

**METRONIDAZOLE - metronidazole gel**  
**Cosette Pharmaceuticals, Inc.**

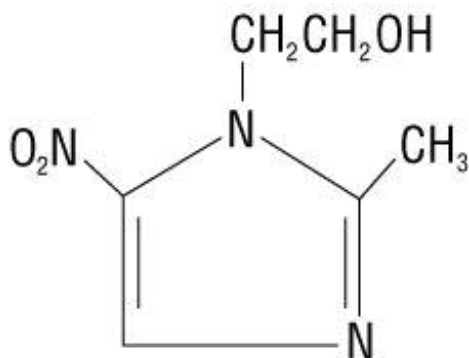
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**Metronidazole Gel USP, 0.75%**

**FOR TOPICAL USE ONLY (NOT FOR OPHTHALMIC USE)**

**Rx Only**

**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 carbomer homopolymer NF, edetate disodium, methylparaben, propylene glycol, propylparaben, purified water, and sodium hydroxide. Metronidazole is classified therapeutically as an antiprotozoal and antibacterial agent. Chemically, metronidazole is named 2-methyl-5-nitro-1*H*-imidazole-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.

**INDICATIONS AND USAGE:**

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

**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.

**To report SUSPECTED ADVERSE REACTIONS, contact Cosette Pharmaceuticals, Inc. at 1-800-922-1038 or FDA at 1-800-FDA-1088 or [www.fda.gov/medwatch](http://www.fda.gov/medwatch).**

## 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.

## HOW SUPPLIED:

Metronidazole Gel USP, 0.75% is supplied in a 45 g tube

NDC 0713-0637-37

**Storage conditions:** Store at 20°-25°C (68°-77°F) [see USP Controlled Room Temperature].

**Manufactured for:**

**Cosette Pharmaceuticals, Inc.**

111 Coolidge Street,  
South Plainfield, NJ 07080

8-0637BFCPLNC1

Iss. 08/2019

GW 7312

**PRINCIPAL DISPLAY PANEL**

**NDC 0713-0637-37**

**Metronidazole Gel USP 0.75%**

**45 g**

**Rx only**

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

**Cosette Pharmaceuticals, Inc.**

Bend Line

The image shows a rectangular product label with a teal and white color scheme. On the left, there is a teal vertical bar containing a white logo with a circle and a square, and the text '45 g' below it. The main white area contains the product name 'Metronidazole Gel USP' in large black and teal letters, with '0.75%' in white on a teal background below it. To the right, 'NDC 0713-0637-37' and 'Rx only' are printed. At the bottom, there is a barcode with 'N 3' to its left and '0713-0637-37 4' below it. A small black square is located in the bottom right corner of the label area.

**NDC 0713-0637-37**

**Rx only**

**45 g**

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

Store at 20°-25°C (68°-77°F) [see USP Controlled Room Temperature].  
**KEEP OUT OF REACH OF CHILDREN.**

**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.

**Each gram contains:** 7.5 mg of metronidazole as the active ingredient with 0.8 mg of methylparaben and 0.2 mg of propylparaben as preservatives in a gel consisting of carbomer homopolymer NF, edetate disodium, propylene glycol, purified water, and sodium hydroxide.

See crimped end for lot number and expiration date.

**Manufactured for:**  
**Cosette Pharmaceuticals, Inc.**  
111 Coolidge Street  
South Plainfield, NJ 07080  
GW 1105 Iss. 08/2019 11-637BF37CPLNC1

N 3 0713-0637-37 4



## METRONIDAZOLE

metronidazole gel

### Product Information

<b>Product Type</b>	HUMAN PRESCRIPTION DRUG	<b>Item Code (Source)</b>	NDC:0713-0637
<b>Route of Administration</b>	TOPICAL		

### Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
<b>Metronidazole</b> (UNII: 140QMO216E) (Metronidazole - UNII:140QMO216E)	Metronidazole	7.5 mg in 1 g

### Inactive Ingredients

Ingredient Name	Strength
<b>Methylparaben</b> (UNII: A2I8C7HI9T)	
<b>Propylparaben</b> (UNII: Z8IX2SC1OH)	
<b>Carbomer Homopolymer Type C</b> (UNII: 4Q93RCW27E)	
<b>Edetate Disodium</b> (UNII: 7FLD91C86K)	
<b>Propylene Glycol</b> (UNII: 6DC9Q167V3)	
<b>Water</b> (UNII: 059QF0KO0R)	

Sodium Hydroxide (UNII: 55X04QC32I)

### Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0713-0637-37	1 in 1 CARTON	04/01/2011	
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	ANDA078178	04/01/2011	

**Labeler** - Cosette Pharmaceuticals, Inc. (116918230)

**Registrant** - Cosette Pharmaceuticals, Inc. (116918230)

### Establishment

Name	Address	ID/FEI	Business Operations
Cosette Pharmaceuticals, Inc.		116918230	analysis(0713-0637) , label(0713-0637) , manufacture(0713-0637) , pack(0713-0637)

Revised: 11/2020

Cosette Pharmaceuticals, Inc.