PATIENT PACKAGE INSERT IN ACCORDANCE WITH THE PHARMACISTS' REGULATIONS (PREPARATIONS) - 1986

This medicine is dispensed with a doctor's prescription only

DANOL 200 mg Capsules

SANOFI

The active ingredient and its quantity:
Each capsule contains: Danazol 200 mg
Inactive ingredients: see section 6.

Read this leaflet carefully in its entirety before using this medicine.

Keep this leaflet; you may need to read it again.

This leaflet contains concise information about the medicine.

If you have further questions, refer to the doctor or pharmacist.

This medicine has been prescribed for you. Do not pass it on to others. It may harm them even if it seems to you that their medical condition is similar.

If a side effect worsens or if a side effect not mentioned in this leaflet occurs, please refer to a doctor or pharmacist.

1. WHAT IS THE MEDICINE INTENDED FOR?

Therapeutic activity: This medicine is intended for treatment of endometriosis amenable to hormonal management and for treatment of hereditary angioedema in both sexes.

Therapeutic group: A synthetic steroid with antiandrogenic properties.

2. BEFORE USING THE MEDICINE

☒ Do not use the medicine if:

- You are sensitive to danazol or to any of the ingredients included in the medicine (see section 6). Signs of an allergic reaction include: rash, swallowing or breathing problems, swelling of the lips, face, throat or tongue.
- You have a rare inherited illness which affects metabolism (porphyria).
- You are pregnant, might become pregnant, or think you may be pregnant.
- You are breastfeeding.
- You have severely impaired function of the kidney, liver or heart.
- You have previously had internal blood clots (thrombosis).
- You have a type of cancer which is affected by hormones.
- You have vaginal bleeding which has not been checked by a doctor.

Special warnings regarding use of the medicine:

Before treatment with Danol, inform your doctor if you are suffering, or have suffered in the past, from the following conditions:

- Impaired function of the liver or kidney/urinary tract.
- An illness which could be aggravated by fluid retention.
- Hypertension or a cardiovascular disease.
- Diabetes.
- A blood system disorder (hypercoagulation or polycythaemia).
- Epilepsy.
- A blood lipid disorder.
- You have had a bad reaction to a hormonal treatment similar to Danol.
- You suffer, or have suffered in the past, from migraines.
- Cancer or suspicion of breast cancer or any other cancer.

This medicine may cause particular sensitivity upon exposure to sun; therefore, avoid exposure to sun and be sure to have proper protection (long clothes, hat, sunscreens, etc.).

Use of this medicine may increase the risk of developing ovarian cancer in women with endometriosis.

This treatment may affect the blood sugar homeostatis in diabetes patients.

❗ If you are taking, or have recently taken, other medicines, including non-prescription medicines and nutritional supplements, inform the doctor or pharmacist. This is because Danol may affect the activity of other medicines and other medicines can affect the activity of Danol. It is especially important to inform the doctor or pharmacist if you are taking:

- Steroids such as testosterone, estrogen, progesterone (including oral contraceptives and hormone replacement therapy).
- Statins such as simvastatin, atorvastatin, and lovastatin. Danol may increase the risk of muscle weakness or rapid muscle breakdown.

Danol may increase the effect of the following medicines:

- Medicines for treatment of epilepsy, fits or convulsions (anti-convulsants).
- Medicines to thin the blood (anticoagulants such as warfarin and coumadin).
- Anesthetics (see section "Operations and tests" below).
- Ciclosporin and tacrolimus – medicines to prevent transplant rejection. Danol may increase the levels of these medicines in the blood and cause kidney damage.
- Alpha-calcidol (a form of vitamin D) used to treat vitamin D deficiency and illnesses where there is not enough calcium in the blood.

Danol may lower the effect of the following medicines:

- Medicines for treatment of diabetes. Sometimes, diabetes patients being treated with insulin have to increase their insulin dosage.
- Medicines for treatment of hypertension.
- Medicines for treatment of migraines.

❗ Operations and tests

If you are due to have an operation with anesthetics, tell the doctor you are taking Danol (Danol can increase the effect of anesthetics).

During the course of treatment with the medicine, you should be under regular medical surveillance and have blood tests and tests for liver function performed.

Before any laboratory test, report that you are taking this preparation, as it may cause changes in certain laboratory test results.

❗ Taking Danol with food and drink/alcohol consumption

Do not drink wines or alcoholic beverages during the course of treatment with Danol. Use of alcohol may cause nausea or shortness of breath.

❗ Pregnancy and breastfeeding

Do not take Danol if you are pregnant, can become pregnant or if you suspect you are pregnant. If you think you may have become pregnant during the course of treatment with Danol, discontinue treatment and consult the doctor.

Do not breastfeed during the course of treatment with Danol, as small amounts of the medicine may pass into the breastmilk.

❗ Important information regarding some of the ingredients of the medicine

The preparation contains lactose. People sensitive to lactose should consult the doctor before use.

3. HOW SHOULD YOU USE THE MEDICINE?

Always use according to the doctor's instructions. Check with the doctor or pharmacist if you are unsure.

Directions for use

Do not chew! Swallow the capsule whole, with water.

If you feel that the effect of the medicine is not strong enough or is too strong, do not change the dosage. Consult the doctor.

During the course of treatment with Danol, prevent pregnancy! It is recommended to begin treatment on the first day of menstruation. If not, perform a pregnancy test to rule out pregnancy before beginning treatment. Use a non-hormonal contraceptive. Do not use oral contraceptives or hormonal contraceptives until completion of treatment with Danol.

Dosage

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

The usual dosage is generally:

For treatment of endometriosis – 200 mg to 800 mg per day for 3-6 months.

For treatment of hereditary angioedema – starting dosage of 200 mg, two to three times a day. Afterwards, the dosage will be reduced in accordance with the doctor's instructions.

The treatment is not recommended for children and the elderly.

Do not exceed the recommended dose.

If you accidentally took a higher dosage, or if a child accidentally swallowed the medicine, immediately refer to a doctor or proceed to a hospital emergency room and bring the package of the medicine with you.

If you forget to take the medicine at the required time, take a dose as soon as you remember. However, if it is almost time for the next dose, skip this dose and take the next dose at the regular time and consult a doctor. Do not take a double dose to compensate for the forgotten dose.

If you stop taking the medicine

Adhere to the treatment regimen recommended by the doctor. Even if there is an improvement in your health, do not discontinue treatment with the medicine without consulting a doctor. Discontinuing treatment with the medicine without consulting a doctor may make the disease worse.

Blood tests

During the course of treatment with the medicine, you should be under regular medical monitoring and have general and liver function blood tests performed.

Taking Danol may affect the results of some blood tests. This includes the following tests:

- Levels of the hormone testosterone.
- Liver and thyroid functions.
- Lipid, sugar and protein levels in the blood.

If you are going to have blood tests, tell your doctor you are taking Danol.

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

If you have further questions regarding use of this medicine, consult the doctor or pharmacist.

4. SIDE EFFECTS

As with any medicine, use of Danol may cause side effects in some users. Do not be alarmed when reading the list of side effects. You may not experience any of them.

Stop taking Danol and immediately refer to a doctor or to a hospital in the following situations:

- Onset of an allergic reaction. The signs may include a rash, swallowing or breathing problems, swelling of the lips, face, throat or tongue.
- Onset of pain or tightness in the chest, jaw or arms - these could be symptoms of a heart attack.

Stop taking Danol and refer to a doctor immediately if you experience any of the following symptoms – you may need urgent medical treatment:

- Severe headache and nausea (feeling sick).
- Enlargement of the clitoris.
- Blurred vision, problems with eyesight, problems wearing contact lenses.
- Liver problems that may cause yellowing of the eyes or skin (jaundice).
- Pain in the liver or liver failure that may cause a swollen abdomen, disorientation and confusion (may be due to hepatic injury or non-cancerous liver tumor).
- Pain when moving arms or legs (may be caused by a blood clot).
- Feeling weak together with numbness in the arms or legs, which you may not be able to move (this may indicate a stroke).
- Bruising that occurs more easily, infections that occur more easily (may indicate a blood problem).
- Any other severe symptoms which you cannot explain.

Inform the doctor immediately if you experience any of the following symptoms:

- Head hair loss.
- Excessive hairiness on the face and body.
- Sore throat, hoarse voice or change in voice (higher or lower than usual).
- Skin rash or blistering, changes in skin color or sensitivity to the sun.
- Blood in the urine.
- Stronger migraine than usual.
- Worsening of epilepsy.
- Pain in the abdomen or chest.

Tell the doctor if one of the following symptoms gets worse or persists for more than a few days:

- Weight gain, or increased appetite.
- Spots on the skin, acne, greasy skin or rash.
- Changes in menstrual cycle, vaginal dryness, change in sex drive.
- High fever with skin rash.
- Backache, muscle cramps and twitching, pain or swelling in joints, arms or legs.
- Headache, feeling tired.
- Flushing.
- Feeling depressed, anxious or more nervous than usual.
- Decreased breast size.
- Retention of fluids or bloating.
- Nausea, dizziness or balance problems (vertigo).
- Palpitations, fast heartbeat, high blood pressure.
- Lowered fertility in men (may be caused by a reduced quantity of sperm).

If you experience any side effects not mentioned in this leaflet, if any of the side effects worsens, or if there is a change in your general health, consult the doctor immediately.

Side effects can be reported to the Ministry of Health by clicking on the link "Report Side Effects of Drug Treatment" found on the Ministry of Health homepage (www.health.gov.il) that directs you to the online form for reporting side effects, or by entering the link:
https://forms.gov.il/globaldata/getsequence/getsequence.aspx?formType=AdversEffectMedic@moh.gov.il

5. HOW SHOULD THE MEDICINE BE STORED?

Avoid poisoning! This medicine and any other medicine must be kept in a closed place out of the reach of children and/or infants to avoid poisoning.

Do not induce vomiting without explicit instruction from the doctor!

Do not use the medicine after the expiry date (exp. date) that appears on the package. The expiry date refers to the last day of that month.

Store at a temperature below 30°C.

Store in the original package.

6. FURTHER INFORMATION

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

Lactose monohydrate, maize starch, talc, magnesium stearate, gelatin, titanium dioxide, red iron oxide, yellow iron oxide.

The ink used to print on the capsules contains:

Shellac glaze 45% (20% esterified) in ethanol, propylene glycol, black iron oxide, ammonium hydroxide 28%.

or:

Shellac, propylene glycol, strong ammonia solution, potassium hydroxide, black iron oxide.

Each capsule contains 76.6 mg lactose monohydrate.

What the medicine looks like and the contents of the package:

Hard white and orange capsules that contain a white or almost-white powder. "D200" is printed on the capsule. Package of 60 capsules is available.

- License holder and address: sanofi-aventis Israel ltd., P.O.B. 8090, Netanya 4250499.
- Manufacturer and address: Zentiva K.S., Prague, Czech Republic or Sanofi Synthelabo Ltd., UK.
- This leaflet does not contain all the information about your medicine. If you have any question or are not sure about anything, please ask your doctor.
- This leaflet was checked and approved by the Ministry of Health in May 2016.
- Registration number of the medicine in the National Drug Registry of the Ministry of Health: 1420822276