Leishmania

Leishmaniasis Is caused by a parasite called Leishmania

Other names and types of lesions caused by leishmania species

- 1- **Baghdad boils** A common name for an infection with *Leishmania* tropica; it is a cutaneous form of leishmaniasis presenting with pus-containing ulcers
- 2- Bay sore A common name for a cutaneous form of infection caused by Leishmania mexicana
- **3-** Chiclero ulcer A form of cutaneous leishmaniasis cased by *L. mexicana*; it is commonly found in Belize, Guatemala, and the Yucatan peninsula in areas where chicle sap is harvested for making chewing gum
- 4- Dum dum fever A common name for the visceral leishmaniasis caused by *Leishmania* donovani
- 5- Espundia Another name for an infection resulting from *Leishmania braziliensis*, the principal cause of mucocutaneous disease in Central and South America, particularly in Brazil
- 6- Forest yaws Another name for an infection with *Leishmania guyanensis*, the principle cause of mucocutaneous leishmaniasis in the Guianas, parts of Brazil and Venezuela; also known as **pian bois**
- 7- Kala-azar Another name for the most severe form of visceral leishmaniasis caused by members of the *Leishmania donovani* complex
- 8- Oriental sore A common reference for the cutaneous leishmaniasis caused by the *Leishmania tropica* complex
- 9- Pian bois Another name for infection with *L. guyanensis*; also known as forest yaws
- 10- Uta A reference to mucocutaneous leishmaniasis in the Peruvian Andes.

Types of Leishmaniasis

1) Leishmania braziliensis complex :

Other names: <u>Mucocutaneous leishmaniasis</u>, chiclero ulcer, espundia, forest yaws, pian bois, uta.

Found in Mexico, Argentina, Panama, Colombia, the Peruvian Andes, Guiana, Brazil, Bolivia, Paraguay, Ecuador, and Venezuela.

Members of Leishmania braziliensis complex are Leishmania braziliensis, Leishmania panamensis, Leishmania peruviana, and Leishmania guyanensis.

This leishmanial complex referred to as **New World leishmaniasis** because of their geographic location in what is commonly considered **the New World.**

Laboratory Diagnosis

- **Microscopic examination** of **Giemsa-stained** preparations **biopsy of the infected ulcer** for identifying the **amastigotes** of *L. braziliensis* complex. **Promastigotes** may be present when the sample is collected immediately after introduction into the patient.
- Culturing the infected material, which often demonstrates the promastigote stage, and
- Serologic testing such as ELISA, and IFA.
- Molecular diagnosis: Restriction analysis of kinetoplast DNA, a technique referred to as schizodeme analysis.
- Culture on NNN medium demonstrates the promastigote stage of these organisms

Epidemiology: The *L. braziliensis* complex is composed of *L. braziliensis* (found from Mexico to Argentina), *L. panamensis* (found in Panama and Colombia), *L. peruviana* (found in the Peruvian Andes), and *L. guyanensis* (found in Guiana, parts of Brazil, and Venezuela), particularly in the rain forest regions, where **chicle sap** for chewing gum is harvested (known under these circumstances as **chiclero ulcer**).

Transmission is generally through the bite of the *Lutzomyia* or *Psychodopygus* sandfly and reservoir hosts, including forest rodents and domestic dogs

Life Cycle Sandflies of the genera *Lutzomyia* and *Psychodopygus* are responsible for transmitting the pro-mastigotes of the species of the *L. braziliensis* complex humans via a blood meal. After skin bite, the promastigotes invade the reticuloendothelial cells and transform into amastigotes, which actively reproduce, causing tissue destruction. Reproduction and invasion of additional cells then occur. Affecting skin and mucous membrane. The amastigote serves as the infective stage for the sandfly. On ingestion, during a blood meal of an infected human, the amastigotes transform back into promastigotes in the fly midgut. These promastigotes multiply and the resulting developed forms eventually migrate into the salivary gland of the fly, where they are ready to be transferred to a new human during a blood meal. Thus, the cycle repeats itself. .

Clinical Symptoms

Mucocutaneous Leishmaniasis. Symptoms occur within a few weeks to months after transmission. Large ulcers in the oral or nasal mucosa areas (mucocutaneous) develop in a number of these patients after the initial invasion of the reticuloendothelial cells. A cutaneous (meaning affecting or relating to the skin) lesion may heal on its own. However, untreated cases of mucosal lesions result in destruction of the nasal septum, lips, and nose. Edema and secondary bacterial infections may cause disfigurement of the patient's face. Death is usually attributed to a secondary bacterial infection.

Treatment :

- Antileishmanial agent with antimony compounds (pentavalent antimonials). However, *L. braziliensis* has shown an increased resistance to pentavalent antimonials such as sodium stibogluconate (Pentosam).
- Alternative treatment include **liposomal amphotericin B** (Ambisome) and oral antifungal drugs such as fluconazole (Diflucan), ketoconazole (Nizoral).

Prevention and Control

- Public awareness through education programs in endemic areas and exercising personal protection against contact with sandflies (e.g., protective clothing, repellents, screening) are preventive measures against infections with *L. braziliensis* complex members.
- Prompt treatment and eradication of infected ulcers.
- Control of the sandfly population and reservoir hosts,
- Vaccine against members of the *L. braziliensis* complex and other *Leishmania* spp. is ongoing, with some vaccines for animals (dogs) presently in experimental trials.

2) Leishmania donovani complex

- Other names: <u>Visceral leishmaniasis</u>, kala-azar, dum dum fever.
- Found in Asia including the middle east and Iraq, North Africa including Egypt, and Central and South America. This group is comprised of *L. donovani, Leishmania infantum,* and *Leishmania chagasi.*
- Referred to as **Old or New World,** depending on the geographic location of the species of *Leishmania* involved.

Laboratory Diagnosis

- The **Montenegro skin test** is a screening test used for screening large populations at risk for infections caused by *Leishmania* spp. Its reliability in detecting exposure to the organisms causing leishmaniasis is related to the patient's disease status.
- **Giemsa-stained slides of blood, bone marrow, lymph node aspirates, and biopsies** of the infected areas to check for **amastigote** forms.
- Culturing of Blood, bone marrow, and other tissues; these samples often show the promastigote forms.
- **Serologic testing** is available using IFA (indirect fluorescent antibody), ELISA (enzymelinked immunosor- bent assay), and DAT (direct agglutination test).
- **Molecular tools** sauch as **schizodeme analysis** (restriction enzymes analysis of kinetoplast DNA (kDNA).

Life Cycle: The life cycle is identical to that of *L. braziliensis*, with only two exceptions. **First**, the specific sandfly species vary with each of the three subspecies. **Second**, *L. donovani* primarily affects the visceral tissue of the infected human.

Epidemiology: The *L. donovani* complex is composed of *L. donovani*, *L. infantum*, and *L. chagasi* (Table below). *L. donovani* and *L. infantum* are known to be endemic in areas of the Middle East, including Iraq, Saudi Arabia, the United Arab Emirates, and Bahrain.

TABLE 5-7 Leishmania donovani Complex: Epidemiology				
Subspecies	Geographic Distribution	Vector	Reservoir Hosts	
L. donovani chagasi	Central America, especially Mexico, West Indies, South America	Lutzomyia sandfly	Dogs, cats, foxes	
L. donovani donovani	Parts of Africa, India, Thailand, Peoples Republic of China, Burma, East Pakistan	Phlebotomus sandfly	India, none; China, dogs	
L. donovani infantum	Mediterranean Europe, Near East, Africa; also in Hungary; Romania, southern region of former Soviet Union, northern China, southern Siberia	Phlebotomus sandfly	Dogs, foxes, jackals, porcupines	

Clinical Symptoms

- Abdominal illness and **hepatosplenomegaly** (enlargement of the spleen and liver, **fever** and **chills**. The incubation period **2 weeks to 18 month**s.
- Diarrhea, as well as anemia, may often be present.
- Additional symptoms, including weight loss and emaciation, occur following parasitic invasion of the liver and spleen.
- Advanced stages of disease result in kidney damage (e.g., glomerulonephritis, inflammation of the glomeruli of the kidney).
- A characteristic darkening of the skin may be noted. This symptom is referred to by the common disease name, kala- azar, which means black fever. Chronic cases usually lead to death in 1 or 2 years.

Treatment:

- Liposomal amphotericin B (Ambisome) is the drug of choice for treating visceral leishmaniasis.
- **Sodium stibogluconate** (**Pentosam**) is also an effective treatment for infections with *L. donovani* complex, but resistance has been demonstrated by organisms in India and the Mediterranean.
- Successful treatment has been accomplished with the use of **gamma interferon** combined with **pentavalent antimony**.

3) Leishmania mexicana complex

Other names: New World cutaneous leishmaniasis, chiclero ulcer, bay sore.

• Rreferred to as **New World** because of the geographic location in **Central** and **South America**.

Life Cycle Notes; Is similar to that of *L. braziliensis*. The vectors are sandfly *Lutzomyia*. And rodents reservoir host.

Epidemiology: The *L. mexicana* complex is composed of *L. mexicana, L. pifanoi*, *L. amazonensis L. venezuelensis*, and *L. garnhami*. See the Table. Members of this complex are often transmitted by the bite of a *Lutzomyia* sandfly, with forest rodents serving as the reservoir host.

TABLE 5-8 Leishmania mexicana Complex: Epidemiology				
Subspecies	Geographic Distribution	Vector	Reservoir Hosts	
L. mexicana	Belize, Guatemala, Yucatan Peninsula	Lutzomyia sandfly for all species comprising this complex	Forest rodents for all species comprising this complex	
L. pifanoi	Amazon River Basin, Brazil, Venezuela			
L. amazonensis	Amazon River Basin, Brazil			
L. garnhami	Venezuelan Andes			
L. venezuelensis	Venezuela			

Clinical Symptoms

- New World Cutaneous Leishmaniasis. Approximately 40% of infections shows puscontaining ulcer affect the ear which is generally self- healing and can cause serious damage to the surrounding cartilage. Infected patients initially develop a small red papule, located at the bite site, which is typically 2 cm or larger in diameter and may cause pruritis (intense itching).
- Diffuse cutaneous leishmaniasis (DCL) In diffuse cutaneous infections with *L. pifanoi*, the initial lesion appears, ulcerates or disappears and, after a period of months to years, appears in local and distant areas from the bite site with lepromatous- appearing lesions.
 L. amazonensis infections have been known to progress to an incurable diffuse cutaneous form of the disease.

Treatment:

- Pentavalent antimonials, such as sodium stibogluconate (Pentosam)
- Antimony combined with pentoxifylline taken orally three times a day for 30 days
- Amphotericin B and liposomal amphotericin B (Ambisome) also proven to be effective.

Prevention and Control: Similar to other forms

4) Leishmania tropica complex

Other names: <u>Old World cutaneous leishmaniasis</u>, oriental sores, Delhi boils, Baghdad boils, dry or urban cutaneous leishmaniasis.

Found in the Mediterranean, Middle East, Armenia, Caspian region, Afghanistan, India, Kenya, Ethiopian highlands, southern Yemen, Turkmenistan deserts, Uzbekistan, Kazakhstan, northern Africa, the Sahara, Iran, Syria, Israel, and Jordan.

- It is comprised of *L. tropica, Leishmania aethiopica,* and *Leishmania major*. This leishmania complex and the diseases for which its organisms are the causative agent may
- Referred to as **Old World leishmaniasis.**

Laboratory Diagnosis

- **Microscopic examination** of Giemsa-stained slides of aspiration of fluid underneath the ulcer bed for the typical amastigotes.
- **Culture** of the ulcer tissue may also reveal the promastigote forms.
- Serologic tests, such as IFA testing, are available.
- Schizodeme analysis.

Life Cycle: Identical to that of *L. braziliensis*. All three of the *L. tropica* subspecies are transmitted by the *Phlebotomus* sandfly. *L. tropica* complex primarily attacks the human lymphoid tissue of the skin.

Clinical Symptoms

- Old World Cutaneous Leishmaniasis. Infected patients initially develop a small red papule 2 cm or larger in diameter, located at the bite site with intense itching,.
- **Diffuse cutaneous leishmaniasis (DCL)** occurs especially on the limbs and face when an immune response fails to take place. **Thick plaques** of skin, along with multiple lesions or nodules, usually result.

Epidemiology: Details in the table below

TABLE 5-9 Leishmania tropica Complex: Epidemiology				
Subspecies	Geographic Distribution	Vector	Reservoir Hosts	
L. aethiopica	Highlands of Ethiopia, Kenya, perhaps Southern Yemen	Phlebotomus sandfly for all species comprising this complex	Rock hyrax	
L. major	Former Soviet Union, Iran, Israel, Jordan, parts of Africa, Syria (esp. in rural areas)		Gerbils, other rodents	
L. tropica	Mediterranean, parts of the former Soviet Union, Afghanistan, India, Kenya, Middle East (especially in urban areas)		Possibly dogs	

Treatment

- As with the other leishmaniases, an effective treatment of *L. tropica* complex is sodium stibogluconate (Pentosam).
- The use of **steroids**, **application** of **heat** to the infected lesions,
- **Meglumine antimonate** (Glucantime), pentamidine, and oral ketoconazole may be indicated for treating *L. tropica* complex infections.
- **Paromomycin ointment** may also be given to aid in healing.

Prevention and Control: Similar to other forms