Patient Leaflet According to the Pharmacists' Regulations (Preparations) – 1986

This medicine is sold with a doctor's prescription only

Daliresp 500 mcg

Film-coated tablets

Active ingredient:

Each tablet of Daliresp 500 mcg contains:

Roflumilast 500 mcg.

For the list of the additional ingredients, see section 6.

Read this entire leaflet carefully before using the medicine.

This leaflet contains concise information about the medicine. If you have any further questions, please refer to your doctor or pharmacist.

This medicine has been prescribed to treat your illness. Do not pass it on to others. It may harm them, even if it seems to you that their illness is similar to yours.

The medicine is not indicated for treatment of children and adolescents under 18 years of age.

1. What is the medicine intended for?

The medicine is indicated for maintenance treatment of severe COPD (Chronic Obstructive Pulmonary Disease) accompanied by chronic bronchitis, in adults with a prior history of recurring flare-ups, in addition to bronchodilators treatment.

Therapeutic group:

Phosphodiasterase 4 inhibitor (PDE-4 inhibitor), a type of anti-inflammatory medicine.

2. Before using the medicine

Do not use the medicine if:

•Do not use if you are sensitive (allergic) to the active ingredient or to any of the other ingredients the medicine contains (for a list of the other ingredients, see section 6).

• Do not use if you suffer from moderate to severe liver disease.

Special warnings regarding the use of this medicine:

- Sudden attack of breathlessness: Daliresp is not intended for the treatment of a sudden attack of breathlessness (acute bronchospasm). Therefore it is very important that your doctor provide you with another medicine to be available to you at all times and to assist you in treating these attacks.
- Body weight: During the course of treatment with Daliresp, you should check your body weight on a regular basis. If, while taking this medicine, you observe an unintentional loss of body weight (not related to a diet or exercise program) - refer to your doctor.
- The doctor should be informed immediately of any behavioral changes, mood changes and of any suicidal thoughts and/or behavior.
- Patients with a body weight of less than 60 kg: these patients have a higher risk of suffering from sleepl disorders.

Before (and during) treatment with Daliresp, tell your doctor if:

- You suffer from any liver problem. You suffer from severe immunological diseases (such as HIV infection/AIDS, multiple sclerosis, lupus erythematosus or viral inflammation of the brain - progressive multifocal leukoencephalopathy).
- You suffer or have suffered in the past from a severe infectious disease (such as tuberculosis, liver infection/hepatitis), herpes or herpes zoster.
- You suffer or have suffered in the past from cancer.
- You suffer from severe impairment of the heart function.
- You suffer or have suffered in the past from from mental problems, depression accompanied by suicidal thoughts or suicidal behavior.
- You suffer from sleeplessness, anxiety, nervousness or depression.

Possible interactions with other medicines taken concurrently with Daliresp:

If you are taking, or have recently taken any other medicines, including non-prescription medicines and nutritional supplements, please tell your doctor or pharmacist. Especially inform your doctor or pharmacist if you are taking the following medicines (it should be noted that the following list indicates the active ingredients in the medicines. If you are unsure whether you are using one of these medicines, please consult with your doctor or pharmacist):

- Theophylline (for the treatment of respiratory diseases).
- Fluvoxamine, enoxacin, cimetidine.
- Rifampicin (antibiotic), phenobarbital, carbamazepine, phenytoin (medicines for the treatment of epilepsy): if they are taken concurrently with Daliresp, the effects of Daliresp may be reduced.
- Medicines used for the treatment of immunological diseases, such as methotrexate, azathioprine, infliximab, etanercept or long-term oral corticosteroids.
- Daliresp may be taken together with other medicines used in the treatment of COPD, such as corticosteroids for short-term treatment or bronchodilators (inhaled or oral). Do not stop taking these medicines or reduce their dose unless explicitly instructed by your doctor.

Use of this medicine and food:

This medicine may be taken with or without food.

Pregnancy and breastfeeding:

- Do not take Daliresp if you are pregnant, plan to become pregnant or if you are breastfeeding.
- Do not become pregnant during treatment with the medicine because of the risk of harm to the fetus. Use an effective contraceptive method during the treatment. The treatment is not recommended for women who could become pregnant and are not using reliable contraceptives.

Use in Children:

The medicine is not indicated for treatment of children and adolescents under 18 years of age.

Important information about some of the medicine's ingredients:

The medicine contains lactose. If you are sensitive to lactose, inform your doctor before taking this medicine (see section 6).

3. How to use this medicine?

Always use according to the doctor's instructions. Check with your doctor or pharmacist if you are not sure.

The dosage and treatment regimen will be determined by the doctor only.

The standard dosage is usually: one tablet once daily.

Do not exceed the recommended dosage.

Swallow the tablets with water.

It is recommended to take the tablets at the same time every day.

It may take a few weeks until you feel the beneficial effect of the medicine.

There is no information on crushing/splitting/chewing.

Tests and follow-up:

During the treatment with Daliresp, you should check your body weight on a regular basis. If you observe an unintentional loss of body weight (not related to a diet or exercise program) - refer to your doctor immediately!

If you accidentally take a higher dosage: if you took an overdose, or if a child has accidentally swallowed the medicine, refer immediately to a doctor or proceed to a hospital emergency room, and bring the package of the medicine with you. Symptoms of an overdose may include: headache, digestive problems such as nausea and diarrhea; dizziness, rapid and strong

heartbeats (palpitations), dizziness, feeling of clamminess, stickiness and cold; low blood pressure.

If you forgot to take the medicine at the proper time take the missed dose as soon as you remember. If one day you have forgotten to take a tablet, on the next day take the tablet at the usual time. Do not take a double dose to make up for a forgotten dose.

Adhere to the treatment as recommended by your doctor.
Even if your state of health improves, do not stop the treatment with this medicine without consulting your doctor or pharmacist.
Do not take medicines in the dark! Check the label and the dose each time you take a medicine. Wear glasses if you need them.

4. Side effects

As for any medicine, the use of Daliresp may cause side effects in some users. If these side effects persist or are bothersome or get worse, please consult your doctor. Do not be alarmed while reading the list of side effects. You may not suffer from any of them.

Stop the treatment and contact your doctor or a hospital emergency room immediately, if an allergic reaction appears:
Allergic reaction (hypersensitivity) is a rare side effect and can affect the skin. In rare cases a severe allergic reaction may occur (including angioedema). This reaction may include symptoms such as: swelling of the face, eyelids, lips, tongue and may cause difficulty in breathing and/or a drop in blood pressure and rapid heartbeats.

The doctor should be informed immediately of any behavioral or mood changes and in any case of any suicidal thoughts and/or behavior (suicides has been reported as a rare side effect). You may also experience sleeplessness, anxiety, nervousness, panic attack or depression (the frequency of the side effects appears below).

Additional side effects (including frequency):
Common side effects (appear in 1-10 users out of 100):
- Weight loss, loss of appetite, sleeplessness.
- Headaches, diarrhea, nausea or stomach aches. You may experience these side effects during the first weeks of treatment. If these side effects persist, refer to your doctor.

Uncommon side effects (appear in 1-10 users out of 1,000):
Anxiety, trembling, dizziness, vertigo, palpitations (sensation of rapid or irregular heartbeat), stomach inflammation, vomiting, reflux of stomach acid to the gullet, indigestion, rash, muscle pain or cramps, muscle weakness, back pain, weakness, tiredness, feeling unwell.

Rare side effects (appear in 1-10 users out of 10,000):
Male breast enlargement, nervousness or depression, panic attack, decreased sense of taste, respiratory tract infections, bloody stools, constipation, elevation of liver or muscle enzymes (seen in blood tests), urticaria.
If you experience side effects not mentioned in this leaflet or if there is a change in your general feeling, consult the doctor immediately!

Side effects may be reported to the Ministry of Health by clicking on the link "Report on side effects following medicinal treatment" on the homepage of the Ministry of Health website (www.health.gov.il) which leads to an online form for reporting side effects, or by entering the link:
htpps://sideeffects.health.gov.il

5. How to store the medicine?

- Avoid poisoning! This medicine, and any other medicine, must be stored in a closed place out of the reach and sight of children and/or infants, to avoid poisoning. Do not induce vomiting unless explicitly instructed to do so by the doctor.
- Do not use the medicine after the expiry date (exp. date) stated on the package. The expiry date refers to the last day of that month.
- Storage conditions: store below 25°C.

6. Additional information

- **In addition to the active ingredient, the medicine also contains the following ingredients:**

Lactose monohydrate, maize starch, macrogol 4000, povidone (K90), hypromellose, magnesium stearate, titanium dioxide (E171), iron oxide yellow (E172).
Each tablet contains approximately 199 mg lactose.

- **What does the medicine look like and what does the package contain?**

Yellow, film-coated, D-shaped tablets embossed with the letter D on one side.
The tablets are marketed in blister packs of 30 or 90 tablets per box.

Manufacturer: Takeda GMBH, Germany

Registration holder: Rafa Laboratories Ltd., P.O. Box 405, Jerusalem 9100301.
Medicine registration number in the National Medicines Registry of the Ministry of Health: 1488033545

Drawn up in January 2021