

# PLAGUE

## Epidemiology

### Source:

Rodents, carnivores, & their fleas

### Transmission

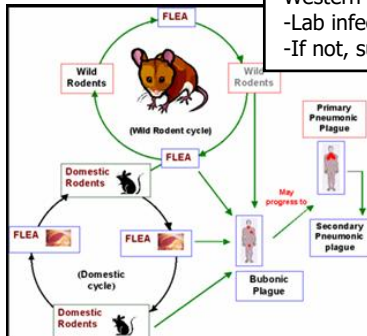
- Bites of infected rodent fleas
- Bites from infected human fleas (*Pulex irritans*)
- Direct contact with tissues/fluid of infected animals (bubonic, septicemic)
- Droplets from infected animal or human (pneumonic)
- Exposure to laboratory aerosols (pneumonic)

### Infectious dose

10-500 organisms

### Not in Louisiana

- Imported from Western US or foreign
- Lab infection
- If not, suspect BT



### Incubation

Bubonic: 2-8 d  
Septicemic: 1-6 d  
Pneumonic: 1-3 d

Contagious up to 48 hrs after Tx (pneumonic)

### Clinical case definition

Abrupt onset of fever, chills, headache, malaise  
**-Bubonic:** swollen lymph nodes (buboes) esp. in inguinal, axillary, cervical regions.  
**-Septicemic:** hypotension, acute respiratory distress, intravascular coagulopathy  
**-Pneumonic:** cough, fever, dyspnea, hemoptysis

### Communicability

- Bubonic: by fleas, infectious for weeks by contact with pus while symptomatic
- Septicemic: blood & body fluid exposure
- Pneumonic: droplet while symptomatic

Mortality: Asia=54%, Africa=34%, Americas=11%.

**Complications:** sepsis with renal failure, acute respiratory distress syndrome, hemodynamic instability, diffuse intravascular coagulation, necrosis of distal extremities

**Morbidity:** 50% bubonic and septicemic, 90% pneumonic if treatment is not started within 18 hrs.



*Xenopsylla cheopis*

A flea infected with *Yersinia pestis*, shown as a dark mass. The foregut of this flea is blocked by a *Y. pestis* biofilm, which is a prerequisite for efficient transmission.



### PNEUMONIC

Central: -Headache

Respiratory: -Cough, -Hemoptysis, -Dyspnea, -Chest pain

Muscular: -Weakness

### BUBONIC

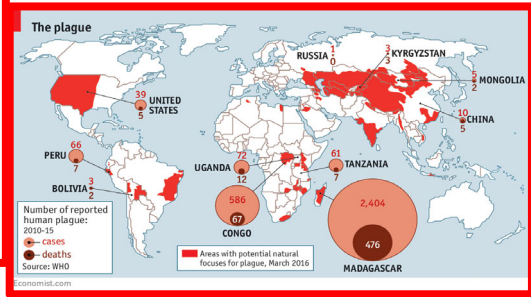
Invasion of a lymph node → Pus formation in the marginal sinuses → Lymph node → enlarged & bursts open

### SEPTICEMIC

- Attacked the blood system (Blood Poisoning)
- Fevers
- Skin turns deep shades of purple due to DIC (disseminated intravascular coagulation)

### ENVIRONMENTAL RESISTANCE

- Low resistance to environmental factors.
- Sunlight, high temperatures and desiccation have a destructive effect,
- Ordinary disinfectants as Lysol® & chlorine preparations kill it within 1 to 10 minutes.



## Diagnosis

*Yersinia pestis* is a nonmotile, gram-negative bacillus that belongs to the *Enterobacteriaceae* family.

### Lab Diagnosis

- **Culture-** Culture of *Y. pestis* confirmed from blood, bubo aspirate (bubonic), sputum or tracheal wash (pneumonic), and CSF.
  - Gram stain shows organism with bipolar (safety-pin) morphology.
  - Direct fluorescent assay (FA) shows bacilli
- **Immunofluorescence** - positive for *Y. pestis* F1 antigen.
- **Serologic test-** 4-fold difference in antibody titer between 2 serum specimens (4 wk apart)
- **Polymerase chain reaction** -for rapid diagnosis.
- Microbiology lab should be informed of suspected cases to minimize risks of transmission.

- Suspect:** Clinically compatible case
- Probable:** Clinically compatible
  - Elevated serum antibody titers to *Y. pestis* F1 antigen,
  - F1 antigen by fluorescent assay
- Confirmed:**
  - Isolation of *Y. pestis*
  - Detection by PCR,
  - Fourfold change in serum antibody titer to *Y. pestis* F1 antigen

## Control

### EPIDEMICS

- ❑ Human infection most frequently occurs when an epizootic develops among synanthropic rats in centers of human population, following contact with infected wild rodents.
- ❑ Commensal rat fleas, including plague-infected fleas, leave the bodies of rats killed by plague seeking a blood meal
- ❑ Following the death of rodents during an epizootic in a natural focus. The fleas can accumulate at the entrance to and the ground surface around burrows. The fleas are not strictly species-specific.
- ❑ Exposure to droplets from a pneumonic plague
- ❑ Bioterrorism possible

### PLAGUE LANDSCAPE

- ❑ Climatic conditions are favorable for a high and stable number of rodent reservoirs and flea vectors of *Y. pestis*.
- ❑ Most natural foci = mountains, low annual precipitation, dry seasons inhibiting growth of thick woody vegetation → formation of deserts, semi-deserts and steppes (savannas, prairies, pampas and so on).



### RODENT SPECIES

Many species of rodents / other small mammals are susceptible to infection but only occasionally infected and not necessarily important reservoirs of infection.

- ❑ **Enzootic** (maintenance) hosts: rodents from genera that are relatively resistant to plague, low mortality, positive antibody rate up to 100%. Die-offs rare.
- ❑ **Epizootic** (amplification) hosts. Introduction into a highly plague-susceptible species to plague, an epizootic may reach considerable magnitude with high mortality

### FLEA SPECIES

To understand epidemiology and transmission it is essential to determine the flea species involved

Fleas specific to commensal rodents with wide distribution found in several endemic areas:

- ❑ *Xenopsylla cheopis* (Oriental rat flea) has a wide distribution,
- ❑ *X. brasiliensis* and *Nosopsylla fasciatus* is more limited
- ❑ Restricted distribution, *X. astia*.
- ❑ Flea species common in the environment of commensal rodents, not specific for them. *Echidnophaga gallinacean*, *Pulex irritans*, and the cat flea, *Ctenocephalides felis*.

### EPIDEMIOLOGIC INVESTIGATION

An epidemiological investigation should be performed for each human case to determine **source of infection and risk of additional human cases**. Reports of these investigations should include:

- Complete history of the patients' activities and travel during the incubation period of the infection;
- Results of field studies to determine which animal and flea species are likely sources of infection or pose a continuing threat to humans;
- Proximity of infected rodents and fleas to human dwellings or workplaces;
- Estimated number of people involved in activities that place them at high risk of plague infection;
- Information on possible exposure to *Y. pestis* infection of patient contacts (especially important for pneumonic plague cases).

## Treatment, Prophylaxis

### Standard & Droplet Precautions (until 24 hrs after treatment)

#### TREATMENT

- **Streptomycine:** 30mg/kg Max 2g/day, IM, div in 2 doses/day, 10 days
- **Tetracycline:** Oral loading dose 15mg/kg, max 1g, then 25-50mg/kg (max 2g), PO
- **Doxycycline:** 200mg/day, 1 or 2 doses, PO
- **Sulfamide:** Higher mortality
- **Gentamycin:** 3mg/kg/day, 3 divided doses, IM or IV
- **Children: streptomycin** (30 mg/kg per day in 2 or 3 divided doses given intramuscularly) or **gentamicin** (6-7.5 mg/day), IM
- Pneumonic plague = Streptomycin
- Alternative drugs: Tetracycline, doxycycline, chloramphenicol, trimethoprim-sulfamethoxazole, ciprofloxacin Alternative drugs.
- Duration 7-10 days or until several days after lysis of fever.
- Drainage of abscessed buboes may be necessary.

**Household contacts** or anyone with face-to-face exposure to a plague patient should report fever >38.5°C or other signs of illness to their physician.

#### VACCINATION

- Worldwide, live attenuated and formalin-killed *Y. pestis* vaccines are variously available for human use.
- Variably immunogenic and moderately to highly reactogenic.
- No protection against primary pneumonic plague.
- In general, vaccinating communities against epizootic and enzootic exposures rarely feasible; little use during human plague outbreaks, since a month or more is required to develop a protective immune response.
- Indicated for persons whose work routinely brings them into close contact with *Y. pestis*, such as laboratory technicians in plague reference and research laboratories and persons studying infected rodent colonies.

#### PROPHYLAXIS

- Indicated for people with close exposure (<2m) to a patient with pneumonic plague
- Anyone exposed to pneumonic plague with a >38.5°C fever or any other symptoms
- Children: trimethoprim-sulfamethoxazole, 40mg/kg, 2 doses/day, PO
- Adults: doxycycline, ciprofloxacin, trimethoprim-sulfa Adult 1-6g/day, 2 doses/day, PO
- Given for 7 days in usual therapeutic doses