PRODUCT INFORMATION

NAME OF THE MEDICINE

Active ingredient: metoprolol tartrate
Chemical name: di-[(±)-1-(isopropylamino)-3-[p-(2-methoxyethyl) phenoxy]-2-propanol] L(+) -tartrate
Structural formula:

Molecular formula: \((C_{15}H_{25}NO_3)_2\cdot C_4H_6O_6\)
Molecular weight: 685
CAS Registry no.: 56392-17-7

DESCRIPTION

Metoprolol tartrate, an aryloxypropanolamine derivative, is a white crystalline powder with a melting point of 120°C. The powder is odourless or almost odourless. It is very soluble in water, soluble in chloroform, methylene chloride and alcohol, and almost insoluble in benzene, diethylether and acetone.

Each Minax 50 tablet contains 50 mg of metoprolol tartrate. The tablets also contain the following inactive ingredients: lactose, microcrystalline cellulose, povidone, colloidal anhydrous silica, sodium starch glycollate and magnesium stearate.

Each Minax 100 tablet contains 100 mg of metoprolol tartrate. The tablets also contain the following inactive ingredients: lactose, microcrystalline cellulose, povidone, purified talc, colloidal anhydrous silica, sodium starch glycollate, carmellose sodium and magnesium stearate.

PHARMACOLOGY

Pharmacodynamics

Metoprolol is a relatively cardioselective β-blocker, i.e. it acts on β₁-receptors mainly located in the heart at lower doses than those needed to influence the β₂-receptors mainly located in the bronchi and peripheral vessels.

Metoprolol has little membrane-stabilising effect, nor does it display partial agonist activity, i.e. intrinsic sympathomimetic activity (ISA), at doses required to produce β-blockade.

Metoprolol reduces or inhibits the stimulant effect of catecholamines on the heart. This leads to a decrease in heart rate, cardiac contractility and cardiac output.
Metoprolol lowers elevated blood pressure in both the standing and lying position. It also reduces the extent of rises in blood pressure occurring in response to physical and mental stress.

In angina pectoris, metoprolol decreases the frequency and severity of the attacks, and the need for glyceryl trinitrate relief, and increases exercise tolerance.

Metoprolol has a regulating effect on the heart rate in cases of supraventricular tachycardia or atrial fibrillation, and in the presence of ventricular extrasystoles.

Metoprolol lowers mortality in patients with a suspected or confirmed myocardial infarction. This effect may possibly be attributable to a decrease in the incidence of severe ventricular arrhythmias, as well as to limitation of infarct size. Metoprolol has also been shown to reduce the incidence of recurrent myocardial infarction.

Due to its β-blocking effect, metoprolol is suitable for the prevention of migraine.

Metoprolol has been shown to reduce diuretic-induced increase in plasma renin activity. It inhibits catecholamine-induced insulin secretion to a far lesser degree than nonselective β-blockers.

Orthostatic reactions or disturbances of electrolyte balance have not been observed.

In therapeutic doses, metoprolol has less effect on peripheral circulation and the bronchial muscles than nonselective β-blockers. However, it should be used with caution in patients with asthma, and concomitant use of an adrenergic bronchodilator, e.g. terbutaline or salbutamol, is recommended. Patients already taking β₂-stimulants for reversible airways obstruction may require adjustment of dosage of these if metoprolol therapy is subsequently introduced.

Metoprolol inhibits catecholamine-induced lipolysis.

The active metabolite of metoprolol, 2-hydroxymetoprolol, does not contribute significantly to the therapeutic effect.

Metoprolol is a relatively lipid soluble compound i.e. less soluble than propranolol and more lipid soluble than atenolol.

**Pharmacokinetics**

**Absorption and Distribution**

Metoprolol is rapidly and almost completely (more than 95%) absorbed from the gastrointestinal tract. It is rapidly and extensively distributed to the extravascular tissue. The volume of distribution is 5.6 L/kg. At therapeutic concentrations approximately 12% of metoprolol is bound to human serum proteins.

**Metabolism and Excretion**

Studies with radioactively labelled drug have shown that more than 90% of the dose is excreted in the urine in 72 hours, mainly in the form of known metabolites. Only about 3% of the administered dose is excreted unchanged in the urine in 72 hours. The rate of renal excretion of metoprolol has a linear relationship to its plasma concentration. Metoprolol is excreted mainly by glomerular filtration.

Long-term studies have shown that metoprolol neither enhances nor inhibits its own metabolism.

The elimination half-life of metoprolol is between 3 and 5 hours.

**Dose-response**

The duration of the β-blocking effects is dose dependent (as measured by reduction of exercise heart rate). For instance, in healthy subjects the effect of 20 mg metoprolol given intravenously is halved after about 6 hours.
Pharmacokinetics in the elderly

Elderly subjects showed no significant differences in the plasma concentrations of metoprolol as compared with young persons, in a study involving eight healthy elderly individuals (mean age 74.5 years) and eight young subjects (mean age 26.3 years).

INDICATIONS

- **Hypertension**: as monotherapy or for use in combination with other antihypertensives.
- **Angina pectoris**: for long-term prophylaxis. Glyceryl trinitrate should be employed if necessary for alleviating acute attacks.
- Confirmed or suspected myocardial infarction
- Prevention of migraine.

CONTRAINDICATIONS

- Bronchospasm.
  
  Beta-adrenergic blockade of the smooth muscle of bronchi and bronchioles may result in an increased airways resistance. These drugs also reduce the effectiveness of asthma treatment. This may be dangerous in susceptible patients.

  Therefore, β-blockers are contraindicated in any patient with a history of airways obstruction or a tendency to bronchospasm. Use of cardioselective β-blockers can also result in severe bronchospasm. If such therapy must be used, great caution should be exercised. Alternative therapy should be considered.

- Hypersensitivity to metoprolol tartrate, related derivatives, or any of the excipients in MINAXs.

- Sensitivity to other β-blockers (cross-sensitivity between β-blockers can occur).

- Second and third degree atrioventricular block.

- Non-compensated congestive heart failure (see PRECAUTIONS).

- Sinus bradycardia (less than 45 to 50 beats/minute).

- Sick-sinus syndrome (unless a permanent, appropriately functioning pacemaker is in place).

- Severe peripheral arterial circulatory disorders.

- Shock (including cardiogenic and hypovolaemic shock).

- Myocardial infarction patients with a heart rate of < 45 beats/minute, a P-R interval of > 0.24 seconds, a systolic blood pressure of <100 mmHg, and/or moderate to severe non-compensated heart failure.

- Right ventricular failure secondary to pulmonary hypertension.

- Significant right ventricular hypertrophy.

- Hypotension.

- Untreated phaeochromocytoma (see PRECAUTIONS).
• Allergic disorders (including allergic rhinitis) which may suggest a predisposition to bronchospasm.

• Continuous or intermittent inotropic therapy acting through beta receptor agonism.

**PRECAUTIONS**

**Bronchospastic disease**

In general, patients with bronchospastic disease should not be given β-blockers of any type (e.g. selective or nonselective). If such therapy must be used, great caution should be exercised. Alternative therapy should be considered.

**Cardiac failure**

Beta-blockade depresses myocardial contractility and may precipitate cardiac failure in some patients with a history of cardiac failure, chronic myocardial insufficiency or unsuspected cardiomyopathy. In patients without a history of cardiac failure, continuing depression of the myocardium may lead to cardiac failure. If signs of cardiac failure are present, the patient should be fully digitalised and/or given a diuretic and carefully monitored. If cardiac failure persists, metoprolol tartrate should be discontinued gradually (see **PRECAUTIONS - Abrupt withdrawal**).

Beta-blockers should not be used in patients with untreated congestive heart failure. This condition should first be stabilised. (Note: Although congestive heart failure has been considered to be a contraindication to the use of β-blockers, there is a growing literature on the experimental use of β-adrenergic blocking drugs in heart failure. As further trials are needed to identify which patients are most likely to respond to which drugs, β-blockers should not normally be prescribed for heart failure outside of specialist centres).

**Prinzmetal angina**

There is a risk of exacerbating coronary artery spasm if patients with Prinzmetal or variant angina are treated with a β-blocker. If this treatment is essential, it should only be undertaken in a Coronary or Intensive Care Unit.

**Conduction disorders**

Very rarely, a pre-existing A-V conduction disorder of moderate degree may become aggravated (possibly leading to A-V block). Minax should be administered with caution to patients with first degree A-V block (see **CONTRAINDICATIONS**).

**Phaeochromocytoma**

In patients with phaeochromocytoma, an alpha-blocker (e.g. phentolamine or phenoxybenzamine) should be administered before the β-blocker to avoid exacerbation of hypertension.

**Diabetes**

Minax should be used with caution in patients with diabetes mellitus, especially those who are receiving insulin or oral hypoglycaemic agents. Diabetic patients should be warned that β-blockers affect glucose metabolism and may mask some important premonitory signs of acute hypoglycaemia, such as tachycardia.

In patients with insulin or non-insulin dependent diabetes, especially labile diabetes, or with a history of spontaneous hypoglycaemia, β-blockade may result in the loss of diabetic control and delayed recovery from hypoglycaemia. The dose of insulin or oral hypoglycaemic agent may need adjustment. Diabetic patients receiving Minax should be monitored to ensure that diabetes control is maintained.
Allergic conditions

Allergic reactions may be exaggerated by β-blockade (e.g. allergic rhinitis during the pollen season and allergic reactions to bee and wasp stings). Beta-blockers should be avoided if there is a risk of bronchospasm.

In patients taking β-blockers, anaphylactic shock assumes a more severe form and may be resistant to normal doses of adrenaline. Whenever possible, β-blockers should be avoided in patients who are at increased risk of anaphylaxis.

Hyperthyroidism

Special care should be exercised in those patients who are hyperthyroid and are also receiving β-blockers because β-blockers may mask the clinical signs of developing or continuing hyperthyroidism, resulting in symptomatic improvement without any change in thyroid status. Where Minax is administered to patients having, or suspected of developing thyrotoxicosis, both thyroid and cardiac function should be monitored closely.

Peripheral vascular disease

Beta-blockade may impair the peripheral circulation and exacerbate the symptoms of peripheral vascular disease (see CONTRAINDICATIONS).

Renal disease

In patients with severe renal disease, haemodynamic changes following β-blockade may impair renal function further. Beta-blockers which are excreted mainly by the kidney may require dose adjustment in patients with renal failure.

Liver disease

Metoprolol is mainly eliminated by means of hepatic metabolism (see PHARMACOLOGY - Pharmacokinetics). Therefore, liver cirrhosis may increase the systemic bioavailability of metoprolol and reduce its total clearance, leading to increased plasma concentrations.

Use in the elderly

Caution is indicated in elderly patients. An excessive decrease in blood pressure or pulse rate may cause the blood supply to vital organs to fall to inadequate levels.

Concomitant therapy with calcium antagonists

The concomitant use of β-blockers and calcium antagonists with myocardial depressant and sinus node activity (e.g. verapamil and to a lesser extent diltiazem) may cause hypotension, bradycardia and asystole. Extreme caution is required if these drugs have to be used together.

A calcium channel blocker of the phenylalkylamine type (e.g. verapamil) should not be administered intravenously to patients receiving metoprolol because there is a risk of cardiac arrest in this situation. Patients taking an oral calcium channel blocker of this type in combination with metoprolol should be closely monitored.

The combination of β-blockers with dihydropyridine calcium channel blockers with a weak myocardial depressant effect (e.g. felodipine, nifedipine) can be administered together with caution. In case excess hypotension develops, the calcium antagonist should be stopped or the dosage reduced.

Antiarrhythmic drugs

Care should be taken when prescribing β-blockers with antiarrhythmic drugs. Interactions have been reported during concomitant β-blocker therapy with the Class IA agents disopyramide, and less frequently quinidine;
Class IB agents, tocainide, mexiletine and lignocaine; Class IC agents, flecainide and propafenone (not available in Australia); the Class III agent amiodarone; and the class IV agents (e.g. verapamil).

**Clonidine**

Concurrent use of β-blockers and clonidine should be avoided because of the risk of adverse interaction and severe withdrawal symptoms. If administered concomitantly, the clonidine should not be discontinued until several days after the withdrawal of the β-blocker.

**Catecholamine-depleting agents**

Concomitant use of drugs such as reserpine and guanethidine requires careful monitoring since the added effect of a β-blocker may produce an excessive reduction of the resting sympathetic nervous tone.

**General anaesthetics**

Beta-blockade may have beneficial effects in decreasing the incidence of arrhythmias and myocardial ischaemia during anaesthesia and the post-operative period. It is currently recommended that maintenance β-blockade be continued peri-operatively. The anaesthetist must be made aware of β-blockade because of the potential for interactions with other drugs, resulting in severe bradyarrhythmias and hypotension, the decreased reflex ability to compensate for blood loss, hypovolaemia and regional sympathetic blockade, and the increased propensity for vagal-induced bradycardia. Incidents of protracted severe hypotension or difficulty restoring normal cardiac rhythm during anaesthesia have been reported.

Acute initiation of high-dose metoprolol to patients undergoing non-cardiac surgery should be avoided, since it has been associated with bradycardia, hypotension and stroke including fatal outcome in patients with cardiovascular risk factors.

Modern inhalational anaesthetic agents are generally well tolerated, although older agents (ether, cyclopropane, methoxyflurane, trichlorethylene) were sometimes associated with severe circulatory depression in the presence of β-blockade. If it is thought necessary to withdraw β-blocker therapy before surgery, this should be done gradually and completed about 48 hours before surgery (see PRECAUTIONS - Abrupt Withdrawal).

**Effects on the heart rate**

If the patient develops increasing bradycardia (heart rate less than 50 to 55 beats/minute), the dosage of Minax should be gradually reduced or treatment gradually withdrawn (see CONTRAINDICATIONS).

**Effects on the thyroid**

The effects of β-blockers on thyroid hormone metabolism may result in elevations of serum free thyroxine (T₄) levels. In the absence of any signs or symptoms of hyperthyroidism, additional investigation is necessary before a diagnosis of thyrotoxicosis can be made.

**Other metabolic effects**

Beta-adrenoceptors are involved in the regulation of lipid as well as carbohydrate metabolism. Some drugs affect the lipid profile adversely although the long-term clinical significance of this change is unknown and the effect appears to be less for drugs with intrinsic sympathomimetic activity.

**Effects on the eye and skin**

Various rashes and conjunctival xeroses have been reported with β-blocking agents. Cross reactions may occur between β-blockers, therefore substitutions within the group may not necessarily preclude occurrence of symptoms.

During long-term treatment with the β-blocking drug, practolol, a specific rash bearing a superficial resemblance to psoriasis was occasionally described. In a number of patients affected, this rash was
accompanied by adverse effects on the eye (xerophthalmia and/or keratoconjunctivitis) of varying severity. This condition is called the oculomucocutaneous syndrome or practolol syndrome. On a few rare occasions, serous otitis media, sclerosing peritonitis, pericarditis and pleurisy have been reported.

The oculomucocutaneous syndrome as reported with practolol has not been reported with metoprolol. However, dry eyes and skin rash have been reported with metoprolol. If such symptoms occur, discontinuation of metoprolol should be considered.

More recently, an association between Peyronie's disease (a fibrosing induration of the penis) and various β-blockers has been suggested but is not proven.

**Abrupt withdrawal**

Care should be taken if β-blockers have to be discontinued abruptly in patients with coronary artery disease. Severe exacerbation of angina and precipitation of myocardial infarction and ventricular arrhythmias have occurred following abrupt discontinuation of β-blockade in patients with ischaemic heart disease.

Therefore, it is recommended that the dosage be reduced gradually over a period of about 8-14 days during which time the patient's progress should be assessed. Minax should be temporarily reinstated if the angina worsens.

If the drug must be withdrawn abruptly in these patients, close observation is required. In the peri-operative period, metoprolol should not be withdrawn unless indicated.

**Use in Pregnancy (Category C)**

Metoprolol has been shown to increase postimplantation loss and decrease neonatal survival in rats at doses up to 55.5 times the maximum daily human dose of 450 mg. Distribution studies in mice confirm exposure of the foetus when Minax is administered to the pregnant animal. These studies have revealed no evidence of impaired fertility or teratogenicity. There are no adequate and well controlled studies in pregnant women. Because animal reproduction studies are not always predictive of human response, this drug should be used during pregnancy only if clearly indicated.

Beta-adrenergic blocking agents may reduce placental perfusion and cause pharmacological effects such as bradycardia in the foetus and newborn infant. Metoprolol crosses the placental barrier in pregnant women; in one study the concentration in the umbilical vein was almost the same as in maternal vein plasma.

During the later stages of pregnancy, these drugs should only be given after weighing the needs of the mother against the risk to the foetus.

The lowest possible dose should be used and treatment should be discontinued at least 2 to 3 days before delivery to avoid increased uterine contractility and effects of β-blockade in the newborn (e.g. bradycardia, hypoglycaemia).

**Use in Lactation**

Metoprolol is excreted in human breast milk. Beta-blockers taken by the mother may cause side-effects, e.g. bradycardia, in the breast fed infant, although when the doses used are within the recommended therapeutic range, the very small amount of drug ingested by the infant renders such effects unlikely. Nevertheless, breastfed infants should be closely observed for signs or symptoms of β-blockade.

Experience suggests that metoprolol only need be discontinued during lactation if the infant's hepatic function is severely impaired.

**Paediatric Use**

The safety and efficacy in children have not been established.
Effects on ability to drive or use machinery

Minax may cause dizziness, fatigue or visual disturbances (see ADVERSE EFFECTS) and, therefore, may adversely affect the patient's ability to drive or use machinery.

INTERACTIONS WITH OTHER MEDICINES

Other anti-hypertensive agents

Metoprolol enhances the effect of other antihypertensive drugs. Particular caution is required when administering a β-blocker and prazosin together for the first time.

Sympathetic ganglion blocking agents, other beta-blockers or monoamine oxidase (MAO) inhibitors

Patients receiving concurrent treatment with sympathetic ganglion blocking agents, other β-blockers (including eye drops), or monoamine oxidase (MAO) inhibitors should be carefully monitored.

Clonidine

If concomitant treatment with clonidine is to be discontinued, the β-blocker medication should be withdrawn several days before clonidine. The rebound hypertension associated with clonidine withdrawal can be exacerbated by the presence of a β-blocker. If both drugs are withdrawn simultaneously, marked rise in blood pressure, and/or arrhythmias may result.

Calcium antagonists

When metoprolol is given together with calcium antagonists of the verapamil and diltiazem type and/or anti-arrhythmic agents, the patient should be monitored for possible negative inotropic and chronotropic effects. Calcium antagonists of the verapamil type should not be given by intravenous administration to patients treated with β-blockers.

Anti-arrhythmic agents

When metoprolol is given together with anti-arrhythmic agents the patient should be monitored for possible negative inotropic and chronotropic effects. The negative inotropic and negative chronotropic effects of antiarrhythmic agents of the quinidine type and amiodarone may be enhanced by beta-blockers.

Prostaglandin synthetase inhibiting agents

Concurrent treatment with indomethacin or other prostaglandin synthetase inhibiting agents may decrease the antihypertensive effect of β-blockers.

Alcohol

Metoprolol may modify the pharmacokinetic behaviour of alcohol when taken concomitantly. The plasma level of metoprolol may be raised by alcohol.

Liver enzyme effects

Enzyme-inducing and enzyme-inhibiting substances may change the plasma concentration of metoprolol. The plasma concentration of metoprolol is lowered by rifampicin and may be raised by cimetidine, alcohol, hydralazine and selective serotonin reuptake inhibitors (SSRIs) e.g. paroxetine, fluoxetine and sertraline.

Oral antidiabetic agents

The dosage of oral antidiabetics may need to be adjusted in patients receiving β-blockers (see PRECAUTIONS).
Anaesthetics

Metoprolol may reduce the clearance of other drugs (e.g. lignocaine). Inhalation anaesthetics enhance the cardiodepressant effect of β-blocker therapy (see PRECAUTIONS). Metoprolol may also reduce the clearance of other drugs (e.g. lignocaine).

Warfarin

A limited number of reports have demonstrated a rise in AUC and concentration of warfarin when taken with another β-blocker. This could potentially increase the anticoagulant effect of warfarin.

Digitalis glycosides

Digitalis glycosides, in association with beta blockers, may increase atroventricular conduction time and may induce bradycardia.

ADVERSE EFFECTS

Occasionally, especially at the start of treatment, beta-blockers may give rise to gastro-intestinal upsets, sleep disturbances, or exertional tiredness. These effects, however, are of a mild nature and seldom necessitate a reduction in the dosage.

The following events have been reported as adverse events in clinical trials or reported from routine use. In many cases a relationship with metoprolol has not been established. The following definitions of frequency are used: very common ≥ 10%; common 1 - 9.9%; uncommon 0.1 - 0.9%; rare 0.01 - 0.09%; very rare < 0.01%.

Cardiovascular

Common: bradycardia, postural disorders (very rarely with syncope), cold hands and feet (Raynaud's phenomenon), palpitations, clinically significant falls in blood pressure after intravenous administration.


Rare: disturbances of cardiac conduction, cardiac arrhythmias.

Very rare: gangrene in patients with pre-existing severe peripheral circulatory disorders.

*Excess frequency of 0.4% compared with placebo in a study of 46000 patients with acute myocardial infarction where the frequency of cardiogenic shock was 2.3% in the metoprolol group and 1.9% in the placebo group in the subset of patients with low shock risk index. The shock risk index was based on the absolute risk of shock in each individual patient derived from age, sex, time delay, Killip class, blood pressure, heart rate, ECG abnormality, and prior history of hypertension. The patient group with low shock risk index corresponds to the patients in which metoprolol is recommended for use in acute myocardial infarction.

Central nervous system

Very common: fatigue

Common: dizziness, headache.

Uncommon: paraesthesia, muscle cramps.

Gastrointestinal

Common: nausea, diarrhoea, constipation, abdominal pain.

Uncommon: vomiting.

Rare: dry mouth.
Haematologic

Very rare: thrombocytopenia.

Hepatic

Rare: liver function test abnormalities.
Very rare: hepatitis

Metabolic

Uncommon: weight gain.

Psychiatric

Uncommon: depression, impaired concentration, somnolence or insomnia, nightmares.
Rare: nervousness, anxiety, impotence / sexual dysfunction.
Very rare: amnesia / memory impairment, confusion, hallucinations.

Respiratory

Common: dyspnoea on exertion.
Uncommon: bronchospasm (which may also occur in patients without a history of obstructive lung disease).
Rare: rhinitis.

Sense organs

Rare: disturbances of vision, dry and/or irritated eyes, conjunctivitis (see PRECAUTIONS).
Very rare: tinnitus, taste disturbances.

Skin

Uncommon: rash (in the form of urticaria, psoriasiform and dystrophic skin lesions), increased sweating.
Rare: loss of hair
Very rare: photosensitivity reactions, aggravated psoriasis.

Miscellaneous

Very rare: arthralgia

DOSAGE AND ADMINISTRATION

It is advisable to individualise the dosage.

Hypertension

Mild: 50 or 100 mg once daily for one week.

Moderate to severe: 50 or 100 mg twice daily for one week.
**Maintenance:** 50 or 100 mg once or twice daily. Some patients may respond to 50 mg once daily. A larger number will respond to 100 mg once daily as initial and maintenance therapy. Response is rarely improved by increasing the dose beyond 200 mg daily.

The maximum daily dose should not exceed 400 mg. Although twice daily dosage is optimal, in those patients whose maintenance dosage is 150 mg daily or less, it may be administered as a single dose.

**Angina pectoris**

50 mg to 100 mg two or three times daily.

**Myocardial infarction**

The recommended dosage can be reduced depending on the haemodynamic status of the patient. Initially, therapy should commence with 50 mg twice daily and be continued for 48 hours.

**Maintenance:** Generally 100 mg twice daily.

**Prevention of migraine**

100 to 150 mg given in divided doses, morning and evening.

**OVERDOSE**

**Signs and symptoms**

An overdosage of Minax may lead to severe hypotension, sinus bradycardia, atrioventricular block, heart failure, cardiogenic shock, cardiac arrest, bronchospasm, impairment of consciousness (or even coma), convulsions, nausea, vomiting and cyanosis.

Concomitant ingestion of alcohol, antihypertensives, quinidine or barbiturates may aggravate the patient’s condition.

The first manifestations of overdosage can appear in 20 minutes but are more commonly seen within 1 to 2 hours after the drug's ingestion. The effects of massive overdosage may persist for several days despite declining plasma concentrations.

**Treatment**

Care should be provided at a facility that can provide appropriate supporting measures, monitoring, and supervision.

Activated charcoal may reduce absorption of the medicine if given within one or two hours after ingestion. In patients who are not fully conscious or have impaired gag reflex, consideration should be given to administering activated charcoal via a nasogastric tube, once the airway is protected.

Syrup of ipecac and gastric lavage are no longer considered to be standard therapy for gut decontamination.

Atropine, adreno stimulating drugs or pacemaker to treat bradycardia and conduction disorders.

Hypotension, acute cardiac failure, and shock to be treated with suitable volume expansion, injection of glucagon (if necessary, followed by an intravenous infusion of glucagon), intravenous administration of adreno stimulating drugs such as dobutamine, with α1 receptor agonistic drugs added in presence of vasodilation. Intravenous use of calcium salts (Ca2+) can also be considered.

Bronchospasm can usually be reversed by bronchodilators.

Contact the Poisons Information Centre on 131126 (Australia) for advice on management of overdosage.
PRESENTATION AND STORAGE CONDITIONS

Minax 50  :  White, round, marked "ML/50" on one side, "α" on the reverse
            Available in blister* and bottle packs of 100's.

Minax 100 :  White, round, marked "ML/100" on one side, "α" on the reverse
            Available in blister* and bottle packs of 60's.

* Not available in Australia.

Store below 30°C.

NAME AND ADDRESS OF THE SPONSOR

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POISON SCHEDULE OF THE MEDICINE

S4 (Prescription Only Medicine)

DATE OF FIRST INCLUSION IN THE AUSTRALIAN REGISTER OF THERAPEUTIC GOODS (THE ARTG)

MINAX 100 tablet blister pack:  23/11/1992
MINAX 50 tablet blister pack:  23/11/1992
MINAX 100 tablet bottle:  24/11/1992
MINAX 50 tablet bottle:  24/11/1992

DATE OF MOST RECENT AMENDMENT

22/12/2015