

**Patient leaflet in accordance with Pharmacist's Regulations (Preparations) 1986**

The medicine is marketed according to doctor's prescription only.

**Rexulti 0.5 mg, Rexulti 1 mg, Rexulti 2 mg, Rexulti 3 mg, Rexulti 4 mg  
film coated tablets**

**Composition**

Each film coated tablet of Rexulti contains the active ingredient:

Rexulti 0.5 mg: Brexpiprazole 0.5 mg

Rexulti 1 mg: Brexpiprazole 1 mg

Rexulti 2 mg: Brexpiprazole 2 mg

Rexulti 3 mg: Brexpiprazole 3 mg

Rexulti 4 mg: Brexpiprazole 4 mg

**Inactive ingredients**

See section 6.

- **Read the leaflet carefully until the end before using the medication.** This leaflet contains concise information about the medicine. If you have further questions, refer to the doctor or the pharmacist.

- This medicine was prescribed for you. Do not pass it on to others. It may harm them even if their ailment/ medical condition seems similar to yours.

**Increased Mortality in Elderly Patients with Dementia-Related Psychosis**

**Elderly patients with dementia-related psychosis treated with antipsychotic drugs are at an increased risk of death. REXULTI is not intended for the treatment in cases of dementia-related psychosis.**

**Suicidal Thoughts and Behaviors**

**Antidepressants increased the risk of suicidal thoughts and behaviors in young patients aged up to 24 years in short-term studies. Monitor closely for clinical worsening and for emergence of suicidal thoughts and behaviors. REXULTI is not intended for children.**

**1. What is the medicine for?**

REXULTI is intended for the treatment of adults with:

- Major depressive disorder (MDD) in combination with antidepressant medicines.
- Schizophrenia.

**Therapeutic group:** atypical antipsychotic drugs

**2. Before using the medicine**

**Do not use the medicine:**

- if you are allergic to the active ingredient (brexpiprazole) or to any of the other ingredients of this medicine (listed in section 6).

Reactions have included rash, facial swelling, urticaria, and anaphylaxis.

**Special warnings regarding the use of the medicine**

**Rexulti may cause serious side effects, including:**

- **Increased mortality in elderly patients with dementia-related psychosis. REXULTI is not intended for the treatment in cases of dementia-related psychosis.** (see framed warning at the beginning of the leaflet)
- **Risk of suicidal thoughts or actions** (see framed warning at the beginning of the leaflet).

- **Antidepressant medicines may increase suicidal thoughts or actions in some children, teenagers, or young adults within the first few months of treatment.**
- **How to watch and try to prevent suicidal thoughts and actions?**
  - Pay close attention to any changes, especially sudden changes, in mood, behaviors, thoughts, or feelings. This is very important when an antidepressant medicine is started or when the dose is changed.
  - Call the doctor right away to report new or sudden changes in mood, behavior, thoughts, or feelings.
  - Keep all follow-up visits with the doctor as scheduled. Call the doctor between visits as needed, especially if you have concerns about symptoms.

**Call a doctor right away if you or your family member has any of the following symptoms, especially if they are new, worse, or worry you:**

- |  |  |
|--|--|
| ○ thoughts about suicide or dying                    | ○ attempts to commit suicide                 |
| ○ new or worsening depression                        | ○ new or worsening anxiety                   |
| ○ feeling very agitated or restless                  | ○ acting on dangerous impulses               |
| ○ panic attacks                                      | ○ trouble sleeping (insomnia)                |
| ○ new or worsening irritability                      | ○ acting aggressive, being angry, or violent |
| ○ an extreme increase in activity or talking (mania) | ○ other unusual changes in behavior or mood  |

- **Stroke (cerebrovascular problems) in elderly people with dementia-related psychosis that can lead to death. REXULTI is not intended for the treatment in cases of dementia-related psychosis.**
- **Neuroleptic Malignant Syndrome (NMS):** Tell your doctor right away if you have some or all of the following symptoms: high fever, stiff muscles, confusion, sweating, changes in pulse, heart rate, and blood pressure. These may be symptoms of a rare but serious condition that can even lead to death.
- **Uncontrolled body movements (tardive dyskinesia):** Rexulti may cause movements that you cannot control in your face, tongue or other body parts. Tardive dyskinesia may not go away, even if you stop taking Rexulti, and might also start after you stop taking Rexulti. Contact your doctor if you have any of these symptoms.
- **Unusual urges.** Some people treated with Rexulti have reported unusual urges, such as gambling, binge eating or eating that you cannot control (compulsive), compulsive shopping and sexual urges. If you or your family members notice that you are having unusual urges or behaviors, inform your Doctor.
- Low white blood cell count. Refer to your Doctor in case of fever or other symptom of infection.
- Decreased blood pressure (orthostatic hypotension). You may feel lightheaded or faint when you rise too quickly from a sitting or lying position.
- Problems controlling your body temperature so that you feel too warm (avoid getting overheated or dehydrated while taking Rexulti: do not over-exercise, stay out of the sun. Do not wear too much or heavy clothing, in hot weather stay inside in a cool place if possible, drink plenty of water).
- Difficulty swallowing that can cause food or liquid to get into your lungs.

**Before taking REXULTI, tell your doctor if you:**

- have diabetes or high blood sugar or a family history of diabetes or high blood sugar. Your doctor should check your blood sugar before you start REXULTI and during your treatment.
- have high levels of cholesterol, triglycerides, LDL-cholesterol, or low levels of HDL cholesterol
- have or had seizures (convulsions)
- have or had low or high blood pressure
- have or had heart problems or a stroke
- have or had a low white blood cell count
- are pregnant or plan to become pregnant. See " Pregnancy and breast-feeding" section below.
- are breastfeeding or plan to breastfeed. See " Pregnancy and breast-feeding" section below.

- **Antidepressants are medicines used to treat depression and other illnesses.** It is important to discuss all the risks of treating depression and also the risks of not treating it. Patients and their families or other caregivers should discuss all treatment choices with the doctor, not just the use of antidepressants.

**Children and adolescents:** REXULTI is not intended for treating children.

**Interactions with other drugs:**

**If you are taking, or have recently taken, other medicines, including nonprescription medicines and nutritional supplements, tell the doctor or pharmacist.**

- strong inhibitors of CYP3A4 such as ketoconazole, itraconazole, clarithromycin
- strong inhibitors of CYP2D6 such as fluoxetine, paroxetine, duloxetine and quinidine
- strong activators of CYP3A4 such as St. John's Wort, rifampicin

**Use of the medicine and food:** REXULTI can be taken with or without food.

**Pregnancy and breast-feeding**

- Tell your doctor before taking Rexulti, if you are pregnant or plan to become pregnant. It is not known if REXULTI may harm your unborn baby. Using REXULTI in the last trimester of pregnancy may cause muscle movement problems, medicine withdrawal symptoms, or both of these in your newborn.
  - If you become pregnant while taking REXULTI, talk to your doctor.
- Tell your doctor before taking Rexulti, if you are breastfeeding or plan to breastfeed. It is not known if REXULTI passes into your breast milk. You and your doctor should decide if you will take REXULTI or breastfeed.

**Driving and using machines:** **Do not** drive a car, operate machinery, or do other dangerous activities until you know how REXULTI affects you. REXULTI may make you feel drowsy.

**Important information on some of the ingredients of the medicine:** Rexulti contains lactose. If you have been told by your doctor that you have an intolerance to some sugars, consult with your doctor before taking this medicine.

**3. How should you use the medicine?**

Always take this medicine exactly as your doctor has told you. Check with your doctor or pharmacist if you are not sure. **The dosage and treatment regimen will be determined by the doctor only.** Rexulti is taken orally.

The usual dose is:

**Major depressive disorder (MDD) in adults:**

The recommended starting dosage for Rexulti as adjunctive treatment is 0.5 mg or 1 mg once daily, taken orally with or without food.

The doctor may gradually increase the dosage to 2 mg once daily. Dosage increases should occur at weekly intervals based on your response and tolerability. The maximum recommended daily dosage is 3 mg.

**Schizophrenia in adults:**

The recommended starting dosage for Rexulti is 1 mg once daily on Days 1 to 4, taken orally with or without food.

The doctor may gradually increase the dosage to 2 mg once daily on Day 5 through Day 7, then to 4 mg on Day 8 based on your clinical response and tolerability. The maximum recommended daily dosage is 4 mg.

**Take Rexulti exactly as your doctor tells you to take it. Do not change the dose or stop taking Rexulti yourself.**

**Do not exceed the recommended dosage.**

- Do not divide the tablet. The tablets do not have a score/dividing line and therefore the tablets cannot be split into equal doses.

- **If you have accidentally taken a higher dosage**, you should refer to your doctor or go to the nearest hospital emergency room. If a child has accidentally swallowed the medicine, proceed immediately to a doctor or to the emergency room of a hospital and bring the package of the medicine with you.

**- If you have forgotten to take the medicine**

You should not miss a dose of Rexulti. If you miss a dose, take the missed dose as soon as you remember. If you are close to your next dose, just skip the missed dose and take your next dose at your regular time. Do not take 2 doses of Rexulti at the same time. If you are not sure about your dosing, call your doctor.

- You should complete the treatment as recommended by the doctor. Even if there is an improvement in your health, do not stop the treatment without first talking to your doctor.

**Stopping an antidepressant medicine suddenly** can cause other symptoms.

- **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.**

- **If you have any further questions on the use of this medicine, ask your doctor or pharmacist.**

**4. Side effects**

As with all medicines, the use of Rexulti may cause side effects in some users. Do not be alarmed at reading the list of side effects. You may not suffer from any of them.

**Rexulti may cause serious side effects, including:**

- **Stroke (cerebrovascular problems) in elderly people with dementia-related psychosis that can lead to death.**
- **Neuroleptic Malignant Syndrome (NMS):** Tell your doctor right away if you have some or all of the following symptoms: high fever, stiff muscles, confusion, sweating, changes in pulse, heart rate, and blood pressure. These may be symptoms of a rare but serious condition that can even lead to death. Contact your doctor if you have any of these symptoms.
- **Uncontrolled body movements (tardive dyskinesia):** Rexulti may cause movements that you cannot control in your face, tongue or other body parts. Tardive dyskinesia may

not go away, even if you stop taking Rexulti, and might also start after you stop taking Rexulti. Contact your doctor if you have any of these symptoms.

- **Unusual urges.** Some people treated with Rexulti have reported unusual urges, such as gambling, binge eating or eating that you cannot control (compulsive), compulsive shopping and sexual urges. If you or your family members notice that you are having unusual urges or behaviors, inform your Doctor.
- Low white blood cell count.
- Decreased blood pressure (orthostatic hypotension). You may feel lightheaded or faint when you rise too quickly from a sitting or lying position.
- Seizures (convulsions)
- Problems controlling your body temperature so that you feel too warm.
- Difficulty swallowing that can cause food or liquid to get into your lungs.
- **High blood sugar (hyperglycemia):** Increases in blood sugar can happen in some people who take Rexulti. Extremely high blood sugar can lead to coma or death. If you have diabetes or risk factors for diabetes (such as being overweight or having a family history of diabetes), your doctor should check your blood sugar before you start taking Rexulti and during your treatment.

Call doctor if you have any of these symptoms of high blood sugar while taking Rexulti: feel very thirsty; feel very hungry; feel sick to your stomach; feel weak or tired; need to urinate more than usual; feel confused, or your breath smells fruity.

- **Increased fat levels (cholesterol and triglycerides) in your blood.**
- **Weight gain:** You and your doctor should check your weight regularly.

#### **Other side effects- frequencies are according to the clinical trials frequencies**

##### **Common side effects – appearing in 1-10 out of 100 users**

- Constipation, fatigue, nasopharyngitis, blood cortisol decreased, increased appetite, sense of inner restlessness such as feeling like you need to move constantly, headache, somnolence, tremor, dizziness, anxiety, restlessness, dyspepsia, diarrhea, blood creatine phosphokinase increased.

##### **Other side effects reported with frequency higher than 1%:**

Vision blurred, salivary hypersecretion, dry mouth, abdominal pain, flatulence, urinary tract infection, blood prolactin increased, myalgia, abnormal dreams, increased sweating.

##### **Additional possible side effects with frequency not yet determined**

- Seizures (convulsions)
- Falls
- Potential for cognitive and motor impairment
- Dystonia (spasm of the neck muscles, and sometimes tightness of the throat, swallowing difficulty, difficulty breathing, and protrusion of the tongue) may occur in susceptible individuals during the first few days of treatment.

**If you get a side effect, if one of the side effects worsens, or when you suffer from side effects not mentioned in this leaflet, consult with the doctor.**

### **Reporting of side effects**

Side effects can be reported to the Ministry of Health by clicking on the link "Adverse Drug Reactions Report" that appears on the home page of the Ministry of Health web site ([www.health.gov.il](http://www.health.gov.il)), which leads to an online form for reporting side effects. Alternatively you can use the following link: <https://sideeffects.health.gov.il/>

### **5. How should the medicine be stored?**

Avoid poisoning! This medicine and all other medicines should be stored in a closed place out of the reach and sight of children and/or infants in order to avoid poisoning.

Do not induce vomiting unless explicitly instructed by a doctor to do so. Do not use this medicine after the expiry date (exp.date) that appears on the package.

The expiry date refers to the last day of that month.

**Storage conditions:** Store below 30°C, in the original package in order to protect from light and moisture.

Do not throw away any medicines via wastewater or household waste. Ask your pharmacist how to throw away medicines you no longer use or expired. These measures will help protect the environment.

### **6. Additional information**

#### **In addition to the active ingredient, the medicine also contains:**

lactose monohydrate, corn starch, low-substituted hydroxypropyl cellulose, microcrystalline cellulose,; Colorants - Rexulti 0.5 mg- OPADRY 03A430000 Orange, Rexulti 1 mg- OPADRY 03A420002 Yellow, Rexulti 2 mg- OPADRY 03A410000 Green, Rexulti 3 mg- OPADRY 03A400000 Purple, Rexulti 4 mg- OPADRY 03A480004 White; hydroxypropyl cellulose, magnesium stearate.

#### **- What does the medicine look like and contents of the pack:**

**Rexulti 0.5 mg:** light orange, round, shallow convex; bevel-edged tablet, marked with "BRX" and "0.5", 7 tablets in package.

**Rexulti 1 mg:** light yellow, round, shallow convex; bevel-edged tablet, marked with "BRX" and "1", 28 tablets in package.

**Rexulti 2 mg:** light green, round, shallow convex; bevel-edged tablet, marked with "BRX" and "2", 28 tablets in package.

**Rexulti 3 mg:** light purple, round, shallow convex; bevel-edged tablet, marked with "BRX" and "3", 28 tablets in package.

**Rexulti 4 mg:** white, round, shallow convex; bevel-edged tablet, marked with "BRX" and "4", 28 tablets in package.

**License Holder:** Lundbeck Israel Ltd., 11 Galgaley Haplada, P.O.B. 13105, Herzliya 4672211

**Manufacturer:** H. Lundbeck A/S, Ottiliavej 9, DK-2500 Valby, Denmark.

Registration numbers of the medicine in the Ministry of Health registry:

Rexulti 0.5 mg: 163-39-35733

Rexulti 1 mg: 163-40-35734

Rexulti 2 mg: 163-41-35735

Rexulti 3 mg: 163-42-35736

Rexulti 4 mg: 163-43-35737

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