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

The medicine is dispensed with a doctor's prescription only

Baclofen Sintetica 0.5 mg/ml

Baclofen Sintetica 2 mg/ml

Solution for intrathecal infusion

Active ingredient
Baclofen Sintetica 0.5 mg/ml: 1 ml contains 0.5 mg baclofen
Each 20 ml ampoule contains 10 mg of baclofen.
Baclofen Sintetica 2 mg/ml: 1 ml contains 2 mg baclofen
Each 5 ml ampoule contains 10 mg of baclofen.
Each 20 ml ampoule contains 40 mg of baclofen.

Inactive ingredients and allergens: See the paragraph in Section 2 "Important information about some of the ingredients of the medicine" and Section 6 "Further information".
Read the leaflet carefully in its entirety before using the medicine. This leaflet contains concise information about the medicine. If you have any 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 ailment is similar.

1. What is this medicine intended for?

Adults:

For the treatment of patients with severe chronic spasticity resulting from trauma, multiple sclerosis or other spinal disorders, who are unresponsive to oral baclofen or other orally administered antispastic agents and/or those patients who experience unreasonable side effects at effective oral doses.

Baclofen Sintetica is effective in patients with severe chronic spasticity of cerebral origin, resulting e.g., from cerebral palsy, brain trauma or cerebrovascular accident (clinical experience with the treatment of those cases is limited).

Children:

For the treatment of children aged 4 years and above with severe chronic spasticity of cerebral origin or of spinal origin (associated with injury, multiple sclerosis or other spinal disorders), who are unresponsive to oral baclofen or other orally administered antispastic agents and/or those patients who experience unreasonable side effects at effective oral doses.

Therapeutic group: muscle relaxant.

2. Before using the medicine

Do not use the medicine if:

- you are sensitive (allergic) to the active ingredient or to any of the other ingredients contained in the medicine (see Section 6).

Special warnings regarding the use of this medicine

- It is important to make sure that there are no problems with the pump. Get urgent medical help if you observe that the pump is not working and you also notice withdrawal symptoms (see Section 3 "How should you use the medicine?"). Treatment with Baclofen Sintetica

must not be stopped suddenly due to the risk of withdrawal symptoms. Care must be taken not to miss visits to the hospital, during which the pump reservoir is refilled.
- Your doctor may ask you to perform a general examination from time to time while you are being treated with Baclofen Sintetica.
- If you are going to undergo an operation of any kind, make sure that the doctor knows that you are being treated with Baclofen Sintetica.
  Anesthetics such as propofol may increase the risk of side effects.

Before the treatment with Baclofen Sintetica, tell your doctor if:

- you receive any other injections into your spine
- you are suffering from any infection
- you had a head injury within the past year
- you have ever had a crisis caused by a condition called autonomic dysreflexia (your doctor will be able to explain this to you)
- you have had a stroke
- you have epilepsy
- you have a stomach ulcer or any other problem with your digestion
- you suffer from mental illness
- you are being treated for high blood pressure
- you have Parkinson's disease
- you suffer from liver, kidney or lung disease
- you have diabetes
- you have difficulties urinating
- you are pregnant or breastfeeding

If you suffer from any of the above, tell your doctor or nurse as Baclofen Sintetica may not be the right medicine for you.
Tell your doctor immediately if you get any of these symptoms during treatment with Baclofen Sintetica:

- **If you have pain** in the back, shoulders, neck and buttocks during the treatment (a type of spinal deformity called scoliosis).
- **If you have** thoughts of harming or killing yourself at any time, contact your doctor or go to a hospital immediately. Also, ask a relative or close friend to tell you if they are concerned about any changes in your behaviour and ask them to read this leaflet.

Children and adolescents
The medicine is intended for adults and children aged 4 years and above.

Drug interactions

If you are taking or have recently taken other medicines, including non-prescription medicines and nutritional supplements, tell the doctor or pharmacist. In particular if you are taking:

- Other medicines for your spastic condition such as tizanidine or diazepam
- Antidepressants such as imipramine or amitriptyline
- Medicines to treat high blood pressure such as diltiazem or moxonidine
- Other medicines which also affect the kidneys, such as ibuprofen
- Medicines to treat Parkinson's disease such as levodopa or carbidopa
- Medicines to treat epilepsy such as carbamazepine or clonazepam
- Opiates for pain relief such as morphine
- Medicines that depress the nervous system, for example antihistamines such as promethazine and sedatives such as temazepam (some can be purchased without a prescription)

Use of the medicine and alcohol consumption
Be careful when drinking alcohol – it may affect you more than usual.

Pregnancy and breastfeeding
If you are pregnant, planning to get pregnant or breastfeeding, do not use this medicine without consulting your doctor before starting treatment.
If you are pregnant or breastfeeding, consult a doctor or pharmacist before taking medicines.

Driving and using machines
Some people may feel drowsy and/or dizzy or have eye problems during treatment with Baclofen Sintetica. In such a case, one should refrain from driving or performing any action that requires alertness (such as operating tools or machines) until these effects have dissipated.

Important information about some of the ingredients of the medicine
Each ampoule of Baclofen Sintetica 0.5 mg/ml contains
69.3 mg of sodium, which is equivalent to 3.5% of the 2-gram recommended maximum daily dietary intake of sodium for an adult.
Each 5 ml ampoule of Baclofen Sintetica 2 mg/ml contains less than 1 mmol (23 mg) of sodium per ampoule, so it is essentially “sodium free”.
Each 20 ml ampoule of Baclofen Sintetica 2 mg/ml contains 69.3 mg of sodium, which is equivalent to 3.5% of the 2-gram recommended maximum daily dietary intake of sodium for an adult.

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 unsure about your dose and how to use this medicine. The dosage and method of use will be determined by the doctor only.

Baclofen Sintetica is administered by intrathecal injection. This means that the medicine is injected directly into the cerebrospinal fluid. The dose needed varies from person to person depending on their condition, and the doctor will decide what dose you need after testing your response to the drug.

First, the doctor will check, by giving you single doses of Baclofen Sintetica, whether the medicine is suitable for you. During this period, the heart and lung functions will be closely monitored. If there is an improvement in the symptoms, a special pump which can deliver the drug continuously will be implanted into your chest or abdominal wall. The doctor will give you all the necessary explanation to use the pump and get the right dosage. Make sure that you fully understand all the instructions.

The final dose of Baclofen Sintetica depends on each patient’s response to the drug. You will first receive a low dose, which will be gradually increased over several days, under the doctor’s supervision, until you receive the right dose for you. If the starting dose is too high, or if the dose is increased too quickly, there is a higher risk of side effects.

To prevent unpleasant side effects which may be serious and even life-threatening, it is important that your pump does not run out. The filling of the pump must always be done by a doctor or nurse, and you must make sure not to miss the clinic appointments scheduled for you.

During long-term treatment, some patients feel a decrease in the effectiveness of Baclofen Sintetica. You may require occasional breaks in treatment. The doctor will advise you what to do.

Do not exceed the recommended dose.

If you overdose on Baclofen Sintetica

It is very important that you and the person caring for you know how to recognize the signs of an overdose.
These signs may appear if the pump is not working properly, and this should be reported to the doctor without delay.
Signs of overdose are:

- Unusual muscle weakness (poor muscle tone)
- Sleepiness

- Dizziness or l light-headedness
- Excessive salivation
- Nausea or vomiting
- Difficulty breathing
- Convulsions
- Loss of consciousness
- Abnormally low body temperature

If you stop using Baclofen Sintetica

Do not stop treatment suddenly. If the doctor decides to stop your treatment, the dose will be reduced gradually to prevent withdrawal symptoms such as muscle spasms and increased muscle rigidity, fast heart rate, fever, confusion, hallucinations, changes in mood and emotion, mental disorders, feeling haunted or convulsions (fits), prolonged and painful erection (priapism). In rare cases, these symptoms can be life-threatening. If you or your caregiver notice any of these symptoms, you should immediately contact your doctor in order to check the integrity of the pump and the delivery system.

Do not take medicine in the dark! Check the label and dose <u>every time</u> you take a medicine. Wear glasses if you need them.

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

4. <u>Side effects</u>

As with any medication, the use of Baclofen Sintetica may cause side effects in some users. Do not be alarmed when reading the list of side effects. You may not suffer from any of them. Implanted drug delivery device or infusion system malfunction can lead to withdrawal symptoms including death.

<u>Very common side effects – effects that appear in more than 1 user out of 10</u>
Feeling tired, drowsy or weak

<u>Common side effects – effects that appear in up to 1 user out of 10</u>
- Feeling lethargic (having no energy)
- Headache, dizziness or stupor
- Pain, fever or chills
- Seizures
- Tingling in the palms of the hands or feet
- Problems with eyesight
- Slurred speech
- Insomnia
- Breathing difficulties, pneumonia
- Feeling confused, anxious, agitated or depressed
- Low blood pressure (fainting)
- Nausea or vomiting, constipation or diarrhea
- Lack of appetite, dry mouth or excessive salivation
- Rash and itching, swelling of the face or hands and feet
- Urinary incontinence, or problems when urinating
- Cramps
- Sexual dysfunction in men, such as impotence

<u>Uncommon side effects – effects that appear in up to 1 user out of 100</u>
- Feeling abnormally cold
- Memory loss
- Mood swings and hallucinations, feeling suicidal
- Stomach pain, difficulty swallowing, loss of taste, dehydration

- Loss of muscle control
- High blood pressure
- Slow heart rate
- Deep vein thrombosis
- Flushed or pale skin, excessive sweating
- Hair loss

Side effects of unknown frequency (effects whose frequency has not yet been determined)

- Restlessness
- Abnormally slow breathing rate
- Increased Increase in sideways curvature of the spine (scoliosis)
- Inability to achieve or maintain an erection (erectile dysfunction)

There have been rare reports of problems associated with the pump and delivery system, for example infections, inflammation of the lining around the brain and spinal cord (meningitis) or the accumulation of immune cells at the tip of the delivery tube.

If any of these symptoms become troublesome, if the side effect worsens, or if you experience a side effect not mentioned in the leaflet, you should consult the doctor. He may want to adjust the dose or give you a different medicine.

Reporting of side effects
You can report side effects to the Ministry of Health (MoH) by following the link "Report side effects due to drug treatment" on the MoH website (www.health.gov.il) which links to an online form for reporting side effects, or by entering the link: https://sideeffects.health.gov.il

5. How should the medicine be stored?

- Prevent poisoning! This medicine and any other medicine should be kept in a closed 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) that appears on the outer packaging and ampoule. The expiry date refers to the last day of that month.

Storage conditions

- Store at a temperature below 25°C. Do not store in a refrigerator or freezer.
- Store in the original package to protect from light.
- After first opening, the product must be used immediately. Any remaining product after filling the pump must be destroyed.

6. Further information

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

Sodium chloride, water for injection.

What does the medicine look like and what are the contents of the package:

Clear and colorless solution in ampoules. The ampoule is made of clear colorless glass.

Baclofen Sintetica 0.5 mg/ml:

Each package contains 1 or 5 ampoules.

Baclofen Sintetica 2 mg/ml:

5 ml ampoules: Each package contains 10 ampoules.

20 ml ampoules: Each package contains one ampoule.

Not all package sizes may be marketed.

Importer and license holder name and address: CTS Ltd., 4 Haharash St., Hod Hasharon, 4524075.

Manufacturer name and address:

Sintetica SA, Via Penate 5, CH-6850 Mendrisio, Switzerland

This leaflet was revised in 02/2023 according to Ministry of Health guidelines.

Registration numbers of the medicine in the National Drug Registry of the Ministry of Health:
Baclofen Sintetica 0.5 mg/ml: 170-36-37175-99
Baclofen Sintetica 2 mg/ml: 156-85-34288-00