

METRONIDAZOLE- metronidazole gel
Fougera Pharmaceuticals Inc.

METRONIDAZOLE GEL USP 0.75%

FOR TOPICAL USE ONLY.
NOT FOR OPHTHALMIC USE.
Rx only

INDICATIONS AND USAGE

Metronidazole gel is indicated for topical application in the treatment of inflammatory papules and pustules of rosacea.

DOSAGE AND ADMINISTRATION

Apply and rub in a thin film of metronidazole gel twice daily, morning and evening, to entire affected areas after washing. Areas to be treated should be cleansed before application of metronidazole gel. Patients may use cosmetics after application of metronidazole gel.

CONTRAINDICATIONS

Metronidazole gel is contraindicated in individuals with a history of hypersensitivity to metronidazole, parabens, or other ingredients of the formulation.

PRECAUTIONS:

General: Metronidazole gel has been reported to cause tearing of the eyes. Therefore, contact with the eyes should be avoided. If a reaction suggesting local irritation occurs, patients should be directed to use the medication less frequently or discontinue use. Metronidazole is a nitroimidazole and should be used with care in patients with evidence of, or history of blood dyscrasia.

Information for patients: This medication is to be used as directed by the physician. It is for external use only. Avoid contact with the eyes.

Drug interactions: Oral metronidazole has been reported to potentiate the anticoagulant effect of coumarin and warfarin resulting in a prolongation of prothrombin time. The effect of topical metronidazole on prothrombin time is not known.

Carcinogenesis, mutagenesis, impairment of fertility:

Metronidazole has shown evidence of carcinogenic activity in a number of studies involving chronic, oral administration in mice and rats but not in studies involving hamsters.

Metronidazole has shown evidence of mutagenic activity in several *in vitro* bacterial assay systems. In addition, a dose-response increase in the frequency of micronuclei was observed in mice after intraperitoneal injections and an increase in chromosome aberrations have been reported in patients with Crohn's disease who were treated with 200-1200 mg/day of metronidazole for 1 to 24 months. However, no excess chromosomal aberrations in circulating human lymphocytes have been observed in patients treated for 8 months.

Pregnancy: Teratogenic effects: Pregnancy category B:

There has been no experience to date with the use of metronidazole gel in pregnant patients. Metronidazole crosses the placental barrier and enters the fetal circulation rapidly. No fetotoxicity was

observed after oral metronidazole in rats or mice. However, because animal reproduction studies are not always predictive of human response and since oral metronidazole has been shown to be a carcinogen in some rodents, this drug should be used during pregnancy only if clearly needed.

Nursing mothers: After oral administration, metronidazole is secreted in breast milk in concentrations similar to those found in the plasma. Even though metronidazole gel blood levels are significantly lower than those achieved after oral metronidazole, a decision should be made whether to discontinue nursing or to discontinue the drug, taking into account the importance of the drug to the mother.

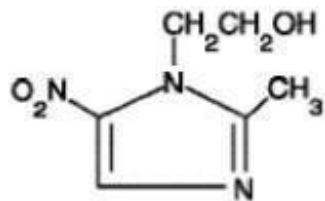
Pediatric use: Safety and effectiveness in pediatric patients have not been established.

ADVERSE REACTIONS

The following adverse experiences have been reported with the topical use of metronidazole: burning, skin irritation, dryness, transient redness, metallic taste, tingling or numbness of extremities and nausea.

DESCRIPTION

Metronidazole Gel USP 0.75% contains metronidazole, USP, at a concentration of 7.5 mg per gram (0.75%) in a gel consisting of purified water, methylparaben, propylparaben, propylene glycol, carbomer 940, sodium hydroxide, and edetate disodium. Metronidazole is classified therapeutically as an antiprotozoal and anti-bacterial agent. Chemically, metronidazole is named 2-methyl-5-nitroimidazole-1-ethanol and has the following structure:



CLINICAL PHARMACOLOGY

Bioavailability studies on the topical administration of 1 gram of metronidazole gel (7.5 mg of metronidazole) to the face of 10 rosacea patients showed a maximum serum concentration of 66 nanograms per milliliter in one patient. This concentration is approximately 100 times less than concentrations afforded by a single 250 mg oral tablet. The serum metronidazole concentrations were below the detectable limits of the assay at the majority of time points in all patients. Three of the patients had no detectable serum concentrations of metronidazole at any time point. The mean dose of gel applied during clinical studies was 600 mg which represents 4.5 mg of metronidazole per application. Therefore, under normal usage levels, the formulation affords minimal serum concentrations of metronidazole. The mechanisms by which metronidazole gel acts in the treatment of rosacea are unknown, but appear to include an anti-inflammatory effect.

HOW SUPPLIED

Metronidazole Gel USP 0.75% is supplied as follows:

45 gram tube NDC 0168-0275-45

Store at 20° to 25°C (68° to 77°F)

[see USP Controlled Room Temperature].

E. FOUGERA & CO.

A division of Fougere Pharmaceuticals Inc.
Melville, New York 11747

I2275D
R07/13
#87

PACKAGE LABEL – PRINCIPAL DISPLAY PANEL – 45 GRAM TUBE

NDC 0168-0275-45

Fougera®

METRONIDAZOLE
GEL USP 0.75%

Rx only

FOR TOPICAL USE ONLY.
NOT FOR OPHTHALMIC USE.
WARNING: KEEP OUT OF
REACH OF CHILDREN.

NET WT 45 grams

<p>NDC 0168-0275-45</p> <p>Metronidazole Gel USP, 0.75%</p> <p>FOR TOPICAL USE ONLY. NOT FOR OPHTHALMIC USE.</p> <p>fougera®</p> <p>Usual Dosage: Apply a thin film to entire affected areas after washing. Use morning and evening or as directed by physician. Avoid application close to the eyes.</p> <p>Store at 20° to 25°C (68° to 77°F) [see USP Controlled Room Temperature].</p> <p>KEEP OUT OF THE REACH OF CHILDREN.</p> <p>TO OPEN: Use cap to puncture seal.</p> <p>IMPORTANT: Do not use if seal has been punctured or is not visible.</p>	<p>R only</p> <p>Each gram contains: 7.5 mg of metronidazole USP as the active ingredient with 0.8 mg of methylparaben and 0.2 mg of propylparaben as preservatives in a gel consisting of purified water, propylene glycol, carbomer 980, sodium hydroxide and edetate disodium.</p> <p>NET WT 45 grams</p> <p>See crimp of tube for lot number and expiration date.</p> <p>E. FOUGERA & CO. A division of Fougera Pharmaceuticals Inc. Melville, New York 11747</p> <p>H5236F R12/19</p>  <p>N 3 0168-0275-45 1</p>	
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PACKAGE LABEL – PRINCIPAL DISPLAY PANEL – 45 GRAM CARTON

NDC 0168-0275-45

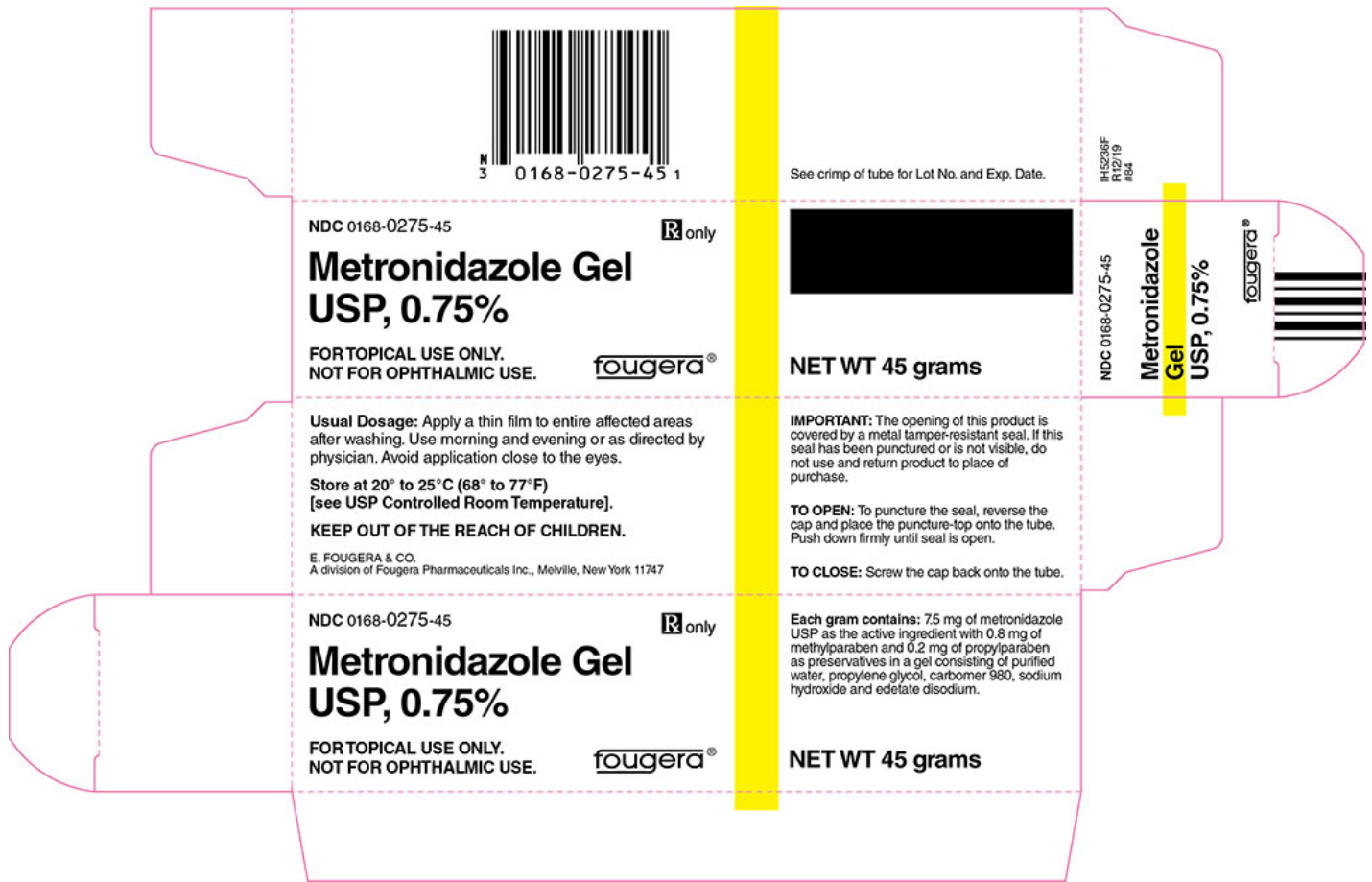
Fougera®

**METRONIDAZOLE
GEL USP 0.75%**

Rx only

**FOR TOPICAL USE ONLY.
NOT FOR OPHTHALMIC USE.**

NET WT 45 grams



METRONIDAZOLE			
metronidazole gel			
Product Information			
Product Type	HUMAN PRESCRIPTION DRUG	Item Code (Source)	NDC:0168-0275
Route of Administration	TOPICAL		
Active Ingredient/Active Moiety			
Ingredient Name		Basis of Strength	Strength
METRONIDAZOLE (UNII: 140QMO216E) (METRONIDAZOLE - UNII:140QMO216E)		METRONIDAZOLE	7.5 mg in 1 g
Inactive Ingredients			
Ingredient Name			Strength

WATER (UNII: 059QF0KO0R)	
METHYL PARABEN (UNII: A2I8C7HI9T)	
PROPYL PARABEN (UNII: Z8IX2SC1OH)	
PROPYLENE GLYCOL (UNII: 6DC9Q167V3)	
SODIUM HYDROXIDE (UNII: 55X04QC32I)	
CARBOMER HOMO POLYMER TYPE C (ALLYL PENTAERYTHRITOL CROSSLINKED) (UNII: 4Q93RCW27E)	
EDETATE DISODIUM (UNII: 7FLD91C86K)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0168-0275-45	1 in 1 CARTON	06/06/2006	
1		45 g in 1 TUBE; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
ANDA	ANDA077018	06/06/2006	

Labeler - Fougera Pharmaceuticals Inc. (043838424)

Revised: 3/2020

Fougera Pharmaceuticals Inc.