

Patient package insert according to Pharmacists' Regulations (Preparations) – 1986

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

CLOPIDEXCEL® 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 the medicine".

Read this entire leaflet carefully before using the 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 had a 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, from 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 is currently causing bleeding, such as: stomach or duodenal ulcer, bleeding within the brain.
- You suffer from severe liver disease.

If any of these apply to you, or if you have any doubt, consult the doctor before taking **Clopidexcel 75**.

Special warnings regarding the use of the medicine

Before the treatment with Clopidexcel 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 blood disorder that makes you prone to internal bleeding (bleeding inside any tissues, organs or joints of your body).
 - You had a recent serious injury.
 - You had a recent 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 bruising under the skin 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 for bleeding to stop. This relates to the way your medicine works as it prevents the formation of blood clots. For minor cuts or 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 to perform blood tests.

Children and adolescents

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

Drug interactions

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

Some other medicines may affect the use of **Clopidexcel 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), usually used to treat pain and/or inflammation of muscles or joints,
 - Heparin or any other injectable medicine, used to reduce blood clotting,
 - Ticlopidine, or other antiplatelet agents,
 - Selective serotonin reuptake inhibitors (SSRIs) (including, but not restricted to fluoxetine or fluvoxamine), medicines usually used to treat depression,
 - Rifampicin (used to treat severe infections)
- Omeprazole or esomeprazole, medicines to treat upset stomach,
- Fluconazole or voriconazole, medicines to treat fungal infections,
- Efavirenz, or other anti-retroviral medicines (used to treat HIV infection),
- Carbamazepine, a medicine to treat some types of epilepsy,
- Moclobemide, a medicine to treat depression,
- Repaglinide, a medicine to treat diabetes,
- Paclitaxel, a medicine to treat cancer,
- Opioids: while you are being treated with clopidogrel, you must inform your doctor before being prescribed any opioid (used to treat severe pains),
- Rosuvastatin (used to lower your cholesterol level).

If you have experienced severe chest pain (unstable angina pectoris or a heart attack), you may be prescribed **Clopidexcel 75** in combination with acetylsalicylic acid, a substance present in many medicines used to relieve pain and lower fever.

Occasional use of acetylsalicylic acid (not more than 1,000 mg in any 24 hour period) should generally not cause problems, but prolonged use in other circumstances should be discussed with your doctor.

Use of the 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 **Clopidexcel 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

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

Important information about some of the ingredients of the medicine

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

3. How to use the medicine?

Always use the 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 **Clopidexcel 75** per day, taken orally, with or without food and at the same time each day.

If you have experienced severe 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 **Clopidexcel 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

Clopidexcel 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 **Clopidexcel 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 **Clopidexcel 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 occur:

- Fever, signs of infection or extreme tiredness. These may be due to rare decrease of some blood cells.
- Signs of a liver problem, such as yellowing of the skin and/or eyes (jaundice), whether or not associated with bleeding which appear under the skin 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 disorders such as rash, itching and blisters of the skin. These may 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 (unusual bleeding or bruising under the skin), 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 Clopidexcel 75:

If you were cut or injured, it may take longer than usual for bleeding to stop. This is linked to the way your medicine works as it prevents the formation of blood clots. For minor cuts and 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 occur in 1-10 out of 100 users):

Diarrhea, abdominal pain, indigestion or heartburn.

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

Vertigo, enlarged breasts in males.

Very rare side effects (effects that occur in less than 1 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; blisters of the skin; skin allergy; inflammation in the mucus membranes of the mouth; decrease in blood pressure; confusion; hallucinations; joint pain; muscular pain; changes in 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 occurs, 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 by clicking the link "דיווח על תופעות לוואי עקב טיפול תרופתי" found on the homepage of the Ministry of Health website (www.health.gov.il) directing to the online form for reporting side effects or via the 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, crospovidone, 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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Drug registration number at the national drug registry of the Ministry of Health:
144-29-33118-01

Registration holder: Dexcel Pharma Technologies Ltd., 21 Haftzadi Nahum St., Jerusalem 9548402, Israel