

This medicine can be sold with a physician's prescription only

Clopidogrel® 75, film coated tablets

Each tablet contains clopidogrel as besylate 75 mg

Inactive ingredients and allergens in the medicine – see section 6 "Additional information" and in section 2 "Important information about some of the ingredients of this medicine".

Read this entire leaflet carefully before using this medicine. This leaflet contains concise information about the medicine. If you have any further questions, ask the doctor or pharmacist. This medicine has been prescribed for you. Do not pass it on to others. It may harm them, even if you think that their medical condition is the same as yours.

1. What is the medicine intended for?

The medicine is used for the secondary prevention of atherothrombotic events within blood vessels (heart attacks, stroke or vascular death) in patients with atherosclerosis, who experienced myocardial infarction, ischemic stroke or are suffering from peripheral arterial disease, and in patients suffering from unstable angina pectoris, including patients undergoing implant of a stent by catheterization and who are taking aspirin (acetylsalicylic acid). The medicine is used for the prevention of atherothrombotic and thromboembolic events (heart attacks, stroke, emboli or vascular death) in combination with aspirin, in patients suffering from atrial fibrillation type arrhythmia, who are not suitable for treatment with anticoagulant of the vitamin K antagonist medications such as warfarin, antithrombin or anti-factor Xa.

Therapeutic group: Platelet aggregation inhibitors of the thienopyridines group.

2. Before using the medicine

Do not use the medicine if:

- you are hypersensitive (allergic) to the active ingredient (clopidogrel) or to any of the other ingredients this medicine contains (see section 6).
 - you suffer from a medical condition that currently causes bleeding, such as: stomach or duodenal ulcer, intracerebral hemorrhage.
 - you suffer from severe liver disease.
- If any of these apply to you, or if you have any doubt, consult the doctor before taking Clopidogrel 75.

Special warnings regarding the use of the medicine

Before treatment with Clopidogrel 75, tell the doctor if:

- You have a risk of bleeding, such as:
 - A medical condition that puts you at risk of internal bleeding (such as stomach ulcer).
 - A problem in the blood system that makes you prone to internal bleeding (bleeding inside tissues, organs or joints of your body).
 - You have recently had a serious injury.
 - You have recently undergone surgery (including dental surgeries).
 - You are about to undergo surgery (including dental surgeries) in the next seven days.
- You have suffered from a clot in an artery in your brain (ischemic stroke) in the last seven days.
- You suffer from kidney or liver disease.
- You have previously suffered from an allergy or

reaction to any medicine that was used to treat your disease.

During treatment with the medicine:

- You must inform your doctor if you have a surgery planned (including dental surgery).
- You must also inform your doctor immediately if you have developed a medical condition (also known as Thrombotic Thrombocytopenic Purpura or TTP) which includes fever and subcutaneous bruising that may appear as red pinpoint dots, with or without unexplained extreme tiredness, confusion, yellowing of the skin or eyes (jaundice) (see section 4 "Side effects").
- If you were cut or injured, it may take longer than usual until the bleeding stops. This relates to the way your medicine works as it prevents the formation of blood clots. As for cuts or minor injuries, such as a cut during shaving, usually there is no problem. However, if you are concerned by the bleeding, you should refer to your doctor immediately (see section 4 "Side effects").
- The doctor may ask you to perform blood tests.

Children and adolescents

The medicine is not intended for children or adolescents under the age of 18 years.

Drug interactions

If you are taking or have recently taken any other medicines, including non-prescription medicines and nutritional supplements, tell the doctor or pharmacist.

Certain other medicines may affect the use of Clopidogrel 75 or vice versa.

Especially inform the doctor if you are taking:

- Medicines that may increase your risk of bleeding such as:
 - Oral anticoagulants, medicines used to reduce blood clotting.
 - Non-steroidal anti-inflammatories (NSAIDs), which are usually used to treat pain and/or inflammation of muscles or joints.
 - Heparin or any other injectable medicine, used to reduce blood clotting.
 - Ticlopidine, another antiplatelet medicine.
 - Selective serotonin reuptake inhibitors (SSRIs) (including, but not restricted to fluoxetine or fluvoxamine), medicines usually used to treat depression.
 - Rifampicin (used to treat serious infections).
- Omeprazole or esomeprazole, medicines to treat abdominal pain.
- Fluconazole or voriconazole, medicines to treat fungal infections.
- Efavirenz or other anti-retroviral medicines (used to treat HIV infection).
- Carbamazepine, a medicine to treat certain types of epilepsy.
- Moclobemide, a medicine to treat depression.
- Repaglinide, a medicine to treat diabetes.
- Pacitaxel, a medicine to treat cancer.
- Opioids: while you are being treated with clopidogrel, you must inform your doctor before receiving a prescription for any opioid (used to treat severe pains).

If you have experienced serious chest pain, (unstable angina pectoris or a heart attack) you

may be prescribed Clopidogrel 75 in combination with acetylsalicylic acid, a substance found in many medicines, used for pain relief and fever reduction. Occasional use of acetylsalicylic acid (not more than 1,000 mg in any 24-hour time period) should generally not cause problems, but prolonged use in other circumstances should be discussed with your doctor.

Use of this medicine and food

The medicine can be taken with or without food.

Pregnancy, breastfeeding and fertility

It is preferable not to take the medicine during pregnancy.

If you are pregnant, or suspect you are pregnant, you should tell the doctor or pharmacist before taking Clopidogrel 75. If you become pregnant while taking the medicine, consult a doctor immediately as it is not recommended to take clopidogrel when you are pregnant.

Do not breastfeed while taking the medicine.

If you are breastfeeding or planning to breastfeed, consult the doctor before taking the medicine. Consult the doctor or pharmacist before taking any medicine.

Driving and using machines

Clopidogrel 75 is unlikely to affect your ability to drive or operate machines.

Important information about some of the ingredients of this medicine

Clopidogrel 75 contains lactose. If you have been told by a doctor that you have an intolerance to certain sugars, consult the doctor before starting treatment with the medicine.

3. How to use the medicine?

Always use this medicine according to the doctor's instructions. Check with the doctor or pharmacist if you are not sure about the dosage and the manner of treatment with the medicine.

The dosage and manner of treatment will be determined by the doctor only. The usual recommended dosage, including for patients with a condition called "atrial fibrillation" (an irregular heartbeat) is: one tablet of Clopidogrel 75 per day, taken orally, with or without food and at the same time each day.

If you have experienced strong chest pain (unstable angina pectoris or a heart attack) your doctor may instruct you to take a 300 mg dosage (4 tablets of 75 mg) one time only, at the beginning of treatment. Afterwards, the recommended dosage is one Clopidogrel 75 tablet per day, taken as described above.

Do not exceed the recommended dose.

You should take the medicine for as long as the doctor prescribes it for you.

Manner of use

Clopidogrel 75 tablets cannot be halved since there is no score-line. There is no information regarding crushing/chewing. Swallow the medicine with water. The medicine can be taken with or without food.

If you have accidentally taken a higher dosage if you have taken an overdose or if a child has accidentally swallowed the medicine, proceed immediately to a doctor or a hospital emergency room, due to the increased risk of bleeding.

Bring the package of the medicine with you.

If you forgot to take the medicine

If you forgot to take a dose of Clopidogrel 75 but remembered within 12 hours of the usual administration time, take a tablet immediately and take the next tablet at the usual time. If you forgot to take the tablet and more than 12 hours have passed, simply take the next single dose at the usual time. Do not take a double dose to make up for the forgotten dose.

Continue with the treatment as recommended by the doctor.

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

Do not take medicines in the dark! Check the label and the dose each time you take a medicine. Wear glasses if you need them. If you have further questions on the use of this medicine, consult the doctor or pharmacist.

4. Side effects

Like any medicine, the use of Clopidogrel 75 may cause side effects in some users. Do not be alarmed when reading the list of side effects. You may not suffer from any of them.

Consult the doctor immediately if the following signs appear:

- Fever, signs of infection or extreme fatigue. These can be a result of a rare decrease in certain blood cells.
- Signs of a liver problem, such as yellowing of the skin and/or eyes (jaundice), whether or not associated with subcutaneous bleeding, appearing as red pinpoint dots, and/or confusion (see in section 2 "Special warnings regarding the use of the medicine").
- Swelling in the mouth or skin problems such as rash, itching and blisters on the skin. These can be signs of an allergic reaction.

The most common side effect reported with clopidogrel is bleeding. The bleeding may occur as bleeding in the stomach or bowels, bruising, hematoma (abnormal bleeding or subcutaneous bruise), nose bleed, or blood in the urine. In a small number of cases, bleeding in the eye, inside the head, in the lungs or in the joints have also been reported.

If you experience prolonged bleeding during treatment with Clopidogrel 75:

If you were cut or injured, it may take longer than usual until the bleeding stops. This is linked to the way your medicine works as it prevents blood clot formation. As for cuts and minor injuries, such as a cut during shaving, usually there is no problem. However, if you are concerned by the bleeding, you should refer to your doctor immediately (see in section 2 "Special warnings regarding the use of the medicine").

Additional side effects:

Common side effects (effects that appear in 1-10 out of 100 users):

Diarrhea, abdominal pain, indigestion or heartburn.

Uncommon side effects (effects that appear in 1-10 out of 1,000 users):

Headache, stomach ulcer, vomiting, nausea, constipation excessive gas in the stomach or intestines, rash, itching, dizziness, sensation of tingling or numbness.

Rare side effects (effects that appear in 1-10 out of 10,000 users):

Vertigo, enlarged breasts in males.

Very rare side effects (effects that appear in less than one out of 10,000 users):

Jaundice; severe abdominal pain with or without backache; fever, breathing difficulties sometimes associated with cough; generalized allergic reactions (for example, overall sensation of heat with sudden general discomfort until fainting); swelling in the mouth; skin blisters; skin allergy; inflammation in the mucus membranes of the mouth; decrease in blood pressure; confusion; hallucinations; joint pain; muscular pain; changes in the sense of taste or loss of taste of food.

Side effects with unknown frequency (effects for which a frequency has not yet been determined):

Hypersensitivity reactions with chest or abdominal pain, persistent low blood sugar symptoms.

In addition, your doctor may notice changes in your blood or urine tests results.

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

Side effects can be reported to the Ministry of Health via the link

"דיווח על תופעות לוואי עקב תרופות" that can be found on the home page of the Ministry of Health website (www.health.gov.il) directing to the online form of adverse events reporting or via the following link:

<https://sideeffects.health.gov.il>

5. How to store the medicine

• **Avoid poisoning!** This medicine and any other medicine must be stored in a closed place out of the reach and sight of children and/or infants to avoid poisoning. Do not induce vomiting unless explicitly instructed to do so by the doctor.

• Do not use the medicine after the expiry date (תאריך תפוגה) stated on the package. The expiry date refers to the last day of that month.

• **Storage conditions:** do not store above 25°C.

• Do not throw away any medicines via wastewater or household waste. Ask the pharmacist how to throw away medicines you no longer use. These measures will help protect the environment.

6. Additional information

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

Microcrystalline cellulose, spray-dried mannitol, hydroxypropylcellulose, stearic acid, lactose monohydrate, citric acid monohydrate, croscopolone, talc, hypromellose, titanium dioxide (E171), triacetin (E1518), macrogol 6000, iron oxide red (E172).

What the medicine looks like and what the package contains:

Pink coated, round tablet.

Approved package size: 30 tablets.

Manufacturer and address: Dexcel Ltd., 1 Dexcel St., Or Akiva 3060000, Israel

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Registration holder:

Dexcel Pharma Technologies Ltd.

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