#### <u>Podcast (Video Recorded Lecture Series)</u>: Soft Tissue Infections for the USMLE Step One Exam



My baby's got gas!





The Flesh Eaters: Necrotizing Fasciitis

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## **MSK Infections**

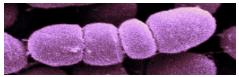
- Bone and Joint Infections
  - Septic Arthritis
  - Osteomyelitis
  - (Reactive Arthritis)
- Tick Borne (and related) Infections
  - Lyme Disease/Anaplasma (Ixodes)
    - Ehrlichiosis (Amblyomma; Lone Star)
  - Babesiosis (Ixodes)
  - RMSF (Dermacentor)
- Soft Tissue Infections
  - Clostridium perfringens  $\rightarrow$  Myonecrosis
  - Necrotizing fasciitis  $\rightarrow$  GAS
- Viral

– Parvovirus (RA like presentation)

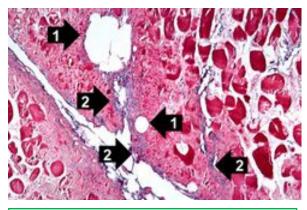
### Clostridium perfringens (Traumatic Gas Myonecrosis)

- Background
  - Spectrum of illness includes: wound contamination, anaerobic cellulitis and myonecrosis (gas gangrene).
  - Perfect storm (myonecrosis): traumatic injury (introduces bug), presence of devitalized tissue (loss of blood supply) and delay in rx.
  - Characterized by: tissue necrosis and absence of inflammatory response (toxin-mediated)



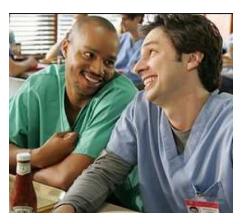


Gram +, spore-forming rod



Devitalized tissue

 $\alpha$ -toxin (PPLase)



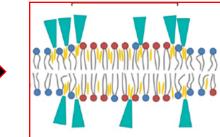
Surgical delay

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- Microbiology
  - Gram (+) anaerobic, spore forming rods that produce exotoxins
  - Does not grow in presence of oxygen (anaerobic); tissue death required.
- Pathology (cellulitis v myonecrosis)
  - Devitalized tissue (anaerobic cellulitis) supports the growth of organism with local growth and gas extending along fascial planes. No bacteremia or myonecrosis (during this phase).
  - Myonecrosis is characterized by the invasion and destruction of healthy muscle tissue. Required: traumatic injury and vascular compromise.

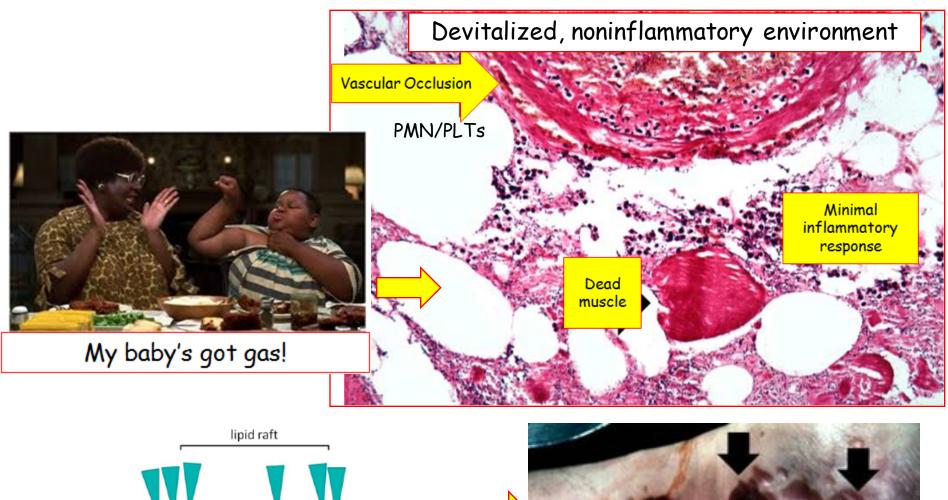
# α-Toxin(s): Phospholipase C

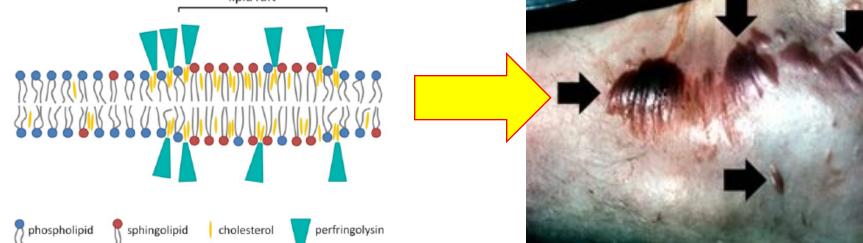
- Alpha (essential) aka Lecithinase (degrades lecithin)
  - Hemolytic toxin with Phospholipase C (PLC) activity
    - 1. Catalyzes splitting of PPL molecules  $\rightarrow$  cell membrane damage and cellular death
    - 2. Creates occlusive vascular aggregates that expand the anaerobic environment.
      - Upregulates adherence molecules on PMN/endothelial cells (they arrive but don't penetrate)
      - Stimulates platelet aggregation via IIb/IIIa activation



Theta (perfingolysin O; nonessential)
– Cytolysin (similar to streptolysin) - pore forming.







### Clostridium perfringens (Gas Myonecrosis)

• Clinical

 $\alpha$ -toxin

- Severe pain at site of trauma (due to tissue ischemia)
- PE (skin):
  - Crepitus may be present (most characteristic); bronze color/discharge
  - Large bullous vesicles that easily rupture
- Systemic toxicity (shock) occurs early (fever, multiorgan dysfunction)
- Diagnosis
  - Gas on imaging
  - Blood/tissue cultures with presence of Gram (+) rods but no pus (or PMN on microscopy)
- Treatment
  - (Repeated) Surgical debridement, antibiotic (PCN plus Clindamycin), shock support, hyperbaric oxygen (?)

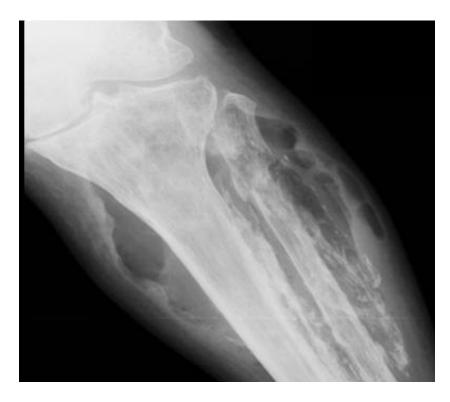
### Clostridium perfringens (Gas Myonecrosis)

They describe a patient with gas myonecrosis and then inquire about other syndrome caused by this organism...

the bait and switch

### • Special Notes:

- May also cause a syndrome of watery diarrhea with spores germinating (in gut); source: meat, poultry or gravy.
- Heat stable enterotoxin is produced in the GI tract after ingestion.
- Symptom delay (unlike ingestion of bacillus and staph toxins)





<u>Clostridium perfringens: spore forming, anaerobic gram (+) rod</u>

- 1. Traumatic injury
- 2. Crepitus +/- Shock
- **3**. Toxin:  $\alpha$ -toxin (aka Phospholipase C, Lecithinase).
  - Destroys cell membrane (catalyzes <u>splitting of PPL</u> molecules) AND
  - Vascular occlusion (PMN and platelet aggregates w/o inflammatory response)
- 4. GI illness: watery diarrhea (enterotoxin-mediated)

### Necrotizing Fasciitis

- Background
  - Fulminant soft tissue destruction w/ shock-like presentation
    - Muscle fascia and subcutaneous fat (c/w myonecrosis)

Similar shock-like presentation to myonecrosis Different organism - S. pyogenes Different tissue - fascia/fat No gas present - aerobic (if crepitus or gas  $\rightarrow C$ . perfringens)



## Necrotizing Fasciitis

- Background
  - Fulminant soft tissue destruction w/ shock-like presentation
    - Muscle fascia and subcutaneous fat (c/w myonecrosis)
  - Infections may be mono- (GAS, SA) or polymicrobial.
    - Tissue injury  $\rightarrow$  monomicrobial; Diabetes  $\rightarrow$  polymicrobial
- Microbiology
  - GAS (Strep pyogenes): catalase negative, bacitracin sensitive (PYR +).
  - Key Virulence Factors:
    - M protein antiphagocytic (blocks opsonization)
    - Pyrogenic exotoxins superantigens associated with TSS
- Gross Pathology
  - Initial spread along fascial planes; the overlying tissue may appear to unaffected.

## Necrotizing Fasciitis

- Clinical
  - Fever and systemic signs of toxicity (multi-organ failure)
  - Pain out of proportion to physical findings (swelling, warmth)
  - Progresses over 3-5 days w/o  $\Delta$  in overlying soft tissue (may appear unaffected)
    - Pain may improve due to involvement of blood vessels (thrombosis) and extension to nerves (akin to thrombangiitis obliterans)
  - Swelling may lead to compartment syndrome with progression  $\rightarrow$  myonecrosis
- Diagnosis: Surgical Visualization
  - There are no diagnostic tests. Studies: oriented toward systemic manifestations of infection and assessing for gas in compartments
    - Blood/Wound (intraoperative) cultures, CBC, CPK, BUN/Cr; Imaging?
- Treatment
  - Surgical debridement and antimicrobials (poly- or monomicrobial coverage)
    - Mortality rate w/o surgery: ~100%
- Special Notes
  - Distinguishing from myonecrosis and identification of responsible organisms is determined via surgery and histopathology.



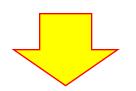
GAS, catalase neg Bacitracin (S), PYR (+)



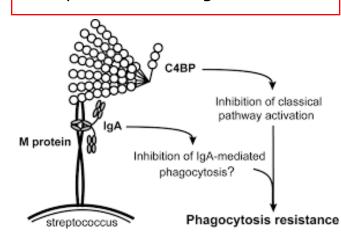
The Flesh Eaters: Necrotizing Fasciitis



Pain out of proportion to physical findings



M-protein: immunogenic, anti $\Phi$ 





Fascia and Fat

#### If gas, think clostridia

