WHAT IS ORKAMBI® (lumacaftor/ivacaftor)?

ORKAMBI is a prescription medicine used for the treatment of cystic fibrosis (CF) in patients age 2 years and older who have two copies of the F508del mutation (F508del/F508del) in their CFTR gene.

ORKAMBI should not be used in patients other than those who have two copies of the F508del mutation in their CFTR gene.

It is not known if ORKAMBI is safe and effective in children under 2 years of age.

All photos are of people with CF who have 2 copies of the F508del mutation in their CFTR gene or their caregivers. The people with CF may or may not be taking ORKAMBI.

Please see Important Safety Information and full Prescribing Information, including Patient Information.
IMPORTANT SAFETY INFORMATION

Your child should not take ORKAMBI® if he or she takes certain medicines or herbal supplements, such as:

- antibiotics: rifampin (Rifamate®, Rifater®) or rifabutin (Mycobutin®)
- seizure medicines: phenobarbital, carbamazepine (Tegretol®, Carbatrol®, and Equetro®), or phenytoin (Dilantin®, Phenytek®)
- sedatives and anti-anxiety medicines: triazolam (Halcion®) or midazolam (Dormicum®, Hypnovel®, and Versed®)
- immunosuppressant medicines: cyclosporine, everolimus (Zortress®), sirolimus (Rapamune®), or tacrolimus (Astagraf XL®, Envarsus XR®, Prograf®, and Protopic®)
- St. John’s wort (Hypericum perforatum)

Talk to your child’s doctor before taking ORKAMBI if he or she takes any of the medicines or supplements listed above.

Before taking ORKAMBI, tell your child’s doctor about all of your child’s medical conditions, including if he or she:

- has or has had liver problems
- has had an organ transplant
- has kidney problems
- is using birth control (hormonal contraceptives, including oral, injectable, transdermal, or implantable forms). Hormonal contraceptives should not be used as a method of birth control when taking ORKAMBI. Talk to the doctor about the best birth control method to use while taking ORKAMBI
- is pregnant or plans to become pregnant. It is not known if ORKAMBI will harm an unborn baby. Patients and their doctor should decide whether ORKAMBI should be taken during pregnancy
- is breastfeeding or planning to breastfeed. It is not known if ORKAMBI passes into breast milk. Patients and their doctor should decide whether ORKAMBI should be taken while breastfeeding

ORKAMBI® (lumacaftor/ivacaftor) may affect the way other medicines work, and other medicines may affect how ORKAMBI works.

Tell your child’s doctor about all the medicines he or she takes, including prescription and over-the-counter medicines, vitamins, and herbal supplements, because the dose of ORKAMBI may need to be adjusted when taken with certain medicines.

Especially tell your child’s doctor if he or she takes:

- antifungal medicines including ketoconazole (such as Nizoral®), itraconazole (such as Sporanox®), posaconazole (such as Noxafil®), or voriconazole (such as Vfend®)
- antibiotics including telithromycin (such as Ketek®), clarithromycin (such as Biaxin®), or erythromycin (such as Ery-Tab®)

Tell your child’s doctor if he or she stops ORKAMBI for more than 1 week. Your child’s doctor may need to change the dose of ORKAMBI or other medicines your child takes.

Please see additional Important Safety Information and full Prescribing Information, including Patient Information.
**IMPORTANT SAFETY INFORMATION (CONTINUED)**

**What are the possible side effects of ORKAMBI®?**

**ORKAMBI can cause serious side effects, including:**

- **Worsening of liver function** in people with severe liver disease. The worsening of liver function can be serious or cause death. Talk to your child’s doctor if you have been told that he or she has liver disease as your child’s doctor may need to adjust the dose of ORKAMBI.

- **High liver enzymes in the blood**, which can be a sign of liver injury in people receiving ORKAMBI. Your child’s doctor will do blood tests to check your child’s liver:
  - before starting ORKAMBI
  - every 3 months during the first year of taking ORKAMBI
  - every year while taking ORKAMBI

Call your child’s doctor right away if he or she has any of the following symptoms of liver problems:

- pain or discomfort in the upper right stomach (abdominal) area
- yellowing of the skin or the white part of the eyes
- loss of appetite
- nausea or vomiting
- dark, amber-colored urine
- confusion

- **Breathing problems** such as shortness of breath or chest tightness in patients when starting ORKAMBI, especially in patients who have poor lung function. If your child has poor lung function, your child’s doctor may monitor him or her more closely when starting ORKAMBI.

- **An increase in blood pressure** in some people receiving ORKAMBI. Your child’s doctor should monitor your child’s blood pressure during treatment with ORKAMBI.

- **Abnormality of the eye lens (cataract)** in some children and adolescents receiving ORKAMBI. Your child’s doctor should perform eye examinations before and during treatment with ORKAMBI to look for cataracts.

**The most common side effects of ORKAMBI® (lumacaftor/ivacaftor) include:**

- gas
- common cold, including sore throat, stuffy or runny nose
- flu or flu-like symptoms
- irregular, missed, or abnormal periods (menses) and increase in the amount of menstrual bleeding

**Side effects seen in children** are similar to those seen in adults and adolescents. Additional common side effects seen in children include:

- cough with sputum
- stuffy nose
- headache
- stomach pain
- increase in sputum

These are not all the possible side effects of ORKAMBI. Call your child’s doctor for medical advice about side effects. You are encouraged to report side effects to FDA at 1-800-FDA-1088.

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Please see additional Important Safety Information and full Prescribing Information, including Patient Information.
Your child should take ORKAMBI, plus all of your child’s other CF therapies, exactly as your child’s healthcare provider says they should be taken.

ORKAMBI oral granules: Designed with a young child in mind

ORKAMBI comes in single-use, oral granule packets for children age 2 through 5 years with cystic fibrosis (CF) who have 2 copies of the F508del-CFTR mutation. The oral granules are similar in size to flour particles and should be mixed with soft food or liquid.

ORKAMBI oral granules are prescribed based on weight and come in 2 different strengths

<table>
<thead>
<tr>
<th>WEIGHT-BASED DOSE</th>
<th>EACH DOSE</th>
<th>TOTAL DAILY DOSE</th>
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<tbody>
<tr>
<td>Less than ~31 lbs (less than 14 kg) lumacaftor 100 mg/ivacaftor 125 mg</td>
<td>1 packet every 12 hours</td>
<td>2 packets per day</td>
</tr>
<tr>
<td>≥31 lbs or more (14 kg or more) lumacaftor 150 mg/ivacaftor 188 mg</td>
<td>1 packet every 12 hours</td>
<td>2 packets per day</td>
</tr>
</tbody>
</table>

Remember, having your child take ORKAMBI every 12 hours is not the same as having him or her take it twice a day.

Your child should take ORKAMBI, plus all of your child’s other CF therapies, exactly as your child’s healthcare provider says they should be taken.

How ORKAMBI is packaged

Every carton has 56 ORKAMBI oral granule packets, organized in 4 weekly wallets. Each wallet has 2 packets for each of the 7 days in a week.

How to give your child ORKAMBI® (lumacaftor/ivacaftor) oral granules

STEP 1: Preparation

- Hold the packet with the cut line on top, shake it gently to settle the ORKAMBI granules, and either tear or cut the packet open along the cut line
- Carefully pour all of the ORKAMBI granules in the packet into 1 teaspoon (5 mL) of soft food such as flavored pudding or yogurt, or liquid such as juice, in a small container. Food or liquid should be at or below room temperature. Then mix the granules with the food or liquid

STEP 2: Administration

- Have your child take ORKAMBI within 1 hour of mixing
- Make sure your child completely finishes the full ORKAMBI mixture at each dose

STEP 3: Fat-containing food before or after the dose

- Make sure to always give your child food that contains fat just before or after the dose of ORKAMBI oral granules
- Some fat-containing foods include eggs, avocados, nuts, butter, and whole milk dairy products

Please see Important Safety Information and full Prescribing Information, including Patient Information.
What if your child misses a dose of ORKAMBI®?

- If your child misses a dose of ORKAMBI and it is **within 6 hours** of when he or she usually takes it, have your child take the dose of ORKAMBI as prescribed with fat-containing food as soon as possible.
- If your child misses a dose and it is **more than 6 hours** after the time he or she usually takes it, have your child **skip that dose only** and take the next dose with fat-containing food when he or she usually takes it. **Do not take 2 doses at the same time to make up for a missed dose**.
- Tell your child’s healthcare provider if your child stops taking ORKAMBI for **more than 1 week**. He or she may need to change your child’s ORKAMBI dose or other medicines your child takes.

Always take ORKAMBI with foods that contain fat

It’s important that your child always take ORKAMBI with a fat-containing meal or snack because these foods help the body better absorb the medicine. ORKAMBI should **always** be taken before or after a meal or snack that contains fat.

Your child’s healthcare provider can help you choose healthy fat-containing meals and snacks.

Looking for recipe ideas? Visit [ORKAMBI.com](http://ORKAMBI.com) for some fun meals to prepare for your family.

Tips for helping your child take ORKAMBI® (lumacaftor/ivacaftor)

Your child may find that ORKAMBI oral granules have a bitter taste. Every dose matters, so it’s important to make sure that your child completely finishes the ORKAMBI mixture at each dose. Here are a few tips that may help.

1. **Give your child ORKAMBI oral granules with 5 mL of soft food or liquid that is sweet or rich** like pudding, yogurt, chocolate sauce, or caramel sauce. Melted ice cream is also a nice treat to mix with granules.
2. **Maple syrup or honey may help with the aftertaste of ORKAMBI** — you can give some to your child after he or she completely finishes the dose of ORKAMBI. It’s worth a sweet shot!
3. **Numb the taste buds before taking ORKAMBI** by having your child eat an ice pop or another cold treat before his or her dose.
4. **Put some control in your child’s hands** by letting him or her decide what fat-containing food to eat before or after the dose. A slice of whole-milk cheese pizza or another favorite fat-containing food after ORKAMBI could be a tasty reward.
5. **Does your child have a favorite doll or toy?** Holding a special toy while taking ORKAMBI may make your child feel more comfortable.
6. **Try creating a rewards system for finishing ORKAMBI, such as collecting stickers.** You know best how to keep your child motivated!

Please see Important Safety Information and full Prescribing Information, including Patient Information.
Meet Cammy and her dog, Sam

Cammy and Sam are here to explain how to prepare and take ORKAMBI® oral granules.

Watch Taking ORKAMBI Oral Granules With Cammy & Sam

For more information about how to take ORKAMBI, visit ORKAMBI.com and watch this fun and informative video.

Color Cammy and Sam’s World

The following pages are included for your child to color. You can tear each of them out to hang his or her latest masterpiece on the wall or refrigerator.

Please see Important Safety Information and full Prescribing Information, including Patient Information.
This is my dog Sam. We’re best friends. Mom helps me go on the computer to learn about cystic fibrosis.
We like to play together.

Sam's a real good boy.
Vertex GPS provides eligible patients with information about reimbursement, potential financial assistance, product educational resources, and more, including:

- **Monthly refill reminders** from your Case Manager
- **Delicious Dishes** videos for tasty recipes

Vertex GPS is here to provide you with product support. You can speak to your dedicated Case Manager at 1-877-752-5933 (press 2), Monday through Friday, from 8:30 AM to 7:00 PM ET.

If your child has been prescribed ORKAMBI® (lumacaftor/ivacaftor) and you are not enrolled, please speak with your child’s healthcare provider about your interest in Vertex GPS. You can also learn more at VertexGPS.com.

Please see full [Prescribing Information](#), including [Patient Information](#).