

# Urogenital Tract Module Sexually Transmitted Diseases (Syphilis) Lecture 6

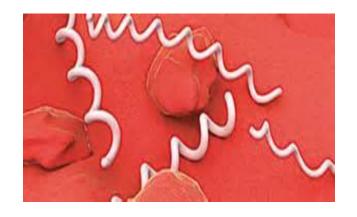
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# **Etiology of syphilis**

#### Caused by *Treponema pallidum* which is:

- A member of the spirochete family
- Spiral-shaped (corkscrew shape)





- Gram-negative, thin (0.1-0.2um in diameter) and 5 to 15um long
- Highly motile bacterium
- Rapidly die on drying
- killed by detergents and disinfectants
- Heat sensitive

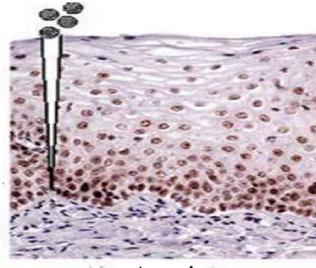
# **Transmission & Epidemiology**

- Direct contact with infectious lesions of skin and mucous membranes
- Most commonly occurs during sexual activity.
- Less commonly occurs during non sexual activity
  - Blood transfusions
  - Congenital (during pregnancy)
  - Direct contact with an infectious lesion
- No spread through contact with toilet seats, doorknobs, swimming pools, shared clothing, or eating utensils
- The infective dose 50 (ID50) is 60 organisms
- Groups at risk are:
  - √ Homosexuals (MSM)
  - ✓ Individuals with HIV
  - ✓ Lower socioeconomic classes in urban areas
  - ✓ Sex workers

# **Entry**

The spirochete can reach the subepithelial tissues through

- 1. Minute abrasions that occurs during sexual intercourse
- 2. Passage between the epithelial cells of mucous membranes



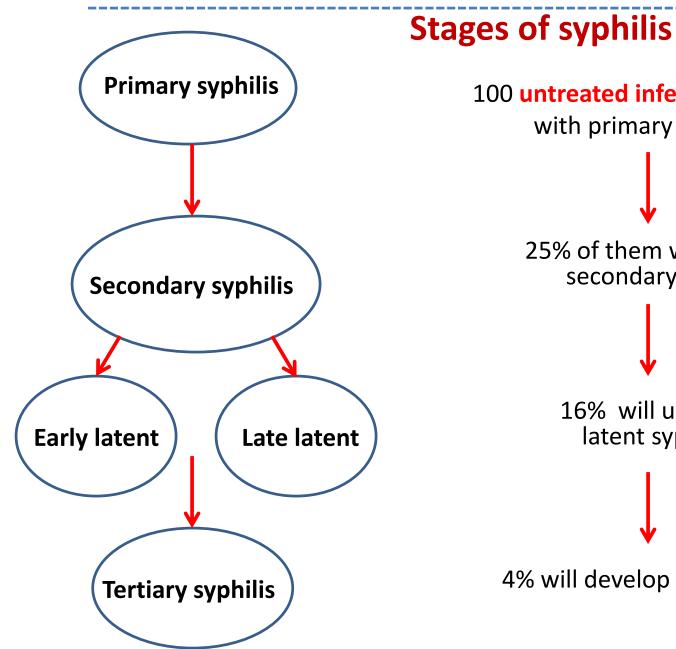
ectocervix



Endocervix

Then they replicate in the subepithelial tissue

From their they can escape to the systemic circulation



100 untreated infected patients with primary syphilis

> 25% of them will undergo secondary syphilis

> > 16% will undergo latent syphilis

4% will develop tertiary syphilis

# Pathogenesis and Clinical Manifestations Stages of syphilis

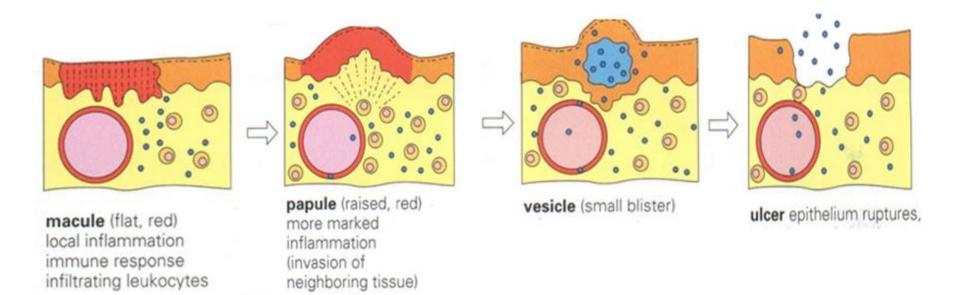
- ☐ *Treponema* causing obliterating endarteritis
- Obliterating endarteritis:

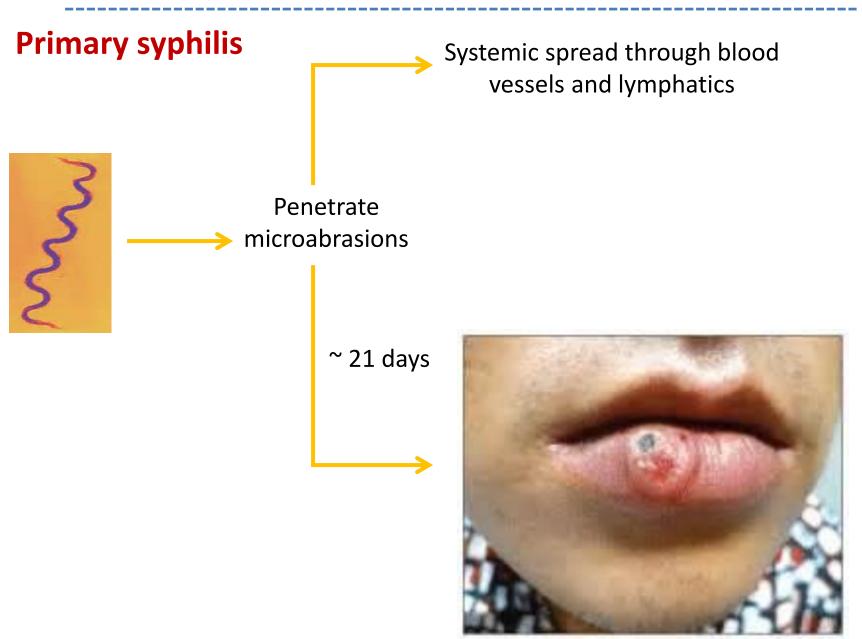
(inflammation of the intima or inner lining of an artery) that results in an occlusion of the lumen. This obstruction eventually causing necrosis, ulcers, and death of the affected tissue

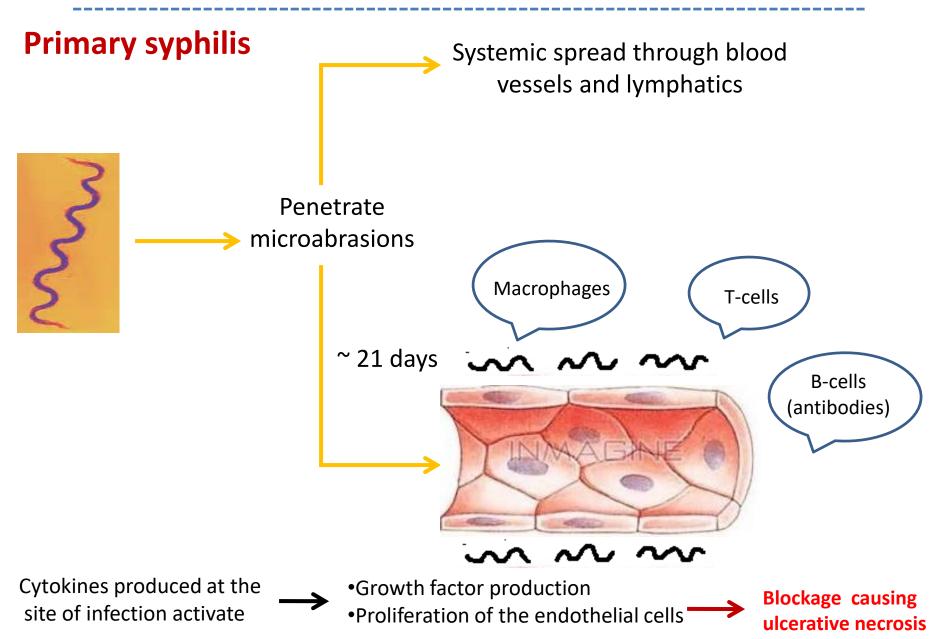


#### **Primary syphilis**

- The first symptom of syphilis is <u>chancre</u> (sore) develops between 10-90 days after exposure (mean of 21 days) at the site of infection
- ➤ The chancre is a painless ulcer, moist base with well defined and indurated margins



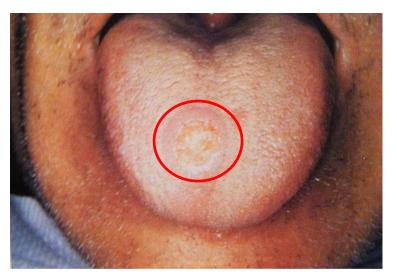




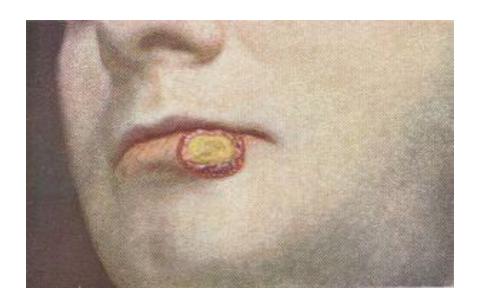
#### **Primary syphilis**

- Highly infectious
- Regional <u>painless</u> and <u>nontender</u> Lymphadenopathy (swollen glands)
- Systemic dissemination continues during this period
- The chancre resolves spontaneously over a period of 4-6 weeks without treatment to form a scar
- > 25% of untreated patients undergo the secondary syphilis

## **Primary syphilis (chancre)**









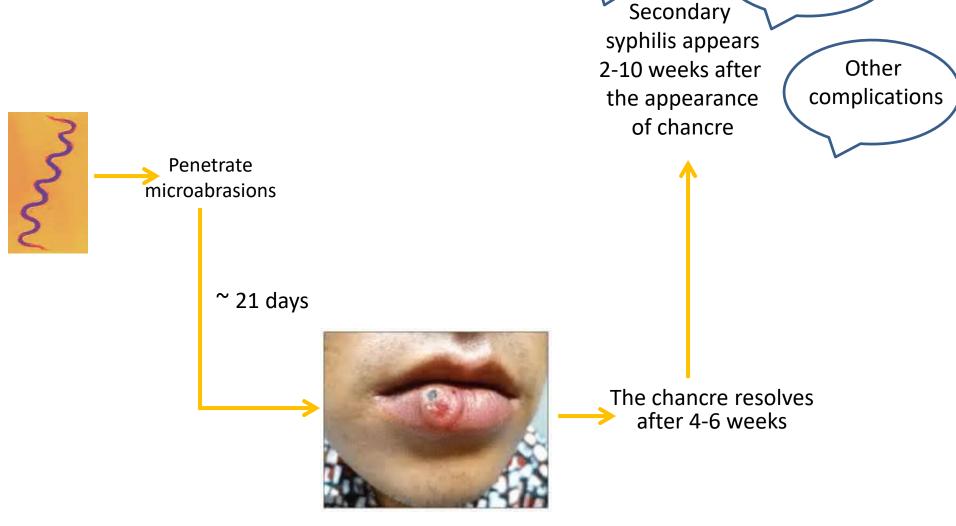
Constitutional

symptoms

maculopapular

skin rash

# **Secondary syphilis**



#### Secondary syphilis

- Secondary or disseminated syphilis develops 2 to 10 weeks after the appearance of the chance
- Infection spreads through the blood and lymph system
- Common sings & symptoms include:
  - generalized nontender lymph node enlargement
  - chancre may still be present
  - fever, malaise and other manifestations of systemic infection
  - mucocutaneous maculopapular skin rash
    - palms & soles
    - Abdomen
    - Face
  - condyloma lata: painless, mucosal, warty like rash, develop in warm, moist sites of the genitals and perineum.
  - May affect other organs

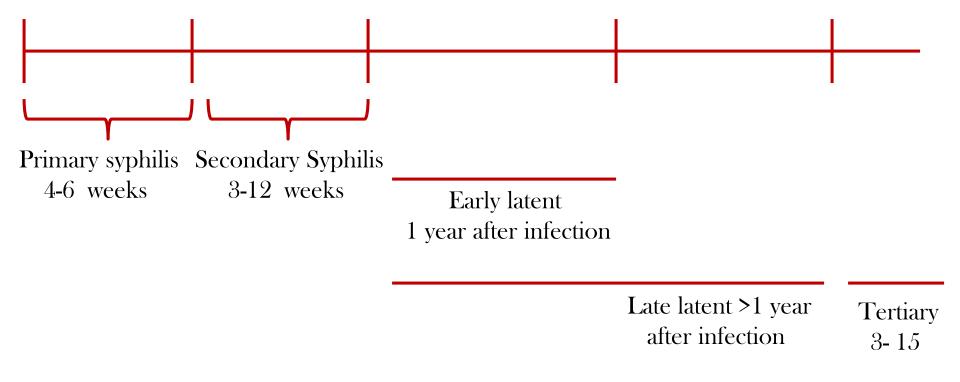
#### **Secondary syphilis**

- Lesions are highly infectious
- >Symptoms resolve spontaneously after 3-12 weeks
- ➤Infection is resolved in only 1/3 of patients
- The illness enters the latent state in the remaining 2/3 of patients









#### **Latent Syphilis (hidden)**

- Latent syphilis is defined as having serologic proof of infection without symptoms of disease
- Divided into:

#### a. Early latent syphilis

- One year or less post-infection.
- Occasional relapses of active lesions
- Infectious
- Transmittable from mother to child, resulting in congenital syphilis

#### b. <u>Late latent syphilis</u>

- is defined <u>as asymptomatic infection of longer than one year post-infection</u>.
- Non infectious period
- Transmittable from mother to child, resulting in congenital syphilis
- One third of patients with untreated latent syphilis develop tertiary syphilis



#### **Tertiary syphilis**

- The stage of end organ damage
- The major manifestations occur after 3 to 15 years of primary infection
- Three major forms

#### 1- Gummatous syphilis:

- an area of necrosis secondary to endarteritis obliterance
- most commonly found in liver (gumma hepatis), but can also be found in brain, heart, skin, bone, testis, and other tissues.
- the most common





#### **Tertiary syphilis**

#### 2- Cardiac syphilis:

aneurysm of ascending aorta, aortitis

#### 3- Neurosyphilis:

a. meningeal

Looks like meningitis

b. Meningeovascular

Looks like ischemic stroke

c. paranchymatous

Actual damage of neural tissues

#### Differential diagnosis of painless genital ulcer

1. Chancre:

(Tryponema pallidum)

- 2. lymphogranuloma venerum: *Chlamydia trachomatis* L1, L2, L3
- 3. Granuloma inguinale: Klebsiela granulomatis

# Pathogenesis and Clinical Manifestations Congenital syphilis

#### > Early congenital syphilis

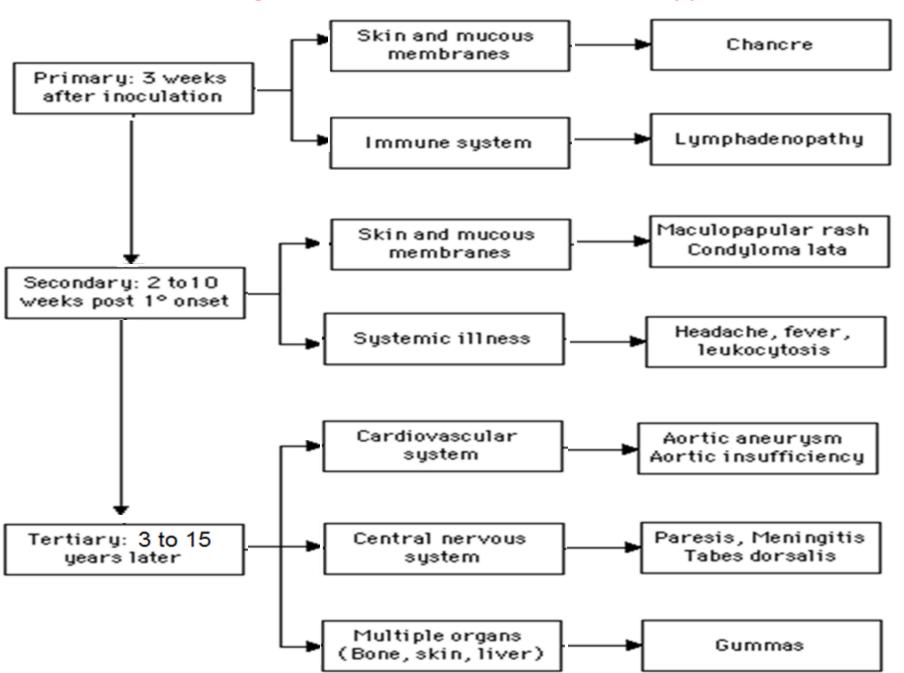
- Early manifestations appearing in the first 2 years of life.
- Resembles severe symptoms of adult secondary syphilis.
- The first symptom seen in up to 50% of newborns with congenital syphilis is snuffles (a nasal discharge in infancy characteristic of congenital syphilis).

#### > Late congenital syphilis

- Late manifestations appearing after 2 years.
- Ocular syphilis, arthropathy, gummas.



#### Stages and manifestations of syphilis



# Diagnosis

#### Syphilis diagnosis relies on:

- Dark-field microscopy:
  - is the most specific technique for diagnosing syphilis when an active chancre or condyloma latum is present
- Nonspecific serological testing (nontreponemal antibodies)
  - VDRL=Venereal Disease Research Laboratory
  - RPR=Rapid Plasma Reagin
- 3. Specific treponemal tests (confirmatory test)
  - FTA Antibody-Fluorescent treponemal antibody absorption
  - TPHA test- Treponemal pallidum haemagglutination assay
  - TPPA test- Treponemal pallidum particle agglutination assay



# **Syphilis**

Methods of laboratory diagnosis of syphilis:

- 2. Indirect detection of spirochetes:
  - A. Venereal Disease Research Laboratory (VDRL)
  - B. Rapid plasma reagin (RPR)

#### **Principle**

T. pallidum infection 

Destruction of cells

Release of cell components including the mitochondrial inner membrane lipid

Production of antibody against cardiolipin (nontreponemal antigens)

The agglutination between the reagin antibody from the patient serum and the cardiolipin (fom the ox heart) in the reagent indicates positive results

Agglutination under microscope

# Diagnosis

# Congenital syphilis:

Testing for IgM and retesting at 6 months of age, by which time maternal antibody levels have waned. Antibody titers remain elevated in babies with congenital syphilis

#### Treatment & Prevention

#### **Treatment:**

- Penicillin G (2.4 million units I.M)
- If allergic, Doxycycline or erythromycin is a good alternative
- In neurosyphilis use penicillin and Doxycycline together

#### **Prevention:**

- No vaccine
- Early diagnosis and treatment of case and contact is important
- Test for syphilis if any STD exists

# Case

Mr. B., a 24 years old homosexual man, came to the clinic with

- ✓ Fever
- √ swollen lymph nodes
- ✓ spotty discoloration of his skin, most notably on the palms and soles.
- ✓ He had recently noted a penny –sized, gray, translucent lesion on the inner aspect of his lower lip.

#### Mr. B reported that he engages in oral sex as well as anal-receptive intercourse

- Doctor recognized the macular rash on his palms and soles and the lesion on his lip as a characteristic of secondary syphilis.

#### The lab reported:

- Corkscrew appearance under darkfield microscope
- Positive serology for specific anti-treponemal antibodies

#### **Treatment**

He was treated with a course of penicillin and his lesions and symptoms abated

#### He was considered cured when the:

RPR drop 4 fold by 6 months(Test of cure)

# **Key Words**

**Spirochete** Treponema pallidum **Syphilis** Chancre **Primary lesion Darkfield microscopy Secondary Lesion Tertiary Lesion VDRL RPR**