

PATIENT INFORMATION LEAFLET

SCHEDULING STATUS: **S4**

INOZI-CO 300 mg/300 mg (film coated tablet)

Isoniazid 300 mg and Rifapentine 300 mg

Sugar free

Read all of this leaflet carefully before you start taking INOZI-CO

- Keep this leaflet. You may need to read it again.
- If you have further questions, please ask your doctor, pharmacist, nurse or other health care provider.
- **INOZI-CO** 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 **INOZI-CO** is and what it is used for
2. What you need to know before you take **INOZI-CO**
3. How to take **INOZI-CO**
4. Possible side effects
5. How to store **INOZI-CO**
6. Contents of the pack and other information

1. What **INOZI-CO** is and what it is used for

INOZI-CO contains isoniazid and rifapentine. It is used to prevent progression of inactive (latent) tuberculosis infection (LTBI) to active tuberculosis disease in adults.

2. What you need to know before you take **INOZI-CO**

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Do not take INOZI-CO:

- If you are hypersensitive (allergic) to isoniazid or rifapentine or to any of the ingredients of **INOZI-CO** listed in section 6.1.
- If you are hypersensitive to medicines which are chemically related to isoniazid such as ethionamide, pyrazinamide (used for TB) and niacin (Vitamin B3).
- If you are pregnant, breastfeeding or planning to become pregnant during treatment.
- If you have acute or chronic liver disease
- If you have a condition called porphyria (a group of disorders that can cause nerve or skin problems (blisters and itching) due to a problem with the production of haem (blood pigment) and red blood cells within the body).
- If you experience fits.
- If you are alcoholic.
- If you have severe kidney problems
- If you are currently on specific antiretroviral therapy belonging to the classes protease inhibitors (e.g. indinavir, atazanavir, lopinavir, ritonavir) and non-nucleoside reverse transcriptase inhibitors (NNRTIs) (e.g. efavirenz, nevirapine)

Warnings and precautions**Talk to your doctor before taking INOZI-CO if you**

Have liver problems. Your doctor may do a blood test to check your liver function before and while you take **INOZI-CO** (see section Do not take **INOZI-CO** if).

Have active TB disease.

Take medicines to treat HIV infection or take oral contraceptives. Using these medicines with **INOZI-CO** may make them less effective (see Other medicines and **INOZI-CO**).

Use a medicine called digoxin (used to make the heart beat stronger and with a more regular rhythm). Your doctor will monitor your dose of digoxin as it could change due to **INOZI-CO**. Even small changes need to be monitored to prevent serious side effects from occurring (also see Other medicines and **INOZI-CO**).

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Wear contact lenses or dentures. **INOZI-CO** may change the normal colour of your skin, mouth and body fluids and cause your skin, teeth, tongue, urine, stools, saliva, sputum, tears, sweat, and breast milk to turn a red-orange colour. Contact lenses or dentures may become permanently stained.

If you experience any of the following after or while you are taking INOZI-CO, tell your doctor or health care provider immediately:

If you experience symptoms of a hypersensitivity (allergic) reaction after taking **INOZI-CO**, e.g. low blood pressure, hives, swelling of the face, lips, tongue or throat, difficulty breathing, red eyes or flu-like syndrome (weakness, extreme tiredness, muscle pain, nausea and vomiting, headache, fever, chills, aches, rash, itching, sweats, dizziness, shortness of breath, chest pain, cough, fainting, fast heartbeat). These may be symptoms of a serious allergic reaction and you must immediately stop taking **INOZI-CO** and go to the casualty department at your nearest hospital (see Possible side effects).

INOZI-CO may cause serious liver problems. Stop taking **INOZI-CO** and call your doctor right away if you have any of the following signs and symptoms of liver problems: nausea, vomiting, stomach pain, loss of appetite, tiredness, yellowing of the skin or whites of your eyes, dark urine (see Possible side effects).

Severe skin problems such as Stevens-Johnson syndrome (SJS) and drug reaction with eosinophilia and systemic symptoms (DRESS) syndrome have been reported with the use of **INOZI-CO**.

SJS symptoms may include blistering, peeling or bleeding on any part of your skin with or without a rash. This includes your lips, eyes, mouth, nose, genitals, hands or feet. You may also have flu-like symptoms at the same time, such as fever, chills or aching muscles.

DRESS symptoms and signs may include flu-like symptoms and a widespread rash with a high body temperature and enlarged lymph nodes. Abnormal blood test results may include increased levels of liver enzymes and an increase in a type of white blood cell (eosinophilia). If you develop any skin reaction, stop treatment immediately and contact your doctor or health care provider.

INOZI-CO may cause a type of diarrhoea called Clostridium difficile-associated diarrhoea (CDAD) which may occur during or after taking **INOZI-CO**. The severity of CDAD can range from mild diarrhoea to severe diarrhoea that may cause death (fatal colitis). Tell your doctor immediately if you have diarrhoea while you are taking or after you have stopped taking **INOZI-CO** (see Possible side effects).

Your doctor may prescribe Vitamin B₆ (pyridoxine) with **INOZI-CO** to minimise adverse reactions if you

are diabetic, alcoholic, HIV infected, malnourished or you have neuropathy (damage to the nerves resulting in poor coordination, muscle weakness, pain in the fingers or toes, or numbness).

Other medicines and INOZI-CO

Always tell your healthcare professional if you are taking any other medicine. (This includes complementary or traditional medicines.)

Tell your doctor or pharmacist if you are taking any of the following medicines:

INOZI-CO may make the following medicines work less well:

- medicines used to treat HIV infection: indinavir, darunavir, lopinavir, saquinavir, ritonavir, rilpivirine, efavirenz, zidovudine
- antifungal medicine (to treat fungal infections): itraconazole, ketoconazole, voriconazole
- pain killers (narcotic analgesics): methadone, alfentanil, buprenorphine
- medicines used for diabetes to lower blood sugar: repaglinide
- medicines to lower blood pressure, called calcium channel blockers: felodipine, diltiazem, verapamil, nifedipine
- medicines to lower blood pressure called alpha or beta adrenergic antagonists: alfuzosin, propranolol
- medicines to treat migraine, sometimes referred to as ergot alkaloid derivatives: ergotamine
- medicines used to prevent the blood from clotting (anticoagulants): warfarin
- hormonal contraceptives for women. If you are using any form of contraception, such as contraceptive tablets, transdermal patches (patches used on the skin) or an implant while taking **INOZI-CO** tablets, you should discuss the need to use an additional non-hormonal means of contraception or to change your contraceptive pill with your doctor or health care provider to make sure that your method of contraception is still effective.
- medicines used to lower your immune system: ciclosporin, tacrolimus, sirolimus
- medicines used as sedatives or to treat anxiety: midazolam.

Other medicines which could be affected by **INOZI-CO**:

- Digoxin (used to make the heart beat stronger and with a more regular rhythm). Your doctor will monitor your dose of digoxin as it could change due to **INOZI-CO**. Even small changes need to

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be monitored to prevent serious side effects from occurring (see Take special care with **INOZI-CO**).

- Medicines used for inflammation (e.g. ibuprofen, diclofenac).
- Medicines used to lower blood sugar in diabetes: glipizide.
- **INOZI-CO** may also interact with the laboratory tests to determine folate and vitamin B12 in your blood. Make sure that you inform your doctor or health care provider that you are using **INOZI-CO** before such tests are performed.

INOZI-CO with food and drink:

INOZI-CO should be taken with food.

Pregnancy, breastfeeding and fertility

If you are pregnant or breastfeeding your baby, please consult your doctor, pharmacist or other healthcare professional for advice before taking **INOZI-CO**.

Safety during pregnancy and breastfeeding has not been established.

Driving and using machines

INOZI-CO may cause dizziness, impaired concentration, and/or drowsiness. If you feel dizzy or drowsy while taking **INOZI-CO** do not drive and do not use any tools or machines.

3. How to take INOZI-CO

Do not share medicines prescribed for you with any other person.

Always take **INOZI-CO** exactly as your doctor has instructed you. You should check with your doctor or pharmacist if you are not sure.

INOZI-CO should be taken once-weekly, for 12 weeks as directly observed therapy (DOT). DOT is a way of helping people during their treatment, e.g. your treatment will be monitored weekly by visiting your local clinic, hospital or nurse and they will make sure you take your treatment correctly.

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Your doctor will calculate the number of tablets you should take weekly based on your weight. This dose will therefore be calculated for you personally.

It is important to take all of your **INOZI-CO**. Do not skip doses. Skipping doses may cause **INOZI-CO** to not work as well and may increase the chance that your TB will not be treatable by **INOZI-CO** or other medicines.

Take **INOZI-CO** with food. This may help prevent nausea, vomiting or stomach upsets due to the medication.

It is preferable to take antacids at least 1 hour before or 2 hours after your dose of **INOZI-CO** tablets. If you cannot swallow **INOZI-CO** tablets whole, they can be crushed and mixed with a small amount of semisolid food. Be sure to take all of the semisolid food with **INOZI-CO** in it immediately.

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

If you take more INOZI-CO than you should

In the event of over dosage, consult your doctor or pharmacist. If neither is available, contact the nearest hospital or poison control centre.

If you forget to take INOZI-CO

Since your weekly dose of **INOZI-CO** will be given to you via directly observed therapy (DOT) which includes a visit to, for example, your local clinic or hospital, it is unlikely that you will forget to take a dose. However, if you have missed the visit to your local clinic or hospital, visit them as soon as possible and do not skip the whole week until the next appointment.

If you stop taking INOZI-CO

If **INOZI-CO** is stopped earlier than your doctor has prescribed, your symptoms of may return or worsen. Always take **INOZI-CO** for as long as your doctor has prescribed.

4. Possible side effects

INOZI-CO can have side effects.

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Not all side effects reported for **INOZI-CO** are included in this leaflet. Should your general health worsen or if you experience any untoward effects while taking **INOZI-CO**, please consult your doctor, pharmacist or other healthcare professional for advice.

If any of the following happens, stop taking INOZI-CO and tell your doctor immediately or go to the casualty department at your nearest hospital:

- Hypersensitivity (allergic) reaction: this may present as a sudden life-threatening allergic reaction and the signs may include low blood pressure, hives, swelling of the face, lips, tongue or throat, difficulty breathing, red eyes or flu-like syndrome (see below).
- Flu-like syndrome (weakness, extreme tiredness, muscle pain, nausea and vomiting, headache, fever, chills, aches, rash, itching, sweats, dizziness, shortness of breath, chest pain, cough, fainting, fast heartbeat) (see Take special care with **INOZI-CO**).
- Hepatitis (inflammation of the liver) and liver problems (symptoms may include: nausea, vomiting, stomach pain, loss of appetite, tiredness, yellowing of the skin or whites of your eyes, dark urine) (see Take special care with **INOZI-CO**).
- Diarrhoea called *Clostridium difficile*-associated diarrhoea (CDAD) which may occur during or after taking **INOZI-CO**. The severity of CDAD can range from mild diarrhoea to severe diarrhoea that may cause death (fatal colitis (see Take special care with **INOZI-CO**).
- Pancreatitis (inflammation of the pancreas) (symptoms may include upper abdominal pain that radiates into the back; swollen and tender abdomen; nausea and vomiting, fever and increased heart rate).
- Pneumonia (infection of the lungs; symptoms may include fever and chills, cough, difficulty breathing, shortness of breath when you climb stairs).
- Seizures (have a “fit” or convulsion).

These are all very serious side effects. If you have them, you may have had a serious reaction to **INOZI-CO**. You may need urgent medical attention or hospitalisation.

Tell your doctor immediately or go to the casualty department at your nearest hospital if you notice any of the following:

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Less frequent:

- influenza (flu)
- upper abdominal pain
- skin reaction
- pain in one or more muscles (myalgia)
- Chills
- Fever
- abnormal physical weakness or lack of energy (asthenia).
- Feeling anxious depressed or disorientated.
- Blurred vision or loss of vision, with or without eye pain (optic neuritis).

Frequency unknown:

- SJS including the following symptoms: blistering, peeling or bleeding on any part of your skin with or without a rash (including your lips, eyes, mouth, nose, genitals, hands or feet), flu-like symptoms (fever, chills or aching muscles).
- DRESS including the following symptoms: flu-like symptoms, a widespread rash with a high body temperature and enlarged lymph nodes. Abnormal blood test results may include increased levels of liver enzymes and an increase in a type of white blood cell (eosinophilia).

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

Tell your doctor if you notice any of the following:

Frequent:

- Nausea, vomiting, constipation, dry mouth, diarrhoea, stomach pain, enlarged salivary glands.

Less frequent:

- Headache
- Fatigue or extreme tiredness
- Irritation of the oesophagus (the tube that carries food from your throat down to your stomach).
- Joint pain, back pain, fever, eating disorder (anorexia).

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

Should your general health worsen or if you experience any untoward effects while taking **INOZI-CO**, please consult your healthcare provider for advice.

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 **INOZI-CO**.

5. How to store INOZI-CO

Store at or below 25 °C in a dry place protected from light.

KEEP OUT OF THE REACH OF CHILDREN.

Keep the blisters in the carton until required for use.

Do not store in a bathroom.

Do not use after the expiry date stated on the blister / label / carton bottle.

Return all unused medicine to your pharmacist.

Do not dispose of unused medicine in drains or sewerage systems (e.g. toilets).

6. Contents of pack and other information

What INOZI-CO contains

The active ingredients are Isoniazid and Rifapentine.

Sugar free

Other ingredients are:

Tablet core contains

Calcium stearate, croscarmellose sodium, disodium EDTA, low-substituted hydroxypropyl cellulose, hypromellose, hydroxypropyl cellulose, iron oxide red, microcrystalline cellulose, povidone, pregelatinised starch, sodium ascorbate, sodium lauryl sulfate, sodium starch glycolate.

The tablet film-coating material contains

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Hypromellose, macrogol, red iron oxide, titanium dioxide,talc

What INOZI-CO looks like and contents of the pack

INOZI-CO

Reddish brown coloured, capsule shaped, biconvex, film-coated tablets debossed with "J" and "21" on either side of breakline on one side and plain on the other side.

The tablet can be divided into equal halves.

Contents of the pack:

Blister pack

Tablets are packed in a blister containing aluminium foil as lidding material and cold form laminate OPA/ aluminium foil / PVC as the forming material.

Blister pack in plain laminated pouch

Tablets are packed in a blister containing aluminium foil as lidding material and cold form laminate OPA/ aluminium foil / PVC as the forming material. Blister is packed in plain laminated pouch.

Strip pack

Tablets are packed in a strip pack containing aluminium foil (soft tempered) laminated with 150 gauge polyethylene film as forming and lidding material.

Pack sizes include 12, 28, 36, 56, 84, 112 or 120 tablets.

Not all pack sizes may be marketed.

7. HOLDER OF CERTIFICATE OF REGISTRATION

MACLEODS PHARMACEUTICALS SA (PTY) LTD
Ground Floor, Block 1,
Bassonia Estate Office Park (East),
1 Cussonia Drive,
Bassonia Rock Ext 12
Alberton
Gauteng

This leaflet was last revised in

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29 August 2023

Registration number

INOZI-CO: 54/20.2.3/0161.160

Access to the corresponding Professional Information

The corresponding professional information can be accessed at <https://www.macleodspharma.com> or email macleodsr@macleodspharma.com or contact 011 682 1169

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