PROPOSED PATIENT INFORMATION LEAFLET FOR LOPIMUNE 40/10 ORAL PELLETS

SCHEDULING STATUS: S4

PROPRIETARY NAME, STRENGTH AND PHARMACEUTICAL FORM:

LOPIMUNE 40/10 ORAL PELLETS

(Lopinavir 40 mg and Ritonavir 10 mg)

ORAL PELLETS

Sugar free

Read all of this leaflet carefully before you start taking LOPIMUNE 40/10 ORAL PELLETS:

- **Keep this leaflet. You may need to read it again.**
- **If you have further questions, please ask your doctor or pharmacist.**
- **LOPIMUNE 40/10 ORAL PELLETS has been prescribed for you personally and you should not share your medicine with other people. It may harm them, even if their symptoms are the same as yours.**

WHAT LOPIMUNE 40/10 ORAL PELLETS CONTAIN:

Each capsule of oral pellets contains lopinavir 40 mg and ritonavir 10 mg as active ingredients.

The other ingredients are colloidal silicon dioxide, copovidone, gelatin capsule, hydroxyl propyl methylcellulose, PEG-6000, sodium stearyl fumarate, sorbitan monolaurate and talc.

Capsule ingredients: ammonia solution, black iron oxide, gelatin, iron oxide yellow, potassium hydroxide, propylene glycol, purified, shellac, sodium dodecyl sulfate and titanium dioxide.

Sugar free.

WHAT LOPIMUNE 40/10 ORAL PELLETS ARE USED FOR:

LOPIMUNE 40/10 ORAL PELLETS are indicated in combination with other antiretroviral agents for the treatment of HIV infection.

BEFORE YOU TAKE LOPIMUNE 40/10 ORAL PELLETS:

Do not take LOPIMUNE 40/10 ORAL PELLETS if:

You are allergic to lopinavir, ritonavir or to any of the other ingredients of **LOPIMUNE 40/10 ORAL PELLETS.**

You have severe liver disease.

You are currently taking any of the following medicines:

- Sedatives such as midazolam or triazolam (for anxiety or sleeping problems).
- Ergot derivatives, such as ergotamine, dihydroergotamine, and methylergonovine (for the treatment of migraines).
- Antipsychotics, such as pimozide (to treat schizophrenia).
- Gastrointestinal motility agents/stomach complications medicines such as cisapride.
- Sildenafil, (used for pulmonary hypertension).
- Alfuzosin (used for treatment of enlarged prostate).
- Antidysrhythmics, used for heart conditions- abnormal heart rhythms like dronedarone.
- Colchicine, used for treating gout.
- *Hypericum perforatum* (St. John's Wort for depression).
- Rifampin, which is used for treating tuberculosis (TB).
- Cholesterol (too much fat in your body) lowering medicines like simvastatin and lovastatin.

Take special care with LOPIMUNE 40/10 ORAL PELLETS:

Combination antiretroviral treatment such as **LOPIMUNE 40/10 ORAL PELLETS** is associated with complications that may need medical attention. Consult your doctor if the following problems arise while on **LOPIMUNE 40/10 ORAL PELLETS** treatment:

- Body fat redistribution indicated by central obesity, dorso-cervical fat, enlargement (buffalo hump), peripheral wasting, facial wasting, breast enlargement.
- You become prone to opportunistic infections including TB, flu, skin and/or eye infections. Fever and generally feeling weak can be a sign of infections.
- Bone death and/or wasting. Your bones may become weakened and painful, this condition may be aggravated by corticosteroid use and alcohol consumption.
- Diabetes mellitus (high blood sugar), signs include unusual thirst and frequent urination.
- Abdominal pain, this can be a sign of liver impairment or impaired function of the pancreas.
- You become prone to bleeding.
- You have a history of or suffer from heart problems like abnormal heart rhythms, this can be life-threatening due to the risk of heart attacks.
- You previously developed resistance to other HIV medicines.
- You have haemophilia type A and B as there may be an increased risk of bleeding.

The risk of HIV transmission to others

The current antiretroviral medicines including **LOPIMUNE 40/10 ORAL PELLETS** do not prevent the risk of transmission of HIV to others through sexual contact or blood contamination. You need to take appropriate measures such as using condoms while having sexual intercourse

or a protective barrier when you are about to be in contact with blood in order to protect yourself and others from the risk of HIV transmission.

Important information if you are using other medicines with LOPIMUNE 40/10 ORAL PELLETS

You must take **LOPIMUNE 40/10 ORAL PELLETS** (and other ARV medicines) every day as prescribed by your doctor.

LOPIMUNE 40/10 ORAL PELLETS may interact with some medicines; therefore, report to your doctor if you are using any other prescription, non-prescription medication or herbal products, particularly St. John's Wort (*Hypericum perforatum*).

If you are taking oestrogen-based hormonal contraceptives, ensure that you use additional or alternate contraceptive measures during therapy with **LOPIMUNE 40/10 ORAL PELLETS** as your oestrogen-based hormonal contraceptive medicine may become ineffective.

Taking LOPIMUNE 40/10 ORAL PELLETS with food and drink:

LOPIMUNE 40/10 ORAL PELLETS must always be taken with food (see '**HOW TO TAKE LOPIMUNE 40/10 ORAL PELLETS'**).

Pregnancy and breastfeeding:

If you are pregnant or breastfeeding your baby, please consult your doctor, pharmacist or other healthcare professional for advice before taking LOPIMUNE 40/10 ORAL PELLETS.

Do not take **LOPIMUNE 40/10 ORAL PELLETS** if you are pregnant.

Do not breastfeed your child while taking **LOPIMUNE 40/10 ORAL PELLETS** or if you have HIV disease.

Driving and using machinery:

LOPIMUNE 40/10 ORAL PELLETS can cause visual disturbances and drowsiness and can therefore impair your ability to drive or operate machines. You need to exercise caution as you should only perform these functions if you are aware that **LOPIMUNE 40/10 ORAL PELLETS** does not have a negative influence on these abilities.

Taking other medicines with LOPIMUNE 40/10 ORAL PELLETS:

Always tell your healthcare professional if you are taking any other medicine.

This includes complementary or traditional medicines.

Do not take product LOPIMUNE 40/10 ORAL PELLETS with the following medicines:

- Alfuzosin (treatment of enlarged prostate).
- Sedatives like midazolam and triazolam (they are used for treating fits and sleeplessness).
- Lovastatin and simvastatin (treatment of high cholesterol).
- Sildenafil (used for pulmonary hypertension).
- Dronedarone (used for abnormal heart rhythms).
- Herbal product, St John's Wort (*Hypericum perforatum*), for treatment of depression.
- Ergot alkaloids (dihydroergotamine, ergotamine and methylergonovine), they are used to treat migraines.
- Colchicine (for treating gout).

- Pimozide (for depression associated with bipolar disorder).
- Cisapride (for treatment of bowel motility).
- Rifampin, used for treating TB.

Tell your doctor if you are taking the following medicines as your dose may need to be adjusted and/or it may be necessary to monitor your health:

- Other medicines for treating HIV disease.
- Anticoagulants, e.g. warfarin and rivaroxaban (medicines to prevent blood clots).
- Anticonvulsants (medicines for fits or seizures such as phenobarbitone, phenytoin, carbamazepine, lamotrigine and valproate).
- Antidepressants (bupropion and trazodone).
- Antiparasitic (medicines like atovaquone used in the prevention and/or treatment of malaria).
- Corticosteroids (dexamethasone, budesonide, prednisone and fluticasone propionate).
- High blood pressure medicines (felodipine and nifedipine).
- Medicines for lowering blood cholesterol such as atorvastatin and rosuvastatin.
- Immuno-suppressants, medicines that compromise your immune system, including ciclosporin, tacrolimus and sirolimus.
- Antifungals including ketoconazole, itraconazole and voriconazole.
- Antibacterials such as clarithromycin (used in the treatment of infections).
- TB medicines such as rifabutin and bedaquiline.
- Pain relief medicines like methadone and fentanyl.
- Oral/ patch contraceptives (ethinyl estradiol).
- Medicines for erectile dysfunction such as tadalafil and vardenafil.

- Salmeterol (used for prevention of asthma).
- Cancer medicines including dasatinib, nilotinib, vincristine and vinblastine.
- Quetiapine, used for mood disorders and depression.

HOW TO TAKE LOPIMUNE 40/10 ORAL PELLETS:

Do not share medicines prescribed for you with any other person.

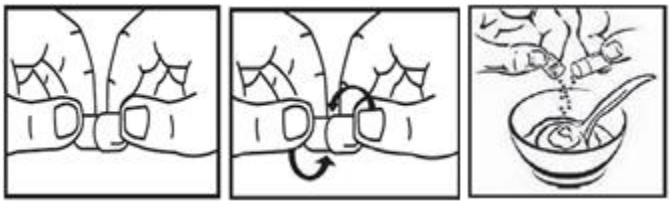
Always take **LOPIMUNE 40/10 ORAL PELLETS** exactly as your doctor has instructed you. You should check with your doctor or pharmacist if you are unsure.

Caregivers, parents or guardians must also give **LOPIMUNE 40/10 ORAL PELLETS** according to the patient's doctor's instructions.

*REMEMBER TO ALWAYS TAKE **LOPIMUNE 40/10 ORAL PELLETS** WITH FOOD.*

Method of administration:

Capsules containing the oral pellets should not be swallowed whole but should be administered with food as shown below (capsule contents should be emptied into ready-to-eat-porridge).



- Place sweetened porridge, which is at room temperature, in a small bowl.
- Obtain the prescribed number of capsules needed for a dose.
- Hold both ends of the capsule between your fingertips as depicted above.
- Twist the ends of the capsule in opposite direction and pull apart so that the entire contents of the capsule are sprinkled over the sweetened porridge.

- Repeat this step for the prescribed number of capsules per dose. Ensure that the entire content of each capsule is sprinkled over the porridge.
- This medicine-porridge mixture should be eaten immediately. The oral pellets should not be chewed or crushed. They should not be stored for future use.
- Administration of the required dose should be followed by drinking water, to ensure that no pellets are left behind in the mouth.
- Repeat above steps for next dose.

The recommended dose must be taken twice daily with food. You doctor will determine the correct dose for you.

Dosage of LOPIMUNE 40/10 ORAL PELLETS in children without nevirapine, efavirenz, amprenavir and nelfinavir using a weight band approach:

Weight of child	**Recommended dose to be taken twice per day**
5 kg to less than 6 kg	80/20 mg (2 capsules)
6 kg to less than 10 kg	120/30 mg (3 capsules)
10 kg to less than 14 kg	160/40 mg (4 capsules)
14 kg to less than 20 kg	200/50 mg (5 capsules)
20 kg to less than 25 kg	240/60 mg (6 capsules)
25 kg to less than 30 kg	280/70 mg (7 capsules)
30 kg to less than 35 kg	320/80 mg (8 capsules)
Equal to and greater than 35 kg	Adult dose, 400/100 mg (10 capsules)

Your doctor will determine the correct dose for you if you are taking other medicines which may include nevirapine, efavirenz, amprenavir or nelfinavir.

If you take more LOPIMUNE 40/10 ORAL PELLETS than you should:

In the event of overdosage, consult your doctor or pharmacist. If neither is available, contact the nearest hospital or poison control centre.

If you miss a dose of LOPIMUNE 40/10 ORAL PELLETS:

Do not take a double dose to make up for forgotten individual missed doses.

POSSIBLE SIDE EFFECTS:

Not all side effects reported for **LOPIMUNE 40/10 ORAL PELLETS** are included in this leaflet. Should your general health worsen or if you experience any untoward effects while taking **LOPIMUNE 40/10 ORAL PELLETS**, please consult your doctor, pharmacist or other healthcare professional for advice.

If any of the following happens, stop taking LOPIMUNE 40/10 ORAL PELLETS and tell your doctor immediately or go to the casualty department at your nearest hospital:

- Irregular (too high or slow) heart rates, this might be risk factor for stroke.
- Heart attack.
- Liver disease or liver failure (symptoms include abdominal pain, nausea and vomiting).
- Yellowing of the skin and/or eyes, also known as jaundice.
- Kidney disease, including kidney stones, signs include severe back pain and pain on the sides of your body and groin area and painful and frequent urination.
- Steven-Johnson syndrome, a life-threatening condition indicated by fever, fatigue and development of painful widespread blisters and ulcers on the skin, mouth area, face and limbs.

- Erythema multiforme, life threatening conditions affecting the skin, symptoms include itchy and/or sore, mostly reddish-looking skin rash, fever, feeling weak, eye pain and joint aches.

These are all very serious side effects. If you have them, you may have had a serious reaction to ***LOPIMUNE 40/10 ORAL PELLETS****. You may need urgent medical attention or hospitalisation.*

Tell your doctor immediately or go to the casualty department at your nearest hospital if you notice any of the following:

Frequent side effects:

- High blood pressure (may be indicated by nosebleeds and shortness of breath).
- Diarrhoea (runny tummy, possibly with nausea and vomiting).
- Any form of infections such as flu-like illness, sore throat, persistent coughs, coughing bloody phlegm, occurrence of boils on your skin, blisters or cold sores in your mouth or groin, urine retention and burning urine (may be a sign of urinary tract infection), ear infection, sneezing and stomach bug. Usually, infections are accompanied by fever and fatigue.
- Diabetes, high blood sugar (excessive thirst, sweating and frequent urination may be a sign of high blood sugar).
- Frequent lightheadedness (also called postural hypotension).

Less frequent side effects:

- Anaemia, you may feel unusually faint and appear pale.

- Blood clots (this condition is asymptomatic but risk factors include tenderness, pain, discolouration and swelling around affected veins).
- Cholecystitis (also known as gall stones, you might experience severe belly pain accompanied by nausea and vomiting).
- Asthma.
- Breathlessness.
- Chest pain.
- Eczema (skin disorder with skin becoming rough and blistered, with bleeding and itchiness).
- Appearance of moles and lumps on the skin.
- Encephalopathy (brain injury/ brain dysfunction, numerous parts of the body are affected by this syndrome, symptoms include loss of cognitive function, inability to concentrate, not being in control of emotions, impaired muscle function/ breathing/ body movements).

These are serious side effects. You may need urgent medical attention.

Tell your doctor if you notice any of the following:

Frequent side effects:

- Headaches
- Insomnia/ sleeplessness
- Rash
- Buffalo hump (accumulation of body fat around neck and shoulder)
- Abnormal state of being exhausted
- Abdominal pain (stomach ache)
- Abnormal stools
- Indigestion

- Nausea
- Vomiting
- Flatulence
- Muscle spasm/ pain
- Muscle weakness
- Back pain
- Heavy menstrual periods
- Lack of menstruation

Less frequent side effects:

- Pain
- General discomfort
- Chills
- Migraine (severe and recurrent headaches)
- Cushing's syndrome
- Weight gain
- Weight loss
- Dehydration
- Increased appetite
- Lactic acidosis (too much acid in your body generally indicated by muscle weakness and rapid breathing)
- Obesity
- Anorexia
- Abnormal dreams
- Agitation

- Anxiety
- Confusion
- Depression
- Emotional lability
- Decreased libido
- Nervousness
- Apathy (detached state of emotions)
- Abnormal thinking
- Amnesia (memory loss)
- Drowsiness
- Dizziness
- Impaired or involuntary muscle movements
- Inability to control body movements
- Increased muscle tension
- Facial paralysis
- Loss of taste
- Taste perversion
- Tingling or a feeling of pins and needles
- Tremors
- Convulsions (fits or seizures)
- Abnormal vision
- Eye disorder
- Being disorientated
- Ringing in the ears
- Varicose vein (enlarged veins, mostly occurring on your legs)

- Enlarged abdomen (inflated belly)
- Constipation
- Dry mouth
- Difficulty swallowing
- Belching
- Heartburn
- Bowel incontinence
- Acne
- Alopecia (hair loss)
- Dry skin
- Skin rash that makes your skin appear scaly
- Nail disorder
- Itchiness
- Seborrhea (increased skin secretions similar to having oily skin problem)
- Skin discolouration
- Increased sweating
- Stretch marks
- Joint pain
- Abnormal ejaculation
- Impotence
- Breast enlargement.

If you notice any side effects not mentioned in this leaflet, please inform your doctor or pharmacist.

STORING AND DISPOSING OF LOPIMUNE 40/10 ORAL PELLETS:

Store in the original container at or below 30 °C.

KEEP OUT OF REACH AND SIGHT OF CHILDREN.

PRESENTATION OF LOPIMUNE 40/10 ORAL PELLETS:

LOPIMUNE 40/10 ORAL PELLETS oral pellets in capsules are packed in pack sizes of 120's in white HDPE containers fitted with white HDPE lids, containing two 1 g silica gel dessicants.

IDENTIFICATION OF LOPIMUNE 40/10 ORAL PELLETS:

White to off-white, circular, biconvex, pellets plain on both sides filled in "size 1" hard gelatin capsules that have a clear, transparent body with "414" sprinted in black ink and yellow caps with "CL" sprinted in black ink.

REGISTRATION NUMBER:

51/20.2.8/0123

NAME AND ADDRESS OF REGISTRATION HOLDER:

CIPLA MEDPRO MANUFACTURING (PTY) LTD.

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