

# ANZARIL TABLET

Clozapine (25mg, 100mg)

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## What Anzaril Tablet is used for

Clozapine is the active ingredient of Anzaril Tablet. It is indicated in people with treatment-resistant schizophrenic only i.e. schizophrenic (a severe brain disorder in which people interpret reality abnormally) people who are non-responsive to or intolerant to classic neuroleptic (antipsychotic) medicines.

Anzaril Tablet is also used to treat people suffering from schizophrenia or schizoaffective disorders (schizophrenia with mood disorders) who might otherwise attempt to commit suicide. Schizophrenia is a mental illness involving disturbances in thinking, emotional reactions, and behavior.

## How Anzaril Tablet works

Clozapine is a psychiatric medication (antipsychotic type) that works by changing the activity of certain natural substances in the brain.

## Before you use Anzaril Tablet

### - When you must not use it

- If you are allergic (hypersensitive) or to clozapine or any other components of the formulations.
- If you have a history of toxic or idiosyncratic granulocytopenia/agranulocytosis (low white blood cell count).
- If you have impaired bone marrow function.
- If you have uncontrolled epilepsy.

- If you have alcoholic and other toxic mental disorder, drug intoxication and comatose conditions.
- If you have circulatory collapse (rapid or sudden development of circulatory failure) and/or CNS depression (physiological depression of the central nervous system that can result in decreased rate of breathing, decreased heart rate, and loss of consciousness possibly leading to coma or death) of any cause.
- If you have severe kidney or heart disorders.
- If you have active liver disease associated with nausea, anorexia (eating disorder), jaundice, progressive liver disease or liver failure.
- If you suffer or have ever suffered from severe constipation, obstruction of the bowel or any other condition which has affected your large bowel.
- If you are unable to undergo regular blood tests.
- Before you start to use it  
Talk to your doctor or pharmacist if:
  - You have allergies to any medicine which you have taken previously to treat your current condition.
  - You are pregnant, trying to become pregnant or breast-feeding.
  - You are taking medicine known to have a substantial potential to depress bone marrow function.
  - Have or are at a risk of having diabetes (e.g being overweight or a family history of diabetes). Your doctor should check your blood sugar before you start taking Anzaril Tablet and regularly during treatment.

### - Taking other medicines

- Please tell your doctor or pharmacist if you are taking or have recently taken any of these medicines:
- Alcohol, monoamine oxidase inhibitors (MAOIs) inhibitors

and central nervous system (CNS) depressants such as antihistamine-may enhance the central effect.

- Benzodiazepine and other psychotropic medicines-increased risk of circulatory collapse which may lead to cardiac arrest (sudden cessation of heartbeat and heart function, resulting in the loss of effective circulation) and/or respiratory arrest (cessation of breathing).
- Medicines which cause constipation (such as anticholinergic which are used to relieve stomach cramps, spasm and travel sickness).
- Lithium or central nervous system (CNS) active agents-increase risk of development of neuroleptic malignant syndrome (NMS). NMS is a life-threatening neurological disorder most often caused by an adverse reaction to neuroleptic or antipsychotic medicines.
- Norepinephrine- may reduce the blood pressure increasing effect of Norepinephrine.
- Valproic acid-onset of seizures in non-epileptic individuals and isolated cases of delirium (a serious disturbance in mental abilities that results in confused thinking and reduced awareness of your environment).
- Carbamazepine, phenytoin and rifampicin omeprazole, nicotine-decrease the blood levels of clozapine.
- Cimetidine, erythromycin and fluvoxamine, azole antimycotics, protease inhibitor, caffeine, paroxetine, sertraline, fluoxetine- increase blood levels of clozapine.

Always inform your doctor and pharmacist if you are taking any other medicines, including herbal

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tonics, supplements and medicines that you buy without prescription. Talk to your pharmacist or doctor if you are not sure about this.

## How to use Anzaril Tablet

Always take this medicine exactly as your doctor has told you. You should check with your doctor or pharmacist if you are not sure. Take this medicine only by mouth.

### - How much to use

#### *Starting therapy*

12.5mg once or twice on the 1<sup>st</sup> day, followed by one or two 25mg tablets on the 2<sup>nd</sup> day. If well tolerated, daily dose may be increased slowly in increments of 25mg to 50mg in order to achieve a dose level of up to 300mg/day within 2-3 weeks.

Thereafter, if required the daily dose may be further increased in increment of 50mg to 100mg at half-weekly or preferably weekly intervals.

#### *Use in elderly*

Initiate treatment at 12.5mg given once and restrict subsequent dose increments to 25mg/day.

#### *Patients with heart disease*

The starting dose should be low (1 x 12.5 mg on the first day) in patients with heart disease. It should be increased slowly and in small increments. It is contraindicated for patients with severe cardiovascular disease.

#### *Patients with renal insufficiency*

The starting dose should be low (1x 12.5 mg on the first day) in patients with mild to moderate renal insufficiency. It should be increased slowly and in small increments.

#### *Patients with hepatic insufficiency*

In patients with hepatic insufficiency, clozapine should only be administered with care and regular monitoring.

#### *Therapeutic dose*

300-450mg per day in divided dose. Some may require up to 600mg per day.

#### *Maximum Dose*

The possibility of increase in adverse reactions (in particular seizures) occurring at doses over 450mg/day must be borne in mind.

#### *Maintenance Dose*

After achieving maximum therapeutic benefit, many patients can be maintained effectively on lower doses. Careful downward titration is therefore recommended. Treatment should be maintained for at least 6 months. If the daily dose not exceed 200 mg, once daily administration in the evening may be appropriate.

#### *Ending Dose*

Gradual reduction over 1-2 week period is recommended.

#### *Restarting Therapy*

In individual whom the interval since the last dose exceeds 2 days, treatment should be re-initiated at 12.5mg given once or twice on the 1<sup>st</sup> day.

#### *Switching from previous neuroleptic therapy to Clozapine*

It is recommended that the other neuroleptics should be discontinued by tapering the dosage downwards over a period of approximately 1 week. Once the neuroleptic has been completely discontinued over 24 hours, clozapine treatment can be started as described above.

### - When to use it

Take your medicine at about the same time each day. Taking your tablets at the same time each day will have the best effect. It will also help you to remember when to take the tablets. It does not

matter if you take this medicine before or after food.

### - How long to use it

Continue taking Anzaril Tablet for as long as your doctor recommends.

### - If you forget to use it

- If you forget a dose, take it as soon as you remember it.
- However, if it is nearly time for the next dose, skip the missed dose.
- Do not take a double dose to make up for a forgotten dose.

### - If you use too much (overdose)

Symptoms: lethargy (tiredness), drowsiness, areflexia (absence of reflexes), coma, confusion, hallucinations (hearing voices or seeing things which are not there), agitation, delirium (confused thinking and reduced alertness), extrapyramidal symptoms (physical symptoms including tremor, slurred speech, anxiety, distress and paranoia), hyper-reflexia (overactive or overresponsive reflexes), convulsions, hypersalivation (excessive production of saliva), mydriasis (great dilatation of the pupil), blurred vision, thermolability, low blood pressure, collapse, rapid heartbeat, cardiac arrhythmias (irregular heartbeat), aspiration pneumonia (inflammatory of the lungs and large airway), dyspnoea (shortness of breath), respiratory depression or failure.

Close medication supervision is necessary for at least 5 days because of the possibility of delayed reactions.

Contact your doctor immediately or go to Emergency Department of your nearest hospital, if you think you or anyone else may have taken too much of this medicine. Do this even if there are no signs of discomfort or

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poisoning. You may need urgent medical attention.

## While you are using it

### - Things you must do

- Contact your doctor if your symptoms worsen or they do not improve.
- Your doctor will check and monitor your white blood count and absolute neutrophil count prior to treatment and weekly for 18 weeks then monthly thereafter.
- Contact your doctor if any kind of infection, fever, sore throat or other flu-like symptoms develop.
- Contact your doctor immediately if you suffer from tachycardia (abnormal rapid heart rate) that persists at rest, accompanied by arrhythmias (an irregular or abnormal rhythm of heart beat), shortness of breath or signs and symptoms of heart failure. Added as per Specific Labelling Requirement-Clozapine
- Contact your doctor or pharmacist if you have experienced constipation, abdominal tenderness, fever, bloating and/or bloody diarrhea. Your doctor will need to examine you.
- You must be monitored for worsening of glucose control if you are suffering from diabetes mellitus.
- You must discontinue the treatment if you develop jaundice.

### - Things you must not do

Do not give Anzaril Tablet to anyone else, even if their symptoms seem similar or they have same condition as you. Your doctor has prescribed Anzaril Tablet for you and your condition

### - Things to be careful of

This medicine will affect your ability to drive or use machines. If the tablets make you feel sick, dizzy or tired, or give you a headache, do not drive or use

machines and contact your doctor immediately.

Tell your doctor immediately if you become pregnant during Anzaril Tablet treatment. Newborn babies of mothers taking antipsychotic drugs during the third trimester of pregnancy may have increased risk of developing stiff limbs, trembling, agitation, muscle stiffness, muscle flaccidity, drowsiness, short and shallow breathing, and feeding disorders following delivery. In some cases, these symptoms may be self-limiting, in other cases, babies may require intensive care unit support or hospitalization.

Tell your doctor or pharmacist if you are breast-feeding. Clozapine, the active ingredient of Anzaril Tablet, may pass into your milk and affect your baby. You should not breast-feed during Anzaril Tablet treatment.

## Side effects

Like all medicines, Anzaril Tablet can have side effects, although not everybody gets them. Talk to your doctor or pharmacist if you experience:

- Blood and lymphatic system-development of granulocytopenia and agranulocytosis (an abnormally low number of granular white blood cells in the blood). Unexplained leucocytosis (an increase in the number of white cells in the blood) and/or eosinophilia (an increase in the number of eosinophils in the blood) may occur especially in the initial weeks of treatment. Very rarely, thrombocytopenia (deficiency of platelets in the blood).
- Central nervous system-fatigue, drowsiness and sedation. Dizziness or headache may also occur. Very rarely, clozapine may cause tardive dyskinesia, obsessive compulsive symptoms. Extrapyramidal effects may also occur.

- Autonomic nervous system-dry mouth, blurred vision, disturbances in sweating and temperature regulation.
- Cardiovascular system- rapid heart beat and postural low blood pressure. High blood pressure has occurred.
- Respiratory system- respiratory depression or arrest has occurred.
- Gastrointestinal system- abdominal pain, cramping, swollen abdomen, vomiting, constipation and failure to pass gas which may be signs and symptoms of bowel obstruction.
- Genitourinary system- urinary incontinence (involuntary loss of urine from the bladder) and urinary retention. Priapism (prolonged erection of the penis) has been reported.
- Miscellaneous- benign hyperthermia (body temperature rises to levels higher than normal), isolated report of skin reactions, severe hyperglycaemia (high blood sugar), increase in creatine phosphokinase (CPK) value, considerable weight gain, sudden and unexplained deaths.
- Increase in blood sugar level and/or symptoms of high blood sugar (e.g. increased thirst, increased hunger, and frequent urination).
- Unpleasant leg sensations and an intense urge to move the legs (restless legs syndrome).
- Trouble breathing during sleep (sleep apnoea).
- Difficulty or inability to pass urine (urinary retention).

If any of the side effects gets serious, or if you notice any side effects not listed in this leaflet, please tell your doctor or pharmacist.

Ask your doctor or pharmacist if you have any questions.

Visit your doctor or pharmacist immediately if you experience any side effects listed or not listed above after taking this medicine.

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You may report any side effects or adverse drug reactions directly to the National Centre for Adverse Drug Reaction Monitoring by visiting the website [npra.gov.my](http://npra.gov.my) [Consumers → Reporting Side Effects to Medicines (ConSERF) or Vaccines (AEFI)].

## Storage and Disposal of Anzaril Tablet

### - Storage

Do not use after the expiry date stated on the pack. The expiry date refers to the last day of that month.

Store below 30 °C in well closed container. Protect from light, heat and moisture.

**Keep out of the reach and sight of children.**

### - Disposal

Medicines should not be disposed of via wastewater or household waste. Ask your pharmacist how to dispose of medicines no longer required.

These measures will help to protect the environment.

## Product Description

### - What it looks like

Anzaril Tablet 25 mg:

A yellow, scored, flat of diameter 7 mm round tablet with “MPI” marked.

Anzaril Tablet 100 mg:

A yellow, flat-bevelled edge tablet with scored at one side, marked “MPI” on the other diameter of 10mm round tablet.

They are supplied in 10 x 10's blister pack.

### - Ingredients

Active ingredient:

Each tablet contain 25 mg or 100mg of the active substance, Clozapine

Inactive ingredients:

*Lactose*

*Corn starch*

*Sodium lauryl sulphate*

*Povidone K-30*

*Magnesium stearate*

*Colloidal silicone dioxide*

### - MAL Number

Anzaril Tablet 25mg

MAL06100580AZ

Anzaril Tablet 100mg

MAL06100579AZ

## Name and Address of Manufacturers

### Packed and released by:

GoodScience Sdn Bhd

No.7, Jalan KPK 4/3,

Kawasan Perindustrian Kundang,

Kundang Jaya, 48020 Rawang,

Selangor, Malaysia

### Manufactured by:

Malaysian Pharmaceutical

Industries Sdn. Bhd.

Plot 14, Lebuhraya Kampung Jawa,

11900 Bayan Lepas, Penang,

Malaysia.

## Product Registration Holder

GoodScience Sdn. Bhd.

No.7, Jalan KPK 4/3,

Kawasan Perindustrian Kundang,

Kundang Jaya 48020 Rawang,

Selangor,

Malaysia

## Date of revision

09/06/2025

## Serial number

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