

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

This medicine can be sold with a doctor's prescription only

Gliolan Powder for oral solution

Active ingredient and its quantity in a dosage unit:

Each vial contains:

5-aminolevulinic acid hydrochloride 1.5gr

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

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

1. What is the medicine intended for?

- visualization of certain brain tumors (called malignant glioma) during brain surgery.

Therapeutic group: Imaging agents

2. Before you take the medicine:

| Do not use the medicine if: |
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| <ul style="list-style-type: none">• you are hypersensitive (allergic) to the active ingredient (5-aminolevulinic acid hydrochloride), or to porphyrin.• in case of porphyria or suspected acute or chronic porphyria (acquired or inherited disorder of certain enzymes in the synthesis pathway of the red blood pigment).• you are pregnant or suspect you are pregnant. |

Special warnings regarding the use of this medicine:

Before treatment with Gliolan tell the doctor if:

You have a heart disease or had heart disease in the past.

- The eyes and skin should be protected from exposure to strong light (such as: direct sunlight or bright lighting) for a period of 24 hours after taking the medicine.
- If you have a heart disease or had heart disease in the past – you should tell your doctor. Use with caution since blood pressure may drop.
- No studies have been performed in patients with impaired liver or renal function - use this medicine with caution in these patients.

Children and adolescents

This drug is not intended for children and adolescents under the age of 18, as there is no experience with Gliolan in this age group.

Tell the doctor or pharmacist if you are taking or have recently taken any other medicines, including non-prescription drugs and nutrition supplements.

Especially inform the doctor or pharmacist if you are taking or plan on taking:

- Medicines or herbs that may cause skin problems when the skin is exposed to light (such as antibiotics or hypericum extracts [St. John's Wort]) - do not take these products for up to two weeks after taking Gliolan.
- Avoid taking other medicines that may harm the liver for 24 hours after taking Gliolan

Use of this medicine and food

The medicine is usually administered once, mainly 2-4 hours before anesthesia for surgery of brain tumors called glioma. Do not drink or eat at least 6 hours before the anesthesia.

Pregnancy and Breastfeeding

- It is unknown whether the medicine is harmful to the fetus. Do not use if you are pregnant.
- It is unknown whether the medicine passes into breast milk. Do not breastfeed for 24 hours after taking the medicine.

Driving and use of machinery:

This medicine does not affect the ability to drive or operate machinery.

3. How to use this medicine

- The medicine should always be used according to the doctor's instructions. Check with the doctor or pharmacist if you are not sure regarding the dosage and manner of treatment with this medicine.

The dosage and administration will be determined by the doctor only.

Usual recommended dosage:

20 mg per kilogram body weight. The exact dosage will be calculated by the nurse or pharmacist.

Do not exceed the recommended dose.

- The medicine contains a powder that needs to be mixed with a glass of water before use. Preparation of the medicine will always be done by a nurse or pharmacist.
- Drink the prepared solution 2-4 hours before anesthesia.
- If the anesthesia/surgery is delayed by a few hours, do not give an additional dose of this medicine.
- If surgery is delayed by a day or more, another dose of this medicine can be taken 2-4 hours before anesthesia.

If you have taken an overdose or if a child has accidentally swallowed the medicine, proceed immediately to a doctor or a hospital emergency room and bring the package of the medicine with you. The doctor will decide on necessary measures to avoid problems, including sufficient protection from strong light (such as direct sunlight).

This medicine is given once on the day of surgery, 2-4 hours before the start of anesthesia. **If you forget to take the medicine** during this time period, it is not recommended to take it immediately before the start of anesthesia. In this case, if possible, postpone the anesthesia and surgery by at least two hours.

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 regarding the use of this medicine, consult a doctor or pharmacist.

4. **Side Effects**

Like all medicines, the use of Gliolan may cause side effects, although not everybody gets them. Do not be alarmed while reading the list of side effects, you may not suffer from any of them.

The most serious side effects include: mild alterations of blood cell counts (red and white blood cells, platelets), neurological disorders (disorders that affect the nervous system such as: hemiparesis [partial paralysis on one side of the body] and thromboembolism (a blood clot that may obstruct blood vessels).

Additional frequently observed side effects are: vomiting, nausea and slight increase of certain enzymes (transaminases, gamma-glutamyl transferase, amylase) or bilirubin (a bile pigment produced in the liver by the breakdown of red blood cells) in the blood.

Tell the doctor immediately if you have any complaints.

Side effects may occur as a result of taking the drug before anesthesia, or as a result of taking the drug along with the anesthesia and removal of the tumor.

After taking Gliolan and before the start of anesthesia, the following side effects may occur:

Uncommon side effects (may affect up to 1/100 in 1/1000 people):

- nausea, decrease of blood pressure, skin reactions (such as rash, sunburn like appearance), brain edema (swelling of the brain).

In combination with anesthesia and removal of the tumor, additional side effects may occur:

Very common side effects (may affect more than 1 in 10 people):

- mild alterations of blood cell counts (red and white blood cells, platelets), slight increase of certain enzymes (transaminases, gamma-glutamyl transferase, amylase) or bilirubin (a bile pigment produced in the liver by the breakdown of red blood cells) in the blood. These changes reach their peak between 7-14 days after surgery. These changes will completely resolve within a few weeks. Usually, you will not experience any symptom when these changes occur.

Common side effects (may affect up to 1/10 in 1/100 people):

- nausea, vomiting, neurological disorders - disorders that affect the nervous system such as: hemiparesis (partial paralysis on one side of the body), aphasia (partial or total loss of the ability to use or understand language), convulsions (seizures) and hemianopia (blindness in half the vision field of one or both eyes), thromboembolism (a blood clot that may obstruct blood vessels).

Very rare side effects (may affect up to 1 in 10,000 people) or **unknown frequency** (cannot be estimated from the available data) :

- hypoesthesia (reduced sense of touch), diarrhea (loose or watery stools).

If a side effect has occurred, or if you experience a side effect not mentioned in the leaflet, consult a doctor.

Reporting of suspected adverse reactions

Reporting suspected adverse reactions after authorisation of the medicinal product is important. It allows continued monitoring of the benefit/risk balance of the medicinal product. Any suspected adverse events should be reported to the Ministry of Health

according to the National Regulation by using an online form:

<https://sideeffects.health.gov.il/>.

4. How to store the medicine

- **Avoid poisoning!** This medicine and all other medicines must be stored in a safe 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.
- Store at temperature that does not exceed 25°C. Keep the vial in the outer package in order to protect from light.
- Store the prepared solution at 25°C and use within 24 hours once it has been prepared.
- Medicines should not be disposed of via wastewater or household waste. Ask the pharmacist how to dispose of medicines no longer required. These measures will help protect the environment.

5. Additional information:

What the medicine looks like and what the package contains:

This medicine is a powder for oral use. The powder is a white - grayish color. After preparation, the solution is clear and colorless to slightly yellowish in color.

Gliolan packages contain 1, 2 or 10 vials.

Not all pack sizes may be marketed.

Registration holder and address: Tzamal Bio-Pharma Ltd.,
20 Hamagshimim St., Kiryat Matalon, Petach-Tikva.

Manufacturer: Medac Gesellschaft Fur Klinische Spes.GmbH,
Theaterstrasse 6, 22880 Wedel, Germany.

6. Drug registration number at the national medicines registry of the Ministry of Health: 150 85 33857

The format of this leaflet was determined by the Ministry of Health and its content was checked and approved in April 2015.

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