

PATIENT INFORMATION LEAFLET

PHOCUSS[®] 10 mg/20 mg film coated tablet

Use orally.

- **Drug Substance:** Each film-coated tablet contains 10 mg donepezil hydrochloride, equivalent to 9.12 mg donepezil, and 20 mg memantine hydrochloride, equivalent to 16,620 mg memantine.
- **Excipients:** Contains bovine lactose monohydrate, microcrystalline cellulose, colloidal silicon dioxide, crospovidone, talc, sodium stearyl fumarate, polyvinyl alcohol, titanium dioxide, polyethylene glycol, red iron oxide, black iron oxide.

• **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. Do not pass it on to others.
- Please tell your doctor that you are taking this medicine if you seek medical assistance during the use of this medicine.
- Follow these instructions carefully. Do not take higher or lower doses than recommended.

In this leaflet:

1. ***What is PHOCUSS[®] and what is it used for?***
2. ***Before using PHOCUSS[®] what should be considered?***
3. ***How to use PHOCUSS[®]?***
4. ***Possible side effects?***
5. ***How to store PHOCUSS[®]?***

Titles are included.

1. What is PHOCUSS[®] and what is used for?

- PHOCUSS[®] are pink, scored, oblong film-coated tablets containing 10 mg donepezil hydrochloride and 20 mg memantine hydrochloride.
- PHOCUSS[®] is offered to the market in transparent PVC/PE/PVDC/Al foil blister packages containing 28 and 84 film-coated tablets.
- PHOCUSS[®] is a combination of donepezil, a piperidine-type reversible acetylcholinesterase inhibitor, and memantine, which belongs to a group of drugs called NMDA receptor antagonists.
- Alzheimer's disease (memory loss due to impaired message signals to the brain) symptoms include increased memory loss, confusion and behavioral changes. As a result, it has been

observed that Alzheimer's patients have difficulty in continuing their normal daily activities. PHOCUSS® is a medication used to treat memory disorders in patients with moderate to severe Alzheimer's disease.

- Only adult patients can use PHOCUSS®.
- PHOCUSS® contains bovine lactose monohydrate.

Before using PHOCUSS® what should be considered? In this following

Do not use PHOCUSS® in the following situations

If,

- **If you are allergic to memantine hydrochloride, donepezil hydrochloride, piperidine derivative drugs or any of the ingredients of PHOCUSS®.**

USE PHOCUSS® WITH CARE in the following situations.

If,

- If you have a stomach or duodenal ulcer,
- If there is a seizure or a severe spasm or sudden contractions in the face, trunk or arms and legs,
- If you have heart disease (irregular or slow heartbeat),
- If you have asthma or another long-term lung disease,
- If you have ever had liver disease or jaundice.
- If you have a history of epileptic seizures,
- If you have recently had a myocardial infarction (heart attack) or if you suffer from congestive heart failure or uncontrolled hypertension (high blood pressure),
- If you have difficulty urinating or have kidney disease,

In these cases, treatment should be carefully controlled and the clinical benefit of PHOCUSS® re-evaluated by your doctor at regular intervals.

If you have kidney failure, your doctor should monitor your kidney function closely and, if necessary, adjust the dose of PHOCUSS® accordingly.

Concomitant use with amantadine (used in the treatment of Parkinson's disease), ketamine (used for anesthesia), dextromethorphan (usually used in the treatment of cough) and other NMDA-antagonists should be avoided.

PHOCUSS® is not recommended for children and adolescents under 18 years of age.

If these warnings are necessary for you, even at any time in the past, please consult your doctor.

Using PHOCUSS® with food and drink

PHOCUSS® can be used on an empty or full stomach.

PHOCUSS® should not be taken with alcohol; because alcohol can change the effect of the medicine.

If you have recently changed your diet too much or are planning to change it (e.g. switching from a normal diet to a strict vegetarian diet) or if you have renal tubular acidosis (RTA,

excess of acid-forming substances in the blood due to kidney dysfunction) or severe urinary tract If you have an infection, you should inform your doctor. The dose of your medicine may need to be adjusted.

Pregnancy

Consult your doctor or pharmacist before using the medicine.

If you are pregnant or planning to become pregnant, inform your doctor. The use of PHOCUSS® is not recommended in pregnant women.

If you notice that you are pregnant during your treatment, consult your doctor or pharmacist immediately.

Braast Feeding

Consult your doctor or pharmacist before using the medicine.

If you are breastfeeding your baby, inform your doctor. Breastfeeding women should not use PHOCUSS®.

Driving and using machines

Alzheimer's disease can impair your ability to drive and use machines. Additionally, PHOCUSS® may cause weakness, dizziness and muscle cramps. If any of these effects occur, you should not drive or use machines. PHOCUSS® may change your reaction ability, making it impractical for you to drive or use machines. Therefore, do such activities only if your doctor allows.

Important information about some excipients contained in PHOCUSS®

PHOCUSS® film kaplı tabletler sığır kaynaklı laktoz monohidrat içermektedir. Eğer daha önceden doktorunuz tarafından bazı şekerlere intoleransınız olduğu söylenmişse bu tıbbi ürünü almadan önce doktorunuzla temasa geçiniz.

Bu tıbbi ürün her film kaplı tablette 1 mmol (23 mg)'dan daha az sodyum ihtiva eder; yani esasında “sodyum içermez”.

Use with other drugs

- If you are using or have just started using any of the following medications, inform your doctor or pharmacist. These may be drugs that you buy from the pharmacy but your doctor has not prescribed for you. The use of these medications may make the effects of PHOCUSS® stronger or weaker in the future, and their doses may need to be adjusted by your doctor.
- • Other acetylcholinesterase inhibitor drugs used in the treatment of Alzheimer's disease (e.g. galantamine),
- • Painkillers or drugs used to treat arthritis (e.g. aspirin, non-steroidal anti-inflammatory drugs, such as ibuprofen or diclofenac sodium),
- Antikolinerjik ilaçlar (genel olarak hareket bozukluklarını veya barsak kramplarını

medicines used to treat it (e.g. tolterodine),

- • Antibiotics (e.g. erythromycin, rifampicin),
- • Drugs used in the treatment of fungi (e.g. ketoconazole),
- • Antidepressants (e.g. fluoxetine),
- • Anticonvulsants (substances used to prevent and relieve seizures, e.g. phenytoin, carbamazepine),
- • Medicines used to treat heart disease (e.g. quinidine, procainamide, beta-blockers [propranolol and atenolol]),
- • Hydrochlorothiazide (or any combination with hydrochlorothiazide, diuretic),
- • Muscle relaxants (e.g. dantrolene, baclofen, diazepam, succinylcholine),
- • Oral anticoagulants (blood thinners; e.g. warfarin),
- • General anesthetics (e.g. ketamine),
- • Medicines available without a prescription, such as herbal treatments (St. John's wort),
- • Barbiturates (substances generally used for sleep),
- • Dopamine agonists (substances used in the treatment of Parkinson's such as L-dopa, bromocriptine),
- • Amantadine (drug used in the treatment of Parkinson's and viral diseases),
- • Dextromethorphan (medicine used in the treatment of cold, cough and flu),
- • Cimetidine, ranitidine (drugs used in ulcer treatment),
- • Quinine (a drug used to treat malaria),
- • Nicotine (contained in smoking cessation preparations),
- • Neuroleptics (drugs used to treat mental disorders).

If you are going to have a surgery that requires general anesthesia, inform your doctor or anesthesiologist that you are using PHOCUSS®. Because PHOCUSS® may affect the amount of anesthesia required.

If you are currently using or have recently used any prescription or non-prescription medication, please inform your doctor or pharmacist about them.

2. How to use PHOCUSS®?

Instructions for proper use and dosage/frequency of application:

Follow your doctor's instructions about when and how to use your medicine and always read the instructions for use. If you are not sure, check with your doctor or pharmacist.

PHOCUSS® is used for the maintenance treatment of Alzheimer's patients treated with 10 mg donepezil per day, when additional memantine is needed, and in patients suitable for combination therapy of 10 mg donepezil + 20 mg memantine per day.

The dose of memantine should be increased gradually according to the titration scheme below.

Dose:

Week 1 (days 1-7):

One 10 mg/5 mg film-coated tablet should be taken once daily for 7 days.

Week 2 (days 8-14):

One 10 mg/10 mg film-coated tablet should be taken once daily for 7 days.

Week 3 (days 15-21):

One 10 mg/15 mg film-coated tablet should be taken once daily for 7 days.

Week 4 (days 21-28):

One 10 mg/20 mg film-coated tablet should be taken once daily for 7 days.

Another pharmaceutical dosage form, the product named PHOCUSS® treatment starter package, easily allows dose adjustment of memantine in patients suitable for donepezil + memantine combination therapy.

Maintenance:

The patient should take one PHOCUSS® 10 mg/20 mg film-coated tablet once daily.

Do not change the dose of your medicine without your doctor's advice.

Always follow your doctor's and pharmacist's instructions about your medicine and how much you should take.

Application route and method:

- PHOCUSS® is for oral use.
- One film-coated tablet should be taken every day.
- Unless directed otherwise by your doctor, take one film-coated tablet of PHOCUSS® every night before going to bed.

Different age groups:

Use in children:

PHOCUSS® is not recommended for use in children under 18 years of age.

Use in the elderly:

PHOCUSS® can be used in elderly patients at the doses stated above.

Special use cases:

Kidney failure:

If you have kidney failure, your doctor will decide on a dose appropriate for your condition. In this case, your kidney functions should be monitored by your doctor at regular intervals.

Liver failure:

If you have liver failure, your doctor will determine the dose appropriate for your condition. The use of PHOCUSS® is not recommended in patients with severe hepatic impairment.

If you have the impression that the effect of PHOCUS is too strong or weak, talk to your doctor or pharmacist.

If you use more PHOCUSS® than you should

If you have used more PHOCUSS® than you should, talk to a doctor or pharmacist.

Do not take more than one film-coated tablet each day. If you take more than the recommended dose, call your doctor immediately. If you cannot reach your doctor, contact a hospital emergency department immediately. When you go, take the medicine box containing the tablets with you. Symptoms of overdose may include nausea, increased salivation, sweating, slow heart rate, low blood pressure (feeling faint or dizzy when standing), difficulty breathing, loss of consciousness, seizure or convulsions.

If you forget to use PHOCUSS®

Do not take a double dose to make up for forgotten doses.

If you forget to use PHOCUSS®, take your next dose on time. If you forget to take PHOCUSS® for more than a week, call your doctor before taking the medicine.

Effects that may occur when treatment with PHOCUSS® is stopped

Do not stop taking PHOCUSS® unless your doctor tells you to. If you stop taking PHOCUSS®, there will be a gradual decrease in the beneficial effects of your treatment.

4.What are the possible side effects?

Like all medicines, side effects may occur in people who are sensitive to the ingredients contained in PHOCUSS®.

If any of these symptoms occur, you may need urgent medical intervention or hospitalization.

Side effects are categorized as follows:

Very common	: Can occur in at least 1 out of 10 patients.
Common	: Can occur in less than 1 out of 10, but more than 1 out of 100 patients.
Uncommon	: Can occur in less than 1 out of 100, but more than 1 out of 1000 patients.
Rare	: Can occur in less than 1 out of 1000, but more than 1 out of 10000 patients.
Very rare	: Can occur in less than 1 out of 10000 patients.
Not known	: Frequency cannot be estimated from the available data.

- If you notice any of the following, tell your doctor immediately or go to the emergency department of your nearest hospital

Uncommon:

- Stomach or duodenal ulcer. Symptoms of an ulcer include stomach pain and discomfort between the navel and the breastbone.

- Stomach or intestinal bleeding. In this case, you may experience effects such as black tar-like stools or blood in the rectum,
- Seizure or contraction (convulsion),
- **Rare:**
 - Liver damage such as hepatitis. Symptoms of hepatitis include feeling sick or vomiting, loss of appetite, feeling generally unwell, fever, itching, yellowing of the eyes and skin, dark urine (rarely observed),
- **Very rare:**
 - Fever accompanied by muscle stiffness, sweating or decreased level of consciousness (a disease called neuroleptic malignant syndrome).
 - Muscle weakness, tenderness or pain (especially if you feel unwell, your body temperature is high or your urine is dark). These can be caused by abnormal muscle disorders (rhabdomyolysis), which can be life-threatening and lead to kidney problems.

Other side effects are given below according to frequency:

- **Very common:**
 - Diarrhea
 - Nausea
 - Headache
- **Widespread:**
 - Muscle cramps
 - Fatigue
 - Insomnia (difficulty sleeping)
 - Cold
 - Loss of appetite
 - Hallucination (delusion, delusion)
 - Abnormal dreams (including nightmares)
 - Agitation (restlessness)
 - Aggressive behavior
 - Fainting
 - Dizziness
 - Stomach upset
 - Rash
 - Itching
 - Urinary incontinence
 - Pain

- **Accidents (patients may become more prone to falls and accidental injuries may occur)**
- Sleepiness
- Constipation
- Headache
- Increase in liver function tests
- Dizziness
- Balance Disorders
- Shortness of breath
- High Blood pressure
- Hypersensitivity

Uncommon:

- Slowing of heart rate
- Increase in saliva secretion
- Fungal infections
- Confusion (mental confusion)
- Vomiting
- Gait abnormality
- Heart failure
- • Venous blood clot (thrombosis/thromboembolism)

Rare:

- • Rigidity, tremors or uncontrollable movements, especially of the face, tongue and also limbs.

Unknown:

- Inflammation of the pancreas,
- Psychotic reactions

Alzheimer's disease has been associated with depression, suicidal ideation, and suicide. These events have been reported in patients treated with memantine, the active ingredient in PHOCUS®.

If you experience any side effects not mentioned in this instruction manual, inform your doctor or pharmacist.

Reporting of side effects

As a result of standard experiments with memantine, no genotoxicity was observed. There was no evidence of carcinogenicity in lifetime studies in mice and rats.

Memantine was not teratogenic in rats and rabbits, even at maternally toxic doses, and no adverse effects on productivity were detected.

Fetal growth was observed to be reduced in rats at doses equivalent to or slightly higher than those administered to humans.

5 PHARMACEUTICAL PROPERTIES

6.1 List of excipients

5.Storage of PHOCUSS®

Keep PHOCUSS® out of sight and reach of children and in its packaging.

Store PHOCUSS® at room temperature below 25°C..

Use in accordance with the expiration date.

Do not use ELEPSİ® after the expiration date printed on the packaging

Do not use ELEPSİ® if you notice any defects in the product or packaging.

Do not dispose of expired or unused medication in the trash! Dispose of them according to the collection system established by the Ministry of Environment and Urbanization.

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This leaflet has been approved <{DD month YYYY}>. [To be completed nationally]