Mycobacterium obuense Bacteremia in a Patient with Pneumonia

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Mycobacterium obuense is a pigmented, rapidly growing mycobacterium. Because it has been considered nonpathogenic, M. obuense is being investigated in clinical trials of cancer immunotherapy and bioremediation. We report a case of bacteremia caused by M. obuense in a patient with pneumonia, showing its potential pathogenicity.

A pproximately 75 species of rapidly growing mycobacteria (RGM) have been isolated from soil, animals, and water (1). The RGM Mycobacterium obuense, an environmental pigmented mycobacterium, is mobile and easily adaptable to the environment and possesses oxygenases that enable it to degrade organic compounds and dechlorinate methoxychlor-based insecticides (2). Until recently, M. obuense has been considered nonpathogenic. We report a case of bacteremia caused by M. obuense.

A 29-year-old man from a rural community in Puebla, Mexico, arrived at an emergency department in Mexico City reporting a 2-day history of chest pain, dyspnea, and fever. On physical examination, his heart rate was 94 bpm, blood pressure 175/88 mm Hg, temperature 38.5°C, and peripheral oxygen saturation 70%. Chest auscultation revealed bibasilar fine crackles and signs of pleural effusion.

The patient was a farmer; had been in close contact with pigs, sheep, and cows; and reported consuming unpasteurized dairy products. He had a history of diabetes mellitus with chronic kidney disease categorized as stage G4 A3 (glomerular filtration rate 16.7 mL/min/1.73 m²; proteinuria >2.8 g/d) of the KDIGO classification (Kidney Disease: Improving Global Outcomes, https://kdigo.org) without replacement therapy. He reported taking metformin, amlodipine, furosemide, and iron sulfate.

At admission, laboratory test results included leukocyte count, 11,900 cells/µL with 88.2% neutrophils; C-reactive protein, 250 mg/L; procalcitonin, 17 ng/mL; creatinine, 4.0 mg/dL; and arterial blood gases, pH 7.24, pO₂ 40.8 mm Hg, pCO₂ 34.8 mm Hg, lactate 2.9 mmol/L, HCO₃⁻ 14.6 mmol/L, and SO₂ 71% on ambient air. Findings of a computed tomography scan of the chest suggested that the patient had a lung infection (Figure).

The patient began empirical treatment for community-acquired pneumonia with ceftriaxone and clarithromycin. No respiratory samples were obtained because the patient was unable to produce sputum. We performed blood cultures in Aerobic/F medium (Becton Dickinson, https://www.bd.com). After a 7-day incubation period, we detected growth and observed gram-positive bacilli in the gram stain (Appendix Figure, panel A, http://wwwnc.cdc.gov/EID/article/25/5/18-0208-App1.pdf). We made subcultures on sheep blood, chocolate, and Sabouraud agar and performed Kinyoun and Ziehl-Neelsen stains, in which we observed partially acidic alcohol-resistant coccobacilli (Appendix Figure, panels B,C). After 2 weeks of incubation at 35°C–37°C, we observed rough, mucoid colonies (Appendix Figure, panels D,E).

We attempted to identify the bacilli by using GenoType Mycobacterium (Hain Lifescience, https://www.hain-lifescience.de) but were unsuccessful because no species-specific probe is available. We performed amplification and sequencing of the 16S rRNA gene (498 bp) and hsp65 gene (400 bp). Pairwise sequence aligned 100% with 16S rRNA and 99.6% with hsp65 to the sequences of M. obuense strain CIP 106803 (GenBank accession no. AF547954.1).

The patient was started on intravenous amikacin, clarithromycin, and moxifloxacin as soon as we notified clinicians of RGM isolated from his blood cultures. We conducted susceptibility testing by broth microdilution, which showed susceptibility to all antimicrobial drugs tested except tobramycin (intermediate susceptibility) (Appendix) (3).

The patient completed 2 weeks of intravenous moxifloxacin, amikacin, and clarithromycin and was discharged. He received 4 additional weeks of oral azithromycin and moxifloxacin and experienced complete resolution.

M. obuense, first described as a scotochromogenic RGM species isolated primarily from soil >40 years ago in Obu, Japan, is a catalase-positive, peroxidase-negative bacillus that can degrade salicylates, forming a black product (4,5). In culture, M. obuense has 2 morphotypes, smooth and rough variants. In the smooth variants, its cell wall contains long-chain saturated fatty acids that enable it to colonize the environment and are responsible for the pleomorphic observation on the surface of solid agars (6). Although this species was later isolated from sputum samples from patients with apparent pulmonary disease, no additional clinical data were reported, and thus, no evidence of pathogenicity was established (4,7).

Phylogenetic analysis of M. obuense shows close association with M. chubuense (81.3% identity) and M. rufum (92.2% identity) (7). Its genome consists of 5,576,960 bp...
Although these mycobacteria are considered nonpathogenic, they have homologous genes to mammalian cell entry that encode proteins involved in virulence and cell invasion. In addition, M. obuense contains genes involved in antimicrobial resistance, such as marA, aminoglycoside-resistance protein kinase, β-lactamases, and monoxygenases, which confer M. obuense with intrinsic rifampin resistance (2,7).

Establishing etiology in this case was challenging because current nucleic acid probe assays cannot identify M. obuense correctly. Clinicians should avoid discarding RGM or misclassifying these isolates as colonizers until definitive species identification confirms etiology.

M. obuense has been evaluated as an adjuvant immunotherapy in phase I and II trials on patients with melanoma, pancreatic cancer, and colorectal cancer, with promising results. This treatment consists of an intradermal application of a suspension of heat-killed whole cell M. obuense (8–10).

The isolation of M. obuense from blood cultures of a patient with community-acquired pneumonia suggests its capacity for virulence and invasiveness in humans. Because M. obuense might become an adjuvant in cancer therapy, researchers should ensure implementation of proper, standardized inactivation protocols.

About the Author

Dr. Luis is a fellow in infectious diseases and Dr. Ponce-de-León is a senior researcher at the National Institute of Medical Science and Nutrition Salvador Zubirán, Mexico City. Their research fields are mycobacterial diseases and clinical microbiology.

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Appendix

We conducted susceptibility testing on the bacilli by broth microdilution, which showed MICs to amikacin, 0.25 µg/mL; clarithromycin, ≤0.03 µg/mL; ciprofloxacin, 0.03 µg/mL; doxycycline, 0.5 µg/mL; imipenem, 2 µg/mL; linezolid, 0.5 µg/mL; moxifloxacin, ≤0.015 µg/mL; tobramycin, 4 µg/mL; and trimethoprim/sulfamethoxazole, 0.25/4.8 µg/mL. According to Clinical Laboratory Standards Institute guidelines (3), all antimicrobial drugs tested were susceptible except tobramycin, which was intermediate.
Appendix Figure. A) Gram stain from positive blood culture showing gram-positive bacilli. Original magnification × 100. B) Kinyoun stain from sheep blood agar with red coccobacilli, some in clusters. Original magnification × 100. C) Ziehl Neelsen stain from sheep blood agar showing partially acid-fast stained coccobacilli. Original magnification × 100. D) *Mycobacterium obuense* on Sabouraud agar showing mucoid and smooth colonies with yellow pigmentation. E) *M. obuense* after several passages showing smaller, dry and rough colonies.