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| Jardiance | Updated Patient Information |
| 10 mg, 25 mg | Feb 2024 |

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

This medicine is dispensed according to a physician's prescription only

Jardiance® 10 mg

Film-coated tablets

Each tablet of Jardiance 10 mg contains 10 mg empagliflozin.

Jardiance® 25 mg

Film-coated tablets

Each tablet of Jardiance 25 mg contains 25 mg empagliflozin.

Inactive ingredients and allergens in the medicine - see section 2 'Important information about some of this medicine's ingredients' and section 6 'Additional information'.

Read the entire leaflet carefully before you start using the medicine. This leaflet contains concise information about the medicine. If you have further questions, contact your physician or pharmacist. This medicine has been prescribed to treat your illness. Do not pass it on to others. It may harm them, even if it seems to you that their illness is similar to yours.

1. What is the medicine intended for?

Jardiance 10 mg and 25 mg is intended:

- to improve blood sugar level control in adults with type 2 diabetes, along with diet and exercise.
- to reduce the risk of cardiovascular death in adult patients with type 2 diabetes and existing cardiovascular disease.

Jardiance 10 mg is intended:

- to reduce the risk of death due to cardiovascular disease and hospitalization for heart failure in adults with heart failure.
- to reduce the risk of further deterioration of kidney function, end-stage kidney disease (ESKD), death due to cardiovascular disease and hospitalization in adults with chronic kidney disease.
- to improve control of blood sugar level in children and adolescents over the age of 10 with type 2 diabetes, along with diet and exercise.

Jardiance is not recommended for patients with type 1 diabetes or for the treatment of diabetic ketoacidosis.

Jardiance is not recommended for use to improve control of blood sugar level in patients with type 2 diabetes and estimated glomerular filtration rate (eGFR) less than 30 ml/min/1.73m².

Jardiance is not recommended for patients with polycystic kidney disease or patients who are taking or have recently taken certain types of immunosuppressive therapy to treat kidney disease. Jardiance is not expected to work if you have these conditions.

Therapeutic group: SGLT2 (sodium-glucose co-transporter 2) inhibitors.

2. Before using this medicine

Do not use this medicine if:

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| <ul style="list-style-type: none"> • You are sensitive (allergic) to the active ingredient empagliflozin or to any of the additional ingredients that the medicine contains (see section 6 – 'Additional |
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|--------------|-----------------------------|
| Jardiance | Updated Patient Information |
| 10 mg, 25 mg | Feb 2024 |

information', for the list of inactive ingredients). Symptoms of a serious allergic reaction to Jardiance may include:

- rash
- raised, red areas on your skin (hives)
- swelling of your face, lips, mouth, and throat that may cause difficulty in breathing or swallowing

If you experience any of these symptoms, stop taking Jardiance and contact your physician right away or go to the nearest hospital emergency room.

Special warnings about using this medicine

Before taking Jardiance, tell your physician about any previous medical condition, including if:

- You have type 1 diabetes. Jardiance should not be used to treat people with type 1 diabetes
- You have a history of ketoacidosis (elevated ketone levels in your blood or urine)
- You have a serious infection
- You have a history of infection of the vagina or penis
- You have a history of amputation
- You have kidney problems
- You have liver problems
- You have a history of urinary tract infections or problems with urination
- You are on a low sodium (salt) diet. Your physician may change your diet or your dose of the medicine
- You are about to have surgery. Your physician may decide to stop use of Jardiance before the surgery. Consult your physician about when to stop taking Jardiance and when to resume your treatment
- You are eating less or there is a change in your diet
- You are dehydrated
- You have ever had an allergic reaction to Jardiance
- You are pregnant or plan to become pregnant - see section 2 'Pregnancy and breastfeeding'.
- You are breastfeeding or plan to breastfeed - see section 2 'Pregnancy and breastfeeding'.

Additionally, before treatment with Jardiance or while taking Jardiance, tell your physician if:

- You have an acute illness such as the flu, a urinary tract infection, you are vomiting, have diarrhea or a fever, or if you are unable to eat or drink, are eating less or there is a change in your diet. Such conditions can lead to dehydration. Your physician can ask you to stop taking Jardiance until you recover to prevent dehydration. Dehydration can damage your kidneys.
- You have or have had problems with your pancreas, including pancreatitis or surgery on your pancreas.
- You drink alcohol very often or drink a lot of alcohol in a short time ("binge" drinking).
- You are being treated at the same time with insulin, and there has been a sharp drop in your insulin dose.

| | |
|--------------|-----------------------------|
| Jardiance | Updated Patient Information |
| 10 mg, 25 mg | Feb 2024 |

These conditions may put you at risk of developing diabetic ketoacidosis. See section 'Side effects'.

Children and adolescents

It is not known whether Jardiance is safe and effective in children under the age of 10.

Tests and follow-up

- Diet and exercise can help your body to use blood sugar better. It is important to maintain the diet and exercise plan recommended by the physician during the treatment with Jardiance.
Like other diabetes treatments, it is advisable to monitor the response to treatment periodically. The physician may recommend treatment with additional medicines to achieve your treatment goals.
- Your physician may refer you for blood tests prior to beginning treatment and during treatment with Jardiance as needed.
- **Urine glucose test** - During the treatment with Jardiance, you may have sugar in your urine, which will show a positive result in urine glucose test.
- **Diabetic foot:** It is important to check your feet regularly according to the recommendations given by your healthcare professional.

Drug interactions

If you are taking or have recently taken other medicines, including non-prescription medicines and dietary supplements, tell your physician or pharmacist, because Jardiance may affect the way certain medicines work, and certain medicines may affect how Jardiance works.

Especially tell your physician or pharmacist if you take any of the following medicines:

- **A medicine used to excrete fluids from the body (diuretic).** Combined use with Jardiance may result in increased urine volume and frequency of urination, which may increase the risk of dehydration.
Possible symptoms of excessive loss of body fluids appear in section 4 – 'Side effects'.
- **Other blood-sugar-lowering medicines (such as insulin or sulfonylurea).** Combined use with Jardiance may increase the risk of low blood sugar levels (hypoglycemia).
Therefore, when used in combination with Jardiance, lower dosages of blood-sugar-lowering medicines (such as insulin or sulfonylurea) may be required to reduce the risk of low blood sugar level.
- **Lithium.** Combined use with Jardiance may reduce lithium concentration in your blood. Therefore, lithium levels in the blood should be monitored more frequently after starting treatment with Jardiance and when dosage changes.

Know the medicines you take and keep a list of them to show your physician and pharmacist when you get a new medicine.

Using the medicine and food

You can take this medicine with or without food.

Pregnancy and breastfeeding

Consult a physician or a pharmacist prior to using this medicine if you are pregnant, think that you may be pregnant, plan to become pregnant, are breastfeeding or plan to

| | |
|--------------|-----------------------------|
| Jardiance | Updated Patient Information |
| 10 mg, 25 mg | Feb 2024 |

breastfeed.

Do not use Jardiance if you are pregnant. Jardiance may harm your unborn baby. If you become pregnant while you are taking Jardiance, tell your physician as soon as possible. Consult your physician about the best way to control your blood sugar levels while you are pregnant.

Do not breastfeed while taking Jardiance. Jardiance may pass into your breast milk and harm your baby. Consult your physician about the best way to feed your baby if you are taking Jardiance.

Driving and using machines

Jardiance has a minor influence on your ability to drive and operate machines.

Taking this medicine, particularly in combination with medicines called sulfonylurea or with insulin may cause low blood sugar levels (hypoglycemia), which may cause symptoms such as shaking, sweating and change in vision, and may affect your ability to drive and operate machines.

Do not drive or use any tool or machine, if you suffer from dizziness during the treatment with Jardiance.

Important information about some of this medicine's ingredients

Jardiance contains lactose (milk sugar). If the physician has told you that you have an intolerance to some sugars, contact the physician before taking the medicine.

This medicine contains less than 1 millimole sodium (23 mg) per tablet so it is considered 'sodium free'.

3. How should you use the medicine?

Always use according to your physician's instructions. Check with your physician or pharmacist if you are not sure about your dosage or about how to take this medicine. Only your physician will determine your dosage and how you should take this medicine.

- The standard initial dosage is usually one Jardiance 10 mg tablet once daily in the morning, with or without food. If you have type 2 diabetes, the physician will decide whether to increase the dosage to Jardiance 25 mg.
- Your dosage will be adjusted by the physician if necessary. Do not change the dosage unless your physician has told you to.

Do not exceed the recommended dose.

Method of use

- Swallow the tablet whole with water. There is no information about crushing/splitting/chewing.
- You may take the medicine with or without food.
- You may take the medicine once a day in the morning; try to take it at the same time of day. This will help you to remember to take the medicine.
- Your physician can tell you to take Jardiance along with other diabetes medicines. A low blood sugar level can happen more often when Jardiance is taken with certain other diabetes medicines. See section 2 'Drug interactions' and section 4 'Side effects', for more information.
- When your body is under certain types of stress, such as: fever, trauma (for example, a car accident), infection or surgery, the amount of diabetes medicine you need may change. Tell your physician right away if you have any of these conditions and follow your physician's instructions.

| | |
|--------------|-----------------------------|
| Jardiance | Updated Patient Information |
| 10 mg, 25 mg | Feb 2024 |

If you have accidentally taken a higher dosage

If you have taken an overdose or a child has accidentally swallowed some medicine, immediately see a physician or go to the nearest hospital emergency room and bring the medicine package with you.

If you forget to take the medicine

Take the dose as soon as you remember and the next dose at the usual time. If it is almost time to take the next dose, skip the forgotten dose and take the next dose at the usual time. Do not take a double dose. Consult with your physician if you have questions about a forgotten dose.

Adhere to the treatment as recommended by your physician. Even if your health has improved, do not stop treatment with the medicine without consulting your physician.

If you stop taking the medicine

Do not stop the treatment with the medicine without consulting your physician. If you stop taking the medicine, your blood sugar levels may increase.

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

If you have any further questions about using this medicine, consult your physician or pharmacist.

4. Side effects

Like with all medicines, using Jardiance may cause side effects in some users. Do not be alarmed by this list of side effects; You may not experience any of them.

Important information regarding Jardiance:

Jardiance can cause serious side effects.

Contact the physician as soon as possible if you experience any of the following serious side effects:

- **Ketoacidosis (elevated ketone levels in your blood or urine)** (frequency unknown): Ketoacidosis appeared in people who have **type 1 diabetes or type 2 diabetes** during treatment with Jardiance. Ketoacidosis has also appeared in people with diabetes who vomited or who had surgery during treatment with Jardiance. Ketoacidosis is a serious medical condition that requires treatment in a hospital. Ketoacidosis can lead to death. **Ketoacidosis may appear during treatment with Jardiance even if your blood sugar levels are below 250 mg/dL.**
Stop using Jardiance and contact your physician right away or go to the nearest emergency room if you experience any of the following symptoms: nausea, vomiting, stomach-area (abdominal) pain, tiredness, and trouble breathing.
If you experience any of these symptoms during treatment with Jardiance, if possible, check for ketones in the urine, even if your blood sugar levels are below 250 mg/dL.
- **Dehydration** (an uncommon side effect - affects 1–10 in 1,000 users):
Jardiance may cause some patients to become dehydrated (the loss of body fluids and salt). Dehydration may cause you to feel dizzy, faint or weak, especially when you shift from sitting/lying down to standing up. There have been reports of sudden worsening of kidney function in people taking Jardiance. You may be at higher chance of dehydration if you:

| | |
|--------------|-----------------------------|
| Jardiance | Updated Patient Information |
| 10 mg, 25 mg | Feb 2024 |

- take medicines to lower your blood pressure, including diuretics
- are on a low sodium (salt) diet
- have kidney problems
- are 65 years of age or older

Consult your physician about what you should do to prevent dehydration, including how much fluid you should drink on a daily basis.

Contact your physician **right away** if you reduce the amount of food you eat or the amount of liquid you drink, for example if you vomit and cannot eat, or start to lose fluids from your body, for example from vomiting, diarrhea or being in the sun too long.

- **Fournier’s gangrene – a rare but life-threatening bacterial infection in the tissues under the skin in the genital area (up to the anus)** (frequency unknown):

Contact your physician as soon as possible if you notice any of the following side effects:

redness, pain, tenderness or swelling in the genital area up to the anus, with fever over 38°C or feeling generally unwell and weak. These side effects may indicate the development of Fournier’s gangrene, a rare but life-threatening bacterial infection in the tissues under the skin in the genital area (up to the anus). There are a number of reports of occurrence of this side effect in women and men who took diabetes medicines that belong to the same group of medicines as Jardiance. This side effect may lead to hospitalization, may require surgery and may even lead to death. The symptoms of Fournier’s gangrene may get worse quickly so it is **important to get medical help quickly if you get any of these symptoms.**

- **Serious urinary tract infections:** (frequency unknown):

Serious urinary tract infections that may lead to hospitalization have happened in people who are taking Jardiance. Tell your physician if you have signs or symptoms of a urinary tract infection such as a burning when passing urine, urinary frequency and urgency, pain in the pelvis, or blood in the urine. Sometimes people also may have a fever, back pain, nausea or vomiting.

- **Low blood sugar level (hypoglycemia)** (affects more than 1 in 10 users):

In adults, if you take Jardiance with another medicine that can cause low blood sugar levels such as insulin or sulfonylurea, your risk of suffering from low blood sugar level is higher.

In children 10 years of age and older, the risk for low blood sugar is higher with Jardiance even if you do not use another medicine that can also lower blood sugar level.

The dosage of your insulin or sulfonylurea may need to be lowered while you take Jardiance.

Signs and symptoms of low blood sugar level may include irritability, sleepiness, dizziness, shaking or feeling jittery, sweating, confusion, fast heartbeat, hunger, headache, weakness.

The physician will instruct you how to treat low blood sugar levels and what to do if any of the above signs appear. If symptoms of low blood sugar level appear, eat glucose tablets, a snack with high sugar content or drink fruit juice. Take your blood sugar level if possible and rest.

| | |
|--------------|-----------------------------|
| Jardiance | Updated Patient Information |
| 10 mg, 25 mg | Feb 2024 |

- **Vaginal yeast infection** (common side effect – affects 1–10 in 100 users).
The symptoms of this are vaginal odor, white or yellowish vaginal discharge (discharge may be lumpy or look like cottage cheese), vaginal itching.

- **Yeast infection of the skin around the penis** (infection of the glans only, or of the glans and foreskin in uncircumcised men) (common side effect – affects 1-10 in 100 users).

Swelling of the penis in uncircumcised patients may develop, which makes it difficult to pull back the skin around the tip of the penis. Other symptoms of yeast infection of the penis include redness, itching or swelling of the penis, rash on the penis, foul smelling discharge from the penis, pain in the skin around the penis.

Talk to your physician about what you should do if you get symptoms of a yeast infection of the vagina or penis. Your physician may suggest you use over-the-counter antifungal medicines. Contact your physician **right away** if the use of these medicines does not help and your symptoms do not go away.

- **Amputations:**

SGLT2 inhibitors may increase your risk of lower limb amputations. You may be at a higher risk of lower limb amputation if:

- you have a history of amputation
- you have had blocked or narrowed blood vessels, usually in your leg
- you have had diabetic foot infection, ulcers or sores

Call your physician right away if you have new pain or tenderness, any sores, ulcers or infections in your leg or foot. Talk to your physician about proper foot care.

- **Serious allergic reactions** (frequency unknown): Serious allergic reactions have occurred in patients taking Jardiance. Symptoms of an allergic reaction can include:
 - rash
 - raised, red areas on your skin (hives)
 - swelling of the face, lips, mouth and throat, which may lead to difficulty in breathing or swallowing

If you experience any symptom of a serious allergic reaction, stop taking Jardiance and contact your physician **immediately** or go to the nearest hospital emergency room.

Additional side effects:

Very common side effects (affect more than 1 in 10 users) include:

- urinary tract infections in women

Common side effects (affect 1-10 in 100 users):

- urinary tract infections in men
- upper respiratory tract infections
- increased urination
- changes in blood lipid profile
- joint pain
- nausea

| | |
|--------------|-----------------------------|
| Jardiance | Updated Patient Information |
| 10 mg, 25 mg | Feb 2024 |

- thirst

Uncommon side effects (affect 1-10 in 1,000 users):

- Decrease in blood volume. The signs of decrease in blood volume are:
 - critical decrease in blood pressure
 - decrease in blood pressure
 - dehydration
 - blood pressure drop upon shifting from sitting/lying down to standing up
 - fainting

Side effects reported after marketing and whose frequency is unknown (the frequency of these effects has not been established yet):

- inflammation of the kidney and renal pelvis (pyelonephritis)
- sepsis following a urinary tract infection (urosepsis)
- skin reactions (such as rash or hives)
- acute kidney injury
- constipation

If you experience any side effect, if any side effect gets worse, or if you experience a side effect not mentioned in the leaflet, consult your physician.

Reporting side effects

You can report side effects to the Ministry of Health by following the link "Reporting side effects of Drug Treatment" on the Ministry of Health home page (www.health.gov.il) which links to an online form for reporting side effects. You can also use this link: <https://sideeffects.health.gov.il>

5. How to store the medicine?

- Store the medicine below 30°C.
- Prevent poisoning! To prevent poisoning, keep this, and all other medicines, in a closed place, out of sight and reach of children and/or infants. Do not induce vomiting unless explicitly instructed to do so by your physician.
- Do not use the medicine after the expiration date (exp. date) which is stated on the carton and blister tray. The expiration date refers to the last day of that month.
- Do not use this medicine if the package is damaged.
- Do not throw away medicines in the household trash or sink. 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:
Lactose monohydrate, cellulose microcrystalline, hydroxypropylcellulose, croscarmellose sodium, colloidal anhydrous silica, magnesium stearate.
The tablet coating is composed of:
Hypromellose 2910, titanium dioxide, talc, macrogol 400, iron oxide yellow.
- What the medicine looks like and contents of the pack:
Jardiance 10 mg are round, pale yellow, biconvex, bevel-edged, film-coated tablets.

| | |
|--------------|-----------------------------|
| Jardiance | Updated Patient Information |
| 10 mg, 25 mg | Feb 2024 |

“S 10” is debossed on one side of the tablet and the Boehringer Ingelheim logo appears on the other side.

Jardiance 25 mg are oval, pale yellow, biconvex, film-coated tablets. “S 25” is debossed on one side of the tablet and the Boehringer Ingelheim logo appears on the other side.

The tablets are packed in blisters. Each pack contains 7, 30, 60 or 90 film-coated tablets. Not all pack sizes may be marketed.

- Registration holder and importer's name and address:
Boehringer Ingelheim Israel Ltd., 89 Medinat Ha-Yehudim St., P.O. Box 4124,
Herzliya Pituach 4676672.
- This leaflet was revised in February 2024.
- Registration number of the medicine in the Ministry of Health's National Drug Registry:
Jardiance 10 mg 154-22-34202-00
Jardiance 25 mg 154-23-34203-00