

SUMMARY OF PRODUCT CHARACTERISTICS

1. NAME OF THE FINISHED PRODUCT

Diabetmin XR 500mg Tablet

2. QUALITATIVE AND QUANTITATIVE COMPOSITION

ACTIVE INGREDIENTS	PER TABLET (MG)
Metformin Hydrochloride	500 mg

Kindly refer to Section 6.1 for excipient.

3. PHARMACEUTICAL FORM

Oblong, white uncoated tablet with deep convex faces and 'HD' embossed on one face.

4. CLINICAL PARTICULARS

4.1 Therapeutic indication

Metformin is used in the treatment of non-insulin-dependent diabetes mellitus (type 2) in adults, not responding to exercise and dietary modification. Diabetmin may be used as monotherapy or in combination with other oral antidiabetic agents, or with insulin.

4.2 Posology and Method of administration

Oral.

Monotherapy and combination with other oral antidiabetic agents:

Usual adult dose:

Diabetmin XR 500mg Tablet Initial dose of one tablet once daily with meals, maximum 4 tablets daily.

Combination with insulin:

Metformin and insulin may be used in combination therapy to achieve better blood glucose control. Metformin is given at the usual starting dose of one tablet 2-3 times daily while insulin dosage is adjusted on the basis of blood glucose measurements.

Usual children dose:

Metformin is not recommended for use in children.

Usual geriatric dose:

Please refer to adult dose. (Due to potential for decreased renal function, the dosage should be adjusted based on renal function and maximum doses are not advised for use in the elderly.)

Contraindication

This medication is contraindicated in patients with the following medical problems:

- Hypersensitivity to metformin.
- Any condition needing close blood glucose control, such as: severe burns, dehydration, diabetic coma, diabetic ketoacidosis, hyperosmolar nonketotic coma, severe infection, major surgery, and severe trauma.
- Conditions associated with hypoxemia, such as: cardiorespiratory insufficiency, cardiovascular collapse, congestive heart failure, acute myocardial infarction.
- Severe, acute, or chronic hepatic disease.
- Active or history of lactic acidosis.
- Renal function impairment or renal disease.
- Diagnostic or medical examinations using intravascular iodinated contrast media such as: angiography, intravenous cholangiography, computed tomography (CT) scan, pyelography and urography.

Warnings and precautions

Lactic Acidosis:

Lactic acidosis is a rare, but serious (high mortality in the absence of prompt treatment), metabolic complication that can occur due to metformin accumulation. Reported cases of lactic acidosis in patients on metformin have occurred primarily in diabetic patients with significant renal failure. The incidence of lactic acidosis can and should be reduced by assessing also other associated risk factors such as poorly controlled diabetes, ketosis, prolonged fasting, excessive alcohol intake, hepatic insufficiency and any condition associated with hypoxia.

Diagnosis:

Lactic acidosis is characterized by acidosis, dyspnea, abdominal pain and hypothermia followed by coma. Diagnostic laboratory findings are decreased blood pH, plasma lactate levels above 5 mmol/L, and an increased anion gap and lactate/pyruvate ratio. If metabolic acidosis is suspected, metformin should be discontinued and the patient should be hospitalized immediately.

Renal Function:

As metformin is excreted by the kidney, serum creatinine levels should be determined before initiating treatment and regularly thereafter:

- at least annually in patients with normal renal function;
- at least 2 to 4 times a year in patients with serum creatinine levels at the upper limit of normal and in elderly subjects.

Decreased renal function in elderly subjects is frequent and asymptomatic. Special caution should be exercised in situations where renal function may become impaired, for example when initiating antihypertensive therapy or diuretic therapy and when starting therapy with an NSAID.

Administration of iodinated contrast agent:

As the intravascular administration of iodinated contrast materials in radiologic studies can lead to renal failure, metformin should be discontinued prior to, or at the time of the test and not

reinstated until 48 hours afterwards, and only after renal function has been re-evaluated and found to be normal.

Surgery:

Metformin should be discontinued before elective surgery with general anaesthesia and should not be usually resumed earlier than 48 hours afterwards.

Other precautions:

- All patients should continue their diet with a regular distribution of carbohydrate intake during the day. Overweight patients should continue their energy- restricted diet.
- The usual laboratory tests for diabetes monitoring should be performed regularly
- Metformin alone never causes hypoglycaemia, although caution is advised when it is used in combination with insulin or sulfonylureas.

Drug Interactions

Concurrent use of this medication with the following may interact with metformin:

- Acute or chronic ingestion of alcohol.
- Cimetidine or other cationic medications excreted by renal tubular transport.
- Furosemide.
- Hyperglycemia-causing and hypoglycemia-causing medications.

4.6 Pregnancy and lactation

For fertility and pregnancy problems, adequate and well-controlled studies in humans have not been done and documented. For patient plans to become pregnant or during pregnancy, control of blood glucose with diet alone or a combination of diet and insulin is recommended, while use of metformin is discouraged. Metformin is distributed into breast milk, but safety for use in nursing mothers has not been established.

4.7 Effects on ability to drive and use machines

Not applicable.

Main Side/ Adverse Effects

Metformin can cause:

- Gastrointestinal adverse effects including anorexia, diarrhea, dyspepsia, flatulence, nausea, vomiting.
- Headache, metallic taste, weight loss.
- Anemia, megaloblastic, hypoglycemia, lactic acidosis.
- Long-term metformin therapy may cause a decrease of vitamin B12 absorption with decrease of serum levels.

4.9 Overdose

Symptoms of overdose

Hypoglycemia and lactic acidosis.

Treatment of overdose

- For hypoglycemia: Treating with immediate ingestion of a source of glucose and counseling patient to obtain emergency medical assistance immediately.
- For lactic acidosis: Hemodialysis with sodium bicarbonate.

5. PHARMACOLOGICAL PROPERTIES

5.1 Pharmacodynamic properties

Metformin is an oral biguanide antidiabetic agent. Its mode of action is thought to be multifactorial and includes delayed uptake of glucose from the gastro-intestinal tract; increased peripheral glucose utilization mediated by increased insulin sensitivity; and inhibition of increased hepatic and renal gluconeogenesis.

5.2 Pharmacokinetic properties

Metformin Hydrochloride is slowly and incompletely absorbed from the gastrointestinal tract. The absolute bioavailability of a single 500mg dose is reported to be about 50 to 60%, although this is reduced somewhat if taken with food. Plasma protein binding is negligible. It is excreted unchanged in the urine. The plasma elimination half-life is reported to range from about 2 to 6 hours after oral administration.

5.3 Preclinical Safety Data

NOT APPLICABLE

6. PHARMACEUTICAL PARTICULARS

6.1 List of excipients

Metformin Hydrochloride
Copovidone
Microcrystalline Cellulose pH 102
Magnesium Stearate
Colloidal Silicon Dioxide
Carboxymethylcellulose Sodium
Methocel K100M CR

6.2 Incompatibilities

NOT APPLICABLE

6.3 Shelf life

3 years from date of manufacture

6.4 Special precaution for storage

Store below 30°C. Protect from light and moisture.

Nature and contents of container

Immediate Container Closure System / Primary Packaging

Primary Packaging

Blister pack

Type

Push-through blister pack; the package consists of a transparent thermoformable plastic material and a heat-sealed, lacquered backing material.

Polyvinylidene Chloride (PVDC)

Description : Clear transparent film

Appearance : Not more than 8.0%

Aluminium blister foil

Description : Aluminium foil with high slip primer on bright surface and heat seal on matt surface

Appearance : Bright surface/Matt surface each side

Outer Container / Secondary Packaging

Outer Container/Packaging

Type: Unit box, Package Insert & Plain Carton for Diabetmin XR Tablet

Instructions for use and handling <and disposal>

NOT APPLICABLE

7. MARKETING AUTHORISATION HOLDER

Name : HOVID Bhd.
Address : 121, Jalan Tunku Abdul Rahman,
(Jalan Kuala Kangsar)
30010 Ipoh, Perak, Malaysia

Manufacturer Name :

Name : HOVID Bhd.
Address : Lot 56442, 7 ½ Miles,
Jalan Ipoh / Chemor,
31200 Chemor,
Perak., Malaysia.

8. NUMBER (S) IN THE NATIONAL REGISTER OF FINISHED PHARMACEUTICAL PRODUCTS

05723/07489/REN/2020

9. DATE OF FIRST AUTHORISATION

2020

10. DATE OF REVISION OF THE TEXT

April 2018