

## **Bijsluiter: informatie voor de gebruiker**

### **Sapropterine DZ 100 mg oplosbare tabletten**

Sapropterin dihydrochloride

**Read all of this leaflet carefully before you start taking this medicine because it contains important information for you.**

- Keep this leaflet. You may need to read it again.
- If you have any further questions, ask your doctor or pharmacist.
- This medicine has been prescribed for you only. Do not pass it on to others. It may harm them, even if their signs of illness are the same as yours.
- If you get any side effects talk to your doctor or pharmacist. This includes any possible side effects not listed in this leaflet. See section 4.

#### **What is in this leaflet**

1. What [Product name] is and what it is used for
2. What do you need to know before you take [Product name]
3. How to take [Product name]
4. Possible side effects
5. How to store [Product name]
6. Contents of the pack and other information

#### **1. What [Product name] is and what it is used for**

[Product name] contains the active substance sapropterin which is a synthetic copy of a body's own substance called tetrahydrobiopterin (BH4). BH4 is required by the body to use an amino acid called phenylalanine in order to build another amino acid called tyrosine.

[Product name] is used to treat hyperphenylalaninaemia (HPA) or phenylketonuria (PKU) in patients of all ages. HPA and PKU are due to abnormally high levels of phenylalanine in the blood which can be harmful. [Product name] reduces these levels in some patients who respond to BH4 and can help increase the amount of phenylalanine that can be included in the diet.

This medicine is also used to treat an inherited disease called BH4 deficiency in patients of all ages, in which the body cannot produce enough BH4. Because of very low BH4 levels phenylalanine is not used properly and its levels rise, resulting in harmful effects. By replacing the BH4 that the body cannot produce, [Product name] reduces the harmful excess of phenylalanine in the blood and increases the dietary tolerance to phenylalanine.

#### **2. What you need to know before you take [Product name]**

##### **Do not take [Product name]**

If you are allergic to sapropterin or any of the other ingredients of this medicine (listed in section 6).

##### **Warnings and precautions**

Talk to your doctor or pharmacist before taking [Product name], particularly:

- if you are 65 years of age or older
- if you have problems with your kidney or liver
- if you are ill. Consultation with a physician is recommended during illness as blood phenylalanine levels may increase
- if you have predisposition to convulsions

When you are treated with [Product name], your doctor will test your blood to verify how much phenylalanine and tyrosine it contains and may decide to adjust the dose of [Product name] or your diet if needed.

You must continue your diet treatment as recommended by your doctor. Do not change your diet without contacting your doctor. Even if you take [Product name], if your phenylalanine blood levels are not well controlled, you can develop severe neurologic problems. Your doctor should continue to monitor your blood phenylalanine levels often during your treatment with [Product name], **to make sure that your blood phenylalanine levels are not too high or too low.**

### **Other medicines and [Product name]**

Tell your doctor or pharmacist if you are taking, have recently taken or might take any other medicines. In particular you should tell your doctor if you are using:

- levodopa (used to treat Parkinson's disease)
- medicines for treatment of cancer (e.g. methotrexate)
- medicines for treatment of bacterial infections (e.g. trimethoprim)
- medicines that cause dilation of blood vessels, (such as glyceryl trinitrate (GTN), isosorbide dinitrate (ISDN), sodium nitroprusside (SNP), molsidomin, minoxidil).

### **Pregnancy and breast feeding**

If you are pregnant or breast-feeding, think you may be pregnant or are planning to have a baby, ask your doctor or pharmacist for advice before taking this medicine.

If you are pregnant your doctor will tell you how to control phenylalanine levels adequately. If these are not strictly controlled before or when you become pregnant, this could be harmful to you and to your baby. Your doctor will monitor the restriction of dietary phenylalanine intake prior and during pregnancy.

If the strict diet does not adequately reduce phenylalanine amount in your blood your doctor will consider whether you must take this medicine.

You should not take this medicine if you are breast-feeding.

### **Driving and using machines**

[Product name] is not expected to affect the ability to drive and use machines.

### **[Product name] contains sodium**

This medicine contains less than 1 mmol sodium (23 mg) per tablet, that is to say essentially 'sodium-free'.

## **3. How to take [Product name]**

Always take this medicine exactly as your doctor has told you. Check with your doctor if you are not sure.

### **Dosing for PKU**

The recommended starting dose of [Product name] in patients with PKU is 10 mg for each kg of body weight. Take [Product name] as a single daily dose with a meal to increase the absorption, and at the same time each day, preferably in the morning. Your doctor may adjust your dose, usually between 5 and 20 mg for each kg of body weight per day, depending on your condition.

### **Dosing for BH4 deficiency**

The recommended starting dose of [Product name] in patients with BH4 deficiency is 2 to 5 mg for each kg of body weight. Take [Product name] with a meal to increase the absorption. Divide the total daily dose into 2 or 3 doses, taken over the day. Your doctor may adjust your dose up to 20 mg for each kg of body weight per day, depending on your condition.

### **The table below is an example of how an appropriate dose is calculated**

Body weight (kg)	Number of 100 mg tablets	Number of 100 mg tablets
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	(dose 10 mg/kg)	(dose 20 mg/kg)
10	1	2
20	2	4
30	3	6
40	4	8
50	5	10

### **Method of administration**

For PKU patients, the total daily dose is taken once a day at the same time each day, preferably in the morning.

For BH4 deficiency patients, the total daily dose is divided into 2 or 3 doses over the day.

### Use in all patients

Place the prescribed number of tablets in a glass or cup of water as accurately described below and stir until dissolved.

It may take a few minutes for the tablets to dissolve. To make the tablets dissolve faster you can crush them. Small particles may be visible in the solution, but they will not affect the effectiveness of the medicine. Drink the dissolved preparation of [Product name] with a meal within 15 to 20 minutes of its preparation.

Do not swallow the desiccant capsule contained in the bottle.

### *Use in patients above 20 kg body weight*

Place the tablets in a glass or cup (120 to 240 ml) of water and stir until dissolved.

### *Use in children up to 20 kg body weight*

The dose is based on body weight. This will change as your child grows. Your doctor will tell you:

- the number of [Product name] tablets needed for one dose
- the amount of water needed to mix one dose of [Product name]
- the amount of solution you will need to give your child for their prescribed dose

Your child should drink the solution with a meal.

Give your child the prescribed amount of solution within 15 to 20 minutes after dissolving. If you are not able to give your child's dose within 15 to 20 minutes after dissolving the tablets, you will need to prepare a new solution as the unused solution should not be used beyond 20 minutes.

### *Supplies needed to prepare and give your child's dose of [Product name]*

- The number of [Product name] tablets needed for one dose
- A medicine cup with graduation markings at 20, 40, 60 and 80 ml
- A glass or cup
- Small spoon or clean utensil for stirring
- Oral syringe (graduated in 1 ml divisions) (10 ml syringe for administration of volumes of  $\leq 10$  ml or 20 ml syringe for administration of volumes of  $>10$  ml)

Ask your doctor for the medicine cup for dissolving the tablets and the 10 ml or 20 ml oral syringe if you do not have these supplies.

### *Steps for preparing and taking your dose:*

- Place the prescribed number of tablets in the medicine cup. Pour the amount of water into the medicine cup, as instructed by your doctor (e.g. your doctor told you to use 20 ml for dissolving one [Product name] tablet). Check to make sure that the amount of liquid lines up with the amount that your doctor tells you. Stir with the small spoon or clean utensil until the tablets dissolve.

- If your doctor told you to administer only a portion of the solution, point the tip of the oral syringe into the medicine cup. Slowly pull back the plunger to withdraw the amount as instructed by your doctor.
- Transfer the solution by pushing on the plunger slowly until all of the solution in the oral syringe is transferred to a glass or cup for administration (e.g. if your doctor told you to dissolve two [Product name] tablets in 40 ml water and administer 30 ml to your child, you would have to use the 20 ml oral syringe two times to draw up 30 ml (e.g. 20 ml + 10 ml) of the solution and transfer it to a glass or cup for administration). Use a 10 ml oral syringe for administration of volumes  $\leq 10$  ml or a 20 ml oral syringe for administration of volumes  $> 10$  ml.
- If your baby is too small to drink from a glass or a cup you may administer the solution via the oral syringe. Draw up the prescribed volume from the solution prepared in the medicine cup and place the tip of the oral syringe into your baby's mouth. Point the tip of the oral syringe towards either cheek. Push on the plunger slowly, a small amount at a time, until all of the solution in the oral syringe is given.
- Throw away any remaining solution. Remove the plunger from the barrel of the oral syringe. Wash both parts of the oral syringe and the medicine cup with warm water and air dry. When the oral syringe is dry, put the plunger back into the barrel. Store the oral syringe and the medicine cup for next use.

### **If you take more [Product name] than you should**

If you take more [Product name] than prescribed, you may experience side effects that could include headache and dizziness. Immediately contact your doctor or pharmacist if you take more [Product name] than prescribed.

### **If you forget to take [Product name]**

Do not take a double dose to make up for a forgotten dose. Take the next dose at the usual time.

### **If you stop taking [Product name]**

Do not stop taking [Product name] without prior discussion with your doctor, as phenylalanine levels in your blood may increase.

If you have any further questions on the use of this medicine, ask your doctor or pharmacist.

## **4. Possible side effects**

Like all medicines, this medicine can cause side effects, although not everybody gets them.

Few cases of allergic reactions (such as skin rash and serious reactions) have been reported. Their frequency is not known (frequency cannot be estimated from the available data).

If you have red, itchy, raised areas (hives), runny nose, fast or uneven pulse, swelling of your tongue and throat, sneezing, wheezing, serious difficulty in breathing or dizziness, you may be having a serious allergic reaction to the medicine. If you notice these signs, contact your doctor immediately.

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

Headache and runny nose.

### Common side effects (may affect up to 1 in 10 people)

Sore throat, nasal congestion or stuffy nose, cough, diarrhoea, vomiting, stomach ache, too low levels of phenylalanine in blood tests, indigestion and feeling sick (nausea) (see section 2: "Warnings and precautions").

### Not known side effects (frequency cannot be estimated from the available data)

Gastritis (inflammation of the lining of the stomach), oesophagitis (inflammation of the lining of the gullet).

## **Reporting of side effects**

If you get any side effects, talk to your doctor or pharmacist or nurse. This includes any possible side effects not listed in this leaflet. You can also report side effects directly via the national reporting system listed in Appendix V. By reporting side effects you can help provide more information on the safety of this medicine.

## 5. How to store [Product name]

Keep this medicine out of the sight and reach of children.

Do not use this medicine after the expiry date which is stated on the label and the carton after “EXP”. The expiry date refers to the last day of that month.

Store below 25°C in the original package to protect from moisture and light.

Do not throw away any medicines via wastewater or household waste. Ask your pharmacist how to throw away medicines you no longer use. These measures will help protect the environment.

## 6. Contents of the pack and other information

### What [Product name] contains

- The active substance is sopropterin dihydrochloride. Each tablet contains 100 mg of sopropterin dihydrochloride (equivalent to 77 mg of sopropterin).
- The other ingredients are mannitol (E421), pregelatinised starch, crospovidone (E1202), riboflavin (E101), ascorbic acid (E300), sodium stearyl fumarate.

### What [Product name] looks like and contents of the pack

[Product name] 100 mg soluble tablets are off white to light yellow coloured, round tablets of diameter about 9.5 mm engraved with “L71” on one side and “T” on the opposite side

It is available in high-density polyethylene (HDPE) bottles with child-resistant closure. Each bottle contains 30, 120 or 240 tablets. The bottles are sealed with an aluminium seal. Each bottle contains a small plastic tube of desiccant (silica gel).

Not all pack sizes may be marketed.

### Houder van de vergunning voor het in de handel brengen en fabrikant

*Houder van de vergunning voor het in de handel brengen*

Day Zero ehf.

Reykjavíkurvegi 62

Hafnarfjörður

220

IJsland

*Fabrikant*

Teva Operations Poland Sp. Z.o.o

ul. Mogilska 80

31-546 Krakow

Polen

### In het register ingeschreven onder:

RVG 127854

**Dit medicijn is geregistreerd in lidstaten van de Europese Economische Ruimte onder de volgende namen:**

Nederland Saproterine DZ 100 mg oplosbare tabletten

Duitsland Saproterin HCl DZ 100 mg Tabletten zur Herstellung einer Lösung zum Einnehmen

**Deze bijsluiter is voor het laatst goedgekeurd in maart 2023.**