1.3 PRODUCT INFORMATION

1.3.1 Summary of Product Characteristics (SmPC)

1. Name of the Medicinal Product

Ketoconazole Cream BP 2 % W/W

"Prescription Only Medicine"

2. Qualitative and Quantitative Composition

Each tube contains:

Ketoconazole BP…….2% W/W

Cream Base……..…..Q.S.

3. Pharmaceutical Form

Nature of the Product: Cream.

Description: A white to off white gelatinous homogenous mass.

Primary Packaging Material: 15 gm Lami-tube

4. Clinical Particulars

4.1 Therapeutic Indications

For the treatment of the following mycotic infections of the skin: tinea pedis, tinea cruris and candidal intertrigo.

4.2 Posology and Method of Administration

Ketoconazole cream is for use in adults.

For the treatment of tinea pedis (athlete's foot) and tinea cruris (dhobie itch) and candidal intertrigo (sweat rash).

Tinea cruris, candidal intertrigo and tinea pedis: It is recommended that Ketoconazole 2% Cream be applied once or twice daily to cover the affected and immediate surrounding area.

The usual duration of treatment is tinea cruris 2-4 weeks, candidal intertrigo 2-4 weeks, tinea pedis 4-6 weeks.

Treatment should be continued, until a few days after disappearance of all symptoms. The diagnosis should be reconsidered if no clinical improvement is noted after 4 weeks of treatment.

Method of administration: Cutaneous use.

Paediatric patients

The safety and efficacy of Ketoconazole 2% Cream in children (17 years and younger) has not been established.

4.3 Contraindications

Hypersensitivity to the active substance or to any of the excipients.

4.4 Special Warnings and Precautions for Use

Ketoconazole 2% cream is not for ophthalmic use.

To prevent a rebound effect after stopping a prolonged treatment with topical corticosteroids it is recommended to continue applying a mild topical corticosteroid in the morning and to apply Ketoconazole 2% cream in the evening, and to subsequently and gradually withdraw the steroid therapy over a period of 2-3 weeks.

This medicine contains cetyl alcohol and stearyl alcohol which may cause local skin reactions (e.g. contact dermatitis). Also contains propylene glycol which may cause skin irritation.

4.5 Interaction with Other Medicinal Products and Other Forms of Interaction

None known.

4.6 Pregnancy and Lactation

There are no adequate and well-controlled studies in pregnant or lactating women. To date, no other relevant epidemiological data are available. Data on a limited number of exposed pregnancies indicate no adverse effects of topical ketoconazole on pregnancy or on the health of the foetus/newborn child. Animal studies have shown reproductive toxicity at doses that are not relevant to the topical administration of ketoconazole.

Plasma concentrations of ketoconazole are not detectable after topical application of Ketoconazole 2% cream to the skin of non-pregnant humans. There are no known risks associated with the use of Ketoconazole 2% cream in pregnancy or lactation.

4.7 Effects on Ability to Drive and Use Machines

This medicine has no influence on the ability to drive and use machines.

4.8 Undesirable Effects

The safety of ketoconazole cream was evaluated in 1079 subjects who participated in 30 clinical trials. Ketoconazole cream was applied topically to the skin.

Based on pooled safety data from these clinical trials, the most commonly reported (≥1% incidence) ADRs were (with % incidence): application site pruritus (2%), skin burning sensation (1.9%), and application site erythema (1%). Including the above-mentioned adverse drug reactions (ADRs), the following table displays ADRs that have been reported with the use of ketoconazole cream from either clinical trial or postmarketing experiences. The displayed frequency categories use the following convention:

Very Common: (≥1/10)

Common: (≥1/100 to <1/10)

Uncommon: (≥1/1,000 to <1/100)

Rare: (≥1/10,000 to <1/1,000)

Very rare: (<1/10,000)

Not Known: (cannot be estimated from the available clinical trial data)

System Organ Class	**Adverse Drug Reactions**		
	Frequency Category		
	Common (≥1/100 to <1/10)	**Uncommon** (≥1/1,000 to <1/100)	**Not Known**
Immune System Disorders		Hypersensitivity	
Skin and Subcutaneous Tissue Disorders	Skin burning sensation	Bullous eruption Dermatitis contact Rash Skin exfoliation Sticky skin	Urticaria

General Disorders and Administration Site Conditions	Application site erythema Application site pruritus	Application site bleeding Application site discomfort Application site dryness Application site inflammation Application site irritation Application site paraesthesia Application site reaction	

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.

4.9 Overdose

Topical application

Excessive topical application may lead to erythema, oedema and a burning sensation, which will disappear upon discontinuation of the treatment.

Ingestion

In the event of accidental ingestion, supportive and symptomatic measures should be carried out.

5. Pharmacological Properties

5.1 Pharmacodynamic Properties

Pharmacotherapeutic Group: Imidazole and triazole derivatives;

ATC code: D01 AC08

Ketoconazole has a potent antimycotic action against dermatophytes and yeasts. Ketoconazole cream acts rapidly on the pruritus, which is commonly seen in dermatophyte and yeast infections. This symptomatic improvement often occurs before the first signs of healing are observed.

A study in 250 patients has shown that application twice daily for 7 days of ketoconazole 2% cream vs clotrimazole 1% cream for 4 weeks on both feet demonstrated efficacy in patients with tinea pedis (athlete's foot) presenting lesions between the toes.

The primary efficacy endpoint was negative microscopic KOH examination at 4 weeks. Ketoconazole 2% treatment showed equivalent efficacy to 4 weeks clotrimazole 1% treatment. There was no evidence of relapse following treatment with ketoconazole cream at 8 weeks.

5.2 Pharmacokinetic Properties

Plasma concentrations of ketoconazole were not detectable after topical administration of ketoconazole cream in adults on the skin. In one study in infants with seborrhoeic dermatitis (n = 19), where approximately 40 g of ketoconazole cream was applied daily on 40 % of the body surface area, plasma levels of ketoconazole were detected in 5 infants, ranging from 32 to 133 ng/mL.

5.3 Preclinical Safety Data

Effects in non-clinical studies were observed only at exposures considered sufficiently in excess of the maximum human exposure indicating little relevance to clinical use.

6. Pharmaceutical Particulars

6.1 List of Excipients

Active Substance:

Sr. No.	Materials	Pharma Status	Quantity/ Batch (In kg)
1.	Ketoconazole	BP	6.000

Other Excipients:

Sr. No.	Materials	Pharma Status	Quantity/ Batch (In kg)
1.	Cetostearyl Alcohol	USP	12.000
2.	Self-Emulsifying Glyceryl Monostearate	BP	12.000
3.	Cetomacrogol Emulsifying Wax	BP	6.000
4.	Silicone Oil	IHS	3.000
5.	White Soft Paraffin	BP	45.000
6.	Light Liquid Paraffin	BP	18.000
7.	Methyl Paraben	BP	0.480
8.	Propyl Paraben	BP	0.240
9.	Propylene Glycol	BP	24.000
10.	Disodium Edetate	BP	0.300
11.	Sodium Acid Phosphate	BP	0.300
12.	Sodium Phosphate	BP	0.300
13.	Purified Water	IHS	175.828 (172.380 + 2%)

6.2 Incompatibilities

Not applicable.

6.3 Shelf Life

36 months

6.4 Special Precautions for Storage

Do not store above 25°C.

6.5 Nature and Contents of Container

Pack Size: 15 gm Lamitube.

Description: 1 NITOZOL CREAM (15 gm Lamitube) packed in inner carton with insert.

6.6 Special Precautions for Disposal and Other Handling

No Special requirements.

Any unused medicinal products or waste material should be disposed of in accordance with local requirements.

7. Marketing Authorisation Holder

Berlin Pharma and Health care Ltd.

No 42, Comfort Oboh Kiri Kiri Industry Area,

Apapa-Lagos, Nigeria.

8. Marketing Authorisation Number(S)

NF-PP-434666

9. Date of First Authorisation/Renewal of the Authorisation

06.2024

10. Date of Revision of the Text

05.2025