

WARNING:

If you are taking **NORVIR** with some medicines used to treat **allergies**, medicines used to relieve anxiety and/or trouble sleeping, medicines used to correct irregular heart beat and medicines used to treat migraine headache they may cause serious adverse events.

SCHEDULING STATUS**S4****PROPRIETARY NAME (strength and dosage form)****NORVIR 80 mg/mL - Solution****Ritonavir****NORVIR SEC 100 mg -Soft Elastic Capsule****Ritonavir****NORVIR 100 mg TABLET – Tablet****Ritonavir****NORVIR ORAL POWDER 100 mg – Powder for Oral suspension****Ritonavir****Read all of this leaflet carefully before you start taking NORVIR**

- Keep this leaflet. You may need to read it again.
- If you have further questions please ask your doctor or pharmacist.
- **NORVIR** 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.

1. WHAT NORVIR CONTAINS

NORVIR 80 mg/mL solution

- The active substance is ritonavir. Each 1 mL solution contains 80 mg ritonavir.
- The other ingredients are alcohol (43.2 % v/v), water, polyoxyl 35 castor oil, propylene glycol, citric acid, saccharin sodium, peppermint oil, creamy caramel flavouring and the dye FD&C Yellow No. 6 (E110).

NORVIR SEC 100 mg soft elastic capsules

- The active substance is ritonavir. Each capsule contains 100 mg ritonavir.
- The other ingredients are alcohol (12.0 % v/v), oleic acid, butylated hydroxytoluene and polyoxyl 35 castor oil.
- The capsule shell components are gelatine, sorbitol, glycerine, titanium dioxide (white colour), medium chain triglycerides, lecithin and black ink containing propylene glycol, black iron oxide, polyvinyl acetate phthalate, polyethylene glycol 400 and ammonium hydroxide.

NORVIR 100 mg Tablet

- The active substance is ritonavir. Each film-coated-tablet contains ritonavir 100 mg
- Other ingredients include copovidone, sorbitan laurate, colloidal anhydrous silica, sodium stearyl fumarate, calcium hydrogen phosphate anhydrous, hypromellose, titanium dioxide E171, macrogols type 400, hydroxypropyl cellulose, talc, macrogols type 3350 and polysorbate 80.

NORVIR ORAL POWDER

- The active substance is ritonavir. Each sachet of oral powder contains 100 mg of ritonavir.
- The other ingredients are copovidone, sorbitan laurate, and colloidal anhydrous silica.

2. WHAT NORVIR IS USED FOR

NORVIR is in a class of medicines called the HIV protease inhibitors. **NORVIR** is used in combination with other anti-HIV medicines to treat human immunodeficiency virus (HIV) infection.

3. BEFORE YOU TAKE NORVIR

Do not take NORVIR:

- If you are hypersensitive (allergic) to ritonavir or any of the other ingredients of **NORVIR** as listed in 'other ingredients' above.
- If you have severe liver disease.
- Do not take **NORVIR** if you are taking any of the following medications
 - amiodarone, bepridil, encainide, flecainide, propafenone, quinidine, digoxin (used to correct irregular heart beat)
 - cisapride (used to relieve some types of stomach problems)
 - dihydroergotamine, ergotamine, (used to treat migraine headache)
 - lovastatin and simvastatin (used to treat elevated cholesterol)
 - midazolam, triazolam (used to relieve anxiety and/or trouble sleeping)
 - pimozone (used to treat schizophrenia)

- rifampicin (used to treat tuberculosis and leprosy)
- salmeterol (used to treat asthma)
- sildenafil (used to treat pulmonary arterial hypertension)
- St. John's Wort (herbal product sold as a dietary supplement)
- Voriconazole (used for treatment of fungal infections)

Take special care with NORVIR:

- If you are taking certain medicines (see **Taking NORVIR with other medicines**). These could cause serious side-effects that could cause death. Before you take **NORVIR**, you must tell your doctor about all the medicines you are taking or are planning to take. These include other prescription and non-prescription medicines and herbal supplements.
- Some patients taking **NORVIR** can develop serious problems with their pancreas (pancreatitis), which may cause death. Tell your doctor as soon as possible if you have nausea, vomiting, or abdominal pain. These may be signs of pancreatitis.
- Blood tests in patients taking **NORVIR** may show possible liver problems. People with liver disease such as Hepatitis B and Hepatitis C who take **NORVIR** may have worsening liver disease. Liver problems including rare cases of death have occurred in patients taking **NORVIR**. It is unclear if **NORVIR** caused these liver problems because some patients had other illnesses or were taking other medicines.
- Some patients have large increases in triglycerides and cholesterol (fats in the blood).

- Diabetes and high blood sugar (hyperglycemia) may occur if you are taking protease inhibitors such as **NORVIR**. You may develop diabetes before starting protease inhibitors. If you are taking **NORVIR** your diabetes medication may need changing.
- If you have liver problems or are infected with Hepatitis B or Hepatitis C (an infection of the liver). Talk to your doctor before taking **NORVIR**.
- If you have haemophilia (a disease that affects the ability of your blood to clot properly and causes increased bleeding) and are taking protease inhibitors such as **NORVIR** you may have increased bleeding. It is not known whether the protease inhibitors caused these problems. Be sure to tell your doctor if you have haemophilia types A and B.
- Changes in body fat have been seen in patients taking medicines such as **NORVIR** to treat HIV. These changes may include increased amount of fat in the upper back and neck ('buffalo hump'), breast and around the trunk. Loss of fat from the legs, arms and face may also happen.
- **NORVIR** does not reduce the risk of passing HIV to others through sexual contact or blood contamination. Continue to practice safe sex and do not use or share dirty needles.
- **NORVIR** does not cure HIV infection or AIDS. People taking **NORVIR** may still get opportunistic infections or other conditions that happen with HIV infection.
- If you are feeling sick (nausea), vomiting (throwing up) or have stomach pain, these may be signs of inflammation of the pancreas (pancreatitis). Some patients taking

Norvir can develop serious problems with their pancreas. Tell your doctor as soon as possible if this applies to you.

- Some patients with advanced HIV infection (AIDS) who then start anti-HIV treatment may develop the symptoms of infections they have had in the past even if they didn't know they had had them. It is believed that this happens because the body's immune response improves and helps the body to fight these infections.

- Autoimmune disorders (a condition that occurs when the immune system attacks healthy body tissue) may also occur after you start taking medicines for the treatment of your HIV infection. Autoimmune disorders may occur many months after the start of treatment. If you notice any symptoms of infection or other symptoms such as muscle weakness, weakness beginning in the hands and feet and moving up towards the trunk of the body, palpitations, tremor or hyperactivity, please inform your doctor immediately to seek necessary treatment.

- Joint stiffness, aches and pains (especially of the hip, knee and shoulder) and difficulty moving, may occur and this may be a sign of a problem that can destroy bone (osteonecrosis).

Taking NORVIR with food and drink:

NORVIR should preferably be taken with food.

NORVIR 80 mg/mL solution contains alcohol. Talk with your doctor if you are taking or planning to take metronidazole or disulfiram. Severe nausea and vomiting can occur if **NORVIR** is taken with these medicines

Pregnancy and Breastfeeding:

- If you are pregnant or planning to become pregnant, you should not be taking **NORVIR**.
The effects of **NORVIR** on pregnant women or their unborn babies are not known.
- If you are breastfeeding: Do not breastfeed your baby if you are taking **NORVIR**. You should not breastfeed if you have HIV. You should be aware that if your baby does not already have HIV, there is a chance that HIV can be transmitted through breastfeeding

Driving and using machinery:

Sleepiness and dizziness are known undesirable effects of **NORVIR** which may interfere with the ability to perform potentially hazardous tasks like driving a car or operating heavy machinery.

Important information about some of the ingredients of NORVIR:

NORVIR 80 mg/mL solution and **NORVIR SEC 100 mg** soft elastic capsules contain alcohol. It could be potentially harmful to you if you are suffering from liver disease, alcoholism, epilepsy, brain injury or disease as well as if you are pregnant.

NORVIR 80 mg/mL solution and **NORVIR SEC 100 mg** soft elastic capsules contain polyoxyl 35 castor oil. This may cause nausea, vomiting, colic and severe purgation at high doses. It should not be given when intestinal obstruction is present or suspected.

NORVIR 80 mg/mL solution contains sodium as saccharin sodium, which may be harmful to you if you are on a low sodium diet.

NORVIR 80 mg/mL solution contains the dye FD&C Yellow No. 6 (E110). This can cause allergic reactions including asthma. Allergy is more common in those people who are allergic to aspirin.

Taking other medicines with NORVIR:

- Always tell your healthcare professional if you are taking any other medicine, this includes complementary or traditional medicines.

- Do not take **NORVIR** with St. John's wort (*Hypericum perforatum*), an herbal product sold as a mild antidepressant or product containing St. John's wort. . Taking St. John's wort may decrease **NORVIR** levels and lead to increased viral load and possible resistance to **NORVIR** or cross-resistance to other antiretroviral medicines.

- Do not take **NORVIR** with the cholesterol-lowering medicines lovastatin or simvastatin because of possible serious reactions. There is also an increased risk of medicine interactions between **NORVIR** and atorvastatin;

- Do not take **NORVIR** with voriconazole. Taking **NORVIR** with voriconazole may reduce the levels of voriconazole.

- Talk to your doctor if you are taking or are planning to take fluticasone propionate. Taking fluticasone propionate and **NORVIR** can increase the levels of fluticasone in your blood and could cause serious side effects.

- If you are taking sildenafil, tadalafil or vardenafil to treat erectile dysfunction, your doctor may lower your dose of these medicines. Before you take sildenafil, tadalafil or

ildenafil with **NORVIR**, talk to your doctor about possible medicine interactions and side effects.

- Do not take **NORVIR** if you are taking sildenafil to treat pulmonary arterial hypertension.
- If you are using birth control pills or birth control patches or implants to prevent pregnancy, your doctor should increase the dose of your birth control treatment or you should use a different type of birth control method since **NORVIR** may reduce the effectiveness of these medicines.
- If you are taking rifabutin, your doctor may lower the dose of rifabutin.
- If you are taking both didanosine and **NORVIR**, the dosage times of Didanosine and **NORVIR** should be separated by at least 2.5 hours.
- Tell your doctor if you are taking rifampin as it may reduce blood levels of **NORVIR**. (see “Do not take **NORVIR**”)

4. HOW TO TAKE NORVIR

- Always take **NORVIR** exactly as your doctor has instructed you. Do not change your treatment or stop treatment without first talking with your doctor. If you are not sure, you should check with your doctor or pharmacist.

Dosage:

- Your doctor will prescribe the correct dose for you.

- Take **NORVIR** oral powder or solution with food if possible.
- Take **NORVIR** capsule or solution with food if possible.
- Take **NORVIR** tablets with food.

- **NORVIR** tablets should be swallowed whole and not chewed, broken or crushed.
- **NORVIR SEC 100 mg** soft elastic capsule should be swallowed whole with a lot of water without opening or splitting it.

- **NORVIR** oral powder can be sprinkled on soft food such as apple sauce or dessert pudding or mixed with a suitable liquid such as water, chocolate milk, or infant formula.
- ***Administering with food***
- The entire contents of each sachet are to be poured over a small amount of soft food (e.g. apple sauce or vanilla pudding). All of the mixed soft food must be administered within 2 hours.
- ***Administering with liquid***
- The entire contents of each sachet should be suspended in 9.4 mL of liquid (water, chocolate milk, or infant formula)
- The oral dosing syringe and mixing cup should be dry prior to use.
- Draw up 9.4 mL of liquid using the provided oral dosing syringe, remove the bubbles, and transfer the liquid to the mixing cup. All measuring should be done in mL using the syringe.
- Pour the entire contents of 1 sachet (100 mg) into the mixing cup.
- Close the lid and shake hard for at least 90 seconds until all the lumps have dissolved.
- Let the liquid stand for 10 minutes in order for most of the bubbles to disappear.

- Use the provided oral dosing syringe to measure and administer the prescribed volume is sure to remove the bubbles prior to dose administration.
- Once the powder is mixed, the dosage must be consumed within 2 hours. Discard any mixture remaining in the mixing cup
- You should always use the measuring cup provided with the medicine to take your **NORVIR 80 mg/mL** solution in order to make sure you take the right dose. Wash the measuring cup after use with dish soap and warm water as soon as you can. The cup should be dry prior to re-use.
- When your **NORVIR** supply starts to run low, get more from your doctor or pharmacy. This is very important because the amount of virus in your blood may increase if the medicine is stopped for even a short time. The virus may develop resistance to **NORVIR** and become harder to treat.
- Be sure to set up a schedule and follow it carefully.
- Only take medicine that has been prescribed specifically for you. Do not give **NORVIR** to others or take medicine prescribed for someone else.

If you take more NORVIR than you should:

If you think that you took more than the prescribed dose of **NORVIR**, contact your doctor or go to the nearest hospital.

NORVIR should be kept out of reach of young children. **NORVIR 80 mg/mL** solution contains a large amount of alcohol. If a toddler or young child accidentally drinks more than the recommended dose of **NORVIR**, it could make him/her sick from too much

alcohol. Contact your doctor, local poison control centre or emergency room immediately if it happens.

If you forget to take / missed a dose of NORVIR:

It is important that you do not miss any dose. If you miss a dose of **NORVIR**, take it as soon as possible and then take your next scheduled dose at its regular time. If it is almost time for your next dose, wait and take the next dose at the regular time. Do not double the next dose. Call your doctor or pharmacist if you have any questions about how to take **NORVIR**.

5. POSSIBLE SIDE EFFECTS

NORVIR can have side effects. Not all side effects reported for **NORVIR** are included in this leaflet. Should your general health worsen while taking **NORVIR**, please consult your doctor, pharmacist or other health care professional for advice.

- The frequent side effects reported are: diarrhoea, vomiting, nausea, abdominal pain, feeling weak/tired, headache and dizziness, loss of appetite, changes in taste and tingling feeling or numbness in hands or feet or around the lips.
- Less frequent side effects are: heart attack, diabetes and kidney failure.
- Allergic reactions ranging from mild to severe have occurred in patients taking **NORVIR**.

Frequent side effects:

Sore throat

Allergic reaction including rash (may be red, raised, and itchy) severe swelling of the skin and other tissues

Weight loss

Dehydration, increase in cholesterol, gout, swelling of hands and feet

Agitation, confusion, mood alteration, nervousness

Tingling sensation or numbness in the hands, feet or around the lips and mouth.

Bad taste in the mouth; dizziness

Blurred vision, painful eye, sensitivity to light

Low blood pressure and feeling faint, when getting up, increase in heart beat/rate

Cough

Stomach ache, vomiting, diarrhoea, nausea, dry mouth, wind, indigestion, mouth ulcer

Inflammation of the liver and yellowing of the skin or whites of the eye

Skin disorders (rash, redness dry skin, inflammation, swelling of the skin, itchiness), acne

Muscular pain

Joint pain and back pain

Increase/decrease in urination, reduced kidney function

Feeling weak/tired, fever

Changes in liver function tests

Flushing, feeling hot

Less frequent side effects:

Changes in blood test results, Low levels of blood platelets

High levels of sugar in the blood

Buzzing/roaring sound in the ear, Pain in the ear, Imbalance

Abnormal dream, difficulty in paying attention, convulsions (fits)

Change in the heart rate, fainting

Shortness of breath, nose bleeding

Decreased sexual drive

Severe diarrhoea, constipation, blood in the stool

If you have questions about side-effects, ask your doctor, or pharmacist.

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

6. STORING AND DISPOSING OF NORVIR

Keep **NORVIR** and all other medicines out of reach of children.

NORVIR 80 mg/mL solution

- Store **NORVIR 80 mg/mL** solution at room temperature. Do not refrigerate **NORVIR 80 mg/mL** solution. Avoid exposing **NORVIR 80 mg/mL** solution to excessive heat or cold.
- Shake **NORVIR 80 mg/mL** solution well before each use.
- The **NORVIR 80 mg/mL** solution dosage cup should be cleaned immediately with hot water and dish soap after use. The dosage cup **must** be dry prior to use.
- Use **NORVIR 80 mg/mL** solution by the expiration date on the bottle.

NORVIR ORAL POWDER

- **NORVIR oral powder** must be stored below 30°C. The reconstituted powder must be consumed within 2 hours.
- Avoid exposure to excessive heat.

- The oral dosing syringe and mixing cup should be cleaned immediately with warm water and soap after use. When cleaned immediately, the residue is removed. The oral dosing syringe and mixing cup should be dry prior to use.

NORVIR SEC 100 mg capsules

- Refrigeration of **NORVIR SEC 100 mg** soft gelatin capsules by the patient is recommended, but not required if used within 30 days and stored below 25 °C. Avoid exposing **NORVIR SEC 100 mg** soft gelatin capsules to excessive heat or cold.
- Store **NORVIR SEC 100 mg** soft gelatin capsules and **NORVIR 80 mg/mL** solution in the original container.

NORVIR 100 mg Tablet

- Should be stored at room temperature (below 30 °C).
- Do not keep medicine that is out of date or that you no longer need. Be sure that if you throw any medicine away, and it is out of reach of children.
- Do not use **NORVIR** if you notice 'visible sign of deterioration'.
- Return all unused medicine to your pharmacist. Do not dispose of unused medicine in drains of sewerage systems (e.g. toilets).

7. PRESENTATION OF NORVIR

NORVIR 80 mg/mL solution comes in amber coloured bottles containing 90 mL and 225 mL. A dosage cup containing graduations at 3.75 mL (300 mg dose), 5 mL (400 mg dose), 6.25 mL (500 mg dose) and 7.5 mL (600 mg dose) is provided.

NORVIR SEC 100 mg capsules come in plastic bottles containing 84 capsules.


NORVIR 100 mg tablets are contained in a white plastic bottle containing either 30 or 60 tablets


NORVIR ORAL POWDER

NORVIR oral powder is available in 30 foil-laminate single-use, child-resistant sachets per carton. The carton also contains a mixing cup and 10 mL calibrated oral dosing syringe.

8. IDENTIFICATION OF NORVIR

NORVIR 80 mg/mL solution is a practically clear, orange solution with a peppermint and caramel odour.

NORVIR SEC 100 mg soft elastic capsules are white oblong capsules printed with Abbott logo , 'DS' and '100' in black ink. The capsules contain a clear liquid essentially free of particles.

NORVIR 100 mg Tablet is a white, oval film-coated tablet debossed with Abbott logo  and "NK" on one side.

NORVIR oral powder: Powder is beige/pale yellow to yellow in colour. Reconstituted powder with water: When reconstituted with water the colour may range from white to yellow.

9. REGISTRATION NUMBER

NORVIR 80 mg/mL : 31/20.2.8/0217

NORVIR SEC 100 mg : 34/20.2.8/0175

NORVIR 100 mg Tablet: 44/20.2.8/0128

NORVIR oral powder: 51/20.2.8/0154

**10. NAME ,BUSINESS ADDRESS AND TELEPHONE NUMBER OF REGISTRATION
HOLDER**

AbbVie (Pty) Ltd

Abbott Place, 219 Golf Club Terrace

1709, Constantia Kloof,

Republic of South Africa

Tel: 011 831 3200

11. DATE OF PUBLICATION

25 September 2018