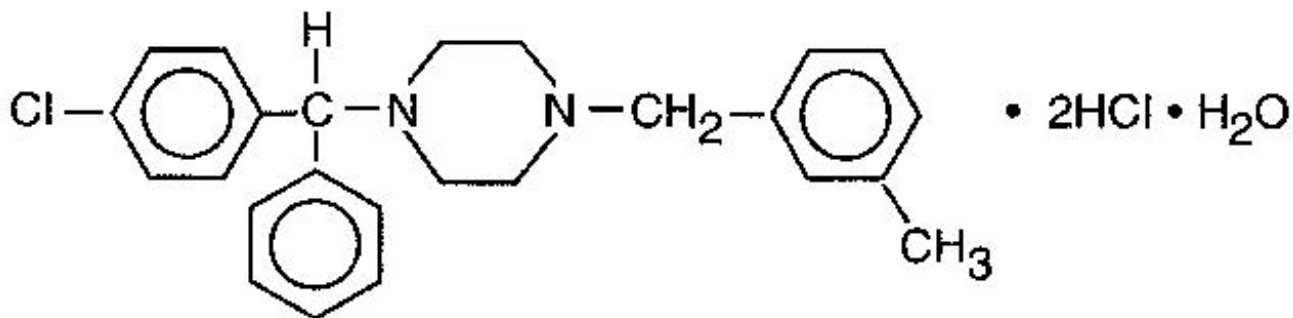


MECLIZINE HYDROCHLORIDE- meclizine hydrochloride tablet
Par Pharmaceutical, Inc.

DESCRIPTION

Meclizine hydrochloride, an oral antiemetic, is a white, slightly yellowish, crystalline powder which has a slight odor and is tasteless. It has the following structural formula:



C₂₅H₂₇ClN₂•2HCl•H₂O M.W. 481.89

The chemical name is 1-(*p*-chloro- α -phenylbenzyl)-4-(*m*-methyl-benzyl) - piperazine dihydrochloride monohydrate.

Meclizine Hydrochloride Tablets are available in 12.5 mg, and *25 mg strengths for oral administration.

*Contains FD&C Yellow #5 (see PRECAUTIONS).

Each tablet contains the following inactive ingredients: colloidal silicon dioxide, lactose, magnesium stearate, microcrystalline cellulose, sodium starch glycolate, starch, and stearic acid. In addition, the 12.5 mg tablet contains FD&C Blue #1; and the 25 mg tablet contains D&C Yellow #10 and FD&C Yellow #5.

CLINICAL PHARMACOLOGY

Meclizine hydrochloride is an antihistamine which shows marked protective activity against nebulized histamine and lethal doses of intravenously injected histamine in guinea pigs. It has a marked effect in blocking the vasodepressor response to histamine, but only a slight blocking action against acetylcholine. Its activity is relatively weak in inhibiting the spasmogenic action of histamine on isolated guinea pig ileum.

Pharmacokinetics

The available pharmacokinetic information for meclizine following oral administration has been summarized from published literature.

Absorption

Meclizine is absorbed after oral administration with maximum plasma concentrations reaching at a median T_{max} value of 3 hours post-dose (range: 1.5 to 6 hours) for the tablet dosage form.

Distribution

Drug distribution characteristics for meclizine in humans is unknown.

Metabolism

The metabolic fate of meclizine in humans is unknown. In an *in vitro* metabolic study using human hepatic microsome and recombinant CYP enzyme, CYP2D6 was found to be the dominant enzyme for metabolism of meclizine.

The genetic polymorphism of CYP2D6 that results in extensive-, poor-, intermediate- and ultrarapid metabolizer phenotypes could contribute to large inter-individual variability in meclizine exposure.

Elimination

Meclizine has a plasma elimination half-life of about 5 to 6 hours in humans.

INDICATIONS AND USAGE

INDICATIONS

Based on a review of this drug by the National Academy of Sciences – National Research Council and/or other information, FDA has classified the indications as follows:

Effective: Management of nausea and vomiting, and dizziness associated with motion sickness.

Possibly Effective: Management of vertigo associated with diseases affecting the vestibular system.

Final classification of the less than effective indications required further investigation.

CONTRAINDICATIONS

Meclizine hydrochloride is contraindicated in individuals who have shown a previous hypersensitivity to it.

WARNINGS

Since drowsiness may, on occasion, occur with use of this drug, patients should be warned of this possibility and cautioned against driving a car or operating dangerous machinery.

Patients should avoid alcoholic beverages while taking the drug. Due to its potential anticholinergic action, this drug should be used with caution in patients with asthma, glaucoma, or enlargement of the prostate gland.

Usage in Children

Clinical studies establishing safety and effectiveness in children have not been done; therefore, usage is not recommended in children under 12 years of age.

Usage in Pregnancy

Pregnancy Category B. Reproduction studies in rats have shown cleft palates at 25 to 50 times the human dose. Epidemiological studies in pregnant women, however, do not indicate that meclizine increases the risk of abnormalities when administered during pregnancy. Despite the animal findings, it would appear that the possibility of fetal harm is remote. Nevertheless, meclizine, or any other medication, should be used during pregnancy only if clearly necessary.

PRECAUTIONS

The Meclizine Hydrochloride Tablets, 25mg contain FD&C Yellow #5 (tartrazine) which may cause allergic-type reactions (including bronchial asthma) in certain susceptible individuals. Although the overall incidence of FD&C Yellow #5 (tartrazine) sensitivity in the general population is low, it is frequently seen in patients who also have aspirin hypersensitivity.

Nursing Mothers

It is not known whether this drug is excreted in human milk. Because many drugs are excreted in human milk, caution should be exercised when meclizine is administered to a nursing woman.

Hepatic Impairment

The effect of hepatic impairment on the pharmacokinetic of meclizine has not been evaluated. As meclizine undergoes metabolism, hepatic impairment may result in increased systemic exposure of the drug. Treatment with meclizine should be administered with caution in patients with hepatic impairment.

Renal Impairment

The effect of renal impairment on the pharmacokinetics of meclizine has not been evaluated. Due to a potential for drug/metabolite accumulation, meclizine should be administered with caution in patients with renal impairment and in the elderly as renal function generally declines with age.

Drug Interactions

There may be increased CNS depression when meclizine is administered concurrently with other CNS depressants, including alcohol, tranquilizers, and sedatives. (see WARNINGS).

Based on *in-vitro* evaluation, meclizine is metabolized by CYP2D6. Therefore there is a possibility for a drug interaction between meclizine and CYP2D6 inhibitors.

ADVERSE REACTIONS

Anaphylactoid reaction, drowsiness, dry mouth, headache, fatigue, vomiting and, on rare occasions, blurred vision have been reported.

DOSAGE AND ADMINISTRATION

Vertigo

For the control of vertigo associated with diseases affecting the vestibular system, the recommended dose is 25 to 100mg daily, in divided dosage, depending upon clinical response.

Motion Sickness: The initial dose of 25 to 50 mg meclizine hydrochloride, should be taken one hour prior to embarkation for protection against motion sickness. Thereafter, the dose may be repeated every 24 hours for the duration of the journey.

HOW SUPPLIED

Meclizine Hydrochloride Tablets, USP 12.5 mg - blue, oval tablets debossed with "034" on one side and "par" on the other side. Tablets may contain characteristic dye spots. They are supplied in bottles of 100 (NDC 49884-034-01) and 1000 (NDC 49884-034-10).

Meclizine Hydrochloride Tablets, USP 25 mg - yellow, oval tablets debossed with "035" on one side and "par" on the other side. They are supplied in bottles of 100 (NDC 49884-035-01) and 1000 (NDC 49884-035-10).

Dispense in tight, light-resistant containers as defined in the USP.

Store at controlled room temperature 15° to 30°C (59° to 86°F).

Manufactured by:

PAR PHARMACEUTICAL COMPANIES, INC.

Spring Valley, NY 10977

Revised: 11/2012

OS034-01-1-12

PRINCIPAL DISPLAY PANEL: 12.5 MG CONTAINER LABEL 100 TABLETS

<p>NDC 49884-034-01</p> <p>Meclizine Hydrochloride Tablets, USP</p> <p>12.5 mg</p> <p>Rx only</p> <p>100 Tablets</p> 	<p>Each tablet contains: Meclizine HCl, USP..... 12.5 mg</p> <p>USUAL DOSAGE: Read Accompanying Literature.</p> <p>KEEP THIS AND ALL DRUGS OUT OF REACH OF CHILDREN.</p> <p>Dispense in a tight, light-resistant container as defined in the USP.</p> <p>Store at controlled room temperature 15° to 30°C (59° to 86°F).</p>	<p>Control No.: Exp. Date:</p> <p>R11/08 LA034-01-1-14</p> <p>Par Pharmaceutical Companies, Inc. Spring Valley, NY 10977</p>	 <p>N 3 49884-034-016</p>
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PRINCIPAL DISPLAY PANEL: 25 MG CONTAINER LABEL 100 TABLETS

<p>NDC 49884-035-01</p> <p>Meclizine Hydrochloride Tablets, USP</p> <p>25 mg</p> <p>Rx only</p> <p>100 Tablets</p> 	<p>Each tablet contains: Meclizine HCl, USP..... 25 mg</p> <p>USUAL DOSAGE: Read Accompanying Literature.</p> <p>CAUTION: This product contains FD&C Yellow No. 5 (tartrazine) which may cause allergic type reactions in certain susceptible individuals.</p> <p>KEEP THIS AND ALL DRUGS OUT OF REACH OF CHILDREN.</p> <p>Dispense in a tight, light-resistant container as defined in the USP.</p> <p>Store at controlled room temperature 15°-30°C (59°-86°F).</p>	<p>Control No.: Exp. Date:</p> <p>R09/08 LA035-01-1-14</p> <p>Par Pharmaceutical Companies, Inc. Spring Valley, NY 10977</p>	 <p>N 3 49884-035-013</p>
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MECLIZINE HYDROCHLORIDE

meclizine hydrochloride tablet

Product Information

Product Type	HUMAN PRESCRIPTION DRUG	Item Code (Source)	NDC:49884-035
Route of Administration	ORAL		

Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
MECLIZINE HYDROCHLORIDE (UNII: HDP7W44CIO) (MECLIZINE - UNII:3L5TQ84570)	MECLIZINE HYDROCHLORIDE	25 mg

Inactive Ingredients

Ingredient Name	Strength
ANHYDROUS LACTOSE (UNII: 3SY5LH9PMK)	
CELLULOSE, MICROCRYSTALLINE (UNII: OP1R32D61U)	
SODIUM STARCH GLYCOLATE TYPE A POTATO (UNII: 5856J3G2A2)	
STARCH, CORN (UNII: O8232NY3SJ)	
STEARIC ACID (UNII: 4ELV7Z65AP)	
WATER (UNII: 059QF0KO0R)	
ISOPROPYL ALCOHOL (UNII: ND2M416302)	
D&C YELLOW NO. 10 (UNII: 35SW5USQ3G)	
MAGNESIUM STEARATE (UNII: 70097M6I30)	
SILICON DIOXIDE (UNII: ETJ7Z6XBU4)	
FD&C YELLOW NO. 5 (UNII: I753WB2F1M)	

Product Characteristics

Color	YELLOW	Score	no score
Shape	OVAL	Size	6mm
Flavor		Imprint Code	Par;035
Contains			

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:49884-035-01	100 in 1 BOTTLE; Type 0: Not a Combination Product	06/03/1981	11/30/2019
2	NDC:49884-035-10	1000 in 1 BOTTLE; Type 0: Not a Combination Product	06/03/1981	11/30/2019

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
ANDA	ANDA087128	06/03/1981	

MECLIZINE HYDROCHLORIDE

meclizine hydrochloride tablet

Product Information

Product Type	HUMAN PRESCRIPTION DRUG	Item Code (Source)	NDC:49884-034
Route of Administration	ORAL		

Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
MECLIZINE HYDROCHLORIDE (UNII: HDP7W44CIO) (MECLIZINE - UNII:3L5TQ84570)	MECLIZINE HYDROCHLORIDE	12.5 mg

Inactive Ingredients

Ingredient Name	Strength
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ANHYDROUS LACTOSE (UNII: 3SY5LH9PMK)	
CELLULOSE, MICROCRYSTALLINE (UNII: OP1R32D61U)	
SODIUM STARCH GLYCOLATE TYPE A POTATO (UNII: 5856J3G2A2)	
FD&C BLUE NO. 1 (UNII: H3R47K3TBD)	
WATER (UNII: 059QF0K00R)	
ISOPROPYL ALCOHOL (UNII: ND2M416302)	
STARCH, CORN (UNII: O8232NY3SJ)	
STEARIC ACID (UNII: 4ELV7Z65AP)	
MAGNESIUM STEARATE (UNII: 70097M6I30)	
SILICON DIOXIDE (UNII: ETJ7Z6XBU4)	

Product Characteristics

Color	BLUE	Score	no score
Shape	OVAL	Size	5mm
Flavor		Imprint Code	Par;034
Contains			

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:49884-034-01	100 in 1 BOTTLE; Type 0: Not a Combination Product	06/03/1981	02/28/2020
2	NDC:49884-034-10	1000 in 1 BOTTLE; Type 0: Not a Combination Product	06/03/1981	08/31/2019

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
ANDA	ANDA087127	06/03/1981	

Labeler - Par Pharmaceutical, Inc. (092733690)

Registrant - Par Pharmaceutical, Inc. (092733690)

Establishment

Name	Address	ID/FEI	Business Operations
Par Pharmaceutical Inc.		092733690	MANUFACTURE(49884-034, 49884-035)