Ativan (lorazepam)

Generic name: Lorazepam
Available strengths: 0.5 mg, 1 mg, 2 mg tablets; 2 mg/mL oral solution; 2 mg/mL, 4 mg/mL injection
Available in generic: Yes
Drug class: Benzodiazepine/anxiolytic; sedative-hypnotic

General Information

Ativan (lorazepam) is a benzodiazepine indicated for management of anxiety disorders or the short-term relief of symptoms of anxiety. The use of a drug for its approved indications is called its labeled use. In clinical practice, however, physicians often prescribe medications for unlabeled (“off-label”) uses when published clinical studies, case reports, or their own clinical experiences support the efficacy and safety of those treatments. Physicians may use Ativan outside its approved indications to treat social phobia, posttraumatic stress disorder, agitation in acute mania and psychosis, acute alcohol withdrawal, and other conditions. Like other benzodiazepines, Ativan is associated with dependence and abuse and is regulated as a controlled substance by state and federal laws.

Ativan’s effectiveness for treating anxiety may be explained by its pharmacological action in the brain at specific receptor sites. Receptors are specific sites on the nerve cell membrane that receive a signal from a neurotransmitter called the neurotransmitter. Once a neurotransmitter locks in on the receptor, the neurochemical signal is changed to an electrical or another chemical signal and travels down the neuron. The receptor sites in which benzodiazepines elicit their action are found in various regions of the brain, and the specific receptors are also known as benzodiazepine receptors. The coupled reaction of benzodiazepines to the receptors facilitates the inhibitory action of the neurotransmitter γ-aminobutyric acid (GABA) in that region of the brain. Benzodiazepines’ action on GABA receptors appears to produce their anxiolytic, sedative, and anticonvulsant actions. Ativan, for example, is an effective anxiolytic, hypnotic, and antiseizure medication.

Dosing Information

The usual starting dosage for Ativan is 0.5 mg three times a day, with an increase to a therapeutic dosage of 4–6 mg/day administered in divided doses. Depending on the severity of symptoms, the dosage may be increased to a maximum of 8 mg/day.
Common Side Effects

The most common side effects reported with Ativan are sedation and drowsiness, especially shortly after initiating therapy. Other frequent symptoms are impaired concentration and memory, feeling of dissociation (“spacey”), and impaired coordination.

Adverse Reactions and Precautions

Ativan affects alertness and coordination, and patients should exercise caution when driving or performing other tasks requiring alertness while taking this medication. Seniors may be more adversely affected, because it may affect their coordination and reflexes and lead to falls and injury. Taking Ativan with other central nervous system (CNS) depressants such as alcohol, narcotics, and barbiturates may compound these CNS effects.

Prolonged use of benzodiazepines can lead to dependence. When the medication is abruptly withdrawn, symptoms of withdrawal may occur. Withdrawal symptoms include headache, vomiting, impaired concentration, confusion, tremor, muscle cramps, and seizures.

Benzodiazepines are centrally acting depressants, and they can depress respiration. This is particularly problematic for patients with chronic obstructive pulmonary disease and emphysema. Patients with sleep apnea—a sleep disorder in which respiration is interrupted by long pauses during the sleep cycle—should not take Ativan or other benzodiazepines. The respiratory depressant effect of benzodiazepines may further suppress the respiratory drive in these patients and put them at risk for respiratory depression and death.

Benzodiazepines may induce paradoxical reactions in susceptible individuals. Instead of the expected depressant effects, the medication produces excitement, aggression, anger, uninhibited behavior, and rage in susceptible individuals. These reactions are more likely to occur in seniors, people with brain damage, and individuals with personality and impulse-control disorders.

Possible Drug Interactions

The drug interactions reported with Ativan are summarized in the table below.

<table>
<thead>
<tr>
<th>Central nervous system (CNS) depressants (e.g., alcohol, narcotics, barbiturates, hypnotics) and antihistamines</th>
<th>Combination of Ativan with another CNS depressant may impair coordination and breathing and increase sedation.</th>
</tr>
</thead>
<tbody>
<tr>
<td>Oral contraceptives</td>
<td>Clearance rate of Ativan may be increased when it is taken concurrently with oral contraceptives, which may decrease the effectiveness of Ativan.</td>
</tr>
<tr>
<td>Lanoxin (digoxin)</td>
<td>Ativan may increase the blood levels of Lanoxin, and toxicity may occur. Patients taking Ativan and Lanoxin should have their Lanoxin levels monitored closely.</td>
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</tbody>
</table>

Patients taking Ativan should not consume alcohol because the combination may increase sedation and drowsiness.
Use in Pregnancy and Breastfeeding: Pregnancy Category D

Benzodiazepines and their metabolites are known to cross the placenta and accumulate in the fetal circulation. They are associated with a risk of congenital malformations when used during pregnancy, causing cleft lip and heart deformities in the fetus. Benzodiazepines should be avoided during pregnancy, particularly in the first trimester. The use of benzodiazepines during pregnancy should be considered only when the need for the medication outweighs its risk and alternative therapies have failed.

Nursing mothers should not take Ativan, because it will pass into breast milk and be ingested by the baby. If stopping the drug is not an alternative, breastfeeding should not be started or should be discontinued.

Overdose

Overdoses from oral ingestion of benzodiazepines alone are generally not fatal. Most fatalities reported with benzodiazepines involve multiple medication ingestion, particularly the combination of a benzodiazepine with CNS depressants, including alcohol, narcotics, and barbiturates.

Mild symptoms of benzodiazepine overdose include drowsiness, confusion, somnolence, tiredness, loss of coordination, clumsiness in walking (ataxia), and slow reflexes. Benzodiazepine overdose, when these agents are taken alone, is rarely fatal. When multiple medications are implicated in benzodiazepine overdose, severe symptoms include difficulty breathing, slowed heart rate, low blood pressure, loss of coordination, and loss of consciousness leading to coma and, potentially, death.

Any suspected overdose should be treated as an emergency. The person should be taken to the emergency department for observation and treatment. The prescription bottle of medication (and any other medication suspected in the overdose) should be brought as well, because the information on the prescription label can be helpful to the treating physician in determining the number of pills ingested.

Special Considerations

- If you miss a dose, take it as soon as possible, but if it is close to the next scheduled dose, skip the missed dose and continue on your regular dosing schedule. Do not take double doses.
- Ativan may be taken with or without food.
- Ativan may cause sedation and drowsiness, especially during initiation of therapy, and may impair your alertness. Use caution when driving or performing tasks that require alertness. Avoid alcohol when taking Ativan, because alcohol may intensify these effects.
- Store the medication in its originally labeled, light-resistant container, away from heat and moisture. Heat and moisture may precipitate breakdown of your medication.
- Keep your medication out of reach of children.

If you have any questions about your medication, consult your physician or pharmacist.