Patient leaflet in accordance with the Pharmacists' Regulations (Preparations) - 1986
This medicine is dispensed with a doctor's prescription only

Decapeptyl Depot 3.75 mg
Powder and solvent for suspension for intramuscular injection

Composition:
Each vial of Decapeptyl Depot 3.75 mg contains 3.75 mg triptorelin (as embonate)

Inactive ingredients and allergens: See section 2 'Important information about some of this medicine's ingredients' and section 6 'Additional information'.

Read the entire leaflet carefully before you start using this medicine. This leaflet contains concise information about this medicine. If you have any further questions, consult 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.

1. What is this medicine intended for?

This medicine is indicated for:
- treatment of advanced hormone-dependent prostate cancer
- reducing the level of sex hormones
- treatment of precocious puberty
- as adjuvant treatment in combination with tamoxifen or aromatase inhibitor in pre-menopausal women with hormone-responsive early-stage breast cancer after completion of chemotherapy and in men with breast cancer at high risk of disease recurrence.

Therapeutic group: Long-acting gonadotropin releasing hormone (GnRH) analog

2. Before using this medicine

Do not use this medicine:

- If you are sensitive (allergic) to triptorelin embonate, gonadotropin releasing hormone (GnRH), other analogs of the gonadotropin releasing hormone (GnRH is also called LHRH), or to any of the other ingredients that this medicine contains (see Section 6).

Men
- This medicine is not indicated for patients with hormone-independent prostate cancer or in case of surgical castration.

Women
- If you are pregnant or breastfeeding.

Special warnings about using this medicine
All patients treated with Decapeptyl Depot
Before treatment with the medicine, tell your doctor if:
- You have diabetes or another disease affecting metabolism. Your doctor will check your sugar levels during treatment.
- You have a cardiac or vascular disease. Your doctor will check your blood pressure during treatment.
- You have any cardiac or vascular diseases, including cardiac arrhythmias or are treated with medicines for these diseases. The risk of cardiac arrhythmias may increase during Decapeptyl Depot use.

- You have a depressed mood. There have been reports on potentially severe depression in patients taking Decapeptyl Depot.
- You are using an anticoagulant, since bruising may occur in the injection area.

Treatment with Decapeptyl Depot, similarly to other **GnRH** analogs, may lead to bone weakening (osteoporosis) and increased risk of fractures, especially if you consume high alcohol quantities, smoke, have a family history of osteoporosis, malnutrition, undergo long-term treatment with medicines reducing bone mineral density (e.g., anticonvulsants, steroids). Your doctor will perform a bone scan before starting treatment to check whether you are at risk of osteoporosis and periodic testing during treatment.

- If you have pituitary hyperplasia (a benign tumor) which has not been known to you, it may be revealed during treatment with Decapeptyl Depot. Symptoms include sudden headaches, vomiting, visual impairment and paralysis of the eye muscles. In such cases, seek medical treatment immediately.
- Tests of pituitary or gonadal function conducted during treatment or after discontinuation of treatment with Decapeptyl Depot may be misleading.

Men:
- At the beginning of treatment, there will be an increase in the level of testosterone in your body. This may cause worsening of cancer symptoms. Contact your doctor if this happens. Your doctor may give you a medicine (anti-androgen) to prevent worsening of the symptoms.
- After surgical castration, triptorelin does not induce any further decrease in blood testosterone levels, therefore it should not be used.
- During the first weeks of treatment, Decapeptyl Depot may, in isolated cases, cause spinal cord compression or urethral (a tube transporting urine out of the body) obstruction. Your doctor will monitor your condition and treat you accordingly, if required.

Women**:**
- Pregnancy should be excluded prior to treatment initiation.
- Menstruation-like bleeding may occur during the first month of treatment. The menstrual bleeding will subsequently stop. Tell your doctor if you have menstrual bleeding after the first month of treatment.
- Menstrual bleeding may resume about 2 months after the last injection. You must use non-hormonal contraceptive methods during the treatment and for 1 month after the treatment, since pills or other hormonal contraceptive methods will not be effective in preventing pregnancy.

In children and adolescents:
- Tell the doctor if the child has a progressive brain tumor. This can affect the decisions the doctor makes about the treatment.
- Girls with precocious puberty may suffer from vaginal bleeding in the first month of treatment.
- After the end of treatment, the child will develop puberty characteristics.
- Information regarding future fertility is still limited. In most girls, regular menstrual bleeding will start, on the average, about one year after the end of treatment.
- Impairment of growth of the superior epiphysis of the femur and a medical condition called slipped capital femoral epiphysis. This condition has been observed after the end of treatment and it may cause limping, pain in the knees, pain in the hip, hip stiffness, legs turned outward, limitation of hip movement.

Tests and follow-up
Your doctor may instruct you to perform blood tests during the treatment to monitor treatment efficacy. In treatment of breast cancer, blood tests for determination of blood hormone levels should be performed prior to treatment initiation and during the treatment.

Drug interactions
If you are taking or have recently taken other medicines, including nonprescription medications and dietary supplements, tell your doctor or pharmacist. Particularly if you are taking:

- Decapeptyl may interfere with some medicines used for treatment of cardiac arrhythmias (e.g. quinidine, procainamide, amiodarone, sotalol) or may increase the risk of arrhythmias when administered together with other medicines (e.g. methadone used for pain relief and treatment of addictions),
  moxifloxacin antibiotic, antipsychotics used for treatment of mental illnesses.

Pregnancy and breastfeeding
If you are pregnant or breastfeeding, think you may be pregnant or are planning to have a baby, consult your doctor about this medicine.
Do not use Decapeptyl Depot if you are pregnant or breastfeeding.
Prior to treatment initiation, women of childbearing age should be tested by the doctor to exclude pregnancy. During the treatment, use non-hormonal contraceptive methods until the menstrual period is resumed. Consult the attending physician regarding this issue.

Driving and using machines
Even when taken as instructed, Decapeptyl Depot may alter your reaction and impair your ability to drive and operate machines. You may experience dizziness, tiredness or visual disturbances such as blurred vision. Some of these effects are side effects of the medicine and some are caused by your disease. If you experience any of these effects, do not drive or use instruments and devices.

Important information about some of this medicine's ingredients
This medicine contains less than 1 mmol sodium (23 mg) per dose, therefore it is considered sodium free.

3. How to use this medicine?

Always use this medicine according to your doctor's instructions. Check with your doctor or pharmacist if you are not sure about your dose or about how to take this medicine. Only your doctor will determine your dose and how you should take this medicine. The recommended dosage is usually one injection every month.

The doctor or nurse will inject the medicine intramuscularly.

Do not exceed the recommended dose.

If you have accidentally injected an overdose or if a child has accidentally swallowed some medicine, immediately see a doctor or go to a hospital emergency room and bring the medicine package with you.
If you forgot to receive an injection at the scheduled time (once a month), consult your doctor.

Adhere to the treatment as recommended by your doctor. Even if your health improves, do not stop taking this medicine without consulting your doctor. This is especially important for breast cancer patients using Decapeptyl Depot together with an aromatase inhibitor since treatment discontinuation may cause an increase in estrogen level. Your doctor will monitor estrogen levels during treatment with Decapeptyl Depot. If you stop the treatment with Decapeptyl Depot, you must also discontinue the treatment with aromatase inhibitor within 1 month after receiving the last dose of Decapeptyl Depot.

Do not take medicines in the dark! Check the label and dose every time you take medicine. Wear glasses if you need them.

If you have any further questions about using this medicine, consult your doctor or pharmacist.

4. Side effects

Like with all medicines, using Decapeptyl Depot 3.75 mg may cause side effects in some users. Do not be alarmed by this list of side effects; you may not experience any of them.

Contact your doctor immediately if you develop symptoms of an allergic reaction or anaphylactic reaction, which may include dizziness, swelling of the face, pharynx or tongue, difficulty swallowing, hives and breathing difficulties (angioedema).
Uncommon cases of post injection tenderness in the injection area in case of subcutaneous injection have been reported with other triptorelin agents.

Men

Very common side effects (effects occurring in more than one in ten users)

- Hot flushes
- Weakness
- Hyperhidrosis
- Back pain
- Sensation of tingling in the legs
- Impotence, decreased libido

Common side effects (effects occurring in 1-10 in 100 users)

- Nausea, dry mouth
- Pain, bruising, redness and swelling at the injection site, edema (accumulation of fluids in body tissues)
- Musculoskeletal pain, pain in the arms and legs, lower abdominal pain
- High blood pressure
- Allergic reactions
- Weight gain
- Dizziness, headache
- Loss of libido
- Depression, mood swings

Uncommon side effects (effects occurring in 1-10 in 1000 users)

- Increased platelet count
- Palpitations
- Tinnitus, vertigo, visual disturbances
- Abdominal pain, constipation, diarrhea, vomiting
- Somnolence, chills accompanied by sweating and fever, pain
- Effect on blood test (including increased liver enzymes), increased blood pressure
- Weight loss
- Loss of appetite, increased appetite, gout (intense pain and swelling in the joints, usually in the thumb), diabetes, increased blood lipids
- Arthralgia, muscle cramps, muscle weakness, myalgia, swelling and tenderness, bone pain
- Sensation of tingling and numbness
- Insomnia, irritability
- Gynecomastia in men, breast pain, testicular atrophy, testicular pain
- Difficulty breathing
- Acne, hair loss, pruritus, rash, skin redness, urticaria
- Nocturia, urination disorders
- Nosebleed

Rare side effects (effects occurring in 1-10 in 10,000 users)

- Red or purple spots on the skin

- Abnormal sensation in the eye, visual disturbances
- Sensation of abdominal fullness, flatulence, changes in the sense of taste
- Chest pain
- Difficulty standing
- Flu-like symptoms, elevated body temperature
- Anaphylactic reactions (a severe allergic reaction which may cause dizziness and difficulty breathing)
- Nasopharyngitis
- Joint stiffness, joint swelling, muscle stiffness, osteoarthritis
- Impaired memory
- Sensation of confusion, decreased activity, sensation of euphoria
- Dyspnea in lying position
- Blisters
- Decreased blood pressure
- Increased liver enzymes

Side effects of unknown frequency (the frequency of these effects has not been established yet):
Anaphylactic reactions (severe allergic reactions with fever, rash, swelling and sometimes decreased blood pressure), ECG changes (QT interval prolongation), general feeling of illness, anxiety, swelling of the face, tongue or pharynx, difficulty swallowing, urticaria and difficulty breathing (angioedema), urinary retention.
If there is a tumor on the pituitary gland, there is an increased risk of bleeding in this area.
Patients receiving long-term treatment with a GnRH analog in combination with radiation may suffer from numerous side effects, particularly in the stomach and gut, relating to the radiation treatment.

Women

Breast cancer

The following side effects have been observed upon administration of Decapeptyl Depot combined with tamoxifen or aromatase inhibitor for treatment of breast cancer

Very common side effects (effects occurring in more than one in ten users)
- Nausea
- Sensation of extreme tiredness
- Myalgia and arthralgia
- Decrease in bone density
- Hot flushes
- Hyperhidrosis
- Sleep disorders
- Depression
- Decreased libido, vaginal dryness, pain during and after sex
- Urinary retention
- Increased blood pressure

Common side effects (effects occurring in 1-10 in 100 users)
- Diabetes
- Hyperglycemia
- Pain
- Redness, swelling and bruising at the injection site
- Allergic reactions
- Bone fractures
- Blood clots in blood vessels

Uncommon side effects (effects occurring in 1-10 in 1000 users)
- Cerebral hemorrhage
- Deficiency in blood supply to the brain or heart

Rare side effects (effects occurring in 1-10 in 10,000 users)
- ECG changes (QT interval prolongation)

Endometriosis
Very common side effects (effects occurring in more than one in ten users)

- Sleep disorders, mood changes
- Headache
- Hot flushes
- Hyperhidrosis, acne, oily skin
- Breast disorders, pain during sexual intercourse, genital bleeding, decreased libido, ovarian hyperstimulation, ovarian hypertrophy, pelvic pain, vaginal dryness

Common side effects (effects occurring in 1-10 in 100 users)
- Nausea, abdominal pain or discomfort
- Arthralgia, muscle cramps, pain in the arms and legs
- Breast pain
- Pain, bruising, redness and swelling at the injection site
- Swelling and tenderness
- Tiredness
- Allergic reaction
- Weight gain
- Depression, irritability
- Dizziness

Uncommon side effects (effects occurring in 1-10 in 1000 users)
- Rapid heartbeats
- Vertigo
- Dry eyes, blurred vision
- Distension, vomiting, flatulence, dry mouth, mouth ulcers
- Weight loss
- Decreased appetite, fluid retention
- Back pain, myalgia
- Changes in the sense of taste, sensory loss, temporary loss of consciousness, memory loss, lack of concentration, prickling or numbness, involuntary shaking
- Mood swings, anxiety, confusion
- Bleeding after sexual intercourse, organ prolapse, irregular menstrual period, heavy and painful periods, small ovarian cysts (swelling) which may be painful, vaginal discharge
- Difficulty breathing, nosebleed
- Hair loss, hirsutism on the body
- Dry skin, fragile nails, pruritus, skin rash

Side effects of unknown frequency (the frequency of these effects has not been established yet):
Anaphylactic reactions (severe allergic reactions with fever, rash, swelling and sometimes decreased blood pressure), confusion, visual disturbances, diarrhea, swelling of the face, tongue or pharynx, difficulty swallowing, urticaria and difficulty breathing (angioedema), muscle weakness, amenorrhea, elevated body temperature, general feeling of illness, change in blood counts (including raised liver function tests), increased blood pressure.
If there is a tumour on the pituitary gland, there is an increased risk of bleeding in this area.
In the treatment of endometriosis, the symptoms of the disease (back pain, period pain) may become worse at the beginning, but should disappear within 1 to 2 weeks. Although this can happen even if the treatment is successful, you should inform your doctor immediately.

Children
Very common side effects (effects occurring in more than one in ten users)
- Vaginal bleeding may occur in girls in the first month of treatment

Common side effects (effects occurring in 1-10 in 100 users)
- Allergic reactions
- Headache
- Hot flushes
- Abdominal pain
- Pain, bruising, redness and swelling at the injection site
- Weight gain
- Acne

Uncommon side effects (effects occurring in 1-10 in 1000 users)
- Blurred vision
- Vomiting, constipation, nausea
- General feeling of illness
- Overweight
- Neck pain
- Mood changes
- Breast pain
- Nosebleed
- Pruritus, rash or urticaria

Side effects of unknown frequency (the frequency of these effects has not been established yet):
Anaphylactic reactions (severe allergic reactions with fever, rash, swelling and sometimes decreased blood pressure), mood changes, depression, irritability, visual disturbances, swelling of the face, tongue or pharynx, difficulty swallowing, urticaria and difficulty breathing (angioedema), myalgia, changes in blood tests, including hormone levels, increased blood pressure

If you experience any side effect, if any side effect gets worse, or if you experience a side effect not mentioned in this leaflet, consult your doctor.

Reporting side effects
You can report side effects to the Ministry of Health by following the link 'Reporting Side Effects of Drug Treatment' on the Ministry of Health home page (www.health.gov.il) which links to an online form for reporting side effects. You can also use this link: https://sideeffects.health.gov.il

5. How to store the medicine?

Prevent poisoning! To prevent poisoning, keep this, and all other medicines, in a closed place, out of the reach and sight of children and/or infants. Do not induce vomiting unless explicitly instructed to do so by a doctor.
Do not use the medicine after the expiry date (exp. date) which is stated on the package. The expiry date refers to the last day of that month.

Storage conditions
Store below 25˚C, use the suspension immediately after reconstitution.

6. Additional information

In addition to the active ingredient, this medicine also contains:
poly (d,l-lactide-co-glycolide), mannitol, carmellose sodium, polysorbate 80.
Solvent composition: water for injection.

What the medicine looks like and contents of the pack:
The medicinal product contains powder and solvent for generation of delayed release suspension for injection.
The powder is white to off-white and the solvent is a clear solution.
The medicinal product package contains one vial of powder, one ampoule of solvent. In addition, the package contains a syringe and two needles.

Registration holder's name and address:
Ferring Pharmaceuticals Ltd., 8 Hashita St., Industrial Park, Caesarea 3088900

Manufacturer's name and address:
Debiopharm Research & Manufacturing S.A., Martigny, Switzerland

This leaflet was revised in February 2022 according to the Ministry of Health guidelines.

Registration number of the medicine in the Ministry of Health's National Drug Registry:
132-28-28860