).

# Laboratory findings

Erythrocyte sedimentation rate (ESR) was accelerated in all cases with a median of 88 (60 - 128 mm). Eight patients had anemia ranging between 6.5 and 11.5 g/dL.

Three cases exhibited hypoalbuminemia (median: 2.55 g/dl, ranges: 2.1-2.7 g/dl), whereas hypergamma globulinemia was seen in 2 other cases. Ascites fluid was haematic or yellow and exudative in 2 cases with a median protein concentrations of 53 g/L (46 and 53 g/L). Adenosine deaminase activity was not measured in our series.

## Bacteriological data

The PPD skin test was positive in 7 cases. Mycobacterium tuberculosis was detected and cultured in 4 patients: ascites liquid in 1, sputum samples in 1, urines in 1 and in the last one urines, sputum and abscess puncture culture were all positive. Mycobacterium tuberculosis was sensitive to all antituberculous antibiotics. No mycobacterium bovis was detected in our series.

Upper fibroscopy was performed in 3 patients, showing the following lesions; stenosis and fistula of the  $3^{rd}$  portion of the duodenum in one case associated with peritoneal and intestinal tuberculosis (patient  $N^{\circ}$  3), atrophic gastritis in one case with intestinal tuberculosis (patient  $N^{\circ}$  4), and an external compression of duodenum in one case of tuberculous adenitis (patient  $N^{\circ}$  9).

Chest radiographs showed patchy alveolar infiltrates in 4 cases, interstitial infiltrate and pleural effusion in 1 patient and mediastinal lymphadenopathies in 2 cases. Urinary tract radiography visualized mesenteric calcifications in 1 patient (patient  $N^{\circ}$  7).

Abdominal CT scan, performed in 3 patients, visualized mesenteric lymphadenopathies in two cases (patients  $N^{\circ}$  2 and 7).

In patient 4, small bowel series showed an irregular narrowing of the distal portion of ileum making Crohn disease diagnosis highly suspected. Colonoscopy showed a narrowing tumoral process of caecum evoking carcinoma of the colon. Mycobacterium tuberculosis was later detected in a culture of sputum samples.

In patient 3, in whom diagnosis of generalized tuberculosis was suspected, colonoscopy with biopsies showed a normal colic mucosa.

Exploratory laparotomy was performed in 3 patients, because of unclear diagnosis in two patients ( $N^{\circ}$  2 and 9) and an acute surgical abdominal tenderness suggesting digestive perforation in patient  $N^{\circ}$  8. In these patients, whitish granulations and peritoneal abscesses were visualized and histology revealed caseating granulomas. No laparoscopy was carried out in this series.

Quadruple tuberculostatic treatment was prescribed in all patients based on the combination of Isoniazid (10-15 mg/kg/day), Rifampin (10-20 mg/kg/day), Pyrazinamide (20-40 mg/kg/day), combined at the start of the treatment with Streptomycin (20-40 mg/kg/day) in 7 cases and Ethambutol (15-25 mg/kg/day) in 3 cases. The median duration of treatment was 10.5 months, (range: 6 and 19 months). No one received corticoids (Table III).

Short term outcome was favorable in all cases with no deaths. No adverse effects of antituberculous chemotherapy were noted. An outcome follow up of a median of 13 months (1-34 months) was favorable in 8 cases. Ascites persisted after 1 month of chemotherapy in patient  $N^{\circ}$  10 who continued his treatment in Algeria. Patient  $N^{\circ}$  9 developed portal hypertension due to portal obstruction by enlarged mesenteric lymph nodes.

# Discussion

Abdominal tuberculosis is rare in children. In our study, its incidence was 2.9 cases per 10 000 hospitalizations. Abdominal tuberculosis represented 20% of all tuberculosis cases collected during the study period. This might reflect an important recruitment bias. In Marroco, peritoneal tuberculosis represents 76.3% of abdominal tuberculosis and 34% of all tuberculosis

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Table III. — Therapeutic regimen for the 10 children with abdominal tuberculosis

Patient	Treatment	Duration
1	Iso-Rif-Pyr-Strep Iso-Rif-Pyr	2 months 6 months
2	Iso-Rif-Pyr-Eth Iso-Rif-Eth Iso-Rif	2 months 2 months 6 months
3	Iso-Rif-Pyr-Strep Iso-Rif-Pyr Iso-Rif	2 months 7 months 2 months
4	Iso-Rif-Pyr-Strep Iso-Rif-Pyr Iso-Rif	2 months 4 <sup>1</sup> / <sub>2</sub> months 2 months
5	Iso-Rif-Pyr-Strep Iso-Rif	2 months 9 months
6	Iso-Rif-Pyr-Strep Iso-Rif	2 months 6 months
7	Iso-Rif-Pyr-Strep Iso-Rif	2 months 7 months
8	Iso-Rif-Pyr-Strep Iso-Rif	2 months 17 months
9	Iso-Rif-Pyr-Strep Iso-Rif	2 months 11 months
10	Iso-Rif-Pyr-Eth Iso-Rif	2 months 6 months

Iso: Isoniazid Rif: Rifampicin Pyr: Pyrazinamide Strep: Streptomycin Eth: Ethambutol

cases in children (4). In developed countries, the number of extra-pulmonary tuberculosis in adults has increased by the late eighties (5,6).

Abdominal tuberculosis consists of 3 forms: peritoneal tuberculosis, intestinal tuberculosis and mesenteric lymph node tuberculosis, which are difficult to distinguish in children.

We collected 8 cases of peritoneal tuberculosis and 2 cases of intestinal tuberculosis. Veeragandham *et al.* (7) reported 26 cases of abdominal tuberculosis in children; 13 cases of intestinal tuberculosis and 9 cases of peritoneal tuberculosis.

The pathophysiology of tuberculous enteritis has been attributed to 4 mechanisms; 1) hematogenous spread from active pulmonary or miliary tuberculosis; 2) swallowing of infected sputum in patients with active pulmonary tuberculosis; 3) ingestion of contaminated milk or food and 4) contiguous spread from adjacent organs. After the tubercle bacillus enters the gastrointestinal tract, it crosses the mucosa to lodge in the submucosa. There, the presence of the bacillus induces inflammatory changes, including serosal and submucosal edema, cellular infiltration, and lymphatic hyperplasia. Eventually, the appearance of granulomata causes small papillary mucosal elevations. Lymphangitis, endarteritis, and fibrosis ensue, which lead to mucosal ulceration, caseating necrosis, and narrowing of the intestinal lumen (8).

Abdominal tuberculosis could be difficult to diagnose. Symptoms are vague and signs are nonspecific. In our study, clinical manifestations were not specific, mimicking several diseases such as celiac disease, periodic disease and Hodgkin lymphoma.

Ileocecal tuberculosis was seen in two of our patients (N° 3 and 4, Table I) and a differential diagnosis including Crohn disease and colon carcinoma was made in one case (patient N°4, Table I). Amebiasis, yersinia infection, gastrointestinal histoplasmosis and periappendiceal abscess were other possibilities (9). Surgery is applied in these cases of uncertain diagnosis.

Our study was characterized by the long delay of confirming diagnosis, more than two months in 6 cases. Abdominal pain was the most frequent observed sign, seen in 6 patients followed by weight loss and fever. In the literature, colicky mid-abdominal and right lower quadrant pain which reflects intermittent partial bowel obstruction is the presenting complaint in 90-100% of patients (5,6,10). PPD is positive in 70 to 86% of cases (7,9,10), while about 48% of patients have abnormality of chest x-ray (7,11). ESR ranged from 25 to 105 mm/h (7).

Table IV summarize the frequency of signs and laboratory findings in our study and literature.

Determination of adenosine deaminase level in the ascites fluid, an enzyme involved in the catabolism of purine bases, is particularly useful in the diagnosis of tuberculosis peritonitis. Its level is increased as a result of stimulation of T-lymphocytes in response to cell-mediated immunity to mycobacterial antigens (12).

Polymerase chain reaction assay, performed on endoscopic biopsy specimens, is a useful technique in the diagnosis of gastrointestinal tuberculosis in the absence of tissue and culture positivity (13). Determination of ascites fluid levels of adenosine deaminase and polymerase chain reaction assay are however not available in Tunisia.

Radiographic findings of tuberculosis enteritis are nonspecific. Abdominal ultrasonography could help positive diagnosis when it showes mesenteric lymph nodes, ileocecal mass or ascites. Barium enema and small bowel series are useful, supporting intestinal tuberculosis diagnosis in 66 per cent of cases (1). The most common findings are a pipe stream colon and a cone shaped retracted caecum (1). A wide gap between thickened ileocaecal valve and narrowed ileum is another suggestive finding (1).

For patients with predominantly colonic symptoms, colonoscopy is helpful in establishing the diagnosis (14). Typical colonoscopic findings include deformation of ileocaecal valve of caecum, segmental nodular and ulcerative lesions, and finally, areas of strictures with nodules and ulcerations (15).

Laparoscopy with direct peritoneal biopsy is an excellent study for diagnosing tuberculosis peritonitis and should be considered in patients with unexplained ascites. Findings include multiple whitish "miliary" nodules scattered over the visceral and parietal

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Table IV. — Frequency of signs	and	laboratory	findings	in
our study and	liter	ature		

	Our study	11, 14, 17	Veeragandham et al. (7)
Fever	40%	54-100%	53.8%
Ascites	20%	51-100%	34.6%
Abdominal pain	60%	36-93%	15.3%
Weight loss	70%	37-87%	61.1%
Abdominal swelling	60%	65-100%	23%
Anemia	80%	48-68%	
Positive PPD skin test	70%	55-100%	88.4%

peritoneum, adhesions between the peritoneum and organs (16). These findings were found in 3 of our patients during laparotomy. Laparoscopically guided peritoneal biopsies will detect caseating granulomas in about 72 to 95% of cases (6,11,16)).

The treatment of abdominal tuberculosis is primarily a medical one. Surgery is reserved for complications (obstruction, perforation, massive hemorrhage, abscess, and fistula). Conservative operations are generally recommended. Bowel resection should be reserved to circumferential or partially obstructing lesions (2).

A common effective treatment regimen is the daily triple anti-tuberculosis therapy (isoniazid, rifampicin, pyrazinamide) for two months followed by the bi anti-tuberculosis therapy daily or twice a week for 7 months (17). During the last few decades, recommendations for treament of tuberculosis in children have focused on shorter, more intense regimens and on increased adherence to treatment. A 6 month short-course chemotherapy regimen has then been used with great success (18). This resulted in reduced cost and a minimization of the development of drug-resistant organisms.

The response to chemotherapy usually is excellent. Deaths in the beginning of treatment due to meningitis, small bowel obstruction or perforation have been reported (2). These severe complications could be prevented by an early diagnosis and treatment.

Long term outcome was complicated in two of our patients by persistent ascites and portal hypertension. In the literature, presentation of abdominal tuberculosis with portal hypertension is rare and reported cases are adult ones (19,20,21). Mechanisms invoked for such portal hypertension are compression of the portal vein by tuberculous lymph nodes at the hepatic hilum and disseminated tuberculosis involving the liver and spleen. In one reported case, after antitubercular therapy, features of portal hypertension disappeared as the nodes regressed (22).

## Conclusion

Despite a large program of BCG vaccination, abdominal tuberculosis remains a non negligible percentage of tuberculosis in Tunisia. Abdominal pain, weight loss, and fever are the most indicative signs but are non specific.

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