

Concern About Possible Stroke, Loss of Left Arm Control, Numbness and Tingling in Both Legs, in Last 90 Minutes

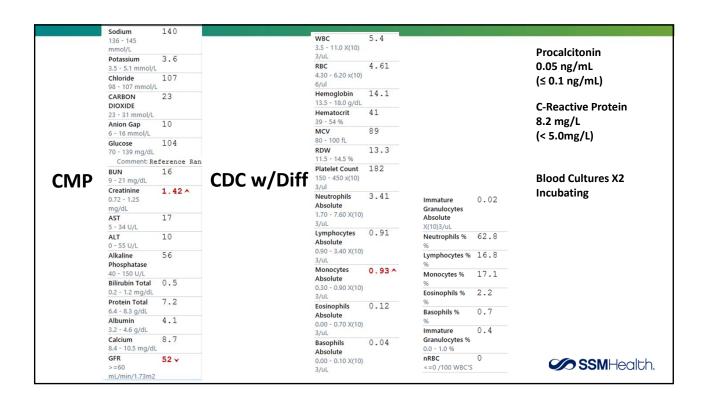
- 64 y/o ♂ presents to the ED
- House painter, lives alone
- Sarcoidosis on hydroxychloroquine, low dose prednisone
- Quit smoking 2018 (after 37 years)
- · Generally in fair health



Patient Workup

- EKG
- Chest x-ray
- CT head
- Procalcitonin
- C-reactive protein
- CMP
- CBC
- Blood Cultures x2





Diagnostic Procedures

Chest X-ray

Questionable bilateral infiltrates c/w edema or infection

EKG

• No evidence of arrhythmia or ischemia

CT of head

3 cm, R temporal lobe brain mass with edema and mass effect



Patient Management

- Admitted
- Decadron IV for cerebral edema



Hospital Course

- ID started meropenem and vancomycin/concern for abscess in brain.
- Neurosurgery to perform craniotomy for evacuation of the mass.

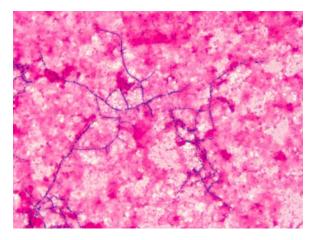


Surgical Outcome

- Temporal mass found to be an abscess
- 3 mL of fluid sent for cultures and stains
 - Bacterial culture and Gram stain
 - AFB culture and Auramine/Rhodamine stain
 - Fungal culture



Gram Stain of Brain Abscess submitted for culture

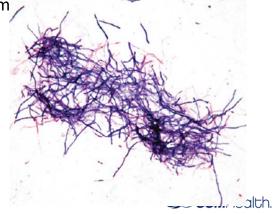


Heavy PMNs Few Gram positive bacilli filamentous and branching



Brain Abscesses Culture

- Small dry looking colonies after 2 days on BAP
 - Gram stain of colonies beaded Gram positive rods



Organism Identification

Routine Spotting for MALDI-TOF MS Identification Using the Vitek MS

Method Name	VITEK® MS / VITEKMSACQ01
Organism Name	Nocardia farcinica 99.9 %
Basaltina Basanata at	0/7/04 0:40 ANA



Nocardia farcinica sub-culture





Hospital Course Summary

- Hospitalized for 5 days
- Blood Cultures no growth after 5 days
- Surgery for right temporal craniotomy, 3 mL of abscess fluid removed
- Abscesses grew *N. farcinica* after 2 days
- Went home



Treatment For Patient's Brain Abscess

After Gram Stain started on

Imipenem and TMP/SMX

Final Report

 Discharged on imipenem and TMP/SMX, treatment duration at least 12 months, first 6-12 weeks IV, guided by repeat brain MRI results.

SEE NOTE



9 days after discharge susceptibility report received.

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Comment: Nocardia farcinica
Organism identified by client
INTERPRETIVE INFORMATION: Gram Positive Rod Susceptibility
Units = micrograms/mL
Susceptibility testing is performed by CLSI-approved broth
microdilution method using custom-made MIC panels.
AFBMIC
Trimethoprim/Sulfamethoxazole
                                     2/38 Suscept
Ciprofloxacin
                                     0.5 Suscept
Moxifloxacin
                                     <=0.25 Suscept
Amikacin
                                     <=1 Suscept
Doxycycline
                                      4 Intermed
Clarithromycin
                                     >=32 Resist
Linezolid
                                     4 Suscept
                                     16 Resist
Amoxicillin/Clavulanate
                                    8/4 Suscept
Ceftriaxone
                                     64 Resist
Minocycline
                                      2 Intermed
Tobramycin
Performed by ARUP Laboratories,
500 Chipeta Way, SLC,UT 84108 800-522-2787
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www.aruplab.com, Tracy I. George, MD, Lab. Director



Treatment For Patient's Brain Abscess

After Gram Stain started on

- Imipenem and TMP/SMX
- Discharged on imipenem and TMP/SMX, treatment duration at least 12 months

After susceptibility test results

Linezolid and TMP/SMX

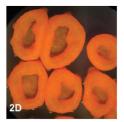


Nocardia species

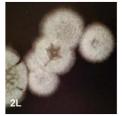
- Grouped in with the aerobic actinomycetes (many genera and species)
- Widely distributed in the environment, soil and aquatic habitats
- Nocardia aerobic actinomycetes most commonly causing infections in humans
- About 100 validly named species of Nocardia
- Direct Gram stain generally appear as very long, branching, thin, and beaded Gram positive rods
- · Weakly positive in modified acid-fast stain
- Infections generally caused by trauma, or from inhalation (particularly in immunocompromised patients), hematogenous spread from lungs to other body sites
- Brain is one of the most common secondary sites of infection



Nocardia spp., Colony Morphology on Sab. Dex. Agar



N. brasiliensis - 10 days



N.cyriacigeorgica – 10 days



N. otitidiscaviarum – 7 days



N. farcinica – 5 days (BAP)



N.carnea – 25 days SSMHealth.

Nocardia, Rhodococcus, Gordonia, Actinomadura, Streptomyces, and Other Aerobic Actinomycetes PATRICIA S. CONVILLE, BARBARA A. BROWN-ELLIOTT, AND PATRICIA S. ONTEISEN, G. WITEISEN,

Chapter 30, Manual of Clinical Microbiology, 2019

Nocardia farcinica

- Old N. asteroides drug pattern V
- Well known to cause disseminated infections
- Lungs are the most common site of involvement, but also found in blood, brain abscesses, keratitis, and muscle abscesses
- Most individuals with this infection are immunocompromised



Brain Abscess

- Most common in 30-50 year old males
- Direct spread from a contiguous site (otitis media, sinuses, mastoiditis, dental infection) accounts for 20 60 percent of cases, single abscess
- Hematogenous spread from lungs, skin, pelvic infections, and other sites, multiple abscesses
- Most frequent causes of brain abscesses are Staphylococcus spp. and Streptococcus spp., S. aureus and viridans Streptococcus being the most common
- Other organisms associated with brain abscesses include *Bacteroides spp.*, *Prevotella spp.*, Gram negative enteric bacteria, rarely *Nocardia spp.*



Shortness of breath (SOB), Leg Rash, Diarrhea, Cardiac Evaluation

- 52 y/o ♂ transfer from OSH
- Nonsmoker
- History of hypertension, anxiety, and heart murmurs (family history of heart murmurs)
- Grew up on a farm
- Works two jobs, single, reports having unprotected sex with men



Physical Examination

- Temp 99.8
- Pulse 104
- Heart, holosystolic murmur
- Rash on left upper leg, resolving



Patient Workup

- TEE
- Chest x-ray
- CT chest
- Enteric Panel PCR, C. difficile toxin, Giardia/Crypto Ag
- CMP
- CBC
- Procalcitonin
- Blood Cultures x 2 (with 2 follow up sets ordered next day)



	Sodium	128 🗸				
	135 - 145 mmol/L					
	Potassium	5.0 ^	Osmolality Calculated	273 ∨		
	3.4 - 4.8 mmol/L		275 - 295 mosm/kg			
	Chloride	97	Globulin	5.3^		
	97 - 108 mmol/L		2.0 - 4.0 g/dL			
	CO2	18 🗸	A/G Ratio 1.3 - 2.8 Creatinine Based eGFR >60 mL/min/[1.73_m2]	0.5~		
	23 - 32 mmol/L			0.01		_
	Glucose	96		47 V	BC	
	70 - 99 mg/dL					
	BUN	33 ^			WBC	9.87
	8 - 23 mg/dL				3.50 - 10.70 10*3/uL	
	Creatinine	1.65 ^			RBC	3.49~
СМР	0.50 - 1.20 mg/dL				4.42 - 5.95 10*6/uL	0.10
	Calcium	7.8 🗸			HGB	9.4~
	8.5 - 10.4 mg/dL	7.8			13.7 - 17.7 g/dL	
	Total Protein	7.8			нст	29.8
	5.8 - 8.3 g/dL Albumin				40.6 - 52.6 %	
	3.6 - 5.4 g/dL	2.5 🗸	Component Ref Range & Units Troponin I <0.05 ng/mL	0.03	MCV	85.4
	Bilirubin Total	0.6			85.0 - 98.4 fL	
	0.3 - 1.3 mg/dL	0.0			МСН	26.9
	Alkaline Phosphatase	175^	Resulting Agency Component Ref Range & Units Procalcitonin <0.10 ng/mL	ST MARY'S HOSPITA	28.5 - 33.5 pg	
	40 - 129 U/L	1/5^			MCHC	31.5 🗸
	AST	350 ^			32.0 - 36.5 g/dL	
	0 - 40 U/L	330 ^			Platelets	158
	ALT	219^			150 - 425 10*3/uL	No. 2 1 2 10
	0 - 41 U/L	219^			RDW-CV	17.6^
	Anion Gap	13			11.5 - 15.0 %	0.04
	7 - 14 mmol/L				NRBC Absolute 10*3/uL	0.04
	BUN/Creatinine Ratio	20.0			MPV	10 7 .
	8.0 - 26.0				8.4 - 12.4 fL	12.7^



Hospital Procedures

Started on ceftriaxone and vancomycin for cellulitis of lower leg.

Chest x-ray showed enlarged cardiac silhouette and mild to moderate airspace opacity in the right lung base with a small right pleural effusion.

CT chest PE protocol: showed bilateral pulmonary emboli with small sub-segmental pulmonary emboli in the left upper lobe and superior segment of the right lower lobe, small to moderate right pleural effusion, bilateral airspace opacities somewhat ground glass and morphology, and hilar/mediastinal lymphadenopathy.

Transesophageal echocardiogram (TEE): showed severe mitral valve regurgitation from prolapse anterior leaflet with apparent torn chordae tendon, severe pulmonary hypertension, significantly dilated IVC, severe bilateral enlargement, thickened aortic valve with mild-to-moderate regurgitation, small mitral and aortic vegetation, and normal ejection fraction at 60%.

Patient diagnosed with acute heart failure secondary to mitral regurgitation and endocarditis



Blood Cultures not growing: Culture negative endocarditis workup

- Q Fever Antibody
- Legionella Antibody Panel
- Tropheryma whipplei DNA
- Aspergillus Galatomannan Antigen Blood
- Beta-D-Glucan (1,3) (Fungitell
- Chlamydia Antibody IgG/IgM Panel
- · Bartonella Antibody Panel
- HIV-1 HIV-2 Antibody + HIV p24 Ag
- Fungal Antibody Panel Immunoduffusion
- Brucella Ab IgG/IgM Panel
- RPR
- Hepatitis Ab Panel ABC



Blood Cultures not growing: Culture negative endocarditis workup

- Q Fever Antibody No Ab detected, < 1:16
- Legionella Antibody Panel Positive
- Tropheryma Whipplei DNA Not detected
- Aspergillus Galatomannan Antigen Blood Neg.
- Beta-D-Glucan (1,3) (Fungitell) 95 pg/mL positive (> 80 pg/mL)
- Chlamydia Antibody IgG/IgM Panel *C. pneumoniae, C. trachomatis, C. psittaci* IgM <1:20 *C. pneumoniae, C. trachomatis, C. psittaci* IgG 1:256
- Bartonella Antibody Panel B. henselae IgG 1:1,024
- HIV-1 HIV-2 Antibody + HIV p24 Ag Reactive HIV-1 Ab (HIV-1 viral load 14,000 copies/mL)
- Fungal Antibody Panel Immunoduffusion None Detected
- Brucella Ab IgG/IgM Panel Neg. IgM, Reactive IgG, Neg. confirmatory test
- RPR non-reactive
- Hepatitis Ab Panel ABC Neg., Neg., Neg.



Hospital Procedure

After *B. henselae* serology patient treated with IV ceftriaxone, doxycycline, and IV gentamicin.

Patient treated with Biktarvy for HIV.

Aortic and Mitral value replacement

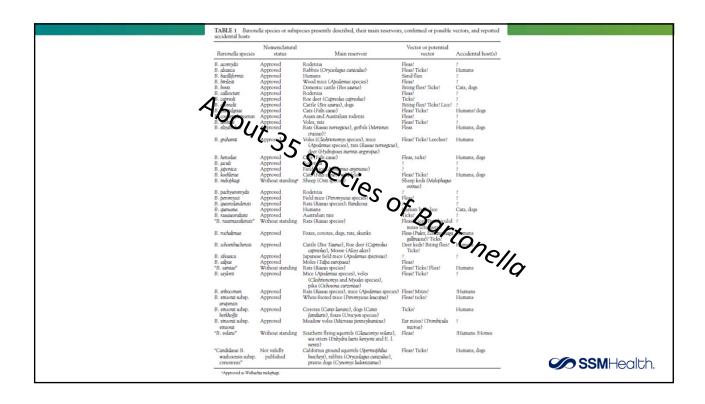
Removed valves sent for Gram stain/culture, fungal culture, AFB culture, and broad-spectrum 16S PCR.

Culture and Gram stain - Heavy PMNs, no bacteria seen, ng after 6 days Fungal culture - no fungus isolated

AFB culture + smear - No AFB seen, no growth after 8 weeks 16S PCR – *B. henselae* DNA detected, both valves

Confirmed B. henselge endocarditis





Bartonella species are short, Gram-negative coccobacilli, fastidious, slow-growing bacteria, and cultures should be held for a minimum of 21 days.

Transmission

Bartonella species are typically transmitted by insect vectors, such as fleas, sand flies, body lice, and potentially ticks, biting flies, and keds (wingless flies).

Transmission also occurs by animal scratches or possibly bites.



Nineteen *Bartonella spp.* and subspecies are infectious to humans and can elicit a wide spectrum of diseases, including fever of unknown origin, cat scratch disease (CSD), cutaneous vasculitis, endocarditis, myocarditis, osteomyelitis, bacillary angiomatosis, bacillary peliosis hepatitis, and granulomatous inflammatory Disease.

Most Well Known Diseases Caused by Bartonella spp.

CSD - Bartonella henselae Trench Fever - Bartonella Quintana Carrión's disease - Bartonella bacilliformis Subacute endocarditis – many Bartonella spp.



Bartonella spp. are the second most common cause of culture-negative endocarditis behind Coxiella burnetii



B. henselae CSD - typically a self-limiting infection in immunocompetent individuals, particularly children, but which can occasionally be complicated by neuroretinitis, granulomatous hepatitis, osteomyelitis, fever of unknown origin, and blood culture-negative endocarditis.

Endocarditis has also been reported to be caused by *B. quintana*, *B. elizabethae*, *B. alsatica*, *B. koehlerae*, "Candidatus Bartonella mayotimonensis".



B. henselae is the predominant cause of CSD in humans, with B. grahamii being suspected in a few cases.

It is estimated that 12,500 human cases of CSD occur yearly in the United States, for an average annual incidence of 4.7 diagnoses/100,000 population.

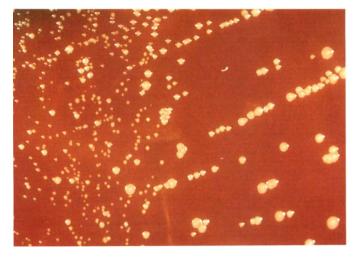
Children ≤ 14 years of age account for 33% of cases.

There is a seasonal pattern that varies by U.S. geographic region, with most cases occurring in fall and winter.

B. henselae and *B. quintana* are associated with 12-28% of culture-negative endocarditis cases worldwide.



Bartonella henselae – BAP, 35 C, 5% CO2, 5 days



Clinical Microbiology and Infection 1997 3677-689DOI: (10.1111/j.1469-0691.1997.tb00478.x)



Diagnosis

- CSD may be diagnosed presumptively in patients with typical signs and symptoms and a compatible exposure history. Serology can confirm the diagnosis, although cross-reactivity may limit interpretation in some circumstances.
- Bartonella endocarditis can be diagnoses using serological testing for B.
 henselae and B. quintana with titers ≥ 1:128 considered significant. Serum for
 plasma PCR testing for Bartonella can be useful as well as PCR testing of
 heart valve tissue.
 - B. henselae has been cultured from lymph node aspirates, and blood.
 - Since *B. henselae* is a fastidious, slow-growing bacterium, cultures should be held for a minimum of 21 days.



Treatment

The use of antibiotics to shorten the course of CSD is subject to debate. Most cases of CSD resolve without treatment. Azithromycin has been shown to decrease lymph node size more rapidly compared to no treatment.

Complicated Bartonella cases require antibiotic treatment. Effective antibiotics include penicillins, tetracyclines, cephalosporins, and aminoglycosides. Aminoglycosides are typically used as first-line treatment for complicated Bartonella infections because they are bactericidal. Complicated infections are commonly treated using combination therapy.



Corneal Surface Swelling – R eye S/P Revision of Operative Wound – R eye Corneal Transplant – R eye

- 94 y/o ♂ presents for eye follow-up visit
- Lives with daughter
- · Generally in good health



Patient Examination by Ophthalmologist

- · Pain right eye
- General vision not great
- Well appearing
- Alert, oriented, answers questions appropriately
- New R eye central corneal epithelial cell defect
- Using corticosteroid eye drops in R eye 2x day
- Possible Infectious Crystalline Keratopathy

Infectious crystalline keratopathy (ICK) is rare.



Past Medical History of R eye

10 months prior to this visit

Descemet's Stripping Automated Endothelial Keratoplasty (DSAEK)

Partial thickness cornea transplant procedure



Diagnostic Procedures

Corneal Scrapping

For Gram stain and bacterial/fungal cultures



Patient Treatment

- Moxifloxacin (0.5%) drops 6x day for R eye
- Fluorometholone drops 2x day (corticosteroid, anti-inflam.)
- Muro 128 drops 3x day (draws fluid out of cornea)
- Ketorolac drops as needed (nonsteroidal anti-inflam.)



Microbiology Test Results

Routine Bacterial Culture and Gram stain
Gram stain – Rare PMNs, No Bacteria Seen
Culture – No Growth after 2 days

Routine Fungal Culture

Heavy Growth on IMA noted after 3 weeks

Wet mount – rod shaped bacteria

Gram Stain – Beaded Gram Positive Rods



Organism Identification

Routine Spotting for MALDI-TOF MS Identification Using the Vitek MS

Method Name / Host(s)

VITEK® MS

VITEKMSACQ01

Organism Name

Mycobacteroides chelonae 99.9%



Infectious Crystalline Keratopathy

- Citrobacter,
- Acinetobacter,
- Alternaria,
- Acanthamoeba,
- Enterobacter,
- •Enterococcus species,
- Candida species,
- Serratia marcescens,
- •Gemella haemolysans
- Actinomyces species

- •alpha hemolytic viridans Streptococcus
- •Streptococcus Pneumoniae,
- •coagulase-negative Staphylococcus,
- Peptostreptococcus,
- •Haemophilus species,
- Mycobacterium species,
- Pseudomonas,
- Stenotrophomonas,



Final Report Mycobacteroides (Mycobacterium) chelonae Organism identified by client Susceptibility Results Organism: Mycobacteroides (Mycobacterium) chelonae Interpretation: RESISTANT Cefoxitin MIC (ug/mL): 128 Ciprofloxacin Interpretation: RESISTANT MIC (ug/mL): 4 Clarithromycin MIC (ug/mL): 0.12 Doxycycline Interpretation: RESISTANT MIC (ug/mL): >=32**Imi penem** Interpretation: RESISTANT MIC (ug/mL): 64 Interpretation: SUSCEPTIBLE Linezolid MIC (ug/mL): 4 Minocycline Interpretation: RESISTANT MIC (ug/mL): 8 Moxifloxacin Interpretation: RESISTANT MIC (ug/mL): 4 Tigecycline MIC (ug/mL): 0.25 Interpretation: SUSCEPTIBLE MIC (ug/mL): 2 S=Susceptible, I=Intermediate, R=Resistant, NonS=Nonsusceptible, IND=Indeterminate, SDD=Susceptibility is dose dependent, None=Interpretive guidelines are not available H=High, L=Low, *=Abnormal, C=Critical SSMHealth.

Myobacterium chelonae

- Nonchromogenic rapidly Growing Mycobacteria
- Found in the environment, tap water, surface water
- Member of M. chelonae/M. abscessus group with about 6 other species
- Causes primarily skin and soft tissue infections by direct inoculation, piercing wounds, contaminated tattoo inks, and can cause disseminated cutaneous disease
- 2012 study* found that for ocular infections due to *Mycobacterieum spp., M. chelonae* (45%) and *M. abscessus* (42%) were the most common causes

^{*} Brown-Elliott BA, Mann LB, Hail D, Whitney C, Wallace RJ Jr. 2012. Antimicrobial susceptibility of nontuberculous mycobacteria from eye infections. Cornea 31:900–906.



Treatment <i>M. chelonae</i> Eye Infectior	Treatment	<i>M.</i> c	chelonae	Eye	Infectior
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Topical therapy used initially in the absence of perforation, moxifloxacin (0.5%), clarithromycin (1%), tobramycin (1.4%).

Combination topical therapy is frequently used.



Questions?

Thank you!

