

READ THIS FOR SAFE AND EFFECTIVE USE OF YOUR MEDICINE
PATIENT MEDICATION INFORMATION

LORBRENA™
Lorlatinib tablets

Read this carefully before you start taking LORBRENA and each time you get a refill. This leaflet is a summary and will not tell you everything about this drug. Talk to your healthcare professional about your medical condition and treatment and ask if there is any new information about LORBRENA

What is LORBRENA used for?

LORBRENA is used to treat adult patients with a type of lung cancer called non-small cell lung cancer (NSCLC). It is used in a special type of NSCLC that is anaplastic lymphoma kinase (ALK)-positive. LORBRENA is used if your cancer has spread to other parts of your body and:

- your cancer has gotten worse after taking crizotinib and at least one other ALK tyrosine kinase inhibitor (TKI) medicine or,
- your cancer has gotten worse on ceritinib or alectinib.

Read the boxed message below for more information.

LORBRENA is not approved for use in children.

For the following indication LORBRENA has been approved *with conditions* (NOC/c). This means it has passed Health Canada's review and can be bought and sold in Canada, but the manufacturer has agreed to complete more studies to make sure the drug works the way it should. For more information, talk to your healthcare professional.

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What is a Notice of Compliance with Conditions (NOC/c)?

A Notice of Compliance with Conditions (NOC/c) is a type of approval to sell a drug in Canada.

Health Canada only gives an NOC/c to a drug that treats, prevents, or helps identify a serious or life-threatening illness. The drug must show promising proof that it works well, is of high quality, and is reasonably safe. Also, the drug must either respond to a serious medical need in Canada, or be much safer than existing treatments.

Drug makers must agree in writing to clearly state on the label that the drug was given an NOC/c, to complete more testing to make sure the drug works the way it should, to actively monitor the drug's performance after it has been sold, and to report their findings to Health Canada.

Serious Warnings and Precautions

LORBRENA should only be prescribed by a healthcare professional experienced in the use of anti-cancer drugs.

LORBRENA can cause serious side effects which may include:

- **High blood lipid levels (cholesterols or triglycerides):** LORBRENA can cause your blood lipid levels to increase. Your healthcare professional will do regular blood tests while you are taking LORBRENA to check your blood lipid levels.
- **Lung problems:** LORBRENA can cause severe or life-threatening swelling (inflammation) of the lungs that can lead to death. Symptoms may be similar to those from lung cancer. Tell your healthcare professional immediately if you have any new or worsening symptoms of lung problems, including trouble breathing, shortness of breath, cough, or fever.
- **Liver problems:** LORBRENA can cause serious liver problems if it is taken with other medicines. Tell your healthcare professional about all the other medicines you take. While you are taking LORBRENA, if you experience yellowing of the skin or eyes, dark urine, abdominal pain, nausea, vomiting and/or loss of appetite, contact your healthcare professional immediately.

How does LORBRENA work?

LORBRENA belongs to a group of anti-cancer medicines called ALK tyrosine kinase inhibitors (TKI). It blocks the action of an enzyme called 'ALK tyrosine kinase'. By blocking this enzyme LORBRENA may slow down or stop the growth of your cancer. It may also help to shrink your cancer.

If you have any questions about how LORBRENA works or why this medicine has been prescribed for you, ask your healthcare professional.

What are the ingredients in LORBRENA?

Medicinal ingredients: lorlatinib

Non-medicinal ingredients: dibasic calcium phosphate anhydrous, ferrousferic oxide/black iron oxide, hydroxypropyl methylcellulose/hypromellose, iron oxide red, lactose monohydrate, macrogol/polyethylene glycol, magnesium stearate, microcrystalline cellulose, sodium starch glycolate, titanium dioxide, triacetin

LORBRENA comes in the following dosage forms:

Tablets, 25 mg, 100 mg

Do not use LORBRENA if:

- you are allergic to lorlatinib or any of the other ingredients of LORBRENA;
- you are taking any of these medicines:
 - rifampicin (used to treat tuberculosis);
 - carbamazepine, phenytoin (used to treat epilepsy);
 - enzalutamide (used to treat prostate cancer);
 - mitotane (used to treat cancer of the adrenal glands);
 - medicines containing St. John's wort (*Hypericum perforatum*, a herbal preparation).

To help avoid side effects and ensure proper use, talk to your healthcare professional before you take LORBRENA. Talk about any health conditions or problems you may have, including if you:

- have high blood lipid levels (cholesterol or triglycerides);
- have heart problems;
- have lung problems;
- have liver problems;
- have any other medical conditions;
- are pregnant, or plan to become pregnant. You should not get pregnant or father a child while you are taking LORBRENA. LORBRENA can harm your unborn baby.
 - **Females** who are able to become pregnant must use effective, non-hormonal birth control during treatment with LORBRENA and for at least 21 days after the final dose of LORBRENA. If hormonal birth control use cannot be avoided, use condoms in addition to hormonal birth control for at least 21 days after the final dose of LORBRENA.
 - **Males** who have pregnant partners or female partners who can become pregnant must use condoms during treatment with LORBRENA and for at least 97 days after the final dose of LORBRENA.
 - Talk to your healthcare professional about birth control methods that may be right for you.
 - If you or your partner becomes pregnant, tell your healthcare professional right away.
 - LORBRENA can cause decreased fertility in both males and females. If you may want to become pregnant or father a child after treatment with LORBRENA talk to your healthcare professional about fertility preservation options that may be right for you.
- are breastfeeding or plan to breastfeed. It is not known if LORBRENA passes into your breast milk. Do not breastfeed during treatment with LORBRENA and for 7 days after the final dose. Talk to your healthcare professional about the best way to feed your baby during this time.

Other warnings you should know about:

- Your healthcare professional will test your cancer before you start taking LORBRENA to make sure it is ALK-positive.
- **High blood lipids levels (cholesterol or triglycerides):**
 - Your healthcare professional will do a blood test to check your blood lipid levels before you start taking LORBRENA. Once you start taking LORBRENA your healthcare professional will do blood tests after 2 weeks, 4 week, and 8 weeks. Your healthcare professional may also do blood tests at other times during your treatment.
 - If your blood lipid levels increase while you are taking LORBRENA, your healthcare professional may need to start you on a lipid-lowering medicine to lower the levels.

- If you are already taking a lipid-lowering medicine, your healthcare professional may need to increase your dose of that medicine.
- **Serious lung problems:** LORBRENA can cause severe or life-threatening swelling (inflammation) of the lungs that can lead to death. Symptoms may be similar to those from lung cancer. Tell your healthcare professional immediately if you have any new or worsening symptoms of lung problems, including trouble breathing, shortness of breath, cough, or fever.
- **Serious liver problems:** LORBRENA can cause serious liver problems if it is taken with other medicines. Tell your healthcare professional about all the other medicines you take. While you are taking LORBRENA, if you experience yellowing of the skin or eyes, dark urine, abdominal pain, nausea, vomiting and/or loss of appetite, contact your healthcare professional immediately.
- **Mental status changes, speech problems and mental health problems:** LORBRENA can cause problems with thinking (such as forgetfulness or confusion), trouble with speech, mood changes and hallucinations (seeing and hearing things that are not real). If you experience any problems with thinking, trouble with your speech, changes in your mood or hallucinations while you are taking LORBRENA, contact your healthcare professional immediately. Your healthcare professional may change your dose of LORBRENA if these symptoms occur. If these symptoms are severe, your healthcare professional may tell you to stop taking LORBRENA.
- **Heart problems:** LORBRENA may cause very slow or abnormal heartbeats. Your healthcare professional may need to check your heart closely while you are taking LORBRENA. Tell your healthcare professional right away if you feel dizzy or faint or have abnormal heartbeats. If you have these symptoms, your healthcare professional may need to change your dose of LORBRENA.

Driving and using machines

LORBRENA can affect your ability to drive and use machines. Avoid driving or using machinery until you know how LORBRENA affects you.

Tell your healthcare professional about all the medicines you take, including any drugs, vitamins, minerals, natural supplements or alternative medicines.

The following may interact with LORBRENA:

- boceprevir, telaprevir, medicines used to treat hepatitis C;
- conivaptan, a medicine used to increase sodium levels in hospitalized patients;
- efavirenz, cobicistat, ritonavir, paritaprevir in combination with ritonavir and ombitasvir and/or dasabuvir, and ritonavir in combination with either danoprevir, elvitegravir, indinavir, lopinavir, saquinavir or tipranavir, medicines used to treat AIDS/HIV;
- ketoconazole, itraconazole, voriconazole, posaconazole, medicines used to treat fungal infections. Also troleandomycin, a medicine used to treat certain types of bacterial infections;
- quinidine, a medicine used to treat irregular heartbeat and other heart problems;
- pimozide, a medicine used to treat mental health problems;
- alfentanil and fentanyl, medicines used to treat severe pain;
- hormonal contraceptives

- ciclosporin, sirolimus, and tacrolimus, medicines used in organ transplantation to prevent transplant organ rejection;
- rifampicin, a medicine used to treat tuberculosis;
- carbamazepine, phenytoin, medicines used to treat epilepsy;
- enzalutamide, a medicine used to treat prostate cancer;
- mitotane, a medicine used to treat cancer of the adrenal glands;
- medicines containing St. John's wort (*Hypericum perforatum*, a herbal preparation);
- grapefruit juice or any products containing grapefruit juice.

How to take LORBRENA:

- Take LORBRENA exactly as your healthcare professional tells you.
- Do not change your dose or stop taking LORBRENA unless your healthcare professional tells you to.
- Swallow LORBRENA tablets whole. Do not chew, crush or split LORBRENA tablets before swallowing them.
- You may take LORBRENA with or without food.
- You should not eat or drink grapefruit products during your treatment with LORBRENA. It may increase the amount of LORBRENA in your blood to a harmful level.
- If you vomit after taking a dose of LORBRENA, do not take an extra dose; just take your next dose at your regular time.

Usual adult dose:

The recommended dose is 100 mg taken orally once daily.

If you have side effects, your healthcare professional may need to change your dose, temporarily stop, or completely stop your treatment with LORBRENA.

Overdose:

If you think you have taken too much LORBRENA contact your healthcare professional, hospital emergency department or regional poison control centre immediately, even if there are no symptoms.

Missed Dose:

If you miss a dose, take it as soon as you remember. If it is close to your next dose (within 4 hours), just take your next dose at your regular time. Do not take two doses at the same time to make up for a missed dose.

What are possible side effects from using LORBRENA?

These are not all the possible side effects you may feel when taking LORBRENA. If you experience any side effects not listed here, contact your healthcare professional.

The most common side effects of LORBRENA include:

- feeling of numbness or pins and needles in the joints, arms or legs (peripheral neuropathy);
- tiredness (fatigue);
- weight gain;
- pain in your joints;
- muscle pain, back pain, pain in your arms or legs;

- diarrhea;
- nausea, vomiting;
- headache;
- dizziness;
- rash.

LORBRENA can cause abnormal blood test results, including high blood lipid levels. Your healthcare professional will decide when to perform blood tests and will interpret the results.

Serious side effects and what to do about them			
Symptom / effect	Talk to your healthcare professional		Stop taking drug and get immediate medical help
	Only if severe	In all cases	
VERY COMMON Changes in mental status and speech problems: confusion, memory loss, trouble with attention, difficulty speaking, such as slurred or slow speech		X	
Mental health problems: changes in mood, irritability, agitation, mood swings, anxiety, depression, hallucinations (seeing or hearing things that aren't real)		X	
Edema: swelling of the legs, ankles, feet and hands		X	
Vision problems: double vision, sensitivity to light, blurred vision, vision loss, floaters, flashes of light		X	
Liver problems if it is taken with other medicines: yellowing of the skin or eyes, dark urine, abdominal pain, nausea, vomiting, loss of appetite		X	
COMMON Lung problems: new or worsening lung problems, trouble breathing, shortness of breath, cough, or fever		X	
Heart problems: feel dizzy or faint or have very slow or abnormal heartbeats		X	

If you have a troublesome symptom or side effect that is not listed here or becomes bad enough to interfere with your daily activities, talk to your healthcare professional.

Reporting Side Effects

You can report any suspected side effects associated with the use of health products to Health Canada by:

- Visiting the Web page on [Adverse Reaction Reporting](https://www.canada.ca/en/health-canada/services/drugs-health-products/medeffect-canada/adverse-reaction-reporting.html) (<https://www.canada.ca/en/health-canada/services/drugs-health-products/medeffect-canada/adverse-reaction-reporting.html>) for information on how to report online, by mail or by fax; or
- Calling toll-free at 1-866-234-2345.

NOTE: Contact your healthcare professional if you need information about how to manage your side effects. The Canada Vigilance Program does not provide medical advice.

Storage:

Store at 15°C to 30°C in the original package to protect from light.

Keep out of reach and sight of children.

If you want more information about LORBRENA:

- Talk to your healthcare professional
- Find the full product monograph that is prepared for healthcare professionals and includes this Patient Medication Information by visiting the [Health Canada website](http://www.canada.ca/en/health-canada.html) (<http://www.canada.ca/en/health-canada.html>); the manufacturer's website (<http://www.Pfizer.ca>), or by calling 1-800-463-6001.

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