

SECTION 1: PATIENT INFORMATION LEAFLET (PIL)

Imipenem+Cilastatin

500 mg/500 mg powder for solution for infusion

- **Read all of this leaflet carefully before you start using 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, pharmacist or nurse.**
- **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, pharmacist or nurse. This includes any possible side effects not listed in this leaflet.**

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1. SERIOUS SIDE EFFECTS

Get emergency medical help if you have any of these signs of an allergic reaction:

- hives
- difficulty breathing, closing of the throat
- swelling of your face, lips, tongue, or throat.

Call your doctor at once if you have a serious side effect such as:

- diarrhea that is watery or bloody
- unusual bleeding or bruising
- white patches or sores inside your mouth or on your lips

- skin rash
- seizure (convulsions)
- being breathless when you do not expect it
- red or brown urine
- unusual tiredness or weakness; or
- severe skin reaction - fever, sore throat, swelling in your face or tongue, burning in your eyes, skin pain, followed by a red or purple skin rash that spreads (especially in the face or upper body) and causes blistering and peeling.

2. WHAT IMIPENEM+CILASTATIN IS AND WHAT IS USED FOR

Imipenem+Cilastatin 500 mg/500 mg powder for solution for infusion (hereinafter referred to as Imipenem+Cilastatin) belongs to a group of medicines called carbapenem antibiotics. It works by killing bacteria, which can cause serious infections:

- Infection affecting the lungs (pneumonia)
- Complicated infections in the abdomen
- Severe genital tract infections
- Infections that you can catch during or after the delivery of your baby
- Complicated urinary tract infections
- Complicated skin and soft tissue infections

Imipenem+Cilastatin may be used in the management of patients with low white blood cell counts (called “neutropenia”), who have fever that is suspected to be due to a bacterial infection.

Imipenem+Cilastatin may be used to treat bacterial infection of the blood (called “sepsis” or “septicemia”) which might be associated with a type of infection mentioned above.

3. WHAT YOU NEED TO KNOW BEFORE YOU TAKE IMIPENEM+CILASTATIN

a. You must not be given Imipenem+Cilastatin if:

- you are allergic (hypersensitive) to imipenem, cilastatin or any of the other ingredients of this medicine (listed in section 6)
- you are allergic (hypersensitive) to other antibiotics such as penicillins, cephalosporins, or carbapenems.

b. Tell your doctor about any medical condition you have or have had including:

- allergies to any medicines including antibiotics (sudden life-threatening allergic reactions require immediate medical treatment)
- inflammatory bowel disease
- kidney or urinary problems, including reduced kidney function (Imipenem+Cilastatin blood levels increase in patients with reduced kidney function. Central nervous system adverse reactions may occur if the dose is not adjusted to the kidney function)
- any central nervous system disorders such as localized tremors or epileptic seizures (fits)
- liver problems.

You may develop a positive test (Coombs test) which indicates the presence of antibodies that may destroy red blood cells. Your doctor will discuss this with you.

If you are not sure if any of the above applies to you, talk to your doctor or nurse before using Imipenem+Cilastatin.

c. Children

Imipenem+Cilastatin must not be given to babies if:

- the baby is newborn infants and infants below the age of 3 months as the safety of use has not yet been established in this group.

d. Tell your doctor if you are taking or might take any other medicines, especially any of the following:

- ganciclovir which is used to treat some viral infections
- valproic acid or sodium valproate (used to treat epilepsy, bipolar disorder, migraine, or schizophrenia). Imipenem+Cilastatin should not be used because it may decrease the effect of sodium valproate,
- any blood thinners such as warfarin (used to treat or prevent blood clots).

Tell your doctor, pharmacist or nurse if you are taking, have recently taken or might take any other medicines. This is because Imipenem+Cilastatin can affect the way some medicines work and some medicines can have an effect on Imipenem+Cilastatin. Your doctor will decide whether you should use Imipenem+Cilastatin in combination with these medicines.

Talk to your doctor or pharmacist or nurse before your child is administered

e. Pregnancy, breast-feeding and fertility

It is important that you tell your doctor if you are pregnant or are planning to become pregnant before receiving Imipenem+Cilastatin. Imipenem+Cilastatin has not been studied in pregnant women. Imipenem+Cilastatin should not be used during pregnancy unless your doctor decides the potential benefit justifies the potential risk to the developing baby.

It is important that you tell your doctor if you are breast-feeding or if you intend to breast-feed before receiving Imipenem+Cilastatin. Small amounts of this medicine may pass into breast milk and it may affect the baby. Therefore, your doctor will decide whether you should use Imipenem+Cilastatin while breast-feeding.

Ask your doctor or pharmacist for advice before taking any medicine.

f. Driving and using machines

There are some side effects associated with this product (such as seeing, hearing, or feeling something that is not there, dizziness, sleepiness, and a spinning sensation) that may affect some patients' ability to drive or operate machinery.

g. Imipenem+cilastatin contains sodium

This medicinal product contains approximately 1.6 mEq (approximately 37.6 mg) of sodium per 500 mg dose which should be taken into consideration by patients on a controlled sodium diet.

If you have a condition which requires you to monitor your sodium intake please inform your doctor, pharmacist or nurse.

4. HOW TO TAKE IMIPENEM+CILASTATIN

Imipenem+Cilastatin will be prepared and given to you by a doctor or another health care professional. Your doctor will decide the correct dose of Imipenem+Cilastatin for you, or how much Imipenem+Cilastatin you need. The dose will depend on the type of infection that you have; where the infection is in the body and how serious the infection is; whether you are on any other antibiotics; your weight and age; how well your kidneys and liver are working.

Use in adults and adolescents

- The recommended dose for adults and adolescents is 500 mg/500 mg every 6 hours or 1,000 mg/1,000 mg every 6 or 8 hours. If you have kidney problems your doctor may lower your dose.

Use in children

- The recommended dose for children one year of age or older is 15/15 or 25/25 mg/kg/dose every 6 hours. Imipenem+Cilastatin is not recommended in children below the age of 3 months and children with kidney problems as the safety of use has not yet been established in these groups.

How to use Imipenem+cilastatin:

- Your doctor or nurse will normally give Imipenem+Cilastatin to you

- Imipenem+Cilastatin is given intravenously (into a vein) over 20-30 minutes for a dose of ≤ 500 mg/500 mg or 40-60 minutes for a dose of >500 mg/500 mg. The rate of infusion may be slowed if you feel sick
- You should normally have your injections at the same times each day.

a. If you forget to take Imipenem+cilastatin

- If you are concerned that you may have missed a dose, contact your doctor or another healthcare professional immediately
- Do not have a double dose (two injections at the same time) to make up for a forgotten dose

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

b. If you take more Imipenem+cilastatin than you should

- Symptoms of overdose may include seizures (fits), confusion, tremors, nausea, vomiting, low blood pressure and slow heart rate
- If you are concerned that you may have been given too much Imipenem+Cilastatin, contact your doctor or another healthcare professional immediately.

c. If you stop using Imipenem+cilastatin:

Do not stop having Imipenem+cilastatin until your doctor tells you to.

If you still feel unwell at the end of your prescribed course of treatment, tell your doctor.

5. POSSIBLE SIDE EFFECTS

Like all other medicines Imipenem+cilastatin can cause side effects although not everybody gets them. If you have a severe allergic reaction (see section 1 of this leaflet), stop having Imipenem+Cilastatin and see a doctor straight away. You may need urgent medical treatment.

The following side effects may happen with this medicine:

Common (may affect up to 1 in 10 people)

- Nausea, vomiting, diarrhoea. Nausea and vomiting appear to occur more frequently in patients with low number of white blood cells
- Swelling and redness along a vein which is extremely tender when touched
- Rash
- Abnormal liver function detected by blood tests
- Increase in some white blood cells.

Uncommon (may affect up to 1 in 100 people)

- Local skin redness

- Local pain and formation of a firm lump at the injection site
- Skin itchiness
- Hives
- Fever
- Blood disorders affecting the cell components of the blood and usually detected by blood tests (symptoms may be tiredness, paleness of skin, and prolonged bruising after injury)
- Abnormal kidney, liver and blood function detected by blood tests
- Tremors and uncontrolled twitching of muscles
- Seizures (fits)
- Psychic disturbances (such as mood swings and impaired judgment)
- Seeing, hearing or feeling something that is not there (hallucinations)
- Confusion
- Dizziness, sleepiness
- Low blood pressure.

Rare (may affect up to 1 in 1 000 people)

- Fungal infection (candidiasis)
- Staining of the teeth and/or tongue
- Inflammation of the colon with severe diarrhoea
- Disturbances in taste
- Inability of the liver to perform normal function
- Inflammation of the liver
- Inability of the kidney to perform normal function
- Changes in the amount of urine, changes in urine colour
- Disease of the brain, tingling sensation (pins and needles), localised tremor

Very rare (may affect up to 1 in 10 000 people)

- Severe loss of liver function due to inflammation (fulminant hepatitis)
- Inflammation of stomach or intestine (gastro-enteritis)
- Inflammation of intestine with bloody diarrhea (hemorrhagic colitis)
- Red swollen tongue, overgrowth of the normal projections on the tongue giving it a hairy appearance, heartburn, sore throat, increase in the production of saliva
- Stomach pain

- A spinning sensation (vertigo), headache
- Ringing in the ears (tinnitus)
- Pain in several joints, weakness
- Irregular heartbeat, the heart beating forcefully or rapidly
- Chest discomfort, difficulty breathing, abnormally fast and superficial breathing, pain in the upper spine
- Flushing, bluish discoloration of the face and lips, skin texture changes, excessive sweating
- Itching of the vulva in women
- Changes in the amounts of blood cells
- Worsening of a rare disease associated with muscle weakness (aggravation of myasthenia gravis)

Frequency not known (cannot be estimated from available data)

- Abnormal movements
- Agitation
- Hearing loss.

Reporting of side effects

If you get any side effects, talk to your doctor or nurse. This includes any possible side effects not listed in this leaflet.

6. HOW TO STORE IMIPENEM+CILASTATIN AND OTHER INFORMATION

Keep out of reach and sight of children

Do not use Imipenem+Cilastatin after the expiry date which is stated on the vial and carton after EXP. The expiry date refers to the last day of that month.

Store the unopened vial below 30°C. Keep container in the outer carton.

After reconstitution:

Diluted solutions should be used immediately. The time interval between the beginning of reconstitution and the end of intravenous infusion should not exceed four hours.

Do not freeze the reconstituted solution.

If the solution is cloudy, it should not be used.

Any unused solution should be discarded.

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

7. FURTHER INFORMATION

a. What Imipenem+cilastatin contains

- The active substances are imipenem and cilastatin. Each vial contains imipenem monohydrate equivalent to 500 mg imipenem and cilastatin sodium equivalent to 500 mg cilastatin.
- The other ingredient is sodium bicarbonate.

b. What Imipenem+cilastatin looks like

Imipenem+Cilastatin is a white to light yellow powder for solution for infusion in a glass vial. Pack sizes of 1, 10 or 25 vials. Not all pack sizes may be marketed.

It is supplied in colorless glass vials sealed with an aluminum seal or flip-off aluminum seal with 1, 10, 1-50 or 50 vials in a carton.

Not all pack sizes may be marketed.

c. Marketing Authorisation Holder and Manufacturer

“KRASPHARMA” OJSC

Russia, 660042, 60 let Oktyabrya st. 2, Krasnoyarsk

Tel.: +7 (391) 261-25-90. Fax: +7 (391) 261-17-44

For any information about this medicinal product, please contact the local representative of the Marketing Authorisation Holder:

“KRASPHARMA” OJSC

Russia, 660042, 60 let Oktyabrya st. 2, Krasnoyarsk

Tel.: +7 (391) 261-25-90. Fax: +7 (391) 261-17-44

d. This leaflet was last approved _____; version number _____.

e. To report any side effect(s):

- _____:

- _____

Fax: _____

Call NPC at _____, Exts: _____.

Toll free phone: _____

E-mail: _____@_____

Website: _____

f. Council of _____

THIS IS A MEDICAMENT

- **A medicament is a product which affects your health and its consumption contrary to instructions is dangerous for you.**
- **Follow strictly the doctor's prescription, the method of use and the instructions of the pharmacist who sold the medicament.**
- **The doctor and the pharmacist are experts in medicine, its benefits and risks.**
- **Do not by yourself interrupt the period of treatment prescribed.**
- **Do not repeat the same prescription without consulting your doctor.**
- **Keep medicament out of reach of children**

g. This patient information leaflet is approved by the _____.