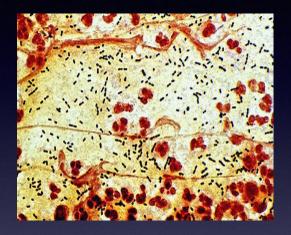
Clinical Microbiology

ACOI Board Review 2019 gerald.blackburn@beaumont.org

"No Disclosures"

S. aureus skin/soft tissue infection:





abscess (rather than diffuse cellulitis), w/ purulent drainage

common in diabetes

"Community-acquired" (CA)
 MRSA more likely scenario

spontaneous

body contact sports

hx of "spider bite"

Rx of MSSA bacteremia?

Nafcillin (side effect: AIN; dx: eosinophils in urine)

Rx if non-anaphylactic allergic rxn (rash) to PCN?

Cephalosporin - cefazolin

Rx if MRSA (if susceptible)?

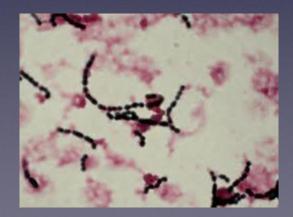
- Vancomycin
- Daptomycin (but <u>NOT pneumonia</u>)
- Linezolid (but <u>NOT bacteremia</u>)
- Trimethoprim/sulfa especially SSTI
- Clindamycin especially SSTI
- Quinolones (?) not Cipro

Rx of acute parotiditis (S. aureus most common); assoc. w/ surgery, dehydration, mouth breathing

Typical streptococcal SSTI (Grp A, B, G):

- Intense erythema
- (Often recurrent) cellulitis rather than abscess
- Lymphangitis
- Often (preceding) systemic symptoms
- Areas of pre-existent lymphedema, venous insufficiency
- Drainage, when present, often watery or serous





Erysipelas

(superficial cellulitis of Streptococcal etiol.)



Rx: penicillins if allergy: cephalosporin; vancomycin; clindamycin <u>NOT</u> trimethoprim/sulfa

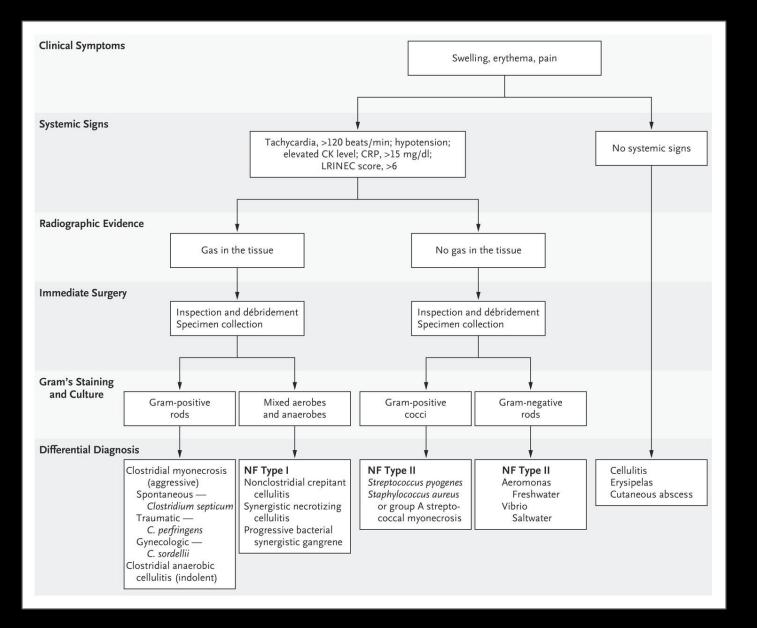


Additional Streptococcal Syndromes:

- "Viridans" strep bacteremia endocarditis
- S bovis/gallolyticus bacteremia GI malignancy
- Grp A strep necrotizing fasciitis "flesh eating bacteria" Rx: Beta-lactams + <u>clindamycin</u>
- TSS source usually obvious; often + blood culture (as can S. aureus - but....source often not obvious; blood cultures <u>negative</u>)
- Note:1/3 of Grp B strep are resistant to clindamycin (e.g. diabetic foot infection)

Necrotizing Soft Tissue Infections

- Group A strep: <u>pain out of proportion</u> to initial clinical findings
- Clostridium perfringens: progression over hours
- Mixed flora, most always including anaerobes
 - Most commonly in diabetics; obesity
 - Wounds involving/crossing mucous membranes
 - Foul odor
 - Mixed flora on gm stain
 - Delayed or no growth on culture



Stevens DL, Bryant AE. N Engl J Med 2017;377:2253-2265.



Diabetic Foot

• Usually polymicrobial, w/ foul odor -

- Anaerobes
- Gm negatives
- Many feel pseudomonas commonly involved

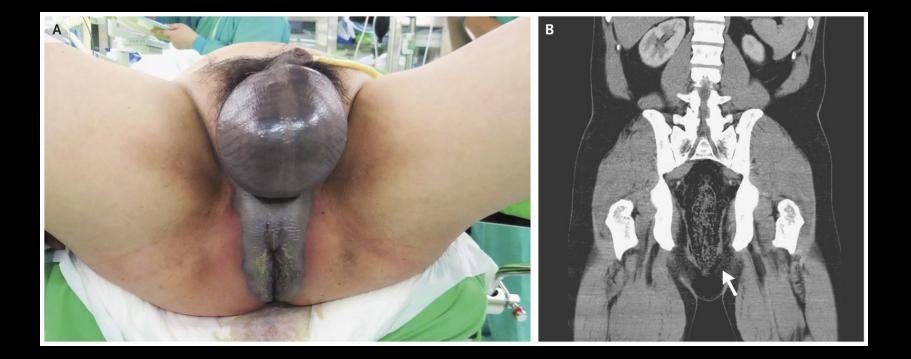
Rx (including pseudomonas coverage)?

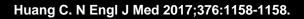




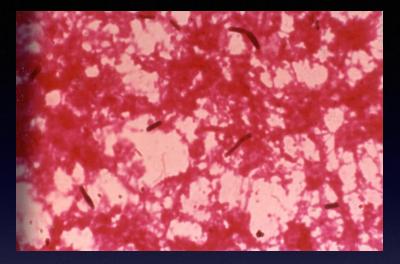


Hx, Gm stain, speed of progression, location of wound, useful in predicting organism(s) Now what?

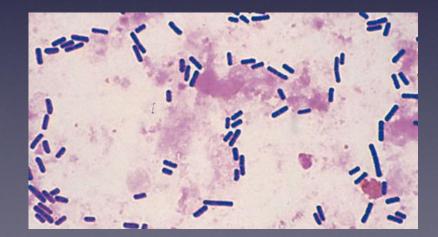








C. perfringens "gas gangrene" Rx: Surgery + Penicillin + Clindamycin + ? IVIG, HBO?



C. perfringens -> gas gangrene

- C. septicum bacteremia/sepsis
 - GI/gyn malignancy (sometimes occult),
 - chemotherapy-induced neutropenia
- C. botulinum -> skin popping w/ "black tar" heroin

Cat Bite

- Pasteurella multocida
- Rapid onset painful, throbbing, cellulitis
- gm neg rod
- RX:
 - amoxicillin +/-clavulanate
 - cefuroxime
 - doxycycline
 - quinolones
 - <u>NOT CEPHALEXIN</u>
 <u>(Keflex®)</u>



Additional "Pearls" re: Bites

- Dog bites/splenectomy: overwhelming sepsis due to Capnocytophaga sp. (DF-2)
- Human bites: Eikenella can't use clindamycin
- Snake bites: gm negs
- Rabies any wild carnivore
 - most common domestic animal (in U.S.)? cat
- Lagomorphs don't get rabies (exception: woodchucks)

Aeromonas hydrophila

- Gm neg rod
- Fresh water injuries, medicinal uses of leeches

Rx:

- fluoroquinolone
- 3rd gen cephalosporin
- trimethoprim/sulfa







"Nodular" lymphangitis

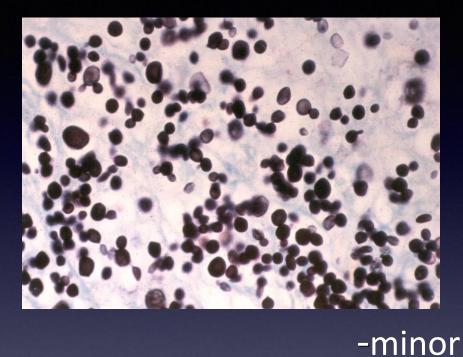
Working in yard: Dx? Tx?



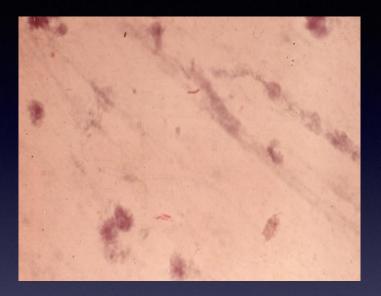
D/D nodular lymphangitis

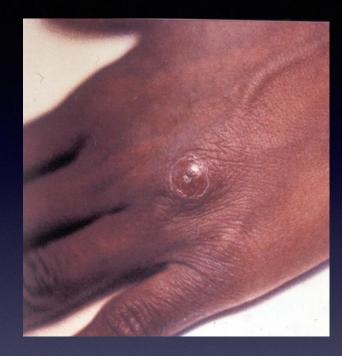
- Staph, strep
- Sporotrichosis: 1° lesion is painless
- Nocardia: 1° lesion is a tender ulceration
- M. marinum: 1° lesion is a tender papule
- Tularemia: 1° lesion is a painful ulceration, w/ <u>systemic symptoms</u> (classically associated w/ skinning rabbits)





Sporotrichosis: trauma from roses or sphagnum moss -variable size yeast cells w/ multiple buds Tx: itraconazole





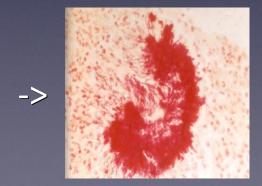
Mycobacterium marinum: cleaning fish tanks, water injuries, fish hooks, splinter from a boat

Rx:

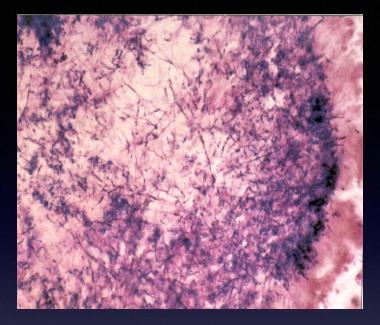
- Clarithromycin
- Doxycycline
- Minocycline
- Rifampin/ethambutol

Actinomycosis

- Spontaneous drainage from head ("lumpy jaw"), neck, or chest
- Often dental or oral mucosal origin
- Indolent, "wooden" mass effect; often confused w/ malignancy
- Assoc. w/ IUD's
- "sulfur granules"







Actinomyces:

- Gm + <u>anaerobic</u>, filamentous, beaded rods
- <u>not</u> acid fast (vs

Nocardia: aerobic, weakly acid fast)

- Tx prolonged course of:
 - ampicillin
 - doxycycline
 - clindamycin



H. Zoster







Type of isolation?Hutchinson's sign

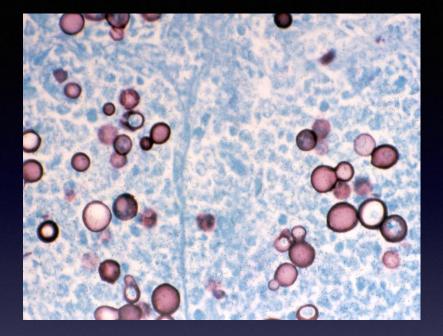
 Ramsey Hunt syndrome (H. zoster oticus), involving facial nerve (VII), w/ facial palsy, otalgia, dermatomal vesicles, occasional hearing loss

Blastomycosis



Mackowiak P A et al. Clin Infect Dis. 2012;55:1390-1391

Blastomycosis



- single, broad-based buds
 decaying vegetation,
 e.g. beaver dams
- Tx: Itraconazole



often involves skin, bone;
GU tract in males
regardless of presentation,
always considered as
disseminated disease, w/
lungs being the primary entry
site

Neutropenia/immunosuppression

....including initial approach to the febrile neutropenic patient as well as the persistently febrile patient

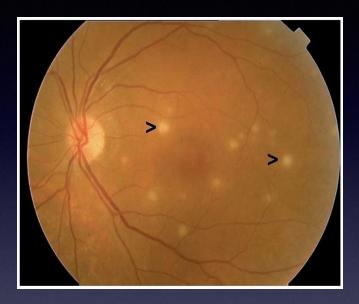
- most common bacteremia: E. coli
- most lethal organism: pseudomonas initial empiric rx always includes antipseudomonas coverage
- when remains febrile: antifungals

Ecthyma gangrenosum

- Most frequently assoc.
 w/ pseudomonas
 bacteremia
- Neutropenia, or other severely impaired immunity
- Erythematous / hemorrhagic pustule, evolving into central necrosis



Candida endophthalmitis



- severe neutropenia; may become apparent as neutrophils are recovering
- other setting/risk factors: ICU, multiple IVs/central lines, multiple antibiotics, TPN
- initial rx: echinocandins



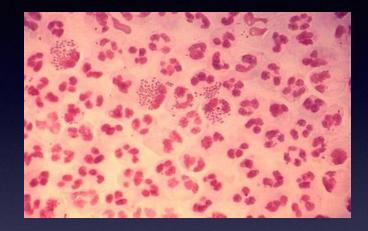


Strongyloides:

- diffuse pulmonary infiltrates in an immunosuppressed host
- NOT grossly visible ...
- Rx: ivermectin / albendazole

CNS Presentations





Neisseria meningitidis:

acute meningitis w/ <u>rash</u>; Rx: ceftriaxone
sepsis syndrome/<u>bacteremia</u> associated w/ terminal compliment deficiencies (C₅ - C₉), as well as splenectomy



Cryptococcus

HIV+, other dx assoc. w/ t-cell deficiencies;
sub-acute headache, mental status changes
tx: amphotericin B + flucytosine / fluconazole

Additional potential CNS questions:

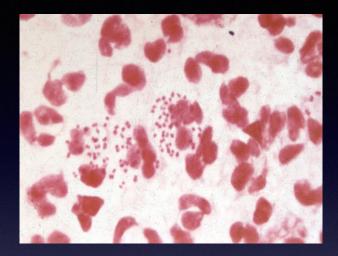
Interpretation of CSF results in a patient with fever, CNS findings

- Bacterial, viral, fungal, TB
- Meningitis vs encephalitis
- HSV encephalitis
- Meningitis w/ highest mortality (pneumococcus)
- Complement deficiency
- Most common cause of lymphocyte-predominant meningitis in a young, otherwise healthy individual
- Listeria scenarios who? CSF results, including gm stain w/ gm + rods? Rx?

D/D of Meningitis

	Bacterial	Viral	TB/Fungal
Cell count	increased; neutrophil predominant	sl. increased; lymphocyte predominant	increased; lymphocyte predominant
Glucose	decreased	normal or sl. decrease	decreased
Protein	elevated	normal or minimally elevated	elevated





N. gonorrhoeae

- gm neg intracellular diplococci
- <u>painful</u> urethral/cervical discharge
- pustular rash
- Late compliment deficiency
- Tx: cefixime (?) / ceftriaxone



Primary Syphilis:

- painless (usually genital) ulcer
- darkfield microscopy; PCR
- serology usually negative











Badri T, Ben Jennet S. N Engl J Med 2011;364:71-71.

T. pallidum

- 2° stages and beyond -> dx by serology
- rash includes palms and soles
- RPR or VDRL to screen
- FTA as confirmatory (though being replaced by TP-PA)



Treatment of Syphilis

- Less than 1 yr's duration (includes primary, secondary and early latent): benzathine PCN 2.4 mill units i.m. x 1
 If HIV +, some treat weekly x 3
 Allergy: doxycycline, ceftriaxone
- Greater than 1 yr's duration (or unknown duration); late latent: benzathine PCN 2.4 mill units weekly x 3 Allergy: ceftriaxone, doxycycline

Treatment of Syphilis

- Neurosyphilis: 10-14 days IV PCN G, 18 -24 mill. units/day
- If PCN allergic: desensitize or...ceftriaxone probably effective
- In pregnancy, if PCN allergy must densensitze (though ceftriaxone probably effective)
- Jarish-Herxheimer rxn

Malaria







- appropriate travel hx
- "black water fever"
- speciation by PCR at the CDC
- D/D Babesiosis

Banana gametocyte: P. falciparum

Babesiosis (B. microti; B. divergens)

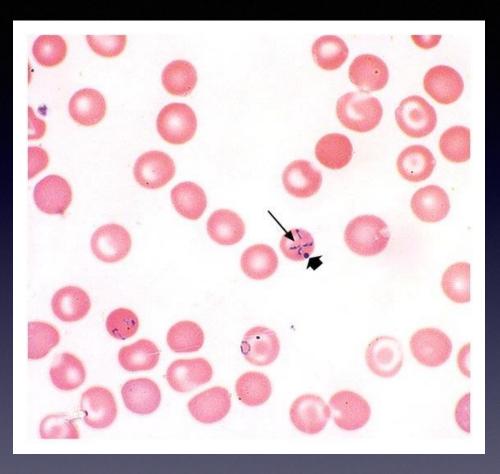
- Non-specific illness w/ headache, myalgia, malaise, fever after travel to coastal northeastern U.S. in late spring, early summer, particularly if hx of tick exposure 1 - 4 weeks earlier; much more severe illness if <u>splenectomized</u>
- Occasionally transmitted by transfusion
- Tick vector: Ixodes scapularis
- Reservoir host: white footed mouse
- Note: this same tick also transmits Lyme dx (Borrelia burgdorferi) and anaplasmosis. Consider if severe dx or poor response to treatment for these other diseases

Babesiosis*

- Hemolytic anemia, thrombocytopenia
- No rash
- Dx: RBC inclusion bodies ~ malaria on blood smear; however, tetrads ("Maltese cross formations") NOT seen in malaria
- Dx: PCR
- Rx: atovaquone + azithromycin if mild; IV clindamycin + p.o. quinine +/- exchange transfusion if severe

*Vannier & Krause. NEJM 2012;366: 2397-2407

Babesiosis



Noskoviak K, Broome E. N Engl J Med 2008;358:e19.



The NEW ENGLAND JOURNAL of MEDICINE

Lyme Disease (Borrelia burgdorferi)





Erythema migrans

Lyme Disease

- > 300,000 cases/yr in U.S.
- Systemic symptoms, rash, joint, CNS involvement
 - erythema migrans > 90%
 - carditis w/ conduction defects <10%
 - various neurologic presentations ~15%
 - cranial neuropathies; esp. bilateral VII nerve palsey; meningitis; radiculopathy
- IF chronic disease exists, may be due to immune dysregulation (elevated IL-23 and/or other)
- Prophylaxis: 200mg doxycycline x 1

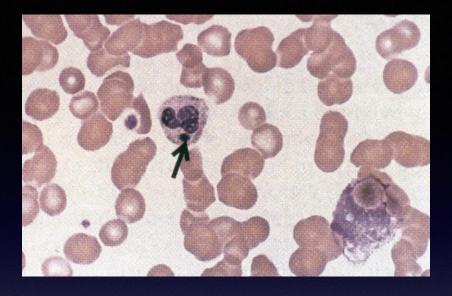
Ehrlichiosis/Anaplasmosis

Human <u>Monocytic</u> <u>Ehrlichiosis:</u>

- E. chaffeensis
- Monocytes
- macrophages of liver, spleen and bone marrow
- S.E, south-central, mid-Atlantic U.S.

Human <u>Granulocytic</u> <u>Anaplasmosis:</u>

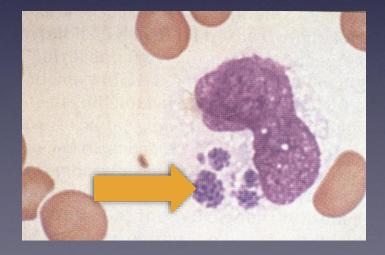
- A. phagocytophilum
- seen on peripheral smear (granulocytes)
- upper-midwest, N.E.,
 California, Europe
- E. ewingii: as above except geography of HME



Ehrlichiosis/Anaplasmosis:

- following tick exposure
- flu-like illness w/ leukopenia,
- thrombocytopenia
- spring/summer
- hyponatremia, elevated LFT's

- Note: "morulae "- cytoplasmic inclusions of "elementary bodies" Common only w/ anaplasma
- Diagnosis by PCR, blood smear
- Tx: doxycycline



Treatment

Warning: Questions about treatment failure

Lyme Dx:

- Amoxicillin/cefuroxime
- Ceftriaxone
- Doxycycline
- Ehrlichiosis/Anaplasmosis:
 - Doxycycline
- <u>Babesiosis:</u>
 - Atovaquone + azithromycin





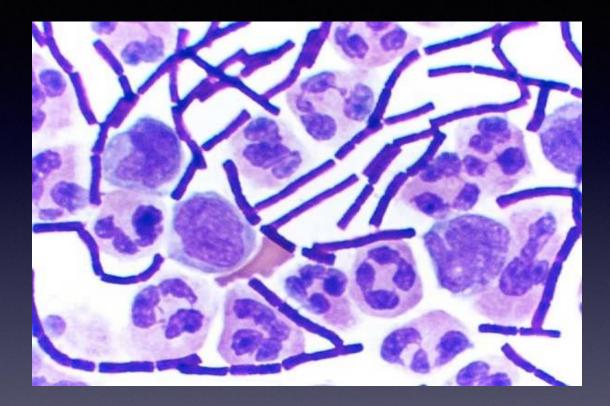
Esfandbod M, Malekpour M. N Engl J Med 2009;361:178-178.

Cutaneous anthrax

- systemically ill
- painless eschar w/ marked, localized edema
- contaminated soil, livestock

Bioterrorism-related Anthrax

- Multiple, previously healthy pts, w/ severe, rapidly fatal, flu-like illness
- "pneumonia" uncommon
- Widened mediastinum
- Large <u>hemorrhagic</u> pleural effusions
- <u>Hemorrhagic</u> meningitis
- Tx: penicillin / ciprofloxicin / doxycycline



CSF Gm stain - Anthrax (JAMA; 2001)

Smallpox

- Severe illness w/ painful, nodular rash
- Severe back pain
- All lesions in the same stage
- Rash most prominent on face and extremities, including palms and soles



