

**MEDICATION GUIDE**  
**SYNJARDY® (sin-JAR-dee)**  
**(empagliflozin and metformin hydrochloride tablets), for oral use**  
**and**  
**SYNJARDY® XR (sin-JAR-dee XR)**  
**(empagliflozin and metformin hydrochloride extended-release tablets), for oral use**

**What is the most important information I should know about SYNJARDY or SYNJARDY XR?**

SYNJARDY or SYNJARDY XR can cause serious side effects, including:

- **Lactic Acidosis. Metformin hydrochloride (HCl), one of the medicines in SYNJARDY and SYNJARDY XR, can cause a rare but serious condition called lactic acidosis (a build-up of lactic acid in the blood) that can cause death. Lactic acidosis is a medical emergency and must be treated in a hospital.**

**Stop taking SYNJARDY or SYNJARDY XR and call your healthcare provider right away or go to the nearest hospital emergency room if you get any of the following symptoms of lactic acidosis:**

- feel very weak and tired
- have unusual sleepiness or sleep longer than usual
- have unusual (not normal) muscle pain
- feel cold, especially in your arms and legs
- have trouble breathing
- feel dizzy or lightheaded
- have unexplained stomach or intestinal problems with nausea and vomiting, or diarrhea
- have a slow or irregular heartbeat

**You have a higher chance of getting lactic acidosis with SYNJARDY or SYNJARDY XR if you:**

- have moderate to severe kidney problems.
- have liver problems.
- drink a lot of alcohol (very often or short-term “binge” drinking).
- get dehydrated (lose a large amount of body fluids). This can happen if you are sick with a fever, vomiting, or diarrhea. Dehydration can also happen when you sweat a lot with activity or exercise and do not drink enough fluids.
- have certain x-ray tests with injectable dyes or contrast agents.
- have surgery or other procedure for which you need to restrict the amount of food and liquid you eat and drink.
- have congestive heart failure.
- have a heart attack, severe infection, or stroke.
- are 65 years of age or older.

Tell your healthcare provider if you have any of the problems in the list above. Tell your healthcare provider that you are taking SYNJARDY or SYNJARDY XR before you have surgery or x-ray tests. Your healthcare provider may need to stop your SYNJARDY or SYNJARDY XR for a while if you have surgery or certain x-ray tests. SYNJARDY or SYNJARDY XR can have other serious side effects. See **“What are the possible side effects of SYNJARDY or SYNJARDY XR?”**

- **Diabetic ketoacidosis (increased ketones in your blood or urine) in people with type 1 and other ketoacidosis.** SYNJARDY or SYNJARDY XR can cause ketoacidosis that can be life-threatening and may lead to death. Ketoacidosis is a serious condition which needs to be treated in a hospital. People with type 1 diabetes have a high risk of getting ketoacidosis. People with type 2 diabetes or pancreas problems also have an increased risk of getting ketoacidosis. Ketoacidosis can also happen in people who: are sick, cannot eat or drink as usual, skip meals, are on a diet high in fat and low in carbohydrates (ketogenic diet), take less than the usual amount of insulin or miss insulin doses, drink too much alcohol, have a loss of too much fluid from the body (volume depletion), or who have surgery. Ketoacidosis can happen even if your blood sugar is less than 250 mg/dL. Your healthcare provider may ask you to periodically check ketones in your urine or blood. **Stop taking SYNJARDY or SYNJARDY XR and call your healthcare provider or get medical help right away if you get any of the following. If possible, check for ketones in your urine or blood, even if your blood sugar is less than 250 mg/dL:**

- nausea
- tiredness
- vomiting
- trouble breathing
- stomach-area (abdominal) pain
- ketones in your urine or blood

- **Dehydration. SYNJARDY or SYNJARDY XR can cause some people to become dehydrated (the loss of body water and salt). Dehydration may cause you to feel dizzy, faint, light-headed, or weak, especially when you stand up (orthostatic hypotension). There have been reports of sudden worsening of kidney function in people who are taking SYNJARDY or SYNJARDY XR. You may be at higher risk of dehydration if you:**

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

Talk to your healthcare provider about what you can do to prevent dehydration including how much fluid you should drink on a daily basis.

Call your healthcare provider right away if you reduce the amount of food or liquid you drink, for example if you are sick

or you cannot eat, or start to lose liquids from your body, for example from vomiting, diarrhea or being in the sun too long.

- **Vaginal yeast infection.** Symptoms of a vaginal yeast infection include:
  - 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 (balanitis or balanoposthitis).** Swelling of an uncircumcised penis may develop that 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 of the penis
  - foul smelling discharge from the penis
  - pain in the skin around penis

**Talk to your healthcare provider about what to do if you get symptoms of a yeast infection of the vagina or penis.** Your healthcare provider may suggest you use an over-the-counter antifungal medicine. Talk to your healthcare provider right away if you use an over-the-counter antifungal medication and your symptoms do not go away.

### **What is SYNJARDY or SYNJARDY XR?**

SYNJARDY or SYNJARDY XR is a prescription medicine that contains 2 diabetes medicines, empagliflozin (JARDIANCE) and metformin HCl.

SYNJARDY can be used along with diet and exercise to improve blood sugar (glucose) in adults and children who are 10 years of age and older with type 2 diabetes.

SYNJARDY XR can be used along with diet and exercise to improve blood sugar (glucose) in adults with type 2 diabetes.

One of the medicines in SYNJARDY and SYNJARDY XR, empagliflozin (JARDIANCE), can also be used in adults with type 2 diabetes:

- who have known cardiovascular disease to reduce the risk of cardiovascular death.
- who have heart failure (when the heart cannot pump enough blood to the rest of your body) to reduce the risk of cardiovascular death and hospitalization for heart failure.
- SYNJARDY or SYNJARDY XR is not for use to lower blood sugar (glucose) in people with type 1 diabetes. It may increase their risk of diabetic ketoacidosis (increased ketones in blood or urine).
- SYNJARDY or SYNJARDY XR is only for use in people with type 2 diabetes, because it contains the prescription medicine metformin HCl.
- It is not known if SYNJARDY is safe and effective in children under 10 years of age.
- It is not known if SYNJARDY XR is safe and effective in children.

### **Who should not take SYNJARDY or SYNJARDY XR?**

#### **Do not take SYNJARDY or SYNJARDY XR if you:**

- have severe kidney problems, end stage renal disease or are on dialysis.
- have a condition called metabolic acidosis or diabetic ketoacidosis (increased ketones in the blood or urine).
- are allergic to empagliflozin (JARDIANCE), metformin HCl, or any of the ingredients in SYNJARDY or SYNJARDY XR. See the end of this Medication Guide for a complete list of ingredients in SYNJARDY and SYNJARDY XR. Symptoms of a serious allergic reaction to SYNJARDY and SYNJARDY XR 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 have any of these symptoms, stop taking SYNJARDY or SYNJARDY XR and call your healthcare provider right away or go to the nearest hospital emergency room.

### **What should I tell my healthcare provider before taking SYNJARDY or SYNJARDY XR?**

**Before taking SYNJARDY or SYNJARDY XR, tell your healthcare provider about all of your medical conditions, including if you:**

- have type 1 diabetes or have had diabetic ketoacidosis.
- have a decrease in your insulin dose.
- have a serious infection.
- have a history of infection of the vagina or penis.
- have a history of amputation.
- have kidney problems.
- have liver problems.
- have heart problems, including congestive heart failure.
- are 65 years of age or older.
- have a history of urinary tract infections or problems with urination.
- are on a low sodium (salt) diet. Your healthcare provider may change your diet or your dose.
- are going to have surgery. Your healthcare provider may stop your SYNJARDY or SYNJARDY XR before you have surgery. Talk to your healthcare provider if you are having surgery about when to stop taking SYNJARDY or SYNJARDY XR and when to start it again.

- are eating less, or there is a change in your diet.
- are dehydrated.
- have or have had problems with your pancreas, including pancreatitis or surgery on your pancreas.
- drink alcohol very often or drink a lot of alcohol in the short term (“binge” drinking).
- have ever had an allergic reaction to SYNJARDY or SYNJARDY XR.
- are going to get an injection of dye or contrast agents for an x-ray procedure. SYNJARDY or SYNJARDY XR may need to be stopped for a short time. Talk to your healthcare provider about when you should stop SYNJARDY or SYNJARDY XR and when you should start SYNJARDY or SYNJARDY XR again. See **“What is the most important information I should know about SYNJARDY or SYNJARDY XR?”**
- have low levels of vitamin B<sub>12</sub> in your blood.
- are pregnant or plan to become pregnant. SYNJARDY or SYNJARDY XR may harm your unborn baby. If you become pregnant while taking SYNJARDY or SYNJARDY XR, tell your healthcare provider as soon as possible. Talk with your healthcare provider about the best way to control your blood sugar while you are pregnant.
- are breastfeeding or plan to breastfeed. SYNJARDY or SYNJARDY XR may pass into your breast milk and may harm your baby. Talk with your healthcare provider about the best way to feed your baby if you are taking SYNJARDY or SYNJARDY XR. Do not breastfeed while taking SYNJARDY or SYNJARDY XR.
- are a person who has not gone through menopause (premenopausal) who does not have periods regularly or at all. SYNJARDY or SYNJARDY XR can cause the release of an egg from an ovary in a person (ovulation). This can increase your chance of getting pregnant. Tell your healthcare provider right away if you become pregnant while taking SYNJARDY or SYNJARDY XR.

**Tell your healthcare provider about all the medicines you take**, including prescription and over-the-counter medicines, vitamins, and herbal supplements.

SYNJARDY or SYNJARDY XR may affect the way other medicines work, and other medicines may affect how SYNJARDY or SYNJARDY XR works. Know the medicines you take. Keep a list of them to show your healthcare provider and pharmacist when you get a new medicine.

#### **How should I take SYNJARDY or SYNJARDY XR?**

- Take SYNJARDY or SYNJARDY XR exactly as your healthcare provider tells you to take it.
- **If you are prescribed SYNJARDY:**
  - take SYNJARDY by mouth 2 times each day with meals. Taking SYNJARDY with meals may lower your chance of having an upset stomach.
- **If you are prescribed SYNJARDY XR:**
  - take SYNJARDY XR by mouth 1 time each day with a meal in the morning. Taking SYNJARDY XR with a meal may lower your chance of having an upset stomach.
  - swallow SYNJARDY XR tablets whole. Do not break, cut, crush, dissolve, or chew SYNJARDY XR tablets. If you cannot swallow SYNJARDY XR tablets whole, tell your healthcare provider.
  - you may see something that looks like the SYNJARDY XR tablet in your stool (bowel movement). This is not harmful and should not affect the way SYNJARDY XR works to control your diabetes.
- Your healthcare provider will tell you how much SYNJARDY or SYNJARDY XR to take and when to take it. Your healthcare provider may change your dose if needed.
- Your healthcare provider may tell you to take SYNJARDY or SYNJARDY XR along with other diabetes medicines. Low blood sugar can happen more often when SYNJARDY or SYNJARDY XR is taken with certain other diabetes medicines. See **“What are the possible side effects of SYNJARDY or SYNJARDY XR?”**
- If you miss a dose, take it as soon as you remember. If it is almost time for your next dose, skip the missed dose and take the medicine at the next regularly scheduled time. Do not take two doses of SYNJARDY or SYNJARDY XR at the same time. Talk with your healthcare provider if you have questions about a missed dose.
- If you take too much SYNJARDY or SYNJARDY XR, call your healthcare provider or Poison Help line at 1-800-222-1222, or go to the nearest hospital emergency room right away.
- When taking SYNJARDY or SYNJARDY XR, you may have sugar in your urine, which will show up on a urine test.
- When your body is under some types of stress, such as fever, trauma (such as a car accident), infection, or surgery, the amount of diabetes medicine you need may change. Tell your healthcare provider right away if you have any of these conditions and follow your healthcare provider’s instructions.
- Your healthcare provider may do certain blood tests before you start SYNJARDY or SYNJARDY XR and during your treatment as needed.

#### **What should I avoid while taking SYNJARDY or SYNJARDY XR?**

Avoid drinking alcohol very often, or drinking a lot of alcohol in a short period of time (“binge” drinking). It can increase your chances of getting serious side effects.

#### **What are the possible side effects of SYNJARDY or SYNJARDY XR?**

**SYNJARDY or SYNJARDY XR may cause serious side effects, including:**

- See **“What is the most important information I should know about SYNJARDY or SYNJARDY XR?”**
- **Serious urinary tract infections.** Serious urinary tract infections that may lead to hospitalization have happened in

people who are taking empagliflozin, one of the medicines in SYNJARDY and SYNJARDY XR. Tell your healthcare provider if you have any signs or symptoms of a urinary tract infection such as a burning feeling when passing urine, a need to urinate often, the need to urinate right away, pain in the lower part of your stomach (pelvis), or blood in the urine. Sometimes people also may have a fever, back pain, nausea or vomiting.

- **Low blood sugar (hypoglycemia).** In adults, if you take SYNJARDY or SYNJARDY XR with another medicine that can cause low blood sugar, such as a sulfonylurea or insulin, your risk of getting low blood sugar is higher. In children who are 10 years of age and older, the risk for low blood sugar is higher with SYNJARDY even if you do not use another medicine that can also lower blood sugar. The dose of your sulfonylurea medicine or insulin may need to be lowered while you take SYNJARDY or SYNJARDY XR.

Signs and symptoms of low blood sugar may include:

- |              |                  |                              |             |
|--------------|------------------|------------------------------|-------------|
| ○ headache   | ○ irritability   | ○ confusion                  | ○ dizziness |
| ○ drowsiness | ○ hunger         | ○ shaking or feeling jittery | ○ sweating  |
| ○ weakness   | ○ fast heartbeat |                              |             |

- **A rare but serious bacterial infection that causes damage to the tissue under the skin (necrotizing fasciitis) in the area between and around the anus and genitals (perineum).** Necrotizing fasciitis of the perineum has happened in people who take empagliflozin, one of the medicines in SYNJARDY and SYNJARDY XR. Necrotizing fasciitis of the perineum may lead to hospitalization, may require multiple surgeries, and may lead to death. **Seek medical attention immediately if you have a fever or you are feeling very weak, tired or uncomfortable (malaise), and you develop any of the following symptoms in the area between and around your anus and genitals:**

- |                      |            |                              |
|----------------------|------------|------------------------------|
| ○ pain or tenderness | ○ swelling | ○ redness of skin (erythema) |
|----------------------|------------|------------------------------|

- **Amputations. SGLT2 inhibitors may increase your risk of lower limb amputations.**

You may be at higher risk of lower limb amputation if you:

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

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

- **Serious allergic reactions.** If you have any symptoms of a serious allergic reaction, stop taking SYNJARDY or SYNJARDY XR and call your healthcare provider right away or go to the nearest hospital emergency room. See **“Who should not take SYNJARDY or SYNJARDY XR?”**.
- **Low vitamin B<sub>12</sub> (vitamin B<sub>12</sub> deficiency).** Using metformin for long periods of time may cause a decrease in the amount of vitamin B<sub>12</sub> in your blood, especially if you have had low vitamin B<sub>12</sub> blood levels before. Your healthcare provider may do blood tests to check your vitamin B<sub>12</sub> levels.

**The most common side effects of SYNJARDY or SYNJARDY XR include:**

- |  |                      |
|--|----------------------|
| • low blood sugar                      | • gas                |
| • urinary tract infections             | • stomach discomfort |
| • stuffy or runny nose and sore throat | • indigestion        |
| • yeast infections in females          | • weakness           |
| • diarrhea                             | • headache           |
| • nausea or vomiting                   |                      |

These are not all the possible side effects of SYNJARDY or SYNJARDY XR. For more information, ask your healthcare provider or pharmacist.

Call your doctor for medical advice about side effects. You may report side effects to FDA at 1-800-FDA-1088.

**How should I store SYNJARDY or SYNJARDY XR?**

- Store SYNJARDY or SYNJARDY XR at room temperature between 68°F to 77°F (20°C to 25°C).
- **Keep SYNJARDY or SYNJARDY XR and all medicines out of the reach of children.**

**General information about the safe and effective use of SYNJARDY or SYNJARDY XR.**

Medicines are sometimes prescribed for purposes other than those listed in a Medication Guide. Do not use SYNJARDY or SYNJARDY XR for a condition for which it was not prescribed. Do not give SYNJARDY or SYNJARDY XR to other people, even if they have the same symptoms that you have. It may harm them.

You can ask your pharmacist or healthcare provider for information about SYNJARDY or SYNJARDY XR that is written for health professionals.

**What are the ingredients in SYNJARDY?**

**Active Ingredients:** empagliflozin and metformin HCl

**Inactive Ingredients:** colloidal silicon dioxide, copovidone, corn starch, and magnesium stearate. In addition, the film coating contains the following inactive ingredients: hypromellose, polyethylene glycol 400, talc, and titanium dioxide. 5 mg/500 mg and 5 mg/1,000 mg tablets also contain ferric oxide yellow. 12.5 mg/500 mg and 12.5 mg/1,000 mg tablets also contain black ferrousferrous oxide and ferric oxide red.

**What are the ingredients in SYNJARDY XR?**

**Active Ingredients:** empagliflozin and metformin HCl

**Inactive Ingredients:** Tablet core contains: hypromellose, magnesium stearate, and polyethylene oxide. The film coatings and printing ink contain: carnauba wax, FD&C blue#2/indigo carmine aluminum lake (12.5 mg/1,000 mg, 25 mg/1,000 mg), ferric oxide red (10 mg/1,000 mg), ferric oxide yellow (5 mg/1,000 mg, 10 mg/1,000 mg, 25 mg/1,000 mg), ferrousferrous oxide, hypromellose, isopropyl alcohol, polydextrose, polyethylene glycol, propylene glycol, purified water, talc, and titanium dioxide.

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For more information about SYNJARDY or SYNJARDY XR, including current prescribing information and Medication Guide, go to [www.synjardy.com](http://www.synjardy.com) or [www.synjardyxr.com](http://www.synjardyxr.com), scan the code, or call Boehringer Ingelheim Pharmaceuticals, Inc. at 1-800-542-6257.



This Medication Guide has been approved by the U.S. Food and Drug Administration.

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