Genito-Urinary System Chlamydia trachomatis, Ureaplasma and Gardnerella

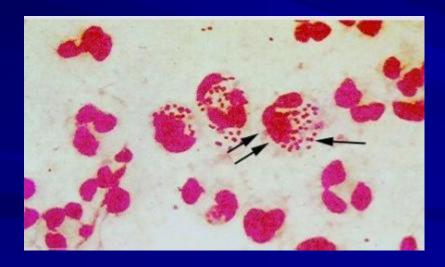
Chlamydia

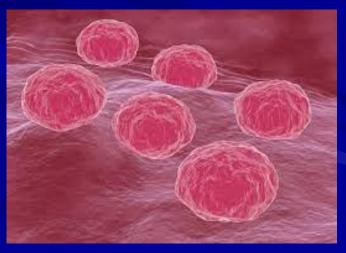
- Three of the nine species cause disease in humans.
 - Chlamydia trachomatis
 - is the most important human pathogen as a major cause of genital infection and conjunctivitis.
 - trachoma, is the leading preventable cause of blindness in the world.
 - Transmition direct contact
 - Chlamydophila pneumoniae
 - Chlamydophila psittaci
 - often infects birds



Chlamydia trachomatis

- C. trachomatis
 - round cells between 0.3 and 1 µm in diameter depending on the replicative stage.
 - lack the peptidoglycan layer
 - obligate intracellular parasites





REPLICATIVE CYCLE

Involves two forms of the organism:

Elementary body (EB)

 a small, hardy metabolically inert infectious form



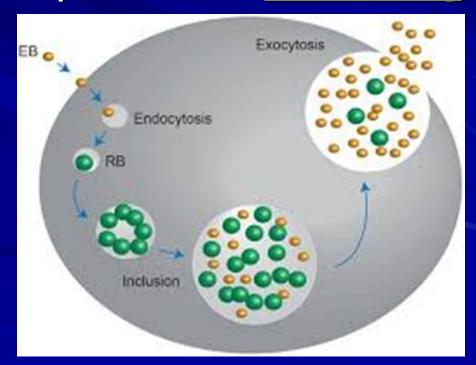
- a larger fragile intracellular replicative form termed the **Reticulate body (RB).**
 - Star shape

REPLICATIVE CYCLE

■ The <u>EB</u> attaches to unknown <u>receptors</u> on the <u>plasma membrane</u> (usually columnar or transitional epithelial cells).

It then enters the cell in an endocytotic vacuole and begins the process of converting

to the replicative RB.



■ As the RBs increase in number, the endosomal membrane expands by fusing with lipids of the Golgi apparatus eventually forming a large inclusion body. After 24 to 72 hours, the process reverses and the RBs reorganize and condense to yield multiple

Endocytosis

Exocytosis

EBs.

- The endosomal membrane then either disintegrates or fuses with the host cell membrane, releasing the EBs to infect adjacent cells.
- C. trachomatis also inhibits apoptosis of epithelial cells, thus enabling completion of its replicative cycle.

Chlamydia trachomatis Diseases EPIDEMIOLOGY

- C. trachomatis causes disease in several sites, including the <u>conjunctiva</u> and <u>genital</u> tract.
- It is the most common sexually transmitted disease.
- Humans are the sole reservoir.
- Neonatal conjunctivitis contracted from maternal genital infection (2 to 6% of newborn infants).

PATHOGENESIS

- Chlamydiae
 - endocervix and upper genital tract of women,
 - the <u>urethra</u>, <u>rectum</u> and <u>conjunct</u>. of both sexes.

IMMUNITY

C. trachomatis infections do not reliably result in protection against reinfection

CLINICAL ASPECTS

Genital Infections

- The clinical spectrum of sexually transmitted infections with *C. trachomatis* is similar to that of *Neisseria gonorrhoeae*. *C. trachomatis*
 - cause <u>urethritis</u> and <u>epididymitis</u> in men
 - cervicitis, salpingitis, and a urethritis in women.

- C. trachomatis urethritis
 - dysuria
 - a thin creamy <u>urethral discharge</u>.

In contrast with N.gonococcal which cause sever pain along with thick purulent discharge

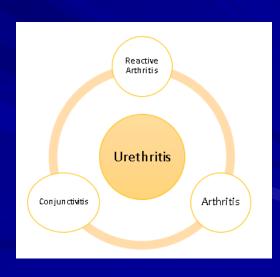
- Infections of the <u>uterine cervix</u> may produce <u>vaginal discharge</u>, usually asymptomatic.
- Ascending infection in the form of <u>salpingitis</u> and <u>pelvic inflammatory disease</u>.
- The <u>scarring</u> produced by chronic or repeated infection is an important cause of <u>sterility</u> and <u>ectopic pregnancy</u>.

- three strains of C. trachomatis cause Lymphogranuloma venereum LGV,
 - L1, L2, or L3.
- It is characterized by
 - transient genital lesions
 - followed by <u>multilocular suppurative involvement</u> of the <u>inguinal lymph nodes</u>.
 - The primary genital lesion is usually a <u>small</u> <u>painless ulcer</u> or <u>papule</u>, which <u>heals</u> in <u>a few days</u>.
 - Abscesses, strictures, fistulas if <u>chronic</u>.

Neonatal chlamydia

- More than 50% of all <u>infants</u> born to mothers excreting *C. trachomatis* during labor show evidence of infection during the first year of life.
- Most develop <u>inclusion conjunctivitis</u>, but 5 -10% develop <u>infant pneumonia syndrome</u>.

- Conjunctivitis(trachoma)
- Urethritis
- Reactive arthritis (Ab attack the joint)
 - Reiter's syndrome
 - You cant see
 - You cant pee
 - You cant climb a tree



DIAGNOSIS

- Epithelial cells from the site of infection are required for detection.
- For genital infections, cervical specimens are preferred in females and urethral scrapings in males.
- Isolation of C. trachomatis has been the "gold standard" for diagnosis.
 - It is achieved in cell culture.

- Ligase chain reaction (LCR) or polymerase chain reaction (PCR)
 - the most sensitive, most specific methods of diagnosis.
- Serodiagnostic methods have <u>little use</u> in diagnosis of chlamydial genital infection
 - difficulty of distinguishing current from previous infection.

non-LGV C. trachomatis infection	Azithromycin
pregnant women and infants	Erythromycin
drug of choice for treating LGV	doxycycline

tetracyclines, macrolides and some fluoroquinolones

Chlamydia coverage: Azithromycin (Zithromax), 1 g PO in a single dose or Doxycycline, 100 mg PO bid x 7 days

Recommended treatment regimens for gonococcal infections: cervicitis, urethritis, proctitis

- Single dose + chlamydia coverages (Azithromycin or Doxycycline)
 - Cefixime (suprax)400mg or oflxacin 400mg or Cipro 500mg
 - Single large dose IM ceftriaxone 125mg
- Chlamydia coverage: Azithromycin (Zithromax), 1 g
 PO in a single dose or Doxycycline, 100 mg PO bid x 7 days

Ureaplasma and mycoplasma

- Smallest free living bacteria
- Atypical bacteria
- No cell wall
- Cell membrane contain sterols
- No cell wall inhibitors
- No microscopic
- No stain
- No fixed shape
- Slow growing
- Many female asymptomatic carrier
- M hominis, M genitalium, Ureaplasma urealyticum (facultatively anaerobic)
- M pneumonia (aerobic)

Ureaplasma urealyticum

- Cause genitourinary tract infections.
- highly pleomorphic,
 - may appear as coccoid bodies, filaments, and large multinucleoid forms.
- Ureaplasma is distinguished from Mycoplasma by its production of urease.

MANIFESTATIONS

- one half of cases of nongonococcal, nonchlamydial urethritis in men may be caused by *U. urealyticum*.
- In women, *Ureaplasma* has been shown to cause <u>chorioamnionitis</u> and <u>postpartum</u> <u>fever</u>.
 - The organism has been isolated from 10% of women with the latter syndrome.
- Pain, discharge, and itching of the genital area

DIAGNOSIS AND TREATMENT

■ Tetracycline is the treatment of choice because it is also active against Chlamydia,

protein synthesis inhibitors (tetracyclines and macrolides), as well as, bactericidal agents that inhibit DNA replication (fluoroquinolones).

- <u>Bacterial vaginosis</u> (BV),
 - formerly known as <u>nonspecific vaginitis</u>, was named because <u>bacteria</u> are the <u>etiologic agent</u> in this infection and an associated <u>inflammatory</u> <u>response</u> is <u>lacking</u>.
- BV is the most common cause of vaginitis.

Non-infectious vaginitis is often caused by an allergic reaction or an irritation from vaginal sprays, vaginal deodorants, spermicidal products, soaps, condoms if there is an allergy to latex

PATHOGENESIS

- Vaginal flora becomes altered, causing an increase in the local pH.
- This may result from a reduction in the H₂O₂ producing lactobacilli.

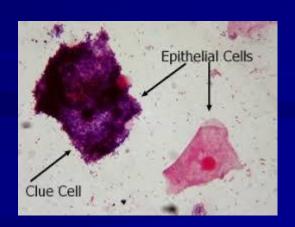
alteration in the vaginal ecology It is overgrowth

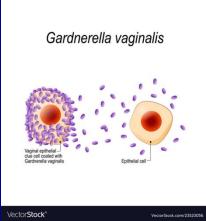
Gardnerella vaginalis

- G. vaginalis
 - facultatively anaerobic
 - gram-variable rod.
 - one of the organisms responsible for <u>bacterial</u> <u>vaginosis.</u>

Gardnerella vaginalis

- It is associated microscopically with <u>clue</u> <u>cells</u>,
 - epithelial cells covered in bacteria.
- Although BV is <u>not considered</u> a <u>sexually</u> <u>transmitted</u> disease,
- sexual activity has been linked to development of this infection.





CLINICAL ASPECTS MANIFESTATIONS

■ Symptoms of infection typically include a gray, thin, and homogeneous vaginal discharge that is adherent to the vaginal mucosa, associated with a "musty" or "fishy" odor.



CLINICAL ASPECTS MANIFESTATIONS

- there is <u>little vulvar</u> or <u>vaginal irritation</u> associated with this infection,
- the pungent odor is usually the chief complaint.

DIAGNOSIS

- A wet mount preparation of physiologic saline mixed with <u>vaginal secretions</u> should be examined under low- and high-power objectives.
- The characteristic "clue cells" are identified as <u>numerous stippled</u> or <u>granulated</u> <u>epithelial cells</u>.
- Cultures are seldom necessary to establish a diagnosis.

TREATMENT

- The treatment of choice for *G. vaginalis* is oral metronidazole, 500 mg twice daily for 6 days.
- The drug is <u>contra-indicated</u> during <u>early</u> <u>pregnancy</u> and <u>lactation</u>.