Chicken & Fish

CID conference Hunter Ratliff 09/05/2024

Ages, dates, and other less-relevant (and identifying) information may have been changed



Case 1: HPI

A **58 y/o M** with PMH including central serous retinopathy, Afib, T2DM, HTN, HLD, OSA, CKD, hypothyroidism, hypogonadism p/w **multiple abscesses**

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- One month ago, had a large, heavy **chicken coup** fall across him
 - Fell on top of his right chest to left thigh
 - Unclear if puncture wound, but coop was covered in chicken wire
- A few days later, **developed SQ abscesses** at site of injury
 - Rx: Keflex & doxy but no improvement in symptoms
- Seen in surgery clinic where he was lethargic, mildly confused, and ill-appearing
 - Sent to ED and found to have sepsis

Case 1: Past medical history

A **58 y/o M** with PMH including central serous retinopathy, Afib, T2DM, HTN, HLD, OSA, CKD, hypothyroidism, hypogonadism p/w **multiple abscesses**

- <u>Central serous retinopathy</u>: Follows with Cleveland clinic, has been on high dose steroids, up to prednisone 60 mg / day, tapering recently
- **<u>Hx of cellulitis</u>**: Treated with IV antibiotics two years ago, unclear history
- Numerous **dental issues** in the past as well as **recurrent sinus infections**
- Hx of a few surgeries (multiple spine surgeries, TKR) w/ hardware

Case 1: Social History, Exposures, Risk Factors

<u>Geographic & Occupational</u>: The patient lives in West Virginia. He denies recent foreign or domestic travel. They are retired, used to sell pharma

<u>Substance</u>: They deny alcohol use and he does not use tobacco . They report no recreational drug use

<u>Sexual History</u>: Sexual activity with only women. Denied using any condoms or barriers. Denied ever testing positive for STIs

<u>Environmental exposures</u>: Denies freshwater exposure. Though he cannot recall if the coop caused a puncture wound, he thinks **if it did cause puncture injury there would have been soil exposure**

Animal Exposures: Cats, dogs, chickens

<u>Tattoos & Piercing</u>: They have previously gotten tattoos (professionally done)

Case 1: Initial exam

Hypotensive requiring low dose levophed

<u>Gen</u>: alert and oriented, NAD, vitals reviewed <u>Head/Neck</u>: NCAT; trachea appears midline, no gross LAD <u>ENT</u>: EOMI grossly, anicteric sclerae; MMM <u>Resp</u>: normal respiratory effort, symmetric chest rise <u>CV</u>: RRR; extremities perfused <u>GI</u>: non-distended; no rebound or guarding <u>Extremities</u>: **3+ BLE edema, + redness LLE**, no tenderness in the calves or thighs, no varicosities <u>Neurologic</u>: gait is normal, DTR intact bilaterally. Alert and oriented x3. **Drowsy on exam**.

Case 1: Initial workup

<u>WBC</u>: 21 (90% ΝΦ)

ESR 27; **CRP** 8.6

<u>Cr</u>: 1.9

Lactate: 4

<u>A1c</u>: 9.7

CT C/A/P

- Multiple probable subcutaneous abscesses overlying (up to 5.7cm) the right anterolateral upper abdominal soft tissues, with overlying cellulitis.
- No extension into the peritoneum evident.

CT LLE

16.2 cm subcutaneous abscess in the soft tissues of the medial left mid to distal thigh with invasion of adjacent musculature

Case 1: Summary

A **58 y/o M** with PMH including central serous retinopathy (on **steroids**), Afib, T2DM, HTN, HLD, OSA, CKD p/w **multiple abscesses** one month after **chicken coop** fell on him.

- Rx: Keflex & doxy but no improvement in symptoms
- Unclear if puncture injury
- Recurrent sinusitis & dental issues

Exam Hypotensive, BLE swelling, ?confused

Labs WBC 21, lactate 4, A1c 9.7

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- Rx: Keflex & doxy but no improvement in symptoms
- Unclear if puncture injury
- Recurrent sinusitis & dental issues

Exam Hypotensive, BLE swelling, **?confused**





Labs WBC 21, lactate 4, A1c 9.7

CT Head

2.3 cm rounded hypodense mass lesion within or adjacent to the **left anterior** frontal lobe possibly representing a cystic neoplasm or even abscess in the appropriate setting



- Taken from ED by surgery for emergent I&D of left leg & abdominal wall abscesses
- Operative cultures
 - Abdomen: Raoultella planticola
 - Leg: No growth at time of chart review
- Admitted to MICU for shock, transferred to Ruby for NSGY evaluation
- ID consulted (on my first day of fellowship)

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- Operative cultures
 - Abdomen: Raoultella planticola, S epi
 - Leg: Nocardia farcinica
 - Brain: Nocardia, S epi
- s/p multiple abdominal & leg washouts
- NSGY I&D of brain abscess



- Uncomplicated first admission
- Discharged on
 - Bactrim + Zyvox (CNS: Nocardia + MRSE)
 - Augmentin (SQ: Raoultella)

Antimicrobial course

- Bactrim / Imi / Linezolid (empiric)
- Discharge:
 - Bactrim + Zyvox (Nocardia + MRSE)
 - Augmentin (Raoultella)

Micro data

- Abdomen: Raoultella planticola, S epi
- Leg: Nocardia farcinica
- Brain: Nocardia, S epi

Case 1: Readmission

- Readmitted for unrelated reasons, but family wanted IV treatment
- Susceptibilities returned. Discharged on
 - CNS nocardia: Moxi + Bactrim
 - CNS MRSE: Vanco (DOT: 4 weeks) d/t TCP
- At discharge, had finished 2 weeks of Raoultella coverage for SQ infection

Antimicrobial courses

- Bactrim / Imi / Linezolid (empiric)
- Discharge #1:
 - Bactrim + Zyvox (Nocardia + MRSE)
 - Augmentin (Raoultella)
- Discharge #2:
 - CNS nocardia: Moxi + Bactrim
 - CNS MRSE: Vanco (DOT: 4 weeks)

Micro data

- Abdomen: Raoultella planticola, S epi
- Leg: Nocardia farcinica
- Brain: Nocardia, S epi

Nocardia susceptibilities

Tobramycin Linezolid Bactrim Doxycycline Ciprofloxacin Amoxicillin/Clavulanate Moxifloxacin Amikacin Imipenem Ceftriaxone Clarithromycin Minocycline 32 Resist 4 Suscept 2/38 Suscept 8 Resist 0.5 Suscept 8/4 Suscept 0.25 Suscept 1 Suscept 32 Resist >=128 Resist >=32 Resist 4 Intermed

Case 1: Clinic #1

- During OPAT, developed AKI so vanco stopped a little early (but likely still had ~4 weeks)
- Having GI intolerance (maybe to Bactrim?)
- Kept on Moxi & Bactrim until 6 weeks

Antimicrobial courses

- Bactrim / Imi / Linezolid (empiric)
- Discharge #1:
 - Bactrim + Zyvox (Nocardia + MRSE)
 - s/p 2 weeks Augmentin (Raoultella)
- Discharge #2:
 - CNS nocardia: Moxi + Bactrim
 - o s/p 4 weeks Vanco
- Clinic #1:
 - CNS nocardia: Moxi + Bactrim

Micro data

- Abdomen: Raoultella planticola, S epi
- Leg: Nocardia farcinica
- Brain: Nocardia, S epi

Nocardia susceptibilities

Tobramycin Linezolid Bactrim Doxycycline Ciprofloxacin Amoxicillin/Clavulanate Moxifloxacin Amikacin Imipenem Ceftriaxone Clarithromycin Minocycline 32 Resist 4 Suscept 2/38 Suscept 8 Resist 0.5 Suscept 8/4 Suscept 0.25 Suscept 1 Suscept 32 Resist >=128 Resist >=32 Resist 4 Intermed

Case 1: Clinic #2

- During OPAT, developed AKI (again) so Bactrim stopped at 6 weeks. GI symptoms resolved
- Added Augmentin for SQ disease

Nocardia susceptibilities

Tobramycin	32 Resist
Linezolid	4 Suscept
Bactrim	2/38 Suscept
Doxycycline	8 Resist
Ciprofloxacin	0.5 Suscept
Amoxicillin/Clavulanate	8/4 Suscept
Moxifloxacin	0.25 Suscept
Amikacin	1 Suscept
Imipenem	32 Resist
Ceftriaxone	>=128 Resist
Clarithromycin	>=32 Resist
Minocycline	4 Intermed

Antimicrobial courses

- Bactrim / Imi / Linezolid (empiric)
- Discharge #1:
 - Bactrim + Zyvox (Nocardia + MRSE)
 - s/p 2 weeks Augmentin (Raoultella)
- Discharge #2:
 - CNS nocardia: Moxi + Bactrim
 - s/p 4 weeks Vanco
- Clinic #1:
 - CNS nocardia: Moxi + Bactrim
- OPAT: Had to stop bactrim 2/2 AKI
- Clinic #2:
 - o c/w Moxi
 - Added Augmentin

Case 1 discussion



Links to articles discussed here

Nocardia

Objectives

- Review microbiology & risk factors
- Describe clinical syndromes
- Review treatment options



Articles

Nocardia: Microbiology

- Delicate filamentous gram-positive branching rods
- Weakly acid fast
- Aerobic growth

Numerous species

• Species can be helpful in predicting susceptibilities (Toyokawa 2021)



Nocardia: Antimicrobial susceptibilities

	N. farcinica	N. nova	N. brasiliensis	N. cyriacigeorgica	N. abscessus	N. otitidiscaviarum
Bactrim	S	S	S	S	S	S
Imipenem	S	S	R	S	S	R
Amikacin	S	S	S	S	S	S
Linezolid	S	S	S	S	S	S
Tobramycin	R	R	S	S	S	S
Augmentin	S	R	S	R	S	R
Ceftriaxone	R	S	R	S	S	R
Ciprofloxacin	R	R	R	R	R	R
Clarithromycin	R	S	R	R	R	R
Doxycycline	R	R	R	R	S	R
Minocycline	R	R	R	R	S	R
Erythromycin	R	S	R	R	R	R

Adapted from UpToDate

Nocardia: Risk factors

- Natural reservoir is **soil & decaying vegetation**
- Transmission via inhalation or direct inoculation
- Immunosuppression is major risk factor (though can occur in immunocompetent)
 - Steroids are most frequently reported risk factor
 - Solid organ transplant

- Subacute infection
- Main sites of infections
 - Lungs
 - o Skin
 - CNS
- Hematogenous spread is common in disseminated disease

- Subacute infection
- Main sites of infections
 - Lungs
 - o Skin
 - CNS
- Hematogenous spread is common in disseminated disease

Pulmonary disease

- Most common site of disease
- Variable presentation:
 - Pneumonia
 - Lung abscess
 - Cavitary lesion
 - Empyema
- More likely to go to CNS (vs skin)
- Can mimic flares of chronic lung disease (e.g. sarcoid patient who is on steroids)

- Subacute infection
- Main sites of infections
 - Lungs
 - o Skin
 - CNS
- Hematogenous spread is common in disseminated disease

Skin / soft tissue

- Often in setting of trauma
- Primary skin infection is most common in immunocompetent

Types

- 1. Localized: Cellulitis or SQ abscess +/- drainage
- 2. **Lymphocutaneous**: Nodular lymphangitis "sporotrichoid nocardiosis"
 - If thinking sporotrichosis (e.g. thorn injury) consider nocardia
- 3. **Mycetoma**: chronic cutaneous infection that coalesces to large necrotic abscess with draining sinus tracts

- Subacute infection
- Main sites of infections
 - Lungs
 - o Skin
 - CNS
- Hematogenous spread is common in disseminated disease

CNS disease

- Nocardia loves to go to the brain
- One or more abscesses
- May or **may not be symptomatic**
- If someone has nocardia, you should scan their head!

Nocardia: Initial treatment

- Bactrim is the mainstay treatment
 - Resistance can occur, so send for susceptibilities
- Initial therapy often **Bactrim + something else**
 - Often imipenem or amikacin
 - If CNS, some use all three

- Disseminated disease (including CNS) induction therapy:
 - At least six weeks of double coverage
 - Frequently IV

Nocardia: Long term treatment

- Always guided by susceptibilities and clinical response
 - **High risk of relapse** / treatment failure
- **PO stepdown**: After 3-6 weeks and clinical response:
 - **Non-CNS**: After **3 weeks**, consider PO monotherapy
 - **<u>CNS &/or immunocompromised</u>**: After **6 weeks**, consider switch to PO combo therapy
- Duration of therapy
 - Not well defined
 - **Isolated cutaneous**: 3-6 months (immunocompetent), 6-12 months (immunocompromised)
 - **CNS or immunocompromised**: At least one year
- Secondary prophylaxis:
 - May be some role for Bactrim PPx (1 DS daily) if immunosuppressed and no expected recovery (e.g. BMT, SOT)

NOCARDIA Spp



💟 @TheIDTrivia



Case 2: HPI

A 25 y/o M with PMH of bipolar (on Lamictal) is referred to ID clinic for a positive PPD

- Feels well, ROS entirely negative
- No known exposure to TB
- Prior TB testing (both TST & IGRA) have all been negative

Case 2: Social History, Exposures, Risk Factors

<u>Geographic</u>: He has **never traveled abroad** (all of his military work was stateside). During his time in the military, was **stationed in Louisiana** for most of the time, but also a few months in North Carolina. When he left the military two years ago, he moved from Louisiana to WV and hasn't traveled out of WV since

<u>Occupational / Hobbies</u>: Joined Army six years ago. Primarily **worked in healthcare** (his base's clinic) doing mostly **administrative work**. Would provide **limited patient care** when needed (e.g. triage) and only off-base work was during EMT training. While in Louisiana, he went out to the **bayous to fish**. Still enjoys **fishing in WV**, but not as good crayfishing up here. Now working in athletics department here, which prompted TB testing.

Case 2: Social History, Exposures, Risk Factors

<u>Infectious PMH</u>: **No history of BCG vaccine**. No known household contacts with TB or who are immunocompromised. Aside from what was mentioned in occupational section, no other worrisome possible exposures to TB (incarceration exposure, homeless shelters)

<u>Animal Exposures</u>: Has pets (cats and fish). Otherwise denies farm animal exposures, bird/reptile exposures, or other animal exposure.

<u>Substance</u>: They only drink socially and he does not use tobacco. They report no recreational drug use

<u>Sexual History</u>: They are are sexually active with only their female partner (of many years). The patient reports consistent condom use. No history of STIs

Tattoos & Piercing: They have previously gotten tattoos (professionally done) and no piercings .

Case 2: TB testing history

- Initial TST when he joined the military was **negative**
- While in the military, had TB testing (both TST and IGRA) **every six months** that were **all negative**
 - Was not told that any of his clinic patients were positive for TB
- He had testing at the end of his time in the service which was **negative**
- Had two step PPD for pre-employment screen

Date	PPD induration	
-3mo	0mm	
-3mo + 9 days	12mm	
1 week ago	15mm	



Case 2: Summary

A **25 y/o M** with PMH of bipolar (on Lamictal) is referred to ID clinic for a **positive PPD**

- Pre-military: TST negative
- q6mo: all negative
- End of service: **negative**

Date	PPD induration	
-3mo	0mm	
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1 week ago	15mm	

Case 2: Conclusion

A **25 y/o M** with PMH of bipolar (on Lamictal) is referred to ID clinic for a **positive PPD**

- Pre-military: TST **negative**
- q6mo: all negative
- End of service: **negative**

Date	PPD induration
-3mo	0mm
-3mo + 9 days	12mm
1 week ago	15mm

HIV screen negative

QuantGold negative

Treated PPD as false positive

• Possibly from NTM exposure given fishing history & time in Louisiana

Case 2 discussion



Links to articles discussed here

NTM infections & Mtb testing

NTM = nontuberculous
mycobacterial
Mtb = Mycobacterium tuberculosis



Objectives

Articles

- Review the classification of mycobacteria
- Review features of NTM
 - Transmission & exposures
 - Clinical syndromes
- Describe geographic distribution of NTM infections in US
- Discuss LTBI screening guidelines
 - Focus on distinguishing NTM from Mtb

Types of mycobacteria

Mycobacterium tuberculosis complex

- *M. tuberculosis*
- M. bovis
- M. africanum
- M. microti
- M. canetti

M. leprae

Rapidly growing NTM

Slow growing NTM

Types of mycobacteria

Mycobacterium tuberculosis complex

M. leprae

Rapidly growing NTM

- *M. fortuitum complex*
 - *M. fortuitum*
 - *M. peregrinum*
 - *M. porcinum*
- M. chelonae
- *M. abscessus complex*
- M. smegmatis
- M. mucogenicum

Slowly growing NTM **Photochromogens** M. kansasii • M. marinum **Scotochromogens** *M. gordonae* M. scrofulaceum **Nonchromogens** *M. avium complex* • *M. avium* • *M. intracellulare* • M. chimaera M. terrae complex M. ulcerans . M. xenopi M. simiae • M. malmoense M. szulgai M. asiaticum M. haemophilum



NON-TUBERCULOUS MYCOBACTERIA (NTM) 🛹

Ubiquitous environmental organisms

Higher rates of isolation and clinical disease in SE US

NTMS often classified as rapidly growing (RGM) and slowly growing mycobacteria

RGM = usually grow within 7 days*

*specifically means growth on solid media after subculture -- growth in liquid media influenced by quantity of inoculum!

Runyon classification can help with phenotypic identification

· Photochromogens: slow growing; yelloworange pigment only with light exposure

- Scotochromogens: pigment +/- light exposure
- Nonchromogens: produce no pigment (all RGMS!)

Transmission via:

 Inhalation of infected aerosols from environment (shower head, hot tub)

 Ingestion of organism (presumed route for disseminated MAC)

 Direct inoculation in skin breaks or trauma

 No human-human or animalhuman transmission (not communicable)

Inoculate clinical specimens in both: Solid media Middlebrook 7H11 media Lowenstein-Jensen media Liquid media MGIT broth

> Episode 7: MACsterclass febrilepodcast.com | @febrilepodcast | @swinndong



Disseminated

infection (ICH)

M.avium, M.kansasii M.abscessus M.chelonae

Where does NTM live?

Habitat	Reference
Natural waters	Falkinham et al. 1980; von Reyn et al. 1993
Drinking water distribution systems	Covert et al. 1999; Falkinham et al. 2001
Biofilms in drinking water distribution systems	Falkinham et al. 2001; Torvinen et al. 2004
Building, hospital, and household plumbing	Du Moulin <i>et al.</i> 1988; Wallace <i>et al.</i> 1998; Nishiuchi <i>et al.</i> 2007; Falkinham <i>et al.</i> 2008
Hot tubs and spas	Embil <i>et al.</i> 1997; Kahana <i>et al.</i> 1997; Mangione <i>et al.</i> 2001; Marras <i>et al.</i> 2005
Natural and household/building aerosols	Falkinham <i>et al.</i> 2008
Boreal forest soils and peats	livanainen <i>et al.</i> 1997, 1999
Acidic, brown-water swamps	Kirschner et al. 1992
Potting soils	De Groote <i>et al.</i> 2006
Metal removal fluid systems	Bernstein <i>et al.</i> 1995; Shelton <i>et al.</i> 1999; Moore <i>et al.</i> 2000

Table 2: Habitats of environmental opportunistic mycobacteriaFalkinham 2009

Mycobacterium avium complex

Exposures:

- Soil
- Water (aerosolized from showerheads; hot tubs, natural surface water)
- Animals
- Higher rates near gulf coast

Disease:

- Lung disease
- Disseminated in immunocompromised

Slow growers

- M. avium complex
- M. kansasii
- M. marinum
- *M. ulcerans*
- M. xenopi
- *M. simiae*
- M. malmoense

- M. abscessus complex
- *M. fortuitum complex*
- *M. chelonae*
- *M. abscessus* subspecies *massiliense*
- *M. abscessus* subspecies *bolletii*

Mycobacterium kansasii

Exposures:

- Also high predisposition for south coast & central plains
- Not found in soil or natural water supply
- Has been found in city water supply

Disease:

- Can cause lung disease like MTB
- Similar to MAC, mostly occurs in those with existing lung disease or immunocompromised

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Mycobacterium abscessus complex

Exposures:

- Similar environmental exposures (soil, water, animals)
- Hospital tap water
- Breast implant outbreak

Disease:

- Lung disease (even in immunocompetent)
- SSTI
- Disseminated in immunocompromised

Slow growers

- M. avium complex
- M. kansasii
- M. marinum
- *M. ulcerans*
- *M. xenopi*
- *M. simiae*
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Very old study of 275,000 Navy recruits (1969)

- I can't access the article but sounds like they all lived their entire lives in one county
- Over 70% of those in southeastern or gulf coast states had been exposed or infected with MAC

Geographic variation in the frequency of reactors to PPD-S (top) and to PPD-B 9 (bottom). Edwards, et al, Am Rev Respir Dis 1969; 99(Suppl):1.





Prevalence and Incidence Rates of NTM

277.6

2016

Cases per 100k patient-years

- Also VA patients (2009 2012)
 - Not restricted to COPD patients
- Identified cases by ICD code and natural language processing of micro data



M. chelonae-abscessus group M. kansasii M. avium complex Other Mycobacteria

Incident rates of NTM infections in VA patients (2009-2012) Jones et al (2018), Figure 1

- Environmental study out of LSU & Tulane (summer of 2011)
- PCR for M. ulcerans
 - So far no human cases in US from M ulcerans

Military base

- Samples collected from 9 sites
 - High rates (47%) found in water samples
 - Similar rates for fresh / brackish / salt water



No. of *M. ulcerans*-positive samples/total no. of samples (%)

Location	Aquatic vegetation	Water	Combined
Area 1	1/6	2/6	3/12 (25)
Area 2	1/8	4/6	5/14 (36)
Area 3	1/6	4/6	5/12 (42)
Area 4	1/9	3/9	4/18 (22)
Area 5	1/7	3/6	4/13 (31)
Area 6	3/6	1/6	4/12 (33)
Area 7	0/6	3/6	3/12 (25)
Area 8	0/5	4/6	4/11 (36)
Area 9	1/6	3/6	4/12 (33)
Total	9/59 (15)	27/57 (47)	36/116 (31)

<u>Top</u>: Sampling sites in Louisiana study of M. ulcerans (Figure 1) <u>Bottom</u>: Sample PCR positivity for M. ulcerans, by type & source (Table 3) Hennigan (2013)

But back to the Cajun food







Mycobacterium marinum

- Infection is less common
- Found in water (fresh & salt): marine environment, swimming pools, and fish tanks
- **Fish tank granuloma**: Sink infection from contaminated fish tank, often when preexisting skin injury

Slow growers

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Mycobacterium marinum

- Infection is less common
- Found in water (fresh & salt): marine environment, swimming pools, and fish tanks
- **Fish tank granuloma**: Sink infection from contaminated fish tank, often when preexisting skin injury
- Can cause **false positive TST test**
- Case series of patients (n=7) who had SSTI infection with M marinum (Lewis 2003)
 - 100% had PPD <u>></u> 10mm
 - 29% had PPD ≥ 15mm
 - Four patients had negative PPD documented before infection

Slow growers

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Mycobacterium marinum

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- Found in water (fresh & salt): marine environment, swimming pools, and fish tanks
- **Fish tank granuloma**: Sink infection from contaminated fish tank, often when preexisting skin injury
- Can cause **false positive TST test** (Lewis 2003)
- Case series of NTM SSTI (n=78) from **UTMB**, which is on the gulf coast (Philips 2019)
 - M. abscessus (47%)
 - M. fortuitum (26%)
 - M. marinum (21%)

Slow growers

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IDSA: LTBI screening

 Persons at low risk for Mtb infection and disease progression NOT be tested for Mtb infection

Risk of Infection

Groups with Increased Likeli- hood of Infection with Mtb	Benefit of Therapy	LTBI Testing Strategy		
Household contact or recent expo- sure of an active case	Yes	Likely to be Infected Lik Low to Intermediate Risk of Progression Hig (TST ≥ 10mM) (TS		Likely to be Infected High Risk of Pro- gression (TST ≥ 5mM)
Mycobacteriology laboratory personnel	Not demonstrated			
Immigrants from high burden countries (>20 / 100,000)	Not demonstrated			
Residents and employees of high risk congregate settings	Yes			
None	Not demonstrated	Unlikely to be Infected (TST > 15mM)		
		Risk of Developing Tuberculosis if Infected		Infected
		Low	Intermediate (RR 1.3 -3)	High (RR 3-10)
		No risk factors	Clinical predisposition Diabetes Chronic renal failure Intravenous drug use	Children age less than 5 HIV infection Immunosuppres- sive therapy Abnormal CXR consistent with prior TB Silicosis
		Benefit of Therapy Not demonstrated Yes		
				Yes

LTBI = latent tuberculosis infection Mtb = Mycobacterium tuberculosis

Paradigm for evaluation of those with LTBI based on risk of infection, risk of progression to tuberculosis, and benefit of therapy. (figure 1, 2017 IDSA guidelines)

IDSA: LTBI screening

- Persons at low risk for Mtb infection and disease progression NOT be teste for Mtb infection
- If you *have* to test them
 - Favor IGRA over TST (if >5 y.o.)
- If **initial test is positive**, suggest a second diagnostic test

	Group	Testing Strategy	Considerations	
te	Likely to be Infected High Risk of Progression (TST ≥ 5mM)	Adults Acceptable: IGRA OR TST Consider dual testing where a positive result from either result would be considered positive Children ≤ 5 years of age Preferred: TST Acceptable: IGRA OR TST Consider dual testing where a positive result from either would be considered positive'	Prevalence of BCG vaccination Expertise of staff and/or labora-	
	Likely to be Infected Low to Intermediate Risk of Progression $(TST \ge 10mM)$	Preferred : IGRA where available Acceptable : IGRA or TST	tory Test availability Patient perceptions Staff perceptions	
	Unlikely to be Infected (TST > 15mM)	Testing for LTBI is not recommended If necessary: Preferred: IGRA where available. Acceptable: Either IGRA OR TST For serial testing: Acceptable: Either IGRA OR TST Consider repeat or dual testing where a nega- tive result from either would be considered negative ²	Programmatic concerns	

Summary of recommendations for testing for LTBI (figure 2, 2017 IDSA guidelines)

LTBI = latent tuberculosis infection Mtb = Mycobacterium tuberculosis IGRA = interferon-y release assay TST = tuberculin skin test

JAMA study with CDC (April 2024)

- Prospective diagnostic trial of 22,000 subjects at high risk for TB infections
- Compared TST & the two IGRA

Table 3. Incremental Value Gained by Second Test: Change in PPV			
Test Aª	Test B	Incremental change estimate in PPV (95% CI) ^b	
TST⁺	QFT [−] GIT ⁺	1.64 (1.40-1.93)	
TST⁺	QFT ⁻ GIT ⁻	0.36 (0.18-0.75)	
TST⁺	TSPOT ⁺	1.94 (1.65-2.27)	
TST⁺	TSPOT ⁻	0.29 (0.13-0.65)	
QFT [_] GIT ⁺	TST ⁺	1.10 (0.98-1.24)	
QFT ⁻ GIT ⁺	TST ⁻	0.54 (0.19-1.59)	
QFT [_] GIT ⁺	TSPOT ⁺	1.24 (1.09-1.41)	
QFT ⁻ GIT ⁺	TSPOT ⁻	0.39 (0.13-1.12)	
TSPOT ⁺	TST⁺	1.10 (1.02-1.18)	
TSPOT ⁺	TST ⁻	0.31 (0.05-2.13)	
TSPOT ⁺	QFT ⁻ GIT ⁺	1.08 (1.00-1.17)	
TSPOT ⁺	QFT ⁻ GIT ⁻	0.34 (0.05-2.33)	

Results of the second test are highlighted in **red** if it was **positive** and **green** if it was **negative** (Table 3, Ayers 2024)

Learning points & take aways

Learning points & take aways

- Nocardia is a G(+) branching rod, often **found in soil**, that can cause infections in immunocompromised hosts
- Common sites of infection are **skin** and **lungs**, with a high predisposition to go to the **CNS**
- **Bactrim** is a key back bone in **multidrug therapy**, which should be **tailored to susceptibilities**
- Despite prolonged treatment (up to one year), high rates of recurrence
- Nontuberculous mycobacterial (NTM) is everywhere, including water and soil
- Epidemiological studies show higher rates in the **south/gulf coast**
- Some NTM (e.g. *M. ulcerans* & *M. marinum*) have unique associations (e.g. fish tanks)
- NTM can cause **false positive PPDs** (*M. marinum*)
- If positive PPD (and low pretest probability), **repeat IGRA** may be of some utility



Sources

