

SUMMARY OF PRODUCT CHARACTERISTICS

1 NAME OF THE MEDICINAL PRODUCT

Lamisil® Once™ 1% cutaneous solution

2 QUALITATIVE AND QUANTITATIVE COMPOSITION

Each gram contains 10 mg terbinafine (as hydrochloride).

Excipient with known effect

Each gram contains 863.75 mg ethanol.

For the full list of excipients, see section 6.1.

3 PHARMACEUTICAL FORM

Cutaneous solution.

Clear to slightly opaque viscous solution.

4 CLINICAL PARTICULARS

4.1 Therapeutic indications

Treatment for tinea pedis (athlete's foot).

Lamisil Once 1% cutaneous solution is indicated for Adults 18 years of age and over.

4.2 Posology and method of administration

For Cutaneous Use

Adults 18 years of age and over: single administration.

Duration and frequency of treatment

Lamisil Once 1% cutaneous solution should be applied once on both feet, even if lesions are visible on one foot only. This ensures elimination of the fungi (dermatophytes) that might be found in areas of the foot where no lesions are visible.

Relief of clinical symptoms usually occurs within a few days. If there are no signs of improvement after one week, patients should see a doctor. There are no data on repeated treatment with Lamisil Once 1% cutaneous solution. Therefore a second treatment cannot be recommended within a particular episode of athlete's foot.

Dosing in special populations

Paediatric population

Lamisil Once 1% cutaneous solution has not been studied in the paediatric population. Its use is therefore not recommended in patients below 18 years of age.

Elderly patients

There is no evidence to suggest that elderly patients require different dosages or experience side effects different from those in younger patients.

Method of Administration

Patients should wash and dry both feet and hands before applying the product. They should treat one foot, then the other.

Starting between the toes, patients should apply a thin layer evenly between and all around the toes, as well as cover the sole and sides of the foot for up to 1.5 cm. The product should be applied in the same way to the other foot, even if the skin looks healthy. The product should be left to dry to a film for 1-2 minutes. Patients should then wash their hands. Lamisil Once 1% cutaneous solution should not be massaged into skin.

For the best results, the treated area should not be washed for 24 hours after application. It is therefore recommended to apply Lamisil Once 1% cutaneous solution after a shower or bath and wait until the same time the following day before washing the feet again.

Patients should use the quantity they need to cover both feet as instructed above. Any unused medication is to be discarded.

4.3 Contraindications

Hypersensitivity to terbinafine or to any of the excipients listed in section 6.1.

4.4 Special warnings and precautions for use

Lamisil Once 1% cutaneous solution is not recommended to treat hyperkeratotic chronic plantar tinea pedis (moccasin type).

Lamisil Once 1% cutaneous solution:

- Should be used with caution in patients with lesions where alcohol could be irritating (after sun exposure or severe skin scaling).
- Should not be used on the face.
- For external use only.

- May be irritating to the eyes. In case of accidental contact with the eyes, rinse eyes thoroughly with running water/

This medicine contains 3316.8 mg alcohol (ethanol) in each daily dose which is equivalent to 863.75 mg/g of 96% ethanol. It may cause burning sensation on damaged skin.

Do not swallow.

Keep out of sight and reach of children.

In the event of allergic reaction, the film should be removed with an organic solvent such as denatured alcohol and the feet washed with warm soapy water.

- Keep away from naked flames.

4.5 Interaction with other medicinal products and other forms of interaction

No drug interactions are known with the use of topical forms of terbinafine.

4.6 Fertility, Pregnancy and lactation

Pregnancy

There is no clinical experience with terbinafine in pregnant women. Foetal toxicity and fertility studies in animals suggest no adverse effects (see section 5.3).

Lamisil Once 1% cutaneous solution should not be used during pregnancy unless clearly necessary.

Lactation

Terbinafine is excreted in breast milk. Lamisil Once 1% cutaneous solution should not be used during breast-feeding.

Fertility

No effect of terbinafine on fertility have been seen on animal studies (see section 5.3).

4.7 Effects on ability to drive and use machines

Lamisil Once 1% cutaneous solution has no or negligible influence on the ability to drive and use machines.

4.8 Undesirable effects

Summary of the safety profile

Undesirable effects include mild and transient reactions at the site of application. In very rare instances, allergic reactions may occur.

Tabulated list of adverse reactions

Adverse reactions are listed below by system organ class and frequency.

Frequencies are defined as: very common ($\geq 1/10$); common ($\geq 1/100$ to $< 1/10$); uncommon ($\geq 1/1,000$ to $< 1/100$); rare ($\geq 1/10,000$ to $< 1/1,000$); very rare ($< 1/10,000$), or not known (can not to be estimated from available data). Within each frequency grouping, adverse reactions are presented in order of decreasing seriousness.

Description of selected adverse reactions

Skin and subcutaneous tissue disorders:

Very rare ($< 1/10,000$, including isolated reports): allergic reactions such as rash, pruritus, dermatitis bullous and urticaria.

General disorders and administration site conditions

Uncommon ($> 1/1,000$, $< 1/100$): application site reactions such as skin dryness, skin irritation or burning sensation.

Reporting of suspected adverse reactions

Reporting suspected adverse reactions after authorisation of the medicinal product is important. It allows continued monitoring of the benefit/risk balance of the medicinal product. Healthcare professionals are asked to report any suspected adverse reactions via the Yellow Card Scheme at: www.mhra.gov.uk/yellowcard.

4.9 Overdose

Symptoms

Overdose is very unlikely to happen since the product is for single dose, cutaneous use, and the tube only contains the necessary quantity for one application.

Accidental ingestion of one 4 g tube of product contains 40 mg terbinafine which is much lower than one 250 mg Lamisil tablet (oral unit dose).

Should several tubes of Lamisil Once 1% cutaneous solution be inadvertently ingested however, adverse effects similar to those observed with an overdose of Lamisil tablets, are to be expected. These include headache, nausea, epigastric pain and dizziness.

In case of accidental oral ingestion, the alcohol content 81.05% w/w of Lamisil Once 1% cutaneous solution has to be considered

Management

If accidentally ingested, the recommended treatment of overdosage consists of eliminating the active substance, primarily by the administration of activated charcoal, and giving symptomatic supportive therapy if needed.

5 PHARMACOLOGICAL PROPERTIES

5.1 Pharmacodynamic properties

Pharmacotherapeutic group: Antifungal for topical use (ATC code D01 A E15)

Terbinafine is an allylamine which has a broad spectrum of antifungal activity in fungal infections of the skin caused by dermatophytes such as *Trichophyton* (e.g. *T. rubrum*, *T. mentagrophytes*, *T. verrucosum*, *T. violaceum*), *Microsporum canis* and *Epidermophyton floccosum*. At low concentrations terbinafine is fungicidal against dermatophytes and moulds. The activity against yeasts is fungicidal (e.g. *Pityrosporum orbiculare* or *Malassezia furfur*) or fungistatic, depending on the species.

Terbinafine interferes specifically with fungal sterol biosynthesis at an early step. This leads to a deficiency in ergosterol and to an intracellular accumulation of squalene, resulting in fungal cell death. Terbinafine acts by inhibition of squalene epoxidase in the fungal cell membrane. The enzyme squalene epoxidase is not linked to the cytochrome P450 system. Terbinafine does not influence the metabolism of hormones or other drugs.

Studies in patients have shown that a single dose application of Lamisil Once 1 % cutaneous solution on both feet demonstrated efficacy in patients with tinea pedis (athlete's foot) presenting lesions between the toes, and extending to adjacent skin areas of the sides and soles of the feet. The rate of relapse/reinfection at 3 months after treatment was low: 1 person out of 8 (12.5%).

5.2 Pharmacokinetic properties

Once applied to the skin, Lamisil Once 1 % cutaneous solution forms a film on the skin. Terbinafine in the film stays on the skin for up to 72 hours. The film quickly delivers terbinafine to the stratum corneum: at 60 minutes after application, 16 to 18% of the applied dose will be present in the stratum corneum. Delivery progressively continues and terbinafine persists in the stratum corneum for up to 13 days, at levels which are in excess of the in vitro Minimum Inhibitory Concentration for terbinafine against dermatophytes.

Systemic bioavailability is very low. An application of Lamisil Once 1 % cutaneous solution on the back, on an area of 3 times the area of both feet, resulted in exposure to terbinafine of less than 0.5% of the exposure following oral administration of a 250 mg tablet.

5.3 Preclinical safety data

In long-term studies (up to 1 year) in rats and dogs no marked toxic effects were seen in either species up to oral doses of about 100 mg/kg a day. At high oral doses, the liver and possibly also the kidneys were identified as potential target organs.

In a two-year oral carcinogenicity study in mice, no neoplastic or other abnormal findings attributable to treatment were made up to doses of 130 (males) and 156 (females) mg/kg a day. In a two-year oral carcinogenicity study in rats at the highest dose level, 69 mg/kg a day, an increased incidence of liver tumours was observed in males. The changes, which may be associated with peroxisome proliferation, have been shown to be species-specific since they were not seen in the carcinogenicity study in mice or in other studies in mice, dogs or monkeys.

During the studies of high dose oral terbinafine in monkeys, refractile irregularities were observed in the retina at the higher doses (non-toxic effect level was 50 mg/kg). These irregularities were associated with the presence of a terbinafine metabolite in ocular tissue and disappeared after drug discontinuation. They were no associated histological changes.

A standard battery of *in vitro* and *in vivo* genotoxicity tests revealed no evidence of a mutagenic or clastogenic potential for the drug.

No adverse effects on fertility or other reproduction parameters were observed in studies in rats or rabbits.

Repeated dermal administration of Lamisil Once 1 % cutaneous solution in rats and minipigs produces plasma terbinafine levels which are at least 50-100 times lower than the no-adverse-effect-levels established in terbinafine animal toxicity studies, so use of the product is not expected to produce any systemic adverse effect. Lamisil Once 1 % cutaneous solution was well tolerated in a variety of tolerability studies and did not cause sensitisation.

6 PHARMACEUTICAL PARTICULARS

6.1 List of excipients

Acrylates/octylacrylamide copolymer;
Hydroxypropylcellulose;
Medium chain triglycerides;
Ethanol.

6.2 Incompatibilities

Not applicable.

6.3 Shelf life

3 years.

6.4 Special precautions for storage

Store in the original package. There is no special temperature precaution for storage.

6.5 Nature and contents of container

4 g aluminium laminated tube (polyethylene-aluminium-polyethylene) with a polyethylene screw cap.

6.6 Special precautions for disposal

No special requirements.

7 MARKETING AUTHORISATION HOLDER

Karo Healthcare AB
Box 16184
103 24 Stockholm
Sweden

8 MARKETING AUTHORISATION NUMBER(S)

PL 50567/0016

9 DATE OF FIRST AUTHORISATION/RENEWAL OF THE AUTHORISATION

24/04/2025

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