



Gram-Negative Bacteria

Lecture Five



Learning Outcomes



Outcomes

To know how to differentiate between Gram-negative cocci and rods

To provide the basic knowledge about *Neisseria gonorrhoeae* and *Moraxella*

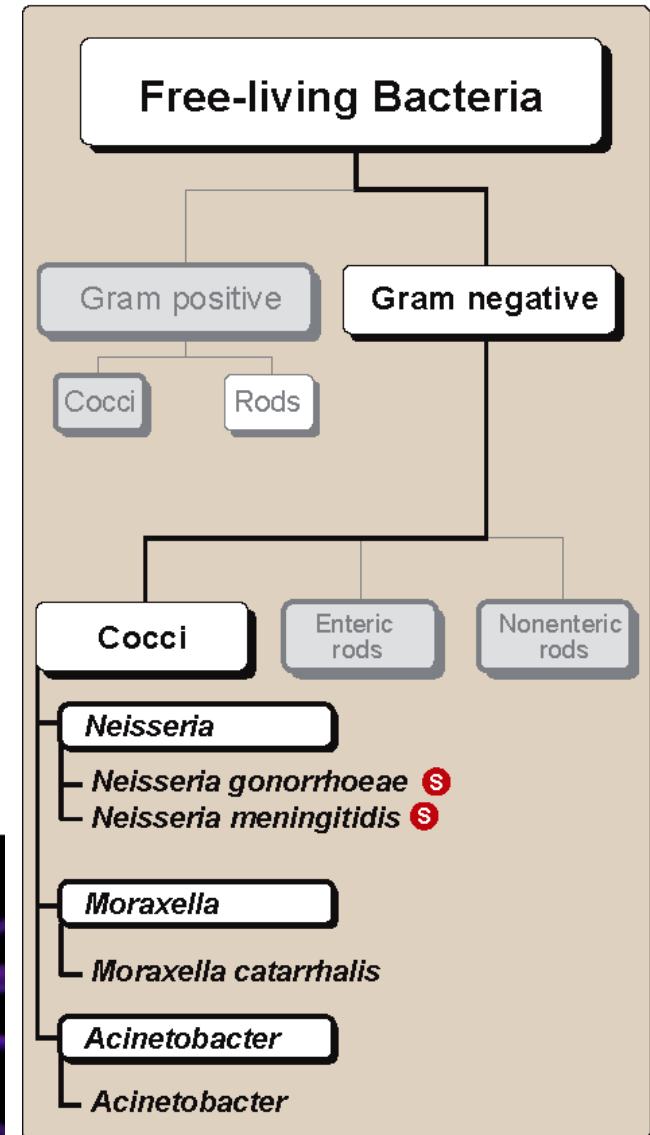
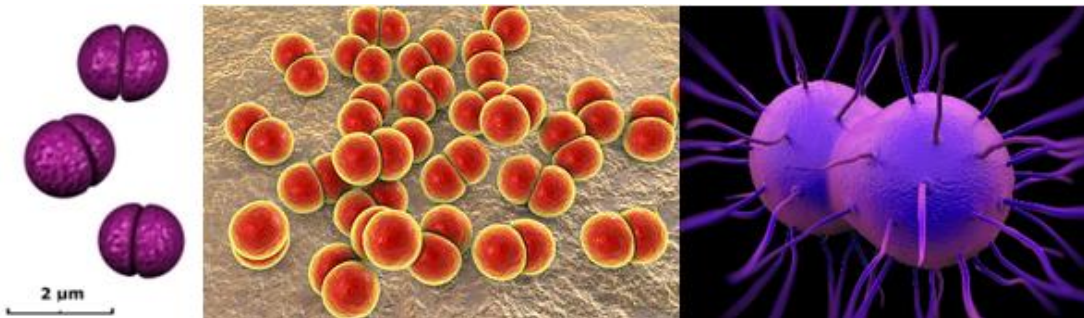
To provide the basic knowledge about *Pseudomonas aeruginosa*

To acquire the basic knowledge about *Haemophilus influenzae*

To learn how to treat diseases caused by Gram-negative bacteria

Gram Negative Cocci

- The genus *Neisseria* consists of Gram-negative, aerobic cocci.
- *Neisseria gonorrhoeae* is a Gram-negative diplococcus that causes gonorrhoea infectious disease.



Neisseria gonorrhoeae

- *Neisseria gonorrhoeae* does not produce any exotoxins but has various virulence factors that help infections to take place.
- Pili
 - It enhance attachment of the organism to the host epithelial cell surface.
 - Lipooligosaccharide (LOS)
 - It causes endotoxic shock.
 - Outer membrane proteins
 - Contribute the virulent effect of bacteria.

Neisseria gonorrhoeae

- Porin Proteins

- The gonococcus expresses a single porin type, known as **PorB**.

- The protein tight binding and invasion of the epithelial cells.

- Also, protein from one bacterium binds to the LOS of an adjacent bacterium, which allows for the construction of a small colony.

- IgA Protease

- It degrades **IgA** (antibody) coating and leads to the invasion of host cell.

Neisseria gonorrhoeae

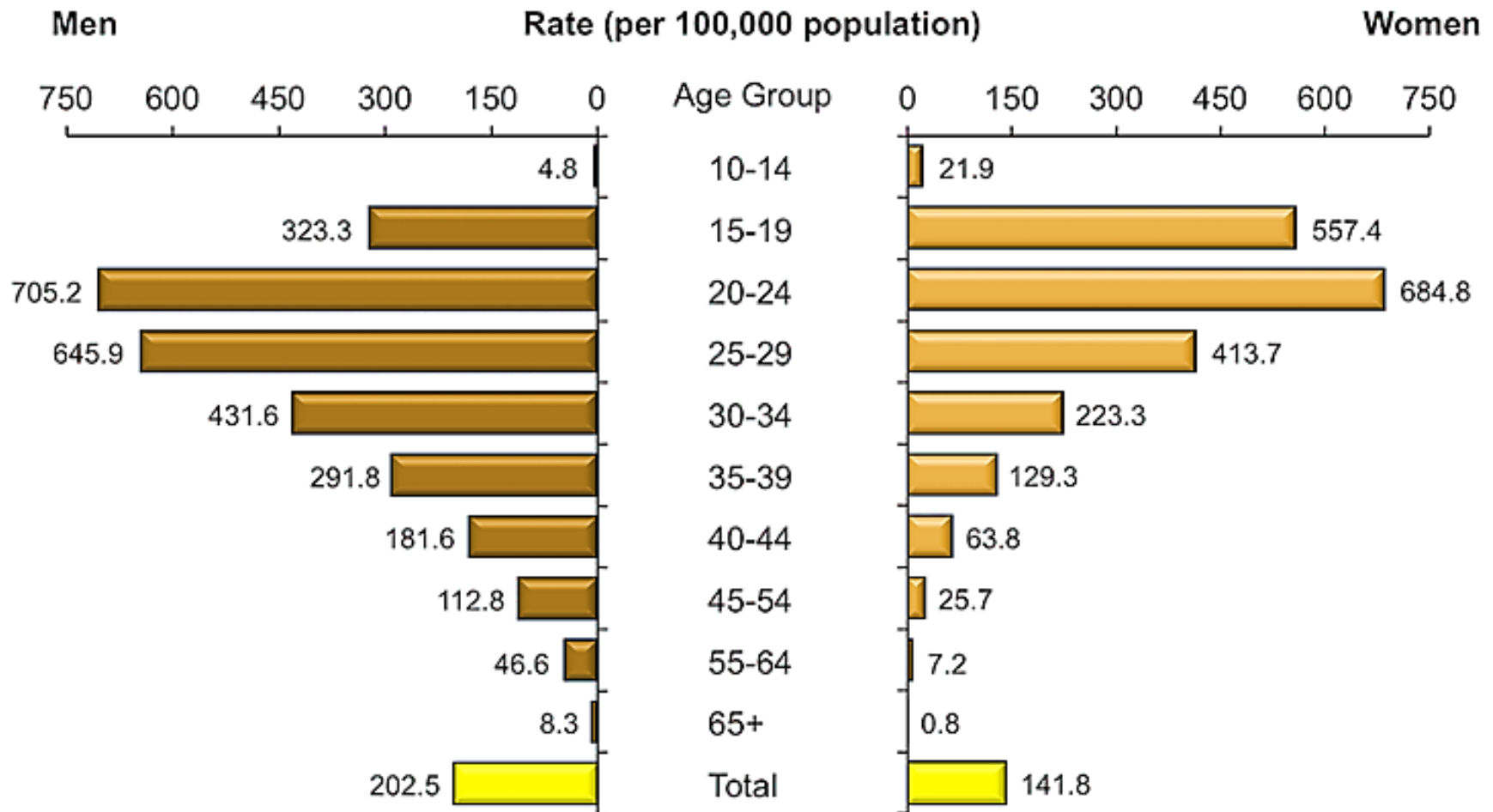
- Pathogenesis of *Neisseria gonorrhoeae*
- **Pili** and **Opa** proteins help adhesion of the gonococcus to the host epithelial cells.
- They make colonization possible.
- Both gonococci and meningococci produce an **IgA protease** that cleaves **IgA1**, helping the pathogen to evade immunoglobulins of this subclass.
- The gonococcus requires **iron** for growth and survival *in vivo*.

Neisseria gonorrhoeae

- Clinical Significance of *N. gonorrhoeae*
- Gonococci often **colonize** the mucous membrane of genitourinary tract.
- It may cause a **localized infection** with the production of pus.
- It may lead to tissue invasion, chronic inflammation and fibrosis.
- Higher proportion of female than male are generally **asymptomatic**, and can transmit gonococcal infections.

Neisseria gonorrhoeae

Gonorrhoea in the United States in 2017



Neisseria gonorrhoeae

The State of STDs in the United States



STDS SURGE FOR THE FIFTH
STRAIGHT YEAR, REACHING
AN ALL-TIME HIGH.



1.8 million
CASES OF CHLAMYDIA
19% rate increase since 2014



583,405
CASES OF GONORRHEA
63% rate increase since 2014



115,045
CASES OF SYPHILIS
71% rate increase of infectious
syphilis since 2014



1,306
CASES OF SYPHILIS
AMONG NEWBORNS
185% rate increase since 2014

LEARN MORE AT: www.cdc.gov/std/

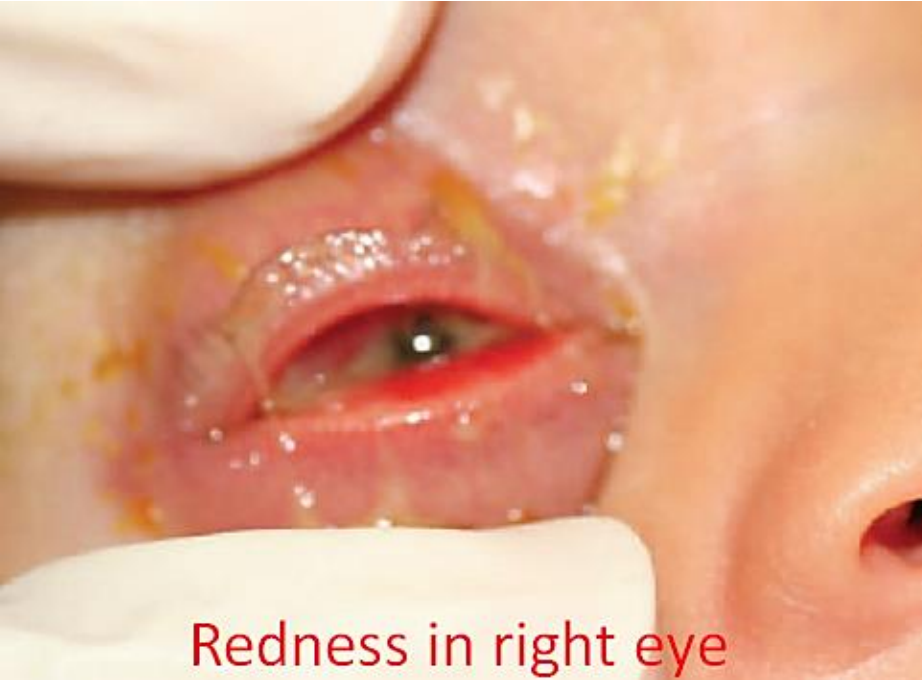
Neisseria gonorrhoeae

- Treatment and Prevention
- More than 20% of *N. gonorrhoeae* isolates are resistant to penicillin, tetracycline, cefoxitin and/or spectinomycin.
- Doxycycline is often used as a part of the treatment regimen for gonorrhea.
- Prevention of gonorrhea involves sexual contacts management of the patients.
- The use of barrier method is also a preventive measure against gonorrhea.

Neisseria gonorrhoeae

- Eye infections caused by *N. gonorrhoeae*
- Conjunctivitis (**ophthalmia neonatorum**) is common in **newborns**.
- Neonatal gonorrhoeal conjunctivitis is contracted when the **infant** is exposed to ***N. gonorrhoeae*** in the birth canal.
- It can lead to **corneal scarring** resulting in blindness in newborns.
- **Silver nitrate** or antibiotics are often applied to eyes to prevent gonorrhoea.

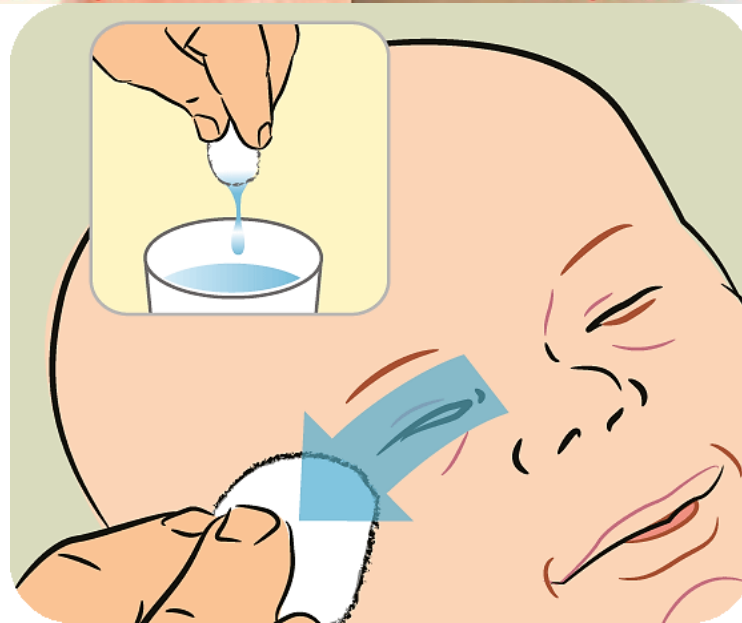
Neisseria gonorrhoeae



Redness in right eye



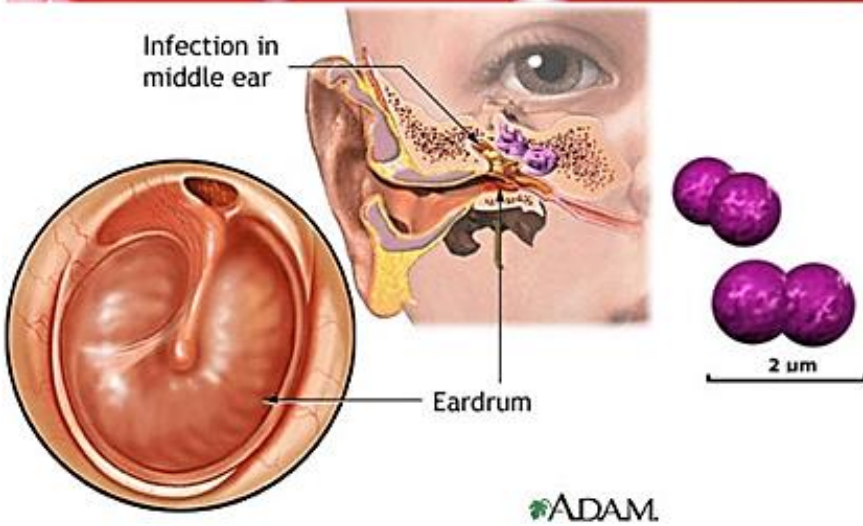
Ophthalmia neonatorum



Moraxella

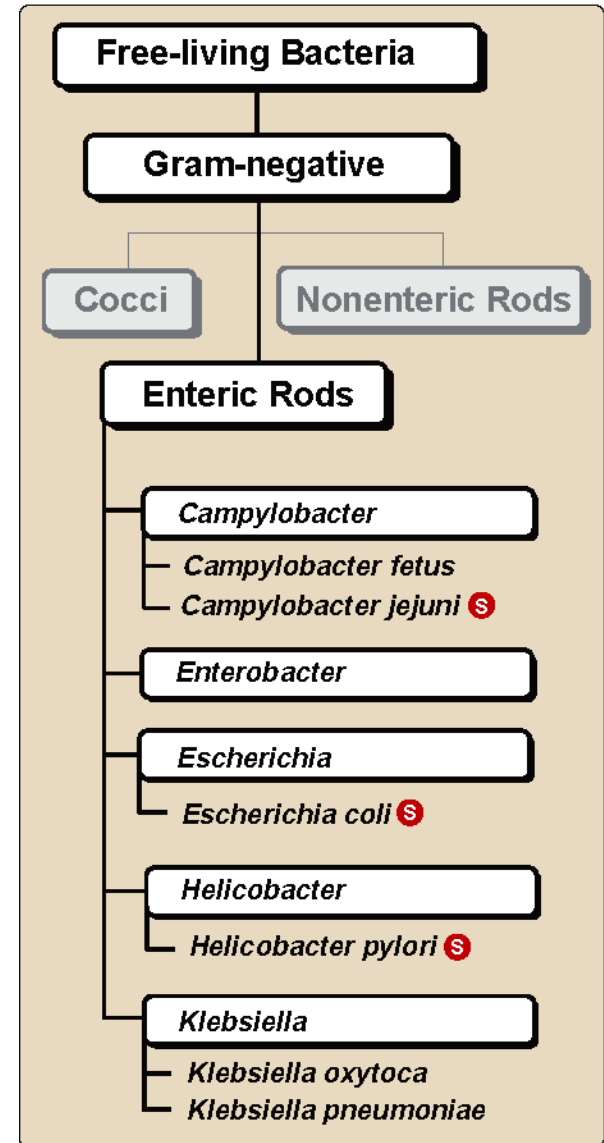
- *Moraxella* are non-motile, Gram-negative diplococci that are generally found in pairs.
- *Moraxella* are aerobic, fastidious organisms that do not ferment carbohydrates.
- It can cause infections of the respiratory system, middle ear, eye, CNS and joints.
- *Moraxella bovis* causes bovine keratoconjunctivitis (new forest eye; in cattle only).
- *Moraxella* can be treated by the use of subconjunctival injection of tetracycline as the antibiotic.

Moraxella



Gram Negative Rods

- **Enteric Bacteria**
- Enteric bacteria are Gram-negative rods.
- They are bacteria of the intestines that are associated with gastrointestinal flora (**Gut flora**) or disease.
- The taxonomic family is **Enterobacteriaceae**.
- *e.g. Escherichia coli*



Gram Negative Rods

- They are found in the **gastrointestinal** (GI) tract of humans or other animals.
- They are **sensitive** to drying and all grow in the presence or absence of oxygen (**facultative anaerobes**).
- They contain **lipopolysaccharide** (LPS), which is both antigenic and an important virulence factor (**endotoxin**).
- **Fecal** contamination is commonly important in the transmission of those organisms that cause GI tract diseases.

Escherichia coli

- *Escherichia coli* is part of the normal flora of the colon in humans and animals.
- It can be pathogenic both within and outside of the GI tract.
- *Escherichia coli* has pili that are important for adherence to host mucosal surfaces.
- It may be motile or non-motile.
- Most *Escherichia coli* strains can ferment lactose and produce acid and gas.
- *E. coli* may cause ophthalmia neonatorum.

Escherichia coli



- Most *Escherichia coli* strains are harmless, but some serotypes can cause serious food poisoning in their hosts.
- *Escherichia coli* can get into meat, raw milk or dairy products during processing.
- *E. coli* spread from one person to another.

Escherichia coli

Enterohemorrhagic bacteria Escherichia coli (EHEC)



Most Escherichia coli (E.coli) strains are harmless.
But some, like enterohemorrhagic **E. coli (EHEC)**, are a hazard to human health and life.

Incubation period:
three to eight days

E. coli (EHEC), once in the human stomach, begins producing toxins that cause serious illnesses

Symptoms caused by E. coli (EHEC)

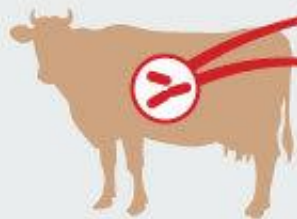
- Stomach muscle spasms
- Diarrhea (sometimes bloody diarrhea)
- Fever
- Vomiting

Complications:

hemolytic uremic syndrome (HUS)

Death rate: 3-5 %

INFECTION SOURCES



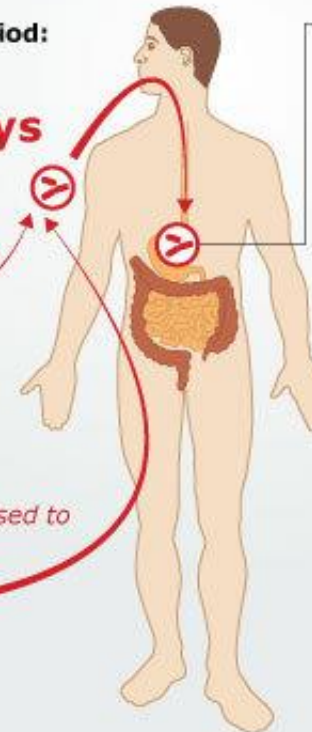
Cattle and other ruminants are the main E. coli (EHEC) carriers



Uncooked meat and raw milk

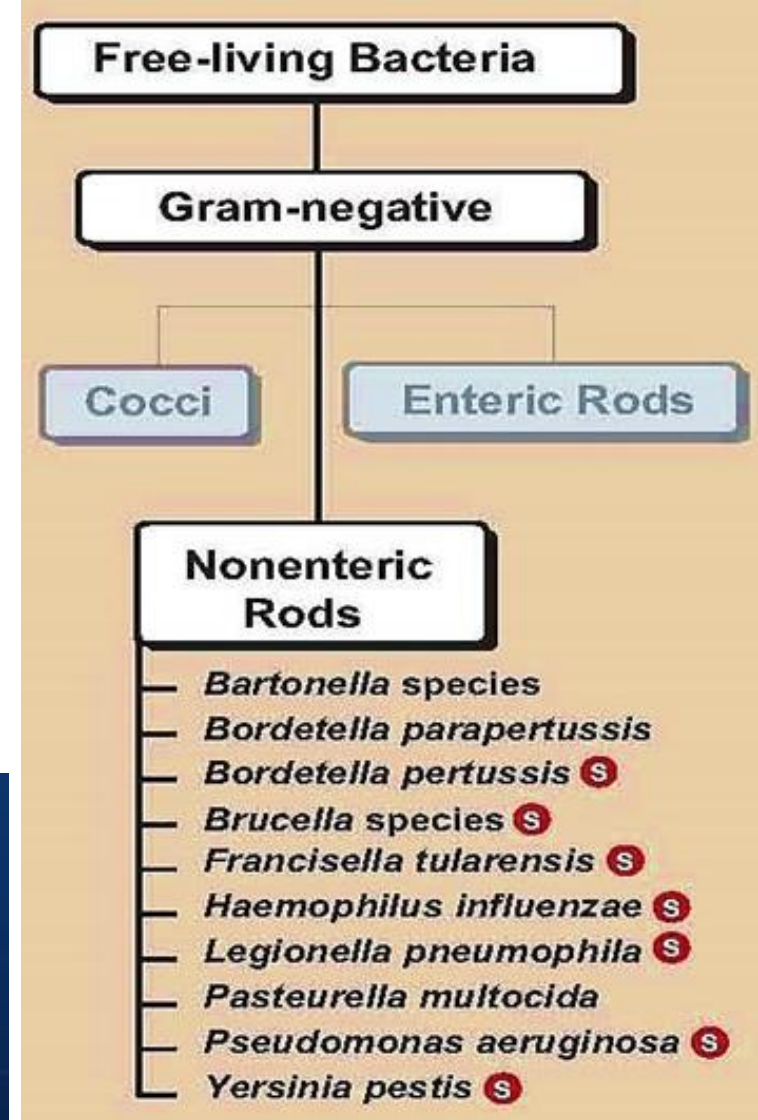
The bacteria die when food is exposed to heat (70°C and higher)

Fruit and vegetables (droppings of sick animals find their way into water bodies that in turn feed the soil)



Gram Negative Rods

- **Nonenteric Bacteria**
- Gram-negative bacilli.
- Do not ferment sugars.
- They resist antibiotics.
- Can be cultured in lab.
- *e.g. Pseudomonas aeruginosa.*



Pseudomonas aeruginosa

- *Pseudomonas aeruginosa* is the primary human pathogen in the genus *Pseudomonas*.
- It is found in soil, water, plants and animals.
- It may colonize healthy humans without causing disease.
- It is an opportunistic pathogen and a major cause of nosocomial infections.
- It can grow in laboratory water baths, hot tubs and other water-containing vessels.
- This explains why it is responsible for so many nosocomial infections.

Pseudomonas aeruginosa

- Pathogenesis of *Pseudomonas aeruginosa*
- *Pseudomonas aeruginosa* disease begins with the **attachment** of bacteria to the host cells followed by the colonization.
- **Pili** on the bacteria mediate adherence.
- The damage of the host tissue facilitates the **adherence** and **colonization**.
- *Pseudomonas aeruginosa* produces numerous types of **toxins** that can promote local invasion and the spread of the microorganism.

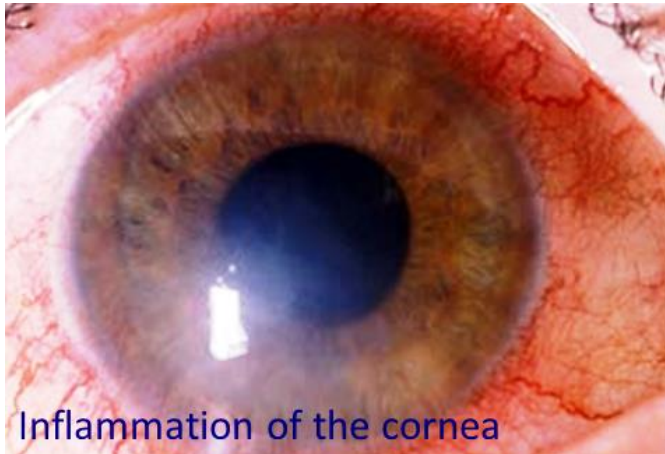
Pseudomonas aeruginosa

- Clinical Significance of *P. aeruginosa*
- *Pseudomonas aeruginosa* causes both localized and systemic illness.
- Localized infection may occur in the eye causing keratitis (inflammation of the cornea and endophthalmitis).
- The systemic infections may include infections to the bones, joints, skin and central nervous system.
- Individuals with impaired immune defences are at a high risk.

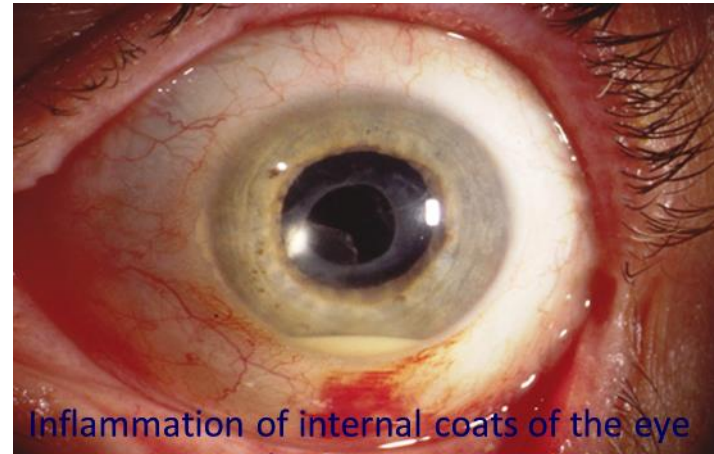
Pseudomonas aeruginosa

- Treatment and Prevention of *P. aeruginosa*
- It is **difficult** to find **antibiotics** that are effective against *P. aeruginosa* because of its rapid development of resistance mutations and its own innate mechanisms of antibiotic resistance.
- *Pseudomonas* infections typically occur in patients with **impaired defenses**.
- An aggressive **antimicrobial** therapy is required which includes the use of a combination of two antibiotics.

Pseudomonas aeruginosa



Inflammation of the cornea



Inflammation of internal coats of the eye

Gram (-) rods

Pseudomonas species

Pseudomonas aeruginosa

- Localized infections
- Systemic infections

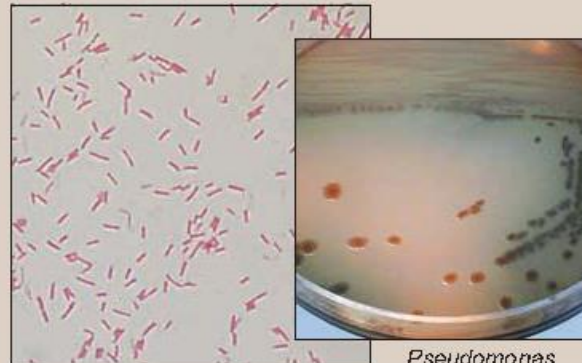
1 Antipseudomonal β -lactams¹

1 Ceftazidime

1 Tobramycin

1 Ciprofloxacin

¹Piperacillin or ticarcillin.



Pseudomonas aeruginosa grown from sputum (Gram stain)

Pseudomonas aeruginosa on MacConkey agar

- Encapsulated, motile rods (polar flagella)
- Aerobic or facultative anaerobe
- Produces diffusible green and blue pigments
- Oxidase positive
- Oxidizes but does not ferment carbohydrates, such as lactose
- Culture on MacConkey agar

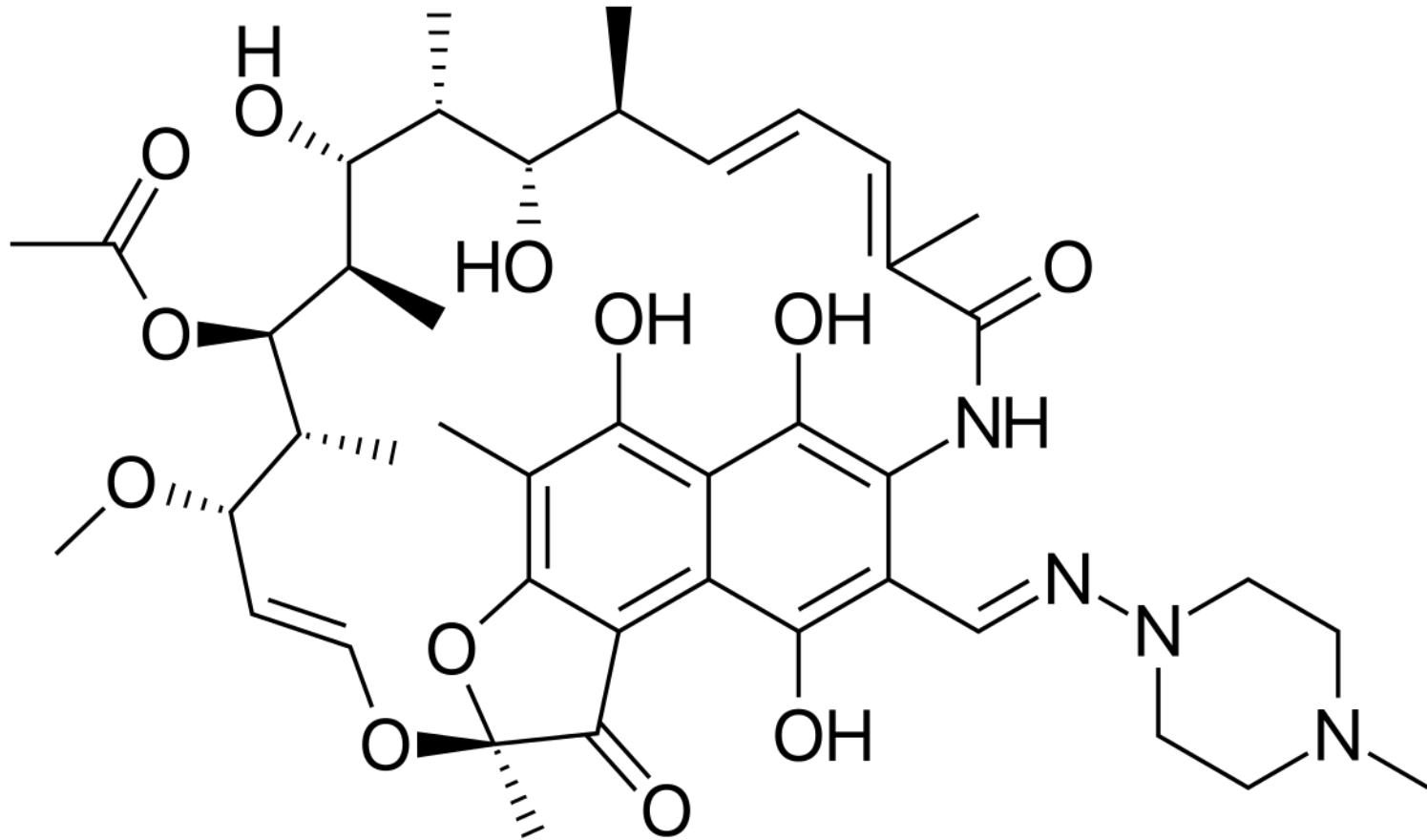
Haemophilus influenzae

- *Haemophilus influenzae* is a normal resident of the human upper respiratory tract.
- *Haemophilus influenzae* causes pneumonia in older adults and immunocompromised individuals.
- The transmission takes place by respiratory droplets.
- After attaching to and colonizing the respiratory mucosa, the infection can become systemic, with bacteria spreading through the blood to the CNS.

Haemophilus influenzae

- *Haemophilus influenzae* was a leading cause of bacterial meningitis, especially in infants and young children.
- Prevention of *Haemophilus influenzae*
- A conjugated vaccine against *H. influenzae* capsular polysaccharide type b is now administered to infants.
- The vaccine has dramatically lowered the number of meningitis infections.
- Rifampin is used to treat the bacterial infection and is given prophylactically.

Haemophilus influenzae



Rifampin

Haemophilus influenzae

- Brazilian purpuric fever (**BPF**) is an illness of children caused by the bacterium *Haemophilus influenzae* biotype aegyptius.
- **BPF** was first recognized in Brazil in **1984**.
- At this time, **young children** between the ages of 3 months and 10 years were contracting a strange illness which was characterized by high fever.
- These cases were all **fatal** due to **sepsis** and thought to be due to **meningitis**.

Haemophilus influenzae

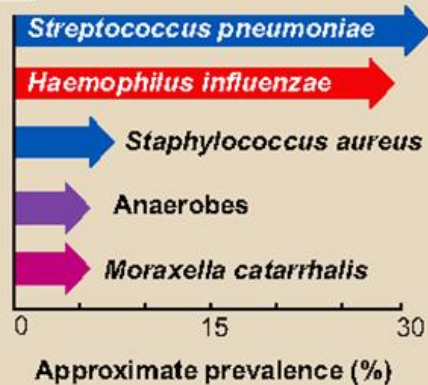
- These **deaths** was confirmed to be due infection by *H. influenzae aegyptius*.
- Although **BPF** was thought to be confined to Brazil, other cases occurred in Australia and the US during 1984–1990.
- The basic method for control of the conjunctivitis includes **proper hygiene** and care for the affected eye.
- It is difficult to treat BPF because of the difficulty obtaining an early **proper diagnosis**.

Haemophilus influenzae

- **Bacterial conjunctivitis** is an inflammation of the bulbar and/or the palpebral conjunctivas due to bacteria.
- Complications developed with extremely pathogenic bacteria.
- Produces a thick and yellow-green eye discharge.
- It may be associated with a respiratory infection.
- It may be associated with a sore throat.

Haemophilus influenzae

A Common pathogens¹



B Classification of pathogens

Gram (+) cocci

Staphylococcus aureus
Streptococcus pneumoniae

Gram (-) cocci

Moraxella catarrhalis

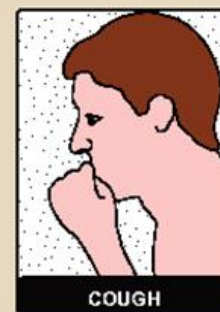
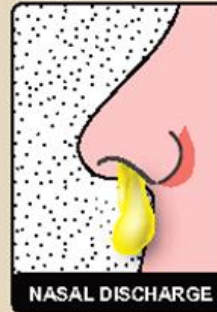
Gram (-) rods

Haemophilus influenzae

Anaerobic organisms

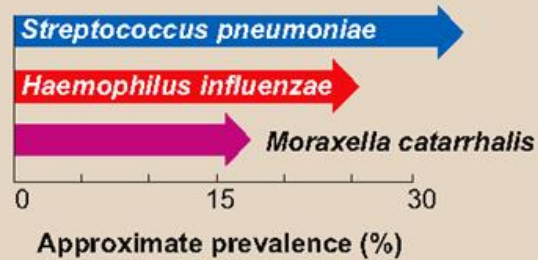
Various

C Common complaints²



Haemophilus influenzae

A Common pathogens¹



B Classification of pathogens

Gram (+) cocci

Streptococcus pneumoniae

Gram (-) cocci

Moraxella catarrhalis

Gram (-) rods

Haemophilus influenzae

C Common complaints²

