

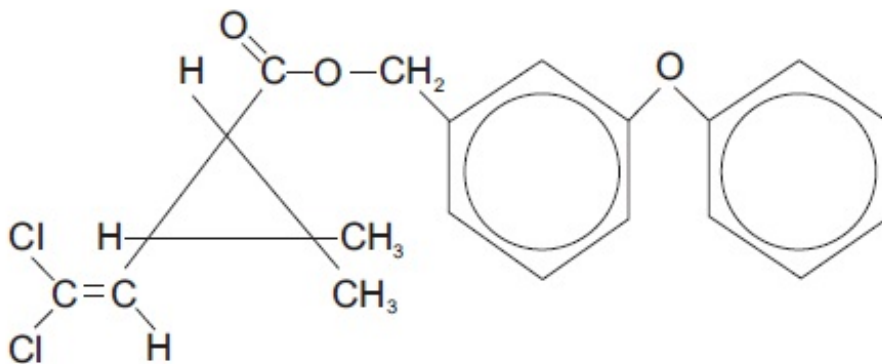
PERMETHRIN - permethrin cream 5% w/w cream
Encube Ethicals Private Limited

PERMETHRIN CREAM 5% W/W

Rx Only

DESCRIPTION

Permethrin Cream 5% w/w is a topical scabicedal agent for the treatment of infestation with *Sarcoptes scabiei* (scabies). It is available in an off-white, vanishing cream base. Permethrin Cream 5% w/w is for topical use only.



Chemical Name - The permethrin used is an approximate 1:3 mixture of the cis and trans isomers of the pyrethroid 3-(2,2-dichloroethenyl)-2,2- dimethylcyclopropanecarboxylic acid, (3-phenoxyphenyl) methyl ester. Permethrin has a molecular formula of $C_{21}H_{20}Cl_2O_3$ and a molecular weight of 391.29. It is a yellow to light orange-brown, low melting solid or viscous liquid.

Active Ingredient - Each gram contains permethrin 50 mg (5%).

Inactive Ingredients - Carbomer homopolymer type B, medium-chain triglycerides, glycerin, mono- and diglycerides, isopropyl myristate, lanolin alcohols, mineral oil, polyoxyethylene (20) cetyl ethers, purified water, and sodium hydroxide. Formaldehyde 1 mg (0.1%) is added as a preservative.

CLINICAL PHARMACOLOGY

Permethrin, a pyrethroid, is active against a broad range of pests including lice, ticks, fleas, mites, and other arthropods. It acts on the nerve cell membrane to disrupt the sodium channel current by which the polarization of the membrane is regulated. Delayed repolarization and paralysis of the pests are the consequences of this disturbance.

Permethrin is rapidly metabolized by ester hydrolysis to inactive metabolites which are excreted primarily in the urine. Although the amount of permethrin absorbed after a single application of the 5% cream has not been determined precisely, data from studies with ^{14}C -labeled permethrin and absorption

studies of the cream applied to patients with moderate to severe scabies indicate it is 2% or less of the amount applied.

INDICATIONS AND USAGE

Permethrin Cream 5% w/w is indicated for the treatment of infestation with *Sarcoptes scabiei* (scabies).

CONTRAINDICATIONS

Permethrin Cream 5% w/w is contraindicated in patients with known hypersensitivity to any of its components, to any synthetic pyrethroid or pyrethrin.

WARNINGS

If hypersensitivity to Permethrin Cream 5% w/w occurs, discontinue use.

PRECAUTIONS

General

Scabies infestation is often accompanied by pruritus, edema, and erythema. Treatment with Permethrin Cream 5% w/w may temporarily exacerbate these conditions.

Information for Patients

Patients with scabies should be advised that itching, mild burning and/or stinging may occur after application of Permethrin Cream 5% w/w. In clinical trials, approximately 75% of patients treated with Permethrin Cream 5% w/w who continued to manifest pruritus at 2 weeks had cessation by 4 weeks. If irritation persists, they should consult their physician. Permethrin Cream 5% w/w may be very mildly irritating to the eyes. Patients should be advised to avoid contact with eyes during application and to flush with water immediately if Permethrin Cream 5% w/w gets in the eyes.

Carcinogenesis, Mutagenesis, Impairment of Fertility

Six carcinogenicity bioassays were evaluated with permethrin, three each in rats and mice. No tumorigenicity was seen in the rat studies. However, species-specific increases in pulmonary adenomas, a common benign tumor of mice of high spontaneous background incidence, were seen in the three mouse studies. In one of these studies there was an increased incidence of pulmonary alveolar-cell carcinomas and benign liver adenomas only in female mice when permethrin was given in their food at a concentration of 5000 ppm. Mutagenicity assays, which give useful correlative data for interpreting results from carcinogenicity bioassays in rodents, were negative. Permethrin showed no evidence of mutagenic potential in a battery of in vitro and in vivo genetic toxicity studies. Permethrin did not have any adverse effect on reproductive function at a dose of 180 mg/kg/day orally in a three-generation rat study.

Pregnancy: Teratogenic Effects: Pregnancy Category B

Reproduction studies have been performed in mice, rats, and rabbits (200 to 400 mg/kg/day orally) and have revealed no evidence of impaired fertility or harm to the fetus due to permethrin. 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 and because of the evidence for tumorigenic potential of permethrin in animal studies, consideration should be given to discontinuing nursing temporarily or withholding the drug while the mother is nursing.

Pediatric Use

Permethrin Cream 5% w/w is safe and effective in pediatric patients two months of age and older. Safety and effectiveness in infants less than two months of age have not been established.

Geriatric Use

Clinical studies of Permethrin Cream 5% w/w did not identify sufficient numbers of subjects aged 65 and over to allow a definitive statement regarding whether elderly subjects respond differently from younger subjects. Other reported clinical experience has not identified differences in responses between the elderly and younger patients. This drug is known to be substantially excreted by the kidney. However, since topical permethrin is metabolized in the liver and excreted in the urine as inactive metabolites, there does not appear to be an increased risk of toxic reactions in patients with impaired renal function when used as labeled.

ADVERSE REACTIONS

In clinical trials, generally mild and transient burning and stinging followed application with Permethrin Cream 5% w/w in 10% of patients and was associated with the severity of infestation. Pruritus was reported in 7% of patients at various times post-application. Erythema, numbness, tingling, and rash were reported in 1 to 2% or less of patients (see PRECAUTIONS - General). Other adverse events reported since marketing Permethrin Cream 5% w/w include: headache, fever, dizziness, abdominal pain, diarrhea and nausea and/or vomiting. Although extremely uncommon and not expected when used as directed (see DOSAGE AND ADMINISTRATION), rare occurrences of seizure have been reported. None have been medically confirmed as associated with Permethrin Cream 5% w/w treatment.

OVERDOSAGE

No instance of accidental ingestion of Permethrin Cream 5% w/w has been reported. If ingested, gastric lavage and general supportive measures should be employed. Excessive topical use (see DOSAGE AND ADMINISTRATION) may result in increased irritation and erythema.

DOSAGE AND ADMINISTRATION

Adults and children - Thoroughly massage Permethrin Cream 5% w/w into the skin from the head to the soles of the feet. Scabies rarely infests the scalp of adults, although the hairline, neck, temple, and forehead may be infested in infants and geriatric patients. Usually 30 grams is sufficient for an average adult. The cream should be removed by washing (shower or bath) after 8 to 14 hours. Infants should be treated on the scalp, temple, and forehead. **ONE APPLICATION IS GENERALLY CURATIVE.** Patients may experience persistent pruritus after treatment. This is rarely a sign of treatment failure and is not an indication for retreatment. Demonstrable living mites after 14 days indicate that retreatment is necessary.

HOW SUPPLIED

Permethrin Cream 5% w/w is available as follows:
60 g tube (NDC 21922-021-07).

STORAGE

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

Manufactured by:

Encube Ethicals Pvt. Ltd.

Plot No. C-1, Madkaim Industrial Estate,

Makaim, Post: Mardol,

Ponda Goa- 403 404- India.

Distributed by:

Encube Ethicals, Inc.

200 Meredith Avenue,

Suite 101A

Durham,

NC 27713

USA

Rev 01

April 2019

PACKAGE LABEL.PRINCIPAL DISPLAY PANEL

PRINCIPAL DISPLAY PANEL- 60 g Container Label

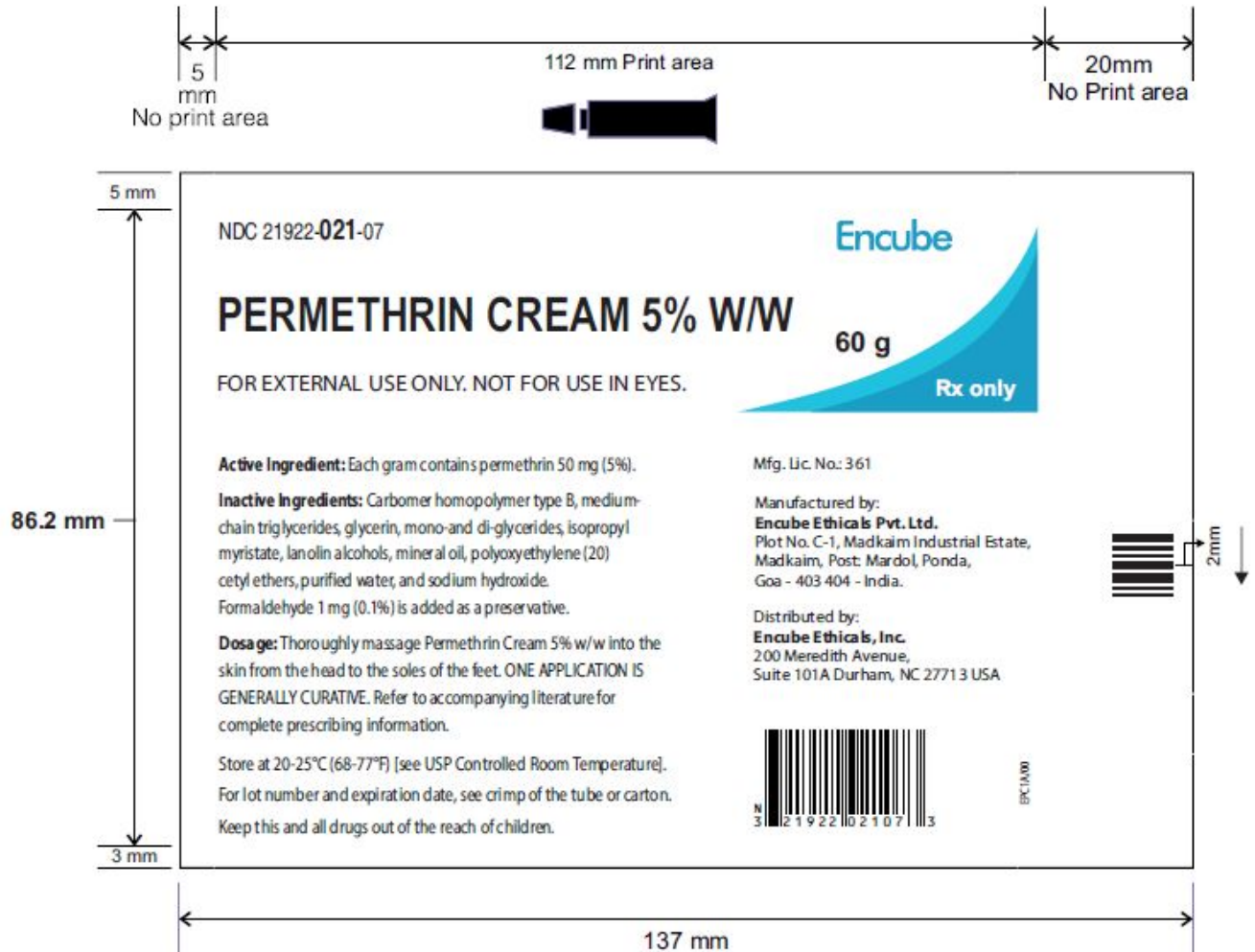
NDC 21922-021-07

Permethrin Cream 5% w/w

Rx Only

60 g

Size: Dia 30mm x 137mm



PRINCIPAL DISPLAY PANEL- 60 g Carton Label


NDC 21922-021-07

Permethrin Cream 5% w/w

Rx Only

60 g

2D Coding and Batch Details to be overcoded
(For Reference Only)



GTIN: XXXXXXXXXX
S.ND: XXXXXXXX
EXP: XXXXX
LOT: XXXXXXXX

Area for batch coding

NDC 21922-021-07

PERMETHRIN CREAM 5% W/W

60 g

FOR EXTERNAL USE ONLY. NOT FOR USE IN EYES.

Rx only



Mfg. Lic. No: 361

Manufactured by:
Encube Ethicals Pvt. Ltd.
Plot No. C-1, Madkaim Industrial Estate,
Madkaim, Post: Mardol, Ponda,
Goa - 403 404 - India.

Distributed by:
Encube Ethicals, Inc.
200 Meredith Avenue,
Suite 101A Durham, NC 27713 USA



3 121922 240210 713

158 x 35 x 40 mm

NDC 21922-021-07

PERMETHRIN CREAM 5% W/W

60 g

FOR EXTERNAL USE ONLY. NOT FOR USE IN EYES.

Rx only



Dosage: Thoroughly massage Permethrin Cream 5% w/w into the skin from the head to the toes of the feet. ONE APPLICATION IS GENERALLY CURATIVE. Refer to accompanying literature for complete prescribing information.

For lot number and expiration date, see crimp of the tube or carton.

Active Ingredient:
Each gram contains permethrin 50 mg (5%).

Inactive Ingredients:
Carbomer homopolymer type B, medium-chain triglycerides, glycerin, mono-and di-glycerides, isopropylmyristate, linolin alcohol, mineral oil, polyoxyethylene (20) cetyl ether, purified water, and sodium hydroxide.
Formaldehyde 1 mg (0.1%) is added as a preservative.

Store at 20-25°C (68-77°F)
[see USP Controlled Room Temperature].
Keep this and all drugs out of the reach of children.

Ph. Code: 152



Material: Permethrin Cream 5% Carton
Artwork Code: EPC3A00
SAP Code: 206251
Pharma code: 152

Specifications:
Size: 158 x 35 x 40 mm
Substrate: FBB Board (ITC Cyber XL)
Grammage: 300 gsm
Varnish: Aqua Varnish
Special requirements: NA

Colours: 05 - PANTONE Process Blue C
PANTONE 639 C, PANTONE 312 C,
PANTONE 311 C & Black

PERMETHRIN			
permethrin cream 5% w/w cream			
Product Information			
Product Type	HUMAN PRESCRIPTION DRUG	Item Code (Source)	NDC:219 22-0 21
Route of Administration	TOPICAL		
Active Ingredient/Active Moiety			
	Ingredient Name	Basis of Strength	Strength
	PERMETHRIN (UNII: 509 F88 P9 S Z) (PERMETHRIN - UNII:509 F88 P9 S Z)	PERMETHRIN	50 mg in 1 g
Inactive Ingredients			
	Ingredient Name		Strength
	CARBOMER HOMOPOLYMER TYPE B (ALLYL PENTAERYTHRITOL CROSSLINKED) (UNII: HHT01ZNK31)		

MEDIUM-CHAIN TRIGLYCERIDES (UNII: C9H2L21V7U)	
GLYCERIN (UNII: PDC6A3C0OX)	
GLYCERYL MONO AND DIPALMITO STEARATE (UNII: KC98RO82HJ)	
ISOPROPYL MYRISTATE (UNII: 0RE8K4LNJS)	
LANOLIN ALCOHOLS (UNII: 884C3FA9HE)	
MINERAL OIL (UNII: T5L8T28FGP)	
CETETH-20 (UNII: I835H2IHHX)	
SODIUM HYDROXIDE (UNII: 55X04QC32I)	
FORMALDEHYDE (UNII: 1HG84L3525)	
WATER (UNII: 059QF0KO0R)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:21922-021-07	1 in 1 CARTON	04/08/2019	
1		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	ANDA211303	04/08/2019	

Labeler - Encube Ethicals Private Limited (915834105)

Registrant - Encube Ethicals Private Limited (915834105)

Establishment

Name	Address	ID/FEI	Business Operations
Encube Ethicals Private Limited		725076298	ANALYSIS(21922-021) , LABEL(21922-021) , MANUFACTURE(21922-021) , PACK(21922-021) , PARTICLE SIZE REDUCTION(21922-021)

Revised: 4/2019

Encube Ethicals Private Limited