

بِسْمِ اللَّهِ الرَّحْمَنِ الرَّحِيمِ



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قَالُوا سُبْحَانَكَ لَا عِلْمَ لَنَا إِلَّا مَا عَلَّمْتَنَا إِنَّكَ أَنْتَ الْعَلِيمُ
الْحَكِيمُ

(البقرة: الآية 32)





ACQUIRED IMMUNODEFICIENCY SYNDROME (AIDS)

By

*Staff Members of Tropical Medicine and
Gastroenterology*

Presented by

Dr. Hammam AbuEl Alameen

Lecturer of Tropical Medicine and Gastroenterology

Sohag Faculty of Medicine

ACQUIRED IMMUNODEFICIENCY SYNDROME (AIDS)

Agenda:

- Definition
- History
- Types of the virus
- Risk groups
- Modes of transmission
- Pathogenesis
- Clinical picture and stages of HIV infection
- Diagnosis
- Treatment and prevention

ACQUIRED IMMUNODEFICIENCY SYNDROME (AIDS)

- **Definition :**

AIDS actually refers to end stage manifestations of a long standing infection with a retrovirus HIV.

- **Cause:**

It is Caused by infection with Human Immunodeficiency Virus (H.I.V) which is a Lentivirus, a subgroup of retroviruses. These have the ability to transcribe their RNA genome to DNA one.

How did the HIV virus originate?

- Origin of virus is not yet certain.
- Evidence suggest that it existed in **chimpanzees** in West Africa and then infected humans possibly through a bite or scratch from a chimpanzee and from there it may have spread into a wider population.

History:

- **In 1981**, cases of a rare lung infection called **Pneumocystis carinii pneumonia (PCP)** were found in five young, previously healthy gay men in Los Angeles.
- At the same time, there were reports of a group of men in New York and California with an unusually aggressive cancer named ***Kaposi's Sarcoma***.
- **In December 1981**, the first cases of PCP were reported in people who inject drugs.
- **By the end of the year**, there were 270 reported cases of severe immune deficiency among gay men - 121 of them had died.

- **In June 1982**, a group of cases among gay men in Southern California suggested that the cause of the immune deficiency was sexual and the syndrome was initially called gay-related immune deficiency (or GRID).
- **Later that month**, the disease was reported in haemophiliacs and Haitians leading many to believe it had originated in Haiti.
- **In September**, the CDC used the term 'AIDS' (acquired immune deficiency syndrome) for the first time, describing it as

Types of the virus

- **There are 2 types of HIV**
 - **HIV-1** : predominant in America and Europe and is more aggressive.
 - **HIV-2**: predominant in central Africa (carrier rate is about 30%).

Risk groups

- 1. Homosexuals and heterosexuals**
- 2. IV drug abusers,**
- 3. addicts**
- 4. Hemophilics**
- 5. Blood transfusion recipients**

Modes of transmission :

- 1. Sexual intercourse:** homosexual or heterosexual intercourse (anal, vaginal)
- 2. Contaminated blood and blood products: -**
Blood transfusion. Blood products e.g. factor VIII.
- 3. Contaminated needles and syringes.**

Modes of transmission :

4. Mother to Child (vertical transmission):

- In utero.
- During childbirth.
- Breast milk.

5. Organ and tissue donation:

- Semen.
- Kidney, skin, and corneas.
- Bone marrow.

- **Sweat, urine, saliva and tears** are non-infectious under ordinary circumstances because the virus is hardly detected in these fluids.
- There is no evidence that HIV is spread by social or household contact nor by blood-sucking insects such as mosquitoes and bed bugs.

Pathogenesis:

The manifestations of HIV disease are related to progressive destruction of the immune system (mainly T-helper cell or CD4) or direct viral effect (CNS).

The virus has a characteristic cytopathic tropism for lymphocytes (mainly T-helper cells).

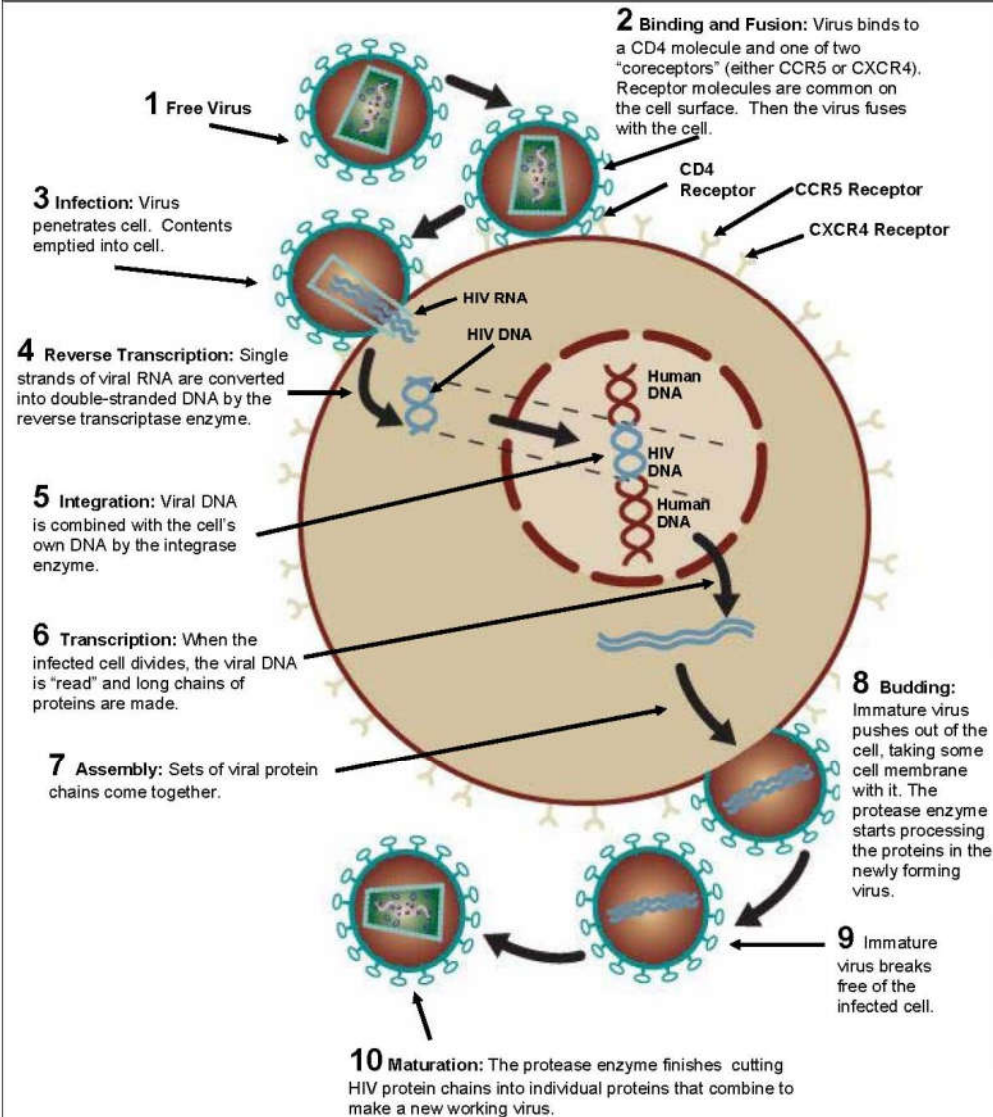
- Once the virus enters a cell, HIV can replicate & cause cell fusion or death by unknown mechanism .

Pathogenesis:

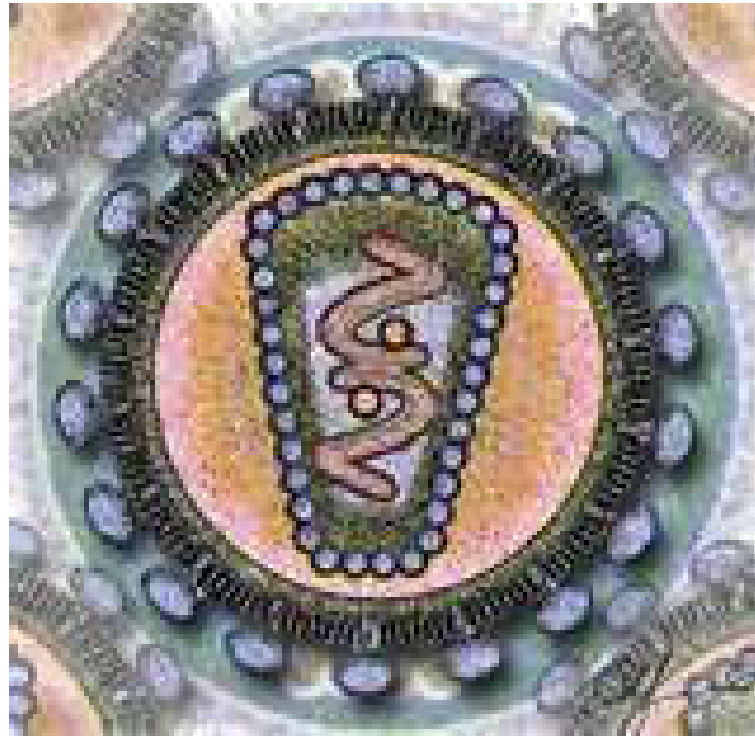
- In many cases, a latent state is established with integration of the HIV genome into the host cell genome .
- With increasing duration of infection, The number of CD4 Lymphocytes fall and with abnormal function.
- Other defects in the immune system includes infection of B-lymphocytes & macrophages .



HIV LIFE CYCLE



Revised April 30, 2005

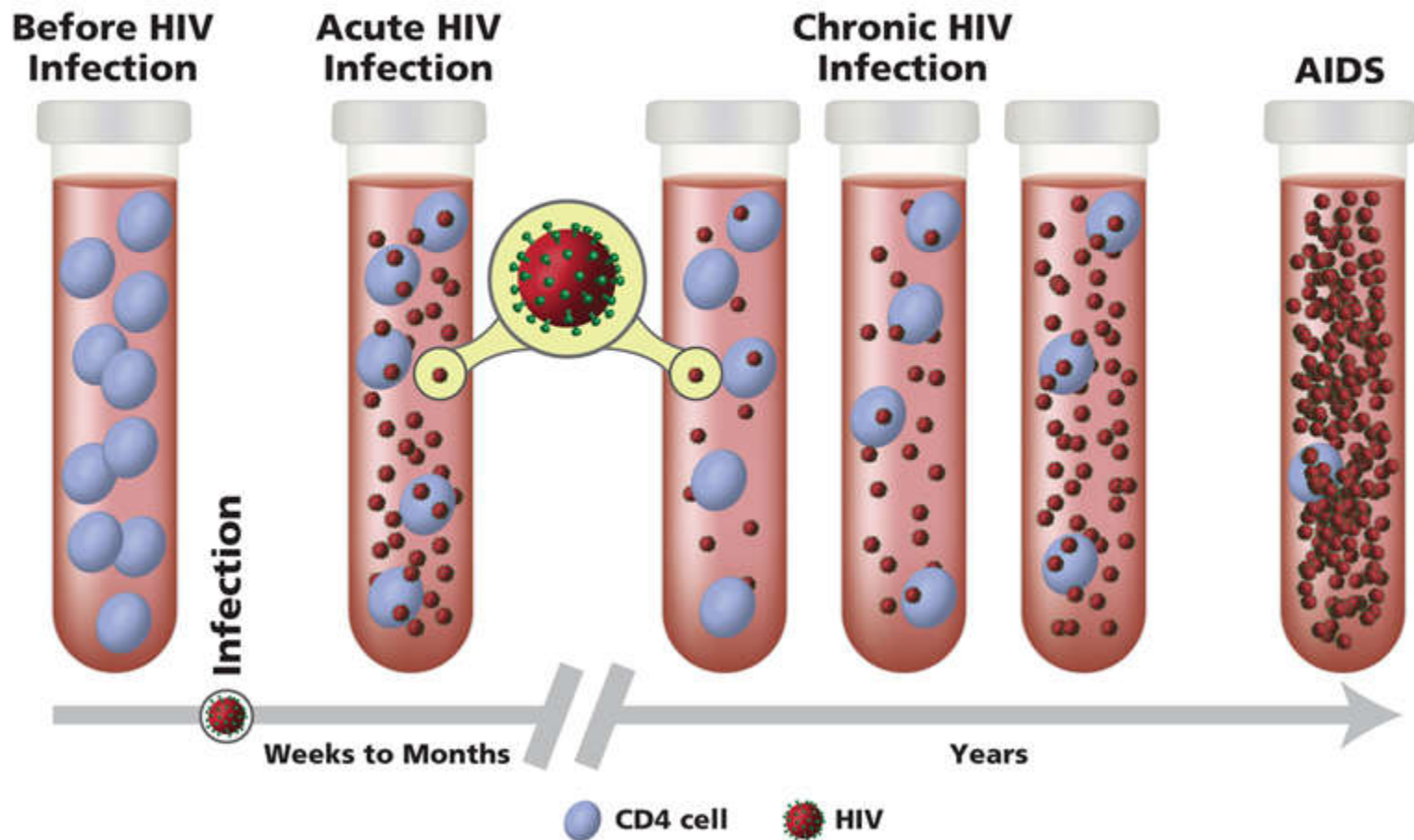


The mature virus

Stages of HIV infection

- There are three stages of HIV infection:
- (1) acute HIV infection, (2) chronic HIV infection, and (3) acquired immunodeficiency syndrome (AIDS).

HIV Progression



Clinical picture

Early stage infection

- 1- (Group I): Acute infection after 3-6 weeks incubation period. usually transient.
 - The earliest clinical presentation may be symptoms are of a glandular fever or flu-like illness with a sore throat, malaise, fever, maculo-papular rash.
 - Lymph node enlargement, diarrhoea, arthralgia
 - Occasionally encephalopathy, neuropathy or meningitis.

2- Group II: Asymptomatic infection.

Symptoms and signs usually resolve, and patients enter a prolonged phase in which they may remain asymptomatic but are potentially infectious.

3- Group III: Persistent generalized lymphadenopathy; after 3 to 10 years.

There is an insidious and variable progression, in which there is lymph node enlargement, tonsillar enlargement, episodes of unexplained fever.



Painless lymph node enlargement in HIV infection.

Advanced HIV diseases

4- **Group IV: Other diseases**, usually after 11 years.

- **Constitutional:**

Major constitutional symptom complexes

These include malaise, fever, night sweats, arthralgia, diarrhoea and weight loss

- **Neurological involvement:**

Myelopathy and peripheral neuropathy, for which the only explanation is HIV infection.

- **Opportunistic infections**

Opportunistic infections

- *Pneumocystis* pneumonia (PCP)
- Chronic cryptosporidiosis
- Toxoplasmosis
- Extra-intestinal strongyloidiasis
- Isosporiasis
- Cryptococcosis
- Candidiasis (oesophageal, bronchial, pulmonary)
- Histoplasmosis
- *Mycobacterium avium* intracellulare or *M. kansasii*
- Cytomegalovirus
- Herpes simplex

- Secondary cancers:
 - Kaposi's sarcoma.
 - Non-Hodgkin's lymphoma.
 - Primary cerebral lymphoma.

HIV-Related Malignancy



Kaposi's Sarcoma

Non-Hodgkin's lymphomas

Primary cerebral lymphoma

Manifestations and effect of HIV on different body systems

1- GIT manifestations:

- **Mouth:**
 - Oral candidiasis.
 - Hairy leukoplakia:
 - Recurrent painful oral ulcers (due to Herpes simplex virus).
 - Papilloma and condylomata (due to Human papilloma virus).
 - Kaposi's sarcoma.
 - Lymphoma.



Extensive oral infection with *Candida albicans* in a patient with HIV infection.

Hairy leukoplakia in HIV



Hairy leukoplakia in HIV



Hairy leukoplakia is specific for HIV. It is asymptomatic whitish lesions found on lateral margin of the tongue.

1- GIT manifestations:

- Oesophageal:

- Infectious esophagitis (Candidiasis, Herpes infection, CMV infection.) dysphagia and odynophagia
- Ulceration.
- Leukoplakia.
- Kaposi's sarcoma.
- Lymphoma.

1- GIT manifestations:

- Gastric:

- Reduced acid secretion.
- Reduced gastric juice volume.
- Reduced pepsin output.

The cause of gastric lesions may be **infections** as cytomegalovirus, campylobacter pylorides, cryptosporidium **or neoplasms** as Kaposi;s sarcoma.

1- GIT manifestations:

- Intestinal manifestations: (AIDS enteropathy):
 - Direct HIV infection to intestinal immune system leading to secretory diarrhea.
 - Partial villous atrophy.
 - Autonomic neuropathy.
 - Bacterial colonisation
 - Parasitic infestation (cryptosporidium, Giardia lamblia, Isospora belli, E. Histolytica).

2- Chest manifestations:

- Pneomcystis pneumonia (PCP):
- Pneumocystis pneumonia (PCP) is caused by Pneumocystis jirovecii.
- The taxonomy of the organism has been changed; Pneumocystis carinii now refers only to the Pneumocystis that infects rats, and P. jirovecii refers to the distinct species that infects humans.
- It is a ubiquitous fungus widely distributed in animals.

2- Chest manifestations:

- Pneomcystis pneumonia (PCP):

The presenting feature in 50% of HIV cases.

- Fever, malaise, fatigue, non productive cough, dyspnea.
- Auscultation of the chest often normal.
- X-ray shows bilateral diffuse shadows.
- The diagnosis is by transbronchial biopsy or lavage and examination using silver staining technique.
-

- **Polymerase chain reaction (PCR)** is an alternative method for diagnosing PCP.
- Trimethoprim-sulfamethoxazole (TMP-SMX) (also named **Co-trimoxazole**) is the recommended prophylactic agent for PCP and also for treatment.

2- Chest manifestations:

- Bacterial pneumonia:

Caused by streptococcus, staphyococcus aureus,

Mycobacterium tuberculosis,

Mycobacterium avium intracellulare.

- Others: as cytomegalovirus, candida, Kaposi's sarcoma, and lymphoma.

2- Chest manifestations:

- **Mycobacterium tuberculosis:**

Main features of TB in HIV:

- Most cases are due to reactivation.
- Extra pulmonary TB is common.
- TB may accelerate HIV disease.
- Tuberculin test is mostly negative (anergy).
- 6-9 months therapy is recommended but INH prophylaxis should continue for life to prevent relapse.

3- Neurological manifestations:

- Encephalopathy: caused by direct HIV infection of the brain).
- Myelopathy and acute transverse myelitis.
- Neuropathy: mostly peripheral sensory neuropathy.
- Encephalitis
- Meningitis: most commonly caused by cryptococcus neophormans.
- Toxoplasmosis.
- CNS lymphoma.

Kaposi's sarcoma

- Kaposi's sarcoma: Common tumor in AIDS patients. It develops as multiple, small dusky purple-red or purple nodular, painless lesions of the skin or buccal mucosa, but may also be found anywhere in the gastrointestinal tract or bronchial mucosa.





Kaposi's sarcoma is a common complication in patients with AIDS (25% of cases).

Main symptoms of **AIDS**

Neurological

- Encephalitis
- Meningitis

Eyes

- Retinitis

Lungs

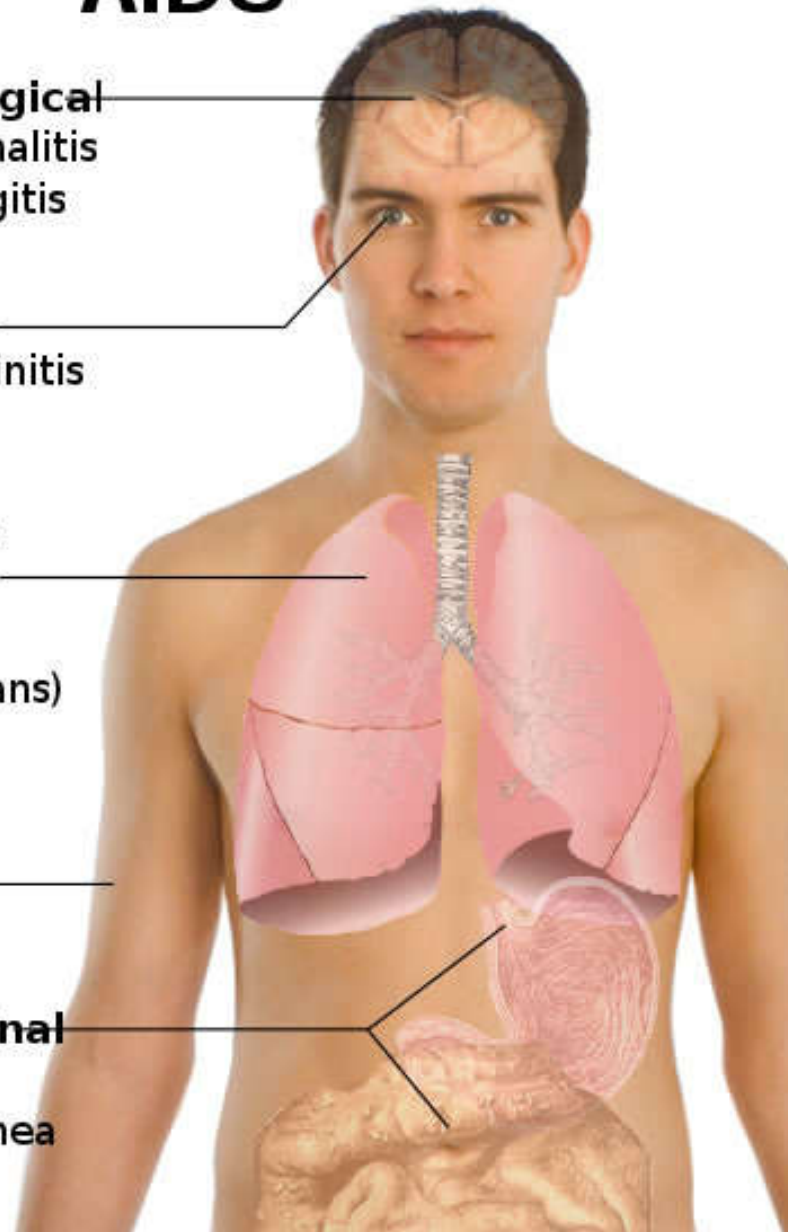
- Pneumocystis pneumonia
- Tuberculosis (multiple organs)
- Tumors

Skin

- Tumors

Gastrointestinal

- Esophagitis
- Chronic diarrhea
- Tumors



Diagnosis:

A) Clinical suspicion:

- The presence of at least 2 major symptoms and at least 1 minor symptom
- **Major symptoms include:**
 - 1- Unexplained fever for more than 1 month.
 - 2- Unexplained diarrhea for more than 1 month.
 - 3- Unexplained weight loss greater than 10% of previous weight.

- Minor symptoms include:

- 1- Presence of maculopapular rash.
- 2- Persistent oral candidiasis.
- 3- Herpes zoster.
- 4- Aggressive uncontrolled herpes simplex.
- 5- Unexplained cough for more than 1 month
- 6- Enlarged lymph nodes in more than one extra-inguinal site.

B) Laboratory

- **CBC which shows** anaemia, neutropenia, thrombocytopenia are common with advanced HIV infection, lymphopenia especially T cells, ↓CD4 /CD8 ratio. Decrease CD 4 count less than 400/mm³. Patients with count less than 200 /mm³ are at very high risk.
- **HIV ELISA : a screening test for HIV**
Sensitivity > 99.9%. To avoid false positive results repeatedly reactive results must be confirmed by **Western blot technique**.
- **Polymerase chain reaction (PCR):** is very specific for DNA replication and confirm the infection by HIV.

Treatment :

- Anti retro viral drugs:

- **reverse transcriptase inhibitors**
 - Nucleoside analogues as → Zidovudine (AZT), Lamivudine.
 - Nonnucleoside analogues as nevirapine
- **Protease inhibitors** as saquinavir, indinavir.
- **Ganciclovir:** is the drug of choice for CMV infection.

- **Acyclovir:** for herpes simplex virus infection topically, orally or intravenously.
- **Desciclovir** (acyclovir analogue) for therapy of hairy leukoplakia.
- **Anti protozoal.**
- **Anti fungal.**
- **Anti TB.**

Prevention:

Prevention of the spread of HIV infection involves:

- 1- Screening of blood donors,
- 2- Advice on safer sexual practices,
- 3- Measures to curtail intravenous drug abuse and the provision of sterile syringe and needle



Thank You