Unit VII – Problem 9 – Pharmacology: Leishmaniasis



- Drug therapy for leishmaniasis:

- Drug of choice: pentavalent antimony compounds
 - Sodium stibogluconate: this is found in English-speaking countries.
 - ✓ <u>Meglumine antimonite</u>: this is found in French-Spanish-speaking countries.
 - Notes for these 2 drugs:
 - They are poorly absorbed from the GIT. Therefore they cause GIT irritation/disturbance.
 - They are prodrugs: which means that they need to undergo biotransformation (metabolism) in the liver to be converted to their active forms.
 - They inhibit the process of glycolysis in parasites (which is necessary to keep them alive!).
 - They can be given intralesionally instead of being given orally.
 - ✓ <u>Liposomal amphotericin-B:</u>
 - It is one of drugs of choice but it is very expensive (50mg vial costs £96!). In contrast, conventional amphotericin-B which is one of the alternative drugs is cheap (50mg vial costs £4).
 - This drug is highly uptaken by macrophages. Therefore, they reach the phagosomes and attack amastigotes which are present and multiplying there.
 - ✤ What are liposomes?
 - They are very effective drug-delivery system transporting the drug to the desired target.
 - They are minute spherical oil droplets.
 - The membrane is composed of a phospholipid bilayer with 2 regions (hydrophilic and hydrophobic).

• Alternative drugs (preserved for antimony failure):

- ✓ <u>Pentamidine isethionate.</u>
- ✓ Conventional amphotericin-B.
- \checkmark <u>Miltefosine</u>.
- Systemic therapy with pentavalent antimony compounds is reserved for:
 - ✓ Multiple, inflamed, ulcerated and large sores.
 - ✓ Sores where scarring would be disfiguring.
 - ✓ Sores that will not heal easily (ex. Lower leg or over a joint).
 - ✓ Sore involving mucosa or cartilage.
 - ✓ Sores that might be due to parasites of the L.brazilinesis group (to avoid the risk of development of mucocutaneous "espundia").
 - ✓ Sores that are due to L.tropica (to reduce transmission since man and domestic dogs seem to be the main host).
- Life cycle of leishmania:
 - A sand fly carries promastigotes in its midgut → it bites the skin of a human → leading to the release of promastigotes which will interact with reticuloendothelial cells (macrophages) → inside macrophages, promastigotes will be converted to amastigotes which will start to multiply intracellularly → amastigotes will be released and they will infect new cells.
- Drug therapy for scabies:
 - Drug of choice:
 - Permethrin 5%:
 - ★ Treatment of choice for pregnant women and infants as young as 2 months old. This topical cream is applied for 8-14 hours followed by thorough bathing. Note that for neonatal scabies in infants less than 1 month → permethrin is applied for 6 hours only.
 - There are other 5 alternative drugs:
 - 1. <u>Ivermectin</u>: the only drug of scabies which is given orally in addition of being given topically.
 - 2. Lindane 1%
 - 3. Crotamiton 10%
 - 4. <u>Benzyl benzoate</u>: it is recommended for:

These 3 drugs are available in health centers



- *Infants*: by mixing it with 3 parts of water (diluting it in a proportion of 1:3)
- *Children*: by mixing it with equal quantity of water (diluting it in a proportion of 1:1).
- Pregnancy.

5. <u>Malathion 0.5%</u>

- **Disinfectants and antiseptics:**
 - Disinfectant: a chemical agent which destroys or inhibits the growth of pathogenic micro-organisms in the non-sporing or vegetative state (الحالة الخامدة). It is used to treat

inanimate objects (الجامادات) and materials.

- Antiseptic: a disinfectant which is used on the skin and other living tissues thereby limiting or preventing infections.
- Types of disinfectants and antiseptics:
 - \checkmark <u>Alcohols</u>: ethanol 70% and isopropanol 90% (skin disinfectants).
 - ✓ <u>Aldehydes</u>:
 - ★ Alkaline glutaraldehyde (2% alkaline solution in 70% isopropanol) \rightarrow instrument disinfectant
 - Methenamine \rightarrow urinary tract antiseptic.
 - \checkmark <u>Acids</u>:
 - ♦ Benzoic acid $0.1\% \rightarrow$ food preservative.
 - ◆ Benzoic acid 6% (fungistatic) + salicylic acid 3% (kerolytic) from white fields ointment for → dermatophytosis نمو الفطريات على الجلد (ex. Athlete's foot).
 - Undecylenic acid (fungistatic for tinea pedis which is causing athlete's foot).
 - Mandelic acid \rightarrow urinary tract antiseptic.
 - ✓ <u>Iodine:</u>
 - ✤ Iodine solution (2% iodine + 2.4% sodium iodide in water).
 - ❖ Iodine tincture (2% iodine + 2.4% sodium iodide in 50% alcohol) → skin disinfectant.
 - Povidine iodide \rightarrow pre-operative skin disinfectant.
 - ✓ <u>Chlorine:</u>
 - ♦ Chlorine 0.25-0.5 ppm \rightarrow water purification.
 - ↔ Halazone (chloramine) \rightarrow water sterilization (4-8 mf/L water).
 - Diluted sodium hypochlorite (modified Dakin's solution for cleansing and disinfecting wounds).
 - ✓ <u>Heavy metals:</u>
 - ✤ Mercury:
 - 4 0.1% thimerosal solution or tincture \rightarrow bacteriostatic antiseptic.
 - 4 2% mebromin \rightarrow bacteriostatic antiseptic.
 - Silver:
 - 4 1% silver nitrate ophthalmic solution \rightarrow for gonococcal ophthalmia.
 - 4 1% silver sulfadiazine cream \rightarrow suppress normal flora in burns.
 - ✓ <u>Phenol and related compounds:</u>
 - ✤ 3% hexachlorophene liquid soap → bacteriostatic, pre-operative scrubs antiseptic.
 - ♦ 4% chlorhexidine gluconate solution (cleansing of wounds) \rightarrow added to soap as skin antiseptic, pre-surgery antiseptic.
 - ♦ 0.2% chlorhexidine gluconate solution \rightarrow oral rinse against teeth plaque and gingivitis.
 - ✓ <u>Cation surface active agents:</u>
 - Benzalconium chloride.