Applicant : Eurolab (Pty) Ltd
Proprietary name : Enzutix
Dosage form and strength : Soft capsules, 40 mg
Date of submission : 12 May 2022
Approval date : 24 January 2023

APPROVED PATIENT INFORMATION LEAFLET FOR ENZUTIX

SCHEDULING STATUS:

S4

ENZUTIX 40 mg soft capsules

Enzalutamide

Contains sorbitol

Read all of this leaflet carefully before you start taking ENZUTIX.

- Keep this leaflet. You may need to read it again.
- If you have further questions, please ask your doctor, pharmacist, nurse or other health care provider.
- **ENZUTIX** 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 is in this leaflet

1. What **ENZUTIX** is and what it is used for
2. What you need to know before you take **ENZUTIX**
3. How to take **ENZUTIX**
4. Possible side effects
5. How to store **ENZUTIX**
6. Contents of the pack and other information

1 What ENZUTIX is and what it is used for

ENZUTIX contains the active substance enzalutamide. It is used to treat adult men with prostate cancer that no longer responds to lowering of testosterone levels by either medical or surgical castration.

ENZUTIX works by blocking the activity of hormones called androgens (such as testosterone). By blocking androgens, enzalutamide stops prostate cancer cells from growing and dividing.

2 What you need to know before you take ENZUTIX

Do not take ENZUTIX:

- if you are hypersensitive (allergic) to enzalutamide or any of the other ingredients of this medicine (listed in section 6).
- if you are pregnant, may become pregnant or breastfeeding a baby (see 'Pregnancy, breast-feeding and fertility'), as **ENZUTIX** is not for use in women.

Warnings and Precautions

Take special care with **ENZUTIX**:

- if you ever had or are at risk of having a seizure (a fit) (see 'Warnings and precautions: seizures' below).
- If you have been diagnosed by your doctor with a condition called posterior reversible encephalopathy syndrome (PRES). (see 'Warnings and precautions: Posterior Reversible Encephalopathy Syndrome' below).
- if you are taking any medicines to prevent blood clots (e.g. warfarin, clopidogrel). Your doctor will regularly send you for blood tests to check that your medicine to prevent blood clots are working.
- if you have problems with your kidneys.
- if you have problems with your liver.

- if you have any heart or blood vessel conditions, including heart rhythm problems (arrhythmia) or uncontrolled high blood pressure, or if are being treated with medicines for these conditions. The risk of heart rhythm problems may be increased when using **ENZUTIX**.
- if you have been diagnosed with changes in heart rhythm (QT prolongation).
- if you currently receive docetaxel chemotherapy.
- if you are hypersensitive (allergic) to enzalutamide or any of the other ingredients of this medicine (listed in section 6).
- if you have been diagnosed with sorbitol intolerance. **ENZUTIX** contains sorbitol.

Seizures

ENZUTIX has been shown to cause seizures. You may have a higher risk of seizures in the following scenarios:

- if you had earlier episodes of seizures.
- if you drink large amounts of alcohol either regularly or from time to time.
- if you have had a serious head injury or a history of head trauma.
- if you have had a stroke.
- if you have had a brain tumour or metastases of cancer in the brain.
- if you are taking a medicine that can cause seizures or medicines that can increase the susceptibility for having seizures. Ask your doctor or pharmacist if you are not sure.

If you have a seizure during treatment

Stop taking **ENZUTIX** and do not take any more capsules. See your doctor, pharmacist or other health care provider as soon as possible.

Posterior Reversible Encephalopathy Syndrome

ENZUTIX can cause a rare reversible condition of the brain called posterior reversible encephalopathy syndrome (PRES). If you have a seizure, worsening headache, confusion, blindness or other vision problems, contact your doctor as soon as possible (see section 4 'Possible side effects').

If any of the above applies to you or you are not sure, talk to your doctor, pharmacist or other health care provider before taking **ENZUTIX**.

Children and adolescents

ENZUTIX is not for use in children and adolescents.

Other medicines and ENZUTIX

Always tell your health care provider if you are taking any other medicine (this includes complementary or traditional medicines).

You need to know the names of the medicines you take. Keep a list of them with you to show to your doctor when you are prescribed a new medicine. You should not start or stop taking any medicine before you talk to your doctor that prescribed **ENZUTIX**.

Tell your doctor if you are taking the following medicines. These medicines may influence the effect of **ENZUTIX**, or **ENZUTIX** may influence the effect of these medicines.

This includes certain medicines used to:

- lower cholesterol (e.g. gemfibrozil, atorvastatin, simvastatin).
- treat pain (e.g. fentanyl, tramadol).
- treat cancer (e.g. cabazitaxel, docetaxel, irinotecan, sunitinib).

- treat epilepsy (e.g. phenobarbitone, carbamazepine, clonazepam, phenytoin, primidone, valproic acid, phenobarbitone).
- treat certain psychiatric disorders such as severe anxiety or schizophrenia (e.g. diazepam, midazolam, haloperidol).
- treat sleep disorders (e.g. zolpidem).
- prevent blood clots (e.g. warfarin, clopidogrel).
- treat heart conditions or lower blood pressure (e.g. bisoprolol, digoxin, diltiazem, felodipine, nicardipine, nifedipine, propranolol, verapamil).
- treat serious disease related to inflammation (e.g. dexamethasone, prednisolone).
- treat HIV infection (e.g. indinavir, ritonavir).
- treat bacterial infections (e.g. clarithromycin, doxycycline, rifampicin).
- treat thyroid disorders (e.g. levothyroxine).
- treat gout (e.g. colchicine).
- treat stomach disorders (e.g. omeprazole).
- prevent heart conditions or strokes (e.g. dabigatran etexilate).
- prevent organ rejection (e.g. ciclosporin, tacrolimus).

ENZUTIX might interfere with some medicines used to treat heart rhythm problems or might increase the risk of heart rhythm problems when used with some other medicines. Tell your doctor if you are taking any of the following medicines:

- medicines used for treatment of abnormal heart rhythm like quinidine, disopyramide, amiodarone and sotalol.
- methadone, a medicine used for pain relief and part of drug addiction detoxification.
- moxifloxacin, an antibiotic.
- antipsychotic medicines used to treat serious mental illnesses.

Tell your doctor if you are taking any of the medicines listed above. The dose of **ENZUTIX** or any other medicines that you are taking may need to be changed.

ENZUTIX with food and drink

ENZUTIX can be taken with or without food.

Pregnancy, breastfeeding and fertility

- **ENZUTIX** is not for use in women.
- **ENZUTIX** may cause harm to the unborn child or potential loss of pregnancy if taken by women who are pregnant. **ENZUTIX** must not be taken by women who are pregnant, may become pregnant, or who are breastfeeding an infant.
- **ENZUTIX** could possibly have an effect on male fertility.
- If you are having sex with a woman who can become pregnant, use a condom and another effective birth control method, during treatment and for 3 months after treatment with **ENZUTIX**.
- If you are having sex with a pregnant woman, use a condom during treatment and for 3 months after treatment with **ENZUTIX** to protect the unborn child.
- Female caregivers see section 3 ‘How to take **ENZUTIX**’ for handling and use.

Driving and using machines

ENZUTIX may influence your ability to drive or use any tools or machines as the side effects of **ENZUTIX** include seizures (fits). If you are at higher risk of seizures (fits), talk to your doctor.

ENZUTIX contains sorbitol

ENZUTIX contains sorbitol (a type of sugar). Sorbitol is a source of fructose. If your doctor has told you that you have an intolerance to some sugars or if you have been diagnosed with hereditary fructose intolerance (HFI), a rare genetic disorder in which a person cannot break down fructose, talk to your doctor before you take **ENZUTIX**.

3 How to take ENZUTIX

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

Always take **ENZUTIX** exactly as your doctor or pharmacist has told you. Check with your doctor or pharmacist if you are not sure.

The usual dose is 160 mg (4 soft capsules), taken at the same time once a day.

Taking **ENZUTIX**:

- Swallow the capsules whole with water.
- Do not chew, dissolve, or open the soft capsules before swallowing.
- **ENZUTIX** can be taken with or without food.
- **ENZUTIX** should not be handled by persons other than the patient and his caregivers.
- **ENZUTIX** should not be handled by women who are or may become pregnant.

Your doctor will tell you how long your treatment with **ENZUTIX** will last. Do not stop treatment early. If you have the impression that the effect of **ENZUTIX** is too strong or too weak, tell your doctor or pharmacist.

If you take more ENZUTIX 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 take more **ENZUTIX** than prescribed, stop taking **ENZUTIX** and contact your doctor. You may have an increased risk of seizure or other side effects.

If you forget to take ENZUTIX

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

- If you forget to take **ENZUTIX** at the usual time, take your usual dose as soon as you remember.
- If you forget to take **ENZUTIX** for the whole day, take your usual dose the following day.
- If you forget to take **ENZUTIX** for more than one day, talk to your doctor without delay.

If you stop taking ENZUTIX

Do not stop taking **ENZUTIX** without talking to your doctor first.

4 Possible side effects

ENZUTIX can have side effects.

Not all side effects reported for **ENZUTIX** are included in this leaflet. Should your general health worsen or if you experience any untoward effects while taking **ENZUTIX**, please consult your health care provider for advice.

If any of the following happens, stop taking **ENZUTIX** and tell your doctor immediately or go to the casualty department at your nearest hospital:

- swelling of the hands, feet, ankles, face, lips and mouth or throat, which may cause difficulty in swallowing or breathing;

- rash or itching.

These are all very serious side effects. If you have them, you may have had a serious reaction to **ENZUTIX**. You may need urgent medical attention or hospitalisation.

If any of the following happens, stop taking **ENZUTIX** and tell your doctor immediately:

- seizure (or fit). **ENZUTIX** can cause seizures. This is more likely to happen if you take more than the recommended dose, if you take certain other medicines, or if you are at higher than usual risk of seizures.
- seizure, worsening headache, confusion, blindness, or other vision problems. This may be due to Posterior Reversible Encephalopathy Syndrome (PRES), a rare, reversible condition involving the brain.

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

Other possible side effects include:

Frequent side effects:

- High blood pressure and hot flushes.
- Fatigue (feeling tired), weakness, broken bones and falls.
- Dry and itchy skin.
- Feeling anxious, headache, difficulty remembering, reduced concentration and forgetfulness.
- Blockage of the arteries in the heart (ischemic heart disease).
- Breast enlargement in men (gynaecomastia).
- Symptom of restless legs syndrome (an uncontrollable urge to move a part of the body, usually the leg).

Less frequent side effects:

- Hallucinations, difficulty thinking clearly and seizure.
- Low white blood cell count.
- Seizures (fits)
- Difficulty with learning

Side effects of an unknown frequency:

- Muscle pain, muscle spasms, muscular weakness, and back pain.
- Changes in ECG (QT prolongation).
- Upset stomach including feeling sick (nausea), being sick (vomiting) and diarrhoea.
- Rash and swelling of the face, lips, tongue and / or throat.
- Reduction in blood platelets (which increases risk of bleeding or bruising).
- Posterior Reversible Encephalopathy Syndrome (PRES), a rare, reversible condition involving the brain.

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

Reporting of side effect

If you get side effects, talk to your doctor or pharmacist. You can also report side effects to SAHPRA via the **6.04 Adverse Drug Reaction Reporting Form**, found online under SAHPRA's publications: https://www.sahpra.org.za/Publications/Index/8. By reporting side effects, you can help provide more information on the safety of **ENZUTIX**.

5 How to store ENZUTIX

Store at or below 30 °C.

Store all medicines out of reach of children.

Do not use after the expiry date stated on the carton.

Return all unused medicine to your pharmacist.

Do not dispose of unused medicine in drains or sewerage systems (e.g. toilets).

6 Contents of the pack and other information

What ENZUTIX contains

The active substance is enzalutamide. Each capsule contains 40 mg of enzalutamide.

The other ingredients of the capsule are butylhydroxyanisole, butylhydroxytoluene, and caprylocaproyl macrogolglycerides.

The ingredients of the capsule shell are gelatin, glycerol, sorbitol (liquid partially dehydrated), titanium dioxide (E171), and purified water.

What ENZUTIX looks like and contents of the pack

ENZUTIX is a white to off-white opaque, oblong-shaped soft gelatine capsule containing colourless to slightly yellow, transparent liquid.

ENZUTIX is packed into blister strips packed in carton boxes containing 112 capsules.

Holder of Certificate of Registration

Eurolab (Pty) Ltd.

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This leaflet was last revised in

24 January 2023

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Registration number

55/26/0125