

Tishk International University
Faculty of Science
Medical Analysis Department



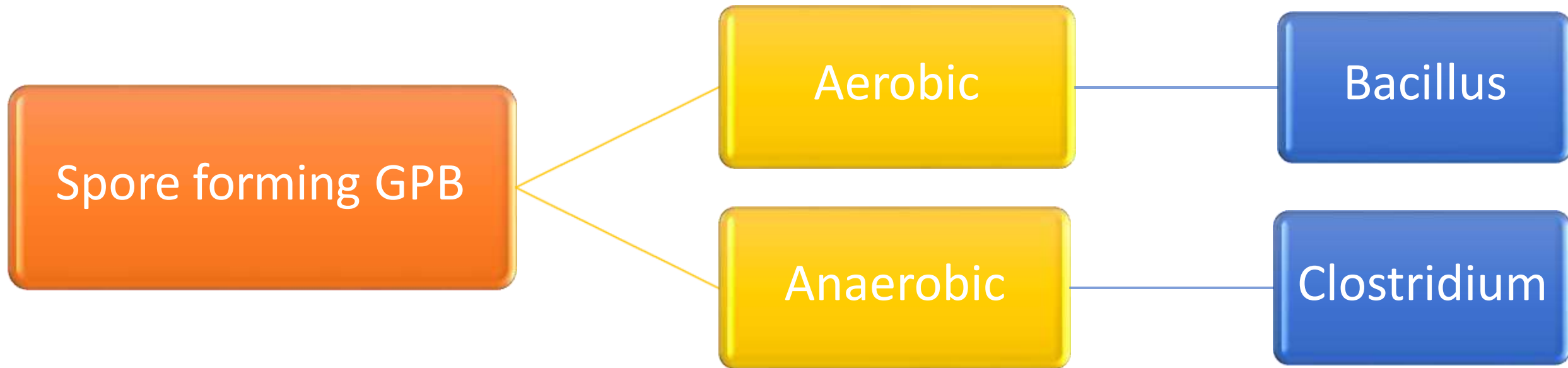
..... Medical Microbiology.....

TOPIC: Spore forming Gram positive bacteria

2nd Grade- Spring Semester 2019-2020



CLASSIFICATION



CLOSTRIDIUM

- Anaerobic, endospore former, gram positive rod
- Clostridia are ubiquitous in soil, water, and sewage
- Part of the normal microbial population in the GI tracts of animals and humans

Medically important species are

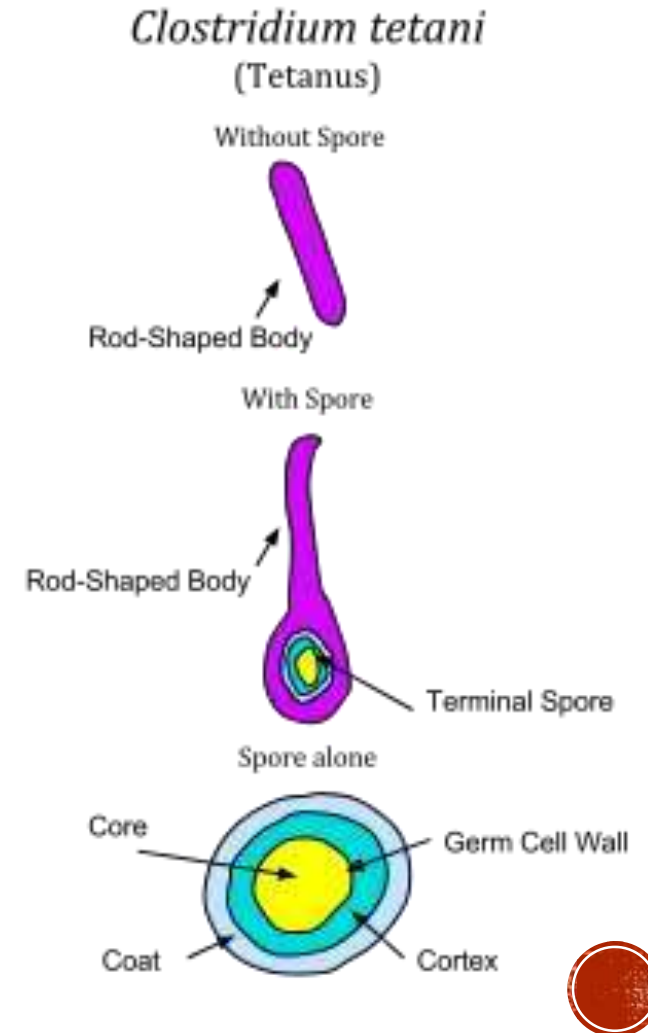
- *Clostridium tetani*
- *Clostridium botulinum*
- *Clostridium erfigen*
- *Clostridium difficile*



CLOSTRIDIUM TETANI

MORPHOLOGY

- Large, motile, spore forming rod
- It produce round terminal spore (drumstick appearance)
- Spores are widespread in soil

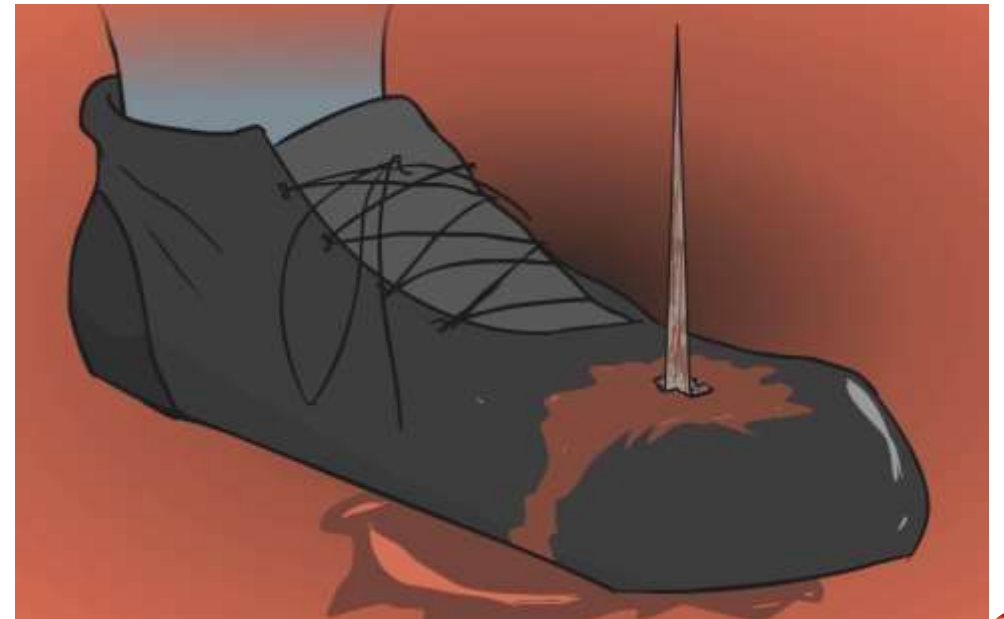


CLOSTRIDIUM TETANI GRAM STAIN



TRANSMISSION

- Portal of entry is wound site
- In neonatal tetanus, it may enter through contaminated umbilicus or circumcision wound.



Two toxins

- Oxygen-labile hemolysin (tetanolysin)
- Heat-labile neurotoxin (tetanospasmin), Plasmid encoded



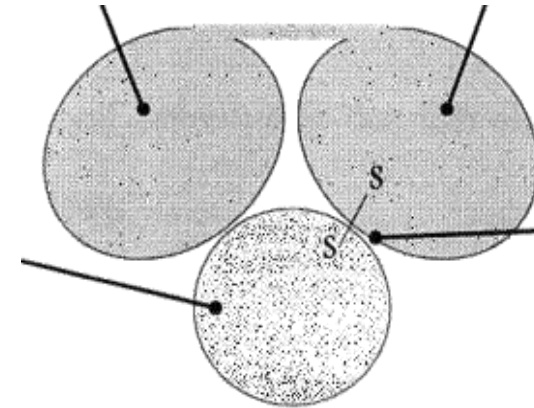
TETANOLYSIN

- Clinical significance is unknown
- Inhibited by oxygen and serum cholesterol.
- serologically related to streptolysin O and the hemolysins.



TETANOSPASMIN

- Released during stationary phase of growth.
- Responsible for clinical manifestations of tetanus.
- Two part (A-B)



■ Heavy chain (100 kDa)
■ Light chan (50 kDa)

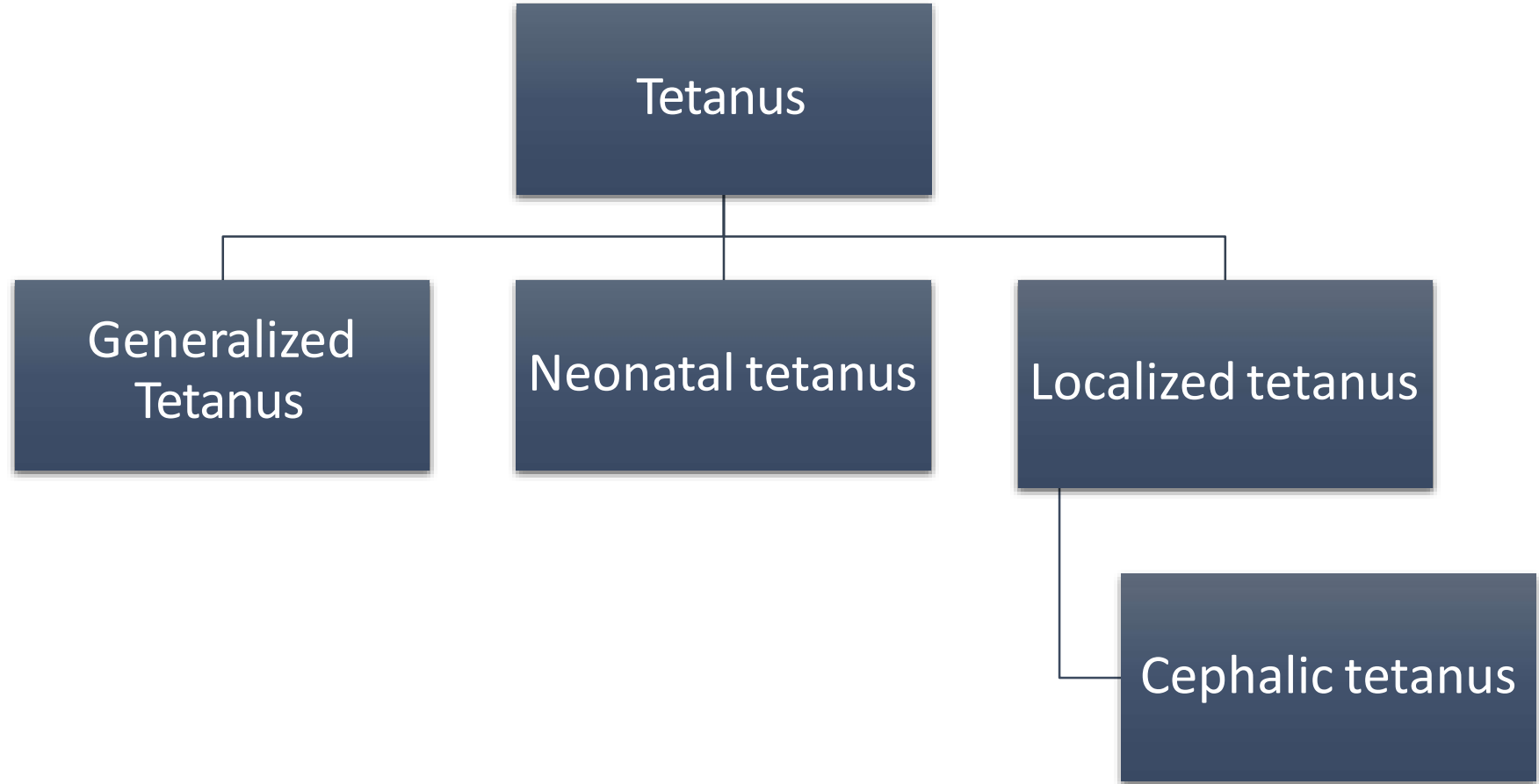


Incubation period

- varies from a few days to weeks.
- The duration of the incubation period is directly related to the distance of the primary wound infection from the central nervous system.



CLINICAL DISEASE



GENERALIZED TETANUS

- Most common form.
- Involvement of the masseter muscles (trismus or lockjaw) is the presenting sign in most patients.
- characteristic sardonic smile that results from the sustained contraction of the facial muscles is known as risus sardonicus
- Other early signs are drooling, sweating, irritability, and persistent back spasms (opisthotonos)



GENERALIZED TETANUS



FIGURE 30-6 Facial spasm and risus sardonicus in a patient with tetanus. (From Cohen J, Powderly WG, Opal SM: *Infectious diseases*, ed 3, Philadelphia, 2010, Mosby.)



FIGURE 30-7 A child with tetanus and opisthotonos resulting from persistent spasms of the back muscles. (From Emond RT, Rowland HAK, Welsby P: *Colour atlas of infectious diseases*, ed 3, London, 1995, Wolfe.)



LOCALIZED TETANUS

- Disease remains confined to the musculature at the site of primary infection.

Cephalic tetanus

- A variant in which the primary site of infection is the head
- The prognosis for patients with cephalic tetanus is very poor.



NEONATAL TETANUS

Neonatal tetanus (tetanus neonatorum)

- Typically associated with an initial infection of the umbilical stump
- progresses to become generalized.
- The mortality in infants exceeds 90%
- developmental defects are present in survivors.
- This is almost exclusively a disease in developing countries.



NEONATAL TETANUS



TREATMENT

- Debridement of the primary wound
- Use of penicillin or metronidazole to kill the bacteria and reduce toxin production

Prevention:

- **Vaccination with tetanus toxoid**



CLOSTRIDIUM BOTULINUM

CLOSTRIDIUM BOTULINUM

- large, spore forming, anaerobic rods.
- Spores are widespread in soil, contaminate vegetables and meat.
- If foods are canned without sterilization, spore will survive and germinate under anaerobic condition and bacteria will produce toxin



- Similar to tetanus toxin, C. botulinum toxin is a protein (A-B toxin) consisting of a small subunit (light, or A chain)



CLINICAL USE OF TOXIN

Botox is a commercial preparation of exotoxin A used to remove wrinkles on the face

Effective in minute amount in certain muscle disorders



CLINICAL DISEASE

Botulism

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graph TD; Botulism --> Wound_botulism[Wound botulism]; Botulism --> Infant_botulism[Infant botulism]; Botulism --- Clinical[• Descending weakness and paralysis<br>• Diplopia<br>• Dysphagia<br>• Respiratory muscle failure<br>• No fever]; Wound_botulism --- Wound_desc[Spores contaminate wound, germinate and produce toxin at site]; Infant_botulism --- Infant_desc[Organism grow in gut and produce toxin. Ingestion of honey containing the organism is the cause];
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- Descending weakness and paralysis
- Diplopia
- Dysphagia
- Respiratory muscle failure
- No fever

Wound botulism

Spores contaminate wound, germinate and produce toxin at site

Infant botulism

Organism grow in gut and produce toxin. Ingestion of honey containing the organism is the cause



TREATMENT

Patients with botulism require the following treatment measures:

- Adequate ventilatory support
- Elimination of the organism from the GI tract through the judicious use of gastric lavage and metronidazole or penicillin therapy,
- Use of trivalent botulinum antitoxin to inactivate unbound toxin circulating in the bloodstream.



PREVENTION

- Disease is prevented by
 - Destroying the spores in food (virtually impossible for practical reasons)
 - Preventing spore germination (by maintaining the food in an acid pH or storage at 4° C or colder)
 - Destroying the preformed toxin (Toxins are inactivated by heating at 60° C to 100° C for 10 minutes).
 - Infant botulism has been associated with consumption of honey contaminated with *C. botulinum* spores, so children younger than 1 year should not eat honey.



CLOSTRIDIUM PERFRINGENS

CLOSTRIDIUM PERFRINGENS

- Most frequent clinical isolate of clostridium
- Inhabits in soil and in intestine of animals and humans.
- Rapidly grows in tissues and in culture
- Large, rectangular, gram positive bacilli
- Capsulated, non motile with subterminal spore



TOXINS AND ENZYMES

Epsilon toxin

- Increase vascular permeability of gastrointestinal wall

Iota toxin

- necrotizing activity
- increase vascular permeability

Enterotoxin

- Produced primarily by type A strain
- Disrupts ion transport in ileum and jejunum by inserting into the cell membrane and altering membrane permeability.



Clostridial myonecrosis (gas gangrene)

- Military settings- Gutshot injuries
- Civilian cases - accidental injuries, surgical complication, injection of medication such as epinephrine.



■ Simple wound contamination

- Care of wound
- removal of necrotic tissue
- cleansing
- Antibiotics - rarely required

Anaerobic cellulitis:

- Opening the involved area
- removing all necrotic tissue
- cleansing thoroughly
- Antibiotics



Clostridial myonecrosis

- Care of wound : surgical removal of all infected and necrotic tissue
- Amputation- rapidly progressive infection involving limb
- Antibiotics - Penicillin G/ Clindamycin or Metronidazole, Gentamicin or Tobramycin (for facultative Gram negative organisms)



CLOSTRIDIUM DIFFICILE

GENERAL CRITERIA

- Part of normal microbial flora of GIT (3 – 5% adult, 40 – 50% healthy neonates)

Morphology:

- large gram positive rod.
- having oval subterminal spore.



VIRULENCE FACTORS

Enterotoxin (Toxin A)

- produce chemotaxis; induce cytokine production, with hypersecretion of fluid;
- produces haemorrhagic necrosis.

Cytotoxin (Toxin B)

- induces depolymerization of actin with loss of cellular cytoskeleton.

Adhesin factor

- binding to colonic cells.



TRANSMISSION

- Exogenous - Person to person in hospital.
- Endogenous -Over growth of toxin producing strains after Rx of Antibiotics.

Antibiotics implicated in Cl.difficile Associated Diarrhoea and Colitis:

- Cephalosporin
- Ampicillin and amoxicillin
- Clindamycin.
- Other Penicillins, Macrolides, Tetracyclines.

