

# STARTING TREATMENT WITH LORBRENA® (lorlatinib)

## Guide to common side effects



LORBRENA is a prescription medicine that is used to treat adults with non-small cell lung cancer (NSCLC) that is caused by an abnormal anaplastic lymphoma kinase (ALK) gene and that has spread to other parts of your body.

Your healthcare provider will perform a test to make sure that LORBRENA is right for you.

It is not known if LORBRENA is safe and effective in children.

### LORBRENA CLINICAL TRIALS

#### Newly diagnosed patients

LORBRENA was evaluated in a clinical trial known as the CROWN trial. The trial studied 296 patients with ALK+ NSCLC whose cancer had spread to other parts of the body. The study included patients with and without cancer that had spread to their brain. Patients in the trial were given one of two medicines, LORBRENA or crizotinib, which is a different treatment for ALK+ NSCLC. All patients were newly diagnosed at enrollment and had not received any previous treatment for NSCLC. Patients with severe mental health conditions were not included in the trial.

#### Previously treated patients

LORBRENA was also evaluated in a trial that studied 215 patients with ALK+ NSCLC that had spread to other parts of the body, including the brain. Their tumors were no longer responding to certain other medicines for ALK+ NSCLC. The patients had previously been treated with at least one other medicine for ALK+ NSCLC. Forty-three percent had been treated with one prior medicine, 35% with two prior medicines, and 22% had been treated with three prior medicines for ALK+ NSCLC before they enrolled in the trial. At the start of the trial, 69% of patients had tumors that had spread to the brain.

To learn more about the LORBRENA clinical trials, visit [lorbrena.com/clinical-study-results](http://lorbrena.com/clinical-study-results).

### SELECTED SAFETY INFORMATION

#### LORBRENA may cause serious side effects, including:

- **Liver problems due to interactions with other medicines.** It is important to know what medicines should not be taken with LORBRENA.
- **Central nervous system (CNS) effects.** LORBRENA may cause CNS effects, including problems with thinking (such as forgetfulness or confusion), changes in mood (such as depression and thoughts about suicide or dying), psychotic effects such as seeing or hearing things that are not real (hallucinations), seizures, changes in speech and changes in sleep. Tell your healthcare provider if you experience new or worsening symptoms of CNS effects.

These are not all of the possible side effects of LORBRENA.

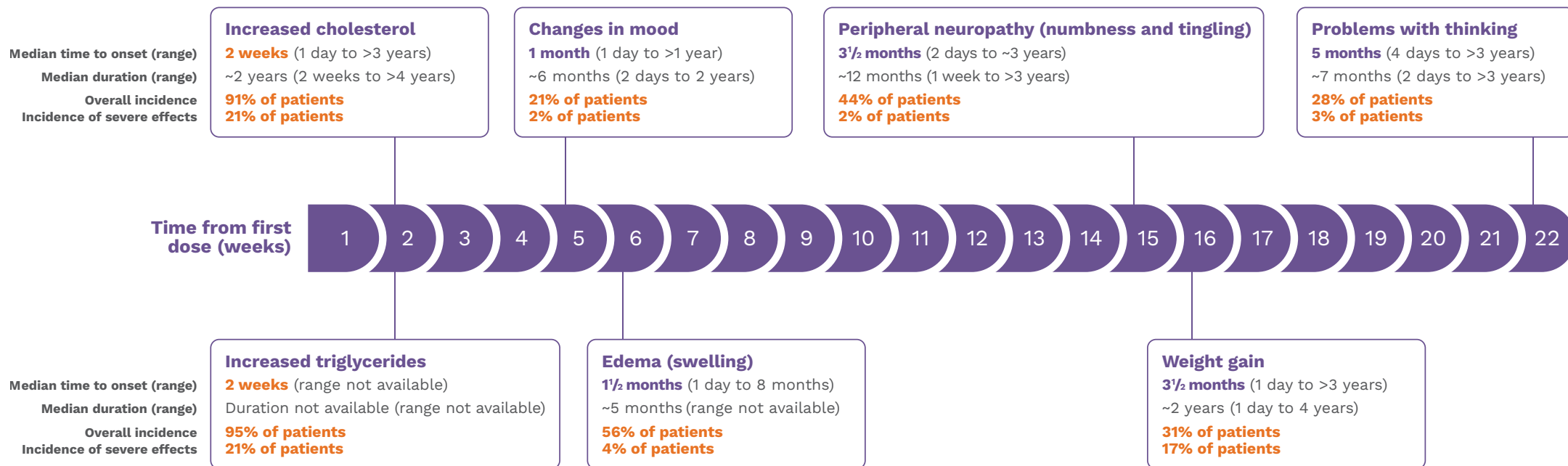
For more information, ask your healthcare provider or pharmacist.

Please see *Important Safety Information* on pages 4-5. Click for the [Full Prescribing Information](#) and [Patient Information](#) or visit [LORBRENA.com](http://LORBRENA.com).

# WHAT ARE THE MOST COMMON SIDE EFFECTS OF LORBRENA® (lorlatinib) AND WHEN ARE THEY LIKELY TO HAPPEN?

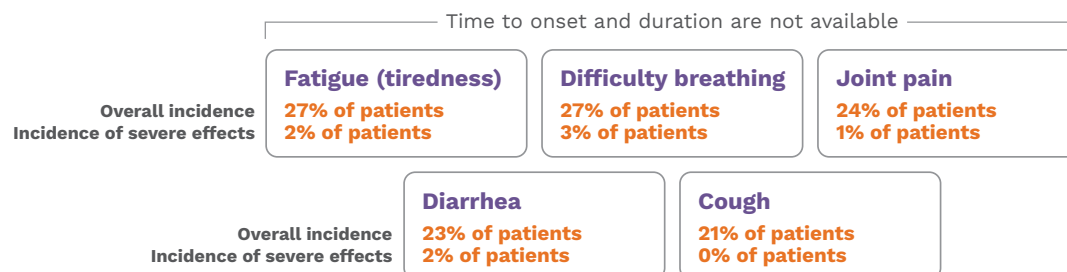
The side effects discussed on this page are not all of the possible side effects of LORBRENA.

All data included below are from what patients experienced in the LORBRENA clinical trials. Refer to page 5 for Important Safety Information. You may or may not experience the side effects described below. If you do experience these side effects, you may not experience them in the time frame or for the duration described.



## UNDERSTANDING THE TIMELINE

- Data that are from the LORBRENA Prescribing Information are in orange; data from published medical journal articles are in black
  - Median is a statistics term. It is the middle value in a set of measurements
  - Changes in mood include, for example, depression and irritability
  - Problems with thinking include, for example, forgetfulness and confusion
  - Severe side effects are medically significant and may mean that you will have to go to the hospital or stay longer in the hospital. They can make daily self-care difficult. Some severe side effects are life-threatening and should be treated right away
- In some patients in the CROWN trial, LORBRENA dosing was adjusted or stopped to manage side effects
- LORBRENA was stopped for a period of time in 49% of patients, the LORBRENA dose was reduced in 21% of patients, and LORBRENA was completely stopped in 7% of patients



To learn more about what to expect while taking LORBRENA, visit [www.LORBRENA.com/what-to-expect](http://www.LORBRENA.com/what-to-expect)



Please see Important Safety Information on pages 4-5. Click for the Full Prescribing Information and Patient Information or visit [LORBRENA.com](http://LORBRENA.com).

# TIPS TO HELP MANAGE SIDE EFFECTS

## Key points to remember



### For high cholesterol or triglycerides (lipids)

Your healthcare provider may need to start you on lipid-lowering medicines (usually a certain statin). If you are already taking lipid-lowering medicines, your healthcare provider may increase the dose of the lipid-lowering medicine or prescribe a different medicine.



### For changes in mood or problems with thinking

Your healthcare provider may adjust your LORBRENA® (lorlatinib) dose or, in some cases, may have you stop taking LORBRENA. In the clinical trial, these side effects were generally reversible by adjusting or withholding the dose of LORBRENA.



### For swelling (edema)

Let your healthcare provider know if you experience swelling in your arms, legs, hands, or feet. Your healthcare provider may adjust your dose of LORBRENA or prescribe a diuretic, which decreases the amount of water in your body through urination. You can also ask your healthcare provider about leg elevation, compression stockings, and lifestyle changes (such as exercising and limiting salt intake).



### For numbness or tingling (peripheral neuropathy)

If you experience feelings of numbness or tingling in the joints or arms and legs, your healthcare provider may start you on a medicine to treat it or change the dose of LORBRENA.



### For weight gain

Dietary advice from a nutritionist and routine exercise may be effective weight management strategies. Speak with your healthcare provider if you are experiencing weight gain.

**If you think you may be experiencing side effects while taking LORBRENA, it's important to tell your healthcare provider right away.** He or she may pause or adjust your LORBRENA dose or suggest additional medicine to take while you are taking LORBRENA. **Do not change your dose or stop taking LORBRENA unless your healthcare provider tells you to.**

# IMPORTANT SAFETY INFORMATION

## LORBRENA® (lorlatinib) may cause serious side effects, including:

- **Liver problems due to interactions with other medicines.** It is important to know what medicines should not be taken with LORBRENA.
- **Central nervous system (CNS) effects.** LORBRENA may cause CNS effects, including problems with thinking (such as forgetfulness or confusion), changes in mood (such as depression and thoughts about suicide or dying), psychotic effects such as seeing or hearing things that are not real (hallucinations), seizures, changes in speech and changes in sleep. Tell your healthcare provider if you experience new or worsening symptoms of CNS effects.
- **Increases in the cholesterol and triglycerides (lipid) levels in your blood.** Most people will have an increase in the lipid levels in their blood during treatment with LORBRENA.
  - o If you have increases in the lipid levels in your blood during treatment with LORBRENA, your healthcare provider may need to start you on a medicine to lower the levels. If you are already taking a medicine to lower the lipid levels in your blood, your healthcare provider may need to increase your dose of that medicine.
  - o Your healthcare provider should do blood tests to check the lipid levels in your blood before starting treatment, 1 to 2 months after starting treatment, and during treatment with LORBRENA.
- **Heart problems.** LORBRENA may cause very slow or abnormal heartbeats. Your healthcare provider should check your heart rhythm (electrocardiogram or EKG) before starting and during treatment with LORBRENA. Tell your healthcare provider right away if you feel dizzy or faint or have abnormal heartbeats. In some people, these problems are severe, and your healthcare provider may need to have you stop taking LORBRENA or have a pacemaker placed.
- **Lung problems.** LORBRENA may cause severe or life-threatening swelling (inflammation) of the lungs during treatment that can lead to death. Symptoms may be similar to those from lung cancer. Tell your healthcare provider right away if you have any new or worsening symptoms of lung problems, including trouble breathing, shortness of breath, cough, or fever.
- **High blood pressure (hypertension).** Your healthcare provider should check your blood pressure before starting treatment, 2 weeks after starting treatment, and then at least every month during treatment with LORBRENA. Your healthcare provider may need to start or change your blood pressure medicine if you have high blood pressure during treatment with LORBRENA. Tell your healthcare provider right away if you get signs or symptoms of high blood pressure, including headaches, dizziness, blurred vision, chest pain or shortness of breath.
- **High blood sugar (hyperglycemia).** LORBRENA may increase your blood sugar levels. Your healthcare provider should do blood tests to check your blood sugar levels before starting and during treatment with LORBRENA. Your healthcare provider may need to start or change your blood sugar medicine to control your blood sugar levels. Tell your healthcare provider right away if you get new or worsening signs and symptoms of high blood sugar, including feeling very thirsty, needing to urinate more than usual, or feeling very hungry, sick to your stomach, weak or tired, or confused.

If you have serious side effects during treatment with LORBRENA, your healthcare provider may change your dose, stop your treatment for a period of time, or completely stop treatment with LORBRENA.

## Before taking LORBRENA, tell your healthcare provider about all of your medical conditions, including if you:

- have kidney problems
- have had episodes of depression or seizures
- have high levels of cholesterol or triglycerides in your blood
- have problems with your heartbeat
- have lung or breathing problems
- have high blood pressure
- have diabetes or high blood sugar

(Continued on page 5)

Click for the [Full Prescribing Information](#) and [Patient Information](#) or visit [LORBRENA.com](http://LORBRENA.com).

# IMPORTANT SAFETY INFORMATION (CONTINUED)

- are pregnant or plan to become pregnant. LORBRENA® (lorlatinib) can harm your unborn baby.
  - Your healthcare provider will do a pregnancy test before you start treatment with LORBRENA.
  - Tell your healthcare provider right away if you become pregnant or think you may be pregnant during treatment with LORBRENA.
    - **Females** who are able to become pregnant should use effective non-hormonal birth control during treatment with LORBRENA and for at least 6 months after the final dose of LORBRENA. Birth control pills (oral contraceptives) and other hormonal forms of birth control may not be effective if used during treatment with LORBRENA. Talk to your healthcare provider about birth control choices that are right for you during this time.
    - **Males** who have female partners who are able to become pregnant should use effective birth control during treatment with LORBRENA and for at least 3 months after the final dose of LORBRENA.
- 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 provider about the best way to feed your baby during this time.

**Tell your healthcare provider about all the medicines you take, including** prescription medicines, over-the-counter medicines, vitamins, and herbal supplements. LORBRENA may affect the way other medicines work and other medicines may affect the way LORBRENA works, causing side effects.

Know the medicines you take. Keep a list of them to show to your healthcare provider and pharmacist when you get a new medicine.

**Do not take LORBRENA if you** take certain other medicines called strong CYP3A inducers. Ask your healthcare provider for a list of these medicines if you are not sure.

**The most common side effects of LORBRENA include:**

- swelling in your arms, legs, hands, and feet (edema)
- numbness and tingling feeling in your joints or arms and legs (peripheral neuropathy)
- weight gain
- problems with thinking, such as forgetfulness or confusion
- tiredness (fatigue)
- difficulty breathing
- pain in your joints
- diarrhea
- changes in mood, such as depression and irritability
- high cholesterol and triglyceride levels in the blood
- cough

LORBRENA may cause decreased fertility in males. In males, this could affect your ability to father a child. Talk to your healthcare provider if you have concerns about fertility.

These are not all of the possible side effects of LORBRENA. 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.

Click for the *Full Prescribing Information and Patient Information* or visit [LORBRENA.com](https://www.lorbrena.com).

**LORBRENA**  
LORLATINIB | 100 mg tablets