LACTULOSE- lactulose solution
PAI Holdings, LLC dba PAI Pharma

Lactulose Solution USP

10 g/15 mL

CI08730222

R02/22

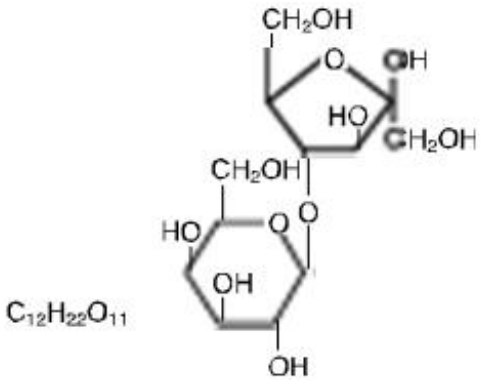
Rx ONLY

DESCRIPTION

Lactulose is a synthetic disaccharide in solution form for oral administration. Each 15 mL of Lactulose Solution contains: 10 g lactulose (and less than 1.6 g galactose, less than 1.2 g lactose, and 1.2 g or less of other sugars).

Lactulose is a colonic acidifier which promotes laxation.

The chemical name for lactulose is 4-0-ß-D-galactopyranosyl-D-fructofuranose. It has the following structural formula:



The molecular weight is 342.30. It is freely soluble in water.

CLINICAL PHARMACOLOGY

Lactulose is poorly absorbed from the gastrointestinal tract, and no enzyme capable of hydrolysis of this disaccharide is present in human gastrointestinal tissue. As a result, oral doses of lactulose solution reach the colon virtually unchanged. In the colon, lactulose is broken down primarily to lactic acid, and also to small amounts of formic and acetic acids, by the action of colonic bacteria, which results in an increase in osmotic pressure and slight acidification of the colonic contents. This in turn causes an increase in stool water content and softens the stool.

Since lactulose does not exert its effect until it reaches the colon, and since transit time through the colon may be slow, 24 to 48 hours may be required to produce the desired bowel movement.

Lactulose solution given orally to man and experimental animals resulted in only small amounts reaching the blood.

Urinary excretion has been determined to be 3% or less and is essentially complete within 24 hours.

INDICATIONS AND USAGE

For the treatment of constipation. In patients with a history of chronic constipation, lactulose solution therapy increases the number of bowel movements per day and the number of days on which bowel movements occur.

CONTRAINDICATIONS

Since lactulose solution contains galactose (less than 1.6 g/15 mL), it is contraindicated in patients who require a low galactose diet.

WARNINGS

A theoretical hazard may exist for patients being treated with lactulose solution who may be required to undergo electrocautery procedures during proctoscopy or colonoscopy. Accumulation of H_2 gas in significant concentration in the presence of an electrical spark may result in an explosive reaction. Although this complication has not been reported with lactulose, patients on lactulose therapy undergoing such procedures should have a thorough bowel cleansing with a non-fermentable solution. Insufflation of CO_2 as an additional safeguard may be pursued but is considered to be a redundant measure.

PRECAUTIONS

General

Since lactulose solution contains galactose (less than1.6 g/15 mL) and lactose (less than 1.2 g/15 mL), it should be used with caution in diabetics.

Information for Patients

In the event that an unusual diarrheal condition occurs, contact your physician.

Laboratory Tests

Elderly, debilitated patients who receive lactulose solution for more than six months should have serum electrolytes (potassium, chloride, carbon dioxide) measured periodically.

Drug Interactions

Results of preliminary studies in humans and rats suggest that nonabsorbable antacids given concurrently with lactulose may inhibit the desired lactulose-induced drop in colonic pH. Therefore, a possible lack of desired effect of treatment should be taken into consideration before such drugs are given concomitantly with lactulose solution.

Carcinogenesis, Mutagenesis, Impairment of Fertility

There are no known human data on long-term potential for carcinogenicity, mutagenicity, or impairment of fertility.

There are no known animal data on long-term potential for mutagenicity.

Administration of lactulose solution in the diet of mice for 18 months in concentrations of 3 and 10 percent (v/w) did not produce any evidence of carcinogenicity.

In studies in mice, rats, and rabbits, doses of lactulose solution up to 6 or 12 mL/kg/day produced no deleterious effects in breeding, conception, or parturition.

Pregnancy - Teratogenic Effects

Reproduction studies have been performed in mice, rats, and rabbits at doses up to 3 or 6 times the usual human oral dose and have revealed no evidence of impaired fertility or harm to the fetus due to lactulose. There are, however, 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 needed.

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 lactulose solution is administered to a nursing woman.

Pediatric Use

Safety and effectiveness in pediatric patients have not been established.

ADVERSE REACTIONS

Precise frequency data are not available.

Initial dosing may produce flatulence and intestinal cramps, which are usually transient. Excessive dosage can lead to diarrhea with potential complications such as loss of fluids, hypokalemia, and hypernatremia.

Nausea and vomiting have been reported.

To report SUSPECTED ADVERSE REACTIONS, contact Pharmaceutical Associates, Inc. at 1-800-845-8210 or FDA at 1-800-FDA-1088 or www.fda.gov/medwatch.

OVERDOSAGE

Signs and Symptoms

There have been no reports of accidental overdosage. In the event of overdosage, it is expected that diarrhea and abdominal cramps would be the major symptoms. Medication should be terminated.

Oral LD_{50}

The acute oral LD_{50} of the drug is 48.8 mL/kg in mice and greater than 30 mL/kg in rats.

Dialysis

Dialysis data are not available for lactulose. Its molecular similarity to sucrose, however, would suggest that it should be dialyzable.

DOSAGE AND ADMINISTRATION

The usual dose is 1 to 2 tablespoonfuls (15 to 30 mL, containing 10 g to 20 g of lactulose) daily. The dose may be increased to 60 mL daily if necessary. Twenty-four to 48 hours may be required to produce a normal bowel movement.

Note: Some patients have found that lactulose solution may be more acceptable when mixed with fruit juice, water or milk.

HOW SUPPLIED

Lactulose Solution, USP 10 g/15 mL is supplied as:

NDC 0121-0873-08: 8 fl oz (237 mL) bottles

NDC 0121-0873-16: 16 fl oz (473 mL) bottles

NDC 0121-0873-32: 32 fl oz (946 mL) bottles

NDC 0121-0873-15: 15 mL unit dose cup. Case contains 40 unit dose cups of 15 mL (NDC 0121-0873-40), packaged in 4 trays of 10 unit-dose cups each and 96 unit dose cups of 15 mL (NDC 0121-0873-06) packaged in a carton of 16 unit dose cups each.

NDC 0121-1746-30: 30 mL unit dose cup. Case contains 40 unit dose cups of 30 mL (NDC 0121-1746-40), packaged in 4 trays of 10 unit-dose cups each, 100 unit dose cups of 30 mL (NDC 0121-1746-00), packaged in 10 trays of 10 unit-dose cups each and 96 unit dose cups of 30 mL (NDC 0121-1746-06) packaged in a carton of 16 unit dose cups each.

Store at controlled room temperature, 20°to 25°C (68° to 77°F). [See USP] Do not freeze.

Under recommended storage conditions, a normal darkening of color may occur. Such darkening is characteristic of sugar solutions and does not affect therapeutic action. Prolonged exposure to temperatures above 30°C (86°F) or to direct light may cause extreme darkening and turbidity which may be pharmaceutically objectionable. If this condition develops, do not use.

Prolonged exposure to freezing temperatures may cause change to a semisolid, too viscous to pour. Viscosity will return to normal upon warming to room temperature.

Dispense in original container or in a tight, light-resistant container with a child-resistant closure.

To the Pharmacist: When ordering this product, include the NDC number in the description.

MANUFACTURED BY

Pharmaceutical Associates, Inc.
Greenville, SC 29605

R02/2022

PRINCIPAL DISPLAY PANEL - 237 mL Bottle Label

NDC 0121-0873-08

Lactulose Solution USP

10 g/15 mL

Each 15 mL contains: 10 g lactulose (and less than 1.6 g galactose, less than 1.2 g lactose, and 1.2 g or less of other sugars).

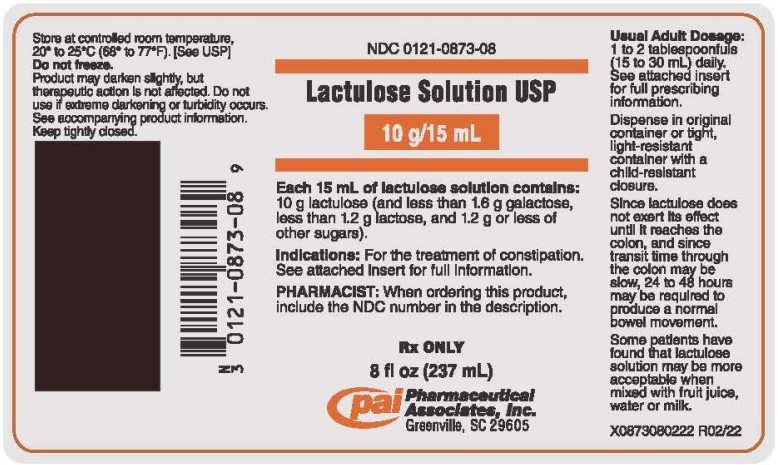
Dispense in original container or tight, light-resistant container with a child-resistant closure.

To the Pharmacist: When ordering this product, include the product number (or NDC) in the description.

Rx ONLY

8 fl oz (237 mL)

Pharmaceutical Associates, Inc.
Greenville, SC 29605



PRINCIPAL DISPLAY PANEL - 473 mL Bottle Label

NDC 0121-0873-16

Lactulose Solution USP

10 g/15 mL

Each 15 mL contains: 10 g lactulose (and less than 1.6 g galactose, less than 1.2 g lactose, and 1.2 g or less of other sugars).

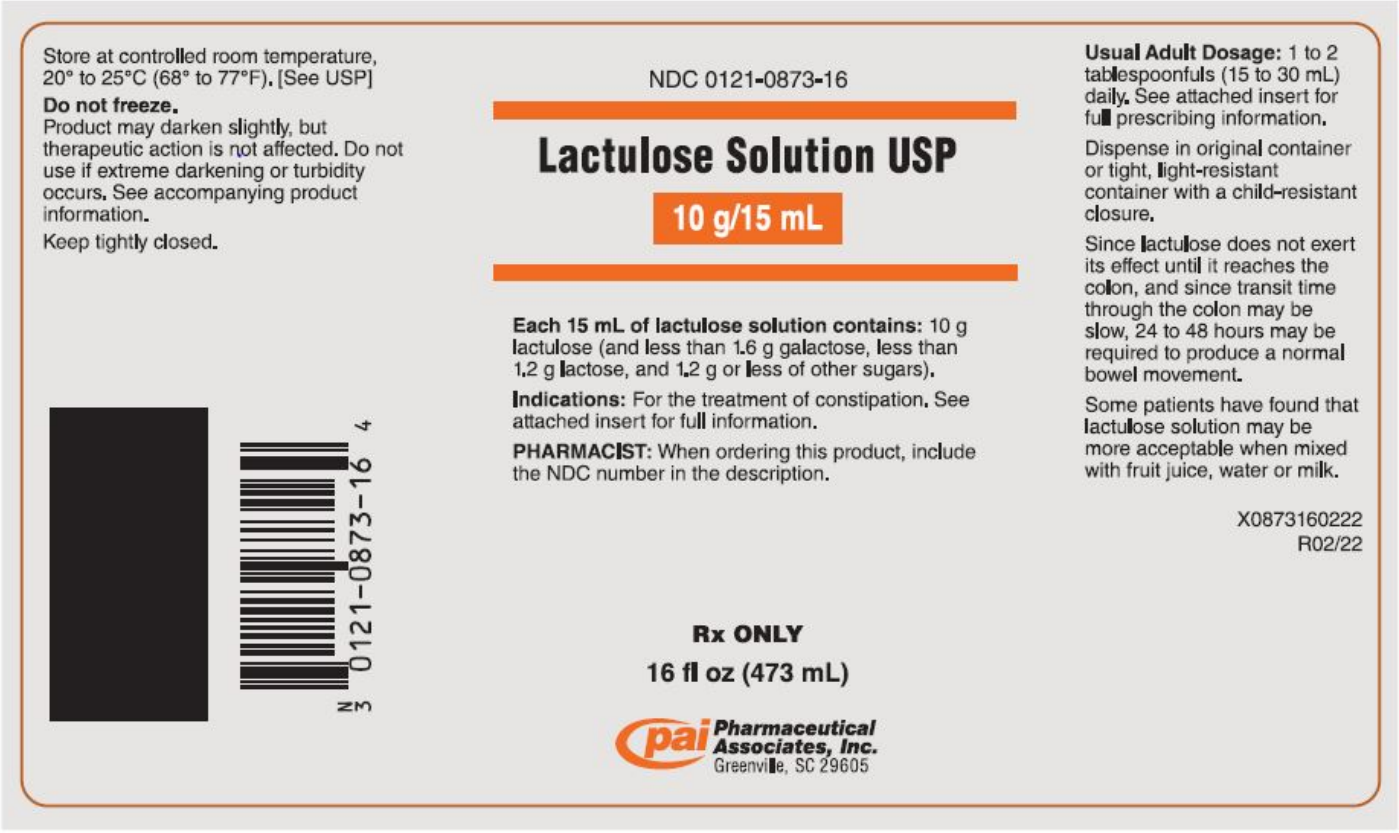
Dispense in original container or tight, light-resistant container with a child-resistant closure.

To the Pharmacist: When ordering this product, include the product number (or NDC) in the description.

Rx ONLY

16 fl oz (473 mL)

Pharmaceutical Associates, Inc.
Greenville, SC 29605



PRINCIPAL DISPLAY PANEL - 15 mL Cup Lid - NDC 0121-0873-15

UNIT DOSE

Delivers 15 mL
NDC 0121-0873-15

LACTULOSE
***SOLUTION USP* 10 g/15 mL**

Indication: For the treatment of constipation. See Insert.

Package Not Child-Resistant

Rx ONLY
PHARMACEUTICAL ASSOCIATES, INC.
GREENVILLE, SC 29605

PRINCIPAL DISPLAY PANEL - 30 mL Cup Lid - NDC 0121-1746-30

UNIT DOSE
Delivers **30 mL**
NDC 0121-1746-30

LACTULOSE
SOLUTION USP 20 g/30 mL

Indication: For the treatment of constipation. See Insert.

Package Not Child-Resistant

Rx ONLY

PHARMACEUTICAL ASSOCIATES, INC.
GREENVILLE, SC 29605

LACTULOSE

lactulose solution

Product Information			
Product Type	HUMAN PRESCRIPTION DRUG	**Item Code (Source)**	NDC:0121-0873
Route of Administration	ORAL		

Active Ingredient/Active Moiety		
Ingredient Name	**Basis of Strength**	**Strength**
LACTULOSE (UNII: 9U7D5QH5AE) (LACTULOSE - UNII:9U7D5QH5AE)	LACTULOSE	10 g in 15 mL

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0121-0873-08	237 mL in 1 BOTTLE; Type 0: Not a Combination Product	12/03/2019	
2	NDC:0121-0873-16	473 mL in 1 BOTTLE; Type 0: Not a Combination Product	12/03/2019	
3	NDC:0121-0873-32	946 mL in 1 BOTTLE; Type 0: Not a Combination Product	12/03/2019	
4	NDC:0121-	4 in 1 CASE	12/03/2019	

4	0873-40	4 in 1 CASE	12/03/2019	
4		10 in 1 TRAY		
4	NDC:0121-0873-15	15 mL in 1 CUP, UNIT-DOSE; Type 0: Not a Combination Product		
5	NDC:0121-0873-06	6 in 1 CASE	12/03/2019	
5		16 in 1 CARTON		
5		15 mL in 1 CUP, UNIT-DOSE; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
ANDA	ANDA074623	12/03/2019	

LACTULOSE

lactulose solution

Product Information

Product Type	HUMAN PRESCRIPTION DRUG	**Item Code (Source)**	NDC:0121-1746
Route of Administration	ORAL		

Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
LACTULOSE (UNII: 9U7D5QH5AE) (LACTULOSE - UNII:9U7D5QH5AE)	LACTULOSE	20 g in 30 mL

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0121-1746-40	4 in 1 CASE	12/03/2019	
1		10 in 1 TRAY		
1	NDC:0121-1746-30	30 mL in 1 CUP, UNIT-DOSE; Type 0: Not a Combination Product		
2	NDC:0121-1746-00	10 in 1 CASE	12/03/2019	
2		10 in 1 TRAY		
2		30 mL in 1 CUP, UNIT-DOSE; Type 0: Not a Combination Product		
3	NDC:0121-1746-06	6 in 1 CASE	12/03/2019	
3		16 in 1 CARTON		
3		30 mL in 1 CUP, UNIT-DOSE; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
ANDA	ANDA074623	12/03/2019	

Labeler - PAI Holdings, LLC dba PAI Pharma (044940096)

Establishment

Name	Address	ID/FEI	Business Operations
PAI Holdings, LLC dba Pharmaceutical Associates, Inc. and dba PAI Pharma		097630693	manufacture(0121-0873, 0121-1746)

Revised: 9/2024 PAI Holdings, LLC dba PAI Pharma