



RICKETTSIAL DISEASES

MEDICAL ACADEMY NAMED BY S.I.GEORGIEVSKY "CFU NAMED BY V.I.VERNADSKY

DEPARTMENT OF MEDICAL BIOLOGY

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INTRODUCTION ABOUT TICKS

| | |
|----------------------------------|--------------------------|
| Domain: | Bacteria |
| Phylum: | Proteobacteria |
| Class: | Alphaproteobacteria |
| Order: | Rickettsiales |
| Family: | Rickettsiaceae |
| Genus: | <i>Rickettsia</i> |
| Species group: | Spotted fever group |
| Species: | <i>R. conorii</i> |
| Binomial name | |
| <i>Rickettsia conorii</i> | |

| Scientific classification | |
|----------------------------------|---------------------------|
| Kingdom: | Bacteria |
| Phylum: | Proteobacteria |
| Class: | Gammaproteobacteria |
| Order: | Legionellales |
| Family: | Coxiellaceae |
| Genus: | <i>Coxiella</i> |
| Species: | <i>C. burnetii</i> |
| Binomial name | |
| <i>Coxiella burnetii</i> | |

GENERAL CHARACTERISTICS

Structurally similar to gram (-) bacilli

DNA S RNA Enzymes for Kreb's cycle Ribosomes for protein synthesis

Inhibited by antibiotics → Tetracycline & Chloramphenicol

Originally thought to be viruses Small size Stain poorly with gram stain

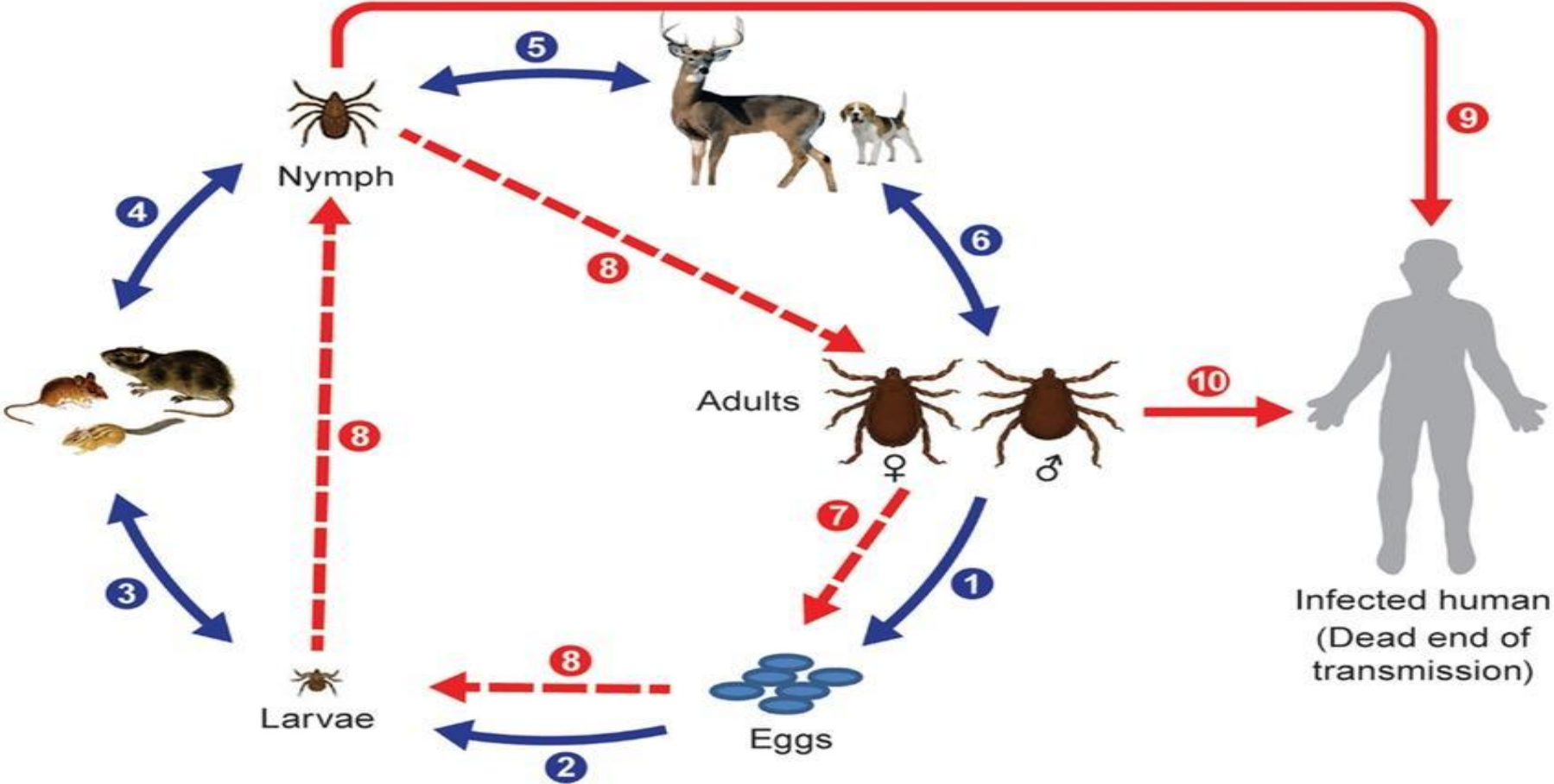
Grows only in cytoplasm of Eukaryotic cells Obligate intracellular parasites

EXCEPT Coxiella Rickettsia survival depends on entry, growth, and replication within the cytoplasm of eukaryotic host cells.

That's why they cannot live in artificial nutrient environments and are grown either in tissue or embryo cultures. Reservoirs animals & arthropods



LIFE CYCLE



Life cycle

The most common hosts are ticks.

Ticks that carry rickettsia fall into the family of Ixodidae ticks also known as “hard bodied” ticks.

Ticks are vectors, reservoirs and amplifiers of this disease.

There are currently three known tick specific that commonly carry rickettsia

American dog tick (*Dermacentor variabilis*)

Rocky Mountain Wood Tick (*Dermacentor andersoni*)

Brown dog tick (*Rhipicephalus sanguineus*). Ticks can contract rickettsia by many means. First,

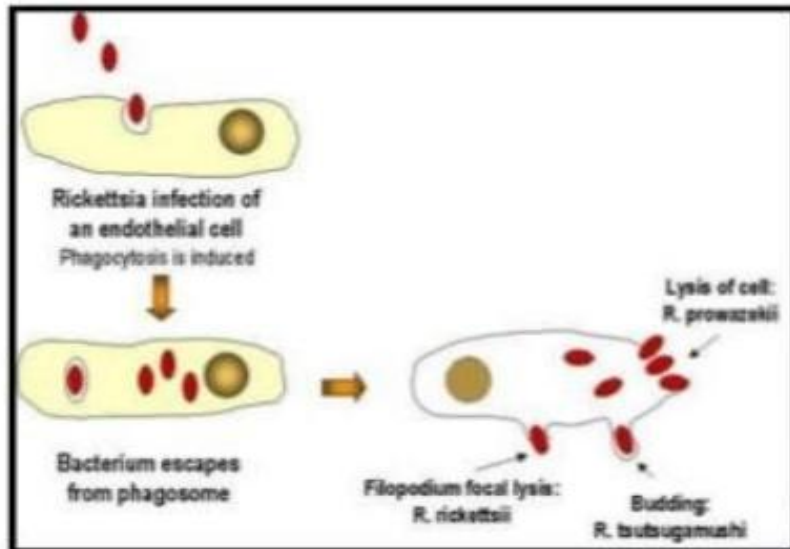
an uninfected tick can become infected when feeding on the blood of an infected vertebrate host; such as a rabbit, during the larval or nymph stages, this mode of transmission called transstadial transmission.

Once a tick becomes infected with this pathogen, they are infected for life. Both the American dog tick and the Rocky Mountain wood tick serve as long-term reservoirs for *Rickettsia rickettsii*, in which the organism resides in the tick posterior diverticulae of the midgut, the small intestine and the ovaries. citation needed In addition, an infected

Etiology

- ▶ Treatment Without waiting for laboratory confirmation antibiotic therapy should be instituted when rickettsial disease is suspected.
 - ▶ Doxycycline is the drug of choice Doxycycline (100 mg bid orally for 7-15 days), or azithromycin (500 mg orally for 3 days) children aged < 8 years and pregnant females-
 - ▶ azithromycin clarithromycin Oral treatment is used unless patient is vomiting or complicated Additional benefits of azithromycin
 - ▶ long tissue half life and the long lasting post-antibiotic effect causes no relapse despite the use of a single dose
2. Safe in child and pregnancy

Pathogenesis:



Obligate I/C organisms

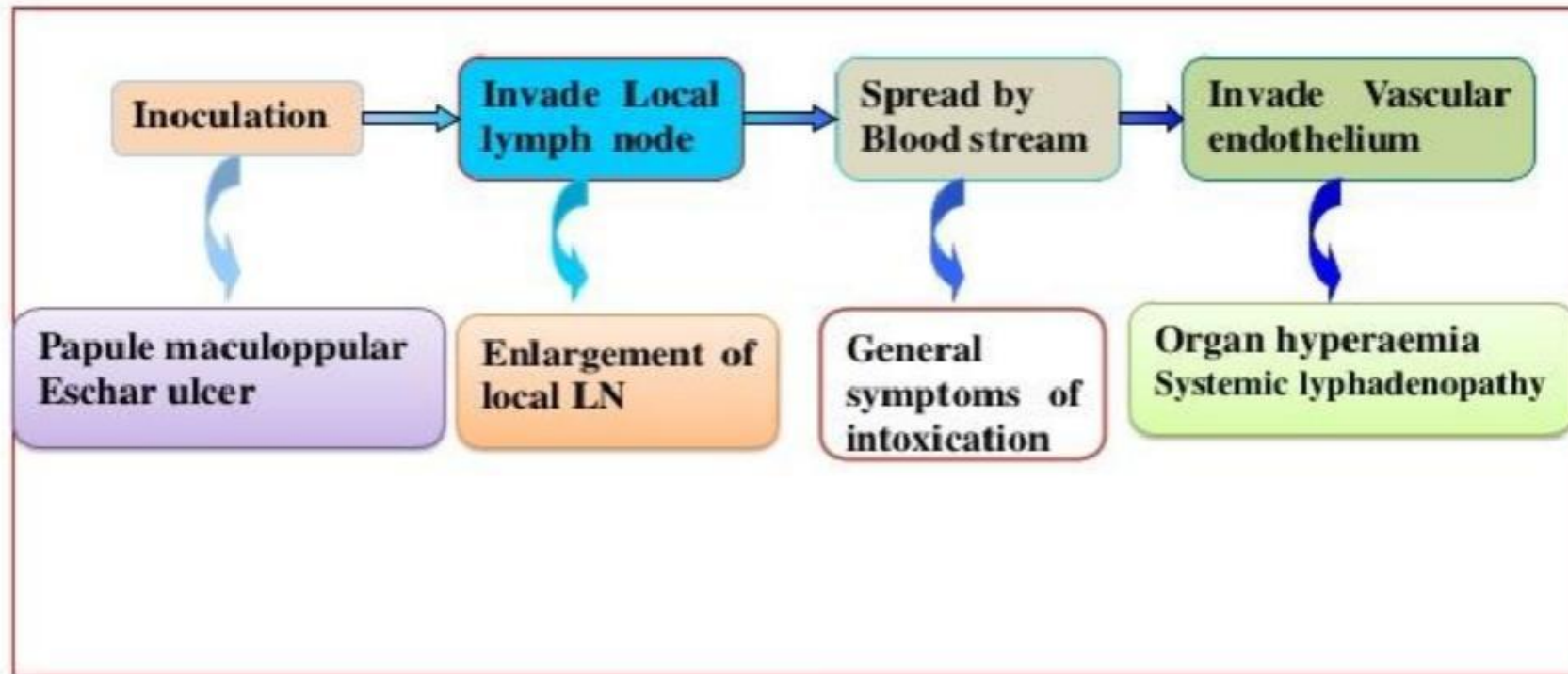
Dermal target cells (fibroblasts, macrophages, dermal dendritic cells, and lymphatic endothelium)

Engulfed by mononuclear phagocytes in the blood stream

Lungs and brain - lethality of rickettsioses

Rickettsia rickettsii is the only organism in the genus that invades beyond the blood vessel lining endothelium; they invade adjacent vascular smooth muscle cells, particularly in arterioles

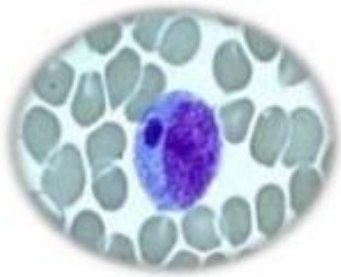
Pathogenesis



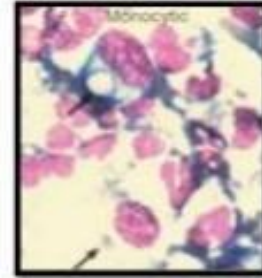
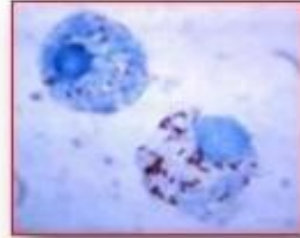
Diagnosis:

Staining:

- Not stained by the Gram's method
- Retain basic fuschin when stained with Gimenez method- Bright red



- **Initial diagnosis**
 - History, clinical signs
 - Hematologic abnormalities
 - Serum chemistry
- **Definitive diagnosis**
 - IFA
 - ELISA
 - PCR
 - Culture



Clinical
Signs

Blood
Smears

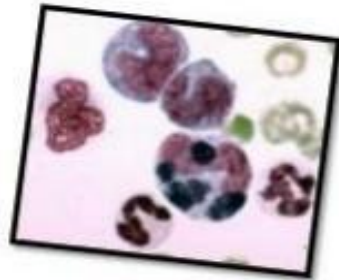
Serological
Tests

Biopsy

Gene
Diagnosis

Diagnosis

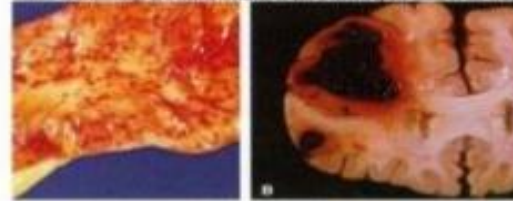
- A dog with fever, enlarged lymph nodes, bleeding, or arthritis in multiple joints.
- Low platelet numbers, high globulin levels, and mild anemia on blood testing



Canine thrombocytopenia

POSTMORTEM LESIONS

if hemorrhages of the colonic mucosa, seen here as a consequence of thrombocytopenia



Pancytopenia

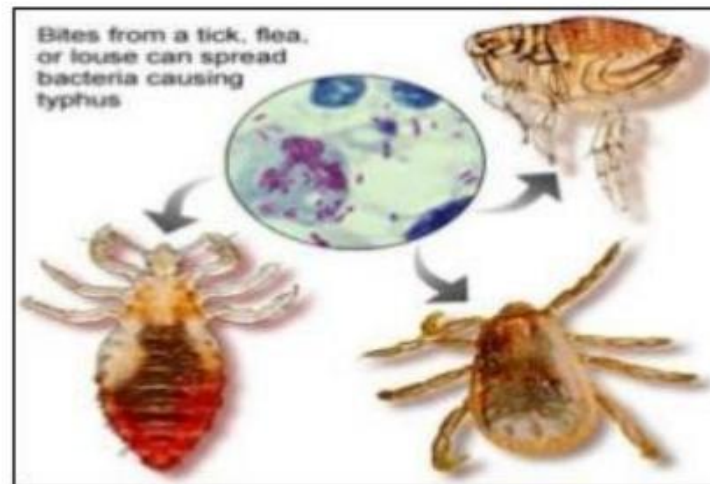


Ehrlichia canis seen in a membrane-bound inclusions (morulae) within the cytoplasm of a monocyte (buffy coat smear, Wright stain)

RICKETTSIAL INFECTIONS

Classified into groups:

- 1 **Typhus Group**
- 2 **Spotted Fever Group**
- 3 **Scrub typhus**



INDIAN SCENERIO: A GLIMPSE

Has been reported
from 11 states in
India:

Jammu and Kashmir

Himachal Pradesh

Uttaranchal

Rajasthan

Assam

West Bengal

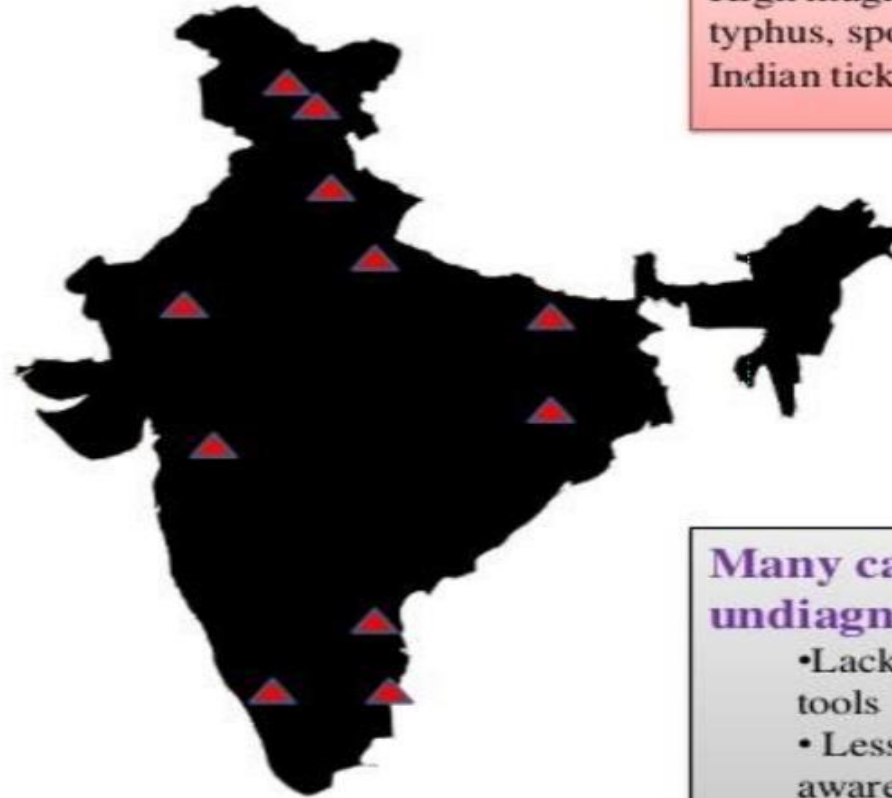
Maharashtra

Pondichery

Kerala

Tamilnadu

(Mahajan S. K., 2012)



Re-emerging

High magnitude of scrub
typhus, spotted fever and
Indian tick typhus

(Batra HV, 2007)

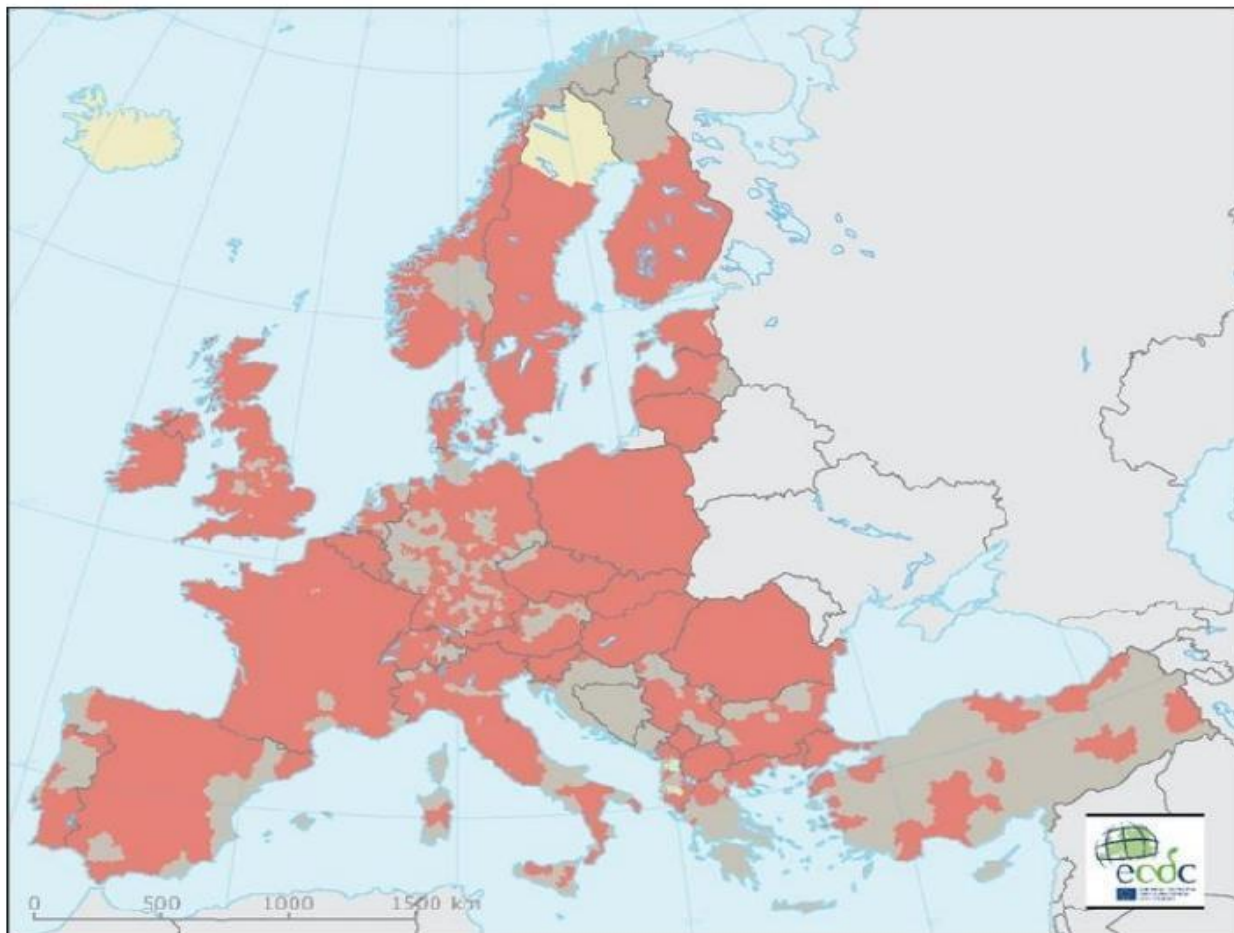
Many cases go undiagnosed ???

- Lack of diagnostic tools
- Less clinical awareness
- Lower index of suspicion

Rickettsia: Treatment and Prevention

- RMSF
 - Doxycycline drug of choice
 - Avoid ticks, wear protective clothing, use insect repellents, insecticides
 - In infested areas, check and remove ticks immediately
- Typhus Fever
 - Doxycycline effective
 - Improve personal hygiene and living conditions, reduce lice by insecticides, control rodent population
 - Inactivated vaccine for epidemic typhus





**Current distribution
of *Ixodes ricinus* ticks in
Europe, 2016**

Presence

- Present
- Introduced
- Anticipated absent
- Unknown
- Outside coverage

| Features | Scrub typhus | Murine typhus | Indian tick typhus | Epidemic typhus | Rickettsial-pox | Rocky mountain fever | Q fever |
|----------------------|-----------------------------------|------------------------------|--------------------|----------------------|---------------------|----------------------|--------------------------|
| Agent | <i>Rickettsia tsutsugamushi</i> | R. moser | <i>R. conorii</i> | <i>R. prowazekii</i> | <i>R. akari</i> | <i>R. rickettsii</i> | Coxiella burnetii |
| Reservoir | Mite | Commensally rats | Ticks | Man | Domestic rodents | Wild rodents | Cattle |
| Insect vector | Mite | Flea | Tick | Louse | Mite | Tick | - |
| IP | 10-12 days | 1-2 weeks | 3-7 days | 1-2 weeks | 7-9 days | 5-7 days | 2-3 weeks |
| Eschar | Present | Absent | Present | Absent | Present | Present | Absent |
| Rash | Centrifugal | Centrifugal | All over body | Centrifugal | All over body | Centripetal | No rash |
| Treatment | Tetracycline | Tetracycline | Doxy | Tetra | Doxy | Doxy | Doxy |
| Remark | Most common rickettsial infection | Also known as endemic typhus | Atypical measles | - | Atypical chickenpox | - | Transmitted by aerosol |

RICKETTSIA IN CRIMEA

- ▶ territory the Crimean peninsula is endemic in a number of transmissible natural focal infections, including the Mediterranean spotted fever (Marseilles fever)
- ▶ the causative agent of which is *Rickettsia conorii*.

In Crimea, a disease similar to the Mediterranean spotted fever was first described in 1938

At presentation, there is a tendency to the more severe clinical presentation of the Mediterranean spotted fever in the territory of Crimea, which is evident from the pre-dominance ones, the intensification of intoxication syndrome

as well as the appearance of various atypical forms of the disease caused by this pathogen. The first lethal case caused by this infection was registered in Crimea in 1996.

The true incidence of the Mediterranean spotted fever in Crimea is probably much higher than of moderately severe forms over the light that officially recorded, since the planned examination of the blood serum of 350 healthy donor individuals revealed antibodies to *R. sibirica* antigen in donors only in 2014

. 14 cases of the Mediterranean spotted fever were recorded 7 of which were laboratory-confirmed by one of the variants of the PCR method. All identified cases of the disease had a clinical form of moderate severity. In Crimea, the territory of depending on the etiological agent, as well as the physical conditions, 16 patients with this infection were registered; one patient had a severe form of the disease



| Species | Disease | Tick species associated (literature data) | Tick species associated (present study) | Fever, % | Diffuse rash, % | Eschar, % | Lymphadenopathy | Lymphangitis | Reported mortality rates,% |
|-----------------------------------|---------------------------------------|--|---|----------|-----------------|---------------------|-----------------|----------------|----------------------------|
| <i>R. conorii conorii</i> | Mediterranean spotted fever | <i>Rh. sanguineus</i> , <i>Rh. simus</i> (?) | <i>Rh. e. evertsi</i> | 100 | 97 | 72 | Rare | No | 1–5 |
| <i>R. africae</i> | African tick-bite fever | <i>A. variegatum</i> , <i>A. hebraeum</i> , <i>Rh. (B.) microplus</i> , <i>Rh. (B.) decoloratus</i> | <i>Rh. e. evertsi</i> , <i>Rh. (B.) annulatus</i> | 92 | 43 | 98%, often multiple | Yes | No | Very low |
| <i>R. sibirica mongolitimonae</i> | Lymphangitis-associated rickettsiosis | <i>H.truncatum</i> , <i>H. asiaticum</i> , <i>H. anatolicum excavatum</i> , <i>Rh. pusillus</i> | <i>H.truncatum</i> | 100 | 78 | 89% | 55% | Yes, up to 44% | 0 |
| <i>R. aeschlimannii</i> | Unnamed | <i>H. m. marginatum</i> , <i>H. m. rufipes</i> , <i>H. aegyptium</i> , <i>Haemaphysalis inermis</i> | <i>H.truncatum</i> , <i>H. m. rufipes</i> , <i>Rh. e. evertsi</i> | Yes | Yes | Yes | ? | ? | 0 |
| <i>R. massiliae</i> | Unnamed (one case) | <i>Rh. sanguineus</i> , <i>Rh. turanicus</i> , <i>Rh. muhsamae</i> , <i>Rh. lunulatus</i> , <i>Rh. sulcatus</i> | <i>Rh. guilhoni</i> | Yes | Yes | Yes | ? | ? | 0 |

Conclusion

- Scrub typhus is a significant public health threat in India, yet it is under-recognised and grossly under-diagnosed
- Doxyxcycline is the drug of choice
- Azithromycin is a good alternative
- Unavailability of IV doxycycline to treat severe infection necessitates urgent advocacy
- Information and advocacy are needed for investing in surveillance, prevention and detection, as well as appropriate management strategies

Video link references

<https://youtu.be/B8D0JeonHiU>

<https://youtu.be/tcg-iShuQt4>

<https://youtu.be/XaLBynjL000>

▶ **STAY HOME**



STAY SAFE

STAY HAPPY