

NYSTATIN AND TRIAMCINOLONE ACETONIDE - nystatin and triamcinolone acetonide cream

Actavis Pharma, Inc.

NYSTATIN AND TRIAMCINOLONE ACETONIDE CREAM, USP

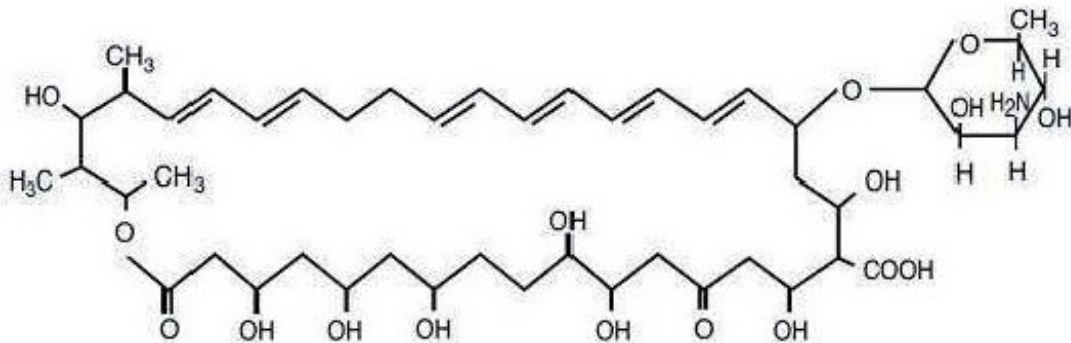
Rx Only

FOR EXTERNAL USE ONLY. NOT FOR OPHTHALMIC USE.

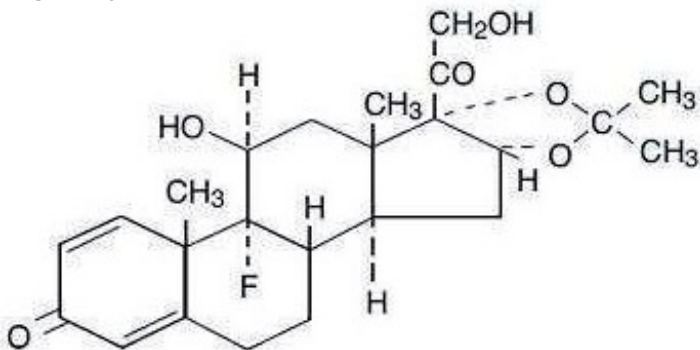
DESCRIPTION

Nystatin and Triamcinolone Acetonide Cream, USP for dermatologic use contains the antifungal agent nystatin and the synthetic corticosteroid triamcinolone acetonide.

Nystatin is a polyene antimycotic obtained from *Streptomyces noursei*. It is a yellow to light tan powder with a cereal-like odor, very slightly soluble in water, and slightly to sparingly soluble in alcohol. It has a structural formula of $C_{47}H_{75}NO_{17}$ and a molecular weight of 926.13.



Triamcinolone acetonide is designated chemically as 9-fluoro-11 β , 16 α , 17, 21-tetrahydroxypregna-1,4-diene-3,20-dione cyclic 16, 17-acetal with acetone. The white to cream crystalline powder has a slight odor, is practically insoluble in water, and very soluble in alcohol. It has a structural formula of $C_{24}H_{31}FO_6$ and a molecular weight of 434.50.



Nystatin and Triamcinolone Acetonide Cream, USP is a soft, smooth cream having a light yellow color. Each gram provides 100,000 Nystatin units and 1 mg Triamcinolone Acetonide in an aqueous perfumed vanishing cream base containing aluminum hydroxide rehydragel, barcroft aluminum hydroxide gel, cetearyl alcohol (and) cetareth-20, glyceryl monostearate, polyethylene glycol monostearate, propylene glycol, simethicone emulsion, sorbic acid, sorbitol solution, titanium dioxide, and white petrolatum.

CLINICAL PHARMACOLOGY

Nystatin

Nystatin exerts its antifungal activity against a variety of pathogenic and nonpathogenic yeasts and fungi by binding to sterols in the cell membrane. The binding process renders the cell membrane incapable of functioning as a selective barrier. Nystatin provides specific anticandidal activity to *Candida* (Monilia) *albicans* and other *Candida* species, but it is not active against bacteria, protozoa, trichomonads, or viruses.

Nystatin is not absorbed from intact skin or mucous membranes.

Triamcinolone Acetonide

Triamcinolone acetonide is primarily effective because of its anti-inflammatory, anti-pruritic and vasoconstrictive actions, characteristic of the topical corticosteroid class of drugs. The pharmacological effects of the topical corticosteroids are well known; however, the mechanisms of their dermatologic actions are unclear. Various laboratory methods, including vasoconstrictor assays, are used to compare and predict potencies and/or clinical efficacies of the topical corticosteroids. There is some evidence to suggest that a recognizable correlation exists between vasoconstrictor potency and therapeutic efficacy in man.

Pharmacokinetics

The extent of percutaneous absorption of topical corticosteroids is determined by many factors including the vehicle, the integrity of the epidermal barrier, and the use of occlusive dressings (see **DOSAGE AND ADMINISTRATION**).

Topical corticosteroids can be absorbed from normal intact skin. Inflammation and/or other disease processes in the skin increase percutaneous absorption. Occlusive dressings substantially increase the percutaneous absorption of topical corticosteroids (see **DOSAGE AND ADMINISTRATION**).

Once absorbed through the skin, topical corticosteroids are handled through pharmacokinetic pathways similar to systemically administered corticosteroids. Corticosteroids are bound to plasma proteins in varying degrees. Corticosteroids are metabolized primarily in the liver and are then excreted by the kidneys. Some of the topical corticosteroids and their metabolites are also excreted into the bile.

Nystatin and Triamcinolone Acetonide

During clinical studies of mild to severe manifestations of cutaneous candidiasis, patients treated with nystatin and triamcinolone acetonide showed a faster and more pronounced clearing of erythema and pruritis than patients treated with nystatin or triamcinolone acetonide alone.

INDICATIONS AND USAGE

Nystatin and Triamcinolone Acetonide Cream, USP is indicated for the treatment of cutaneous candidiasis; it has been demonstrated that the nystatin-steroid combination provides greater benefit than the nystatin component alone during the first few days of treatment.

CONTRAINDICATIONS

This preparation is contraindicated in those patients with a history of hypersensitivity to any of its components.

PRECAUTIONS

General

Systemic absorption of topical corticosteroids has produced reversible hypothalamic-pituitary-adrenal (HPA) axis suppression, manifestations of Cushing's syndrome, hyperglycemia, and glucosuria in some patients. Conditions that augment systemic absorption include application of the more potent steroids, use over large surface areas, prolonged use, and the addition of occlusive dressings (see **DOSAGE AND ADMINISTRATION**).

Therefore, patients receiving a large dose of any potent topical steroid applied to a large surface area should be evaluated periodically for evidence of HPA axis suppression by using the urinary free cortisol and ACTH stimulation tests, and for impairment of internal homeostasis. If HPA axis suppression or elevation of the body temperature occurs, an attempt should be made to withdraw the drug, to reduce the frequency of application, or substitute a less potent steroid.

Recovery of HPA axis function and thermal homeostasis are generally prompt and complete upon discontinuation of the drug. Infrequently, signs and symptoms of steroid withdrawal may occur, requiring supplemental systemic corticosteroids.

Children may absorb proportionally larger amounts of topical corticosteroids and thus be more susceptible to systemic toxicity (see **PRECAUTIONS, Pediatric Use**).

If irritation or hypersensitivity develops with the combination nystatin and triamcinolone acetonide, treatment should be discontinued and appropriate therapy instituted.

Information for the Patient

Patients using this medicine should receive the following information and instructions.

1. This medication is to be used as directed by the physician. It is for external use only. Avoid contact with the eyes.
2. Patients should be advised not to use this medication for any disorder other than for which it was prescribed.
3. The treated skin area should not be bandaged or otherwise covered or wrapped as to be occluded (see **DOSAGE AND ADMINISTRATION**).
4. Patients should report any signs of local adverse reactions.
5. When using this medication in the inguinal area, patients should be advised to apply cream sparingly and to wear loose fitting clothing.
6. Parents of pediatric patients should be advised not to use tight-fitting diapers or plastic pants on a child being treated in the diaper area, as these garments may constitute occlusive dressings.
7. Patients should be advised on preventative measures to avoid reinfection.

Laboratory Tests

If there is a lack of therapeutic response, appropriate microbiological studies (e.g., KOH smears and/or cultures) should be repeated to confirm the diagnosis and rule out other pathogens, before instituting another course of therapy.

A urinary free cortisol test and ACTH stimulation test may be helpful in evaluating hypothalamic-pituitary-adrenal (HPA) axis suppression due to corticosteroids.

Carcinogenesis, Mutagenesis, and Impairment of Fertility

Long-term animal studies have not been performed to evaluate the carcinogenic or mutagenic potential or possible impairment of fertility in males or females.

Pregnancy: Teratogenic Effects—Pregnancy Category C

There are no teratogenic studies with combined nystatin and triamcinolone acetonide. Corticosteroids are generally teratogenic in laboratory animals when administered systemically at relatively low dosage levels. The more potent corticosteroids have been shown to be teratogenic after dermal application in

laboratory animals. Therefore, any topical corticosteroid preparation should be used during pregnancy only if the potential benefit justifies the potential risk to the fetus.

Topical preparations containing corticosteroids should not be used extensively on pregnant patients, in large amounts, or for prolonged periods of time.

Nursing Mothers

It is not known whether any component of this preparation is excreted in human milk. Because many drugs are excreted in human milk, caution should be exercised during use of this preparation by a nursing woman.

Pediatric Use

In clinical studies of a limited number of pediatric patients ranging in age from two months through 12 years, nystatin and triamcinolone acetonide cream formulation cleared or significantly ameliorated the disease state in most patients.

Pediatric patients may demonstrate greater susceptibility to topical corticosteroid-induced hypothalamic-pituitary-adrenal (HPA) axis suppression and Cushing's syndrome than mature patients because of a larger skin surface area to body weight ratio.

HPA axis suppression, Cushing's syndrome, and intracranial hypertension have been reported in children receiving topical corticosteroids. Manifestations of adrenal suppression in children include linear growth retardation, delayed weight gain, low plasma cortisol levels, and absence of response to ACTH stimulation. Manifestations of intracranial hypertension include bulging fontanelles, headaches and bilateral papilledema.

Administration of topical corticosteroids to children should be limited to the least amount compatible with an effective therapeutic regimen. Chronic corticosteroid therapy may interfere with the growth and development of children.

ADVERSE REACTIONS

A single case (approximately one percent of patients studied) of acneiform eruption occurred with use of combined nystatin and triamcinolone acetonide in clinical studies.

Nystatin is virtually nontoxic and nonsensitizing and is well tolerated by all age groups, even during prolonged use. Rarely, irritation may occur.

The following local adverse reactions are reported infrequently with topical corticosteroids (reactions are listed in an approximate decreasing order of occurrence): burning, itching, irritation, dryness, folliculitis, hypertrichosis, acneiform eruptions, hypopigmentation, perioral dermatitis, allergic contact dermatitis, maceration of the skin, perioral secondary infection, skin atrophy, striae and miliaria.

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.

OVERDOSAGE

Topically applied corticosteroids can be absorbed in sufficient amounts to produce systemic effects (see **PRECAUTIONS, General**); however, acute overdosage and serious adverse effects with dermatologic use are unlikely.

DOSAGE AND ADMINISTRATION

Nystatin and Triamcinolone Acetonide Cream is usually applied to the affected areas twice daily in the morning and the evening by gently and thoroughly massaging the preparation into the skin. The cream should be discontinued if symptoms persist after 25 days of therapy (see **PRECAUTIONS, Laboratory Tests**).

Nystatin and Triamcinolone Acetonide Cream should *not* be used with occlusive dressings.

HOW SUPPLIED

Nystatin and Triamcinolone Acetonide Cream, USP is supplied in 15 g, 30 g, and 60 g tubes.

STORAGE

Store at 20-25°C (68-77°F) [See USP Controlled Room Temperature]. Avoid freezing.

Manufactured by:

Cosette Pharmaceuticals, Inc.
111 Coolidge Street
South Plainfield, NJ 07080

Distributed by:

ActavisPharma, Inc.
Parsippany, NJ 07054 USA

Rev. 09/2019
8-0677ACT4

PRINCIPAL DISPLAY PANEL

NDC 0472-0150-15

Nystatin and Triamcinolone Acetonide Cream, USP

15 g

Rx Only

For Dermatologic Use Only. Not for Ophthalmic Use

Cosette Pharmaceuticals, Inc.

Glue Flap

Each Gram Contains: nystatin 100,000 units and triamcinolone acetonide 1 mg in an aqueous, perfumed, vanishing cream base containing aluminum hydroxide rehydrated, barcroft aluminum hydroxide gel, cetearyl alcohol (and) ceteareth-20, glyceryl monostearate, polyethylene glycol monostearate, propylene glycol, simethicone emulsion, sorbic acid, sorbitol solution, titanium dioxide, white petrolatum, and purified water.

Usual Dosage: Apply to affected areas twice daily. For complete information see accompanying insert.

Warning: Keep out of the reach of children.

Store at 20°-25°C (68°-77°F) [see USP Controlled Room Temperature]. Avoid freezing.

Important: The opening of this product is covered by a metal tamper-evident seal. If this seal has been punctured or is not visible, do

not use and return product to place of purchase.

To Open: To puncture the seal, reverse the cap and place the puncture top onto the tube. Push down firmly until seal is open. To close, screw the cap back onto the tube.

See carton or crimp of tube for lot number and expiration date.

Manufactured by:
Cosette Pharmaceuticals, Inc.
111 Coolidge Street
South Plainfield, NJ 07080
Distributed by:
Actavis Pharma, Inc.
Parsippany, NJ 07054 USA
10-67715ACT4 Rev. 08/2019

GTIN 00304720150152



Clear Code Area



NDC 0472-0150-15

Nystatin and Triamcinolone Acetonide Cream, USP

Rx Only

For Dermatologic Use Only – Not for Ophthalmic Use

15 g (0.53 oz)



NDC 0472-0150-15

Nystatin and Triamcinolone Acetonide Cream, USP

Rx Only

For Dermatologic Use Only – Not for Ophthalmic Use

15 g (0.53 oz)



NDC 0472-0150-15

Nystatin and Triamcinolone Acetonide Cream, USP

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For Dermatologic Use Only – Not for Ophthalmic Use

15 g (0.53 oz)



NDC 0472-0150-15

Nystatin and Triamcinolone Acetonide Cream, USP

Rx Only

For Dermatologic Use Only
Not for Ophthalmic Use

15 g (0.53 oz)

Each Gram Contains: nystatin 100,000 units and triamcinolone acetonide 1 mg in an aqueous, perfumed, vanishing cream base containing aluminum hydroxide rehydrated, barcroft aluminum hydroxide gel, cetearyl alcohol (and) ceteareth-20, glyceryl monostearate, polyethylene glycol monostearate, propylene glycol, simethicone emulsion, sorbic acid, sorbitol solution, titanium dioxide, white petrolatum, and purified water.

Usual Dosage: Apply to affected areas twice daily. For complete information see accompanying insert.

Warning: Keep out of the reach of children.

Store at 20°-25°C (68°-77°F) [see USP Controlled Room Temperature]. Avoid freezing.

Important: Do not use if seal has been punctured or is not visible.

To Open: Use cap to puncture seal.

For lot number and expiration date see crimp of tube.

Manufactured by: Cosette Pharmaceuticals, Inc., 111 Coolidge Street, South Plainfield, NJ 07080

Distributed by: Actavis Pharma, Inc., Parsippany, NJ 07054 USA

Rev. 04/2020 11-67715ACT6



NDC 0472-0150-30

Nystatin and Triamcinolone Acetonide Cream, USP

30 g

Rx Only

For Dermatologic Use Only. Not for Ophthalmic Use

Cosette Pharmaceuticals, Inc.



Actavis

NDC 0472-0150-30

Nystatin and Triamcinolone Acetonide Cream, USP

Rx Only

**For Dermatologic Use Only
Not for Ophthalmic Use**

30 g (1.1 oz)

Each Gram Contains: nystatin 100,000 units and triamcinolone acetonide 1 mg in an aqueous, perfumed, vanishing cream base containing aluminum hydroxide rehydrigel, barcroft aluminum hydroxide gel, cetearyl alcohol (and) cetearth-20, glyceryl monostearate, polyethylene glycol monostearate, propylene glycol, simethicone emulsion, sorbic acid, sorbitol solution, titanium dioxide, white petrolatum, and purified water.

Usual Dosage: Apply to affected areas twice daily. For complete information see accompanying insert.

Warning: Keep out of the reach of children.

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

Avoid freezing.

Important: Do not use if seal has been punctured or is not visible.

To Open: Use cap to puncture seal.

For lot number and expiration date see crimp of tube.



Manufactured by: Cosette Pharmaceuticals, Inc., 111 Coolidge Street, South Plainfield, NJ 07080

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Rev. 08/2019 11-67731ACT5

NDC 0472-0150-60

Nystatin and Triamcinolone Acetonide Cream, USP

60 g

Rx Only

For Dermatologic Use Only. Not for Ophthalmic Use

Cosette Pharmaceuticals, Inc.



Actavis NDC 0472-0150-60

Nystatin and Triamcinolone Acetonide Cream, USP

Rx Only

**For Dermatologic Use Only
Not for Ophthalmic Use** 60 g (2.2 oz)

Each Gram Contains: nystatin 100,000 units and triamcinolone acetonide 1 mg in an aqueous, perfumed, vanishing cream base containing aluminum hydroxide rehydrated, barcroft aluminum hydroxide gel, cetearyl alcohol (and) cetareth-20, glyceryl monostearate, polyethylene glycol monostearate, propylene glycol, simethicone emulsion, sorbic acid, sorbitol solution, titanium dioxide, white petrolatum, and purified water.

Usual Dosage: Apply to affected areas twice daily. For complete information see accompanying insert.

Warning: Keep out of the reach of children.

Store at 20°-25°C (68°-77°F) [see USP Controlled Room Temperature]. Avoid freezing.

Important: Do not use if seal has been punctured or is not visible.

To Open: Use cap to puncture seal.

For lot number and expiration date see crimp of tube.

Manufactured by: Cosette Pharmaceuticals, Inc.
111 Coolidge Street, South Plainfield, NJ 07080

Distributed by: Actavis Pharma, Inc.
Parsippany, NJ 07054 USA

Rev. 08/2019
11-67760ACT5

NYSTATIN AND TRIAMCINOLONE ACETONIDE

nystatin and triamcinolone acetonide cream

Product Information

Product Type	HUMAN PRESCRIPTION DRUG	Item Code (Source)	NDC:0472-0150
Route of Administration	TOPICAL		

Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
NYSTATIN (UNII: BDF1O1C72E) (NYSTATIN - UNII:BDF1O1C72E)	NYSTATIN	100000 [USP ^U] in 1 g
TRIAMCINOLONE ACETONIDE (UNII: F446C597KA) (TRIAMCINOLONE ACETONIDE - UNII:F446C597KA)	TRIAMCINOLONE ACETONIDE	1 mg in 1 g

Inactive Ingredients

Ingredient Name	Strength
ALUMINUM HYDROXIDE (UNII: 5QB0T2IUN0)	
CETOSTEARYL ALCOHOL (UNII: 2DMT128M1S)	
POLYOXYL 20 CETOSTEARYL ETHER (UNII: YRC528SWUY)	
GLYCERYL MONOSTEARATE (UNII: 230OU9XXE4)	
PROPYLENE GLYCOL (UNII: 6DC9Q167V3)	
DIMETHICONE (UNII: 92RU3N3Y1O)	
SORBIC ACID (UNII: X045WJ989B)	
SORBITOL (UNII: 506T60A25R)	
TITANIUM DIOXIDE (UNII: 15FIX9V2JP)	

Product Characteristics

Color	YELLOW (light yellow)	Score	
Shape		Size	
Flavor		Imprint Code	
Contains			

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0472-0150-15	1 in 1 CARTON	06/19/2015	
1		15 g in 1 TUBE; Type 0: Not a Combination Product		
2	NDC:0472-0150-30	1 in 1 CARTON	06/19/2015	
2		30 g in 1 TUBE; Type 0: Not a Combination Product		
3	NDC:0472-0150-60	1 in 1 CARTON	06/19/2015	
3		60 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	ANDA062367	06/19/2015	

Labeler - Actavis Pharma, Inc. (119723554)

Registrant - Cosette Pharmaceuticals, Inc. (116918230)

Establishment

Name	Address	ID/FEI	Business Operations
Cosette Pharmaceuticals, Inc.		116918230	analysis(0472-0150) , label(0472-0150) , manufacture(0472-0150) , pack(0472-0150)

Revised: 11/2020

Actavis Pharma, Inc.