

Package leaflet: Information for the patient

Efavirenz Mylan 600 mg Film-coated Tablets

(efavirenz)

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, pharmacist or nurse. This includes any possible side effects not listed in this leaflet. See section 4.

What is in this leaflet:

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

1. What Efavirenz Mylan is and what it is used for

Efavirenz Mylan contains the active substance efavirenz, which belongs to a class of antiretroviral medicines called non-nucleoside reverse transcriptase inhibitors (NNRTIs). It is an antiretroviral medicine that fights human immunodeficiency virus (HIV) infection by reducing the amount of the virus in blood. It is used by adults, adolescents and children 3 months of age and older and weighing at least 3.5 kg.

Your doctor has prescribed Efavirenz Mylan for you because you have HIV infection. Efavirenz Mylan taken in combination with other antiretroviral medicines reduces the amount of the virus in the blood. This will strengthen your immune system and reduce the risk of developing illnesses linked to HIV infection.

2. What you need to know before you take Efavirenz Mylan

Do not take Efavirenz Mylan:

- if you are allergic to efavirenz or any of the other ingredients of this medicine (listed in section 6). Contact your doctor or pharmacist for advice.
- if you have severe liver disease.
- if you are currently taking any of the following medicines:
 - astemizole or terfenadine (used to treat allergy symptoms)
 - bepridil (used to treat heart disease)
 - cisapride (used to treat heartburn)
 - ergot alkaloids (for example, ergotamine, dihydroergotamine, ergonovine, and methylergonovine) (used to treat migraine and cluster headaches)
 - midazolam or triazolam (used to help you sleep)
 - pimozone (used to treat certain mental conditions)
 - St. John's wort (*Hypericum perforatum*) (a herbal remedy used for depression and anxiety)

If you are taking any of these medicines, tell your doctor immediately. Taking these medicines with Efavirenz Mylan could create the potential for serious and/or life-threatening side-effects or stop Efavirenz Mylan from working properly.

Warnings and precautions

Talk to your doctor or pharmacist before taking Efavirenz Mylan

- Efavirenz Mylan must be taken with other medicines that act against the HIV virus. If Efavirenz Mylan is started because your current treatment has not prevented the virus from multiplying, another medicine you have not taken before must be started at the same time.
- You can still pass on HIV when taking this medicine, although the risk is lowered by effective antiretroviral therapy. Discuss with your doctor the precautions needed to avoid infecting other people. This medicine is not a cure for HIV infection and you may continue to develop infections or other illnesses associated with HIV disease.
- You must remain under the care of your doctor while taking Efavirenz Mylan.

Tell your doctor:

- if you have a history of mental illness, including depression, or of substance or alcohol abuse. Tell your doctor immediately if you feel depressed, have suicidal thoughts or have strange thoughts (see section 4, Possible side effects).
- if you have a history of convulsions (fits or seizures) or if you are being treated with anticonvulsant therapy such as carbamazepine, phenobarbital and phenytoin. If you are taking any of these medicines, your doctor may need to check the level of anticonvulsant medicine in your blood to ensure that it is not affected while taking Efavirenz Mylan. Your doctor may give you a different anticonvulsant.
- if you have a history of liver disease, including active chronic hepatitis. Patients with chronic hepatitis B or C and treated with combination antiretroviral agents have a higher risk for severe and potentially life-threatening liver problems. Your doctor may conduct blood tests in order to check how well your liver is working or may switch you to another medicine. If you have severe liver disease, do not take Efavirenz Mylan (see section 2, “Do not take Efavirenz Mylan”).

Once you start taking Efavirenz Mylan, look out for:

- signs of dizziness, difficulty sleeping, drowsiness, difficulty concentrating or abnormal dreaming. These side effects may start in the first 1 or 2 days of treatment and usually go away after the first 2 to 4 weeks.
- any signs of skin rash. If you see any signs of a severe rash with blistering or fever, stop taking Efavirenz Mylan and tell your doctor at once. If you had a rash while taking another NNRTI, you may be at a higher risk of getting a rash with Efavirenz Mylan.
- any signs of inflammation or infection. In some patients with advanced HIV infection (AIDS) and a history of opportunistic infection, signs and symptoms of inflammation from previous infections may occur soon after anti-HIV treatment is started. It is believed that these symptoms are due to an improvement in the body's immune response, enabling the body to fight infections that may have been present with no obvious symptoms. If you notice any symptoms of infection, please tell your doctor immediately. In addition to the opportunistic 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.

- changes in body fat. Redistribution, accumulation or loss of body fat may occur in patients receiving combination antiretroviral therapy. Tell your doctor if you notice changes in your body fat.
- bone problems. Some patients taking combination antiretroviral therapy may develop a bone disease called osteonecrosis (death of bone tissue caused by loss of blood supply to the bone). The length of combination antiretroviral therapy, corticosteroid use, alcohol consumption, severe immunosuppression, higher body mass index, among others, may be some of the many risk factors for developing this disease. Signs of osteonecrosis are joint stiffness, aches and pains (especially of the hip, knee and shoulder) and difficulty in movement. If you notice any of these symptoms please inform your doctor.

Children and adolescents

Efavirenz Mylan is not recommended for children under the age of 3 months or weighing less than 3.5 kg because it has not been adequately studied in these patients.

Other medicines and Efavirenz Mylan

You must not take Efavirenz Mylan with certain medicines. These are listed under “Do not take Efavirenz Mylan”, at the start of section 2. They include some common medicines and a herbal remedy (St. John's wort) which can cause serious interactions.

Tell your doctor, pharmacist or nurse if you are taking, have recently taken or might take any other medicines.

Efavirenz Mylan may interact with other medicines, including herbal preparations such as Ginkgo biloba extracts. As a result, the amounts of efavirenz or other medicines in your blood may be affected. This may stop the medicines from working properly, or may make any side effects worse. In some cases, your doctor may need to adjust your dose or check your blood levels. It is important to tell your doctor or pharmacist if you are taking any of the following:

- Other medicines used for HIV infection:
 - protease inhibitors: darunavir, indinavir, lopinavir/ritonavir, ritonavir, ritonavir boosted atazanavir, saquinavir or fosamprenavir/saquinavir. Your doctor may consider giving you an alternative medicine or changing the dose of the protease inhibitors.
 - maraviroc
 - the combination tablet containing efavirenz, emtricitabine and tenofovir, which is currently known as Atripla. Efavirenz Mylan should not be taken with Atripla unless recommended by your doctor since it contains efavirenz, the active ingredient of Efavirenz Mylan.
- Medicines used to treat infection with the hepatitis C virus: boceprevir, telaprevir, simeprevir.
- Medicines used to treat bacterial infections, including tuberculosis and AIDS-related mycobacterium avium complex: clarithromycin, rifabutin, rifampicin. Your doctor may consider changing your dose or giving you an alternative antibiotic. In addition, your doctor may prescribe a higher dose of Efavirenz Mylan.
- Medicines used to treat fungal infections (antifungals):
 - voriconazole. Efavirenz may reduce the amount of voriconazole in your blood and voriconazole may increase the amount of efavirenz in your blood. If you take these two medicines together, the dose of voriconazole must be increased and the dose of efavirenz must be reduced. You must check with your doctor first.
 - itraconazole. Efavirenz may reduce the amount of itraconazole in your blood.
 - posaconazole. Efavirenz may reduce the amount of posaconazole in your blood.

- Medicines used to treat malaria:
 - artemether/lumefantrine: Efavirenz may reduce the amount of artemether/lumefantrine in your blood
 - atovaquone/proguanil: Efavirenz may reduce the amount of atovaquone/proguanil in your blood.
- Medicines used to treat convulsions/seizures (anticonvulsants): carbamazepine, phenytoin, phenobarbital. Efavirenz can reduce or increase the amount of anticonvulsant in your blood. Carbamazepine may make efavirenz less likely to work. Your doctor may need to consider giving you a different anticonvulsant.
- Medicines used to lower blood fats (also called statins): atorvastatin, pravastatin, simvastatin. Efavirenz can reduce the amount of statins in your blood. Your doctor will check your cholesterol levels and will consider changing the dose of your statin, if needed.
- Methadone (a medicine used to treat opiate addiction): your doctor may need to change your dose of methadone.
- Sertraline (a medicine used to treat depression): your doctor may need to change your dose of sertraline.
- Bupropion (a medicine used to treat depression or to help you stop smoking): your doctor may need to change your dose of bupropion.
- Diltiazem or similar medicines (called calcium channel blockers, which are medicines typically used for high blood pressure or heart problems): when you start taking Efavirenz Mylan, your doctor may need to adjust your dose of the calcium channel blocker.
- Immunosuppressants such as cyclosporine, sirolimus, or tacrolimus (medicines used to prevent organ transplant rejection): when you start or stop taking Efavirenz Mylan, your doctor will closely monitor your plasma levels of the immunosuppressant and may need to adjust its dose.
- Hormonal contraceptive, such as birth control pills, an injected contraceptive (for example, Depo-Provera), or a contraceptive implant (for example, Implanon): you must also use a reliable barrier method of contraception (see “Pregnancy, breast-feeding and fertility”). Efavirenz may make hormonal contraceptives less likely to work. Pregnancies have occurred in women taking efavirenz while using a contraceptive implant, although it has not been established that the efavirenz therapy caused the contraceptive to fail.
- Warfarin or acenocoumarol (medicines used to reduce clotting of the blood): your doctor may need to adjust your dose of warfarin or acenocoumarol.
- Ginkgo biloba extracts (a herbal preparation)

Taking Efavirenz Mylan with food and drink

Taking Efavirenz Mylan on an empty stomach, may reduce the undesirable effects. Grapefruit juice should be avoided when taking this medicine.

Pregnancy and breast-feeding

Women should not get pregnant during treatment with Efavirenz Mylan and for 12 weeks thereafter. Your doctor may require you to take a pregnancy test to ensure you are not pregnant before starting treatment with Efavirenz Mylan.

Pregnancy

If you could get pregnant while receiving Efavirenz Mylan, you need to use a reliable form of barrier contraception (for example, a condom) with other methods of contraception including oral (pill) or other hormonal contraceptives (for example, implants, injection). Efavirenz may remain in your blood for a time after therapy is stopped. Therefore, you should continue to use contraceptive measures, as above, for 12 weeks after you stop taking Efavirenz Mylan.

Tell your doctor immediately if you are pregnant or intend to become pregnant. If you are pregnant, you should take Efavirenz Mylan only if you and your doctor decide it is clearly needed. Ask your doctor or pharmacist for advice before taking any medicine.

Breast-feeding

Serious birth defects have been seen in unborn animals and in the babies of women treated with efavirenz or a combination medicine containing efavirenz, emtricitabine and tenofovir during pregnancy. If you have taken efavirenz or the combination tablet containing efavirenz, emtricitabine, and tenofovir during your pregnancy, your doctor may request regular blood tests and other diagnostic tests to monitor the development of your child.

You should not breast feed your baby if you are taking Efavirenz Mylan.

Driving and using machines

Efavirenz may cause dizziness, impaired concentration, and drowsiness. If you are affected, do not drive and do not use any tools or machines.

Efavirenz Mylan contains lactose

If you have been told by your doctor that you have an intolerance to some sugars contact your doctor before taking this medicine.

3. How to take Efavirenz Mylan

Always take this medicine exactly as your doctor or pharmacist has told you. It is recommended that the tablet be swallowed whole with water. Check with your doctor or pharmacist if you are not sure. Your doctor will give you instructions for proper dosing.

- The recommended dose for adults is 600 mg once daily.
- The dose for Efavirenz Mylan may need to be increased or decreased if you are also taking certain medicines (see section 2, “Other medicines and Efavirenz Mylan”).
- Efavirenz Mylan is for oral use. Efavirenz Mylan is recommended to be taken on an empty stomach preferably at bedtime. This may make some side effects (for example, dizziness, drowsiness) less troublesome. An empty stomach is commonly defined as 1 hour before or 2 hours after a meal.
- It is recommended that the tablet be swallowed whole with water. Efavirenz Mylan must be taken every day.
- Efavirenz Mylan should never be used alone to treat HIV.
- Efavirenz Mylan must always be taken in combination with other anti-HIV medicines.

Use in children and adolescents

- Efavirenz Mylan film-coated tablets are not suitable for children weighing less than 40 kg.
- The dose for children weighing 40 kg or more is 600 mg once daily.

Other formulations of efavirenz are available for patients who need some dose adjustment.

If you take more Efavirenz Mylan than you should:

If you take too much Efavirenz Mylan, contact your doctor or nearest emergency department for advice. Keep the medicine container with you so that you can easily describe what you have taken.

If you forget to take Efavirenz Mylan:

Try not to miss a dose. If you do miss a dose, take the next dose as soon as possible, but do not take a double dose to make up for a forgotten tablet. If you need help in planning the best times to take your medicine, ask your doctor or pharmacist.

If you stop taking Efavirenz Mylan:

When your Efavirenz Mylan supply starts to run low, get more from your doctor or pharmacist. This is very important because the amount of virus may start to increase if the medicine is stopped for even a short time. The virus may then become harder to treat.

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

4. Possible side effects

Like all medicines, this medicine can cause side effects, although not everybody gets them. When treating HIV infection, it is not always possible to tell whether some of the unwanted effects are caused by efavirenz or by other medicines that you are taking at the same time, or by the HIV disease itself.

During HIV therapy there may be an increase in weight and in levels of blood lipids and glucose. This is partly linked to restored health and lifestyle, and in the case of blood lipids sometimes to the HIV medicines themselves. Your doctor will test for these changes.

The most notable unwanted effects reported with efavirenz in combination with other anti-HIV medicines are skin rash and nervous system symptoms.

You should consult your doctor if you have a rash, since some rashes may be serious; however, most cases of rash disappear without any change to your treatment with efavirenz. Rash was more common in children than in adults treated with efavirenz.

The nervous system symptoms tend to occur when treatment is first started, but generally decrease in the first few weeks. In one study, nervous system symptoms often occurred during the first 1-3 hours after taking a dose. If you are affected your doctor may suggest that you take Efavirenz Mylan at bedtime and on an empty stomach. Some patients have more serious symptoms that may affect mood or the ability to think clearly. Some patients have actually committed suicide. These problems tend to occur more often in those who have a history of mental illness. Always notify your doctor immediately if you have these symptoms or any side effects while taking Efavirenz Mylan.

If you notice any of the following side effects, contact your doctor or go to the nearest hospital casualty department straight away:

Uncommon (may affect up to 1 in 100 people)

- serious allergic reaction that may cause shortness of breath, wheezing or difficulty breathing; swelling of the face, lips, tongue or other parts of the body; rash, itching or hives on the skin
- severe skin reactions which may cause red (moist), itchy and irregular spots similar to the rash of measles or widespread rash with blisters and peeling skin, particularly around the mouth, nose, eyes and genitals. The spots may progress to form raised, red, pale-centered marks. Those affected may have fever, sore throat, headache and/or diarrhoea (erythema multiforme, Stevens-Johnson syndrome)
- pain in the abdomen (stomach) which spreads to the back caused by inflammation of the pancreas
- suicidal thoughts or attempting to kill yourself

Rare (may affect up to 1 in 1,000 people)

- liver failure, in some cases leading to death or liver transplant, has occurred with efavirenz. Most cases occurred in patients who already had liver disease, but there have been a few reports in patients without any existing liver disease
- suicide

Other possible side effects

Very common (may affect more than 1 in 10 people)

- skin rash

Common (may affect up to 1 in 10 people)

- abnormal dreams, difficulty concentrating, dizziness, headache, difficulty sleeping, drowsiness, problems with coordination or balance
- stomach pain, diarrhoea, feeling sick (nausea), being sick (vomiting)
- itching
- tiredness
- feeling worried, feeling depressed

Tests may show:

- increased liver enzymes in the blood
- increased triglycerides (fatty acids) in the blood

Uncommon (may affect up to 1 in 100 people)

- nervousness, forgetfulness, confusion, fitting (seizures), abnormal thoughts
- blurred vision
- a feeling of spinning or tilting (vertigo)
- yellow skin or eyes, itching, or pain in the abdomen (stomach) caused by inflammation of the liver
- breast enlargement in males
- angry behaviour, feeling of extreme happiness or contentment, seeing or hearing things that are not really there (hallucinations), mania (mental condition characterised by episodes of overactivity, elation or irritability), paranoia
- severe mental condition in which the person loses contact with reality and is unable to think and judge clearly
 - whistling, ringing or other persistent noise in the ears
 - tremor (shaking)
 - flushing.

Tests may show:

- increased cholesterol in the blood

Rare (may affect up to 1 in in 1, 000 people)

- itchy rash caused by a reaction to sunlight
- unexplained feelings of distress not associated with hallucinations, but it may be difficult to think clearly or sensibly
- fixed, irrational idea not shared by others

Not known (frequency cannot be estimated from the available data)

- 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 (autoimmune disorders)
- combination antiretroviral therapy may change your body shape, by changing the way body fat is distributed. You may lose fat from your legs, arms and face, gain fat around the

abdomen (tummy) and other internal organs, get larger breasts or fatty lumps on the back of the neck ('buffalo hump'). The cause and long-term health effects of these conditions are not yet known. combination antiretroviral therapy may develop death of bone tissue caused by loss of blood supply to the bone (osteonecrosis). You may experience joint stiffness, aches and pains (especially of the hip, knee and shoulder) and difficulty in movement. If you notice any of these symptoms please inform your doctor.

- combination antiretroviral therapy may also cause raised lactic acid and sugar in the blood, increased fat levels in the blood (hyperlipaemia) and resistance to insulin. Your doctor will test for these changes.

Reporting of side effects

If you get any side effects, talk to your doctor, pharmacist or nurse. This includes any possible side effects not listed in this leaflet. You can also report side effects directly via HPRAs Pharmacovigilance, Earlsfort Terrace, IRL – Dublin 2; Tel: +353 1 6764971; Fax: +353 1 6762517. Website: www.hpra.ie; E-mail: medsafety@hpra.ie. By reporting side effects you can help provide more information on the safety of this medicine.

5. How to store Efavirenz Mylan

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

Do not use this medicine after the expiry date, which is stated on the blister or bottle after EXP. The expiry date refers to the last day of that month.

This medicinal product does not require any special storage conditions.

The in-use shelf-life of the product when stored in HDPE bottles is 100 days.

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. Contents of the pack and other information

What Efavirenz Mylan contains

The active substance is efavirenz.

Each film-coated tablet contains 600 mg of efavirenz.

The other ingredients are cellulose microcrystalline, croscarmellose sodium, hydroxypropyl cellulose, lactose monohydrate (see section 2 “Efavirenz Mylan contains lactose”), silica colloidal anhydrous, magnesium stearate, hypromellose (E464), macrogol 400, titanium dioxide (E171), iron oxide yellow (E172) and iron oxide red (E172).

What Efavirenz Mylan looks like and contents of the pack

Efavirenz Mylan tablets are yellow, capsule shaped biconvex bevelled edge film-coated tablets marked with “M” on one side and “EV6” on the other side.

Efavirenz Mylan is available in blisters of 30, 60, 90, 100 film-coated tablets, perforated unit dose blisters of 30 x 1 film-coated tablets, and in plastic bottles of 30, 500 film-coated tablets.

Not all pack sizes may be marketed.

Marketing Authorisation Holder

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This medicinal product is authorised in the Member States of the EEA under the following names:

BE: Efavirenz Mylan 600 mg Filmomhulde Tabletten
ES: Efavirenz Mylan 600 mg Comprimidos Recubiertos Con Pelicula EFG
FR: EFAVIRENZ MYLAN 600 mg, comprimé pelliculé
IE: Efavirenz Mylan 600 mg Film-coated Tablets
IT: Efavirenz Mylan
NL: Efavirenz Mylan 600 mg, filmomhulde tabletten
PL: Efavirenz Mylan
PT: Efavirenz Mylan
UK: Efavirenz 600 mg Film-coated Tablets

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