

# Training Module On Extrapulmonary Tuberculosis

STANDARD TREATMENT WORKFLOW

## ADULT PERIPHERAL LYMPHADENOPATHY

ICD-10-A18.2

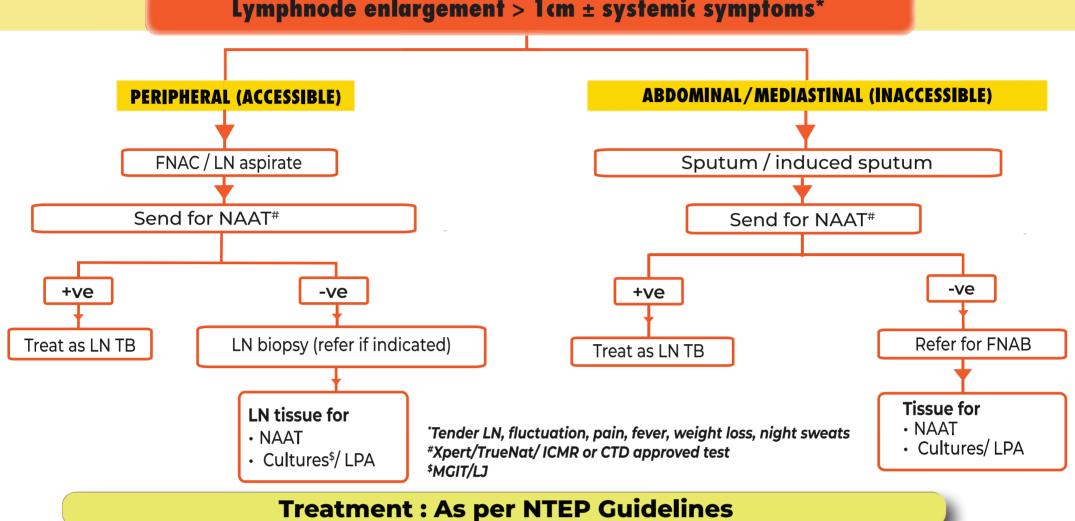
# SUSPECT:

#### WHEN TO SUSPECT?

- · Swelling (>1 cm) in neck, armpit or groin (>2 cm) +/- redness, fluctuation, sinus discharge
- · May or may not be associated with fever, weight loss, night sweats or cough
- · History of similar swelling in the past / past history of tuberculosis
- · History of contact with a patient with a diagnosis of TB

## **DIAGNOSTIC ALGORITHM**

## Lymphnode enlargement > 1cm ± systemic symptoms\*



## **ASSESS RESPONSE TO THERAPY AT 3-4 MONTHS**

- · Resolution: Decrease in size of LN with settling of systemic symptoms
- Delayed response -Paradoxical reaction: Increase in size of LN or new signs of inflammation (up to 3 months of starting treatment) OR appearance of new LN at same/other site
- · May require tissue cultures, if not done, to rule out treatment failure/resistance
- Therapeutic drug monitoring to ensure adequate drug levels
- · If cultures reveal susceptible TB it is likely due to paradoxical worsening: May require anti-inflammatory agents (inaccessible)/ surgical removal (accessible)

#### **COMPLICATIONS**

- Abscess formation
- Rupture may lead to sinus formation

#### REFER TO HIGHER CENTRE IF

- Non responders
- Needs treatment for Drug Resistance
- Large Nodal Mass/Abscess requiring surgical intervention

#### Response to therapy at 3 months No Yes Symptom resolution Constitutional Increase in size of LN ± signs of inflammation symptoms persist **Continue ATT** Check old culture report Send new cultures if not R/o alternate diagnosis available Susceptible TB Drug resistant TB Paradoxical Therapeutic drug Refer to DOTS worsening monitoring plus centre Refer to higher centre

#### **BCG LYMPHADENITIS**

- · Age is usually < 2 years
- · Axillary and/or supraclavicular LN on same side as BCG vaccination (usually given on left)
- No systemic symptoms in immunocompetent children
- · Treatment:
  - · Wait & watch if small
  - · If large & suppurative, repeated aspiration or rarely incision & drainage is required

NAAT/AFB smear positivity can not differentiate between BCG & MTB

#### **ABBREVIATION**

**ATT:** Anti Tubercular Treatment **BCG:** Bacille Calmette Guerin

CTD: Central TB Division DOT: Directly Observed Treatment LJ: Lowenstein Jensen Short-course LN: Lymph Node

**FNAC:** Fine Needle Aspiration Cytology

© Indian Council of Medical Research and Department of Health Research, Ministry of Health & Family Welfare, Government of India.

FNAB: Fine Needle Aspiration Biopsy LPA: Line Probe Assay **MGIT**: Mycobacterial Growth Indicator Tube

> MTB: Mycobacterium Tuberculosis **NAAT:** Nucleic Acid Amplification Test

**NTEP**: National TB Elimination **Programme PCR**: Polymerase Chain Reaction **TB**: Tuberculosis

## REFERENCES

- 1. National TB Elimination Programme, Central TB Division. Training Modules for Programme Managers & Medical Officers. Ministry of Health & Family Welfare, Government of India.
- https://tbcindia.gov.in/index1.php?lang=1&level=1&sublinkid=5465&lid=3540 Last accessed on 11 March, 2022. 2. Guidelines for programmatic management of drug resistant tuberculosis in India March 2021. National TB Elimination Programme, Central TB Division, Ministry of Health & Family Welfare, Government of India. https://tbcindia.gov.in/showfile.php?lid=3590 Last accessed on 11 March, 2022.
- 3. Gaikwad P, Samuel VM, Rupali P. Tb or not Tb. Paradoxical response and the role of selective lymphadenectomy in tuberculous cervical lymphadenitis. Indian J of Applied Research, October 2016; Vol 6(10): 40-43. 4. Alsultan A, Peloquin CA. Therapeutic drug monitoring in the treatment of tuberculosis: an update. Drugs, 2014 Jun; 74(8):839-54. Erratum in: Drugs. 2014 Jun; 74(9):2061.

This STW has been prepared by national experts of India with feasibility considerations for various levels of healthcare system in the country. These broad guidelines are advisory, and are based on expert opinions and available scientific evidence. There may be variations in the management of an individual patient based on his/her specific condition, as decided by the treating physician. There will be no indemnity for direct or indirect consequences. Kindly visit our web portal (stw.icmr.org.in) for more information.

## **ADULT PLEURAL TUBERCULOSIS**

ICD-10-A15.6

**HISTORY** 

**EXAMINATION** 



#### **ESSENTIAL**

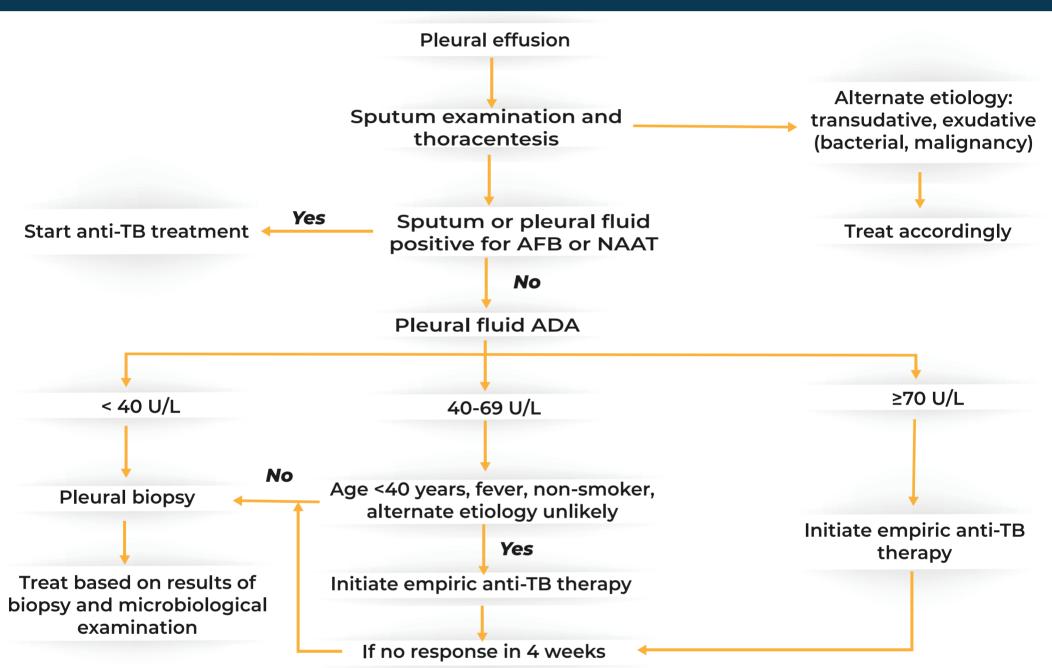
**INVESTIGATIONS** 

- · CXR (to confirm pleural effusion)
- Sputum for AFB/NAAT
- Refer immediately for pleural tap
- Thoracentesis (ultrasound-assisted)
- Pleural fluid analysis:
- Cell count (total and differential)
- → Protein
- → Glucose
- → Gram stain
- → Bacterial cultures
- → Stain for acid-fast bacilli
- Adenosine deaminase (ADA)
- → NAAT
- Cytology evaluation

#### DESIRABLE

- CT chest (before pleural biopsy)
- Pleural biopsy (image-guided/thorac oscopic) (If diagnosis is uncertain)
- Histopathology
- MGIT

## **DIAGNOSTIC**



## **MANAGEMENT**

#### TREATMENT AND RESPONSE

- As per NTEP
- Therapeutic pleural tap can be done under ultrasound assistance if the effusion is large, and the patient is breathless

#### WHEN TO REFER?

- Facility for ultrasound assistance is not available
- Diagnosis is not established after thoracentesis and facilities for pleural biopsy is not available
- Drug-resistant TB is detected: according to NTEP
- Worsening pleural effusion on follow up

## **FOLLOW UP**

- Most patients who respond to treatment will have improvement in their general condition by 2 weeks, and significant improvement in pleural effusion by 4-8 weeks
- Disappearance of constitutional symptoms with decrease in pleural effusion suggests responsiveness to treatment
- Increase in pleural effusion can suggest
  - › Paradoxical reaction or
  - Drug-resistant TB or
  - Alternative etiology
- · A follow up CXR at 4-8 weeks after starting ATT is useful to assess progress

#### **ABBREVIATIONS**

**ADA**: Adenosine Deaminase **AFB**: Acid-fast Bacilli **ATT**: Anti Tubercular Treatment **CT**: Computed Tomography

**CXR**: Chest Radiograph **MGIT**: Mycobacterial Growth Indicator Tube **NAAT:** Nucleic Acid Amplification Test

**NTEP**: National Tuberculosis Elimination Programme

**TB**: Tuberculosis

## REFERENCES

- 1. National TB Elimination Programme, Central TB Division. Training modules for Programme Managers & Medical Officers. Ministry of Health & Family Welfare, Government of India
- https://tbcindia.gov.in/index1.php?lang=1&level=1&sublinkid=5465&lid=3540 Last access on 11 March, 2022. 2. Guidelines for programmatic management of drug resistant tuberculosis in India March 2021. National TB Elimination Programme, Central TB Division, Ministry of Health & Family Welfare, Government of India. https://tbcindia.gov.in/showfile.php?lid=3590 Last access on 11 March, 2022.
- 3. Sharma SK, Ryan H, Khaparde S, et al. Index-TB guidelines: Guidelines on extrapulmonary tuberculosis for India. Indian J Med Res. 2017;145(4):448-463. doi:10.4103/ijmr.IJMR\_1950\_16 4. Lo Cascio CM, Kaul V, Dhooria S, Agrawal A, Chaddha U. Diagnosis of tuberculous pleural effusions: A review. Respir Med. 2021;188:106607. doi:10.1016/j.rmed.2021.106607.
- 5. Light RW. Update on tuberculous pleural effusion. Respirology. 2010 Apr;15(3):451-8. doi: 10.1111/j.1440-1843.2010.01723.x.

## Standard Treatment Workflow (STW) for the Management of **PERICARDIAL TUBERCULOSIS**

ICD-10-A18.84

#### WHEN TO SUSPECT

#### **SYMPTOMS**

- · Cough, fever, breathlessness or pleuritic chest pain
- May be associated with weight loss, night sweats or difficulty lying down
- Past history or a history of contact with a patient with a diagnosis of tuberculosis
- Examination reveals tachycardia, increased jugular venous pressure, hepatomegaly, ascites, & peripheral edema
- A pericardial friction rub and distant heart sounds present on cardiovascular examination
- If clinical picture +/- heart US suggest pericarditis or pericardial effusion refer for echo-cardiogram

#### **COMPLICATIONS**

**Constrictive pericarditis:** Clinical signs for recognition include

- · Kussmaul's sign (lack of an inspiratory decline in jugular venous pressure)
- Elevated & distended jugular veins with a prominent Y descent (second inward deflection of internal jugular pulse due to diastolic inflow of blood into the right ventricle)
- Pericardial knock (rare)

## Cardiac tamponade: Clinical

- signs include Sinus tachycardia
- Hypotension with a narrow pulse pressure
- Elevated JVP jugular venous pressure
- Muffled heart sounds
- Pulsus paradoxus (a decrease in systolic blood pressure by >10 mmHq on inspiration)
- Ascites

## **Essential tests:**

- Chest X-ray
- · ECG
- Echocardiogram

#### **INVESTIGATION**

#### Desirable:

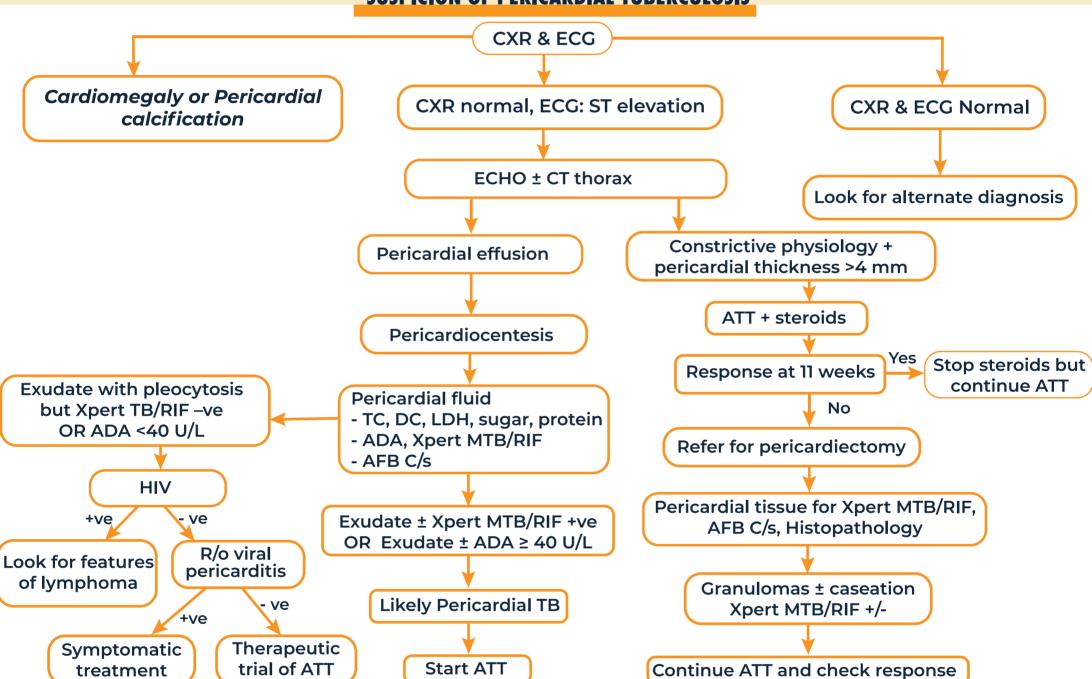
- Cardiac enzymes
- · CT/MRI of Thorax
- Pericardiocentesis Pericardial biopsy

## Other complications:

- · Myopericarditis: Abnormal ejection fraction with evidence of myocarditis and pericarditis (elevated cardiac enzymes & ST elevation on ECG)
- Effusive constrictive pericarditis: Mixed clinical picture. Main clue is elevated JVP clinically & right atrial pressure on ECHO in spite of removal of pericardial fluid

## **DIAGNOSIS**

## SUSPICION OF PERICARDIAL TUBERCULOSIS



## **MANAGEMENT**

#### **TREATMENT**

- Antitubercular therapy is advised as per NTEP
- Steroids are recommended in large pericardial effusions, prominent pleocytosis & pericardial fluid with high inflammatory markers or early constriction
- Give Prednisolone 60 mg/day for 4 weeks, 30 mg/day for 4 weeks, 15 mg/day for 2 weeks & 5 mg/day for 1 week
- Total duration of systemic steroids is 11 weeks depending on the regimen may be used

## NON RESPONSE TO STEROIDS & ATT

- Should prompt a referral to a specialist center for confirmation of diagnosis
- Non response of cardiac symptoms to anti-tuberculous therapy cardiac surgical evaluation may be required
- · Should prompt an evaluation for alternative causes of effusio-constrictive pericarditis
- Common alternate diagnosis include viral infections, Systemic lupus erythematosus, primary effusion lymphomas or pericardial malignancies

## **ABBREVIATION**

**ADA: Adenosine Deaminase ATT: Antituberculous Therapy** 

CXR: Chest X-ray ECG: Electrocardiogram **ECHO: Echocardiogram** 

JVP: Jugular Venous Pressure

**NTEP: National Tuberculosis Elimination Programme** 

**TB: Tuberculosis** 

- 1. National TB Elimination Programme, Central TB Division. Training Modules for Programme Managers & Medical Officers. Ministry of Health & Family Welfare, Government of India accessed at https://tbcindia.gov.in/index1.php?lang=1&level=1&sublinkid=5465&lid=3540 Last access on 15 March, 2022.
- 2. Guidelines for programmatic management of drug resistant tuberculosis in India March 2021. National TB Elimination Programme, Central TB Division, Ministry of Health & Family Welfare, Government of India https://tbcindia.gov.in/showfile.php?lid=3590 Last access on 15 March, 2022.
- 3. Nahid P, Dorman SE, Alipanah N, et al. Official American Thoracic Society/Centers for Disease Control and Prevention/Infectious Diseases Society of America Clinical Practice Guidelines: Treatment of Drug-Susceptible Tuberculosis. Clin Infect Dis. 2016 Oct 1;63(7):e147-e195. doi: 10.1093/cid/ciw376. Epub 2016 Aug 10.

## **ADULT TUBERCULAR MENINGITIS**

ICD-10-17.0

#### SUSPECT TBM WITH **FOLLOWING CLINICAL FEATURES**

Fever (Duration of 5 days or more<sup>#†</sup>)

- Headache
- Vomiting
- · Altered sensorium
- Cranial nerve palsy
- · Hemiparesis/any limb weakness
- Seizures
- Neck pain and stiffness



- · Active TB elsewhere Past history of TB & ATT
- Contact with TB patient
- HIV seropositivity
- · Low socio-economic status
- High endemic area



**ALWAYS ENQUIRE FOR ASSOCIATED FEATURES** 

\*This is to increase sensitivity for diagnosis of TBM. The duration could be variable from days to weeks to months. †Clinical judgement & evaluation of other conditions is also required as fever can be associated with headache in other medical conditions. Delaying work up for meningitis is not recommended.

#### IF TBM SUSPECTED

Refer to a centre where facility of evaluation (at least Lumbar puncture & CT scan) is available.

### **EVALUATION AT CENTRE OF CARE**

#### **CLINICAL HISTORY & EXAMINATION**

- Symptoms type & duration, onset & progression
- · Headache, altered sensorium, focal deficits
- Neck rigidity, Kernig's sign
- Cranial nerve palsy
- · Fundus examination papilledema

#### **LABORATORY EVALUATION**

- · CBC, ESR, CRP
- LFT, RFT, Electrolytes
- · Blood sugar
- · HIV
- Chest X Ray- PA view
- USG whole abdomen
- Mantoux (optional)

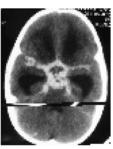
#### **IMAGING**

- NCCT/CECT head- Preferred as initial investigation
- MRI brain (and spine if indicated) in selective cases

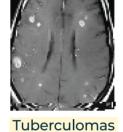
#### **CSF**

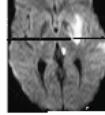
- Mandatory- Should be sent for essential analysis (Box 1)
- Prudent to perform CT head prior to CSF in presence of papilledema & /or focal deficits

#### **COMMON NEUROIMAGING FINDINGS IN TBM**

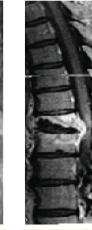








Infarction



**Arachnoiditis** 

Pott's spine

### **CSF EVALUATION\***

02

01

Basal exudates and Hydrocephalus

#### **ESSENTIAL**

- · Cell count & type
- Protein
- Sugar (& Corresponding blood sugar)
- GenXpert / TrueNat
- · Grams stain
- Bacterial culture
- AFB stain
- AFB culture/sensitivity
- ullet India Ink $^*$
- Cryptococcal antigen\*

**DESIRABLE** 

- · Fungal smear & culture
- Cytopathology\*

**OPTIONAL** 

- Wet mount
- VDRL
- Toxoplasma PCR<sup>†</sup>

03

Viral PCR

If some tests are not available at site, store sample in sterile container, keep in refrigerator & transport in icebox to other facility

'CSF samples should be sent to the lab as soon as possible for examination of cells, protein, sugar and cytology. "Cryptococcal meningitis should be excluded wherever possible as it is a close differential diagnosis of TBM. \*In ideal settings, it may be prudent to exclude a diagnosis of carcinomatous meningitis. †Especially in patients with HIV.

## **CSF FINDINGS IN TBM AND OTHER MENINGITIS**

MENINGITIS TYPE	CELL COUNT	PREDOMINANT CELL TYPE	PROTEIN	SUGAR	SPECIFIC TESTS FOR CONFIRMATION
Tubercular	Usually <500	Lymphocytic Neutrophilic in some acute cases	High	Low	AFB smear and culture  *Xpert / TrueNat
Pyogenic	In thousands	Neutrophilic	Moderately High	Very low	Gram stain, culture
Fungal	Variable	Lymphocytic	High	Low	India Ink, Fungal Culture, Cryptococcal antigen
Viral	50-500	Lymphocytic	Normal to marginally high	Normal	PCR for specific virus

\*A negative Xpert / TrueNat result does not rule out TBM. The decision to give ATT should be based on clinical features and CSF profile.

## **MANAGEMENT**

#### ANTI-TUBERCULAR TREATMENT

 Intensive Phase: 2 months of RHZE or RHZS · Continuation phase: 3 drugs: RHZ for at least 10 months

#### **STEROIDS**

- Preferably Dexamethasone 0.4 mg/kg/day intravenously in 3-4 divided doses during hospital stay
- · If not feasible, give oral Dexamethasone 0.4 mg/kg/day in divided doses or oral Prednisolone 1 mg/kg/day in a single morning dose
- Discharge on oral steroids on tapering doses for a total duration of 8-12 weeks

#### treatment duration may be increased in some cases as per the clinician decision

#### **FOLLOW UP**

- Regular follow up is essential every month for at least first 3 months & can be increased thereafter till treatment is
- Monitor liver function tests & any other features of drug toxicity
- Observe for clinical improvement or any deterioration
- Closely observe for development of any complications

© Indian Council of Medical Research and Department of Health Research, Ministry of Health & Family Welfare, Government of India.

#### SUSPECT COMMON COMPLICATIONS

Hydrocephalus and raised ICP: Worsening of headache with vomitings and/or altered sensorium

Optico-chiasmatic arachnoiditis: Complaints of vision loss in one or both eyes with or without headache

Myelitis and or arachnoiditis: Development of paraparesis or quadriparesis with/without sensory disturbances, bladder

**Epidural abscess/Pott's spine:** Complaints of back pain and/or weakness in one/ both lower limbs/ bladder/ bowel disturbances Tuberculoma: Seizures, new onset focal focal deficits, worsening headache

**Seizures:** Consider tuberculoma/electrolyte or metabolic imbalance/ cerebral infarction

Cerebral infarction and stroke: Sudden onset weakness of one half of body, new onset confusion, altered mental status, seizures Hyponatremia, SIADH: Persistent or worsening mental status

**ATT**: Antitubercular therapy **CBC**: Complete Blood Count **CECT**: Contrast Enhanced CT **CRP**: C Reactive Protein CSF: Cerebrospinal Fluid

E: Ethambutol ESR: erythrocyte sedimentation rate H: Isoniazid ICP: Intracranial pressure

**MRI**: Magnetic resonance imaging **NCCT**: Non-contrast CT

**ABBREVIATIONS** 

**NTEP:** National TB Elimination Programme PCR: Polymerase Chain Reaction

**RFT**: Renal function tests S: Streptomycin SIADH: Syndrome of inappropriate antidiuretic hormone **TBM**: Tubercular meningitis

LFT: Liver function tests R: Rfimapicin Z: Pyrazinamide REFERENCES

1. Thwaites G, Fisher M, Hemingway C, Scott G, Solomon T, Innes J; British Infection Society. British Infection Society guidelines for the diagnosis and treatment of tuberculosis of the central nervous system in adults and children. J Infect. 2009;59:167-87.

- 2. Thwaites GE, Chau TT, Stepniewska K, Phu NH, Chuong LV, Sinh DX, White NJ, Parry CM, Farrar JJ. Diagnosis of adult tuberculous meningitis by use of clinical and laboratory features. Lancet. 2002;360:1287-92. 3. Vibha D, Bhatia R, Prasad K, Srivastava MV, Tripathi M, Kumar G, Singh MB. Validation of diagnostic algorithm to differentiate between tuberculous meningitis and acute bacterial meningitis. Clin Neurol Neurosurg. 2012;114:639-44.
- 4. Modi M, Prabhakar S. Fever and altered sensorium. In Singh MB, Bhatia R eds. Emergencies in Neurology. 2nd edition. 2021. 91-106. 5. INDEX-TB guidelines. https://tbcindia.gov.in/showfile.php?lid=3245 Last access on 05/03/2022.

This STW has been prepared by national experts of India with feasibility considerations for various levels of healthcare system in the country. These broad guidelines are advisory, and are based on expert opinions and available scientific evidence. There may be variations in the management of an individual patient based on his/her specific condition, as decided by the treating physician. There will be no indemnity for direct or indirect consequences. Kindly visit our web portal (stw.icmr.org.in) for more information.

# **ADULT ABDOMINAL TUBERCULOSIS**

ICD-10-A18.3

## WHEN TO SUSPECT

Any organs in abdominal cavity, including gut lumen & peritoneum may be affected						
PERITONEAL	INTESTINAL	ESOPHAGEAL	GASTRO-DUODENAL	PERIANAL	PANCREATIC	HEPATO-BILIARY
<ul><li>Abdominal distension</li><li>Pain abdomen</li><li>Fever</li></ul>	<ul> <li>Recurrent intestinal colic</li> <li>Partial/incomplete intestinal obstruction</li> <li>Chronic diarrhoea</li> <li>Weight loss</li> <li>Palpable mass abdomen</li> <li>Lower gastrointestinal</li> </ul>	<ul><li>Dysphagia</li><li>Odynophagia</li><li>Hematemesis</li><li>Constitutional symptoms</li></ul>	<ul><li>Gastric outlet obstruction</li><li>Gastrointestinal bleed</li></ul>	<ul> <li>Simple/ Complex peri-anal fistula</li> <li>Persistent discharge</li> <li>Fistulae which recur after multiple surgeries</li> </ul>	<ul> <li>Abdominal pain</li> <li>Obstructive jaundice</li> <li>Dilated pancreatic or bile duct with (peri)-pancreatic mass or cyst</li> <li>Constitutional symptoms</li> </ul>	<ul><li>FUO</li><li>Hepatomegaly</li><li>Jaundice</li><li>Elevated ALP</li><li>SOL</li><li>Hepatic abscess</li></ul>

## **EVALUATION FOR SUSPECTED ABDOMINAL TUBERCULOSIS**

#### **Pancreatic Peritoneal TB Perianal TB Gastrointestinal TB Hepatobiliary TB** Ascitic tap · Imaging: Malignancy more >CT or MR enterography (preferred): bowel Uncommon Cytology (lymphocyte) important concern thickening, strictures, necrotic LN predominance, absence · Imaging: MRI Needs imaging/ of malignant cells) OCT chest may also be done pelvis/Endoanal Endoscopic SAAG: <1.1 gram/dL · Endoscopic biopsies: UGIE for ultrasound/EUS ultrasound guided esophageal/gastroduodenal/ileo-colonoscopy Culture, NAAT · Tissue acquisition tissue acquisition for intestinal TB is mandatory to acquire tissue Ascitic ADA: >39 U/L for histology & · Refer to higher **Histology:** Caseating granuloma specific; **Imaging** microbiology centre confluent granuloma & ulcers lined by Ultrasound: Ascites, histiocytes suggestive lymph nodes, bowel thickening, for tissue Microbiology: Culture, NAAT • Barium studies: if endoscopy/CT unavailable acquisition for cytology & microbiology If diagnosis unclear If diagnosis unclear Refer for further evaluation (differentiating Crohn's disease is challenging) **Refer for Laparoscopy**

HIV & blood sugar test should be done in all suspected patients as per NTEP guidelines

Trial of Anti tubercular therapy with careful follow-up

#### FOLLOW UP

## **Definitive Abdominal TB**

(Microbiological positive, Caseating granulomas)

bleeding

Clinically Diagnosed Abdominal TB

(Consistent clinical/radiological picture, Elevated ascitic ADA, granuloma on histology, exclusion of differential diagnosis)

**START ATT (2HRZE, 4HRE)** 

#### Clinical Assessment (4 & 8 weeks)

- No fever/pain, disappearance of abdominal distension, weight gain
- Assess ATT induced hepatotoxicity with serial LFTs

# Improvement + Assessment for clinically diagnosed cases • Peritoneal TB: Ultrasound for ascites (4-8 weeks) Intestinal TB: Ileo-colonoscopy (8-12 weeks) Improvement

Refer Refer **Complete ATT Complete ATT** (Exclude alternative diagnosis) (Exclude MDR)

- End of therapy: Document mucosal healing in colonic & upper GI TB
- Imaging in small bowel disease to document healing

## TREATMENT:

- Start treatment & follow-up as per NTEP guidelines
- · 1st line treatment for adults & children with abdominal TB: 2HRZE/RHRE
- Extend duration of treatment in cases of inadequate response
- Refer for surgical management for complications [intestinal obstruction (due to strictures), perforation]. Consider endoscopic dilatation for treatment for accessible strictures
- Refer for biliary drainage in case of Jaundice due to biliary obstruction (hepatobiliary obstruction/pancreatic TB)

#### **ABBREVIATIONS**

**ADA**: Adenosine Deaminase **ALP:** Alkaline phosphatase **ATT**: Anti-Tubercular treatment CT: Computed Tomography EUS: Endoscopic ultrasound

**FUO:** Fever of Unknown Origin **GI:** Gastro-intestinal

HRZE: Isoniazid; Rifampicin; Pyrazinamide; Ethambutol

LFT: Liver function tests MDR: Multi-drug resistance **MR:** Magnetic Resonance

Mtb: Mycobacterium Tuberculosis

Improvement +

NTEP: National TB Elimination

Programme

Rif: Rifampicin

**SOL:** Space occupying Lesion

NAAT: Nucleic Acid Amplification Test SAAG: Serum Ascites Albumin Gradient **UGIE:** Upper gastrointestinal endoscopy

#### REFERENCES

- 1. National TB Elimination Programme, Central TB Division. Training Modules for Programme Managers & Medical Officers. Ministry of Health & Family Welfare, Government of India. https://tbcindia.gov.in/index1.php?lang=1&level=1&sublinkid=5465&lid=3540 Last access on 08 March, 2022.
- 2. Guidelines for programmatic management of drug resistant tuberculosis in India March 2021. National TB Elimination Programme, Central TB Division, Ministry of Health & Family Welfare, Government of India accessed at https://tbcindia.gov.in/showfile.php?lid=3590 Last access on 08 March, 2022.

## **ADULT MUSCULOSKELETAL TUBERCULOSIS**

ICD-10-A18.0



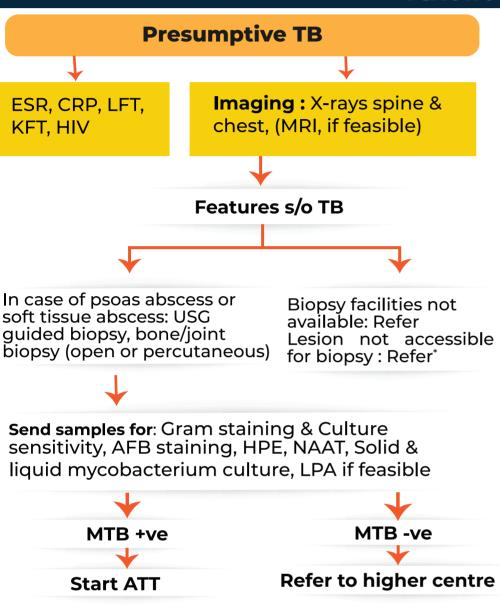
## SPINE TUBERCULOSIS

- Persistent localized pain in spine region >6 weeks, night pains
- · Local tenderness/cold abscess
- · Recent onset deformity in the back
- Recent neurological deficit (better to refer\*)
- Persistent heaviness around the waist/Girdle pain
- Fever, cough, weight loss & night pains
- · History of close contact with TB

## OTHER JOINTS/BONES

- Persistent localized pain & swelling >6 weeks
- Mono-articular joint involvement
- Discharging sinus (+/-)
- Fluctuant swelling with or without inflammation
- Painful restriction of involved joint movements
- · Wasting around the area
- Fever, cough, weight loss & night pains
- History of close contact with TB

## **DIAGNOSTIC ALGORITHM**





Paravertebral shadow

Obliterated disc T<sub>1</sub>WI and T<sub>2</sub>WI images space & bone loss bone edema with VB in X-rays destruction

T<sub>2</sub>WI septate pre/para vertebral abscess in MRI

#### Findings S/o TB

- X-ray findings(spine):
- > Regional Osteopenia
- Decreased/obliterated disc space
- > Vertebral erosions +/- reduced vertebral height
- → Paravertebral shadow
- MRI findings (Spine):
- > Contagious VB involvement with relatively preserved disc
- > Pre & paravertebral septate collection (Abscess)
- > Epidural encroachment +/- intraosseous abscess
- X-ray & MRI Finding (extraspinal):
- Regional osteoporosis with bone destruction on X-rays
- > Inflammation of bone(T1WI & T2WI) +/- abscess on MRI

## **TREATMENT**

Treatment should be started & follow-up should be conducted as per NTEP guidelines

The following algorithm provides additional guidance for follow-up

TB +ve on any test

TB -ve on all tests

NAAT sensitive to Rifampicin: Start 4 drug first line ATT

GeneXpert resistant to
Rifampicin: refer\*/culture for TB
and sensitivity to other drugs

Index of suspicion high ESR, CRP raised: Refer\* Index of suspicion low CRP normal:
Reassurance

- · Clinical symptoms improvement
- CRP decreasing continue for standard 12 months regime
- · Intensive 4 drug regime (not more than 4 months)
- Stop ATT after 12 months if all three parameters clinical, Lab(ESR, CRP) & radiological return to normal
- In case of spine decision to stop ATT to be taken by evaluating healed status on contrast MRI
- · Mildly elevated ESR, CRP (non specific tests) can be ignored
- Follow up every month with CRP, LFT during intensive phase
- $\cdot$  Follow up every 3 months during continuation phase with CRP/LFT
- On treatment worsening of symptoms
  - > Early (<3 months): Paradoxical
  - > Late (>4 months): ?drug resistance
- · Any aberrance in course such as appearnce of nerual deficit: Refer\*

Clinical symptoms not improving ESR, CRP increasing: Refer\*

?Suspected drug resistance

\*Refer to higher centre where advanced diagnostic, & therapeutic facilities including surgical procedures are available.

#### **ABBREVIATIONS**

AFB: Acid-fast Bacillus
ATT: Anti-Tubercular Treatment
CRP: C-Reactive Protein
ESR: Erythrocyte Sedimentation Rate

**HIV**: Human Immunodeficiency Virus **HPE**: Histopathological examination

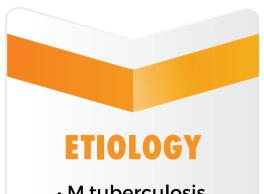
KFT: Kidney Function Tests LFT: Liver Function Tests LPA: Line Probe Assay MRI: Magnetic Resonance Imaging
NTEP: National TB Elimination Programme
TB: Tuberculosis
USG: Ultrason bography
VB: Vertebral body
WNL: Within Normal Limits

#### REFERENCES

- 1. National TB Elimination Programme, Central TB Division. Training Modules for Programme Managers & Medical Officers. Ministry of Health & Family Welfare, Government of India https://tbcindia.gov.in/index1.php?lang=1&level=1&sublinkid=5465&lid=3540 Last accessed on 10 March, 2022.
- 2. Guidelines for programmatic management of drug resistant tuberculosis in India March 2021. National TB Elimination Programme, Central TB Division, Ministry of Health & Family Welfare, Government of India https://tbcindia.gov.in/showfile.php?lid=3590 Last accessed on 10 March, 2022.

## **CUTANEOUS TUBERCULOSIS**

ICD-A18.4



- M.tuberculosis
- M.bovis
- BCG (rarely)

#### WHEN TO SUSPECT

- Presence of ulcer or discharging sinus over lymph node, bone & ioints
- Persistent asymptomatic reddish/ reddish brown lesion of >6 months duration which may show scarring
- Persistent warty or verrucous lesion of >6 months duration

#### OTHER FEATURES

- Extracutaneous involvement
- Lymph node & lungs involvement
- Other organ systems involvement (bones, GIT & CNS)

#### **TYPES OF CLINICAL DISEASE**

- Primary Inoculation tuberculosis
- · Tuberculosis verrucosa cutis
- Lupus vulgaris
- Scrofuloderma
- Acute miliary tuberculosis
- Orificial tuberculosis Metastatic tuberculous
- Abscess (tuberculousgumma)
- Normal primary complex-like reaction
- Postvaccination
- · lupus vulgaris Perforating regional adenitis
- Lichen scrofulosorum and
- Papulonecrotictuberculid
- Facultative tuberculids
- Nodular vasculitis and Erythema nodosum













**INVESTIGATION** 

#### SCREENING FOR SYSTEMIC INVOLVEMENT

- Examination:
  - › Lymph node to be examined (FNAC)
  - Other organ system can be done if indicated
- · Essential:
  - → Chest X-ray
  - > FNAC from theindurated part of lesion
- Desirable:
  - → Histopathology
  - Culture from biopsy sample (Not swab)

#### **INVESTIGATIONS**

- Histopathology: Granulomas with epithelioidhistiocytes & Langerhans - type giant cells
- FNAC: If indicated
- IGRA /NAAT/PCR: Not recommended for diagnosis

#### **CASE DEFINITION**

#### A) Confirmed case:

- M.tuberculosis complex identified by either culture or PCR or histology shows typical morphology
- > Full course of ATT which led to complete clearance of lesions

#### B) Probable case:

> Typical skin lesion with no positive features/investigation as mentioned above (A)

## **DIAGNOSTIC ALGORITHM**

#### Suspected TB case based on presence of clinical signs

- Ulcers/discharging sinuses over sites of LN, bones & joints
- · Persistent, asymptomatic raised reddish/reddish brown lesion of >6 months' which may show scarring at one end
- Persistent, warty skin lesion of >6 months'

### Skin biopsy\* for histopathology ± culture\*\* + CXR\*\*\* Refer to dermatologist with expertise in cutaneous TB Histology diagnostic and/or Histology equivocal Histology negative & culture +ve and culture -ve culture -ve Treat for cutaneous TB Do not Treat Positive >22 mm Negative <5 mm **Equivocal**

#### Strong clinical suspicion > Start ATT

\*FNAC can be done if facilities for skin biopsy are not available

\*\*PCR may also be done if facilities are available but cannot be recommended as a routine due to the low level of evidence regarding its usefulness in diagnosis of cutaneous tuberculosis

\*\*\* if CXR shows signs suggestive of respiratory TB, treat accordingly

## **MANAGEMENT**

#### **TREATMENT**

- As in Pulmonary TB. Extend if needed
- DR -TB to be kept in mind
- No role of steroids oral or topical in management of CT

#### **FOLLOW UP**

- 1st follow-up after 4-6 weeks; majority improves; to look for ADRs
- If no response after 8 weeks
- No response
   – alternate diagnosis/DR-TB; refer to higher centre

#### **ABBREVIATION**

**ATT:** Anti-Tubercular treatment **BCC:** Bacille Calmette Guerin vaccine

CNS: Central Nervous System CT: Cutaneous Tuberculosis

**CXR:** Chest X-ray

**DR-TB:** Drug resistant Tuberculosis **FNAC:** Fine needle aspiration cytology **GIT:** Gastro-intestinal tract

IGRA: Interferon Gamma Release assay LN: Lymph node

NAAT: Nucleic acid amplification test **NTEP:** National TB Elimination Programme NTM: Non-Tuberculous Mycobacterium PCR: Polymerase chain reaction test

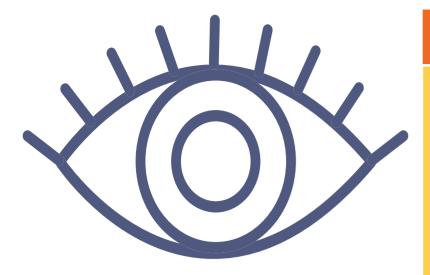
**TB:** Tuberculosis

#### REFERENCES

- 1. National TB Elimination Programme, Central TB Division. Training Modules for Programme Managers & Medical Officers. Ministry of Health & Family Welfare, Government of India https://tbcindia.gov.in/index1.php?lang=1&level=1&sublinkid=5465&lid=3540 Last access on 15 March, 2022.
- 2. Guidelines for programmatic management of drug resistant tuberculosis in India March 2021. National TB Elimination Programme, Central TB Division, Ministry of Health & Family Welfare, Government of India https://tbcindia.gov.in/showfile.php?lid=3590 Last access on 15 March, 2022.

## Standard Treatment Workflow (STW) for the Management of **INTRAOCULAR TUBERCULOSIS**

ICD-10-A18.3



# When to suspect

## **Ocular Symptoms**

- Blurred vision
- Redness
- Photophobia
- Pain in the eye
- Floaters
- Flashes of lights

#### Refer to Ophthalmologist for detailed examination

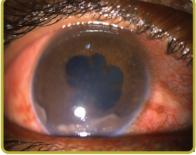
#### **Eye Care facility should have:**

Mandatory: Slit lamp, ophthalmoscope (direct or indirect), intraocular pressure assessment device

Preferred: Fundus camera, Fundus fluorescein angiongram(FFA), Optical Coherence Tomography (OCT)

#### Granulomatous anterior uveitis





# **Examination of the eyes**

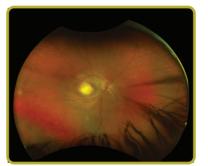
## **Clinical signs**

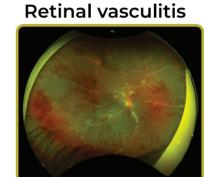
- Assess visual acuity
- · Anterior chamber cells, Keratic precipitates, Synechiae, Irregular pupil, RAPD
- Complicated cataract, high or very low intraocular pressure
- · Vitritis, Pars plana exudates, Retinal vasculitis, Retinitis, Choroiditis, Optic nerve head swelling

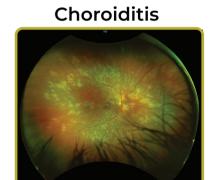
Intermeditate uveitis











**INVESTIGATIONS** 

**Essential:** CXR for healed/ active **pulmonary TB** 

Desirable: **Mantoux Test** (standardised tuberculin units): 10 mm induration considered positive

**Optional:** CT Chest (if available) for healed/active pulmonary TB

Imaging of eye: Ascertaining diagnosis, extent of disease & follow up, teleconsultation

Retinal photographs using fundus tomography camera

**Optical** coherence scans (if available)

Fluorescein angiograms (if available)

Investigations to rule out other causes of clinical presentation

## **MANAGEMENT**

#### **TREATMENT**

- ATT: 2 months of RHEZ + 7 months of RH depending on clinical response & side effects to treatment
- Add pyridoxine 10 mg/day
- · Corticosteroids: Topical steroids eye drops for severe/anterior chamber inflammation
- · For treatment in children refer to paediatrician
- Systemic corticosteroids for severe inflammation in consultation with Uveitis expert

#### REFERRAL TO HIGHER CENTRE

- Not confident to treat
- Vision threatening
- Non-response to treatment
- · Side effects due to treatment
- Atypical reaction

#### MONITORING AND FOLLOW UP

- · Frequency of follow up: 1-2 weeks in 1st month followed by monthly for 3 months & then 3 monthly
- · Eye: Clinical grading of inflammation using fundus photographs & OCT scans (if available)
- Steroids:

E: Ethambutol

H: Isoniazid

- Topical: Monitor IOP, cataract and any signs of bacterial/ fungal infection
- > Systemic steroids: Monitor body weight, blood sugar & blood pressure

#### **ABBREVIATIONS**

**ATT:** Antitubercular treatment

**IOP:** Intraocular pressure

**OCT:** Optical coherence tomography

R: Rifampicin

Z: Pyrazinamide

REFERENCES

**RAPD:** Relative Afferent Pupilary Defect

1. Agrawal R, et al.; Collaborative Ocular Tuberculosis Study Consensus Group. Collaborative Ocular Tuberculosis Study Consensus Guidelines on the Management of Tubercular Uveitis-Report 1: Guidelines for Initiating Antitubercular Therapy in Tubercular Choroiditis. Ophthalmology. 2021 Feb;128(2):266-276. doi: 10.1016/j.ophtha.2020.01.008. Epub 2020 Jan 11. PMID: 32115264. 2. Agrawal R, et al.; llaborative Ocular Tuberculosis Study Consensus Group, Collaborative Ocular Tuberculosis Study Consensus Guidelines on the Management of Tubercular Uveitis-Report 2: Guidelines for Initiating Antitubercular Therapy in Anterior Uveitis, Intermediate Uveitis, Panuveitis, and Retinal Vasculitis. Ophthalmology. 2021 Feb;128(2):277-287. doi: 10.1016/j.ophtha.2020.06.052. Epub 2020 Jun 27. PMID: 32603726.

## **GENITOURINARY TUBERCULOSIS**

**ICD-A18.10** 



#### PRESENTING SYMPTOMS

#### **Urinary Tuberculosis**

- · LUTS (frequency, urgency and nocturia) with dysuria and/or haematuria for at least 2 weeks
- Not responding to a 3–7 day course of antibiotics

#### **Male Genital Tuberculosis** (MGTB)

- Scrotal pain or swelling for 2 weeks or more
- Not responding to a 7-14 day course of antibiotics, or
- Discharging sinuses in the scrotum

Refer to higher

centre/Urologist

Rarely infertility

### **INVESTIGATION**



- (early morning, first void)
- NAAT assay (GeneXpert MTB/RIF)
- USG KUB\*\*
- USG Scrotum (for MGTB only)

\*Deranged renal function \*\*HDN/HDUN

#### **Specific Investigations**

- · If normal renal function
  - > IVP/CT urography
- If deranged Renal Function
  - MR Urogram
  - > Retrograde Pyelography
  - Nephrostogram

#### Optional (If other tests are inconclusive with high suspicion of GUTB)

- FNAC/ Biopsy- from accessible mass lesions or fluid collections
- Cystoscopic biopsy of Genitourinary tract

#### **TREATMENT**

TYPE OF TB	TYPE OF REGIMEN	DRUGS	EXTENSION CRITERIA
DRUG SUSCEPTIBLE TB	DS-TB REGIMEN	2 MONTHS H,R,E,Z 4 MONTHS H,R,E	Extension packets of infection, concurrent smear positive cavitary pulmonary disease, CNS involvement, Delay in positive cultures converting to negative Duration can be increased up to 9 to 12 months
MDR/RR OR XDR-TB	TREATMENT AS PER NTEP GUIDELINES		

## **FOLLOW UP**

At 8 weeks: Resolution of systemic symptoms, improved urinary symptoms, repeat culture if baseline culture positive

**After completion of ATT:** Repeat culture if baseline culture positive

Repeat imaging: If partial or impending ureteric stricture

#### Watch for the following complications at each Follow-up visit:

- Severe LUTS suggestive of small capacity bladder
- Deteriorating renal function

**CT**: Computed Tomography

**CBC**: Complete Blood Count

**CXR**: Chest X- Ray

**DJS**: Double J Stent

## **ABBREVIATIONS**

**ESR**: Erythrocyte Sedimentation Rate **MDR**: Multi Drug Resistant **ATT**: Anti-tubercular treatment

> H: Isoniazid **MTB**: Mycobaterium Tuberculosis

**NAAT**: Nucleic Acid Amplification Test **HDUN**: Hydroureteronephrosis

**IVP**: Intravenous Pyelogram **NTEP**: National Tuberculosis Elimination Programme **Z**: Pyrazinamide

**DS-TB**: Drug Susceptible Tuberculosis **LFT**: Liver Function Test **RFT**: Renal Function Test

E: Ethambutol **LUTS**: Lower Urinary Tract Symptoms **R**: Rifampicin

**HDN**: Hydronephrosis

**URINE AFB:** Urine for Acid-fast Bacillus **MR**: Magnetic Resonance

**RR**: Rifampicin Resistant

**XDR**: Extensively Drug Resistant

**USG KUB**: Ultrasonography Kidney, Ureter and Bladder

#### References

- 1. National TB Elimination Programme, Central TB Division. Training modules for Programme Managers & Medical Officers. Ministry of Health & Family Welfare, Government of India https://tbcindia.gov.in/index1.php?lang=1&level=1&sublinkid=5465&lid=3540 Last access on 11 March, 2022.
- 2. Guidelines for programmatic management of drug resistant tuberculosis in India March 2021. National TB Elimination Programme, Central TB Division, Ministry of Health & Family Welfare, Government of India. https://tbcindia.gov.in/showfile.php?lid=3590 Last access on 11 March, 2022.
- 3. Partin, A. W. et al. Edited (2020). Campbell Walsh Wein Urology (12th ed.). Elsevier.

## Standard Treatment Workflow (STW) for the Management of **FEMALE GENITAL TUBERCULOSIS**

ICD-10-A18.17



#### **SUSPECT**

#### **Consider following symptoms** in history:

- H/O infertility (primary or secondary)
- · Chronic lower abdominal or pelvic pain
- · Amenorrhoea or other menstrual disturbances
- Abnormal vaginal discharge
- Constitutional symptoms of TB (low grade fever, weight loss etc.)
- Other symptoms related to extra-genital TB (abdominal, CNS, bone and lymph nodes etc.)

In addition, standard investigations for TB to be carried out

#### **EXAMINE AND INVESTIGATE**

#### **Clinical Examination**

- General Physical Examination
- Pelvic Examination (cervical growth, uterine) size and mobility, adnexal tenderness & mass)

#### **Abdominal and Pelvic USG (TVS)**

- · Uterus, adnexa & pelvis to be evaluated preferably by transvaginal scan
- Endometrial cavity & vascularity to be looked carefully with colour Doppler

#### **Specific Investigations**

- · Endometrial sampling or biopsy with Pipelle device or Karman cannula (4 mm) for microbiological & histopathological examination
- Endoscopy:
  - › Hysteroscopy & laparoscopy to evaluate uterus, adnexa & other pelvic organs along with lower abdomen
  - > Laparoscopic biopsy from peritoneum or abdominal/pelvic lesions
- MTB diagnosis from biopsy specimen (endometrium & other tissues) by
  - > Smear microscopy (AFB smear) & culture
  - Gene Xpert or other NAAT
  - > HPE of biopsy specimen

## **DIAGNOSIS**

#### SUGGESTIVE FINDINGS IN FGTB **Imaging and Radiological**

- · HSG: to be avoided in acute phase Findings: blocked fallopian tubes, usually cornual; tobacco pouch appearance of the tubes; beaded tubes; filling defect in the uterine cavity (Asherman syndrome)
- USG: cogwheel appearance of tubes; uterine cavity may show thin diffuse endometrium with irregular borders
- · CT/MRI: can be used for tubo-ovarian mass

#### **Endoscopy**

- Hysteroscopy: To look for tubercles, pale endometrium & endometrial adhesions
- · Laparoscopy: Direct visualization of tubercle like lesions on the uterus, tubes and other pelvic organs including peritoneum, & caseous nodules

# **FEMALE GENITAL TB (STEPWISE DECISION)**

- Clinical history
- · General physical and pelvic examination
- Pelvic ultrasound
- HSP as indicated in infertility HSG

#### Definite FGTB needing ATT if any of the following tests are positive

- AFB microscopy positive
- AFB culture positive
- · Gene Xpert or other NAAT +ve
- · Histopathological demonstration of epithelioid granuloma

#### Probable FGTB needing ATT if any of following positive

- · Clinical findings/suspicion of TB with tubo-ovarian masses on imaging studies
- · Clinical findings/suspicion of TB with laparoscopic findings of beaded tubes, caseous nodules, tubercles, adhesions, hydrosalpinx & pyosalpinx etc.
- Clinical findings/suspicion of TB with hysteroscopic findings of tubercles, caseous nodules, pale endometrium, intrauterine adhesions etc.

#### **Negative FGTB: No ATT**

 No microbiological, histological, radiological, laparoscopic & hysteroscopic evidence of FGTB

Menstrual blood should not be used for NAAT.

## **MANAGEMENT**

#### **TREATMENT**

- Treatment of FGTB should be as per NTEP
- Patients requiring specific treatment such as infertility, Asherman syndrome & tubo-ovarian mass etc. should be referred to higher centres

#### **FOLLOW UP**

#### Follow-up of the patient should be flexible depending on the clinical presentation and response to ATT

- 1 month: Clinical Evaluation (General & Gynaecological)
- · 3 months: Clinical Evaluation (General & Gynaecological)
- · 6 months: Clinical Evaluation & Investigations (endometrial biopsy, hystero-laparoscopy & USG as needed)

## **ABBREVIATION**

**AFB**: Acid-Fast Bacilli **ATT**: Anti-Tuberculosis Therapy **CNS**: Central Nervous System **CT:** Computed Tomography

**FGTB**: Female Genital TB

**FNAC**: Fine-needle Aspiration Cytology

**HSE:** Histopathology Examination

**HSG:** Hysterosalpingography

**MRI** - Magnetic Resonance Imaging MTB: Mycobacterium Tuberculosis **NAAT**: Nucleic Acid Amplification Test **PCR**: Polymerase Chain Reaction

**TB**: Tuberculosis TVS: Transvaginal Scan

**NTEP**: National Tuberculosis Elimination **USG**: Ultrasonography **Programme** 

## REFERENCES

- 1. National TB Elimination Programme, Central TB Division. Training Modules for Programme Managers & Medical Officers. Ministry of Health & Family Welfare, Government of India. https://tbcindia.gov.in/index1.php?lang=1&level=1&sublinkid=5465&lid=3540 Last access on 08 March, 2022.
- 2. Guidelines for programmatic management of drug resistant tuberculosis in India March 2021. National TB Elimination Programme, Central TB Division, Ministry of Health & Family Welfare, Government of India accessed at https://tbcindia.gov.in/showfile.php?lid=3590 Last access on 08 March, 2022.
- 3. Sharma JB, Sharma E, Sharma S, Dharmendra S. Recent advances in diagnosis and management of female genital tuberculosis. J Obstet Gynaecol India. 2021;71:1-12.

This STW has been prepared by national experts of India with feasibility considerations for various levels of healthcare system in the country. These broad guidelines are advisory, and are based on expert opinions and available scientific evidence. There may be variations in the management of an individual patient based on his/her specific condition, as decided by the treating physician. There will be no indemnity for direct or indirect consequences. Kindly visit our web portal (stw.icmr.org.in) for more information.

© Indian Council of Medical Research and Department of Health Research, Ministry of Health & Family Welfare, Government of India.

#### **Standard Treatment Workflow (STW) of**

#### MICROBIOLOGICAL WORK-UP FOR ADULT EXTRAPULMONARY TUBERCULOSIS

#### LOGISTICS INVOLVED IN SAMPLE COLLECTION AND TRANSPORTATION

- Collect Samples for Microbiological work-up in sterile containers before treatment is started. (Mention date & time of collection)
- · Specimens to be sent in sterile saline (NOT in formalin)
- Establish linkages between peripheral centres, District centres and Tertiary centre/medical colleges/ IRL. Specify details of person to be contacted, department and contact number during referrals
- · Transportation at 2-8 °C
- Maximum time for transportation in cold chain should be 5 days from time of collection
- Quantity of sample mentioned is only for microbiological work-up. Tests like histopathology, cytology, ADA, glucose, protein, etc will require additional sample
- Microbiological tests for TB (smear, molecular tests, culture) will be performed as per availability and preparedness of site
- PHC and CHC should perform smear microscopy and molecular diagnostic tests. If sample less than 500 µl, refer directly to Tertiary centre/medical colleges/IRL for culture. Residual sample in the needle and syringe used to collect the specimen can be used for smear
- MGIT to be used for culture. However, if MGIT is not available, LJ medium should be used

#### **REJECTION OF SAMPLES**

- Unlabelled samples (All specimens MUST be labelled & have a unique patient identifier)
- · Have no collection date indicated
- Insufficient quantity No specimen in container
- Damaged Specimen leaked or broken in transit
- Samples greater than 3 days old at room temperature and more than 5 days in refrigeration are unreliable specimens for testing

Precious samples should be transported to IRL.

Diagnostic algorithm of NTEP to be followed in the Microbiology labs

#### MICROBIOLOGICAL GUIDANCE FOR COMMON TYPES OF EXTRAPULMONARY TUBERCULOSIS

# OSTEOARTICULAR/ MUSCULOSKELETAL

- · Sample: Tissue, pus, synovial fluid
- Sample amount: Biopsy: Specimen material 1 cm x 1 cm biopsies. Any caseous area should be sampled. Add 0.5-2 ml sterile saline to biopsy depending on its size to avoid drying of tissue specimen
- · Optimum fluid/pus: 2-3ml.
- Swabs are sub-optimal samples

PLEURAL

- · Sample: Pleural fluid
- · Sample amount: 10-15 ml

MENINGITIS

- · Sample: CSF:
- · Sample amount: 3-5 ml

YMPHADENITIS

- · Sample: FNA/ Biopsy
- Sample amount: Specimen material 1 cm x 1 cm biopsy. Add 0.5-2 ml sterile saline to biopsy depending on its size to avoid drying of tissue specimen
- · Optimum FNA sample: 2 ml

UROGENITAL

- · Sample: urine
- Sample amount: Entire early morning urine sample (3-5 days)

FEMALE GENITAL

- · Sample: Endometrial curettage/biopsy
- Sample amount: Biopsy: Specimen material 1cm x 1 cm biopsies. Any caseous area should be sampled. Add 0.5-2 ml sterile saline to biopsy depending on its size to avoid drying of tissue specimen

GASTROINTESTINAL

- · Sample: Tissue, pus, peritoneal fluid
- Sample amount: Biopsy: Specimen material 1 cm X 1 cm biopsy (Atleast 6 biopsies for microbiological diagnosis including any caseous area). Any caseous area should be sampled. Add 0.5-2 ml sterile saline to biopsy depending on its size to avoid drying of tissue specimen
- · Optimum fluid/pus: 5-10ml

#### **Processing:**

- Preferably immediately. If not possible- store/transport at 2-8 °C
- If sample is adequate, attempt molecular testing at that site
- If biopsy is not possible or at an inaccessible site, refer patient to the next higher centre immediately where appropriate test can be done
- If sample obtained at a centre is inadequate, send directly to nearest Tertiary centre/medical colleges/IRL

# Microbiological procedures:

- AFB Smear Microscopy except in GITB
- NAAT
- Culture (MGIT. If MGIT is not available LJ medium should be used)
- Drug susceptibility testing, if culture is positive

### **ABBREVIATIONS**

ADA: Adenosine Deaminase AFB: Acid fast bacilli CHC: Community Health Centre FNA: Fine needle aspirate
LJ medium: Lowenstein Jensen medium

MGIT: Mycobacteria Growth Indicator tube (Liquid culture medium for mycobacteria) IRL: Intermediate Reference NAAT: Nucleic Acid Amplification Tests-Xpert MTB/RIF/TrueNat laboratory

PHC: Primary health Centre TB: Tuberculosis IRL: Intermediate Reference laboratory

#### REFERENCES

- 1. National TB Elimination Programme, Central TB Division. Training Modules for programme managers & Medical Officers. Ministry of Health & Family Welfare, Government of India.
- https://tbcindia.gov.in/index1.php?lang=1&level=1&sublinkid=5465&lid=3540 Last access on 15 March, 2022.

  2. Guidelines for programmatic management of drug resistant tuberculosis in India March 2021. National TB Elimination Programme, Central TB Division, Ministry of Health & Family Welfare, Government of India accessed at https://tbcindia.gov.in/showfile.php?lid=3590 Last access on 15 March, 2022.

## Standard Treatment Workflow (STW) for the Management of **ANTITUBERCULAR THERAPY RELATED HEPATITIS**

#### PATIENT TO BE STARTED ON ATT

#### **Risk factors for ATT Hepatitis**

- History of underlying liver disease (jaundice, ascites, GI bleeding)
- Physical findings suggestive of liver disease (Splenomegaly, ascites, icterus, edema)
- Alcoholism
- · Hypoalbuminemia and Malnutrition
- · Elevated aminotransferases at baseline
- ·HIV
- · IV drug abuse
- Elderly age

Evaluate for underlying liver disease HBsAq, Anti-HCV, Ultrasound

**Chronic Liver** disease +

· TB meningitis or CNS TB

- Intensive education & counselling
- Modified ATT may be needed based on Child Pugh Status
- LFT monitoring

Pericardial TB

· Ocular TB

Any form that is lif

Intestinal TB with

Joint or Spinal TB

intestinal obstruction

threatening, eg.,

· Start ATT

No CLD or

Cirrhosis

 Counsel about symptoms of ATT Hepatitis

No

## REINTRODUCTION OF ATT HEPATOXIC DRUGS

alternative causes of liver disease

Refer to higher center immediately

week & reintroduce (see later)

**START ATT** 

Diagnosis of ATT hepatitis Clinical symptoms

fatigue, yellowing of sclera, altered sensorium)

AST/ALT increased to 3 times of baseline/ULN

AST/ALT increased to 5 times of baseline/ULN

**Exclude viral hepatitis (HBsAg,** 

Get PT/INR, Ultrasound liver

Need urgent ATT: Change to non-hepatotoxic

Non-resolution of LFT abnormalities: exclude

Jaundice and coagulopathy/encephalopathy

drugs (Fluroquinolones, ethambutol &

• No need for urgent ATT: repeat LFT after a

Anti-HCV, IgM- antiHAV, IgM-AntiHEV,

· Jaundice (Bilirubin 2 ULN)

No clinical symptoms

present (abdominal pain, vomiting, unexplained

- Reintroduce only if ALT and AST < 2 ULN & normal</li> bilirubin
- · Start one drug at time: helps identify the culprit

Stop all hepatotoxic drugs

aminoglycosides)

- Rifampicin may be introduced at 10 mg/kg dose
- After one week add Isoniazid 5 mg/kg if LFT normal
- · After one week add pyrazinamide 25 mg/kg if LFT is normal
- If ATT hepatitis severe (liver failure, coagulopathy or altered sensorium): Pyrazinamide reintroduction may be avoided
- · Another approach could be low dose of one drug followed by full dose after three days
- Duration of ATT: count only when full ATT is started

#### No need for Urgent **Urgent ATT: life or organ** threatening **ATT** · Sputum + Pulmonary TB

- Sputum –ve Pulmonary
- TB lymphadenitis
- · Tubercular pleural effusion
- Tubercular ascites
- · Intestinal TB
- Genitourinary TB
- Bone TB

## REINTRODUCTION OF ATT: IF AST AND ALT < 2 ULN

#### **SEQUENTIAL**

Initiate one at a time Rifampicin 10 mg/kg

1 week: repeat LFT

1 week: repeat LFT

Initiate Pyrazinamide 25 mg/kg

Initiate Isoniazid 5 mg/kg

#### **INCREMENTAL**

Initiate Rifampin 150 mg/day Gradually increase dose by day 4

Initiate Isoniazid 100 mg/day at day 8 **Gradually increase dose by day 11** 

Initiate Pyrazinamide 500 mg/day on day 15 Gradually increase dose by day 18

## **CHILD PUGH (CTP) SCORE**

	Score 1	Score 2	Score 3
Bilirubin	< 2 mg/dl	2-3 mg/dl	>3 mg/dl
Albumin	>3.5 gm/dl	2.8-3.5 gm/dl	<2.8 gm/dl
INR	<1.7	1.7-2.2	>2.2
Ascites	Absent	Slight	Moderate
Encephalopathy	Absent	Grade 1-2	Grade 3-4

## **HEPATIC ENCEPHALOPATHY GRADE**

- Grade 0: normal consciousness, personality & neurological examination
- · Grade 1: restless, disturbances in sleep, irritability or agitated, tremors, handwriting affected
- · Grade 2: lethargy, disorientation to time, asterixis, ataxia
- · Grade 3: somnolent & stuporous, disoriented to place, hyperactive reflexes, rigidity
- Grade 4: unrousable coma, decerebrate

ATT SELECTION FOR UNDERLYING LIVER DISEASE

Child Status	Suggested ATT
Child A Cirrhosis (Score 1-6) Stable Liver disease	9 months of therapy with HRE <b>OR</b> 2 months of therapy with HRE followed by 7 months of HR
Child B Cirrhosis (Score 7-10) Advanced Liver Disease	One hepatotoxic drug regimen can be used: Two months of therapy with INH (or) RIF with ETH & aminoglycoside, followed by 10 months of therapy with INH/RIF & ETH
Child C Cirrhosis (Score 11-15) Very advanced liver disease	No hepatotoxic drug 18 to 24 months treatment using a combination of ETH, FQL, cycloserine & aminoglycoside/ capreomycin
In Acute hepatitis	Avoid hepatotoxic drugs ATT with non-hepatotoxic drugs if urgent ATT required Wait till improvement in liver function if no urgent need of ATT

#### **ABBREVIATIONS**

**ALT:** Alanine transaminase **AST:** Aspartate transaminase **ETH:** Ethambutol

**FQL:** Fluoroguinolone

**GI:** gastro-intestinal **HAV:** Hepatitis A virus

ATT: Anti-tubercular treatment HBsAg: Hepatitis B surface Antigen INH: Isoniazid **HCV:** Hepatitis C virus

**HEV:** Hepatitis E virus

HRE: Isoniazid, Rifampicin, Pyrazinamide LFT: Liver function tests IgM: Immunoglobulin M

**INR:** International normalized ratio IV: Intravenous

PT: Prothrombin time **RIF:** Rifampicin **TB:** Tuberculosis

**ULN:** Upper limit of normal

#### REFERENCES

- 1. National TB Elimination Programme, Central TB Division. Training modules for programme managers & Medical officers. Ministry of Health and Family Welfare, Government of India accessed at https://tbcindia.gov.in/index1.php?lang=1&level=1&sublinkid=5465&lid=3540 on 24 February, 2022.
- 2. Guidelines for programmatic management of drug resistant tuberculosis in India March 2021. National TB elimination programme, Central TB division, Ministry of Health and Family Welfare, Government of India accessed at https://tbcindia.gov.in/showfile.php?lid=3590 on 10 February, 2022.
- 3. Satsangi S., Randev S., Taneja S. (2022) Antitubercular Therapy-Related Hepatitis. In: Sharma V. (eds) Tuberculosis of the Gastrointestinal system. Springer, Singapore. https://doi.org/10.1007/978-981-16-9053-2\_23

## **Standard Treatment Workflow (STW) Guidelines for** DRUG SENSITIVE-TB TREATMENT AS PER NTEP

- · For all TB patients whether being treated in public or private sector, clinicians should follow Standards for TB care in India guidelines
- · In NTEP, the principle of TB treatment (except confirmed DR-TB) is to administer daily FDC of 1st line ATT in appropriate weight bands, under direct observation
- · For patients being treated in private sector, FDCs may be provided by NTEP whenever requested

#### Regimen for Drug-Sensitive TB cases: 2HRZE/4HRE

- · This regimen is for H & R sensitive TB cases and cases where the sensitivity pattern can not be established
- Treatment is given in two phases:
- 1. Intensive phase consists of 8 weeks (56 doses) of isoniazid (H), rifampicin (R), pyrazinamide (Z) and ethambutol (E) given under direct observation in daily dosages as per weight band categories
- 2. Continuation phase consists of 16 weeks (112 doses) of isoniazid, rifampicin and ethambutol in daily dosages. Only pyrazinamide will be stopped in the continuation phase. The CP needs to be extended upto 24 weeks in certain forms of TB like CNS TB, Skeletal TB. In disseminated TB or slow response treating physician may extend on case to case basis.

Regimen for DS-TB	IP	СР
Drugs	2 HRZE	4 HRE
Doses	56	112

## **ADULT TB TREATMENT**

#### **Drug dosages for first-line anti- TB drugs Drugs Doses** Isoniazid (H) 5 mg/kg daily (4 to 6 mg/kg) Rifampicin (R) 10 mg/kg daily (8 to 12 mg/kg) 25 mg/kg daily (20 to 30 mg/kg) Pyrazinamide (Z) Ethambutol (E) 15 mg/kg daily (12 to 18 mg/kg) 15 mg/kg daily (15 to 20 mg/kg) Streptomycin (S)\*

\*Streptomycin is administered only in certain situations, like TB meningitis or if any first line drug need to be replaced due to ADR as per weight of the patient

Pyridoxine may be given at a dosage of 10 mg per day

	Number of tablets (FDCs)			
Weight category	Intensive Phase H: 75mg; R: 150 mg; Z: 400 mg; E: 275 mg)	Continuation Phase H: 75mg; R: 150 mg; E: 275 mg)		
25 to 34 kg	2	2		
35 to 49 kg	3	3		
50 to 64 kg	4	4		
65 to 75 kg	5	5		
> 75 kg	6	6		
E: 15 0 1: :: (EDO.) (				

- Fixed Dose Combinations (FDCs) refer to products containing two or more active ingredients in fixed doses, used for a particular indication(s)
- · In NTEP, for Adults: 4-FDC (given in IP) consists of HRZE and 3-FDC (given in CP) consists of HRE
- During treatment if weight of the patient increases by > 5 kg and crosses the next weight band then patient should be given the next higher weight band FDC drugs

#### **Special considerations for Adult TB** Meningitis

- · Intensive Phase: 2 months of RHZE or **RHZS**
- Continuation phase: 3 drugs-RHE for at least 10 months\*
- STEROIDS
  - > Preferably Dexamethasone 0.4 mg/kg/day intravenously in 3-4 divided doses during hospital stay
  - > If not feasible, give oral Dexamethasone 0.4 mg/kg/day in divided doses or oral Prednisolone 1 mg/kg/day in a single morning dose
  - Discharge on oral steroids on tapering doses for total duration of 8-12 weeks
  - > Regular follow up is essential every month for at least first 3 months & can be increased thereafter till treatment is stopped
- Monitor liver function tests & any other features of drug toxicity
- Observe for clinical improvement or any deterioration
- > Closely observe for development of any complications
- \*treatment duration may be increased in

#### **Special considerations for Adult** abdominal TB

- · Extend duration of treatment in cases of inadequate response
- · Refer for surgical management for complications [intestinal obstruction (due to strictures), perforation]
- · Consider endoscopic dilatation for treatment for accessible strictures
- · Refer for biliary drainage in case of Jaundice due to biliary obstruction (hepatobiliary obstruction/pancreatic TB)

#### **Special considerations for** intra-ocular TB

- ATT: 2 months of RHEZ + 7 months of RH depending on clinical response & side effects to treatment
- Add pyridoxine 10 mg/day
- · Corticosteroids: Topical steroids eye drops for severe/anterior chamber inflammation
- · For treatment in children refer to paediatrician
- · Systemic corticosteroids for severe inflammation in consultation with **Uveitits** expert

## PAEDIATRIC TB TREATMENT

- Paediatric cases are to be treated under NTEP in daily dosages as per 6 weight band categories
- Children & adolescents up to 18 years of age weighing less than 39 kg, are to be treated using paediatric weight bands. Those weighing more than 39 kg to be treated with adult weight bands.
- Available paediatric dispersible FDCs and loose drugs

Drug dosages for first-line anti- TB drugs

7-15 mg/kg

10-20 mg/kg

15-25 mg/kg

- 1. Dispersible FDC, flavoured
  - · Rifampicin 75 mg + Isoniazid 50 mg + Pyrazinamide 150 mg
  - · Rifampicin 75 mg + Isoniazid 50 mg
- 2. Dispersible Loose drugs
  - Ethambutol 100 mg
  - · Isoniazid 100 mg

Isoniazid (H)

Rifampicin (R)

Ethambutol (E)

	Number of tablets (dispersible FDCs)				
Weight	Intensive phase		Continuation phase		
Band	HRZ	Е	HR	E	
	50/75/150	100	50/75	100	
4-7 kg	1	1	1	1	
8-11 kg	2	2	2	2	
12 -15 kg	3	3	3	3	
16 -24 kg	4	4	4	4	
25 - 29 kg	3 + 1A *	3	3 + 1A *	3	
30-39 kg	2 + 2A *	2	2 + 2A *	2	

- \*A=Adult FDC (HRZE = 75/150/400/275; HRE = 75/150/275). It is added in higher weight band categories i.e. > 25 kg as these children may be able to swallow tablets
- Pyridoxine may be given at a dosage of 10 mg per day

Special considerations for paediatric TB

meningitis

→ 2 HRZE and 10 HRE (in appropriate doses)

> Prednisolone 2 mg/kg/day for 4 weeks &

Slower taper needed in some patients

ATT for **paediatric** TB Meningitis

then taper over 4 weeks\*

be used either injectable/oral

Corticosteroids

#### **Special considerations for** paediatric Abdominal TB

Special considerations for

paediatric osteoarticular TB

Follow up every month during treatment & subsequently every

with X-ray or MRI & Tubercular

dactylitis or arthritis with plain

Regimen: 2HRZE + 10HRE

3 months: Potts spine

Steroids- Not recommended Supportive treatment-Management of SAM/Malnutrition as per national guidelines Surgical treatment:

- > Acute intestinal obstruction, Bowel perforation
- > Persistence of obstructive symptoms despite conservative management & ATT
- DO NOT start Empirical ATT with isolated:
- → Recurrent/Chronic abdominal pain without danger signs
- > Chronic diarrhoea without proper evaluation

## **ABBREVIATIONS**

\*Equivalent dose of another steroid formulation may

**ADR:** Adverse drug reaction **ATT:** Anti-Tubercular treatment CNS: Central Nervous system

**CP:** Continuation phase

Pyrazinamide (Z) 30-40 mg/kg

**DS-TB:** Drug sensitive Tuberculosis **IP:** Intensive phase

FDC: Fixed dose combination

NTEP: National TB Elimination Programme

R: Rifampicin **S:** Streptomycin **SAM:** Severe acute malnutrition Z: Pyrazinamide

DR-TB: Drug resistant Tuberculosis H: Isoniazid E: Ethambutol

(maximum dose 300 mg/day)

(maximum dose 600 mg/day)

(maximum 2000 mg/day)

(maximum 1500 mg/day)

MRI: Magnetic Resonance imaging TB: Tuberculosis

## REFERENCES

- 1. National TB Elimination Programme, Central TB Division. Training modules for programme managers & Medical officers. Ministry of Health and Family Welfare, Government of India accessed at https://tbcindia.gov.in/index1.php?lang=1&level=1&sublinkid=5465&lid=3540 on 24 February, 2022.
- This STW has been prepared by national experts of India with feasibility considerations for various levels of healthcare system in the country. These broad guidelines are advisory, and are based on expert opinions and available scientific evidence. There may be variations in the management of an individual patient based on his/her specific condition, as decided by the treating physician. There will be no indemnity for direct or indirect consequences. Kindly visit our web portal (stw.icmr.org.in) for more information. © Indian Council of Medical Research and Department of Health Research, Ministry of Health & Family Welfare, Government of India.

## Standard Treatment Workflow (STW) for the Management of PAEDIATRIC ABDOMINAL TUBERCULOSIS

ICD-10-A18.31

#### WHEN TO SUSPECT?

Presence of Fever >2 weeks

Anorexia

ESSENTIAL

Ultrasound abdomen

measuring >15 mm in

short axis, conglomerate

and/or central necrosis

thickening >15 mm with

increased echogenicity

Omental/mesenteric

**SPECIFIC FINDINGS** 

· Abdominal LN:

Ileocaecal wall

thickening

· Chest X Ray

culture

- Unexplained weight loss\* or no weight gain in last 3 months
- · History of contact with TB patient
- · With one or more of following
  - > Recurrent/chronic abdominal pain in presence of red flag signs
  - Abdominal distension/mass
  - > Altered bowel habits

\*Constitutional symptoms may or may not be present in suspected case of Abdominal TB



**NON SPECIFIC FINDINGS** 

Inter-loop ascites

Ascitic fluid with

SAD <15 mm in

signs

sputum/GA/IS (If CXR abnormal) for NAAT, culture

Ascitic fluid (If present) for cytology, protein & albumin

Peripheral LN-FNA (If size >2 cm) for cytology, NAAT,

Intraabdominal fluid

(free or loculated) or

multiple septations

absence of red flag

Abdominal LAP with

#### **CLINICAL FEATURES SPECIFIC TO TYPE OF ABDOMINAL TB**

#### **PERITONEAL TB**

- · Abdominal pain, distension
- Fever
- Weight loss

#### **NODAL TB**

- Pain abdomen
- Fever
- · Palpable abdominal lump

#### **INTESTINAL TB**

- Recurrent intestinal colic
- Altered bowel habits
- · Chronic diarrhoea
- Partial/complete intestinal obstruction
- Weight loss, anorexia
- Palpable abdominal lump
- Lower gastrointestinal bleeding

#### **VISCERAL TB (LIVER, SPLEEN, PANCREAS)**

- Abdominal pain
- Fever
- Jaundice
- Weight loss
- Anorexia
- Hepatomegaly
- Splenomegaly Hepatic abscess
- Palpable
- abdominal lump
- Abnormal LFTs

#### **EXAMINATION FINDINGS**

- Anthropometry
- General physical & systemic examination
- Look for peripheral LAP. ascites. hepatosplenomegaly, doughy feel of abdomen. palpable abdominal lump, visible peristalsis

#### **RED FLAGS**

- Pain abdomen waking child from sleep
- Chronic, severe, or
- nocturnal diarrhea Gastrointestinal blood
- Localized distension or mass

## INVESTIGATION

#### Ascites

#### **Essential**

- · If exudative ascites, ascitic fluid for NAAT, culture
- · No role of ADA

#### **Enlarged Abdominal LN** (SAD>15 mm)

#### Desirable

· USG guided Abdominal LN-FNA for cytology, NAAT, culture

#### **Optional**

 USG/CT guided biopsy of LN for histology, NAAT, culture

## **Intestinal involvement**

#### **Desirable**

- If Ileocaecal thickening with enlarged nearby LN
- · USG guided Abdominal LN-FNA for cytology, NAAT, culture

· Isolated ileocaecal thickening/LN sampling

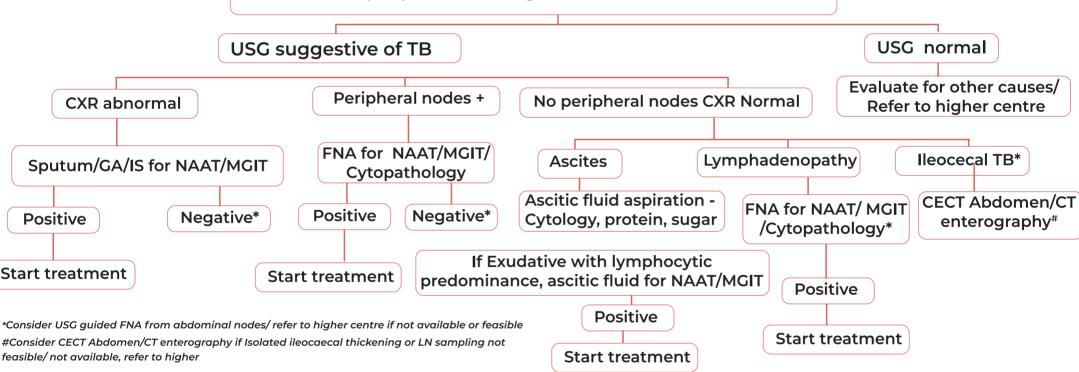
- not feasible
- · CECT Abdomen/CT enterography

## **Optional**

- · Ileocolonoscopy, tissue biopsy (HPE, NAAT, culture)
- Laparoscopy, tissue biopsy for HPE, NAAT, culture

#### **DIAGNOSIS ALGORITHM**

# If abdominal TB suspected Look for peripheral nodes, get CXR and Abdominal USG



## TREATMENT

### Start treatment & follow-up as per NTEP

- ATT for 6 months (2HRZE + 4HRE)
- Pyridoxine supplementation- 10 mg/day
- Steroids- not useful & not recommended
- Supportive treatment- Management of SAM/Malnutrition as per national guidelines
- Surgical treatment:
- Acute intestinal obstruction, Bowel perforation > Persistence of obstructive symptoms
- despite conservative management & ATT • DO NOT start Empirical ATT with isolated:
- > Recurrent/Chronic abdominal pain without red flag signs
- Chronic diarrhoea without proper evaluation

## **MANAGEMENT**

#### MONITORING

- Assessment of response to treatment:
- > Clinical follow up every month during treatment & after that every 3 months
- > Radiologically by USG At the end of treatment or if worsening or non response to treatment
- > Microbiological If worsening or non response to treatment
- Pointers towards DR TB investigation:
- > Non response to treatment or Worsening or deterioration of constitutional symptoms after initial improvement
- Obstructive symptoms may persist or worsen despite treatment with ATT
- **Monitor for**
- Adherence to treatment (ATT)
- Adverse drug reactions- ATT induced

#### WHEN TO REFER?

- Diagnosis is uncertain & additional investigations are required
- Acute intestinal obstruction or bowel perforation
- DR TB
- No response to appropriate treatment
- Oral drug (ATT) intolerance/cannot be given

ATT- Antitubercular treatment **CECT- Contrast Enhanced Computed Tomography** 

CT- Computed Tomography CXR- Chest X-Ray **DR-TB-** Drug Resistant tuberculosis E- Ethambutol FNA- Fine Needle Aspiration **GA-** Gastric Aspirate

H- Isoniazid HPE- Histopathological Examination LAP: Lymphadenopathy LN- Lymph Node MGIT- Mycobacteria Growth Indicator Tube

NAAT- Nucleic Acid Amplification Test

SAD-Sagittal Abdominal Diameter

SAM- Severe Acute Malnutrition USG- Ultrasonography

Z- Pyrazinamide

R- Rifampicin

- 1. National TB Elimination Programme, Central TB Division. Training Modules for Programme Managers & Medical Officers. Ministry of Health & Family Welfare, Government of India. https://tbcindia.gov.in/index1.php?lang=1&level=1&sublinkid=5465&lid=3540 Last access on 10 March, 2022.
- 2. Guidelines for programmatic management of drug resistant tuberculosis in India March 2021. National TB Elimination Programme, Central TB Division, Ministry of Health & Family Welfare, Government of India accessed at https://tbcindia.gov.in/showfile.php?lid=3590 Last access on 10 March, 2022.

## PAEDIATRIC INTRATHORACIC TUBERCULOSIS

(PULMONARY, PLEURAL, MEDIASTINAL)

**ICD-10-A15** 

Documented, persistent unexplained fever for 2 weeks or more Unremitting cough for 2 weeks or more Unexplained documented weight loss of  $\geq$  5% in last 3 months WHEN TO

No weight gain despite adequate

nutrition

Unexplained loss of appetite Chest pain (specific to

pleural effusion)

Contact with TB patients in past

• Temperature, Weight, Mid Arm Circumference (MAC), Lymphadenopathy, cold abscess, discharging sinus

**EXAMINATION** 

· Chest examination findings depend upon underlying pathology like consolidation, pleural effusion etc.

#### **INVESTIGATIONS**

#### **Essential**

- Chest x-ray
- TB suggestive: Hilar/paratracheal lymph nodes, fibrocavitory disease, Miliary pattern
- > Non Specific: effusion, consolidation, bronchopneumonia, other shadows etc.
- Sputum/Induced Sputum/Gastric Lavage/ Aspirate /pleural fluid for NAAT
- Smear examination (if NAAT unavailable)
- · If facilities exist, send aliquot of sample for culture, if NAAT negative for MTB
- · Pleural tap": Gross, Cytology, Biochemistry, NAAT, MGIT/LJ, ZN if NAAT not available \*\*If can't be done at primary level then refer

Do HIV testing for all cases with TB







# Desirable

Chest x-ray of family members

#### **Optional** (to be done in institutions) Pleural Biopsy (caseating granulomas with Langerhans

giant cells, epithelioid cells &

- lymphocytes) Image guided (USG/CT)
- mediastinal LN biopsy
- **CECT scan** Bronchoscopy & BAL

#### DO NOT DO

- TST/Mantoux test
- Overemphasized, supportive
- Not to diagnose TB or to start ATT on basis of +ve TST ONLY
- Serological tests- IGRA
- (Quantiferon/Quantiferon-Gold etc)
   Pleural fluid ADA

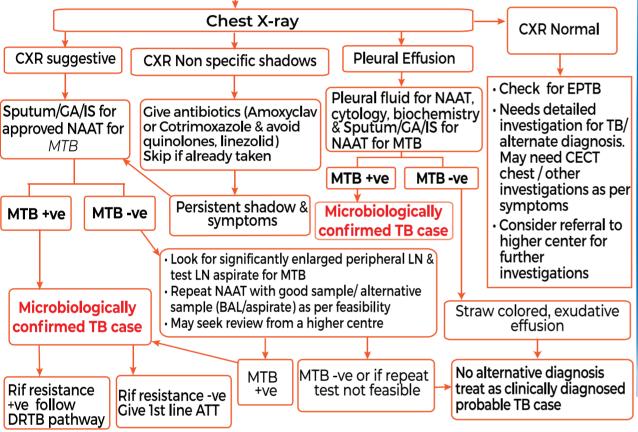
## DIAGNOSTIC ALGORITHM

#### ALGORITHM FOR PEDIATRIC INTRATHORACIC TB AMONG CHILDREN WITH NO RISK FACTORS FOR DRUG RESISTANCE

- Persistent Fever ≥2weeks, without a known cause and/or
- Unremitting cough for ≥2weeks and/or

SUSPECT

- Weight loss ≥5%; or no weight gain in past 3 months despite adequate nutrition; or failure of nutritional rehabilitation in babies with SAM
- With or without contact with patient with Pulmonary TB in past 2 years



#### PEDIATRIC TB FURTHER WORK-UP ALGORITHM UNDER U-DST **Presumed TB NAAT** cases MTB +ve MTB -ve Clinically diagnosed Rif resistance +ve\$ Rif resistance -ve TB case with no risk factors for DRTB FL-LPA **Repeat NAAT** SL - LPA\* **Treat with** Reinvestigate standard Rif resistance -ve for nonresponse regime 2HRZE/4RHE including for **DRTB** Add Pyridoxine FQ and/or SLI 10 mg/day Resistance No response in 4 weeks H Resistant (inhA and/or KatG mutation) **H** Resistant FL-LPA\*\* Eto resistance (inhA Not detected mutation)

\$RR detected in new case with no risk factors for DRTB needs retesting if only MTB detected is very low as it makes Rif resistance detection less reliable SLLPA may be done directly if smear +ve else send for MGIT followed by SLLPA or LC DST (Mfx 2.0, Km, Cm, Lzd)

\*\*LPA may be done directly if smear +ve else send for MCIT followed by FLPA to evaluate for H (inhA and/ or KatG mutn) and Eto (inhA) resistance

#### LK EVILWENT

WEIGHT

**BAND** 

4-7 kg

8-11 kg

12-15 kg

16-24 kg

25-29 kg

**Number of tablets (dispersible FDCs)** 

Continuation

HR

50/75

1

2

3

4

3 + 1A\*

 $2 + 2A^*$ 

Interruption up to 4 weeks

Resume therapy

(Restart if missed within 1st 4 weeks)

Rif resistance not detected

Retreat with 1st line drugs and

Check for INH resistance and treat

Ε

100

1

2

3

4

3

2

Intensive phase

HRZ

50/75/15

1

2

3

4

3 + 1A\*

Ε

100

1

2

3

4

3

2

TYPE OF	TB TREATMENT REGIMENS			
Microbiological Pulmonary TB	lly confirmed RS			
Clinically diagn	2HRZE + 4HRE			
Drug sensitive previously treated TB (recurrent, failure, treatment after default)				
*DR TB algorithm-DST				
ISONIAZID (H)	7-15 mg/kg (maximum dose 300mg/day)			

RIFAMPICIN (R) 10-20 mg/kg (maximum dose 600mg/day) **PYRAZINAMIDE (Z)** 30-40 mg/kg (maximum 2000mg/day) ETHAMBUTOL (E) 15-25 mg/kg (maximum 1500mg/day)

30-39 kg  $2 + 2A^*$ \*A=Adult FDC (HRZE = 75/150/400/275; HRE = 75/150/275) **MONITORING** 

 Then every month till completion of treatment What to assess

When to assess

- Appropriateness of therapy: Correct combination, acceptance/tolerance
- · Counsel about need to complete & not miss on doses (Inform, if doses are missed) Response to therapy:

15th day: For adherence, correct dose, combination & tolerance to drugs

- · Clinical (symptoms, adverse effects, weight, dose revision)
- · X-ray at end of therapy
- Do X-ray for worsening at any time OR slow resolution OR persistent symptoms at end of IP • NAAT is not appropriate follow up tool for monitoring progress of disease
- Smear examination at end of treatment (to declare outcome)
- · Repeat microbiological test (smear, MGIT, NAAT) at end of IP & at end of therapy, if still
- symptomatic or any deterioration/failure to respond
- After treatment completion, follow up patients clinically at end of 6, 12, 18 & 24 months

#### **ABBREVIATIONS**

**ADA**: Adenosine Deaminase BAL: Broncho-alveolar lavage **CBNAAT**: Cartidge-based Nucleic Acid Amplification test **CECT**: Contrast enhanced CT **CP**: Continuation phase

CT: Computed tomography

**DRTB**: Drug resistant TB **DST**: Drug sensitivity test **EPTB**: Extra-pulmonary TB **FTO**: Ethionamide **FDC**: Fixed dose combination FL-LPA: First line - Line probe assay FQ: Fluoroquinolones **GA**: Gastric aspirate H: Isoniazid

HIV: Human Immunodeficiency virus HRZE: Isoniazid; Rifampicin; Pyrazinamide; Ethambutol IGRA: Interferon Gamma Release assay

**IS**: Induced sputum **LN**: Lymph node MAC: Mid Arm Circumference MTB: Mycobacterium Tuberculosis **NAAT**: Nucleic acid amplification test

PPD: Purified Protein Derivative

**RIF**: Rifampicin **SAM**: Severe acute malnutrition SLI: Second line injectables **SL-LPA**: Second line - Line probe assay TST: Tuberculin skin test **USG**: Ultrasonography **ZN**: Ziehl Neelson

Consider steroids in miliary

· Prednisone dose 2 mg/kg

mg/kg/day for 4 weeks

daily or Dexamethasone 0.6

Reduce dose gradually over

Pyridoxine 10 mg/day for 6

Treat co-morbid conditions:

Interruption over 4 weeks

Reinvestigate for DRTB

Rif resistance detected

**Treat as MDRTB** 

Nutritional support

next 4 weeks before stopping

TB with hypoxia,

disseminated TB

months

HIV, SAM

MANAGING TREATMENT INTERRUPTIONS (NON-ADHERENCE)

In case interruption happens in CP & on retrieval, the patient has

Resons for interruption should always be evaluated & addressed

no clinical evidence of active disease & tests for DRTB are

negative, the remaining treatment course to be completed

in all cases (Educate & address myths/fear or any intolerance)

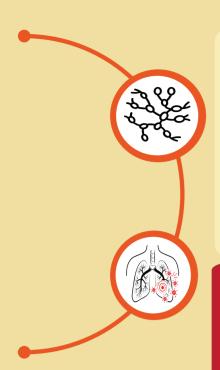
Endobronchial TB,

## REFERENCES

- 1. National TB Elimination Programme, Central TB Division. Training Modules for Programme managers & Medical Officers. Ministry of Health & Family Welfare, Government of India https://tbcindia.gov.in/index1.php?lang=1&level=1&sublinkid=5465&lid=3540 Last access on 06 March, 2022.
- Guidelines for Programmatic Management of Drug Resistant Tuberculosis in India March 2021. National TB elimination programme, Central TB Division, Ministry of Health & Family Welfare, Government of India https://tbcindia.gov.in/showfile.php?lid=3590 Last access on 06 March, 2022

#### PAEDIATRIC LYMPH NODE TUBERCULOSIS

ICD-10-A18.2



#### WHEN TO SUSPECT?

- Persistent enlargement of lymph node for >2 weeks in one or more areas in cervical/axillary/ inguinal regions
  - > 2 cm size or matted lymph nodes ± chronic sinus
- With/without associated systemic symptoms: fever, cough, poor appetite, weight loss
- With no evidence of recent scalp/skin lesions of draining area

农

TB is unlikely if: the lymphnode are few, small (< 2 cm) and may be persisting for a long time (months to years) without any systemic symptoms

#### **INVESTIGATIONS**

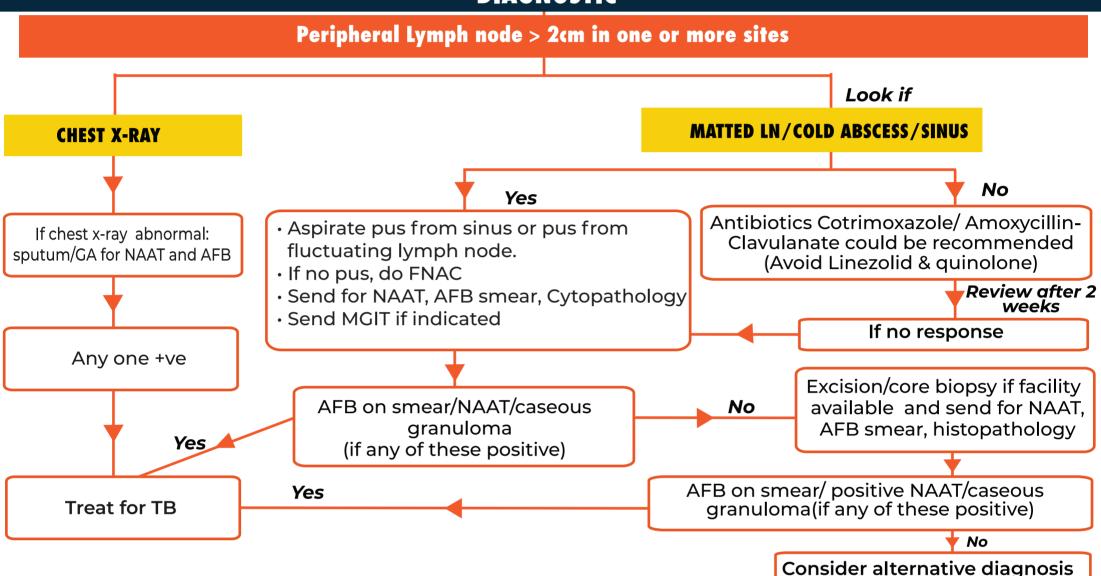
#### **Essential**

- Lymphnode aspirate:
- · Send for NAAT (+/- MGIT)
- · Smear for AFB
- Cytopathology

#### Desirable

- Lymphnode cytopathology (If NAAT and smear negative)
- Lymphnode Biopsy (Core/Excision)
- Chest X-ray
- Hemogram with peripheral smear

## **DIAGNOSTIC**



## TREATMENT AND MANAGEMENT

#### TREATMENT AND RESPONSE

- Treatment should be started and follow-up should be conducted as per NTEP guidelines
- 2 HRZE + 4 HRE (standard doses)
- Disappearance of constitutional symptoms with decrement or no increment in lymph node size suggests response to treatment
- Increment in lymph node size with disappearance of constitutional symptoms may suggest paradoxical reaction, provided drug resistance has been ruled out
- Increment in lymph node size without disappearance of constitutional symptoms suggests drug resistant TB/ alternate cause



- Do not treat for TB based on only positive mantoux test or FNAC suggestive of reactive lymph node with negative NAAT/AFB on smear
- Children with disappearance of constitutional symptoms with no increase in lymphnode size at the end of 6 months therapy,can be kept in follow-up

#### WHEN TO REFER TO AN EXPERT?

- Diagnosis is not established with FNAC/NAAT
- Surgical facility is not available to do excision or core biopsy
- DR is suspected due to any reason including non-response and the facility for DRTB testing are not available
- If there is any pointer towards possible malignancy e.g. skin or mucosal bleed or significant pallor or generalised adenopathy irrespective of the size or associated hepato-splenomegaly

#### **BCG LYMPHADENITIS**

- Age is usually < 2 years</li>
- Axillary and or supraclavicular lymphnode on the same side as BCG vaccination (usually given on the left)
- No systemic symptoms in immunocompetent children
- · Treatment:
  - · Wait and watch if small
  - If large and suppurative, repeated aspiration or rarely incision and drainage is required

\*NAAT or AFB smear positivity can not differentiate between BCG and MTB

#### **ABBREVIATIONS**

AFB: Acid fast bacillus
BCG: Bacille Calmette Guerin vaccine
DR: Drug resistant

FNAC: Fine needle aspiration cytology
HRZE: Isoniazid; Rifampicin; Pyrazinamide; Ethambutol
MGIT: Mycobacteria Growth Indicator Tube

NAAT: Nucleic acid amplification test
NTEP: National TB Elimination Programmet
TB: Tuberculosis

## REFERENCES

- 1. National TB Elimination Programme, Central TB Division. Training modules for programme managers & Medical officers. Ministry of Health and Family Welfare, Government of India accessed at https://tbcindia.gov.in/index1.php?lang=1&level=1&sublinkid=5465&lid=3540 on 24 February, 2022.
- 2. Guidelines for programmatic management of drug resistant tuberculosis in India March 2021. National TB elimination programme, Central TB division, Ministry of Health and Family Welfare, Government of India accessed at https://tbcindia.gov.in/showfile.php?lid=3590 on 10 February, 2022.

## PAEDIATRIC OSTEOARTICULAR TUBERCULOSIS

ICD-10-18.0

#### **POTT'S SPINE** (COMMONEST, 50% OF OSTEOARTICULAR TB)

- Insidious onset back pain for >6 weeks (Commonest thoracic > lumbar >cervical)
- Localized/Referred root pain
- TB Symptoms: Fever/anorexia/weight loss
- · CNS complications like Paraparesis (20-50%), cauda equina syndrome, paraspinal muscle wasting, severe pain
- · Examination: Local tenderness/deformity

#### **DACTYLITIS** (SHORT BONES)

- Swelling of short tubular bones of hands & feet (Proximal phalanx or metacarpals of index/middle/ring fingers are commonly affected)
- In children multiple or consecutive bones are involved, compared to a single bone in adults
- May present without pyrexia or signs of inflammation

#### **ARTHRITIS** (LARGE JOINTS-HIP/KNEE COMMONEST)

- · Insidious onset joint pain, swelling
- Monoarticular arthritis
- · Commonly associated with pulmonary or lymph node TB

#### **ESSENTIAL**

WHEN TO SUSPECT

#### · X-ray Spine

- → In early stage X-ray may be normal
- > May show end plate erosions, joint space narrowing/collapse, decreased vertebral height, paravertebral soft tissue shadow
- MRI Spine preferred if not feasible do CT
- > Marrow edema
- > Destruction of intervertebral disc, adjacent vertebral bodies & opposing end plates
- > Pre/para vertebral or epidural abscess
- Sputum/GA for NAAT, MGIT/LJ (if CXR abnormal)
- FNAC (if peripheral lymphnodes enlarged) for Cytology, NAAT & MGIT/LJ

#### **DESIRABLE**

**BOX A: Risk factors for TB** 

Contact history with TB case

Immunocompromised

·HIV

Image guided (USG/CT) aspiration of abscess (if feasible) for NAAT & MGIT/LJ. (No need of biopsy)

#### **ESSENTIAL**

- · Plain X-ray of involved parts
- › Diaphyseal expansile lesion
- > Periosteal reaction is uncommon
- > Healing is by sclerosis (usually gradual)
- · X-ray film of chest
  - → Sputum/GA for NAAT & MGIT/LJ, if CXR abnormal
- FNAC (if peripheral lymphnodes enlarged) for Cytology, NAAT & MGIT/LJ

#### **DESIRABLE**

· Image guided (USG/CT) aspirate from involved bones for NAAT & MGIT/LJ. (No need of biopsy)

#### **ESSENTIAL**

- Plain X-ray: A triad of X-ray abnormalities (Phemister's triad) includes
- › Peri-articular osteoporosis
- > Peripherally located osseous erosion
- Gradual joint space narrowing
- · Early stage synovitis & arthritis imaging may show wide joint space due to effusion
- · Bony ankylosis development is rare in TB arthritis in contrast to Pyogenic arthritis
- · USG/ MRI of joint
- · X-ray film of chest
- Sputum/GA for NAAT, MGIT(if CXR abnormal)
- · FNAC (if peripheral lymphnodes enlarged) for Cytology, NAAT & MGIT/LJ

#### **DESIRABLE**

· Image guided (USG/CT) aspirate from joint fluid for NAAT & MGIT/LJ. (No need of biopsy)

### **DIAGNOSTIC ALGORITHM**

#### BOX B: Clinical manifestation of Spinal TB (STB) /TB arthritis (TBA)

- Insidious onset back pain for >6 weeks (STB)
- · Spine deformity/Kyphoscoliosis/Gibbus/Paraplegia/Sensory loss/Autonomic dysfunction (STB)
- Insidious onset pain and swelling in joints for >6 weeks (TBA)
- TB Symptoms: Persistent Fever, Anorexia, Weight loss (>5% in last 3 months)

Suspect if symptoms of STB/TBA present with/without constitutional symptoms of TB (Box B) and/or risk factors of TB (Box A)

- X-ray of Spine (AP/Lateral): May show end plate erosions, narrow /collapsed joint space, reduced vertebral height, paravertebral soft tissue shadow. Early stage X-ray may be normal
- X-ray joints (AP/Lateral): Erosions, sclerosis, calcification or narrow joint space
- USG abdomen for Iliopsoas Abscess
- USG joints for joint effusion and diagnostic aspiration
- Chest X-ray, ESR, blood sugar, HIV

- · MRI Spine(100% sensitive): indicated in all cases: may show
  - › Marrow edema
- Destruction of intervertebral disc, adjacent vertebral bodies & opposing end plates
- > Prevertebral, paravertebral and/or epidural abscesses
- $\cdot$  MRI Joints: Synovial proliferation with periarticular picture s/o TBA
- · Radiological findings s/o osteoarticular TB and/or TB at additional site- can be labeled as clinically diagnosed Osteoarticular TB
- · Refer patients to higher centre for biopsy (Percutaneous CT-guided biopsy-preferred or open biopsy) of lesion for NAAT/culture to confirm diagnosis & DST and to rule out other diseases. (Microbiological confirmed osteoarticular TB) (Risks and benefits of obtaining a biopsy must be considered)

## MANAGEMENT

#### TREATMENT & MONTORING

- · Start treatment for microbiologically /Lab confirmed TB and probable TB
- Regimen: 2HRZE + 10HRE (Standard doses) + Pyridoxine 10 mg/day
- Follow up every month during treatment & subsequently every 3 months: Pott's spine with X-ray or MRI & Tubercular dactylitis or arthritis with plain X-ray
- Monitor on each visit :

AFB: Acid fast bacillus

CT: Computed Tomography

AP: Antero-Posterior

DR: Drug Resistant TB

CXR: Chest X-ray

- a. Symptomatic improvement, weight gain, side effects of medicines
- b. Microbiology: sputum/GA if CXR abnormal at end of IP. Site samples like aspiration of pus from lesions including psoas abscess (if worsening of symptoms/poor response)
- · Imaging: MRI/CT/X ray of affected parts: at end of treatment or early if worsening

### **Surgical Indications in Potts Spine**

- · Progressive neurological deficit
- Paraplegia of recent onset or severe paraplegia
- Persistent pain with spinal instability
- Spinal deformity-severe kyphotic deformity at presentation, or in children (<10 years of age) at high risk of progression of kyphosis with growth after healing of disease

HIV: Human Immunodeficiency Virus

#### WHEN TO REFER

- · Suspected osteoarticular disease if essential investigations are not available
- · Diagnosis (microbiological or probable) not established by investigations
- · Surgery needed: imaging suggest compressive myelopathy, motor deficits
- · No improvement with appropriate treatment
- · DR TB: diagnosed or high suspiscion

## Confirm microbiologically in all cases, if possible, before ATT

#### OTHER INFORMATION

- · In case of synovial fluid or cold abscess aspiration (against gravity), send samples for confirmation of TB in following 3 ways
  - > Two dry slide for demonstration of AFB (ZN staining)
  - > Two samples in formalin for histopathological examination > Two samples in saline for culture' followed by DST and/or NAAT
- Confirmed cases to undergo HIV/blood sugar testing/parent counselling \*MGIT/LJ (if MGIT not available)

**DST**: Drug Sensitivity Test ESR: Erythrocyte Sedimentation Rate IP: Intensive Phase **FNAC**: Fine Needle Aspiration Cytology LJ: Lowenstein Jensen **GA**: Gastric Aspirate MGIT: Mycobacteria Growth Indicator Tube

**NAAT**: Nucleic Acid Amplification Test HRZE: Isoniazid; Rifampicin; Pyrazinamide; Ethambutol s/o: Suggestive of **STB**: Spinal TB **TBA**: TB Arthritis **USG**: Ultrasonography

**ZN**: Ziehl Neelson

#### REFERENCES

MRI: Magnetic Resonance Imaging

- 1. National TB Elimination Programme, Central TB Division. Training Modules for Programme Managers & Medical Officers. Ministry of Health & Family Welfare, Government of India.
- https://tbcindia.gov.in/index1.php?lang=1&level=1&sublinkid=5465&lid=3540 Last access on 12 March, 2022. 2. Guidelines for programmatic management of drug resistant tuberculosis in India March 2021. National TB Elimination Programme, Central TB Division, Ministry of Health & Family Welfare, Government of India accessed at https://tbcindia.gov.in/showfile.php?lid=3590 Last access on 12 March, 2022.
- 3. Sharma SK, Ryan H, Khaparde S, Sachdeva KS, Singh AD, Mohan A, et al., Index-TB guidelines: guidelines on extrapulmonary tuberculosis for India, Indian J Med Res. 2017;145(4):448–6

This STW has been prepared by national experts of India with feasibility considerations for various levels of healthcare system in the country. These broad guidelines are advisory, and are based on expert opinions and available scientific evidence. There may be variations in the management of an individual patient based on his/her specific condition, as decided by the treating physician. There will be no indemnity for direct or indirect consequences. Kindly visit our web portal (stw.icmr.org.in) for more information.

© Indian Council of Medical Research and Department of Health Research, Ministry of Health & Family Welfare, Government of India.

#### **PAEDIATRIC TUBERCULAR MENINGITIS**

ICD-10-A17.0

#### WHEN TO SUSPECT?

- Fever with one or more of the following
- → Headache
- → Vomiting
- → Seizures
- Irritability/Lethargy/Drowsiness
- Loss of function e.g.
   recent onset deviation of eyes/mouth and/or
   weakness of arm/leg
   and/or altered mentation
- → Malaise, Anorexia, Weight loss

**Criterion 1** 

>3 of the following

>5 days of symptoms

• TLC < 15,000/ cumm

CSF sugar < 50% of</li>

of (one or more):

Basal exudates

> Hydrocephalus

> Tuberculoma

> Infarct

blood sugar

· CSF WBC 10-500/ cumm

CSF lymphocytes >50%

Neuroimaging finding

 Symptoms are usually of 5 to 7 days duration with insidious onset, particularly with history of exposure to infectious TB in past 2 years

#### **EXAMINATION**

- Assessment of sensorium\*
- Full/bulging anterior fontanelle
- Meningeal irritation-Neck stiffness, Kernig's sign & Brudzinski's sign
- Examine eye, if feasible for papillodema/ choroid tubercles/ optic atrophy
- · Cranial nerves
- Motor system including power, reflexes & plantar responses
- Peripheral lymph nodes
- Chest examination for signs of pulmonary involvement

\*Use any standardized scale including Glasgow Coma scale/ AVPU scale

## INVESTIGATIONS

#### **Essential**

- · CBC
- CSF examination
  - Cell count and differential
  - Sugar (with simultaneous blood sugar)
  - → Protein
  - → NAAT\*
  - MGIT culture
  - > Bacterial culture
- ·HIV
- · Contrast enhanced CT scan of head
- · CXR
- Gastric lavage/ Induced sputum in patients where CXR is abnormal and CSF NAAT is negative

\*ICMR/NTEP approved NAAT test, use 3-5 ml CSF if possible

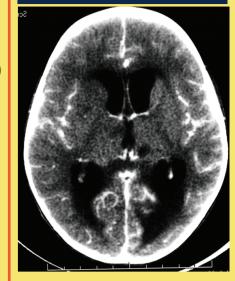
#### Desirable

 MRI brain with contrast when CECT is not contributory

# Optional

- · CSF cryptococcal antigen
- Contrast CT chest/abdomen to look for extracranial sites of infection

## NEUROIMAGING IN TB



#### **CECT** showing

 Hydrocephalus (ventricular dilatation)

AFB seen/CSF NAAT +ve

(Microbiologically

confirmed TBM)

Start treatment

- Thick basal exudates
- Tuberculoma

## **DIAGNOSTIC ALGORITHM**

• Immediate investigations
• CBC, HIV
• CECT head
• CSE: Cell count including

No

TREATMENT

- CSF: Cell count including differential, CSF sugar (with blood sugar), protein, NAAT, bacterial culture
- · CXR

Criterion 2- 2 or more risk factors for TBM

- · HIV infection
- Severe acute malnutrition
- Recent contact with infectious TB
- Evidence of TB elsewhere

Yes

**Criterion 3** 

2 or more criteria met

No

Continue investigations & management for partially treated bacterial meningitis

#### IF NOT BETTER

- · Repeat LP after 48-72 hours
- Expand search for TB elsewhere

Re-review criterion 1, 2 & 3 and see if > 2 criterion fulfilled

Does patient have falling CSF glucose/dropping sensorium?

No

Consider MRI contrast if not done earlier

Continue evaluation

· Anti TB drug regimen

› Pyridoxine 10 mg/day

Corticosteroids

to be done as per NTEP guidelines

2 HRZE and 10 HRE (in appropriate doses)

> Prednisolone 2 mg/kg/day for 4 weeks &

Slower taper needed in some patients

\*Equivalent dose of another steroid formulation

# • Treatment should be started & follow-up • Other supportive therapy

Have new focal deficit?

- Care of unconscious child
- Nasogastric feeding, if indicated
- Anti edema measures (mannitol/ hypertonic saline/glycerol/ acetazolamide)
- Anticonvulsants, if seizures
- Surgical therapy, if indicated
- > External ventricular drain
- VP shunt

# REFERRAL

Yes

Yes

- Cases should be managed at least at a district hospital
- Early referral to Medical College/ higher centre to be considered if
  - Unresponsive child/rapid deterioration indicating need for intensive care
  - No diagnosis after initial evaluation
  - Surgical treatment needed
  - MDR TB meningitis
  - No improvement/deterioration after2-4 weeks of treatment
- Need for ICU care

#### **ABBREVIATIONS**

AFB: Acid-fast Bacillus
CBC: Complete Blood Count
CECT: Contrast Enhanced Computed Tomography
CSF: Cerebro-spinal Fluid
CT: Computed Tomography

then taper over 4 weeks\*

may be used either injectable/oral

CXR: Chest X-ray
HIV: Human Immunodeficiency Virus
HRZE: Isoniazid; Rifampicin; Pyrazinamide; Ethambutol
ICU: Intensive Care Unit
LP: Lumbar Puncture

MDR: Multi-drug Resistant
MGIT: Mycobacteria Crowth Indicator Tube
MRI: Magnetic Resonance Imaging
NAAT: Nucleic Acid Amplification Test
NTEP: National TB Elimination Programme

TB: Tuberculosis TBM: Tubercular Meningitis TLC: Total Leucocyte Count VP: Ventriculo-peritoneal WBC: White Blood Cells

# REFERENCES

- 1. National TB Elimination Programme, Central TB Division. Training Modules for Programme Managers & Medical Officers. Ministry of Health & Family Welfare, Government of India https://tbcindia.gov.in/index1.php?lang=1&level=1&sublinkid=5465&lid=3540 Last access on 05 March, 2022.
- 2. Guidelines for Programmatic Management of Drug Resistant Tuberculosis in India March 2021. National TB Elimination Programme, Central TB Division, Ministry of Health & Family Welfare, Government of India https://tbcindia.gov.in/showfile.php?lid=3590 Last access on 05 March, 2022.

