

Patient leaflet in accordance with the Pharmacists' Regulations  
(Preparations) - 1986

The medicinal product is dispensed with a doctor's prescription only

**Ezator S.K. 10 mg/10 mg**  
**Ezator S.K. 10 mg/20 mg**  
**Ezator S.K. 10 mg/40 mg**  
**Ezator S.K. 10 mg/80 mg**

## Film-coated tablets

Each film-coated tablet contains:

Ezetimibe 10 mg and atorvastatin (as calcium trihydrate) 10 mg, 20 mg, 40 mg or 80 mg.

For a list of inactive ingredients, please see Section 6, "Further information". See also Section 2.8, "Important information about some of the ingredients of **Ezator S.K.**."

### **Read all of the leaflet carefully before using the medicinal product.**

- This leaflet contains concise information about **Ezator S.K.**. If you have more questions, ask your doctor or pharmacist.
- This medicinal product has been prescribed for your treatment. Do not pass it on to others. It may harm them even if it seems that their medical condition is similar to yours.
- **Ezator S.K.** is intended for the treatment of adults over the age of 18 years.

### **1. What Ezator S.K. is and what it is used for**

**Ezator S.K.** is used, in combination with diet,

- for the treatment of primary heterozygous familial and non-familial] hypercholesterolemia or to reduce high levels of blood lipids (combined hyperlipidemia) in adults patients who are:
  - not balanced using statin only; or
  - already treated using statin and ezetimibe
- for the treatment of adult patients with hereditary disease (homozygous familial hypercholesterolemia). The treatment may be administered in combination with other treatments.
- in the event of heart disease, to reduce the risk of cardiovascular events (such as heart attack, stroke, surgery to improve blood flow to the heart, or hospitalization due to chest pain).

**Therapeutic group:** Ezetimibe belongs to a group of medicines called *CAIs* (cholesterol absorption inhibitors), which delay the absorption of cholesterol in the intestines. Atorvastatin belongs to a group of medicines called statins, which are inhibitors of the enzyme HMG-CoA reductase.

**Ezator S.K.** works to reduce your cholesterol in two ways. It reduces the absorption of cholesterol in your digestive system, as well as reducing the amount of cholesterol your body produces by itself.

Cholesterol is one of a number of fatty substances found in the blood circulation. Your total cholesterol is mostly made of HDL and LDL cholesterol.

LDL cholesterol is often referred to as "bad" cholesterol, because it can accumulate in your arterial walls creating plaque. Ultimately, the accumulation of this plaque can lead to arterial stenosis. This narrowing might slow down or block the flow of blood to vital organs such as the heart and the brain. This blockage could cause a heart attack or stroke.

HDL cholesterol is often referred to as “good” cholesterol, because it helps to prevent the accumulation of bad cholesterol in the arteries and protects against heart disease.

Triglycerides are another form of fat in your blood, which could increase your risk of getting heart disease.

**Ezator S.K.** does not help you lose weight.

## 2. Before use of Ezator S.K.

### 2.1 Do not use Ezator S.K. if:

- You are sensitive (allergic) to ezetimibe, atorvastatin or any of the other ingredients of **Ezator S.K.** (for a list of inactive ingredients, please refer to Section 6, “Further information”)
  - You [currently have] or have ever had a disease affecting the liver
  - You have had unexplained abnormal liver function blood tests
  - You are a woman of childbearing potential and you do not use reliable contraception
  - You are pregnant, trying to conceive, or are nursing
  - You are using a combination of glecaprevir/pibrentasvir for the treatment of hepatitis C

### 2.2 Special precautions regarding the use of Ezator S.K.

#### Before treatment with Ezator S.K., tell your doctor if:

- you have had a previous stroke involving intracerebral hemorrhage, or you have small pockets of fluid in the brain from previous cerebrovascular events
- you have kidney problems
- your thyroid gland function is reduced (hypothyroidism)
- you have had unexplained recurring aches or pains in the muscles, personal or familial history of muscle problems
- you have had previous muscle problems during treatment with other medicines to reduce cholesterol (e.g., other “statins” or “fibrate” medicines)
- you regularly consume a large amount of alcohol
- you have a history of liver disease
- you are over the age of 70 years
- you have been told by your doctor that you have an intolerance for certain sugars, contact your doctor before using this preparation
- you are currently taking or have taken within the last 7 days a medicine called fusidic acid (a medicine to treat bacterial infection) either orally or by injection. The combination of fusidic acid and **Ezator S.K.** may lead to severe muscle problems (rhabdomyolysis).

**Contact your doctor immediately if you experience unexplained pain in a muscle, sensitivity or weakness while taking Ezator S.K.. This is because, in rare cases, muscle problems might be severe, including muscle breakdown causing kidney damage. Atorvastatin is known to cause muscle problems, and cases of muscle problems have also been reported with ezetimibe.**

Please also tell your doctor or pharmacist if you have permanent muscle weakness. Further tests and medicines may be necessary to diagnose and treat this.

Check with your doctor or pharmacist before taking **Ezator S.K.:**

- if you have severe respiratory failure.

If any of the above applies to you (or if you are not certain), ask your doctor or pharmacist before taking **Ezator S.K.**, because your doctor will need to perform a blood test prior to, and potentially during, the treatment with **Ezator S.K.**, in order to detect your risk of muscle-related side effects. The risk of muscle-related side effects, such as muscle breakdown (rhabdomyolysis) is known to increase when certain

medicines are taken simultaneously (see Section 2.4, “Interactions with other medicines”).

While you are taking this medicine, your doctor will carefully monitor your condition if you have or are at risk of developing diabetes. You are expected to be at risk of developing diabetes if your blood sugar and lipid levels are high, you suffer from obesity and have high blood pressure.

Inform your doctor of all your medical conditions, including allergies.

Avoid the combined use of **Ezator S.K.** and fibrates (medicines to reduce cholesterol) because the combination of **Ezator S.K.** and fibrates has not been investigated.

### 2.3 Children and adolescents

**Ezator S.K.** is not intended for children and adolescents.

### 2.4 Interactions with other medicines

**If you are taking, or have recently taken, other medicines, including non-prescription medicines and food supplements, tell the attending doctor or pharmacist.** Certain medicines may change the effect of **Ezator S.K.** or their effect may be altered by **Ezator S.K.** (see Section 3 “How to use **Ezator S.K.**”). This kind of interaction could reduce the efficacy of one or both medicines. Alternatively, this could increase the risk or severity of side effects, including the severe side effects of muscle breakdown known as “rhabdomyolysis”, outlined in Section 4 “Side effects”.

- Cyclosporine (a medicine often used in post-transplantation patients)
- Erythromycin, clarithromycin, telithromycin, fusidic acid\*\*, rifampicin (medicines for treating bacterial infections)
- Ketoconazole, itraconazole, voriconazole, fluconazole, posaconazole (medicines for treating fungal infections)
- Gemfibrozil, other fibrates, nicotinic acid, colestipol, cholestyramine (medicines to regulate lipid levels)
- Several calcium channel blockers used for the treatment of angina or high blood pressure, such as amlodipine, diltiazem
- digoxin, verapamil, and amiodarone (medicines to regulate the heart rhythm)
- HIV medicines, such as ritonavir, lopinavir, atazanavir, indinavir, darunavir, the combination of tipranavir/ritonavir, etc. (AIDS medicines drugs)
- Certain hepatitis C drugs, for example, telaprevir, boceprevir and a combination of elbasvir/grazoprevir.
- Daptomycin (a drug used in complicated skin and skin-structure infections and bacteremia)

**\*\* If you need to take fusidic acid orally to treat a bacterial infection, you will need to temporarily stop treatment with this drug. Your doctor will tell you when it is safe for you to start retaking Ezator S.K.. Taking Ezator S.K. along with fusidic acid could, in rare cases, lead to weakness, sensitivity or muscle aches (rhabdomyolysis). See further information about rhabdomyolysis in Section 4 “Side effects”.**

- Other drugs known to interact with **Ezator S.K.**
  - Oral contraceptives (drugs to prevent pregnancy)
  - Stiripentol (anticonvulsant drug for epilepsy)
  - Cimetidine (a drug used to treat heartburn and stomach ulcers)
  - Phenazone (analgesic)
  - Antacids (aluminum- or magnesium-containing drugs for digestion problems)
  - Warfarin, phenprocoumon, acenocoumoral or fluindione (drugs to prevent blood clots)
  - Colchicine (used to treat gout)
  - St. John’s wort (a drug to treat depression)

### 2.5 Taking Ezator S.K. with food, drink, and alcohol

Please see Section 3 for instructions on how to take **Ezator S.K.**. Please note the following:

#### *Grapefruit juice*

Do not drink more than 1 or 2 small glasses of grapefruit juice per day, because large quantities of grapefruit

juice could change the effects of **Ezator S.K.**.

#### *Alcohol*

Avoid drinking too much alcohol while taking this medicine. For details please see section 2.2, “**Special precautions regarding the use of Ezator S.K.**”.

### **2.6 Pregnancy and breast-feeding**

Do not take **Ezator S.K.** if you are pregnant, trying to conceive or think that you might be pregnant. Do not take **Ezator S.K.** if you are a woman of childbearing potential unless you are using reliable contraception methods. If you have become pregnant while taking **Ezator S.K.**, stop taking it immediately and inform your doctor.

Do not take **Ezator S.K.** if you are breast-feeding.

The safety of **Ezator S.K.** during pregnancy and nursing has yet to be proven.

Consult your doctor or pharmacist before taking this medicine.

### **2.7 Driving and using machinery**

**Ezator S.K.** is not expected to affect your ability to drive or operate machinery. However, please bear in mind that some people may feel dizzy after taking **Ezator S.K.**

### **2.8 Important information about some of the ingredients in Ezator S.K.**

- **Ezator S.K.** tablets contain a sugar called lactose. If you have been told by your doctor that you have an intolerance for certain sugars, contact your doctor before taking this medicine (also see Section 6, “Further information”).
- This medicine contains mannitol. Mannitol could have a mild diuretic effect.
- This medicine contains less than 1 mmol sodium (23 mg) per tablet, i.e., it is actually ‘sodium-free’.

## **3. How to use Ezator S.K.**

Always use **Ezator S.K.** according to the doctor’s instructions. Check with the doctor or pharmacist if you are uncertain about the dosage and how to use **Ezator S.K.** Your doctor will decide the tablet strength suitable for you, based on your current treatment and your personal risk status.

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

- Before starting treatment with **Ezator S.K.**, you must be on a diet to lower your cholesterol.
- Maintain this cholesterol-lowering diet while taking **Ezator S.K.**

The standard dosage is usually:

1 tablet of **Ezator S.K.** orally once daily.

**Ezator S.K.** can be taken at any time of the day. It can be taken with or without food.

**Do not exceed the recommended dose.**

There is no information on crushing/splitting/chewing the tablets.

If your doctor prescribed you **Ezator S.K.** in combination with cholestyramine or another bile acid sequestrant (a medicine to lower cholesterol), you must take **Ezator S.K.** at least 2 hours before or 4 hours after taking the bile acid sequestrant.

### **If you have taken more Ezator S.K. than you should have**

Please contact your doctor or pharmacist. If you have taken an overdose or if a child has accidentally swallowed the medicine, refer immediately to a hospital emergency room and bring the medicine’s packaging with you.

### **If you have forgotten to take Ezator S.K.**

Do not take another dose; take your usual amount of **Ezator S.K.** at the usual time on the next day.

Always adhere to the treatment as recommended by the doctor.

Even if the state of your health improves, do not stop treatment with the medicine without consulting your doctor.

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

**If you have further questions regarding use of the medicine, please ask your doctor or pharmacist.**

#### **4. Side effects**

**As with any medicine, the use of Ezator S.K. 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.**

**If you experience any of the following severe side effects or symptoms, stop taking your tablets and inform your doctor immediately or refer to your nearest hospital emergency room.**

- A severe allergic reaction causing swelling of the face, tongue and throat which could cause serious breathing difficulties
- A severe disease with severe skin peeling and swelling, blisters on the skin, mouth, eyes, genitalia, and fever; skin rash with pink-red spots, especially on the hands or feet, which could become blisters
- Muscle weakness, sensitivity, muscle pain or rupture, red-brown urine discoloration, and especially if you are simultaneously feeling unwell or have a high fever, this could be caused by abnormal muscle breakdown, which could be life-threatening and lead to kidney problems.
- Lupus-like syndrome (including rash, joint disorders and effects on blood cells). Please consult your doctor as soon as possible if you are experiencing unexpected or abnormal bleeding or bruising problems, as this could indicate a liver problem.

Additional side effects:

Common side effects (could affect up to 1 in 10 users):

- Diarrhea
- Muscular pain

Uncommon side effects (could affect up to 1 in 100 users):

- Influenza
- Depression; sleeping difficulties; sleep disorder
- Dizziness; headache; tingling sensation
- Slow heart rate
- Hot flashes
- Shortness of breath
- Abdominal pain; abdominal swelling; indigestion; flatulence; frequent bowel movements; stomach inflammation; nausea; stomach discomfort; irritable bowel
- Acne; urticaria
- Joint pains; back pain; leg cramps; tiredness, cramps or muscle weakness; pain in arms and legs
- Abnormal weakness; feeling tired or sick; swelling, especially in the ankles (edema)
- Elevated values in certain liver function or muscle function lab tests (CK).
- Weight gain

In addition, the following side effects have been reported in individuals who have taken **Ezator S.K.**, or ezetimibe tablets or atorvastatin tablets:

- Allergic reactions including swelling of the face, lips, tongue and/or throat, which could cause difficulties with breathing or swallowing (requiring immediate care)
- Raised red rash, sometimes with target-shaped lesions
- Liver problems

- Cough
- Heartburn
- Reduced appetite; loss of appetite
- High blood pressure
- Skin rash and itch; allergic reactions including rash and urticaria
- Tendon damage
- Gallbladder stones or cholecystitis (which might cause abdominal pain, nausea, vomiting)
- Pancreatitis, often with severe abdominal pain
- Reduced blood cell count, which could cause bruising/bleeding (thrombocytopenia)
- Sinusitis; nosebleed
- Pain in the neck; pain; chest pain; sore throat
- Elevated and reduced blood sugar levels (if you have diabetes, you must continue to monitor your blood sugar levels carefully)
- Nightmares
- Numbness or tingling in the fingers and toes
- Reduced sensation of pain or touch
- Altered sense of taste; dry mouth
- Loss of memory
- Tinnitus; hearing loss
- Vomiting
- Belching
- Hair loss
- Increased body temperature
- White blood cell-positive urine tests
- Blurred vision; vision disorders
- Gynecomastia (enlarged breasts in men and women)

Potential side effects reported with use of statin medicines:

- Sexual performance difficulties
- Depression
- Respiratory issues including prolonged cough and/or shortness of breath or fever
- Diabetes. This is more likely if your blood sugar and lipid levels are elevated, you are overweight and have high blood pressure. Your doctor will monitor you while you are taking this medicine.
- Pain, sensitivity or permanent muscle weakness, especially if you are simultaneously feeling unwell or have a high fever, which may not resolve after stopping **Ezator S.K.** (incidence unknown).

**If a side effect appears, if one of the side effects gets worse, or when you experience a side effect that was not mentioned in this leaflet, please consult a doctor.**

#### Reporting side effects

Side effects can be reported to the Ministry of Health by clicking the link “Reporting side effects due to medication treatment” on the Ministry of Health homepage ([www.health.gov.il](http://www.health.gov.il)) that refers to the online form for reporting side effects, or by clicking the following link:

[/https://sideeffects.health.gov.il](https://sideeffects.health.gov.il)

#### **5. How to store Ezator S.K.**

- Prevent poisoning! This medicine, like any other medicine, should be kept in a secure place out of the reach and sight of children and/or babies, thereby preventing poisoning. Do not induce vomiting without explicit instructions from the doctor.
- Do not use **Ezator S.K.** after the expiry date (exp. Date) which is stated on the packaging. The expiry date refers to the last day of that month.
- **Storage conditions:** There are no special storage conditions for this medicine - please store in the

original packaging to protect from oxygen.

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

## **6. Additional information**

### **In addition to the active ingredients, the medicine also contains:**

mannitol, calcium carbonate, microcrystalline cellulose, croscarmellose sodium, hydroxypropyl cellulose, sodium lauryl sulfate, magnesium stearate, povidone, polysorbate 80, iron oxide yellow.

The film coating contains:

Ezator S.K. 10/10 mg, Ezator S.K. 20/10 mg, Ezator S.K. 40/10 mg contains: lactose monohydrate, hypromellose, titanium dioxide, macrogol.

Ezator S.K. 80/10 mg contains: hypromellose, titanium dioxide, talc, macrogol, iron oxide yellow.

### **What Ezator S.K. film-coated tablets look like and contents of the pack**

Ezator S.K. 10 mg/10 mg white, biconvex, round, film-coated tablets

Ezator S.K. 10 mg/20 mg white, biconvex, oval, film-coated tablets

Ezator S.K. 10 mg/40 mg white, biconvex, capsule-shaped, film-coated tablets

Ezator S.K. 10 mg/80 mg yellow, oblong, biconvex, film-coated tablets

Pack sizes: 10, 30 tablets per tray.

Not all pack sizes may be marketed.

### **Registration holder and its address:**

K.S. Kim International Ltd., 94 Yigal Alon Street, Tel-Aviv. 6789139

### **Name and address of the manufacturer:**

ELPEN Pharmaceutical Co., Inc

Marathonos Ave. 95, Pikermi Attiki, 19009, Greece

The brochure was prepared in November 2022 in accordance with Ministry of Health guidelines.

### **The medicinal product registration numbers in the Israel National Drug Registry at the Ministry of Health:**

**Ezator S.K. 10 mg/10 mg: 170-85-36878**

**Ezator S.K. 10 mg/20 mg: 170-86-36879**

**Ezator S.K. 10 mg/40 mg: 170-87-36880**

**Ezator S.K. 10 mg/80 mg: 170-88-36881**