### VISCERAL LEISHMANIASIS

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# **Objectives**

- Scientific names of causative organisms
- Life cycle- morphological forms, mode of transmission, vectors & reservoir hosts
- Epidemiology of VL
- Clinical manifestations
- Diagnosis
- Management
- Prevention & control

# LEISHMANIASES

- About 100 million people affected in Africa, Asia, Middle East, Europe Central & South America (about 2 million new cases each year)
- Sri Lanka
  - A newly established infectious disease
  - Previously only cutaneous form was seen
  - Recently few cases of mucocutaneous and visceral forms diagnosed

## Leishmaniases.....

 Caused by flagellate protozoan parasites of the order 'Kinetoplastida'

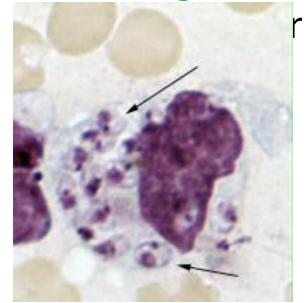
- Species of Leishmania causing VL
  - Leishmania donovani
  - Leishmania infantum

 VL is also known as 'Kala-azar' (black disease), or dumdum fever

# Morphology of Leishmania sp.

2 morphological forms

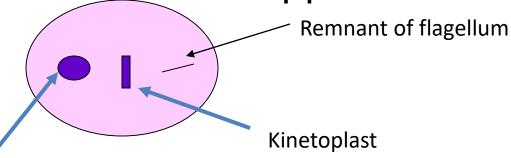
Amastigote (Leishman-Donovan bodies)



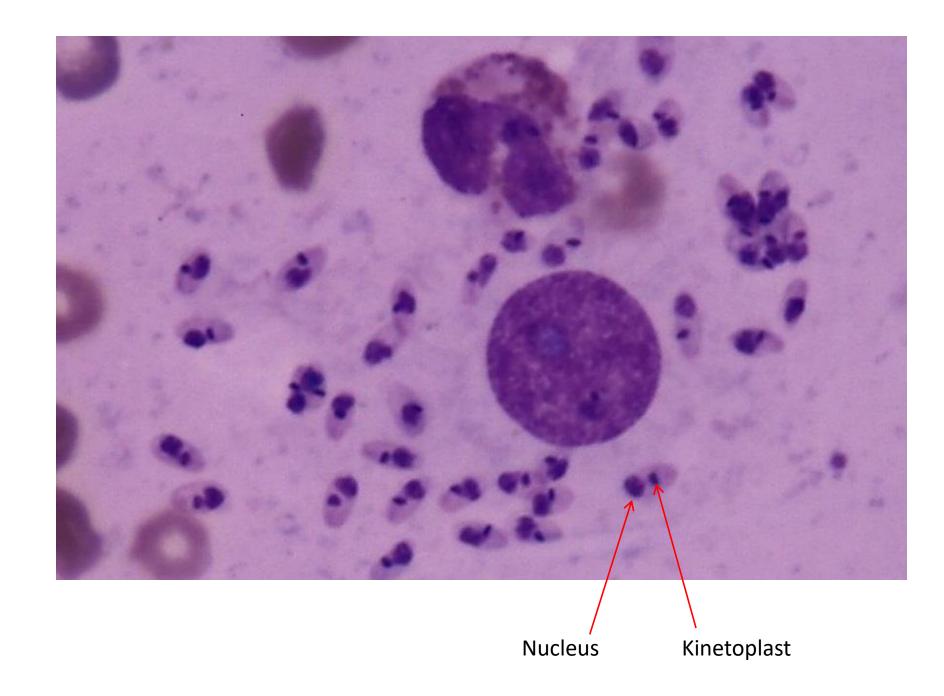
2-6 microns in size

r (within macrophages)

- -found in vertebrate host
- -'dot and dash' appearance



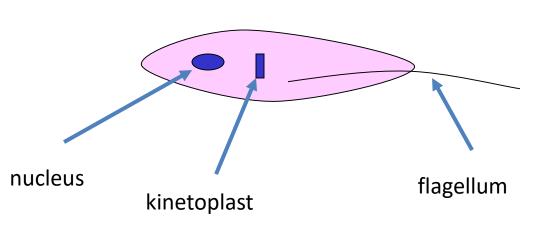
**Nucleus** 



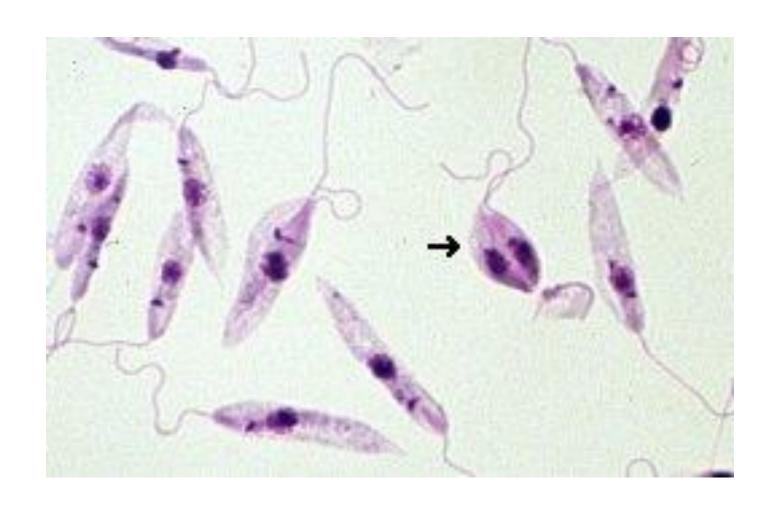
# Morphology......

### **Promastigote**

- Extracellular form
- Found in invertebrate vector and in culture
- Long and slender body with anterior flagellum
- Surface proteins bind C3 and attach to C3b
  receptors on macrophage surface







Leishmania - promastigotes

### **Life Cycle**

Promastigotes multiply in midgut and hind gut of sandfly

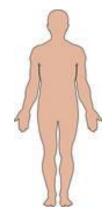


Amastigotes released from macrophages convert into promastigotes

SANDFLY VECTOR

Taken up by sandfly while feeding

Promastigotes regurgitated while feeding



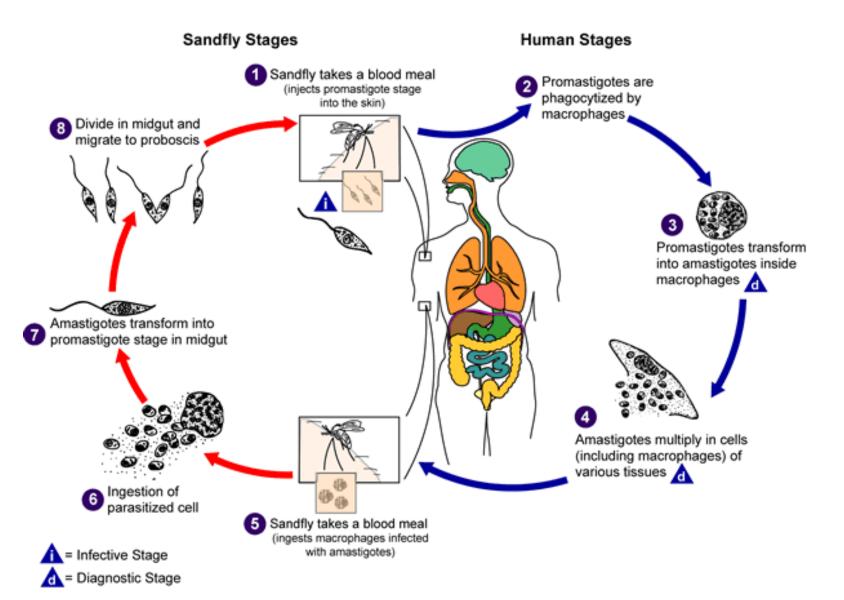
Promastigotes ingested by skin macrophages

MAMMALIAN HOST Multiply inside macrophage as amastigotes

Macrophage ruptures to release amastigotes

Amastigotes invade new macrophages

#### Leishmania life cycle



# Pathology

 Metacyclic promastigotes inoculated by sandfly bite, escape complement activation

Phagocytosed by macrophages and transformed into amastigotes

 Amastigotes resists digestion by lysosomal enzymes and multiplies within macrophages

## **Epidemiology of Visceral leishmaniasis**

- Found in Northern India, Mediterranean, African and South American regions
- VL may be zoonotic or non-zoonotic
- Usually zoonotic in Africa and the Mediterranean region
- Reservoir hosts
  - Africa- rats, ground squirrels etc.
  - Mediterranean- dogs
- Largely human-human transmission in India

#### Geographic distribution of leishmaniases



## Vector



#### Phlebotomine sandflies

- Small, hairy, brown flies like mosquitoes
- Feed at night
- Found in relatively dry habitats because larvae live in cracks & crevices of trees, rocks, houses, etc. (places with high humidity and moisture) - do not require pools of water

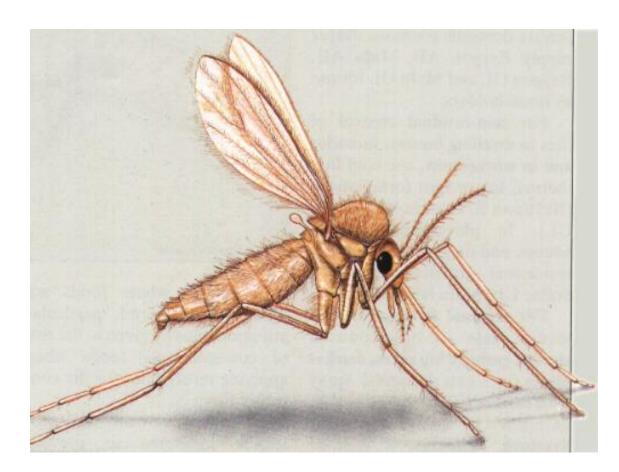
## Phlebotomine sandfly

Uniformly brown in colour

Generally hairy appearance

Wings held upright at rest

Long stilt-like legs





## Clinical Manifestations

- · Severe illness, often fatal if left untreated
- Clinical features
  - Long incubation period (4 6 months)
  - Insidious onset (maybe sudden in visitors)
  - Fever intermittent or remittent, with sweats
  - Massive splenomegaly
  - Moderate hepatomegaly
  - lymphadenopathy
  - Skin pigmentation (blackish discolouration 'kala azar')
  - Cachexia

## Clinical manifestations......

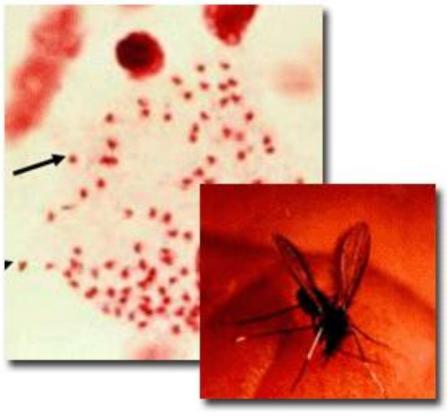
Anaemia, leucopaenia and thrombocytopaenia

Dysproteinaemia – reversal of albumin: globulin

Majority of deaths due to intercurrent infections

### Massive splenomegaly with mildmoderate hepatomegaly with mild-





# Lab diagnosis & treatment of VL

 Parasitological diagnosis: demonstrate amastigotes in bone marrow or spleen (by biopsy) as for CL( i.e. by smear or culture)

 Serological diagnosis: demonstrate specific antibodies or antigens (eg.rK 39 antibodies)

Molecular diagnosis: PCR assays

### **Treatment**

#### First line

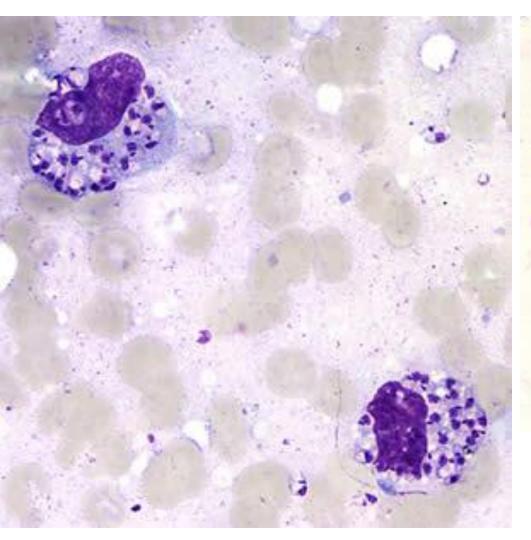
 IV pentavalent antimonials (sodium stibogluconate or meglumine antimonate) for several weeks

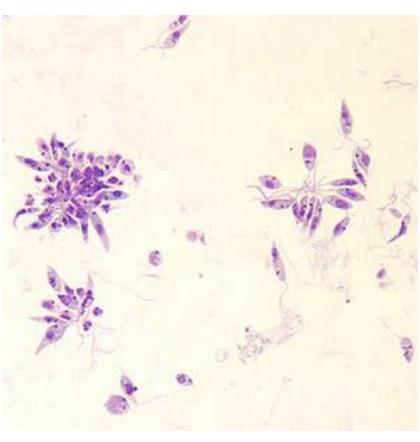
#### Second line

- Amphotericin B
- Oral miltefosine

#### In smears and H & E stained sections

#### In culture





#### Prevention and control of leishmaniasis

Primary preventive measures

- Avoiding outdoor activities during peak sandfly biting hours
- Proper clothing
- Using insect repellents
- Using bed nets

Vector control: usually only if malaria vector control is also being carried out

# Summary

- Leishmaniases are transmitted by sandfly vectors
- Life cycle; promastigote (I) & amastigote (D)
- VL- L. donovani & L. infantum
- Presents with fever, massive spenomegaly, moderate hepatomegaly
- Diagnosis-amastigotes in splenic & Bone marrow aspirate, culture (in-vitro & in-vivo), PCR
- Treatment; Pentavalent antimonials, miltefosine

# Thank You!