PATIENT LEAFLET IN ACCORDANCE WITH THE PHARMACISTS' REGULATIONS (PREPARATIONS) – 1986

The medicine is dispensed with a doctor's prescription only

Teriflunomide Teva

Film-coated tablets

Composition

Each tablet contains:

Teriflunomide 14 mg

For information regarding inactive ingredients and allergens, see section 2 - 'Important information about some ingredients of the medicine' and section 6 - 'Additional information'.

Read the entire leaflet carefully before using the medicine.

This leaflet contains concise information about the medicine. If you have additional questions, refer to the doctor or the pharmacist.

This medicine has been prescribed for you only, do not pass it on to others. It may harm them even if it seems to you that their medical condition is similar.

Patient safety information card

In addition to the leaflet, there is a patient safety information card for Teriflunomide Teva. This card contains important safety information that you must know before starting treatment with Teriflunomide Teva and during the treatment, and act accordingly. Please review the patient safety information card and the patient leaflet before using the medicine. You should keep the card for further review, if necessary.

1. What is the medicine intended for?

Teriflunomide Teva is intended for the treatment of adults with relapsing-remitting multiple sclerosis in order to reduce the frequency of relapses and to delay the progression of disability.

Therapeutic group:

medicines that suppress the immune system (immunosuppressants), inhibitors of the enzyme dihydroorotate dehydrogenase.

Teriflunomide Teva contains the active substance teriflunomide which is a substance that affects the immune system by limiting its attack on the nervous system.

What is multiple sclerosis?

Multiple Sclerosis (MS) is a chronic illness that affects the central nervous system (CNS). The central nervous system is composed of the brain and the spinal cord.

In multiple sclerosis, inflammation destroys the protecting cover (called Myelin) around nerve fibers in the central nervous system. Loss of Myelin is called demyelination and it impairs the normal function of the nerves.

People with relapsing multiple sclerosis will have recurrent attacks of physical symptoms caused by improper functioning of the nerves.

These symptoms differ between patients, but usually include:

- Difficulty walking.
- Vision problems.
- Balance disorders.

Symptoms may completely resolve after the attack is over, but over time, certain problems may remain between attacks, and can cause physical disabilities which may impede your daily activities.

How does Teriflunomide Teva work?

Teriflunomide Teva helps to protect the central nervous system from attacks of the immune system by limiting the increase in the level of some white blood cells (lymphocytes). This reduces the inflammation that causes neural damage in multiple sclerosis.

2. Before using the medicine

Do not use this medicine if:

- You are sensitive (allergic) to the active ingredient, teriflunomide, or any of the other ingredients this medicine contains (see section 6 – 'Additional information').
- You have ever developed a severe skin rash or skin peeling, blistering and/or mouth sores after taking teriflunomide or leflunomide.
- You have serious liver problems.
- You are pregnant, think you may be pregnant or are breastfeeding.
- You have a serious problem that affects your immune system, for example acquired immunodeficiency syndrome (AIDS).
- You have a serious problem in your bone marrow, or if you have a significant decline in red or white blood cells levels or a decline in platelets levels.
- You have a serious infection.
- You have serious kidney problems that require dialysis.
- You have very low protein levels in the blood (hypoproteinemia).

If you are unsure, or if you have any questions about the use of this medicine, speak with your doctor or the pharmacist before taking this medicine.

Special warnings regarding the use of the medicine

Before taking Teriflunomide Teva, tell your doctor if:

- You have liver problems and/or if you drink large amounts of alcohol; your doctor will order blood tests before and during the treatment in order to check liver function. If the test results will show that there is a liver problem, your doctor may stop the treatment with Teriflunomide Teva. Please read section 4: 'Side effects'.
- You have high blood pressure (hypertension), whether controlled with medicines or not. Teriflunomide Teva can cause a rise in blood pressure. Your doctor will check your blood pressure before you start treatment and regularly during the treatment. Please read section 4: 'Side effects'.
- You have an infection. Before taking Teriflunomide Teva, your doctor will make sure you have enough white blood cells and platelets in your blood. This is because Teriflunomide Teva decreases the number of white blood cells, which may affect your ability to fight the infection. If you think you have any infection, your doctor can order blood tests to check the levels of white blood cells. Herpes virus infections, including oral herpes or Herpes zoster (shingles) may occur with teriflunomide treatment. In some cases, serious complications have occurred. You should inform your doctor immediately if you suspect you have any symptoms of herpes virus infections. Please read section 4: 'Side effects'.
- You have serious skin reactions.
- You have respiratory symptoms.
- You feel weakness, numbness, and pain in your hands and feet.
- You are about to be vaccinated.
- You are taking leflunomide with Teriflunomide Teva.
- You are switching to Teriflunomide Teva treatment or from Teriflunomide Teva treatment.
- You are about to have a specific blood test (calcium levels). The test results may falsely indicate low calcium levels.

Respiratory reactions:

Tell your doctor if you have unexplained cough and shortness of breath. Your doctor may perform additional tests.

Children and adolescents

Teriflunomide Teva is not intended for children and adolescents under 18 years of age.

Tests and follow-up

Before starting this medicine, your doctor will refer you to blood tests and (for women) a pregnancy test.

Drug interactions

If you are taking or have recently taken other medicines, including non-prescription medicines and food supplements, tell the doctor or the pharmacist. Particularly if you are taking any of the following medicines:

- Leflunomide, methotrexate and other medicines that affect the immune system (usually called immunosuppressants or immunomodulators).
- Rifampicin (a medicine used for the treatment of tuberculosis and other infections).
- Carbamazepine, phenobarbital, phenytoin (for treatment of epilepsy).
- Hypericum perforatum (St. John's Wort - a medicinal herb for treatment of depression).
- Repaglinide, pioglitazone, nateglinide or rosiglitazone for treatment of diabetes.
- Daunorubicin, doxorubicin, paclitaxel or topotecan for treatment of cancer.
- Duloxetine for treatment of depression, urinary incontinence or kidney disease in diabetics.
- Alosetron for treatment of acute diarrhea.
- Theophylline for asthma.
- Tizanidine, a muscle relaxant.
- Warfarin, an anticoagulant used for blood thinning (to make it more liquid) in order to prevent blood clots.
- Contraceptive pills (containing ethinylestradiol and levonorgestrel).
- Cefaclor, benzylpenicillin (penicillin G), ciprofloxacin for infections.
- Indomethacin, ketoprofen for pain and inflammation.
- Furosemide for heart disease.
- Cimetidine for decreasing gastric acidity.
- Zidovudine for HIV infection.
- Rosuvastatin, simvastatin, atorvastatin, pravastatin for hypercholesterolemia (high cholesterol).
- Sulfasalazine for inflammatory bowel disease or for arthritis.
- Cholestyramine for high cholesterol or for relief of itching in liver disease.
- Activated charcoal for reduction of absorption of medicines or other substances.

Use of the medicine and food

The medicine may be taken with or without food.

Pregnancy and breastfeeding

Pregnancy

Do not take Teriflunomide Teva if you are pregnant or think you might be pregnant. If you are pregnant or have become pregnant while taking Teriflunomide Teva, the risk of congenital malformations in the baby is increased. Women of childbearing age should not take this medicine without using reliable contraceptives.

Refer to the doctor if you are planning to become pregnant after discontinuing treatment with Teriflunomide Teva, because you need to make sure that most of the active ingredient in this medicine has been eliminated from the body before you try to conceive. Natural elimination of the active ingredient from the body may take two years. This period of time can be reduced to several weeks by taking certain medicines that accelerate the elimination of Teriflunomide Teva from the body. In any case, a blood test is required to confirm that a sufficient amount of the active ingredient has been eliminated from your body, and to receive confirmation from your doctor that your blood Teriflunomide Teva level is sufficiently low to allow you to become pregnant.

For more information about your lab tests, refer to your doctor.

If you suspect that you might be pregnant while taking Teriflunomide Teva, or during the two years following discontinuation of the treatment, you must stop taking Teriflunomide Teva and contact your doctor **immediately** to undergo a pregnancy test. If the test confirms that you are pregnant, your doctor may suggest treatment with certain medicines in order to eliminate teriflunomide quickly and sufficiently from your body, as this may reduce the risk to your baby.

Contraceptives

You must use efficient contraceptives during and following treatment with Teriflunomide Teva. Teriflunomide remains in your blood for a long time after you stop taking the medicine.

Continue to use efficient contraceptives after stopping the treatment.

- Keep doing this until teriflunomide levels in your blood will be low enough - your doctor will check this.
- Consult the doctor about the best contraception method for you and about any possible need to change the contraception method.

Breastfeeding

Do not take Teriflunomide Teva while you are breastfeeding, since teriflunomide passes into breastmilk.

Driving and operating machinery

Teriflunomide Teva may cause dizziness, which may impair your ability to concentrate and react. Therefore, if you are feeling dizzy, do not drive a vehicle or operate machinery during the treatment.

Important information about some ingredients of the medicine

Teriflunomide Teva contains a sugar called lactose. If you have been told by your doctor that you have an intolerance to certain sugars, consult your doctor before taking this medicine.

This medicine contains less than 23 mg of sodium in a tablet and is therefore considered sodium-free.

3. How should you use the medicine?

Treatment with Teriflunomide Teva should be done under supervision of a doctor who is experienced in treating multiple sclerosis.

Always use the preparation according to the doctor's instructions.

Check with the doctor or pharmacist if you are uncertain about the dosage and how to use the preparation.

Dosage

The dosage and treatment regimen will be determined only by the doctor.

The standard dosage is usually one film-coated tablet (14 mg) per day.

Do not exceed the recommended dose.

Method of use

- Teriflunomide Teva is a medicine that should be taken orally, one tablet a day, at any time of the day.
- The tablet may be taken with or without food.
- Do not chew! The tablet should be swallowed whole with water.
- The tablets should not be halved when there is no score line.

If you accidentally take a higher dosage

If you took an overdose or if a child swallowed this medicine by mistake, go to the doctor or the emergency room of the hospital immediately and take the package of the medicine with you. You may experience side effects like the ones described in section 4: 'Side effects'.

If you have forgotten to take the medicine

If you forgot to take this medicine at the required time, do not take a double dose. Take the next dose at the scheduled time and consult a doctor.

Follow the treatment as recommended by the doctor.

Even if there is an improvement in your health, do not stop treatment with the medicine without consulting the doctor.

If you stop taking the medicine

Do not stop taking Teriflunomide Teva or change the dosage without consulting your doctor first.

Do not take medicines in the dark! Check the label and the dose every time you take the medicine. Wear glasses if you need them.

If you have any other questions regarding use of the medicine, consult the doctor or the pharmacist.

4. Side effects

As with any medicine, using Teriflunomide Teva may cause side effects in some users. Do not be alarmed when reading the list of side effects. You may not experience any of them.

Severe side effects

Some side effects could be or could become severe, **contact your doctor immediately** if you observe any of the following severe side effects:

Uncommon (may affect up to 1 in 100 people)

- Inflammation of the pancreas that can include symptoms of pain in the abdominal area, nausea, or vomiting.
- Allergic reactions that can include symptoms of rash, hives, swelling of the lips, tongue or face, or a sudden breathing difficulty.
- Severe skin reactions that can include symptoms of skin rash, blisters, fever or mouth sores.
- Severe infections or sepsis (a type of infection which may be life-threatening) that can include symptoms of high fever, tremors, chills, decreased urinary flow, or confusion.
- Inflammation of the lungs that can include symptoms of shortness of breath or prolonged cough.

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

- A severe liver disease that can include symptoms of yellowing of the skin or the white part of the eyes, darker urine than usual, unexplained nausea and vomiting or abdominal pain.

Additional side effects may occur with the following frequencies:

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

- Headache
- Diarrhea, nausea
- Rise in ALT values (rise in blood levels of certain liver enzymes) that is observed in blood tests
- Hair thinning

Common side effects (may affect up to 1 in 10 people):

- Influenza, upper respiratory tract infection, urinary tract infection, bronchitis, sinusitis, sore throat and discomfort while swallowing, cystitis, viral gastroenteritis, infection of the teeth (pulpitis), pharyngitis, fungal foot infection
- Herpes virus infections, including oral herpes and herpes zoster (shingles), with symptoms such as blisters, burning sensation, itching, numbness or pain of the skin, typically on one side of the upper body or face, and other symptoms like fever and weakness.
- Lab values: decline in red blood cells count (anemia), changes in the results of liver function tests and white blood cells count (see section 2), as well as a rise in muscle enzyme (creatine phosphokinase) have been observed
- Moderate allergic reactions
- Feeling of anxiety
- Feeling of numbness (as needle pricks), feeling of weakness, loss of sensation, tingling or pain in the lower back or leg; loss of sensation, burning, tingling or pain in the palms and fingers (carpal tunnel syndrome)
- Sensation of heart beats
- Rise in blood pressure
- Nausea (vomiting), toothaches, upper abdominal pain
- Rash, acne
- Tendon, joint, bone, muscle pain (musculoskeletal pain)
- Needing to urinate more frequent than usual
- Menstrual cycles with massive bleeding
- Pain
- Lack of energy or feeling weak
- Weight loss

Uncommon side effects (may affect up to 1 in 100 people):

- Decline in the number of platelets (thrombocytopenia)
- Enhanced sensation or hypersensitivity, especially in the skin, pricking or throbbing pain along one or more neural pathways, problems in the nerves of the arms or legs (peripheral neuropathy)
- Nail problems, severe skin reactions
- Post-traumatic pain
- Psoriasis
- Inflammation of mouth or lips
- Abnormal levels of fats (lipids) in the blood
- Inflammation of the intestine (colitis)

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

- Inflammation or injury of the liver

Side effects with unknown frequency (frequency cannot be estimated from the available data):

- Respiratory hypertension

If a side effect occurs, if one of the side effects worsens, or if you suffer from a side effect not mentioned in this leaflet, consult your doctor.

Reporting side effects

Side effects may be reported to the Ministry of Health by clicking on the link 'report side effects due to medicinal treatment' found on the Ministry of Health website homepage (www.health.gov.il), which will direct you to the online form for reporting side effects, or by clicking on the following link: https://sideeffects.health.gov.il

5. How to store the medicine?

- Avoid poisoning! This medicine and any other medicine must be kept in a closed place out of the reach and sight of children and/or infants to avoid poisoning. Do not induce vomiting without an explicit instruction from the doctor.
- Do not use the medicine after the expiry date (Exp. Date) appearing on the package. The expiry date refers to the last day of that month.
- **Store in a dry place, under 25°C.**
- When using the bottle, after opening the bottle for the first time, the medicine can be used up to the expiry date.
- Do not dispose of any medicine in the sewage or household waste. Ask your pharmacist how to discard medicines you no longer use. These measures will help to protect the environment.

Note: The bottle contains a desiccant. Do not swallow! The desiccant should be left inside the bottle, and the bottle should be closed tightly after each use!

6. Additional information

In addition to the active ingredient, the medicine also contains:

Lactose monohydrate, starch, microcrystalline cellulose, sodium starch glycolate, sodium stearyl fumarate, hypromellose, hydroxypropyl cellulose, titanium dioxide, talc, polyethylene glycol, colloidal silicone dioxide, FD&C Blue#2 / indigo carmine aluminum lake

What does the medicine look like and what are the contents of the package:

A blue, round, film-coated tablet, embossed on one side with "TV" and on the other side with "Y12".

The pack contains 28 or 30 film-coated tablets.

Not all package sizes may be marketed.

Name and address of registration holder and manufacturer:

Teva Israel Ltd.,

124 Dvorah HaNevi'a St., Tel Aviv 6944020.

Contact for queries: 03-6864212

The leaflet was revised in April 2025.

This leaflet does not contain all of the information about the preparation. If you have any question or if you are unsure about anything, please contact the doctor.

Registration number of the medicine in the National Drug Registry of the Ministry of Health: 161.57.35259