PATIENT PACKAGE INSERT IN ACCORDANCE WITH THE PHARMACISTS' REGULATIONS (PREPARATIONS) - 1986
The medicine is dispensed with a doctor's prescription only

Granupas®
Gastro-resistant granules

The active ingredient and its quantity per dosage unit
Each sachet contains: 4 grams of para-aminosalicylic acid
Inactive and allergenic ingredients in the medicine are detailed in section 6 "Further information".

Read this leaflet carefully in its entirety before using the medicine. This leaflet contains concise information about the medicine. If you have further questions, refer to the doctor or pharmacist.
This medicine has been prescribed to treat your ailment. Do not pass it on to others. It may harm them, even if it seems to you that their medical condition is similar.

1. WHAT IS THE MEDICINE INTENDED FOR?
Granupas is intended to treat adults and children over 28 days of age with tuberculosis, as part of a combination medicinal treatment of tuberculosis which is resistant to several medicines. This is when an effective treatment regimen cannot be achieved without using Granupas, due to resistance or intolerability.

Therapeutic group: anti-mycobacterials.

2. BEFORE USING THE MEDICINE
Do not use the medicine if:

- you are sensitive (allergic) to the active substance para-aminosalicylic acid, or to any of the additional ingredients contained in the medicine (see section 6 in the leaflet).
- you are suffering from severe kidney disease.

Special warnings regarding use of the medicine
Before beginning treatment with Granupas, tell the doctor if:
- you have liver problems or a mild or moderate kidney disease
- you have a stomach ulcer
- you are infected with HIV

Children
This preparation is not intended for newborn babies (under 28 days of age).

Drug interactions
If you are taking, or have recently taken, other medicines, including non-prescription medicines and nutritional supplements, tell the doctor or pharmacist, especially if you are taking:
- Antituberculosis medicines or ethionamide (another treatment against tuberculosis)
- Vitamin B12
- Digoxin (for heart disease)
- Diphenylhydramine (for allergic reactions)

Pregnancy and breastfeeding
If you are pregnant or breastfeeding, think you may be pregnant or are planning to have a baby, consult the doctor or pharmacist before taking this medicine.
- Granupas is not recommended for use during pregnancy. Only use Granupas during pregnancy if advised by your doctor.
- Do not breastfeed whilst taking Granupas. This is because small amounts of the medicine can pass into breast milk.

Driving and operating machinery
Use of this medicine requires caution. Granupas may affect your ability to drive and operate machinery.

3. HOW SHOULD YOU USE THE MEDICINE?
Always use the medicine according to the doctor's instructions. Check with the doctor or pharmacist if you are uncertain about the dosage and treatment regimen of the preparation.
The dosage and treatment regimen will be determined by the doctor only. The usual dosage is generally:
Adults:
The recommended dosage is one sachet, three times a day, with a schedule of one sachet every 8 hours. The doctor may recommend to start the treatment with a lower dose to prevent possible side effects.
Do not take more than 3 sachets per day. This treatment is usually given for two years (24 months).
Method of administration:
- Add the contents of the sachet to tomato or orange juice.
- Drink it straight away.
- If some granules are left in the glass, add a little juice and drink it straight away.

Infants, children and adolescents:
The dose for infants, children and adolescents will be calculated by the doctor based on the patient's body weight. The recommended total dose per day is 150 mg for each kg of body weight. This daily amount is divided into two doses administered through the day.
Method of administration:
- Use the measuring spoon included in the package of the preparation to measure the dose.
- To measure the dose:
  - The lines on the measuring spoon indicate the amount (in milligrams of para-aminosalicylic acid). Take the exact amount prescribed by the doctor.
  - Put the granules directly onto the spoon.
  - Tap the spoon once on a table to horizontally level the granules and continue filling if necessary.
- Sprinkle the granules onto applesauce or yogurt.
- Make sure your child eats it straight away.

Do not exceed the recommended dosage.

When taking this medicine:
- Do not crush or chew the granules. Swallow the granules whole. It is important that you do not dissolve, crush or chew the granules as this may interfere with their absorption, as well as causing stomach ache or bleeding.
- Do not use the sachet if it is swollen or if the granules have lost their light brown color.
- You may notice granules in your stools; this is normal.

If you accidentally took 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.

If you forget to take the medicine
If you forgot to take this medicine at the scheduled time, do not take a double dose. Wait until the time for the next dose and then take your usual dose.
Adhere to the treatment regimen as recommended by the doctor.

Do not take medicines in the dark! Check the label and the dose <u>each time</u> you take medicine. Wear glasses if you need them.
If you have further questions regarding use of the medicine, consult the doctor or pharmacist.

4. SIDE EFFECTS
As with any medicine, use of Granupas may cause side effects in some users. Do not be alarmed by the list of side effects. You may not suffer from any of them.
During the first 3 months of your treatment with Granupas, you must be attentive to any sign of an allergic reaction (for example skin eruption, itchy red spots on the skin, itching, rash, itchy and watery eyes or a stuffy nose) or hepatitis (for example, fever, fatigue, dark urine, pale stools, abdominal pain, yellowing of the skin and eyes).
If you experience any of these symptoms, refer to a doctor immediately.

<u>Common side effects</u> (may affect more than 1 in 100 patients):
Dizziness, abdominal pain, vomiting, nausea, bloating, diarrhea, soft stools, skin redness or rash, disturbance of gait and balance.

<u>Uncommon side effects</u> (may affect more than 1 in 1,000 patients):
Loss of appetite (anorexia).

<u>Rare side effects</u> (may affect more than 1 in 10,000 patients):
Thyroid gland problems*, reduced ability to absorb nutrients from food, ulcer, bleeding in the gut, yellowing of the skin or eyes (jaundice), metallic taste, itchy rash.
(*) Thyroid gland problems and specifically an underactive thyroid or low levels of thyroid hormones, are very common side effects (can occur in more than 1 in 10 people) in patients who are also HIV carriers. Regular monitoring of thyroid function is indicated for HIV carriers.

<u>Very rare side effects</u> (may affect less than 1 in 10,000 patients):
Reduction in the number of red blood cells, reduction in the number of white blood cells, reduction in the number of blood platelets, red spots on the skin, reduction of the ability of red blood cells to release oxygen, low levels of blood sugar, tendon pain, headache, vision abnormalities, nerve damage in the hands and feet, dizziness, prolonged bleeding time, destruction of liver cells, elevated liver enzymes, weight loss, crystals in urine.

<u>Side effects of unknown frequency</u> (effects whose frequency has not yet been determined):
Hepatitis.

If a side effect occurs, if one of the side effects worsens, or if you are suffering from a side effect not mentioned in the leaflet, consult with the doctor.

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://sideeffects.health.gov.il

5. HOW SHOULD THE MEDICINE BE STORED?
- Avoid poisoning! This medicine and any other medicine must be kept in a safe place out of the reach and sight of children and/or infants in order 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) that appears on the package. The expiry date refers to the last day of that month.

Storage conditions:
- Do not store above 25°C.
- Shelf-life after first opening the sachet: The sachet can be stored below 25°C for up to 24 hours after first opening.
- Do not use the medicine if the sachets are swollen or if the granules are dark brown or purple.

6. FURTHER INFORMATION
- In addition to the active ingredient, the medicine also contains:
  microcrystalline cellulose, methacrylic acid – ethyl acrylate copolymer (1:1) dispersion 30%, talc, dibutyl sebacate, hypromellose, colloidal hydrated silica.
- What the medicine looks like and the contents of the package:
  Light brown gastro-resistant granules, packaged in sachets. Each box contains 30 sachets and a measuring spoon.
- Registration Holder and address: Truemed Ltd., 10 Benny Gaon Street, Netanya 4250499.
- Manufacturer and address: Eurocept International B.V. (Lucane Pharma) Ankeveen, Netherlands.
- This leaflet was revised in December 2021 according to MOH guidelines.
- Registration number of the medicine in the National Drug Registry of the Ministry of Health: 160-83-34974

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