

## PACKAGE LEAFLET

### **DOLUTEGRAVIR 50 mg, LAMIVUDINE 300 mg and TENOFOVIR DISOPROXIL FUMARATE 300 mg TABLETS**

**Read all of this leaflet carefully before you start taking this medicine because it contains important information for you.**

- Keep this leaflet. You may need to read it again.
- If you have any further questions, ask your doctor or pharmacist.
- This medicine has been prescribed for you only. Do not pass it on to others. It may harm them, even if their signs of illness are the same as yours.
- If you get any side effects, talk to your doctor or pharmacist. This includes any possible side effects not listed in this leaflet. See section 4.

#### **In this leaflet**

1. What Dolutegravir, Lamivudine and Tenofovir Disoproxil Fumarate Tablets is and what it is used for
2. Before you take Dolutegravir, Lamivudine and Tenofovir Disoproxil Fumarate Tablets
3. How to take Dolutegravir, Lamivudine and Tenofovir Disoproxil Fumarate Tablets
4. Possible side effects
5. How to store Dolutegravir, Lamivudine and Tenofovir Disoproxil Fumarate Tablets
6. Further information

#### **Dolutegravir, Lamivudine and Tenofovir Disoproxil Fumarate Tablets**

Dolutegravir, Lamivudine and Tenofovir Disoproxil Fumarate Tablets contains the active ingredient dolutegravir, lamivudine & tenofovir disoproxil fumarate. Dolutegravir belongs to a group of anti-retroviral medicines called *integrase inhibitors (INIs)*. Lamivudine and Tenofovir is a nucleo(t)side reverse transcriptase inhibitor.

Dolutegravir, Lamivudine and Tenofovir Disoproxil Fumarate Tablets does not cure HIV infection; it reduces the amount of virus in your body, and keeps it at a low level. As a result of that, it also increases the CD4 cell count in your blood. CD4 cells are a type of white blood cells that are important in helping your body to fight infection.

Dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets are always used in combination with other anti-retroviral medicines (*combination therapy*). To control your HIV infection, and to stop your illness from getting worse, you must keep taking all your medicines, unless your doctor tells you to stop taking any.

The Marketing Authorisation Holder for **Dolutegravir, Lamivudine and Tenofovir Disoproxil Fumarate Tablets** is:

M/s Aurobindo Pharma Ltd.,

Plot No.: 2, Maitrivihar  
Ameerpet, Hyderabad-500 038  
India.

**Dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets** are manufactured by:  
AUROBINDO PHARMA LTD.,  
Unit VII, SEZ, TSIC, Plot. No.SI,  
Survey No's: 411/P, 425/P, 434/P, 435/P & 458/P,  
Green Industrial Park, Polepally Village,  
Jedcherla Mandal, Mahaboobnagar District,  
Telangana State, India.

### **1. What Dolutegravir, Lamivudine and Tenofovir Disoproxil Fumarate Tablets are and what it is used for**

Dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets are a prescription medicine that are used together with other antiretroviral medicines to treat Human Immunodeficiency Virus (HIV-1) infection.

Dolutegravir, Lamivudine and Tenofovir Disoproxil Fumarate Tablets contains the active ingredient dolutegravir, lamivudine & tenofovir disoproxil fumarate. Dolutegravir belongs to a group of anti-retroviral medicines called *integrase inhibitors (INIs)*. Lamivudine and Tenofovir is a nucleo(t)side reverse transcriptase inhibitor.

Dolutegravir, Lamivudine and Tenofovir Disoproxil Fumarate Tablets does not cure HIV infection; it reduces the amount of virus in your body, and keeps it at a low level. As a result of that, it also increases the CD4 cell count in your blood. CD4 cells are a type of white blood cells that are important in helping your body to fight infection.

### **2. Before you take Dolutegravir, Lamivudine and Tenofovir Disoproxil Fumarate Tablets**

#### **Do not take Dolutegravir, Lamivudine and Tenofovir Disoproxil Fumarate Tablets:**

- have ever had an allergic reaction to a medicine that contains Dolutegravir.
- take dofetilide. Taking Dolutegravir and Dofetilide can cause side effects that may be serious or life-threatening.
- are allergic to lamivudine or any of the ingredients in dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets. See the end of this patient information leaflet for a complete list of ingredients in dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets.

- if you are currently taking St. John's wort (*Hypericum perforatum*) (a herbal remedy used for depression and anxiety) or voriconazole (a drug used against fungal infections).

**Before you take dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets, tell your doctor if you:**

- have ever had an allergic reaction to dolutegravir.
- If you are taking another medicine called dofetilide (to treat heart conditions).
- have had liver problems, including hepatitis B or C infection.
- have kidney problems
- have bone problems
- have any other medical condition. Including HIV infection.
- are pregnant or plan to become pregnant. It is not known if dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets will harm your unborn baby. Tell your doctor if you are pregnant or plan to become pregnant while taking dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets.
- are breastfeeding or plan to breastfeed. **Do not breastfeed if you take dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets.**
- you should not breastfeed if you have HIV-1 because of the risk of passing HIV-1 to your baby. Talk with your doctor about the best way to feed your baby.

**Tell your doctor about the medicines you take**, including prescription and over-the-counter medicines, vitamins, and herbal supplements.

Some medicines interact with dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets. Keep a list of your medicines and show it to your doctor and pharmacist when you get a new medicine. You can ask your product for a list of medicines that interact with dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets.

Over-the-counter medicines that interact with dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets include:

- St. John's wort (*Hypericum perforatum*). Avoid use of St. John's wort with dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets.
- antacids, laxatives, or other medicines that contain aluminum, magnesium, sucralfate, or buffered medicines. Dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets should be taken at least 2 hours before or 6 hours after you take antacids, laxatives, or other medicines that contain aluminum, magnesium, sucralfate, or buffered medicines.
- iron or calcium supplements taken by mouth. Supplements containing calcium or iron

may be taken at the same time with dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets if taken with food. Otherwise, dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets should be taken at least 2 hours before or 6 hours after you take these medicines.

- other medicines that contain tenofovir
- adefovir

Especially tell your doctor if you take the following medications.

- didanosine
- atazanavir
- darunavir
- lopinavir with ritonavir
- ledipasvir with sofosbuvir

**Do not start taking a new medicine without telling your doctor.** Your doctor can tell you if it is safe to take dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets with other medicines.

### **Pregnancy**

If you are pregnant, if you become pregnant, or if you are planning to have a baby:

- **Talk to your doctor** about the risks and benefits of taking Dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets.

### **Breast-feeding**

**Women who are HIV-positive must not breast feed** because HIV infection can be passed on to the baby in breast milk.

If you are breast-feeding, or thinking about breast-feeding:

- **Talk to your doctor immediately.**

### **Driving and using machines**

Dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets can make you dizzy and have other side effects that make you less alert.

- Don't drive or operate machinery unless you are sure you're not affected.

### **3. How to take Dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets**

Always take this medicine exactly as your doctor has told you. Check with your doctor or pharmacist if you are not sure. See possible side effects, section 4 before you take Dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets.

- **Take dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets exactly as your doctor tells you to take them.**

- If you miss a dose of dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets, take it as soon as you remember. Do not take 2 doses at the same time or take more than what your doctor tells you to take.
- Stay under the care of a doctor during treatment with dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets.
- Dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets may be taken with or without food.
- Do not run out of dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets. The virus in your blood may increase and the virus may become harder to treat. When your supply starts to run low, get more from your doctor or pharmacy.
- If you take too much dolutegravir, lamivudine and tenofovir disoproxil fumarate, call your doctor or go to the nearest hospital emergency room right away.

#### **Use in children and adolescents**

Dolutegravir: Safety and efficacy of dolutegravir have not been established in pediatric patients weighing less than 30 kg or in any pediatric patients who are INSTI-experienced.

Lamivudine: The safety and effectiveness of lamivudine in combination with other antiretroviral agents have been established in pediatric patients aged 3 months and older.

Tenofovir disoproxil fumarate: Safety and effectiveness of tenofovir disoproxil fumarate in pediatric patients younger than 2 years of age with HIV-1 infection have not been established.

#### **If you take more Dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets than you should**

If you take too many tablets of Dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets, **contact your doctor or pharmacist for advice**. If possible, show them the Dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets pack.

#### **If you forget to take Dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets**

If you miss a dose, take it as soon as you remember.

→ **Don't take a double dose** to make up for a missed dose.

#### **Don't stop taking Dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets without advice from your doctor**

Take Dolutegravir, Lamivudine and Tenofovir Disoproxil Fumarate Tablets for as long as your doctor recommends. Don't stop unless your doctor advises you to.

If you have any further questions on the use of this medicine, ask your doctor or pharmacist.

#### **4. Possible side effects**

Like all medicines, this medicine can cause side effects, but not everybody gets them.

- **Dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets can cause**

**serious side effects including:**

- **Changes in liver tests.** People with a history of hepatitis B or C virus may have an increased risk of developing new or worsening changes in certain liver tests during treatment with dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets. Your doctor may do tests to check your liver function before and during treatment with dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets.
- **Risk of inflammation of the pancreas (pancreatitis).** Children may be at risk for developing pancreatitis during treatment with dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets if they:
  - have taken nucleoside analogue medicines in the past
  - have a history of pancreatitis
  - have other risk factors for pancreatitis

**Call your doctor right away if your child develops signs and symptoms of pancreatitis including severe upper stomach-area pain, with or without nausea and vomiting.** Your doctor may tell you to stop giving dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets to your child if their symptoms and blood test results show that your child may have pancreatitis.

- **Changes in body fat can happen in people who take HIV-1 medicines.** These changes may include increased amount of fat in the upper back and neck (“buffalo hump”), breast, and around the middle of your body (trunk). Loss of fat from the legs, arms, and face may also happen. The exact cause and long-term health effects of these problems are not known.
- **Changes in your immune system (Immune Reconstitution Syndrome)** can happen when you start taking HIV-1 medicines. Your immune system may get stronger and begin to fight infections that have been hidden in your body for a long time. Tell your doctor right away if you start having new symptoms after you start taking dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets.
- **New or worse kidney problems, including kidney failure,** can happen in some people who take dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets. Your doctor should do blood tests to check your kidneys before you start treatment with dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets. If you have had kidney problems in the past or need to take another medicine that can cause kidney problems, your doctor may need to do blood tests to check your kidneys during your treatment with dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets.

- **Bone problems** can happen in some people who take dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets. Bone problems include bone pain, softening or thinning (which may lead to fractures). Your doctor may need to do additional tests to check your bones.
- **Build-up of an acid in your blood (lactic acidosis).** Lactic acidosis can happen in some people who take dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets. Lactic acidosis is a serious medical emergency that can cause death.
  - Lactic acidosis can be hard to identify early, because the symptoms could seem like symptoms of other health problems. **Call your doctor right away if you get any of the following symptoms which could be signs of lactic acidosis:**
    - feel very weak or tired
    - have unusual (not normal) muscle pain
    - have trouble breathing
    - have stomach pain with
      - nausea (feel sick to your stomach)
      - vomiting
    - feel cold, especially in your arms and legs
    - feel dizzy or light-headed
    - have a fast or irregular heartbeat
- **Severe liver problems.** Severe liver problems can happen in people who take dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets or similar medicines. In some cases these liver problems can lead to death. Your liver may become large (hepatomegaly) and you may develop fat in your liver (steatosis).

**Talk to your doctor right away if you get any of the following signs or symptoms of liver problems:**

- your skin or the white part of your eyes turns yellow (jaundice)
- dark or “tea-colored” urine
- light-colored stools (bowel movements)
- loss of appetite for several days or longer
- nausea
- pain, aching, or tenderness on the right side of your stomach area

**You may be more likely to get lactic acidosis or severe liver problems if you are female, very overweight (obese), or have been taking dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets or a similar medicine for a long time.**

- **Allergic reactions.** Talk to your doctor right away if you develop a rash with dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets. **Stop taking**

**dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets and get medical help right away if you:**

- **develop a rash with any of the following signs or symptoms:**
  - fever
  - generally ill feeling
  - extreme tiredness
  - muscle or joint aches
  - blisters or sores in mouth
  - blisters or peeling of the skin
  - redness or swelling of the eyes
  - swelling of the mouth, face, lips, or tongue
  - problems breathing
- **Worsening of hepatitis B virus in people who have HIV-1 infection.** If you have HIV-1 (Human Immunodeficiency Virus type 1) and hepatitis B virus (HBV) infection, your HBV may get worse (flare-up) if you stop taking dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets. A “flare-up” is when your HBV infection suddenly returns in a worse way than before. Worsening liver disease can be serious and may lead to death.
  - Do not run out of dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets. Refill your prescription or talk to your doctor before your dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets are all gone.
  - Do not stop dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets without first talking to your healthcare provider.
  - If you stop taking dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets, your doctor will need to check your health often and do blood tests regularly for several months to check your liver. Tell your doctor about any new or unusual symptoms you may have after you stop taking dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets.
- **Resistant Hepatitis B Virus (HBV).** If you have HIV-1 and hepatitis B, the hepatitis B virus can change (mutate) during your treatment with dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets and become harder to treat (resistant).
- **Use with interferon and ribavirin-based regimens.** Worsening of liver disease that has caused death has happened in people infected with both HIV-1 and hepatitis C virus who are taking antiretroviral medicines and are also being treated for hepatitis C with interferon with or without ribavirin. If you are taking dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets and interferon with or without ribavirin, tell your doctor if you have any new symptoms.
- **Talk to your doctor about taking an HIV test before starting treatment with dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets for chronic hepatitis B. You should also get a test for HBV if you are taking dolutegravir,**



## **lamivudine and tenofovir disoproxil fumarate tablets for treatment of HIV.**

**The most common side effects of dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets include:**

- trouble sleeping
- tiredness
- headache
- nausea
- generally not feeling well
- nasal signs and symptoms
- itching
- dizziness
- diarrhea
- cough
- rash
- pain
- depression
- weakness
- vomiting
- fever

Tell your doctor about any side effect that bothers you or that does not go away.

These are not all the possible side effects of dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets. Call your doctor for medical advice about side effects.

### **5. How to store Dolutegravir, Lamivudine and Tenofovir Disoproxil Fumarate Tablets**

Keep this medicine out of the sight and reach of children.

- Do not store above 30°C. Store in the original package.
- Keep dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets in the original container.
- Do not use dolutegravir, lamivudine and tenofovir disoproxil fumarate tablets if the seal over the bottle opening is broken or missing.
- Keep the bottle tightly closed. Protect from moisture.
- The bottle contains a desiccant packet to help keep your medicine dry (protect it from moisture). Keep the desiccant packet in the bottle. Do not remove the desiccant packet.

Do not throw away any medicines via wastewater or household waste. Ask your pharmacist how to throw away medicines you no longer use. These measures will help protect the environment.

### **6. Further information**

#### **What Dolutegravir, Lamivudine and Tenofovir Disoproxil Fumarate Tablets contains**

- The active substance is Dolutegravir, lamivudine and tenofovir disoproxil fumarate. Each film coated tablet contains Dolutegravir sodium equivalent to 50 mg Dolutegravir, Lamivudine USP 300 mg, Tenofovir disoproxil fumarate 300 mg equivalent to 245 mg of tenofovir disoproxil.
- The other ingredients in the tablet core are Mannitol, Microcrystalline Cellulose, Povidone, Ferric Oxide Red, Sodium Starch Glycolate, Sodium Stearyl Fumarate,

Croscarmellose sodium, Hypromellose, Magnesium stearate, Colloidal Silicon Dioxide.

- The other ingredients in the tablet film coating are Polyvinyl alcohol, Titanium dioxide, Polyethylene Glycol, Talc, Iron oxide red and Iron oxide black.

**What Dolutegravir, Lamivudine and Tenofovir Disoproxil Fumarate Tablets looks like and contents of the pack**

Dolutegravir, lamivudine and tenofovir disoproxil fumarate Tablets are Pink colored, Oval, biconvex, film coated tablets debossed with 'N33' on one side and plain on the other side.

The film-coated tablets are provided in bottles containing 30 & 90 tablets.

**Reporting of side effects**

For any information about this medicinal product, please contact the local representative of the Marketing Authorisation Holder.

“If you notice any side effect (s) with the use of this drug, please report it immediately via internet to the following e-mail address: [pharmacovigilance@aurobindo.com](mailto:pharmacovigilance@aurobindo.com)”.