

CLOBETASOL PROPIONATE- clobetasol propionate solution
Micro Labs Limited

PRODUCT INFORMATION

Clobetasol Propionate Topical Solution, USP

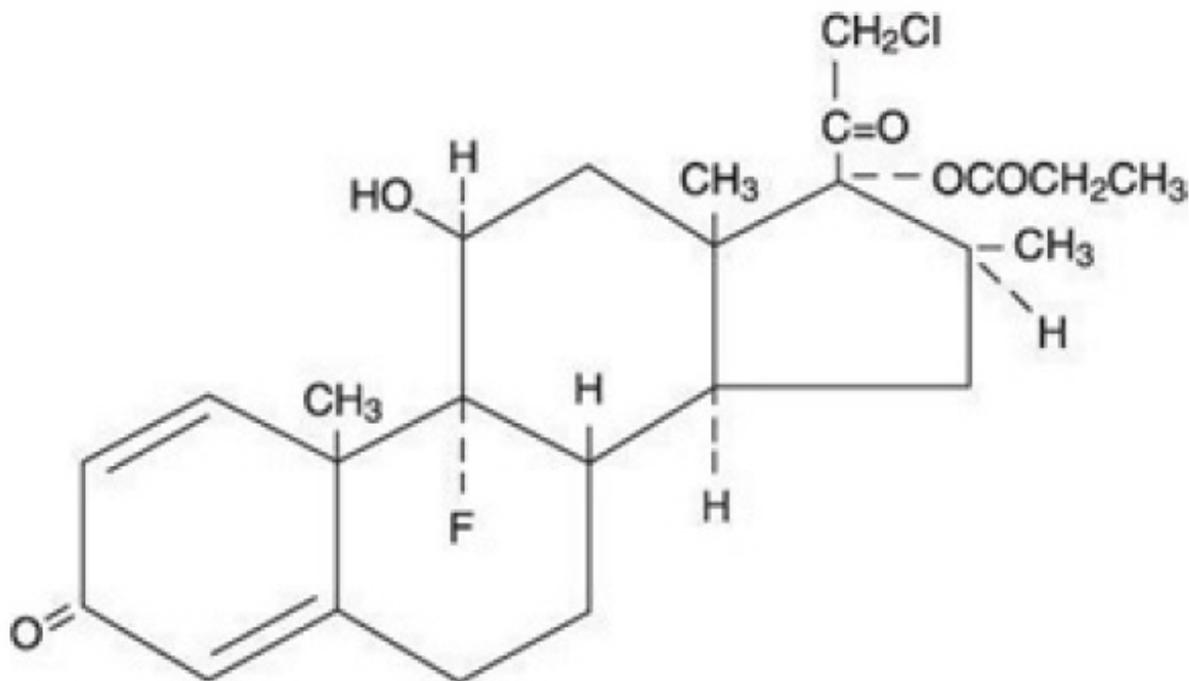
Rx only

FOR TOPICAL DERMATOLOGIC USE ONLY-NOT FOR OPHTHALMIC, ORAL, OR INTRAVAGINAL USE

DESCRIPTION

Clobetasol propionate topical solution, USP contains the active compound clobetasol propionate, USP a synthetic corticosteroid, for topical dermatologic use. Clobetasol, an analog of prednisolone, has a high degree of glucocorticoid activity and a slight degree of mineralocorticoid activity.

Chemically, clobetasol propionate, USP is (11 β ,16 β)-21-chloro-9-fluoro-11-hydroxy-16-methyl-17-(1-oxo-propoxy) pregna-1,4-diene-3,20-dione, and it has the following structural formula:



Clobetasol propionate, USP has the molecular formula C₂₅H₃₂ClFO₅ and a molecular weight of 466.97. It is a white to cream-colored crystalline powder, soluble in acetone, in dimethyl sulfoxide, in chloroform, in methanol, and in dioxane. Sparingly soluble in ethanol; slightly soluble in benzene and diethyl ether; practically insoluble in water.

Clobetasol propionate topical solution, USP contains clobetasol propionate, USP 0.5 mg/g in a base of carbomer 974P, isopropyl alcohol (40% w/w), purified water, and

sodium hydroxide.

CLINICAL PHARMACOLOGY

The corticosteroids are a class of compounds comprising steroid hormones secreted by the adrenal cortex and their synthetic analogs. In pharmacologic doses, corticosteroids are used primarily for their anti-inflammatory and/or immunosuppressive effects. Topical corticosteroids such as clobetasol propionate are effective in the treatment of corticosteroid-responsive dermatoses primarily because of their anti-inflammatory, antipruritic, and vasoconstrictive actions. However, while the physiologic, pharmacologic, and clinical effects of the corticosteroids are well known, the exact mechanisms of their actions in each disease are uncertain.

Clobetasol propionate, a corticosteroid, has been shown to have topical (dermatologic) and systemic pharmacologic and metabolic effects characteristic of this class of drugs.

Pharmacokinetics

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

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

Once absorbed through the skin, topical corticosteroids enter pharmacokinetic pathways similarly 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, including clobetasol propionate and its metabolites, are also excreted into the bile.

Following repeated nonocclusive application in the treatment of scalp psoriasis, there is some evidence that clobetasol propionate topical solution has the potential to depress plasma cortisol levels in some patients. However, hypothalamic-pituitary-adrenal (HPA) axis effects produced by systemically absorbed clobetasol propionate have been shown to be transient and reversible upon completion of a 2-week course of treatment.

INDICATIONS AND USAGE

Clobetasol propionate topical solution, USP is indicated for short-term topical treatment of inflammatory and pruritic manifestations of moderate to severe corticosteroid-responsive dermatoses of the scalp. Treatment beyond 2 consecutive weeks is not recommended, and the total dosage should not exceed 50 mL/week because of the potential for the drug to suppress the HPA axis.

This product is not recommended for use in pediatric patients under 12 years of age.

CONTRAINDICATIONS

Clobetasol propionate topical solution is contraindicated in patients with primary infections of the scalp, or in patients who are hypersensitive to clobetasol propionate, other corticosteroids, or any ingredient in this preparation.

PRECAUTIONS

General

Clobetasol propionate is a highly potent topical corticosteroid that has been shown to suppress the HPA axis at doses as low as 2 g (of ointment) per day. Systemic absorption of topical corticosteroids has resulted in reversible HPA axis suppression, manifestations of Cushing's syndrome, hyperglycemia, and glucosuria in some patients.

Conditions that augment systemic absorption include the application of the more potent corticosteroids, use over large surface areas, prolonged use, and the addition of occlusive dressings. Therefore, patients receiving a large dose of a 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. If HPA axis suppression is noted, an attempt should be made to withdraw the drug, to reduce the frequency of application, or to substitute a less potent steroid.

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

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

If irritation develops, topical corticosteroids should be discontinued and appropriate therapy instituted. Irritation is possible if clobetasol propionate topical solution contacts the eye. If that should occur, immediate flushing of the eye with a large volume of water is recommended.

If the inflammatory lesion becomes infected, the use of an appropriate antifungal or antibacterial agent should be instituted. If a favorable response does not occur promptly, the corticosteroid should be discontinued until the infection has been adequately controlled.

Although clobetasol propionate topical solution is intended for the treatment of inflammatory conditions of the scalp, it should be noted that certain areas of the body, such as the face, groin, and axillae, are more prone to atrophic changes than other areas of the body following treatment with corticosteroids. Frequent observation of the patient is important if these areas are to be treated.

As with other potent topical corticosteroids, clobetasol propionate topical solution should not be used in the treatment of rosacea and perioral dermatitis. Topical corticosteroids in general should not be used in the treatment of acne or as sole therapy in widespread plaque psoriasis.

Information for Patients

Patients using clobetasol propionate topical solution should receive the following

information and instructions:

1. This medication is to be used as directed by the physician and should not be used longer than the prescribed time period. It is for external use only. Avoid contact with the eyes.
2. This medication should not be used for any disorder other than that for which it was prescribed.
3. The treated skin area should not be bandaged or otherwise covered or wrapped so as to be occlusive.
4. Patients should report any signs of local adverse reactions to the physician.

Laboratory Tests

The following tests may be helpful in evaluating patients for HPA axis suppression:

Urinary free cortisol test

ACTH stimulation test

Carcinogenesis, Mutagenesis, Impairment of Fertility

Long-term animal studies have not been performed to evaluate the carcinogenic potential of clobetasol propionate.

Studies in the rat following subcutaneous administration at dosage levels up to 50 mcg/kg per day revealed that the females exhibited an increase in the number of resorbed embryos and a decrease in the number of living fetuses at the highest dose.

Clobetasol propionate was non-mutagenic in 3 different test systems: the Ames test, the *Saccharomyces cerevisiae* gene conversion assay, and the *E. coli* B WP2 fluctuation test.

Pregnancy

Teratogenic Effects

Pregnancy Category C.

Corticosteroids have been shown to be teratogenic in laboratory animals when administered systemically at relatively low dosage levels. Some corticosteroids have been shown to be teratogenic after dermal application to laboratory animals.

Clobetasol propionate has not been tested for teratogenicity when applied topically; however, it is absorbed percutaneously, and when administered subcutaneously it was a significant teratogen in both the rabbit and mouse. Clobetasol propionate has greater teratogenic potential than steroids that are less potent.

Teratogenicity studies in mice using the subcutaneous route resulted in fetotoxicity at the highest dose tested (1 mg/kg) and teratogenicity at all dose levels tested down to 0.03 mg/kg. These doses are approximately 1.4 and 0.04 times, respectively, the human topical dose of clobetasol propionate topical solution. Abnormalities seen included cleft palate and skeletal abnormalities.

In rabbits, clobetasol propionate was teratogenic at doses of 3 and 10 mcg/kg. These doses are approximately 0.02 and 0.05 times, respectively, the human topical dose of clobetasol propionate topical solution. Abnormalities seen included cleft palate, cranioschisis, and other skeletal abnormalities.

There are no adequate and well-controlled studies of the teratogenic potential of

clobetasol propionate in pregnant women. Clobetasol propionate topical solution should be used during pregnancy only if the potential benefit justifies the potential risk to the fetus.

Nursing Mothers

Systemically administered corticosteroids appear in human milk and could suppress growth, interfere with endogenous corticosteroid production, or cause other untoward effects. It is not known whether topical administration of corticosteroids could result in sufficient systemic absorption to produce detectable quantities in human milk.

Because many drugs are excreted in human milk, caution should be exercised when clobetasol propionate topical solution is administered to a nursing woman.

Pediatric Use

Use of clobetasol propionate topical solution in pediatric patients under 12 years of age is not recommended.

Pediatric patients may demonstrate greater susceptibility to topical corticosteroid-induced 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, linear growth retardation, delayed weight gain, and intracranial hypertension have been reported in children receiving topical corticosteroids. Manifestations of adrenal suppression in children include low plasma cortisol levels and an absence of response to ACTH stimulation. Manifestations of intracranial hypertension include bulging fontanelles, headaches, and bilateral papilledema.

Geriatric Use

A limited number of patients at or above 65 years of age (n = 65) have been treated with clobetasol propionate topical solution in US and non-US clinical trials. While the number of patients is too small to permit separate analysis of efficacy and safety, the adverse reactions reported in this population were similar to those reported by younger patients. Based on available data, no adjustment of dosage of clobetasol propionate topical solution in geriatric patients is warranted.

ADVERSE REACTIONS

Clobetasol propionate topical solution is generally well tolerated when used for 2-week treatment periods.

The most frequent adverse events reported for clobetasol propionate topical solution have been local and have included burning and/or stinging sensation, which occurred in 29 of 294 patients; scalp pustules, which occurred in 3 of 294 patients; and tingling and folliculitis, each of which occurred in 2 of 294 patients. Less frequent adverse events were itching and tightness of the scalp, dermatitis, tenderness, headache, hair loss, and eye irritation, each of which occurred in 1 of 294 patients.

The following local adverse reactions are reported infrequently when topical corticosteroids are used as recommended. These reactions are listed in an approximately decreasing order of occurrence: burning, itching, irritation, dryness, folliculitis, hypertrichosis, acneiform eruptions, hypopigmentation, perioral dermatitis,

allergic contact dermatitis, maceration of the skin, secondary infection, skin atrophy, striae, and miliaria. Systemic absorption of topical corticosteroids has produced reversible HPA axis suppression, manifestations of Cushing's syndrome, hyperglycemia, and glucosuria in some patients. In rare instances, treatment (or withdrawal of treatment) of psoriasis with corticosteroids is thought to have exacerbated the disease or provoked the pustular form of the disease, so careful patient supervision is recommended.

To report SUSPECTED ADVERSE REACTIONS, contact Saptalis Pharmaceuticals, LLC at 1-833-727-8254 or FDA at 1-800-FDA-1088 or www.fda.gov/medwatch.

OVERDOSAGE

Topically applied clobetasol propionate can be absorbed in sufficient amounts to produce systemic effects (see **PRECAUTIONS**).

DOSAGE AND ADMINISTRATION

Clobetasol propionate topical solution should be applied to the affected scalp areas twice daily, once in the morning and once at night.

Clobetasol propionate topical solution is potent; therefore, **treatment must be limited to 2 consecutive weeks and amounts greater than 50 mL/week should not be used.**

Clobetasol propionate topical solution is not to be used with occlusive dressings.

Geriatric Use

In studies where geriatric patients (65 years of age or older, see **PRECAUTIONS**) have been treated with clobetasol propionate topical solution, safety did not differ from that in younger patients; therefore, no dosage adjustment is recommended.

HOW SUPPLIED

Clobetasol propionate topical solution, USP 0.05% is a clear, colorless liquid supplied in plastic squeeze bottles, 25 mL (NDC 42571-362-81) and 50 mL (NDC 42571-362-99).

Store at 20° to 25°C (68° to 77°F), with excursions permitted between 15° to 30°C (59° to 86°F) [See USP Controlled Room Temperature]. Do not refrigerate. Keep tightly closed.

Do not use near an open flame.

Distributed by:
Micro Labs USA, Inc.
Somerset, NJ 08873

Manufactured by:
Saptalis Pharmaceuticals, LLC

Hauppauge, NY 11788

Revised: January 2022-R2

Package/Label Display Panel - 25 mL Label

NDC 42571-362-81

Clobetasol Propionate Topical Solution, USP

0.05%

SCALP APPLICATION

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

Rx only

25 mL

Front Label

NDC 42571-362-81

**Clobetasol
Propionate
Topical
Solution, USP**

0.05%

SCALP APPLICATION

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

Rx only 25 mL

Back Label

Contains: clobetasol propionate, USP 0.05% (0.5 mg/g) in a base composed of carbomer 974P, isopropyl alcohol (40% w/w), purified water, and sodium hydroxide.

USUAL DOSAGE: See package insert for full prescribing information.

Store at 20° to 25°C (68° to 77°F), with excursions permitted between 15° to 30°C (59° to 86°F) [See USP Controlled Room Temperature].
Do not refrigerate. Keep tightly closed.

Do not use near an open flame.

WARNING: Keep out of reach of children.

Manufactured by: Saptalis Pharmaceuticals, LLC
Hauppauge, NY 11788

Distributed by: Micro Labs USA, Inc.
Somerset, NJ 08873

MADE IN USA 01/22-H2



N 3 1 4 2 5 7 1 3 6 2 8 1 2

Unvarnished area for batch details
LOT: XXXXXXX
EXP: MM/YYYY
will be printed online

OR

Unvarnished area as per DGFT
2D Matrix: EXP: (17) XXXXXXX
LOT: (10) XXXXXXX
S.No.: (21) Alphanumeric Serialization No.
will be printed online *

* The batch coding given as per regular practice. Once after the Plant is ready for DGFT printing, the batch coding details will be revised as mentioned above to adhere the Directorate General of Foreign Trade Notification

Package/Label Display Panel - 25 mL Carton

NDC 42571-362-81

Clobetasol Propionate Topical Solution, USP

0.05%

SCALP APPLICATION

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

Rx only

25 mL

* The batch coding given as per regular practice. Once after details will be revised as mentioned above to adhere the Directorate General of Foreign Trade Notification.

CD NUMBER: CDX1 (7) 000000
 LOT: XXXXXXX
 MFG: 11/11/11
 EXP: 11/11/11
 OR
 CD NUMBER: CDX1 (7) 000000
 LOT: XXXXXXX
 MFG: 11/11/11
 EXP: 11/11/11

Unshaded area for batch details will be printed online.



Package/Label Display Panel - 50 mL Label
NDC 42571-362-99
Clobetasol Propionate Topical Solution, USP

0.05%

SCALP APPLICATION

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

Rx only

50 mL

Front Label

NDC 42571-362-99

Clobetasol Propionate Topical Solution, USP

0.05%

SCALP APPLICATION

For Dermatologic Use Only—
Not for Ophthalmic Use.

Rx only 50 mL




Back Label

Contains: clobetasol propionate, USP 0.05% (0.5 mg/g) in a base composed of carbomer 974P, isopropyl alcohol (40% w/w), purified water, and sodium hydroxide.

USUAL DOSAGE: See package insert for full prescribing information.

Store at 20° to 25°C (68° to 77°F), with excursions permitted between 15° to 30°C (59° to 86°F) [See USP Controlled Room Temperature]. Do not refrigerate. Keep tightly closed.

Do not use near an open flame.

WARNING: Keep out of reach of children.

Manufactured by: **Saptalis Pharmaceuticals, LLC**
Hauppauge, NY 11788

Distributed by: **Micro Labs USA, Inc.**
Somerset, NJ 08873

MADE IN USA 01/22-R2



N 3 4 2 5 7 1 3 6 2 9 9 7

Unvarnished area for batch details
 LOT: XXXXXXX
 EXP: MM/YYYY
 will be printed online

OR

Unvarnished area as per DGFT *****
 GTIN: (21) Global Trade Item No.
 2D Matrix : EXP: (17) YYMMDD
 LOT: (18) XXXXXXX
 S.No.: (21) Alphanumerical Serialization No.
 will be printed online

***** The batch coding given as per regular practice. Once after the Plant is ready for DGFT printing, the batch coding details will be revised as mentioned above to adhere the Directorate General of Foreign Trade Notification

Package/Label Display Panel - 50 mL Carton
NDC 42571-362-99
Clobetasol Propionate Topical Solution, USP
0.05%

SCALP APPLICATION

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

Rx only

50 mL

clobetasol propionate solution

Product Information

| | | | |
|--------------------------------|-------------------------|---------------------------|---------------|
| Product Type | HUMAN PRESCRIPTION DRUG | Item Code (Source) | NDC:42571-362 |
| Route of Administration | TOPICAL | | |

Active Ingredient/Active Moiety

| Ingredient Name | Basis of Strength | Strength |
|--|-----------------------|----------------|
| CLOBETASOL PROPIONATE (UNII: 779619577M) (CLOBETASOL - UNII:ADN79D536H) | CLOBETASOL PROPIONATE | 0.5 mg in 1 mL |

Inactive Ingredients

| Ingredient Name | Strength |
|---|----------|
| CARBOMER HOMOPOLYMER TYPE B (ALLYL PENTAERYTHRITOL CROSSLINKED) (UNII: HHT01ZNK31) | |
| ISOPROPYL ALCOHOL (UNII: ND2M416302) | |
| SODIUM HYDROXIDE (UNII: 55X04QC32I) | |
| WATER (UNII: 059QF0KO0R) | |

Packaging

| # | Item Code | Package Description | Marketing Start Date | Marketing End Date |
|---|------------------|--|----------------------|--------------------|
| 1 | NDC:42571-362-81 | 1 in 1 CARTON | 10/02/2019 | |
| 1 | | 25 mL in 1 BOTTLE; Type 0: Not a Combination Product | | |
| 2 | NDC:42571-362-99 | 1 in 1 CARTON | 10/02/2019 | |
| 2 | | 50 mL in 1 BOTTLE; Type 0: Not a Combination Product | | |

Marketing Information

| Marketing Category | Application Number or Monograph Citation | Marketing Start Date | Marketing End Date |
|--------------------|--|----------------------|--------------------|
| ANDA | ANDA211494 | 10/02/2019 | |

Labeler - Micro Labs Limited (862174955)

Registrant - Saptalis Pharmaceuticals, LLC (080145868)

Establishment

| Name | Address | ID/FEI | Business Operations |
|-------------------------------|---------|-----------|------------------------|
| Saptalis Pharmaceuticals, LLC | | 080145868 | MANUFACTURE(42571-362) |

