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

The medicine is dispensed with a doctor's prescription only Depalept solution 200 mg/ml

Sodium valproate 200 mg/1 ml

is for the solution

Attention: Depalept solution is 5 times more concentrated than Depalept syrup. Before administering the medicine, make sure that the dosage prescribed by the doctor

nactive ingredients and allergens in the preparation – see section 6 and section 2 Important information about some of the ingredients of the medicine"

Read the entire leaflet carefully before using | patient leaflet before starting to use the the medicine. This leaflet contains concise information about the medicine. If you have any | further review, if necessary. other questions, refer to the doctor or the

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

The medicine is not intended for children veighing less than 17 kg.

Depalept may severely harm the fetus when taken during pregnancy

Neonates born to mothers who have taken

valproate during pregnancy are at an increased risk of serious developmental disorders (mental and physical) and behavioral disorders (approximately 30-40% of cases) and/or congenital malformations (approximately 11% of cases). If you are a woman of childbearing age or if you are pregnant, the doctor will prescribe

valproate for you only if other treatments are

Women of childbearing age should use

effective contraception while taking this medicine. If despite using contraception you disease). doctor immediately in order to discuss the options for alternative therapy, if possible. Do not stop using the medicine without

Huttenlocher syndrome. Have a metabolic disorder, such as urea

· Are currently taking any of the following

depression).

This card contains important safety information that you must know before starting treatment with Depalept and during the treatment, and act accordingly. Please review the patient information card and the preparation. You should keep the card for

What is the medicine intende This medicine is an anticonvulsant, and it is

Therapeutic class: The active ingredient

2. Before using the medicine:

- Or someone in your family have had or the eyes (jaundice).
- especially if it was caused by the use of

- Have hepatic porphyria (a hereditary live)
- become pregnant unintentionally, contact your | | Have a genetic problem causing a mitochondrial disorder, e.g. Alpers
 - cycle disorder.
 - medicines:

Special warnings regarding the use of the

- Women of childbearing age who are
- this medicine plans to become pregnant, she should consult her doctor regarding the possibility of receiving an alternative administered for treatment of certain types of

belongs to the group of anticonvulsants.

consulting the treating doctor, because you

preparation has a patient safety information

In addition to the leaflet, the Depalept

condition may worsen.

- Are sensitive (allergic) to the active ingredient or to any component of the medicine (see section 6 – Additional information).

- currently has a serious liver disease.

- Do not give this medicine to girls, adolescent | check that you are not pregnant and that you months of treatment with this medicine, you girls, women of childbearing age and are using contraception. pregnant women, unless alternative treatments have been found to be unsuitable.
- effective contraception. If a woman taking immediately. maintaining normal body weight.
- ☑ Do not use this medicine if you:
- Are sensitive to another medicine of the valproate family (valproate semisodium.
- Have a liver disease (acute or chronic

- St. John's Wort (for treatment of

Before starting treatment, the doctor will

- this medicine may worsen the seizures or increase their frequency. You may even treated with this medicine should use experience a different kind of seizures.
- This medicine may, in very rare cases, cause | the risk for suicidal actions or thoughts. damage to the liver (hepatitis) or pancreas (pancreatitis), which may be severe and life-threatening.
- Your doctor will refer you to perform blood tests for evaluation of liver function, especially during the first 6 months of treatment. You should consult a doctor immediately if the following effects appear:
- Sudden tiredness, loss of appetite, fatique. drowsiness, swelling in the legs, general weakness.
- · Repeated vomiting, nausea, abdominal pain, yellowing of the skin or the whites of
- Recurrence of epileptic seizures, even though you are taking the medicine correctly.

- Before treatment with Depalept, inform vour doctor if:
- (renal insufficiency), systemic lupus or disturbances of memory or cognitive erythematosus or hereditary enzyme function. The doctor will examine whether deficiency, especially enzyme deficiency in they are caused by an existing condition or the urea cycle which may cause elevation by Depalept. Stopping the treatment may be effect of nimodipine. of ammonium blood levels, or a genetic problem that causes a mitochondrial When treating children, tell your doctor if your certain types of birth control pills) disorder (including members of your
- Before undergoing any type of surgery, inform severe forms of epilepsy. the medical staff that you are taking Depalept.

 Tests and follow-up
- As with other antiepileptic medicines, taking **II Drug interactions** If these effects occur, consult a doctor following medicines:
- This medicine may cause weight gain. Consult your doctor regarding methods fo
- Taking anticonvulsants may increase You and your family members must pay attention to changes in mood, behavior patterns and actions. Watch for signs indicating risk of suicide, such as: talking or thinking about wanting to hurt yourself,
- Medicines containing acetazolamide (for If you suffer from carnitine lowering intraocular pressure or carbon palmitovltransferase (CPT) type II enzyme dioxide levels in the blood). deficiency (hereditary metabolic disease), Antibiotics (medicines containing aztreonam there is an increased risk of muscle or rifampicin). breakdown when taking Depalept.

- You suffer from a kidney disease difficulty walking (extrapyramidal disorders)

 - Certain medicines affect the activity of Do not take Depalept if you are taking the
 - treatment of depression Unless the doctor has instructed you

symptoms such as tremor, limb rigidity and

- following medicines: Lamotrigine (another medicine for the treatment of epilepsy). Medicines of the penem group (carbapenems) (a group of antibiotics for treatment of bacterial infections)
- introversion and withdrawal from family and If you are taking or have recently taken friends, depression or worsening of existing other medicines, including non-prescription • If you are a woman of childbearing age, depression, preoccupation with the subject medicines and nutritional supplements, tell of death, abandoning or giving away prized the doctor or the pharmacist. Especially if possessions. If thoughts of this kind occur. vou are taking:
- refer to the doctor immediately.

Tell your doctor if you are experiencing

- child is taking additional treatment for epilepsy Propofol (anesthetic medicine). or has a neurological or metabolic disease or
- Before starting treatment and during the first six stabilization). should perform blood tests for liver functions. (for treatment of fever and pain). Salicylates (including aspirin).
- Especially in children under the age of 3, avoid the use of medicines containing salicylates • St. John's Wort – an herbal remedy for the (including aspirin) during the treatment with the
- Use of the medicine and food otherwise, do not take Depalept if you are It is best to take the medicine during meals taking, have recently taken or may take the Use of the medicine and alcohol consumption

treatment of HIV).

other diseases).

- Drinking alcoholic beverages is not recommended during treatment with this
- medicine. ■ Pregnancy, breastfeeding and fertility Valproate is harmful to the fetus if taken during pregnancy, therefore:
- the doctor will prescribe this medicine for you only if alternative treatments were found unsuitable.

If you are pregnant, unless alternative

- Read the patient safety information card. Do not take Depalept:
- treatments have been found to be Other medicines for the treatment of epilepsy (medicines containing carbamazepine,

- felbamate, phenytoin, fosphenytoin If you are a woman of childbearing age, th primidone, phenobarbital, rufinamide. doctor will prescribe this medicine for you only if alternative treatments were found
- topiramate or zonisamide). Nimodipine - used for prevention of complications that may occur following cerebral bleeding. Depalept may increase the
- soon as their daughter starts menstruating. Medicines containing estrogen (including Certain types of birth control pills (estrogen-
- Medicines containing zidovudine (for the be the most suitable for you. Medicines containing lithium (for mood
- Risks of valproate when taken during Medicines containing metamizole/dipyrone Valproate may harm the fetus when taken
- during pregnancy. The risk increases with the increased risk of developing symptoms dosage, but it exists with all dosages, even of attention deficit hyperactivity disorders contraception Cannabidiol (for treatment of epilepsy and when valproate is taken in combination with

valproate in utero.

valproate in utero.

- other antiepileptic medicines. Children exposed to valproate in utero are at high risk for severe congenital malformations and developmental disorders.
- reported include spina bifida (a developmental malformation in which the spine does not develop properly), facial, upper lip, palate and skull malformations; your treatment, if possible. malformations of the heart, kidneys, urinary tract and genitals: limb deformities and involvement of multiple other malformations, prescribed for you, the doctor will explain to you inform the doctor before you stop taking your which affect various organs and body parts.
- Congenital malformations may lead to disabilities which may be severe. Hearing disorders and hearing impairment have been reported in children exposed to
- Eve deformities which can affect vision Important issues: have been reported in children exposed to
- Taking valproate during pregnancy increases treatments suitable for you. the risk of giving birth to a child with congenital malformations that will require medical treatment. It has been found that in

- women taking valproate, around 11 babies out of 100 are born with congenital A routine follow-up (at least annually) should the risks to you and to the fetus as much as
- Approximately 30-40% of pre-school Parents or caregivers of girls that are treated children whose mothers have taken with valproate should inform the doctor as problems with early childhood
- containing birth control pills) may reduce the walk and talk, lower intellectual ability. levels of valproate in the blood. Consult the language and memory difficulties. doctor to determine which contraception will Autistic spectrum disorders are more often diagnosed in children exposed to

valproate in utero.

- There is some evidence that children exposed to valproate in utero are at an
- Before giving you the medicine, the doctor will explain to you the possible risks to your baby in the case of exposure to valproate during pregnancy. If you later
- The most common congenital malformations decide that you wish to become pregnant, do not stop taking the medicine or your contraception before consulting with the doctor and considering the option of changing will make sure that you are aware of and Starting treatment with Depalep If this is the first time Depalept has been the possible risks for your baby in the case of
 - exposure to valproate during pregnancy. If you Consult the doctor immediately if you are are a woman of childbearing age, you should use effective contraception while using the medicine. Consult a gynecologist about effective contraception
 - Before starting the treatment, the doctor will and developmental disorders. make sure that there are no other alternative. To the extent possible, consult your treating doctor, as your condition may worsen. Stopping the treatment. doctor before becoming pregnant.
 - Pregnancy should be ruled out before starting Do not stop using the medicine or your contraception before consulting with your doctor. Your doctor will make every effort to find

- Make sure that you are using effective alternative treatments so that the pregnancy • The doctor will instruct you further. will be as normal as possible, while reducing
- out of 100 in the general population. be carried out by a doctor. The doctor will make sure that you are aware of and Your doctor will make every effort to stop the In exceptional cases, when Depalept is the only understand the risks and recommendations treatment with Depalept as early as possible possible treatment during pregnancy: before you become pregnant, to ensure that • Your doctor will be able to refer you to a If you are planning to become pregnant. your disease is stabilized. In exceptional cases. specialist who can provide consultation development, such as: delay in starting to inform the doctor before you stop taking your when this is impossible, read the section **Pregnancy during treatment with Depalept**" • The doctor will try to reduce the dosage of
 - Consult the doctor immediately if you are Consult your doctor regarding taking folic acid valproate pregnant or think you might be pregnant. Continuing Depalept treatment when you are not planning a pregnancy miscarriages, that exists with all pregnancies. development of the fetus.
 - Make sure you are using effective contraception throughout the treatment **period.** Consult a gynecologist about effective

If you are planning to become pregnant.

pregnant or think you might be pregnant.

Treatment with Depalept if you are planning

Children exposed to valproate in utero are at

high risk for severe congenital malformations

 The treating doctor should routinely (at least doctor told you to. congenital malformations associated with annually) evaluate if there are more suitable Do not stop using contraception before you valproate use. treatment alternatives for you. have consulted with your doctor, and worked Before delivery: The doctor will prescribe Make sure that you are using effective together on a treatment plan that will enable

Consult the doctor immediately if you are

of congenital malformations associated with

Do not stop taking Depalept unless your

you to control your epilepsy and reduce the risk to the fetus. A routine follow-up (at least annually) should Consult with your doctor. The doctor will be carried out by a doctor. The doctor understand the risks and recommendations understand the risks and recommendations related to taking valproate during pregnancy.

valproate use.

Important issues:

were treated with valproate during your related to taking valproate during pregnancy. Your doctor will make every effort to stop the pregnancy. The child will be closely monitored maximum recommended daily intake in adults treatment with Depalept as early as possible for neurological development in order to provide the appropriate treatment for him, as before you become pregnant

However, it is unlikely that it will reduce the risk • Consult your doctor regarding taking folic

- pregnant or think you might be pregnant. Important issues: Pregnancy during treatment with Depalept pregnant or think you might be pregnant Children exposed to valproate in utero are at high risk for severe congenital malformations Do not stop taking Depalept unless your and developmental disorders. Do not stop
- The dosage and treatment regimen using the medicine before consulting with the will be determined by the doctor only The doctor may recommend to divide the If you are taking Depalept and you think that • The doctor will give you the full information required dose to twice a day

The doctor will make every effort to

evaluate all alternative treatments.

acid. Taking folic acid can reduce the risk

that exists with all pregnancies. However,

it is unlikely that it will reduce the risk of

vour baby's life or bone deformations.

After delivery: Your baby may receive a

vitamin K injection to prevent bleeding.

The child: Inform the pediatrician that you

of spina bifida and early miscarriages,

you might be pregnant, contact your doctor regarding the risks involved with using Depalept during pregnancy, including the ri for congenital defects and developmental

early as possible, as needed.

- disorders (cognitive, physical and behavioral) It is best to take the medicine during meals. This medicine should be used at set intervals
- discontinue treatment with Depalept and to Make sure you are referred to a specialist for as determined by the treating docto prenatal monitoring in order to detect possible **Do not exceed the recommended dose.** This medicine is not intended for children Inform the pediatrician that you were treated weighing less than 17 k
- regarding pregnancy under valproate. development. Method of administration Breastfeeding The solution should be diluted with a small
- Do not breastfeed while taking Depalept. when planning a pregnancy. Taking folic acid

 • You will be closely monitored to manage unless the doctor recommends otherwise. can reduce the risk of *spina bifida* and early your health condition and to check the Consult the doctor or pharmacist before starting reduced the number of poisoning incidents treatment with any medicine.

oreparation.

Patients with kidney failure: the doctor ma

decide to adjust the dosage.

- Driving and operating machinery you find it difficult to open the package, you ca Use of this medicine may impair alertness, refer to a pharmacist to ask to have the safety especially if taken in combination with other mechanism removed and to turn the cap into a antiepileptic medicines or medicines that cause regular, easy-to-open cap. If you have accidentally taken a higher
- If you experience this effect or if your epilepsy dosage you may suffer from the following is not vet under control and you continue to symptoms: coma, muscle weakness. suffer from seizures, do not drive or operate decreased reflexes, constriction of the pupils certain vitamins for you, so that this medicine dangerous machinery breathing impairment, metabolic acidosis will not cause bleeding during the first days of Children should be cautioned against riding a decreased blood pressure and shock.
 - If you took an overdose or if a child bicycle or playing near a road etc. I Important information about some of the swallowed this medicine by mistake, go to the ingredients of the medicine doctor or the emergency room of the hospital Amount of sodium in each 1 ml: approximately immediately and take the package of the 28 mg, which is equivalent to 1.4% of the
 - If you have forgotten to take this medicine at the required time, do not take a double dos Take the next dose at the scheduled time and blood cells (eosinophils). consult a doctor. Always use the preparation according to the Follow the treatment as recommended by the

Do not stop treatment with the medicine withou

consulting the doctor. Discontinuation of

amount of a non-carbonated drink.

- doctor's instructions. doctor. Even if there is an improvement in you Check with the doctor or pharmacist if you are health, do not stop treatment with Depalept uncertain about the dosage and how to use th without consulting the doctor. If you stop taking Depalept
 - Johnson syndrome). treatment should be done gradually. Additional possible side effects: If you discontinue Depalept treatment abruptly or not according to your doctor's instruction, you might be at an increased risk for seizures.

- discuss the consequences with your doctor. Do not take medicines in the dark! Check Additional side effects: the label and the dose every time you take Very common side effects – side effects that the medicine. Wear glasses if you need
- Duration of treatment: Do not stop treatment If you have any other questions regarding child will be closely monitored for neurological with the medicine without consulting the doctor. use of the medicine, consult the doctor or the pharmacist.

As with any medicine, using Depalept may

Child-proof safety caps have significantly cause side effects in some users. Do not be alarmed when reading the list of side effects. caused by medicines each year. However, if ou may not experience any of them. Memory impairment experience the following side effects:

- Confusion, aggressiveness, nervousness, Damage to the liver (hepatitis) or pancreas attention disorders, hallucinations (seeing or (pancreatitis), which may be severe and life
- nearing non-existing things) threatening. These effects can start abruptl Extrapyramidal disorders (symptoms include: with tiredness, loss of appetite, fatique, tremor, limb rigidity and difficulty walking)* somnolence, nausea, vomiting and intestinal Urinary incontinence
- Quick and involuntary eve movements An allergic reaction which includes:

that may involve the mouth area (erythema

multiforme), emergence of blisters with

- Hearing impairment Sudden swelling of the face and/or neck Gum problems, especially overgrowth of the that causes difficulty breathing and is lifegums (gingival hyperplasia) threatening (angioedema).
 - Pain and swelling in the mouth, ulcers in the Severe allergic reaction that includes mouth and a burning sensation in the mouth symptoms such as: fever, skin rash, (stomatitis) enlarged lymph nodes, liver damage
 - kidney damage, abnormal blood test Irregular menstrual periods
 - results, such as: increase in certain white Nausea or dizziness
- Emergence of a rash on the skin, sometimes Nails and nail bed disorders accompanied by the appearance of blisters Decrease in red blood cell count (anemia
 - and platelet count (thrombocytopenia Low levels of sodium in the blood
- detachment of the skin that can rapidly (hyponatremia, a symptom of improper spread to the entire body and be lifesecretion of antidiuretic hormone) threatening (Lvell's syndrome, Stevens-Uncommon side effects - side effects that occur Do not discontinue the treatment without
- n 1-10 out of 1,000 users: · Alertness impairment which may develop into · Impaired functioning of the ovaries (polycystic Health by clicking on the link "Report side Congenital malformations and physical a temporary coma, with remission after dose ovaries)
- and mental development disorders (see reduction or treatment cessation

- Before discontinuing the treatment you should the section "Pregnancy, breastfeeding and Difficulty in movement synchronization Reversible parkinsonism* Sensation of numbness/tingling in the hands skin rash and fever (systemic lupus
 - color or growth)

At the start of the treatment: vomiting.

abdominal pain, diarrhea

Somnolence

- Common side effects side effects that Hyperandrogenism (symptoms include 1-10 out of 100 users: excessive hairiness, especially in women.
 - woman, acne) Decrease in body temperature (hypothermia)

Breathing difficulties and pain, due to

Decrease in the numbers of all blood cells:

treatment with epilepsy medicines, if you

Rare side effects - side effects that occur i

development of masculine features in a

 Swelling of the extremities (edema) Lack of menstrual period

the lungs (pleural effusion)

white blood cells (leukopenia)

if you are taking corticosteroids

Inflammation of blood vessels

consulting with the doctor first

Behavioral disturbances, increased

1-10 out of 10,000 users:

Rash or hives on the skin

- Worsening and increased frequency of convulsions. Onset of a different type of
- inflammation of the protective membranes of biotinidase Increased ammonium blood levels
- white blood cells, red blood cells and platelets Dementia and cognitive disorders* that appear gradually and recede several weeks What does the medicine look like and what are (pancytopenia), decrease in the number of the contents of the package to several months after stopping treatment

hyperactivity and learning disorders.

Side effects may be reported to the Ministry of

effects due to medicinal treatment" found on

the Ministry of Health website homepage

- Cases of bone damage such as bones that
 Do not discontinue the treatment without. become more fragile (osteopenia), decreased consulting with the doctor first *Symptoms may include signs in brain imaging Name and address of the marketing
- bone density (osteoporosis) and fractures, have been reported. Consult your doctor or (atrophy) Additional side effects in children: pharmacist if you are receiving long-term

Sterility in men, generally reversible 3 months If a side effect occurs, if one of the side

at least after treatment is discontinued, and effects worsens, or if you suffer from a side

potentially reversible after dosage reduction. effect not mentioned in this leaflet, consult

- have or have previously had osteoporosis or frequently in children or are more severe This leaflet was revised in 07/2023 in compared to adults. These effects include liver accordance with the Ministry of Health
 - disorders, behavior that is not within the norm, national drug registry of the Ministry of Health:



online form for reporting side effects, or by clicking on the following link: https://sideeffects.health.gov.il/ 5. How to store the medicine?

medicine must be kept in a closed place out of

the reach and sight of children and/or infants to

avoid poisoning. Do not induce vomiting without

Store at a temperature below 25°C. Can be

used for up to 60 days after opening.

In addition to the active ingredient, the

Urea, diluted hydrochloric acid or sodium

A transparent, amber-colored glass bottle

authorization holder/manufacturer: CTS

6. Additional information

containing a yellowish solution.

medicine also contains:

hydroxide, purified water.

an explicit instruction from the doctor.

(www.health.gov.il), which will direct you to the

erythematosus) Hair problems (changes in the hair structure, Underactive thyroid gland (hypothyroidism) Muscle pain, muscle weakness which m Avoid poisoning! This medicine and any other

- severe (rhabdomyolysis
- Kidnev damage (kidnev failure. tubulointerstitial nephritis. Fanconi syndror Increase in the size of red blood cells

Decreased and irregular blood cell production

psychomotor activity, learning disabilities

An autoimmune reaction with joint pain.

- Do not use the medicine after the expiry date (macrocytosis), a significant decrease in EXP) appearing on the package. The expire white blood cell count (agranulocytosis) date refers to the last day of that month.
- Decrease in coagulation factors, abnormal coagulation test results (such as: increase in partial thromboplastin time and INR time

Decrease in the levels of vitamin B8 (bioting)

- Chemical Industries Ltd., 3 Hakidma St., Kirvat Some side effects of valproate occur more
- damage, inflammation of the pancreas, aggressiveness, nervousness, attention Registration number of the medicine in the