PATIENT INFORMATION LEAFLET

SCHEDULING STATUS:

S4

CISPLATIN 50 mg/50 ml OETHMAAN, solution for infusion

1,0 mg/ml Cisplatin

Contains approximately 180 mg (7.8 mmol) of sodium per 50 ml.

Contains sugar (Mannitol 50 mg per 50 ml).

Read all of this leaflet carefully before you are given CISPLATIN 50 mg/50 ml OETHMAAN.

- Keep this leaflet. You may need to read it again.
- If you have further questions, please ask your medical practitioner, pharmacist, nurse or other healthcare provider.
- CISPLATIN 50 mg/50 ml OETHMAAN has been prescribed for you personally and you should not share your medicine with other people. It may harm them, even if their symptoms are the same as yours.

What is in this leaflet

1. What CISPLATIN 50 mg/50 ml OETHMAAN is and what it is used for
2. What you need to know before you take CISPLATIN 50 mg/50 ml OETHMAAN
3. How to take CISPLATIN 50 mg/50 ml OETHMAAN
4. Possible side effects
5. How to store CISPLATIN 50 mg/50 ml OETHMAAN
6. Contents of the pack and other information

1. What CISPLATIN 50 mg/50 ml OETHMAAN is and what it is used for

CISPLATIN 50 mg/50 ml OETHMAAN contains a medicine called **cisplatin**, which belongs to a group of medicines called cytostatics used in the treatment of cancer. Cisplatin can be used alone but more commonly cisplatin is used in combination with other cytostatics. Cisplatin can destroy cells in your body that may cause certain types of cancer (tumour of testis, tumour of ovary, tumour of the bladder, head and neck tumours, tumour of the endometrium, lung cancer, cancer of the lymphatic system (lymphomas) and for some childhood cancers).

2. What you need to know before you are given CISPLATIN 50 mg/50 ml OETHMAAN

Do not receive CISPLATIN 50 mg/50 ml OETHMAAN if you:

- are known to be hypersensitive (allergic) to cisplatin or to any of the other ingredients of CISPLATIN 50 mg/50 ml OETHMAAN (listed in **See what CISPLATIN 50 mg/50 ml OETHMAAN contains** below).
- have severe kidney disease.
- have hearing difficulties.
- have very low numbers of blood cells (called ‘bone marrow depression’), (your doctor will check this with a blood test).
- are pregnant or breastfeeding.
- need to have a vaccine for chickenpox, measles, mumps, rubella (MMR) or yellow fever.

Warnings and precautions

CISPLATIN 50 mg/50 ml OETHMAAN should be given to you only under the supervision of a medical practitioner experienced in cancer treatment.

Females are advised to use reliable contraception if they are of child-bearing age, since treatment with CISPLATIN 50 mg/50 ml OETHMAAN may harm their unborn babies.

Tell your doctor or health care provider before you are given CISPLATIN 50 mg/50 ml OETHMAAN if you:

- developed complications from treatment with other anticancer medicines, such as mouth ulcers, low blood cell or platelet counts, or generalised infections, you should first recover from these before you begin treatment with CISPLATIN 50 mg/50 ml OETHMAAN.
- have or have ever had kidney disease.
- have hearing problems.
- are currently taking or have recently taken antibiotics, such as cephalosporins, aminoglycosides and amphotericin B and some substances used in medical imaging, may make the side effects of cisplatin worse; particularly kidney problems.
- are currently taking or have recently taken some water tablets called loop diuretics, antibiotics called aminoglycosides and an anti-cancer medicine called ifosfamide, may make the hearing loss side effect of cisplatin worse.

CISPLATIN 50 mg/50 ml OETHMAAN may also cause the following:

- Symptoms of nerve damage (peripheral neuropathy) such as pins and needles, numbness or poor sense of touch; pain and temporary vision loss (optic neuritis); swelling of the optic nerve; blindness and seizures. Tell your doctor if you experience any of the above as your doctor will stop further treatment with CISPLATIN 50 mg/50 ml OETHMAAN.
- Damage to the kidneys. You will have blood tests before and during treatment to check how well your kidneys are working. You will be given extra fluids through the drip before and after treatment with CISPLATIN 50 mg/50 ml OETHMAAN to protect your kidneys. You may also have a medicine called mannitol. The fluids and mannitol can help prevent

kidney damage. Your doctor or nurse will also give you advice about how much fluid to drink for the next 24 hours and the rest of your treatment cycle.

- Decreased blood cells and bone marrow function including blood cancer (leukaemia). If you develop a fever, bleeding or think you may be suffering from an infection, please inform your doctor immediately.
- Hearing changes. CISPLATIN 50 mg/50 ml OETHMAAN may cause hearing changes, including hearing loss. You may have ringing in the ears. This is called tinnitus. You may also become unable to hear some high-pitched sounds. Hearing changes usually get better after the treatment ends. But some can be permanent. Tell your doctor if you notice any changes in your hearing.
- High levels of uric acid in the blood leading to kidney damage.
- Severe nausea and vomiting can occur 1 to 4 hours after a dose and may persist for up to a week. Your doctor will prescribe medicines to help with the nausea and vomiting.

Other medicines and CISPLATIN 50 mg/50 ml OETHMAAN

Always tell your health care provider if you are taking any other medicine. (This includes all complementary or traditional medicines.)

In particular, tell your doctor or pharmacist if you are taking any of the following medicines:

- Medicines that may make side effects of CISPLATIN 50 mg/50 ml OETHMAAN worse:
    - antibiotics, such as cephalosporins, aminoglycosides and amphotericin B; some water tablets called loop diuretics (e.g. furosemide, ethacrynic acid), hydralazine (for high blood pressure) and propranolol (for high blood pressure) may particularly cause kidney problems.
    - some water tablets called loop diuretics (e.g. furosemide, ethacrynic acid), antibiotics called aminoglycosides and an anti-cancer medicine called ifosfamide

may make the hearing loss side effect of CISPLATIN 50 mg/50 ml OETHMAAN worse.

- use of certain antihistamines, buclizine, cyclizine, meclizine, trimethobenzamide (for nausea and vomiting), loxapine, phenothiazines, thioxanthenes (for mental illness) may hide the symptoms of balance changes (such as dizziness or tinnitus and therefore hearing loss).
- Anti-gout medicines e.g. allopurinol, colchicine, probenecid or sulfinpyrazone can increase the risk of kidney damage.
- Anti-cancer medicine Adriamycin (doxorubicin), radiotherapy or killed live vaccines will increase side effects.

- Effect of CISPLATIN 50 mg/50 ml OETHMAAN on other medicines:
  - bleomycin (anti-cancer medicine), methotrexate (used to treat cancer or arthritis) and paclitaxel (anti-cancer medicine) may produce more side effects if CISPLATIN 50 mg/50 ml OETHMAAN is also being used
  - lithium (for mental illnesses) levels may be reduced
  - the effectiveness of medicines used for the treatment of epilepsy (e.g. phenytoin) may be reduced, so blood levels may need to be checked
  - CISPLATIN 50 mg/50 ml OETHMAAN may make the side effects of the anti-cancer medicine ifosfamide worse (particularly effects on the kidneys)
  - When a blood thinner such as warfarin is taken with CISPLATIN 50 OETHMAAN, the combination may increase the likelihood of bleeding or of an unwanted blood clot. Your doctor will monitor your INR blood levels regularly.

CISPLATIN 50 mg/50 ml OETHMAAN with food and drink and alcohol

Taking CISPLATIN 50 mg/50 ml OETHMAAN with alcohol beverages should be avoided.

Pregnancy and breastfeeding and fertility

If you are pregnant or breastfeeding, think you may be pregnant or are planning to have a baby, please consult your doctor, pharmacist or other healthcare provider for advice before taking this medicine.

CISPLATIN 50 mg/50 ml OETHMAAN is contraindicated during pregnancy and breastfeeding as safety in pregnant women and breastfeeding mothers has not been established.

Pregnancy

There is a risk that use of CISPLATIN 50 mg/50 ml OETHMAAN during pregnancy may cause birth defects in your child. If you are sexually active, you are advised to use effective birth control to prevent pregnancy during treatment, whether you are male or female, both during treatment with CISPLATIN 50 mg/50 ml OETHMAAN and for at least six months after treatment has ended.

Breastfeeding

You should stop breastfeeding before starting treatment with CISPLATIN 50 mg/50 ml OETHMAAN as some of the medicine may get into your breast milk and possibly harm your child.

Fertility

Treatment with CISPLATIN 50 mg/50 ml OETHMAAN can potentially cause permanent sterility in men. It is recommended that those who wish to become fathers in the future discuss the possibility of cryoconservation (freezing) of their sperm prior to treatment. Tell your doctor if you have any concerns.

Driving and using machines

Your ability to drive or use machinery may be affected depending on how well you tolerate the medicine. Do not drive or use machines if you experience any side effect like dizziness and blurred vision which may lessen your ability to do so.

CISPLATIN 50 mg/50 ml OETHMAAN contains:

CISPLATIN 50 mg/50 ml OETHMAAN contains sugar (mannitol) which may have a mild laxative effect.

CISPLATIN 50 mg/50 ml OETHMAAN contains 180 mg of sodium (main component of cooking/table salt) in 50 ml vial. This is equivalent to 9.1% of the recommended maximum daily dietary intake of sodium for an adult.

3. How to receive CISPLATIN 50 mg/50 ml OETHMAAN

You will not be expected to give yourself CISPLATIN 50 mg/50 ml OETHMAAN. It will be given to you by a person who is qualified to do so.

CISPLATIN 50 mg/50 ml OETHMAAN will be given to you by a doctor or nurse through a drip (infusion) into a vein.

The usual dose is:

Your doctor will decide what dose to give and the number of days treatment you will receive depending on your condition and if CISPLATIN 50 mg/50 ml OETHMAAN is to be used in combination with other anticancer medicines. The dose is decided by taking into account the condition you have, your height and weight. From your height and weight, the doctor will work out your body surface area; and it is this that your dose is calculated from.

During treatment your doctor will be making regular checks of your:

- Blood: To check for low blood cell counts that may need treatment.
- Kidney function: Kidney damage can occur when high doses of cisplatin are given. This may not be detected for several weeks; so regular tests may be required during this period.
- Hearing: Your doctor will check that this medicine is not affecting the way it functions in a harmful way.
- Blood uric acid levels: Cisplatin may increase uric acid levels in the blood which might cause gout.

People with kidney problems and elderly

You may be given a different dose to the usual dose. Your doctor will decide how much CISPLATIN 50 mg/50 ml OETHMAAN you will need.

If you are given more CISPLATIN 50 mg/50 ml OETHMAAN than you should

Since a healthcare provider will administer CISPLATIN 50 mg/50 ml OETHMAAN, he/she will control the dosage. However, in the event of overdosage your doctor will manage the overdosage. Signs of overdosage may include kidney failure, liver failure, deafness, eye problems including detachment of the retina, damage to the bone marrow (resulting in low blood cells and bleeding disorder), severe nausea and vomiting and nerve damage.

If you forget to take CISPLATIN 50 mg/50 ml OETHMAAN

Since a healthcare provider will administer CISPLATIN 50 mg/50 ml OETHMAAN, it is unlikely that the dose will be missed.

If you stop receiving CISPLATIN 50 mg/50 ml OETHMAAN

While one course of treatment may sometimes be enough, more often your doctor will advise further courses in either one, three or four weeks time. It may take several courses before your illness is under control and you feel better.

4. Possible side effects:

CISPLATIN 50 mg/50 ml OETHMAAN can have side effects.

Not all side effects reported for CISPLATIN 50 mg/50 ml OETHMAAN are included in this leaflet. Should your general health worsen or if you experience any untoward effects while taking CISPLATIN 50 mg/50 ml OETHMAAN, please consult your doctor, pharmacist or other healthcare professional for advice.

If any of the following happens, stop taking CISPLATIN 50 mg/50 ml OETHMAAN and tell your doctor immediately:

- Sudden signs of allergy – including skin rash, itching or hives; swelling of the face, lips or tongue, shortness of breath or difficulty breathing.

These are all very serious side effects. If you have them, you may have had a serious reaction to CISPLATIN 50 mg/50 ml OETHMAAN. You may need urgent medical attention.

Tell your doctor immediately if you notice any of the following:

- Blood poisoning - symptoms include fever, difficulty breathing, low blood pressure, fast heart rate and mental confusion.
- Acute leukemia - a type of blood cancer that starts in white blood cells in your bone marrow. Symptoms may include enlarged lymph nodes, bruising, fever, bone pain, bleeding from the gums and frequent infections.
- Low white blood cell counts. Symptoms may include fever or chills, cough or hoarseness, lower back or flank pain, painful or difficult urination or a sore throat.

- High blood uric acid levels causing attacks of gout, which usually presents with severe pain, swelling and redness of a joint, such as those of the big toe or knee. This occurs most commonly during initial treatment of patients as a result of rapid cell breakdown that leads to elevated blood uric acid concentrations.
- Stroke - symptoms include dizziness, numbness, weakness on one side of the body, and problems with talking, writing, or understanding language
- Brain dysfunction (confusion, slurred speech, sometimes blindness, memory loss and paralysis)
- Problems with your eyesight (loss of vision or eye pain)
- Ringing in the ears or deafness
- Fast, irregular or slow heart beats, chest pain which could indicate heart problems, heart attack, heart arrest.
- Heart failure – Symptoms may include swelling of the feet or legs, or shortness of breath accompanied by an inability to lie down or coughing of pink-coloured sputum.
- Inflammation of veins, blockage of a blood vessel by a clot (thromboembolism).
- Problems with your kidneys or urine

These are all serious side effects. You may need urgent medical attention.

Tell your doctor if you notice any of the following:

Frequent side effects

- Low red blood cell counts (presenting with tiredness or shortness of breath) and low platelet counts (presenting with easy bruising, pinpoint red spots on the skin, or bleeding from the gums).
- Reduced magnesium levels in the blood
- Tingling, numbness or tremor in your hands, feet, arms or legs

- Nausea and vomiting
- Problems with your kidney or urine

Less frequent side effects

- Inappropriate release of vasopressin hormone (ADH) which may lead to low sodium in the blood and water retention
- Reduced levels of magnesium and sodium in the blood
- Ringing in the ears
- Painful swelling and sores in the mouth, loss of appetite
- Changes in liver enzyme levels (as detected by a blood test can determine if the medicine is having an abnormal effect on your liver.)
- Hair loss
- Muscle pain
- Absence or decrease in sperm count, abnormal sperm production
- Fever
- Pain, swelling or red streaks at the injection site.

Frequency unknown

- Infection
- Early destruction of red blood cells leading to anaemia
- Destruction of red blood cells leading to kidney failure
- Blood amylase (enzyme) increased
- Dehydration, reduced level of calcium, phosphate, potassium in the blood, high level of uric acid in the blood, muscle cramps
- A dysfunction of the nerves that regulate non-voluntary body functions, such as heart rate, blood pressure and sweating, seizures, slurred speech, memory loss, spinal disease which may cause a sensation of electric shocks passing into your limbs

- Problems with your eyesight (blurred vision, odd colours, loss of vision or eye pain)
- Unusually cold or white hands and feet
- Hiccups, diarrhoea, discolouration of the gum line
- Increased bilirubin levels in the blood
- Difficulty breathing
- Rash
- Extreme tiredness/weakness

If you notice any side effects not mentioned in this leaflet, please inform your doctor or pharmacist.

Reporting of side effects

If you get 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 to SAHPRA via the "**6.04 Adverse Drug Reaction Reporting Form**", found online under SAHPRA's publications: https://www.sahpra.org.za/Publications/Index/8. By reporting side effects, you can help provide more information on the safety of CISPLATIN 50 mg/50 ml OETHMAAN.

5. How to store CISPLATIN 50 mg/50 ml OETHMAAN

STORE ALL MEDICINES OUT OF REACH OF CHILDREN.

Prior to first use: Store below 30 °C. Do not refrigerate. Protect from light. Keep vial in the outer carton until required for use. Discard any unused portion.

After dilution: After dilution with the recommended intravenous fluids, **CISPLATIN 50 mg/50 ml OETHMAAN** is chemically and physically stable for 12 hours at 20-25 ˚C. The diluted solution should be protected from light. Do not store diluted solutions in the refrigerator or freezer.

6. Contents of the pack and other information

What CISPLATIN 50 mg/50 ml OETHMAAN contains

CISPLATIN 50 mg/50 ml OETHMAAN: Each 1,0 ml solution for infusion contains 1,0 mg cisplatin as the active substance. Contains approximately 180 mg (7.8 mmol) of sodium per 50 ml.

Contains sugar (mannitol 50 mg/vial).

The other ingredients are: Sodium chloride, mannitol pyrogen free, hydrochloric acid and water for injection.

What CISPLATIN 50 mg/50 ml OETHMAAN looks like and contents of the pack

A clear colourless to pale yellow solution.

CISPLATIN 50 mg/50 ml OETHMAAN is packaged in a 50 ml, type 1 amber glass vial with a grey rubber stopper and a 20 mm aluminium flip-off transparent violet seal, which is further packed in an outer cardboard carton.

Holder of Certificate of Registration

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This leaflet was last revised in

Not applicable.

Registration number

57/26/0700