

## Non-venereal diseases due to Treponema

- transmitted through person to person contact or via contaminated articles
- 1. bejel or non-venereal or endemic syphilis
  - caused by *T. pallidum* subspecies *endemicum*
  - usually begins in childhood as a small patch in the the mucous membranes
    - can involve the mouth
  - gradually develops into chronic inflammatory swellings on the limbs and trunk
    - including gummas which may develop on the skin bones and nasopharynx
  - encountered in tropical and subtropical areas of Africa and other continents

## 2. yaws

- causative agent is *T. pallidum pertenue*
  - infects through broken skin
- numerous manifestations which vary according to the stage of the illness include
  - painless papular nodules initially
  - later develops into destructive lesions involving
    - a. skin may form ulcers
    - b. lymph nodes
    - c. bones and joints
    - d. soft tissues
- encountered mostly in various areas of S. America  
Central Africa SE Asia

### 3. pinta

- causative agent is *Treponema carateum* or *T. pallidum carateum*
- primarily restricted to skin
- clinically characterized by
  - initial lesions as small pruritic papules
  - later lesions consist of enlarged plaques which persist for months to years
  - disseminated illness is characterized by recurrent hypopigmentation or depigmentation of skin lesions and extensive scar formation

## Borrelia

- relatively broader than other spirochaetes
- some species are stainable although not readily and observable microscopically
  1. by modified Gram's stain as Gram negative cells
  2. on dried blood films by Giemsa or Wright's stain

### Culture in the laboratory

- do not grow easily on cultures
- not requested for or performed routinely
- some species may grow in
  - a. enriched media specially formulated to provide necessary nutritional requirements
  - b. incubation under microaerophilic conditions

## Species of *Borrelia* which infect humans

### 1. *Borrelia recurrentis*

- causative agent of louse-borne relapsing fever or louse-borne borreliosis or epidemic relapsing fever

### 2. *Borrelia* transmitted by blood sucking ticks

- several species include
  - a. *Borrelia duttoni*
  - b. *Borrelia* species with species name corresponding with that of the tick responsible for transmission
- causative agents of tick-borne relapsing fever or tick-borne borreliosis or endemic relapsing fever

c. *Borrelia burgdorferi*

- isolated in some countries in the temperate regions
- transmitted by hard ticks from mammals including rodents
- causative agent of febrile illness with a wide spectrum of manifestations
  - illness referred to as Lyme disease

3. *Borrelia vincentii* normal flora in the mouth

- significant as a causative agent of ulcerative lesions in the mouth in association with other abnormalities including
  - a. trauma
  - b. malnutrition
  - c. other organisms in the mouth

## Transmission of *Borrelia* species associated with relapsing fever

### *Borrelia recurrentis*

- transmitted by lice from person to person
- infected humans with the organisms in the blood form the reservoir
  - lice acquire the organisms while feeding on infected human blood
- enters susceptible person through broken skin
- rubbing or crushing the infected lice releases fluid containing spirochaetes and facilitates entry
- over-crowding in places with scarce facilities for cleanliness favours transmission

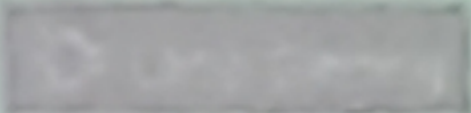
## *Borrelia duttoni* and related tick-borne species

- humans acquire the organisms through broken skin
  - from a tick bite or from crushed infected tick
- ticks
  - transmit the borreliae between animals and from animals to humans
  - infected animals involved in transmission include
    - a. rodents b. pigs c. armadillos d. porcupines
    - together with infected ticks constitute the reservoir
  - infection in ticks
    - a. persists throughout its life
    - *D. duttoni* is passed on via the ova to next generations



## Manifestations of relapsing fever

- louse-borne or tick-borne      similar manifestations
    - characterized by febrile and afebrile periods
  - febrile period
    - lasts approximately 3 to 5 days
    - spirochaetes are abundant in the blood
    - immune system responds by production of antibodies to specific antigenic components
    - together with other immune defense mechanisms antibodies recognize and act to eliminate the spirochaetes in the circulation at the time
  - overall effects
    - marked reduction of borreliae in the blood
- fever subsides

- fever recurs after 4 to 10 days and spirochaetes reappear in the blood causing a relapse
- several recurrences or relapses occur
- during each febrile episode
  - few borreliae undergo mutation
    - acquire new antigenic determinants which differ structurally from the previous ones
    - phenomenon of antigenic variation
  - organisms with new antigenic determinants
    - constitute a new strain which is not recognized by the circulating antibodies at the time and therefore are not eliminated
-  multiply and cause the subsequent relapse

## Laboratory investigation of relapsing fever

1. detection of Borrelia species in peripheral blood
  - chances of detection are high during the febrile periods by
    - a. direct microscopic examination for motile spiral organisms
    - b. examination of dry smears stained by Giemsa or Wright's stain
    - c. dark field or phase contrast microscopic examination for motile spiral organisms
      - more reliable procedure
2. other methods of investigation

## Antibiotic susceptibility of Borrelia

- sensitive to several agents including penicillin

## Leptospira

- narrower than other spirochaetes
  - possesses numerous closely set coils
- recognized by its characteristic vigorous motility
- visualized by dark field or phase contrast or electron microscopes
- stainable by special methods including fluorescent staining methods
- culturable in specifically formulated enriched media
  - liquid or semi-solid preparations as obligate aerobes
  - slow growth especially in primary isolation
    - may take 6 to 14 days or longer
- also capable of growing on embryonated eggs

## Species of Leptospira

- composed of saprophytic organisms and potential pathogens
- two species morphologically indistinguishable
  - each species is composed of numerous serotypes
- 1. *L. interrogans*
  - one of the serotypes is a parasite of rodents and a potential human pathogen serotype *icterohaemorrhagiae*
- 2. *L. biflexa*
  - saprophytes in the environment

## *Leptospira interrogans* serotype associated with human disease

- parasitic to small animals including rodents
  - found in their kidneys
- no disease manifestations in the animals
  - can also infect other animals including domestic animals
  - the infected small animals provide the reservoir
- passed out in the urine and contaminates the environment
  - saprophytes in water or damp soil
- transmission to humans is by accidental exposure through contact with infected urine of the animal

- directly or through contaminated soil or water

### Clinical implication

- causative agent of human leptospirosis
  - entry is through broken skin or mucous membranes
  - infection is associated with invasion into the blood circulation and septicaemia
  - associated with haematogenous spread to various internal organs including the kidneys
    - may be excreted in urine
  - increased risk of exposure for people who work in damp places populated by rodents

## Manifestations of human leptospirosis

- vary from asymptomatic to severe illness
- symptomatic leptospirosis
  - initially characterized by septicaemia constituting the septicaemic phase
    - manifestations include mild non-specific signs and symptoms including fever
  - may develop into a severe illness involving an internal organ
    - resulting in manifestations due to abnormal functions of the organ



## Laboratory investigation of leptospirosis

- involves 1. detection of spirochaetes in specimens  
2. serological tests
  - specimens include 1. blood 2. urine 3. tissue
- procedures
1. culture
  2. dark field microscopy
    - chances for positive results on blood are higher during the septicaemic phase
  3. serology
    - detection and demonstration of rising antibody titre in serum
  4. DNA detection and amplification in specimen

## Antimicrobial susceptibility of *L. interrogans*

- effective antimicrobial agents include penicillin and tetracycline

Prevention and control of *L. interrogans* infection may involve

- reduction of infection in wild and domestic animals
- vaccination of domestic animals in areas where *L. interrogans* is detected
- protective clothing for those at higher risk of exposure