A Hare Unusual: Francisella tularensis Peritonitis Following Orthotopic Liver Transplantation

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INTRODUCTION

Francisella tularensis

- Causes the rare zoonotic disease tularemia
- Transmitted following arthropod bite, inhalation, or ingestion of contaminated water or soil.¹

Only a couple hundred cases are reported annually by the CDC, with very few reported cases in transplant recipients.¹

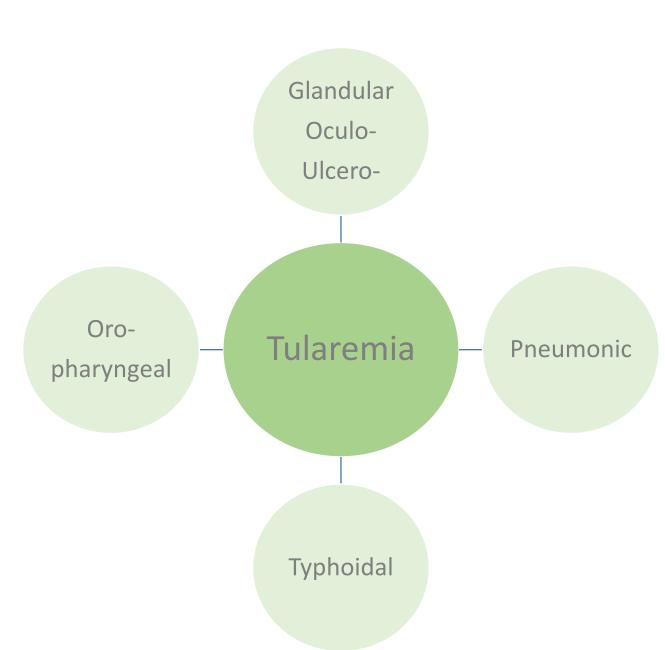


Figure 1: Types of Tularemia.

CASE PRESENTATION

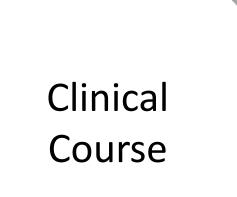
A 59-year-old male who was 79 days post orthotopic liver transplant (OLT) secondary to non-alcoholic steatohepatitis cirrhosis presents with acute onset generalized weakness, abdominal pain, and diarrhea.

Liver Transplant & Presentation

<u>Liver Transplant</u>: Donor - PHS increased risk given (+) Hepatitis C status Complications: underlying CKD → required hemodialysis mmunosuppression: Tacrolimus, prednisone taper, mycophenolate mofetil Symptoms: acute onset of weakness, abdominal pain, diarrhea, fever (103.2F) Physical Exam: abdominal tenderness, distended, fluid wave, (-) jaundice, (-) icterus

Diagnostic Information Labs: WBC 8.2x10³/μL (96% neutrophils), Hepatic panel normal Imaging: CT Abdomen with diffuse distension & small foci of air adjacent to small bowel with pneumatosis, right-sided adenopathy within abdomen **Endoscopy**: EGD with esophageal candidiasis

Paracentesis: 2,850 PMNs, Gram-negative coccobacillus



- → Empiric broad-spectrum antimicrobials cefepime, vancomycin, acyclovir, micafungin
- → Other infectious workup negative (blood & urine cultures, LP, fungal & viral
- \rightarrow No improvement after 72 hours, persistent fever \rightarrow repeat paracentesis (+) stain
- → Transitioned to ciprofloxacin + metronidazole with clinical improvement

FINALIZING THE DIAGNOSIS & TREATMENT

Ascitic fluid culture later speciated by the Ohio Department of Health as Francisella tularensis, confirmed by microagglutination and direct fluorescence antibody staining.

When asked about exposures, the patient revealed he was a farmer. He had recently runover rabbits with his tractor, requiring handling and sanitization of contaminated equipment.

for 21 days, both fevers and ascites resolved.



LITERATURE REVIEW & CASE COMPARISON

Table 1: Literature review of tularemia in solid organ transplantation including liver transplant recipients, modified from Bahuaud et al.²

	Transislant			Antibiotic	
Patient	Transplant History	Clinical Presentation	Diagnosis	Antibiotic Therapy	Ref.
Male, 50 yrs old	3 yrs post-liver transplant for Hep C & EtOH cirrhosis.	 Fever, pneumonia, myalgias for 72 hours 	 Blood cultures initially negative Positive for FT at 72 hours 	Levofloxacin 500mg/day for 21 days	Limaye, Hooper (1999) ³
Male 69 yrs old USA Handled hay	4 yrs post- kidney transplant for polycystic kidney disease	 Fevers, chills, fatigue, neck stiffness, vomiting, diarrhea X-ray with left-sided pneumonia 	 Blood cultures positive initially misidentified Corrected to FT at 7 days 	Doxycycline for 14 days	Khoury et al (2005) ⁴
59 yrs old USA	11 yrs post- kidney transplant for polycystic kidney disease	 Persistent fever Chest CT with multiple nodules 	 BAL with lymph node biopsy: cultures grew FT in 7 days 	Fluoroquinolone (unknown dosage and duration)	Mittal- henkle, Norman (2005) ⁵
Male 69 yrs old USA Hunting Dogs	15 yrs post- kidney transplant for IgA nephropathy	 Fevers, chills, cough, sputum production X-ray bilateral infiltrates 	 Blood cultures positive for FT after 10 days 	Levofloxacin 500mg/day for 14 days	Faucon et al (2011) ⁶
24 yrs old Turkey	12 months post-kidney transplant	 Fever, cervical adenopathy Lymph node pathology: chronic necrotizing granulomatous inflammation 	 PCR of lymph node positive for FT Confirmed by serology later. 	Doxycycline 100mg BID for 4 weeks	Ozkok et al (2012) ⁷
Male 51 yrs old France	7 yrs post-liver transplant	 Septic shock, ARDS, ketoacidosis CT-scan mediastinal adenopathy + bilateral nodular lesions 	 Blood culture positive after 5 days FT strain confirmed with sequencing 	Ciprofloxacin 500 mg BID for 14 days	Bahuaud et al (2015) ⁸
Male 64 yrs old	4 years post- heart transplant	 Fever, chills, night sweats, cough, respiratory distress CT-scan: pleural effusion & mediastinal adenopathy PET-scan: increased uptake pulmonary nodules 	 Pleural fluid cultures negative PCR and cultures of lymph node biopsies positive for FT 	Ciprofloxacin 750 mg BID + gentamicin 300 mg for 7 days, then ciprofloxacin for 14 days	Bahuaud et al (2016) ⁸

KEY POINTS FROM TABLE 1

- 1) Pneumonia from FT is the most common post-transplantation
- 2) Fever & mediastinal or abdominal lymphadenopathy frequently seen
- 3) Abnormal features include small bowel pneumatosis, necrotizing granulomatous

CONCLUSIONS

We present the reported first case of *Francisella tularensis* peritonitis postorthotopic liver transplantation.

The presentation of Tularemia amongst immunocompromised hosts is often atypical, which makes diagnosis challenging. Per literature review (Table 1), tularemia usually presents in the pneumonic form in patients who have undergone solid organ transplantation. However, our patient with a liver transplant presented with the typhoidal form with primarily gastrointestinal symptoms.

Consider Tularemia in your differential diagnosis when:

- Persistent fevers and infectious symptoms in immunocompromised hosts without clear diagnosis
- Social history supporting animal exposures, particularly hares, rodents, or ticks
- Common gastrointestinal manifestations: abdominal pain, nausea/vomiting, diarrhea
- Unique features: abdominal lymphadenopathy on imaging, pneumatosis
- Lack of improvement on broad-spectrum antibiotics, particularly cephalosporins

Why? Two native β-lactamase genes (bla1 and bla2)9

Antimicrobial management: Choose effective antibiotics

- Aminoglycosides streptomycin, gentamycin (10-14 days)
 - preferred for severe cases
- Fluoroquinolones (10-14 days)
 - multiple case reports of success in SOT patients (Table 1)
- Doxycycline (14-21 days)¹

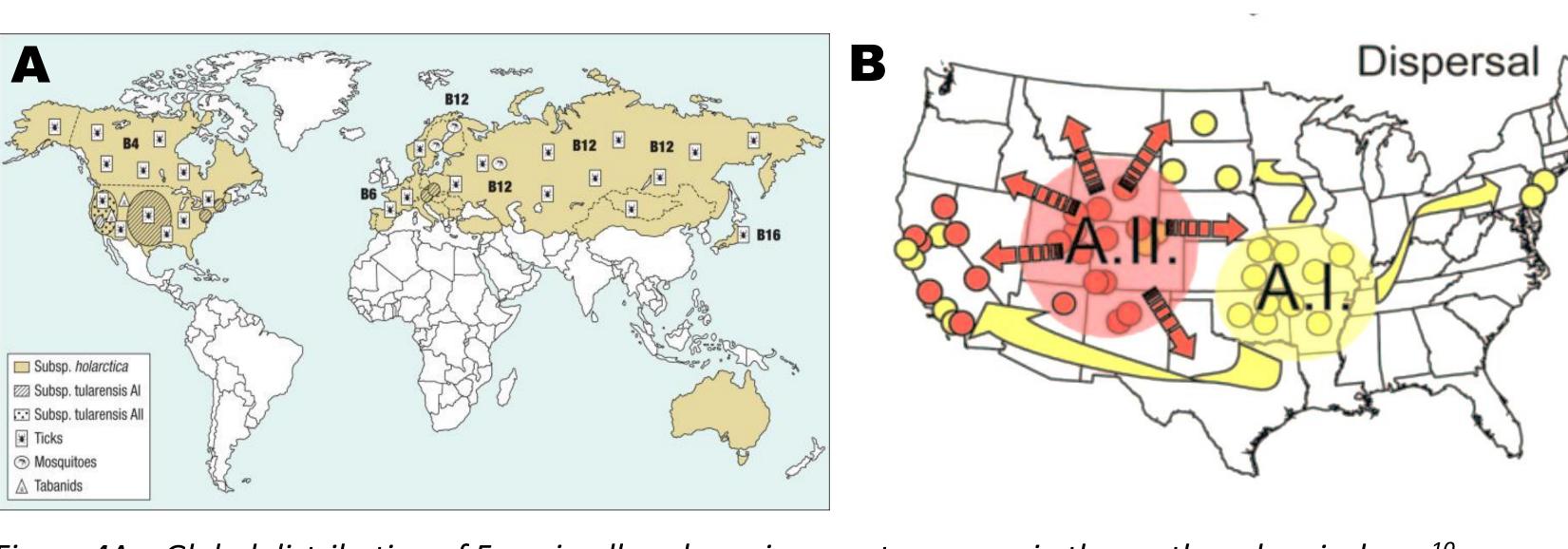


Figure 4A – Global distribution of Franciscella subspecies, most common in the northern hemisphere. 10 4B – National dispersal F. tularenesis subpopulations AI and AII, the two most common in the United States, demonstrating spread to previously unidentified locations. 11

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Disclosures: All authors have nothing to disclose.



inflammation With transition to ciprofloxacin 500mg twice daily