PATIENT INFORMATION LEAFLET

WARNING: WARNING: TENDINITIS AND TENDON RUPTURE (inflammation and rupture of the muscles that bind the muscles to the bones), PERIPHERAL NEUROPATHY (disturbances in the distant nerves for any reason - loss of sensation), CENTRAL NERVOUS SYSTEM EFFECTS (central nervous system) and MYASTHENIA GRAVIS SERIOUS UNWANTED EFFECTS, INCLUDING EXACERBATION

Fluoroquinolones, including RAVIVO, can cause disabling and irreversible undesirable effects such as:

- Tendonitis and tendon (the ligaments that attach the muscles to the bones) rupture
- Peripheral neuropathy (disorders of distal nerves for any reason – loss of sensation)
- Central nervous system effects

Ravivo should be discontinued immediately and fluoroquinolones should be avoided in patients who experience any of these reactions.

Fluoroquinolones, including RAVIVO, may exacerbate muscle weakness in patients with myasthenia gravis. The use of RAVIVO should be avoided in patients with a known history of myasthenia gravis.

Since fluoroquinolone group drugs, including RAVIVO, are known to be associated with serious side effects, they can be used in the following indications if there is no other alternative.

- Acute bacterial sinusitis (acute inflammation of the air void inside the facial bones caused by bacteria)
- Uncomplicated urinary tract infection
- Acute bacterial exacerbation of chronic bronchitis (re-exacerbation of persistent inflammation of the bronchial tubes-membranes of the lung)

RAVIVO 750 mg Film Coated Tablet

It is taken orally.

- ***Active substance:*** 768,69 mg levofloxacin hemihydrate equivalent to 750 mg levofloxacin
- ***Excipients:*** Microcrystalline cellulose (Avicel PH102), crospovidone, hydroxypropyl methyl cellulose, sodium stearyl fumarate, polyvinyl alcohol, titanium dioxide, polyethylene glycol, talc.

Please read these LEAFLET carefully before you start using this medicine because it contains important information for you.

- *Keep these leaflet. You can need to read again.*
- *If you have other questions, please talk your doctor or pharmacist.*
- *This drug has been prescribed for you personally, do not give it to others.*
- *If you go to the doctor or hospital during the use of this medicine, tell your doctor that you are taking it.*
- *Follow the informations in these leaflet. Do not use* ***high or low doses*** *other than the recommended dose*

In this instructions for use:

Headlines are included.

1. What is RAVIVO and what is it used for?

RAVIVO is in the form of white colored, oblong, notched film-coated tablets containing levofloxacin, a broad spectrum antibacterial fluoroquinolone derivative, as active ingredient. RAVIVO is available as 7 film tablets containing levofloxacin hemihydrate equivalent to 750 mg of levofloxacin in a white opaque PVC / PVDC aluminum foil blister pack.

In acute bacterial sinusitis, acute bacterial exacerbation of chronic bronchitis, and uncomplicated urinary infections should not be used because of the risk of serious side effects if alternative treatment options are available.
In these indications, it can only be used with the approval of an infectious diseases specialist in cases where it is proven with an antibiogram and other alternative treatments cannot be applied.
RAVIVO is used in the treatment of lung, sinus, urinary tract, skin and soft tissue infections.

RAVIVO is used to treat the following conditions*:*

- Community-acquired pneumonia
- Hospital-acquired (nosocomial) pneumoni
- Acute exacerbation of chronic bronchitis (long-term respiratory infections that cause difficulty in breathing)
- Acute bacterial sinusitis (Sinus inflammation),Uncomplicated and complicated (severe) urinary tract infections (Urinary tract infections)
- Chronic bacterial prostatitis (long-standing prostate inflammation)
- Acute pyelonephritis (Kidney infection)
- Uncomplicated and complicated (severe) skin and soft tissue infections (skin or subcutaneous infections)
- In case of possible ingestion of the germ of anthrax through the respiratory tract (After exposure, inhaled anthrax)

2. Things to consider before using RAVIVO

DO NOT use RAVIVO in the following cases:

If :

- If you are allergic to levofloxacin or any substance in the composition of RAVIVO, or other fluoroquinolone group antibacterial drugs (moxifloxacin, ciprofloxacin, gemifloxacin, ofloxacin),
- Florokinolon grubu antibiyotik kullanırken bir tendon (kasları ve kemikleri birleştiren bağ) problemi yaşadı iseniz.
- If you have ever had a tendon (ligament connecting muscles and bones) problem while using fluoroquinolone antibiotics.

Fluoroquinolones, including RAVIVO, have been associated with serious disability and potentially

irreversible adverse reactions. Common adverse reactions include musculoskeletal and peripheral nervous system (tendinitis, tendon rupture, tendon swelling or inflammation, tingling or numbness, numbness of arms and legs, muscle pain, muscle weakness, joint pain, joint swelling), arthralgia (joint pain), myalgia (muscle pain), peripheral neuropathy, and central nervous system effects (hallucination, anxiety, depression, suicidal tendency, insomnia, severe headache, and confusion) (see Chapter 4. "What are the possible side effects?").

These reactions can occur within hours or weeks after initiation of treatment of RAVIVO. Patients of all age groups or patients without pre-existing risk factors have experienced these adverse reactions.

RAVIVO should be discontinued immediately if the first signs or symptoms of any serious adverse reactions occur. Furthermore, the use of fluoroquinolones, including RAVIVO, should be avoided in patients experiencing any of these serious adverse reactions associated with fluoroquinolones.

- If you have had epilepsy crisis,
- If tendon inflammation and pain develop during RAVIVO treatment, especially in elderly patients or patients taking corticosteroids (cortisone and similar drugs),
- If severe, persistent and / or bloody diarrhea develops during or immediately after RAVIVO treatment,
- If you are pregnant, suspect of pregnancy or are planning to become pregnant,
- If you are breastfeeding your baby, do not use this drug.
- This medicine is for adults only. Do not give to children under the age of 18.

USE RAVIVO CAREFULLY in the following cases

If:

- If you have a known heart disease or a family history of QT prolongation (problem with the rhythm of the heart),
- If you have diabetes and you are using antidiabetic drugs,
- If you have a nervous system or mental illness,
- If you have suffered paralysis or other brain diseases that cause brain damage,
- If you are 65 years or older,
- If you have glucose-6-phosphate dehydrogenase deficiency disease,
- If you have kidney problems (your doctor may lower your medication dose),

- If you have liver problems,
- If you are directly exposed to sunlight and artificial ultraviolet lamps such as solarium,
- If you have had a seizure (attack) before,
- If you are using medicines containing corticosteroid,
- Patients with a history of myasthenia gravis (impaired nerve-muscle conduction and consequently abnormal fatigue of myasthenia),

Use RAVIVO with caution.

If you are exposed to sunlight while using RAVIVO, your skin may develop sensitivity and sunburn. For this reason, use sunscreen with a high protection factor, wear clothes that will cover your arms and legs, and a hat, avoid sunbathing. If these warnings apply to you, even at any time in the past, please consult your doctor.

Use of RAVIVO with food and drink

Take RAVIVO without chewing, with a sufficient amount of liquid. You can take the tablets during or between meals.

Pregnancy

Consult your doctor or pharmacist before using this medication.

There are no adequate and well-controlled studies in pregnant women, therefore RAVIVO should not be used during pregnancy.

If you notice that you are pregnant during treatment, consult your doctor immediately.

Breast-feeding

Consult your doctor or pharmacist before using this medication.

RAVIVO tablet should not be used during breastfeeding period.

Vehicle and machine use

RAVIVO can cause undesirable effects such as dizziness, daze and drowsiness. ome of these side effects can affect concentration and reflex speed. Therefore, care should be taken while driving and using machinery.

Important information about some of the excipients contained in RAVIVO

This medicinal product contains less than 1 mmol (23 mg) sodium per tablet; No sodium-related side effects are expected at this dose.

Concomitant use with other drugs

- When RAVIVO is taken together with iron salts (used for anemia), zinc-containing multivitamins, magnesium or aluminum-containing antacids (used for heartburn) and sucralfate (used in stomach ulcers), since the absorption of RAVIVO is significantly reduced, use these drugs at least two hours before or two hours after RAVIVO administration.
- If applied together with calcium carbonate, there will be no clinically significant change in its pharmacokinetics
- Since an increase in theophylline (used in shortness of breath) levels is detected with quinolones, theophylline levels should be monitored when used with RAVIVO. The risk of epileptic seizures may be increased if theophylline is used with RAVIVO.
- Prothrombin time and bleeding symptoms should be monitored during the concomitant use of RAVIVO and the vitamin K antagonist warfarin.If nonsteroidal anti-inflammatory drugs (aspirin, ibuprofen, fenbufen, ketoprofen and indomethacin) used for rheumatic pain and rheumatic inflammation are taken together with RAVIVO, the risk of epileptic seizures may increase.
- Increase or decrease in blood sugar (hyperglycemia-hypoglycemia) has been reported during the concomitant use of RAVIVO and antidiabetic drugs. Therefore, blood sugar levels should be monitored during concomitant use.
- Probenecid (used in gout) and cimetidine (used in heartburn) may reduce the excretion of RAVIVO from the kidneys.
- RAVIVO may prolong the effect of cyclosporine (used in organ transplants).
- n case of use with RAVIVO; Antiarrhythmics (quinidine and amiodorone; used to treat abnormal heart rhythm), drugs used to treat depression (tricyclic antidepressants; amitriptyline and imipramine), and some drugs used to treat bacterial infections (macrolide antibiotics; erythromycin, azithromycin, and clarithromycin) can affect your heart rhythm.

- If RAVIVO is used with corticosteroids, the likelihood of tendon inflammation is higher
- Using strong painkillers called opiates use together with RAVIVO may result in "false positive" results in urine tests. Therefore, tell your doctor if you use this type of medication.

If you are currently using or have recently used any prescription or non-prescription drugs, please inform your doctor or pharmacist about them.

3. How to use RAVIVO?

Instructions for proper use and dose/frequency of administration:

- Your doctor will determine the amount of dose you will take and the duration of treatment with RAVIVO.
- RAVIVO is administered once a day.
- The dosage will be adjusted by your doctor depending on the type and severity of the infection and the susceptibility of the causative pathogen.
- The duration of treatment varies depending on the course of the disease.
- As with all antibiotic treatments in general, RAVIVO treatment should be continued for 48- 72 hours after the patient is afebrile or bacterial eradication is achieved.

Application route and method:

- RAVIVO is taken orally. It should be taken without chewing with a sufficient amount of liquid.
- The tablets can be taken during or between meals.
- Take the tablets at the same times of the day

Different age groups:

Use in children:

It should not be used in children under the age of 18

Use in the elderly:

In elderly patients, no dosage adjustment is required if renal function is adequate.

Special use cases

Kidney failure:

The dose of RAVIVO needs to be adjusted in renal impairment. n the use of RAVIVO, the dose will be adjusted by your doctor to prevent accumulation. Hemodialysis and continuous ambulatory peritoneal dialysis have no effect on the removal of Ravivo from the body.

Liver failure:

Dosage adjustment is not required in liver failure.

If you have the impression that the effect of RAVIVO is too strong or too weak, talk to your doctor or pharmacist.

If you have used more RAVIVO than you should:

If you have used more than you should use from RAVIVO, talk to a doctor or pharmacist.

In case of overdose, central nervous system symptoms such as confusion, dizziness, loss of consciousness and epileptic seizures, gastrointestinal system reactions such as nausea and heartburn, and cardiac symptoms such as disruption of heart beat rhythm may occur.

In case of overdose, contact your doctor or a hospital immediately.

If you forget to use RAVIVO:

If you forget a dose, do not worry. If the next dose is not near, take one as soon as you remember. Take your next dose at the usual time.

Do not take a double dose to make up for forgotten doses.

Effects that may occur when treatment with RAVIVO is terminated:

Not available. However, even if you feel better, use your medications until your treatment period is over.

4. What are the possible side effects?

Like all medicines, there may be side effects in people who are sensitive to the ingredients of RAVIVO.

Side effects are classified as shown in the following categories:

Very common : It can be seen in at least one of 10 patients.

Common : Less than one in 10 patients, but more than one in 100 patients.

Uncommon : less than one in 100 patients, but more than one in 1,000 patients.

Rare : Less than one in 1,000 patients may be seen.

Very rare : Less than one in 10,000 patients can be seen.

Unknown : Cannot be estimated from the available data.

If any of the following occur, stop using RAVIVO and IMMEDIATELY inform your doctor or go to the nearest hospital emergency department:

Very rare (It can be seen in less than 1 in 10000 patients.)

- You have an allergic reaction. Symptoms may include: redness, swallowing or breathing problems, swelling of lips, face, throat or tongue

These are all very serious side effects.

If you have one of these, you have a serious allergy to RAVIVO. You may need urgent medical attention or hospitalization. All of these very serious side effects are very rare

If you notice any of the following, tell your doctor immediately or go to the nearest hospital emergency department:

Rare (less than 1 in 1000 patients, but more than 1 in 10000 patients)

- Watery diarrhea, which may also be bloody, possibly stomach cramps and high fever. These could be signs of a serious bowel problem.
- Tendon inflammation and pain. The Achilles tendon is most commonly affected, and in some cases, the tendon may rupture.
- Contractions (convulsions).

Very rare (may be seen in less than 1 in 10000 patients)

- Burning, tingling, pain or numbness. These can sometimes be symptoms of a condition called "neuropathy".

Unknown

- Severe skin rashes around the eyes, lips, mouth, nose and genitals area involving swelling or spilling of the skin
- Loss of appetite, yellow discoloration of the eyes and skin, dark urine, itching, or pain in the abdomen. These could be signs of liver problems.

It may require immediate medical attention. Serious side effects are very rare.

If you notice any of the following, tell your doctor:

Common (less than 1 in 10 patients, but more than 1 in 100 patients)

- Feeling sick (nausea) and diarrhea
- Increased levels of some liver enzymes in the blood.

Uncommon (may affect less than 1 in 100 patients, but more than 1 in 1000 patients)

- Itching and skin rash.
- Loss of appetite, stomach problem or indigestion (dyspepsia), vomiting or abdominal pain, bloating or constipation.
- Headache, lightheadedness, dizziness, insomnia or nervousness.
- Abnormal results in blood tests due to liver or kidney problems
- Changes in the number of white blood cells seen in the results of some blood tests.
- Weakness.
- Changes in the number of other bacteria or fungi that may need to be treated.

Rare (less than one in 1000 patients, but more than one in 10000 patients.)

- Tingling sensation (paraesthesia) or tremors in your hands and feet.
- Anxiety, distress, depression, restlessness or confusion
- Palpitations or low blood pressure.
- Joint pain or muscle pain.
- Bruising and bleeding due to a decrease in the number of platelets (takes part in blood clotting)
- Decrease in the number of white blood cells (white blood cells)
- Difficulty breathing or wheezing (bronchospasm)
- Shortness of breath (dyspnea)
- Serious itching or skin rash

Very rare (less than one in 10000 patients may be seen.)

- Increased skin sensitivity to the sun and ultraviolet light.
- Low blood sugar (hypoglycemia). This is important for people with diabetes.
- Visual and hearing impairments or taste and smell disturbances.
- Psychotic reactions with self-harm, including seeing or hearing things that are not real (hallucinations), suicidal thoughts and behaviors.
- Circulatory impairment (anaphylactic-like reaction)
- Tendon rupture (this undesirable effect may occur within the first 48 hours of treatment and on both sides), muscle weakness. This is important for people with myasthenia gravis (nerve-muscle conduction disorder).
- Hepatic inflammation, renal disorders and an allergic kidney reactions renal failure due to interstitial nephritis,
- Fever, sore throat and a general feeling of sickness that does not go away. This may be due to a decrease in the number of white blood cells.Ateş ve alerjik akciğer reaksiyonları.
- Aggravation of myasthenia gravis

Unknown

- A decrease in the number of red blood cells (anaemia). This condition can turn the skin pale or yellow due to damage to red blood cells and a decrease in the number of all blood cell types.
- Excessive immune response (hypersensitivity).Aşırı terleme
- Pain in the back, chest, arms and legs.
- Difficulty in moving and walking.
- Attacks of porphyria in people with porphyria (a very rare metabolic disease).
- Inflammation of blood vessels due to an allergic reaction.

These are mild side effects of RAVIVO.

Inform your doctor or pharmacist if you encounter any side effects not mentioned in these leaflet.

Reporting of side effects

Talk to your doctor, pharmacist or nurse in case of any side effects that are included or not in the Instructions for Use. Also, report the side effects you encounter to the Turkish Pharmacovigilance Center (TUPC) by clicking on the "Drug Side Effects Reporting" icon on www.titck.gov.tr or by calling the side effect reporting line at 0 800 314 00 08. By reporting

the side effects that occur, you will contribute to obtaining more information about the safety of the drug you are using.

5. Storage of RAVIVO

Keep RAVIVO out of the reach of children and in its packaging.

Store at room temperature below 25˚C.

Use in accordance with the expiry date.

Do not use RAVIVO after the expiry date on the packaging.

Do not throw away expired or unused medicines! Give it to the collection system determined by the Ministry of Environment and Urbanization.

License owner:

Drogsan İlaçları San. ve Tic. A.Ş.
Oğuzlar Mah. 1370. Sok 7/3
06520 Balgat- ANKARA

Manufacturing Site:

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06760 Çubuk/ANKARA

These instructions for use were approved on 28/05/2019.