



Extrapulmonary Tuberculosis

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Disclosures: None





Objectives

- 1) Describe the Epidemiology of Extrapulmonary Tuberculosis (EPTB) in the United States
- 2) Discuss the pathophysiology of EPTB and risk factors for EPTB
- 3) Discuss diagnosis and management of EPTB including Pleural, Lymph node, Bone (Potts disease), Central nervous system (CNS), and Genitourinary sites





Definitions

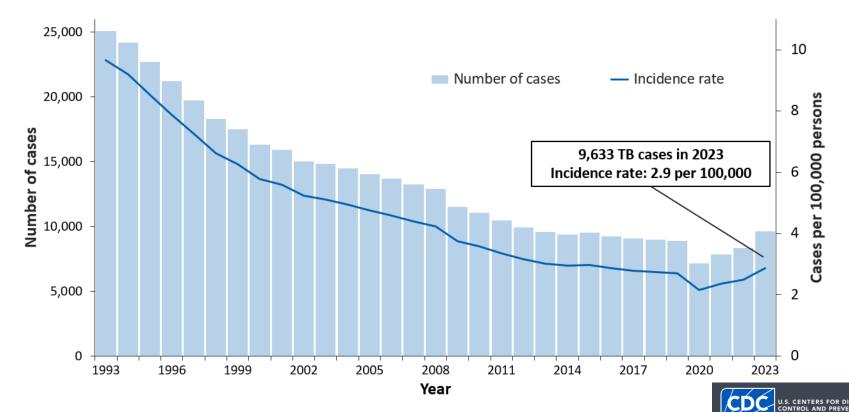
- EPTB : TB disease in any organ system other than pulmonary parenchyma
 - Pleural TB is extrapulmonary TB
- Disseminated TB: involvement of at least two noncontiguous body organs, hematogenous /bone marrow

Miliary is form of hematogenous dissemination and radiographic pattern

Paucibacillary : low bacterial burden

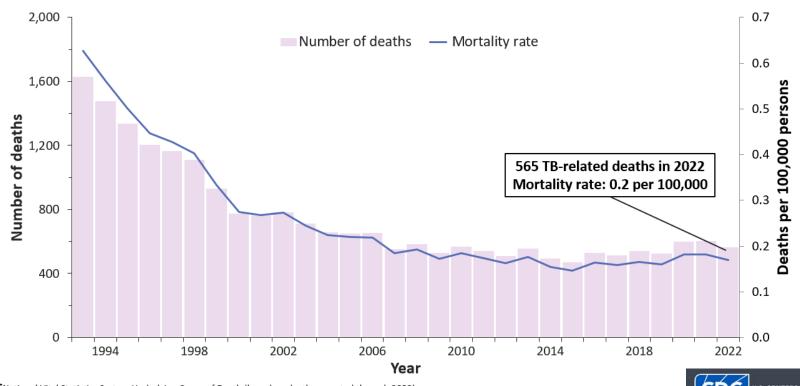


TB Cases and Incidence Rates, United States, 1993–2023





TB-Related Deaths* and Mortality Rates, United States, 1993–2022

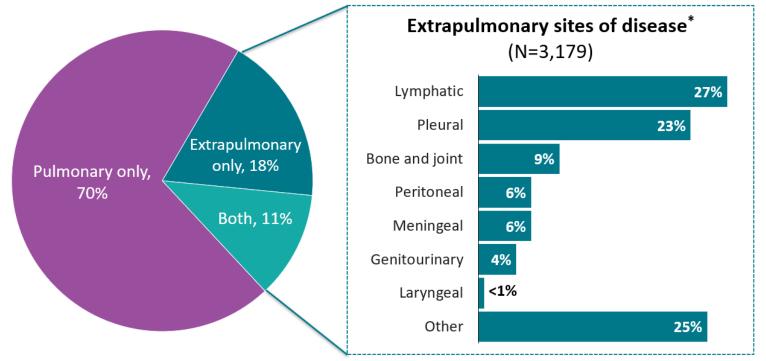


^{*}National Vital Statistics System Underlying Cause of Death (based on deaths reported through 2022)





Percentage of TB Cases by Site of Disease, United States, 2023



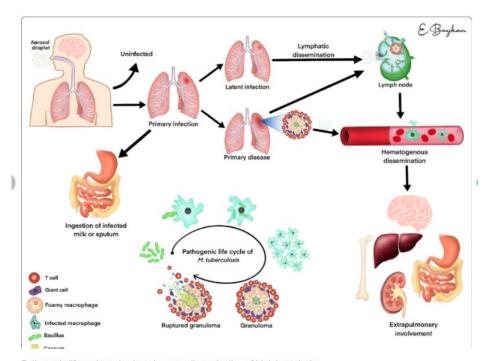
^{*}Persons might have more than one extrapulmonary site of disease.







Pathophysiology of TB



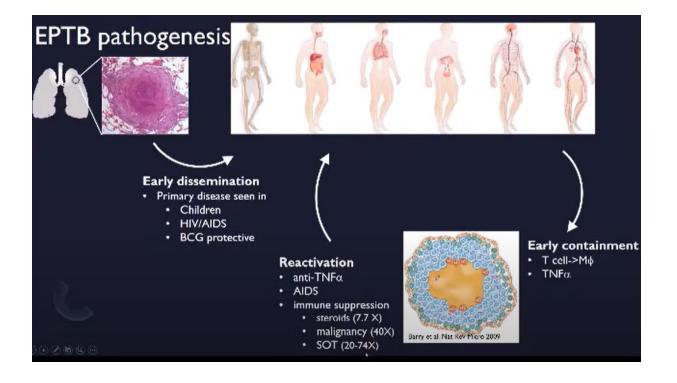
Pathogenic life cycle and extrapulmonary dissemination of M. tuberculosis

- Inhaled /Inspired to reach the alveoli
- Phagocytosed by alveolar macrophages into lysosomes
- some bacilli multiple and enter lymph nodes/blood stream
- Body's immune system usually intervenes to halt multiplication and prevent spread





EPTB Pathogenesis







EPTB Risk factors for US population

- 2 Major US studies
 - O Rieder et al in 1990
 - o Peto et al in 2009

- Female
- Born outside of United States
- Belonging to racial and ethnic minorities
- HIV
- Immunosuppression
 - Tissue allografts
- End stage renal disease (ESRD)





EPTB Risk factors for US population

Open Forum Infectious Diseases



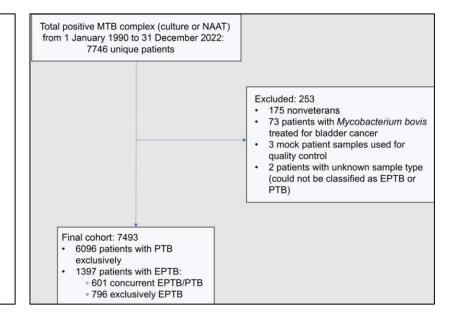




Risk Factors for Extrapulmonary Tuberculosis Among US Veterans, 1990–2022

Gina Oda, 1.0 Cynthia Lucero-Obusan, 1.0 Patricia Schirmer, 1.0 Joyce Chung, 1.0 and Mark Holodniy 1.2.0

¹US Department of Veterans Affairs, Public Health National Program Office, Washington, DC, and Palo Alto, California, USA, and ²Division of Infectious Diseases and Geographic Medicine, Stanford University, Stanford, California, USA







EPTB Risk factors for US population

Table 2. Characteristics of US Veterans With Extrapulmonary Tuberculosis (EPTB) Exclusively or Any EPTB, Compared With Pulmonary Tuberculosis Exclusively, 1990–2022 (N = 7493)

| Characteristic | PTB Exclusively, No. (%) ^a (n = 6096) | EPTB Exclusively, No. (%) ^a (n = 796) | P Value ^b | Any EPTB, No. (%) ^{a,c} (n = 1397) | P Value |
|---|---|---|----------------------|--|----------------------|
| Age, mean; median (IQR), y | 58.7; 58 (20) | 58.9; 59 (24) | .75 ^d | 56.7; 55 (23) | <.001 ^d |
| Sex | | | | | |
| Male | 6041 (99.1) | 779 (97.9) | .001" | 1375 (98.4) | .02° |
| Female | 55 (.9) | 17 (2.6) | | 22 (1.6) | |
| Race/ethnicity | | | | | |
| Hispanic/Latino | 260 (4.3) | 43 (5.4) | <.001° | 62 (4.4) | <.001 |
| Non-Hispanic white | 1544 (25.3) | 130 (16.3) | | 217 (15.5) | |
| Non-Hispanic black | 1535 (25.2) | 263 (33.0) | | 422 (30.2) | |
| Non-Hispanic Asian | 91 (1.5) | 12 (1.3) | | 17 (1.2) | |
| Non-Hispanic Al/AN or NH/OPI | 56 (0.9) | 9 (1.1) | | 14 (1.0) | |
| Unknown | 2610 (42.8) | 339 (42.6) | | 665 (47.6) | |
| Residence type | | | | | |
| Urban | 4652 (76.3) | 639 (80.3) | .03 ^{e,f} | 1133 (81.1) | <.001 |
| Bural | 1423 (23.3) | 154 (19.3) | | 257 (18.4) | |
| Unknown | 21 (0.3) | 3 (0.4) | | 7 (0.5) | |
| Birth country | | | | | |
| United States | 5789 (95.0) | 762 (95.7) | .11 | 1331 (95.3) | .10 |
| Other country | 222 (3.6) | 30 (3.8) | | 56 (4.0) | |
| Unknown | 85 (1.4) | 4 (0.5) | | 10 (0.7) | |
| Period of military service ⁹ | | 1 (2.2) | | , | |
| World War I or II | 1257 (20.6) | 188 (23.6) | <.001* | 267 (19.1) | 03* |
| Vietnam War | 3436 (56.4) | 418 (52.5) | | 826 (59.1) | |
| Korean conflict | 1173 (19.2) | 133 (16.7) | | 234 (16.8) | |
| Gulf War | 209 (3.4) | 50 (6.3) | | 63 (4.5) | |
| Other or unknown | 21 (0.3) | 7 (0.9) | | 7 (0.5) | |
| Area deprivation index | 21 (0.3) | 7 (0.0) | | 7 (0.0) | |
| Group 1 (lowest disadvantage) | 2004 (32.9) | 279 (35.1) | .20 | 517 (37.0) | .007* |
| Group 2 (highest disadvantage) | 3906 (64.1) | 500 (62.8) | .20 | 842 (60.3) | .007 |
| Not coded | 186 (3.1) | 17 (2.1) | | 38 (2.7) | |
| Antimicrobial susceptibility | 100 (3.1) | 17 (2.1) | | 30 (2.7) | |
| MDR | 60 (1.1) | 7 (0.9) | .14 | 13 (0.9) | .62 |
| Not MDR | 69 (1.1) 4591 (75.3) | 7 (0.9) 577 (72.5) | .14 | 13 (0.9) | .62 |
| | | | | | |
| Not tested | 1436 (23.6) | 212 (26.6) | | 317 (22.7) | |
| Tuberculosis risk factors | 4744 (00.0) | 454 (40.0) | 0048 | 000 (00.4) | <.001 |
| Alcohol dependence | 1744 (28.6) | 151 (19.0) | <.001° | 323 (23.1) | |
| Head/neck carcinoma | 253 (4.2) | 23 (2.9) | .09 | 40 (2.9) | .02° |
| Diabetes mellitus | 1332 (21.9) | 218 (27.4) | <.001* | 347 (24.8) | .02° |
| Hepatitis C | 873 (14.3) | 109 (13.7) | .63 | 225 (16.1) | .09 |
| HIV infection | 340 (5.6) | 74 (9.3) | <.001* | 188 (13.5) | <.001 |
| Homelessness | 1124 (18.4) | 89 (11.2) | <.001" | 206 (14.7) | .001 |
| Incarceration | 258 (4.2) | 18 (2.3) | .008° | 46 (3.3) | .11 |
| Malnutrition/low BMI | 417 (6.8) | 42 (5.3) | .10 | 83 (5.9) | .22 |
| Nicotine dependence | 2426 (39.8) | 195 (24.5) | <.001° | 400 (28.6) | <.001 |
| Organ transplant | 10 (0.2) | 2 (0.3) | .64 | 3 (0.2) | .72 |
| Silicosis | 20 (0.3) | 1 (0.1) | .50 | 3 (0.2) | .79 |
| Severe kidney disease | 459 (7.5) | 96 (12.1) | .001° | 159 (11.4) | <.001 |
| Substance abuse disorder | 1059 (17.4) | 99 (12.4) | .001° | 219 (15.7) | .13 |
| Outcomes | | | | | |
| Hospitalization within 30 d of MTB sample | 1662 (27.3) | 231 (29.0) | .30 | 427 (30.6) | .01° |
| LOS, mean; median (IQR) | 7.0; 0 (4) | 7.2; 0 (5) | .50 ^d | 7.9; 0 (6) | .058 |
| Death after tuberculosis diagnosis | | | | | |
| Characteristic | PTB Exclusively, No. (%) ^b (n = 6096) | EPTB Exclusively, No. (%) ^a (n = 796) | P Value ^b | Any EPTB, No. (%) ^{a,c} (n = 1397) | P Value ^b |
| Within 30 d | 467 (7.7) | 113 (14.2) | <.001° | 164 (11.7) | <.001° |
| Within 90 d | 754 (12.2) | 171 (21.5) | <.001° | 277 (19.8) | <.001° |

1245 (20.4)

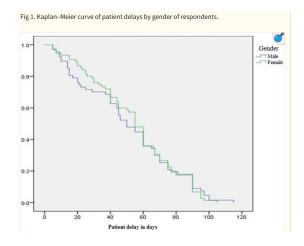
402 (28.8)

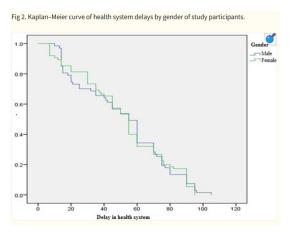




EPTB: Challenges of Early Diagnosis

- Need to have clinical suspicion with atypical presentation
- Acquire tissue/pathology
 - Clinical samples not easily accessible always
- Paucibacillary
 - Decreasing sensitivity of diagnostic tests





Knowledge of symptoms and delays in diagnosis of extrapulmonary tuberculosis patients in North Shewa zone, Ethiopia

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Editor: Frederick Quinn³

► Author information ► Article notes ► Copyright and License information

PMCID: PMC9202887 PMID: 35709224





Case 1

79-year-old male from Vietnam, with history of HTN, asthma, Hep C S/P treatment with chronic cough (> 3months), dyspnea since March 2025. Sputum smears and cultures were negative (X3). Had a thoracentesis in May 2025 and pleural fluid was smear negative but grew +MTB on culture. Started on TB medications in June 2025 after sent to LHD.





Pleural TB

Pathophysiology

- Rupture of subpleural caseous focus/granuloma into the pleural space
- Delayed hypersensitivity reaction to M.tb in pleural space resulting in inflammation/pleuritis

Clinical Manifestations

- Acute or subacute
- Cough (nonproductive)
- Chest pain
- Fever
- Shortness of breath
- Might vary in HIV patients (weight loss, night sweats etc)





Pleural TB

Imaging Findings



Rutgers

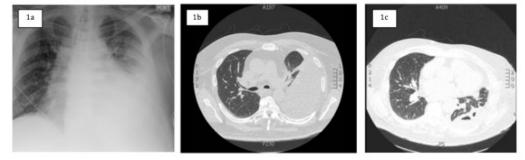


Figure 1. a) Chest X-Ray at presentation to our institution demonstrating a large left-sided pleural effusion. Figure 1b) Chest CT prior to chest tube placement demonstrating large left-sided pleural effusion. Figure 1c) Chest CT several days after chest tube placement demonstrating a partially loculated residual pleural effusion.

Brown Hospital Medicine

Lipman K, Lin C, Kennelly M, Patel S. The Diagnostic Challenges of Pleural Tuberculosis: A Case Report. Brown Hospital Medicine. 2024;3(1). doi:10.56305/001c.91176





Pleural TB

Diagnosis

- PPD/IGRA
- Imaging for effusion
- Pleural Fluid analysis
 - Exudate
 - Lymphocytic predominance mostly unless very early presentation
 - O Adenosine deaminase (ADA)>40
 - O NAAT/PCR
 - Xpert MTB/RIF
 - O Pleural fluid stain/culture
- Pleural biopsy

Treatment

- Treated like pulmonary TB
- 6 months usually
- No evidence to support role for adjunctive steroids





Pleural TB Clinical Pearl

- Without treatment TB
 pleuritis/effusion likely to resolve
 spontaneously in months
- However, majority will develop active TB within years indicating importance of diagnosis and treatment of pleural TB

Think of Pleural TB in the setting of TB infection (+ IGRA/TST), exudative pleural effusion with lymphocyte predominance.....





Case 2

40 year old lady from Honduras referred to LHD after + IGRA in Jan 2025. CXR was normal. Lost to follow up.

Had a mammogram outside and had some concerns leading to a diagnostic workup— with Ultrasound of Left breast.

US --multiple prominent /enlarged lymph nodes in L axilla
U guided biopsy with "suppurative inflammation/necrotic debris/ consistent with abscess".

Seen in March 2025 at which time complained of fever, night sweats, weight loss and started on RIPE/B6

Culture from biopsy + for Mtb





TB Lymphadenitis

Pathophysiology

- Oral route (drinking unpasteurized milk M bovis)
- Hematogenous seeding

Diagnosis
Usually a tissue diagnosis/culture from aspirate, biopsy, surgical, histology, pathology

Treatment
Like Pulmonary disease (usually 6 months)



Articles from Journal of International Oral Health: JIOH are provided here courtesy of International Society of Preventive and Community Dentistry



Historical photo demonstrating advanced scrofula (an older term for tuberculous lymphadenitis).

Author: Scrofula of the neck. From: Bramwell, Byrom Edinburgh, Constable, 1893 Atlas of Clinical Medicine. Source: National Library of Medicine, National Institutes of Health, USA.





TB Lymphadenitis

Analysis of the histopathological findings of lymph node biopsies at a tertiary care centre

Table 1: Diagnosis of lymph node biopsies.

| Diagnosis | Count | % | Male | Female | Ratio |
|---------------------------------------|-------|-------|-------------|-------------|-------|
| Tuberculous lymphadenitis | 305 | 65.03 | 100 | 205 | 1:2 |
| Cancer | 83 | 17.70 | 39 | 44 | 1:1.1 |
| Reactive | 68 | 14.50 | 27 | 41 | 1:1.5 |
| Bacterial infection | 7 | 1.49 | 1 | 6 | 1:6 |
| Histiocytic necrotizing lymphadenitis | 5 | 1.07 | 2 | 3 | 1:1 |
| Cat scratch | 1 | 0.21 | 1 | 0 | 1:0 |
| Total | 469 | | 170 (36.25) | 299 (63.75) | 1:1.7 |

Importance of a good physical Exam /imaging when appropriate to assess for lymphadenopathy

Table 2: Diagnosis of lymph node biopsies based on location.

| Location | Count (%) | Tubercul- ous lympha- denitis (%) | Cancer (%) | Reactive (%) | Bacterial infection (%) | Histiocytic necrotizing lymphadenitis (%) | Cat scratch (%) |
|--------------------|-------------|---|------------|--------------|-------------------------|---|-----------------------|
| Cervical | 316 (67.38) | 225 (47.97) | 39 (8.32) | 44 (9.38) | 4 (0.85) | 5 (1.07) | 0 (0.00) |
| Axillary | 69 (14.71) | 30 (6.4) | 19 (4.05) | 16 (3.41) | 2 (0.43) | 0 (0.00) | 1 (0.21) |
| Neck | 28 (5.97) | 15 (3.2) | 6 (1.28) | 7 (1.49) | 0 (0.00) | 0 (0.00) | 0(0.00) |
| Inguinal/ groin | 26 (5.54) | 11 (2.35) | 14 (2.99) | 0 (0.00) | 1 (0.21) | 0 (0.00) | 0 (0.00) |
| Abdomen | 26 (5.54) | 20 (4.26) | 5 (1.07) | 1 (0.21) | 0 (0.00) | 0 (0.00) | 0 (0.00) |
| Chest thorax | 4 (0.85) | 4 (0.85) | 0 (0.00) | 0 (0.00) | 0 (0.00) | 0 (0.00) | 0 (0.00) |
| Total | 469 | 305 (65.03) | 83 (17.70) | 68 (14.50) | 7 (1.49) | 5 (1.07) | 1 (0.21) |





TB Lymphadenitis Clinical Pearl

- Painless lesion
- Epidemiology is helpful
- Good physical exam
- Appropriate referrals for tissuebased diagnosis/cultures

Not uncommon for the lesion to enlarge, fluctuate in size/drain/new nodules

This does not mean that therapy is ineffective

Need to counsel patient and care team regarding this





Case 3

71-year-old HIV negative lady from Bolivia with Rheumatoid Arthritis (remote hx of methotrexate), Diabetes, with back/hip pain for months.

-11/8 MRI of pelvis: Destructive process involving the left SI joint with erosion of adjacent sacral wing and ileum demonstrating ring enhancement of this process, similar ring enhancing large fluid collection and gluteus maximus in close proximity to SI joint, these findings strongly suggestive of septic arthritis with adjacent large abscess and gluteus on the left, associated left iliac and inguinal adenopathy.

Sputum smears and cultures negative 11/2024 Gluteal abscess aspirate: No growth for aerobic and anaerobic bacteria 11/2024 Left SI joint aspirate: no growth for aerobic and anaerobic bacteria

Treated with Daptomycin + Ertapenem for a month





Case 3

- not improving on broad spectrum antibiotics
- MRI with 4.7 X 3.2 X 5.7 cm L sacroiliac joint abscess with osteomyelitis, improved gluteal abscess 7.4 x 5.5 x 3.0 cm in comparison to 11/8/24 MRI
- Underwent SI L biopsy and cultures sent for AFB (+ for M tb) and pathology + for granuloma

 RIPE started >1.5 months after presentation. Prior to this had rounds of PO antibiotics and then IV antibiotics





Musculoskeletal TB

Pathophysiology Hematogenous dissemination Possibly also direct trauma

Clinical Manifestations Bone/Joint Pain, erythema, soft tissue swelling, abscess

Diagnosis
Aspirated culture/tissue based, histology, pathology
Surgical bone culture and histology

Table 3 Localization of bone and joint tuberculosis

| References: | Mateo [45] | Enache [18] | Houshian [31] | Davies [14] |
|----------------------|---------------|-------------|------------------|----------------|
| Localization (%) | | | | |
| Spine | 64 | 0^{a} | 49 | 36 |
| Knee | 8 | 11 | 7 | 11 |
| Ankle | 8 | 8 | 0 | 8 |
| Elbow-arm | 2 | 11 | 5 | 5 |
| Wrist-hand | 6 | 26 | 4 | 11 |
| Hip | 0 | 10 | 6 | 9 |
| Rib | 2 | 26 | 2 | 1 |
| Sternoclavicular | 6 | 0 | 3 | 2 |
| Sacroiliac | 9 | 0 | 0 | 4 |
| Foot | 0 | 0 | 5 | 3 |
| Tibia | 0 | 0 | 2 | 2 |
| Trochanter- femur | 6 | 10 | 5 | 2 |
| Others | 0 | 5 | 3 | 5 |

^a The low incidence of spinal tuberculosis is explained by the fact that vertebral osteomyelitis is usually treated in specialized referral centers in the country of the study

Eur Spine J (2013) 22 (Suppl 4):S556-S566





Spinal Tuberculosis Clinical

Table 2 Clinical, analytical data and outcome of patients with spinal tuberculosis

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| Country: Year: No. of cases: | France [52] 1999 103 | Spain [10] 2004 78 | Taiwan [72] 2010 58 | France [47] 2006 24 | USA [58] 1995 20 | UK [37] 2009 17 | Turkey [67] 2001 694 | Spain [20] 2002 14 | Turkey [66] 2007 13 | Saudi Arabia [2] 2001 69 |
|------------------------------------|----------------------------|--------------------------|---------------------------|---------------------------|------------------------|-----------------------|----------------------------|--------------------------|---------------------------|--------------------------------|
| Age (mean) | 41 (17-84) | 48 (14-84) | 68 (36-86) | 61 | 49 | 29 | 32 | 58 | 68 | 53 (15-80) |
| Sex (men %) | 67 | 51 | 60 | 38 | 80 | 47 | 50 | 64 | 69 | 54 |
| % immigrants | 68 | 5 | - | 29 | 50 | 92 | _ | - | - | - |
| Underlying disease (%) | | | | | | | | | | |
| HIV | 0 | 15 | 0 | 0 | 25 | - | _ | 7 | 0 | _ |
| Immunosuppression | 6 | 8 | 5 | 8 | - | - | _ | - | - | - |
| Diabetes | 1 | 11 | 18 | 8 | - | - | _ | 13 | 23 | _ |
| Chronic renal failure | 2 | 0 | 18 | 0 | - | - | _ | 0 | 30 | _ |
| Clinical features | | | | | | | | | | |
| Previous tuberculosis | - | - | - | 25 | - | - | _ | 43 | - | 7 |
| Time to diagnosis (months) | 4 | 6 | 2 | 4 | 3 | 7 | _ | _ | 4 | _ |
| Pain (%) | 97 | 83 | 100 | 100 | 100 | 86 | _ | 100 | 100 | 84 |
| Fever (%) | 32 | 35 | 16 | 36 | - | 35 | 1.4 | 7 | 77 | 32 |
| Constitutional symptoms (%) | 40 | 36 | 8 | 36 | 35 | - | _ | _ | - | - |
| Neurological symptoms (%) | 50 | 45 | 53 | _ | 50 | 6 | 67 | 64 | 61 | 28 |
| Spinal deformity (%) | _ | 35 | - | _ | 55 | - | 57 | 50 | 50 | 17 |
| Tuberculin test (+) (%) | _ | 83 | - | 91 | 95 | - | _ | 50 | - | _ |
| Extraspinal involvement (%) | 20 | _ | 44 | _ | 35 | 47 | 5 | 50 | - | _ |
| Pulmonary involvement (%) | 15 | 28 | 32 | _ | 65 | 35 | 2.3 | 23 | _ | _ |





Pott's Disease



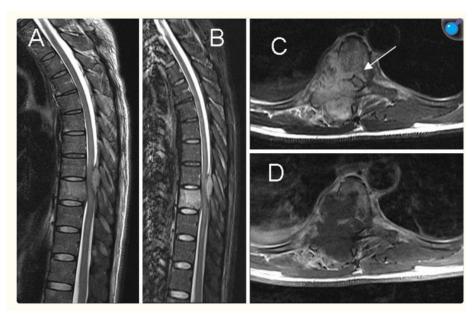


Sagittal T1-postcontrast weighted and STIR imaging show signs of advanced cervical infection with a large prevertebral abscess (*thin arrow*) and early skip lesion with epidural enhancement involving T6 and T7 levels (*thick arrow*)

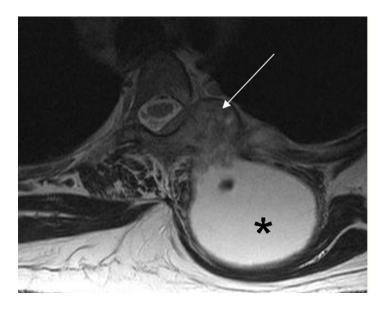




Pott's Disease



Paradiskal pattern. **a**, **b** Sagittal T2-weighted imaging and STIR show hyperintense signal in the T9 vertebral body, suggesting bone marrow edema. **c**, **d** Axial T2- and T1-postcontrast weighted imaging show paradiskal involvement and infection spreading to the epidural space (*arrow*) with spinal cord compression



Posterior pattern. Axial T2-weighted imaging shows a large cold abscess (*asterisk*) located in soft tissues and associated bone infection involving vertebral arch and costovertebral joint (*arrow*)

Eur Spine J. 2013 Jun;22 Suppl 4(Suppl 4):567-78.





Musculoskeletal TB

Medical Treatment

- Longer duration than pulmonary
 TB
 - 9-12 months in straight forward cases without resistances, hardware etc
 - O Longer regimens 12-18 months
 - Rifampin less regimen
 - Hardware association

Surgical Treatment

- Usually indicated for neurological deficits
 - relief of cord compression
 - recurrence of neurologic deficits;
 - instability of the spine
- Situations
 - there is poor response to chemotherapy
 - ongoing infection or ongoing deterioration
- Children with vertebrae affected in thoracic spine, kyphosis is likely to progress with growth





Musculoskeletal TB Clinical Pearls

- Indolent presentation
- "Cold abscess" vs "Hot abscess"
- Delayed culture/tissue diagnosis
- Needs a high index of suspicion to send for correct cultures to avoid delayed diagnosis
- Timely detection and treatment needed to prevent permanent damage and disability

- Radiographic findings lag behind clinical response and not the sole decision driver
- Drug interactions with Rifamycin and Pain medications
 - Rifampin can cause increased metabolism of narcotics





Case 4

67-year-old male with history of Hepatitis B, hypertension, gout, L ureteral mass S/P stenting with recurrent fevers, hematuria. Diagnosed with pyelonephritis and treated with rounds of antibiotics in the preceding months. Imaging with renal and bladder abnormalities led to biopsy of renal pelvis and bladder wall subsequently.

Micro/Pathology
Urine AFB culture *M.Tuberculosis* complex DNA by PCR
Renal Pelvis granulomatous inflammation
Urinary bladder wall granulomatous inflammation





GU TB INH /PZA resistance Intolerance to quinolones Numbness, tingling much improved and resolved with stopping quinolones Initial RIPE Rifampin, Ethambutol, Quinolone (Moxifloxacin to Levofloxacin)

Case 4

Started Cycloserine 250 mg PO Continued Rifampin , Ethambutol Vitamin B 6 100 mg PO daily

- Checked peak and in range
- Monitored for neurotoxicity and depression closely
- Tolerated well overall

- Monitored and counseled about skin discoloration and side effects such as GI, qt prolongation, Neuro
- Frequent check in's and clinical visits at least (monthly if not more)

Started Clofazimine 100 mg PO daily (0btained medication through UVA)

Recommended Duration was 9-12 months given complicated course, resistance and risk for relapse

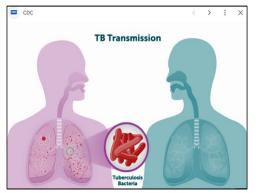
He completed close to 11 months therapy and wanted to travel to Philippines

He and his family were counseled extensively about risk for relapse and symptoms to monitor for

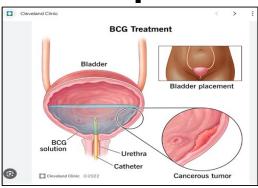




Genitourinary Tuberculosis Routes of Exposure







▶ One Health. 2020 Aug 11;10:100156. doi: 10.1016/j.onehlt.2020.100156 🗷

Risk factors of tuberculosis in human and its association with cattle TB in Nepal: A one health approach

<u>Tulsi Ram Gompo</u> ^{a,*}, <u>Asmita Shrestha</u> ^b, <u>Eliza Ranjit</u> ^c, <u>Bhanu Gautam</u> ^d, <u>Khim Ale</u> ^d, <u>Swoyam Shrestha</u> ^c, <u>Diker Dev Bhatta</u> ^a

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Genitourinary Clinical Presentation

Common

- Dysuria
- Urinary hesitancy
- Urinary frequency
- Gross hematuria (about 10%)
- Microhematuria (about 50%)

Uncommon

- Fever
- Weight loss
- Night sweats

Other Findings

- Infertility
- Bladder stenosis, stricture, vesicoureteral reflex, hydroureteronephrosis
- Penile nodules
- Increased PSA, or Ca 125
- Abnormal UA, sterile pyuria



Case Report

Tuberculosis at Vulva and Vagina

Pranom Buppasiri MD*, Teerayut Temtanakitpaisan MD*, Woraluk Somboonporn MD*

* Department of Obstetrics and Gynecology, Faculty of Medicine, Khon Kaen University, Khon Kaen, Thailand

J Med Assoc Thai 2010; 93 (5): 613-5 Full text. e-Journal: http://www.mat.orth/journal





Genitourinary Tuberculosis Diagnostic Tools

- Microscopic Urinalysis for AFB: + results in up to 90% of renal TB but not diagnostic as it will also appear + for nontuberculous mycobacteria If adequate bacteria present, specificity 97%, sensitivity 42-52%
- Molecular Testing: PCR is faster, detects even if few bacilli, not FDA approved for non pulmonary, sensitivity 80-90%
- Culture: gold standard, sensitivity 65%, specificity 100%
- Additional value with histology, pathology from tissue showing granuloma and AFB stains

CXR: In general, 50% or more of patients with genitourinary TB will have negative CXR

US: masses, wall thickness, vesicoureteral reflux, scarring, adhesions, granulomas,

CT: cortical masses, granulomas, calcifications throughout urinary tract, urinary strictures, bladder wall thickening, mucosal ulcers

Other Modalities:

- MRI
- Cystoscopy and ureteroscopy
- Hysterosalpingography
- IV Pyelogram





Genitourinary Tuberculosis Treatment

Mainstay is medical treatment: Same as pulmonary treatment with modifications based on resistances etc

Surgical treatment: needed in obstructive uropathy

- Abscess formation
- Hydronephrosis
- Infertility due to stricture formation of the fallopian tubes or vas deferens
- Intractable urinary symptoms such as "thimble" bladder
- Need for a tissue sample (biopsy) for diagnostic purposes
- Kidney failure with obstructive uropathy, especially if progressive and unresponsive to pharmacotherapy
- Nonfunctioning kidneys with symptoms not relieved by medical therapy
- Persistent symptoms unresponsive to medical treatment, such as pain or bleeding
- Recurrent endometrial tuberculosis or uterine bleeding resistant to medical therapy
- Strictures of the urethra resulting in voiding dysfunction
- Suspicion of an underlying malignancy





Genitourinary TB Clinical Pearls

- Cure rates are lower for drug resistant strains
- Lifetime risk of relapse has been reported as high as 22%, which is higher than pulmonary *Mtb* disease

- Latency period of up to 20 years or more between primary infection and evidence of GU disease
- Cause of infertility in especially developing countries





Pathophysiology

- Hematogenous dissemination
- Deep seated tubercular granulamotous foci
- Contiguous spread from vertebral osteomyelitis
- Latent infection reactivation

- Meningitis (most common form)
- Tuberculoma
- Spinal Arachnoiditis





Diagnostic features of tuberculous meningitis

Clinical

fever and headache (for more than 14 days)

vomiting

altered sensorium or focal neurological deficit

CSF

pleocytosis (more than 20 cells, more than 60% lymphocytes)

increased proteins (more than 100 mg/dl)

low sugar (less than 60% of corresponding blood sugar)

India ink studies and microscopy for malignant cells should be negative

Imaging

exudates in basal cisterns or in sylvian fissure hydrocephalus infarcts (basal ganglionic)

gyral enhancement

tuberculoma formation

Evidence of tuberculosis elsewhere

Basal cistern/meninges enhancement Hydrocephalus Parenchymal enhancement Cerebral infarct

Focal findings
Edema

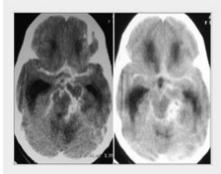


Figure 1: Contrast-enhanced computed tomography reveals the thick basilar exudate with the "spider leg" appearance





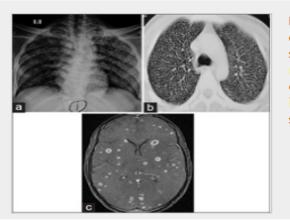


Figure 5: Roentgenogram of the chest (a) and contrast-enhanced computed tomography of the thorax (b) depict the miliary shadowing observed in a patient having definite tuberculous meningitis. Axial T1-weighted fat-suppressed spoiled gradient-echo (SPGR) gadolinium (GAD)-enhanced magnetic resonance image of the brain (c) shows multiple tuberculomas of various sizes involving different parts of the brain parenchyma

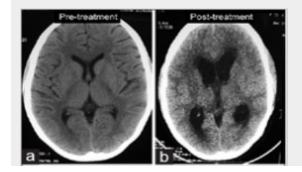


Figure 9: Computed tomography of the brain of two patients with definite tuberculous meningitis depicts the occurrence of hydrocephalus, as part of paradoxical reaction, in week 5 and week 6 after the initiation of anti-tuberculous treatment (a and b: Patient 1, c and d: Patient 2)





CSF

Opening pressure, Protein Glucose, Cell Differential, AFB smear, AFB and other appropriate cultures, X pert

Diagnostic accuracy of Xpert MTB/RIF Ultra for tuberculous meningitis in HIV-infected adults: a prospective cohort study

Nathan C Bahr ^{a,b}, Edwin Nuwagira ^c, Emily E Evans ^c, Fiona V Cresswell ^{d,e}, Philip V Bystrom ^a, Adolf Byamukama ^c, Sarah C Bridge ^{a,c}, Ananta S Bangdiwala ^a, David B Meya ^{a,d}, Claudia M Denkinger ^f, Conrad Muzoora ^c, David R

In 2013, WHO endorsed the Xpert MTB/RIF assay (Cepheid, Sunnyvale, CA, USA) as the preferred initial test to investigate tuberculous meningitis after a systematic review of 13 studies. The Xpert is cartridge-based fully-automated PCR test. Of three large cohorts, the first reported 67% sensitivity using Xpert in microbiologically proven tuberculous meningitis in HIV-infected South Africans. This study initially tested 1 mL of CSF but later found higher sensitivity (82%: 22 of 27 positive cases) when centrifuging 3 mL of CSF.

The diagnostic performance of GeneXpert MTB/RIF in tuberculosis meningitis: A multicentre accuracy study

Chenyuan Wang a †, Lina Davies Forsman b c †, Shanshan Wang a †, Sainan Wang a, Ge Shao a Haiyan Xiong a, Ziwei Bao d ‡ 久 図 , Yi Hu a ‡

The diagnostic sensitivity of Xpert was 71.1 % for definite TBM, and 5.5 % for probable/possible TBM. The positive rate of Xpert was improved with cerebrospinal fluid (CSF) increasing volume and was associated with <u>CSF</u> color (yellow). The additional indicators obtained by CART were CSF lactate and glucose and increased the sensitivity to 96.1 % (definite TBM) and 84.6 % (probable/possible TBM).

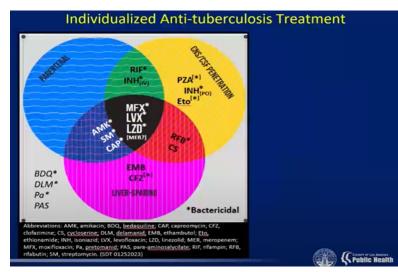




TB Meningitis Treatment

Treatment

- Timing is crucial !!!! High mortality
- Treatment is longer duration (a year) with good CSF penetrating drugs (9-12 months)
- Steroids mortality benefit (dexamethasone/prednisolone tapered over 6-8 weeks)
- Repeat LP's



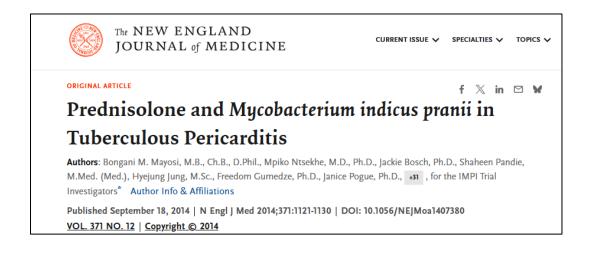
Neurosurgery: hydrocephalus, abscess, paraplegia, complications





Pericardial TB and Steroids

Adjunctive corticosteroids NOT routinely used in tuberculous pericarditis







Complications of EPTB

- High mortality for CNS, pericardial disease
- Mobility Complications with musculoskeletal TB
- Infertility in pelvic /GU/Fallopian Tube TB
- Cosmetic effect (cervical lymph node)
- Delayed diagnosis (long latency period such as GU TB)





Extrapulmonary TB: Draining Lesions

| Tuberculosis (M. tuberculosis) Extrapulmonary, draining lesion | Airborne + Contact + Standard | | Discontinue precautions only when patient is improving clinically, and drainage has ceased or there are 3 consecutive negative cultures of continued drainage [1025, 1026]. Examine for evidence of active pulmonary tuberculosis. |
|--|-------------------------------------|--|--|
|--|-------------------------------------|--|--|



2007 Guideline for Isolation Precautions: Preventing Transmission of Infectious Agents in Healthcare Settings

Last update: September 2024





Take Home Points -- EPTB

- High clinical index of suspicion
- Disease manifestations insidious onset/overlapping symptoms with other comorbid conditions
- Often Paucibacillary and Diagnostics can be challenging (Tissue diagnosis /delayed diagnosis)
- Treatment likely involves multidisciplinary approach/referrals/consults
 - Surgery in some cases
 - Treatment duration, choice of medications, steroids
 - May need recurrent procedures (LP, thoracentesis, paracentesis etc to monitor disease/symptom control)



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