Introduction to Management of Latent TB infections (LTBI)

Latent TB is a state of persistent immune response to stimulation by *M. tuberculosis* antigens with no evidence of clinically manifested active TB, meaning, signs and symptoms of TB are not present.

**TB preventive treatment** is the treatment offered to individuals who are considered at risk of TB disease in order to reduce that high risk.

Provision of preventive treatment has proven to be an effective intervention to avert the development of active TB disease, with efficacy ranging from 60% to 90%, hence it is a principle strategy in achieving the global agenda of Ending TB by 2035.

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**Latent TB Infection**

Latent TB Infection means TB germs are in the body, but not enough to cause sickness or spread germs to others.

**TB Disease**

If TB germs become active and multiply, latent TB infection can turn into TB disease.

**1 in 10**

Without treatment, 1 in 10 people with latent TB infection will develop TB disease.

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To End TB, we need to stop Latent TB!
Pathophysiology of Latent TB Infections

When a person inhales air droplets with *M. tuberculosis* bacilli, most of the larger droplets become lodged in the upper respiratory tract (the nose and throat).

However, the smaller droplet nuclei may reach the small air sacs of the lung (the alveoli), where infection may begin.

In the alveoli, some of the tubercle bacilli are killed, but a few multiply in the alveoli and enter the bloodstream and spread throughout the body. Bacilli may reach any part of the body.

Within 2 to 8 weeks, however, the body’s immune system usually intervenes, halting multiplication and preventing further spread.

The immune system is the system of cells and tissues in the body that protect the body from foreign substances.

At this point, the person does not manifest any signs or symptoms of TB hence said to have latent TB infection (LTBI).

Most people with LTBI are unaware of the danger that may have already quietly settled in them.
Populations to be offered Tuberculosis Preventive Therapy (TPT)

The most at-risk populations for whom systematic LTBI testing and treatment is recommended for the country include:

1. People Living with HIV
   1. Aged more than 12 months regardless of TB exposure history
   2. Aged less than 12 months exposed to bacteriologically confirmed TB.


3. Health care workers and support staff working in health care settings.

4. Prisons (Inmates and prison staff).

5. Others population at risk.
   1. Patients initiating chemotherapy or those who are taking certain immunosuppressive drugs.
   2. Patients receiving dialysis
   3. Patients preparing for an organ or haematological transplant
   4. Patients who have silicosis.
How to rule out active TB

Before initiating TB preventive therapy, the health care workers should rule out active TB using the symptoms screening questions:

- Cough of any duration
- Weight loss
- Night sweats
- Fever.

For children, screen for:

- Cough of any duration
- Reduced playfulness/ lethargy
- Failure to thrive/ poor weight gain
- Fever/ night sweats
- History of contact with a person with TB

Diagnosis of Latent TB infections

The goal of testing for LTBI is to identify persons who are at increased risk for developing TB and who would benefit from treatment of the infection.

Either a tuberculin skin test (TST) or Interferon-Gamma Release Assay (IGRA) can be used to test LTBI.

NOTE

All the at-risk populations found to have none of these symptoms should be tested using IGRA/ TST before initiating TB Preventive Therapy except people living with HIV and children under the age of five years who are household contacts of bacteriologically-confirmed TB.

Any client found to have any of the symptoms suggestive of TB should not be offered TPT instead they should be evaluated further for active TB disease.
Treatment options for Latent TB Infections

**3RH: Given once daily for 3 months**
- Availability of child-friendly formulation (75/50mg)
- Offered as preventive treatment for children and adolescents aged <15 years
- Better adherence (shorter regimen)
- Do not give to patients on PI or NVP based ART
- Less costly
- Available in fixed dose combination

**3HP: Given once a week for 12 weeks**
- Very promising regimen
- 1-month daily treatment option (1HP) being evaluated in PLHIV
- Better adherence (shorter regimen)
- FDC now available
- Child-friendly formulation not available yet
- No evidence for use in children <2 years
- Do not give to patients on PI or NVP based ART
- Safe to use in PLHIV on EFV and RAL
- Studies on use with DTG- so far no interactions

**6 H: Given daily for 6 months**
- Longer treatment duration
- Higher rates of hepatotoxicity than other regimens
- Still regimen of choice for PLHIV on PI based ART regimen and on HIV exposed children on Nevirapine (NVP) prophylaxis.
- Lowest cost (uncoated tab), high cost dispersible tab
- A syrup formulation is available for children

**NOTE**

The treatment of Latent TB infection is given based on the patient’s weight

Full patient dose should be available for the entire treatment period before initiating treatment in patient.
Follow Up of Patients on TB Preventive Therapy (TPT)

Patients on TPT should be followed up on monthly basis and clinic appointments harmonized with any other routine clinic schedule. During each clinic visit, the health care worker will conduct the following:

- Conduct symptom based TB screening at every clinic visit for patients on TPT and update TB status
- Assess and reinforce adherence of the patients at every visit to ascertain compliance and completion of doses
- If a patient screens positive for TB while on TPT, stop TPT and manage according to National TB guidelines
- Assess for any adverse drug reactions at each visit and intervene appropriately.
Algorithm for children (below 15 years) who are contacts of Bacteriologically confirmed TB patients

NOTE
1. If TB is ruled out, antibiotics have been given and they are now asymptomatic, offer TPT according to their age and weight.
2. Contra-indications include active hepatitis (Look for jaundice, tender abdomen), Symptoms of peripheral neuropathy (Look for inability to walk... etc.)
3. Screen for TB symptoms at every visit to rule out active TB.
4. Repeat symptomatic screening and physical examination to be sure that they are not a TB case.
Algorithm for all other risky population - Adults

NOTE

1. If TB is ruled out, antibiotics have been given and they are now asymptomatic, offer TPT according to their age.

2. Contra-indications include active hepatitis (Look for jaundice, tender abdomen), Symptoms of peripheral neuropathy (Look for inability to walk etc).

3. Repeat symptomatic screening and physical examination to be sure that they are not a TB case.

4. Screen for TB symptoms at every visit to rule out active TB.