

MEDICATION GUIDE
KOMBIGLYZE XR (kom-be-glyze X-R)
(saxagliptin and metformin HCl extended-release)
tablets

Read this Medication Guide carefully before you start taking KOMBIGLYZE XR and each time you get a refill. There may be new information. This information does not take the place of talking with your healthcare provider about your medical condition or treatment. If you have any questions about KOMBIGLYZE XR, ask your healthcare provider.

What is the most important information I should know about KOMBIGLYZE XR?

Serious side effects can happen in people taking KOMBIGLYZE XR, including:

1. **Lactic Acidosis.** Metformin hydrochloride, one of the medicines in KOMBIGLYZE XR, can cause a rare, but serious, side effect 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 KOMBIGLYZE XR and call your healthcare provider right away if you get any of the following symptoms of lactic acidosis:

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

You have a higher chance of getting lactic acidosis if you:

- have kidney problems. People whose kidneys are not working properly should not take KOMBIGLYZE XR.
- have liver problems.

- have congestive heart failure that requires treatment with medicines.
 - 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.
 - have a heart attack, severe infection, or stroke.
 - are 80 years of age or older and have not had your kidney function tested.
2. **Inflammation of the pancreas (pancreatitis)** which may be severe and lead to death.

Certain medical problems make you more likely to get pancreatitis.

Before you start taking KOMBIGLYZE XR:

Tell your healthcare provider if you have ever had

- inflammation of your pancreas (pancreatitis)
- stones in your gallbladder (gallstones)
- a history of alcoholism
- high blood triglyceride levels

It is not known if having these medical problems will make you more likely to get pancreatitis with KOMBIGLYZE XR.

Stop taking KOMBIGLYZE XR and contact your healthcare provider right away if you have pain in your stomach area (abdomen) that is severe and will not go away. The pain may be felt going from your abdomen through to your back. The pain may happen with or without vomiting. These may be symptoms of pancreatitis.

What is KOMBIGLYZE XR?

- KOMBIGLYZE XR is a prescription medicine that contains saxagliptin and metformin hydrochloride. KOMBIGLYZE XR is used with diet and exercise to help control high blood sugar (hyperglycemia) in adults with type 2 diabetes.
- KOMBIGLYZE XR is not for people with type 1 diabetes.
- KOMBIGLYZE XR is not for people with diabetic ketoacidosis (increased ketones in your blood or urine).

- If you have had inflammation of the pancreas (pancreatitis) in the past, it is not known if you have a higher chance of getting pancreatitis while you take KOMBIGLYZE XR.

It is not known if KOMBIGLYZE XR is safe and effective in children younger than 18 years old.

Who should not take KOMBIGLYZE XR?

Do not take KOMBIGLYZE XR if you:

- have kidney problems.
- are allergic to metformin hydrochloride, saxagliptin, or any of the ingredients in KOMBIGLYZE XR. See the end of this Medication Guide for a complete list of ingredients in KOMBIGLYZE XR.

Symptoms of a serious allergic reaction to KOMBIGLYZE XR may include:

- swelling of your face, lips, throat, and other areas on your skin
- difficulty with swallowing or breathing
- raised, red areas on your skin (hives)
- skin rash, itching, flaking, or peeling

If you have these symptoms, stop taking KOMBIGLYZE XR and contact your healthcare provider right away.

- have a condition called metabolic acidosis or diabetic ketoacidosis (increased ketones in your blood or urine).

What should I tell my healthcare provider before taking KOMBIGLYZE XR?

Before you take KOMBIGLYZE XR, tell your healthcare provider if you:

- have type 1 diabetes. KOMBIGLYZE XR should not be used to treat type 1 diabetes.
- have a history or risk for diabetic ketoacidosis (high levels of certain acids, known as ketones, in the blood or urine). KOMBIGLYZE XR should not be used for the treatment of diabetic ketoacidosis.
- have kidney problems.
- have liver problems.
- have heart problems, including congestive heart failure.
- are older than 80 years. If you are over 80 years old you should not take KOMBIGLYZE XR unless your kidneys have been checked and they are normal.

- drink alcohol very often, or drink a lot of alcohol in short-term “binge” drinking.
- are taking insulin.
- are going to get an injection of dye or contrast agents for an x-ray procedure or if you are going to have surgery and will not be able to eat or drink much. In these situations, KOMBIGLYZE XR will need to be stopped for a short time. Talk to your healthcare provider about when you should stop KOMBIGLYZE XR and when you should start KOMBIGLYZE XR again. See **“What is the most important information I should know about KOMBIGLYZE XR?”**
- have any other medical conditions.
- are pregnant or plan to become pregnant. It is not known if KOMBIGLYZE XR will harm your unborn baby. If you are pregnant, talk with your healthcare provider about the best way to control your blood sugar while you are pregnant.
- are breast-feeding or plan to breast-feed. It is not known if KOMBIGLYZE XR passes into your breast milk. Talk with your healthcare provider about the best way to feed your baby while you take KOMBIGLYZE XR.

Tell your healthcare provider about all the medicines you take, including prescription and nonprescription medicines, vitamins, and herbal supplements. Know the medicines you take. Keep a list of them to show your healthcare provider and pharmacist when you get a new medicine.

KOMBIGLYZE XR may affect the way other medicines work, and other medicines may affect how KOMBIGLYZE XR works.

Tell your healthcare provider if you will be starting or stopping certain other types of medicines, such as antibiotics, or medicines that treat fungus or HIV/AIDS, because your dose of KOMBIGLYZE XR might need to be changed.

How should I take KOMBIGLYZE XR?

- Take KOMBIGLYZE XR exactly as your healthcare provider tells you.
- KOMBIGLYZE XR should be taken with meals to help lessen an upset stomach side effect.
- Swallow KOMBIGLYZE XR whole. Do not crush, cut, or chew KOMBIGLYZE XR.
- You may sometimes pass a soft mass in your stools (bowel movement) that looks like KOMBIGLYZE XR tablets.
- 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 that you need may change. Tell your healthcare provider right away if you have any of these problems.

- Your healthcare provider should do blood tests to check how well your kidneys are working before and during your treatment with KOMBIGLYZE XR.
- Your healthcare provider will check your diabetes with regular blood tests, including your blood sugar levels and your hemoglobin A1C.
- Follow your healthcare provider's instructions for treating blood sugar that is too low (hypoglycemia). Talk to your healthcare provider if low blood sugar is a problem for you. See **"What are the possible side effects of KOMBIGLYZE XR?"**
- Check your blood sugar as your healthcare provider tells you to.
- Stay on your prescribed diet and exercise program while taking KOMBIGLYZE XR.
- If you miss a dose of KOMBIGLYZE XR, take your next dose as prescribed unless your healthcare provider tells you differently. Do not take an extra dose the next day.
- If you take too much KOMBIGLYZE XR, call your healthcare provider, local Poison Control Center, or go to the nearest hospital emergency room right away.

What are the possible side effects of KOMBIGLYZE XR?

KOMBIGLYZE XR can cause serious side effects, including:

- See **"What is the most important information I should know about KOMBIGLYZE XR?"**
- **Allergic (hypersensitivity) reactions**, such as:
 - swelling of your face, lips, throat, and other areas on your skin
 - difficulty with swallowing or breathing
 - raised, red areas on your skin (hives)
 - skin rash, itching, flaking, or peeling

If you have these symptoms, stop taking KOMBIGLYZE XR and contact your healthcare provider right away.

- **Low blood sugar (hypoglycemia).** If you take KOMBIGLYZE XR with another medicine that can cause low blood sugar, such as sulfonylureas, you have a higher chance of having low blood sugar. Tell your healthcare provider if you take other diabetes medicines. If you have symptoms of low blood sugar, you should check your blood sugar and treat if low, then call your healthcare provider. Symptoms of low blood sugar include:
 - shaking
 - sweating
 - rapid heartbeat
 - change in vision

- hunger
- headache
- change in mood

Common side effects of KOMBIGLYZE XR include:

- upper respiratory tract infection
- stuffy or runny nose and sore throat
- urinary tract infection
- headache
- diarrhea
- nausea and vomiting

Taking KOMBIGLYZE XR with meals can help lessen the common stomach side effects of metformin. If you have unexplained stomach problems, tell your healthcare provider. Stomach problems that start later during treatment may be a sign of something more serious.

Tell your healthcare provider if you have any side effects that bother you or that do not go away.

These are not all of the possible side effects of KOMBIGLYZE 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 the FDA at 1-800-FDA-1088.

How should I store KOMBIGLYZE XR?

Store KOMBIGLYZE XR between 68°F and 77°F (20°C and 25°C).

Keep KOMBIGLYZE XR and all medicines out of the reach of children.

General information about the use of KOMBIGLYZE XR

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

This Medication Guide summarizes the most important information about KOMBIGLYZE XR. If you would like more information, talk with your healthcare provider. You can ask your pharmacist or healthcare provider for information about KOMBIGLYZE XR that is written for healthcare professionals.

For more information, go to www.kombiglyzexr.com or call 1-800-664-5992.

What are the ingredients of KOMBIGLYZE XR?

Active ingredients: saxagliptin and metformin hydrochloride.

Inactive ingredients in each tablet: carboxymethylcellulose sodium, hypromellose 2208, and magnesium stearate.

The 5 mg/500 mg tablet also contains: microcrystalline cellulose and hypromellose 2910.

Tablet film coat contains: polyvinyl alcohol, polyethylene glycol 3350, titanium dioxide, talc, and iron oxides.

What is type 2 diabetes?

Type 2 diabetes is a condition in which your body does not make enough insulin, and the insulin that your body produces does not work as well as it should. Your body can also make too much sugar. When this happens, sugar (glucose) builds up in the blood. This can lead to serious medical problems.

The main goal of treating diabetes is to lower your blood sugar so that it is as close to normal as possible. High blood sugar can be lowered by diet and exercise, and by certain medicines when necessary.

Talk to your healthcare provider about how to prevent, recognize, and take care of low blood sugar (hypoglycemia), high blood sugar (hyperglycemia), and problems you have because of your diabetes.

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

KOMBIGLYZE XR (saxagliptin and metformin HCl extended-release) tablets

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