ERTACZO- sertaconazole nitrate cream
Valeant Pharmaceuticals North America LLC

HIGHLIGHTS OF PRESCRIBING INFORMATION
These highlights do not include all the information needed to use ERTACZO cream safely and effectively. See full prescribing information for ERTACZO cream.

ERTACZO® (sertaconazole nitrate) cream, 2%, for topical use

Initial U.S. Approval: 2003

INDICATIONS AND USAGE

ERTACZO cream 2% is an azole antifungal indicated for the topical treatment of interdigital tinea pedis in immunocompetent patients 12 years of age and older, caused by: *Trichophyton rubrum, Trichophyton mentagrophytes,* and *Epidermophyton floccosum.* (1)

DOSAGE AND ADMINISTRATION

ERTACZO cream should be applied to the affected and immediate surrounding area(s) twice daily for 4 weeks. (2)
Not for ophthalmic, oral, or intravaginal use. (2)

DOSAGE FORMS AND STRENGTHS

Cream, 2%. (3)

CONTRAINDICATIONS

None. (4)

ADVERSE REACTIONS

Most common adverse reactions observed in clinical trials (incidence >2%) were contact dermatitis, dry skin, burning skin, application site skin tenderness. (6.1)
To report SUSPECTED ADVERSE REACTIONS, contact Valeant Pharmaceuticals North America LLC at 1-800-321-4576 or FDA at 1-800-FDA-1088 or www.fda.gov/medwatch.
See 17 for PATIENT COUNSELING INFORMATION and FDA-approved patient labeling.

Revised: 1/2014

FULL PRESCRIBING INFORMATION: CONTENTS*

* Sections or subsections omitted from the full prescribing information are not listed.

FULL PRESCRIBING INFORMATION

1INDICATIONS AND USAGE

ERTACZO® (sertaconazole nitrate) cream, 2%, is indicated for the topical treatment of interdigital tinea pedis in immunocompetent patients 12 years of age and older, caused by: *Trichophyton rubrum*, *Trichophyton mentagrophytes*, and *Epidermophyton floccosum* [see *Clinical Studies (14)*].

2DOSAGE AND ADMINISTRATION

In the treatment of interdigital tinea pedis, ERTACZO cream, 2%, should be applied twice daily for 4 weeks. Sufficient amount of ERTACZO cream, 2%, should be applied to cover both the affected areas between the toes and the immediately surrounding healthy skin of patients with interdigital tinea pedis.

Not for ophthalmic, oral, or intravaginal use.

3DOSAGE FORMS AND STRENGTHS

Cream, 2%. Each gram of ERTACZO cream, 2%, contains 17.5 mg of sertaconazole (as sertaconazole nitrate, 20 mg) in a white cream base.

4CONTRAINDICATIONS

None.

5WARNINGS AND PRECAUTIONS

5.1Local Adverse Reactions

If irritation develops, treatment should be discontinued and appropriate therapy instituted.

Physicians should exercise caution when prescribing ERTACZO cream, 2%, to patients known to be sensitive to imidazole antifungals, since cross-reactivity may occur.

6ADVERSE REACTIONS

6.1Clinical Trials Experience

Because clinical trials are conducted under widely varying conditions, adverse reaction rates observed in the clinical trials of a drug cannot be directly compared to rates in the clinical trials of another drug, and may not reflect the rates observed in practice.

In clinical trials, cutaneous adverse events occurred in 7 of 297 (2%) subjects (2 of them severe) receiving ERTACZO® cream, 2%, and in 7 of 291 (2%) subjects (2 of them severe) receiving vehicle. These reported cutaneous adverse events included contact dermatitis, dry skin, burning skin, application site skin tenderness.

In a dermal sensitization trial, 8 of 202 evaluable subjects tested with ERTACZO® cream, 2%, and 4 of 202 evaluable subjects tested with vehicle, exhibited a slight erythematous reaction in the challenge phase. There was no evidence of cumulative irritation or contact sensitization in a repeated insult patch test involving 202 healthy volunteers.

6.2Postmarketing Experience

The following adverse reactions have been identified during post-approval use of ERTACZO cream, 2%. Because these reactions are reported voluntarily from a population of uncertain size, it is not always possible to reliably estimate their frequency or establish a causal relationship to drug exposure.

In post-marketing surveillance for ERTACZO cream, 2%, the following were reported:

Cutaneous adverse events: erythema, pruritus, vesiculation, desquamation, and hyperpigmentation.

8USE IN SPECIFIC POPULATIONS

8.1Pregnancy

Pregnancy Category C.

There are no adequate and well-controlled studies conducted with ERTACZO cream in pregnant women. ERTACZO cream should be used during pregnancy only if the potential benefit justifies the potential risk to the fetus.

Reproduction studies have not been performed with ERTACZO cream. Sertaconazole nitrate did not produce any evidence of maternal toxicity, embryotoxicity or teratogenicity in rats and rabbits at an oral dose of 160 mg/kg/day (40 times (rats) and 80 times (rabbits) the maximum recommended human dose based on a body surface area comparison). A reduction in live birth indices and an increase in the number of still-born pups were seen at doses of 80 and 160 mg/kg/day sertaconazole nitrate in an oral peri- and post-natal development study in rats.

8.3Nursing Mothers

It is not known if sertaconazole is excreted in human milk. Because many drugs are excreted in human milk, caution should be exercised when prescribing ERTACZO cream, 2%, to a nursing woman.

8.4Pediatric Use

The efficacy and safety of ERTACZO cream, 2%, have not been established in pediatric patients below the age of 12 years.

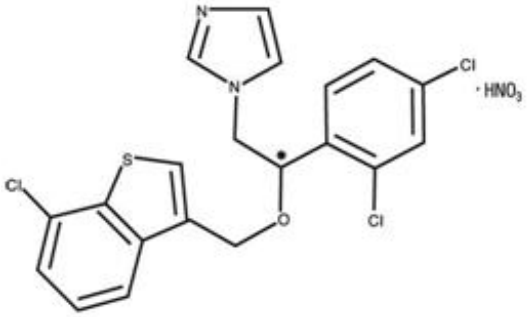
8.5Geriatric Use

Clinical trials of ERTACZO cream, 2%, did not include sufficient numbers of subjects aged 65 and over to determine whether they respond differently from younger subjects.

11DESCRIPTION

ERTACZO (sertaconazole nitrate) cream, 2%, is for topical application. It contains the azole antifungal, sertaconazole nitrate. Sertaconazole nitrate contains one asymmetric carbon atom and exists as a racemic mixture of equal amounts of R and S enantiomers.

Sertaconazole nitrate is designated chemically as (±)-1-[2,4-dichloro-β-[(7-chlorobenzo-[*b*]thien-3-yl)methoxy]phenethyl]imidazole nitrate. It has a molecular weight of 500.8. The molecular formula is $C_{20}H_{15}Cl_3N_2OS \cdot HNO_3$, and the structural formula is as follows:



Sertaconazole nitrate is a white or almost white powder. It is practically insoluble in water, soluble in methanol, sparingly soluble in alcohol and in methylene chloride. Each gram of ERTACZO cream, 2%, contains 17.5 mg of sertaconazole (as sertaconazole nitrate, 20 mg) in a white cream base of ethylene glycol, glyceryl isostearate, glycolized saturated glycerides, light mineral oil, methylparaben, polyethylene glycol palmitostearate, polyoxyethylened saturated glycerides, sorbic acid and purified water.

12CLINICAL PHARMACOLOGY

12.1Mechanism of Action

ERTACZO cream is an azole antifungal [see *Clinical Pharmacology (12.4)*].

12.3Pharmacokinetics

In a multiple dose pharmacokinetic trial that included 5 male subjects with interdigital tinea pedis (range of diseased area, 42 - 140 cm^2; mean, 93 cm^2), ERTACZO cream, 2%, was topically applied every 12 hours for a total of 13 doses to the diseased skin (0.5 grams sertaconazole nitrate per 100 cm^2). Sertaconazole concentrations in plasma measured by serial blood sampling for 72 hours after the thirteenth dose were below the limit of quantitation (2.5 ng/mL) of the analytical method used.

12.4Microbiology

Mechanism of Action:
Sertaconazole, an azole antifungal agent, inhibits fungal cytochrome P-450-mediated 14 alpha-lanosterol demethylase enzyme. This enzyme functions to convert lanosterol to ergosterol. Ergosterol is a key component of fungal cell membranes and lack of this component leads to fungal cell injury by leakage of key constituents in the cytoplasm from the cell.

Activity *In Vitro* and in Clinical Infections:
Sertaconazole nitrate has been shown to be active against isolates of the following microorganisms in clinical infections [see *Indications and Usage (1)*].

Trichophyton rubrum
Trichophyton mentagrophytes
Epidermophyton floccosum

13NONCLINICAL TOXICOLOGY

13.1Carcinogenesis, Mutagenesis, Impairment of Fertility

In a rat dermal carcinogenicity study, topical administration of sertaconazole nitrate cream for up to 102 weeks did not increase the number of neoplastic lesions compared to control animals, at sertaconazole nitrate doses of up to 800 mg/kg/day (approximately 200 times the maximum recommended human dose

based on a body surface area comparison).

No clastogenic potential was observed in a mouse micronucleus test. Sertaconazole nitrate was considered nonclastogenic in the *in vivo* mouse sister chromatid exchange assay. There was no evidence that sertaconazole nitrate induced unscheduled DNA synthesis in primary rat hepatocyte cultures.

At oral doses up to 60 mg/kg/day (16 times the maximum recommended human dose based on a body surface area comparison), sertaconazole nitrate exhibited no toxicity or adverse effects on reproductive performance or fertility in male or female rats.

14CLINICAL STUDIES

In two randomized, double-blind, clinical trials, subjects 12 years and older with interdigital tinea pedis applied either ERTACZO cream, 2%, or vehicle, twice daily for four weeks. Subjects with moccasin-type (plantar) tinea pedis and/or onychomycosis were excluded from the trial. Two weeks after completion of therapy (six weeks after beginning therapy), subjects were evaluated for signs and symptoms related to interdigital tinea pedis.

Treatment outcomes are summarized in the table below.

Treatment Outcomes as Percent (%) of Total Subjects

	Trial 1		**Trial 2**	
	Sertaconazole	**Vehicle**	**Sertaconazole**	**Vehicle**
Complete Cure[*] (Primary Efficacy Variable)	13/99 (13.1%)	3/92 (3.3%)	28/103 (27.2%)	5/103 (4.9%)
Effective Treatment[†]	32/99 (32.3%)	11/92 (12.0%)	52/103 (50.5%)	16/103 (15.5%)
Mycological Cure[‡]	49/99 (49.5%)	18/92 (19.6%)	71/103 (68.9%)	20/103 (19.4%)

[*] **Complete Cure – Subjects who had complete clearing of signs and symptoms and Mycological Cure.**

[†] **Effective Treatment – Subjects who had minimal residual signs and symptoms of interdigital tinea pedis and Mycological Cure.**

[‡] **Mycological Cure – Subjects who had both negative microscopic KOH preparation and negative fungal culture.**

In clinical trials, complete cure in sertaconazole treated subjects was achieved in 32 of 160 (20%) subjects with *Trichophyton rubrum*, in 7 of 28 (25%) subjects with *Trichophyton mentagrophytes* and in 1 of 13 (15%) subjects with *Epidermophyton floccosum*.

16HOW SUPPLIED/STORAGE AND HANDLING

ERTACZO cream, 2%, is white in color and supplied in tubes in the following size:

60-gram tube	NDC 0187-5115-60

Store at 20°C-25°C (68°F-77°F); excursions permitted to 15°-30°C (59°-86°F) [see USP Controlled Room Temperature].

17PATIENT COUNSELING INFORMATION

See FDA-approved Patient Labeling (Patient Information)

The patient should be instructed to:

- Use ERTACZO cream, 2%, as directed by the physician. The hands should be washed after applying the medication to the affected area(s). Avoid contact with the eyes, mouth, vagina and other mucous membranes. ERTACZO cream, 2%, is for external use only.
- Dry the affected area(s) thoroughly before application, if you wish to use ERTACZO cream, 2%, after bathing.
- Use the medication for the full treatment time recommended by the physician, even though symptoms may have improved.
- Inform the physician if the area of application shows signs of increased irritation, redness, itching, burning, blistering, swelling or oozing.
- Avoid the use of occlusive dressings unless otherwise directed by the physician.
- Do not use this medication for any disorder other than that for which it was prescribed.

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Rev. 01/14

PATIENT INFORMATION

ERTACZO (er-tack-zo)
(sertaconazole nitrate) cream, 2%

Important information: ERTACZO cream is for use on skin only. Do not use ERTACZO cream in your eyes, mouth, or vagina.

What is ERTACZO cream?

ERTACZO cream is a prescription medicine used on the skin (topical) to treat athlete's foot that is between the toes (interdigital tinea pedis) in people 12 years of age and older with normal immune systems.

It is not known if ERTACZO cream is safe and effective in children under 12 years of age.

What should I tell my healthcare provider before using ERTACZO cream?

Before using ERTACZO cream, tell your healthcare provider about all of your medical conditions, including if you:

- have any allergies
- are pregnant or plan to become pregnant. It is not known if ERTACZO cream will harm your unborn baby.
- are breastfeeding or plan to breastfeed. It is not known if ERTACZO cream passes into your breast milk.

Tell your healthcare provider about all of the medicines you take, including prescription and over-the-counter medicines, vitamins, and herbal supplements.

How should I use ERTACZO cream?

- Use ERTACZO cream exactly as your healthcare provider tells you to use it.
- Use ERTACZO cream for the full treatment time, even if your symptoms improve.
- If you take a bath or shower, dry the affected skin areas well before you apply ERTACZO cream.
- Apply ERTACZO cream 2 times a day for 4 weeks to the affected skin areas between your toes

and to the healthy skin around the affected areas.
- Wash your hands after you apply ERTACZO cream
- Do not cover the treated skin areas with bandages unless your healthcare provider tells you to.

What are the possible side effects of ERTACZO cream?

The most common side effects of ERTACZO cream include: redness, itching, dry skin, burning, blistering, swelling, drainage and skin tenderness at the treated skin areas. Tell your healthcare provider if you have any of these skin reactions.

These are not all the possible side effects of ERTACZO cream. Call your doctor for medical advice about side effects. You may report side effects to FDA at 1-800-FDA-1088.

How should I store ERTACZO cream?

- Store ERTACZO cream at room temperature, between 68°F to 77°F (20°C to 25°C).
- Keep ERTACZO cream and all medicines out of the reach of children.

General information about the safe and effective use of ERTACZO cream

Medicines are sometimes prescribed for purposes other than those listed in a Patient Information leaflet. You can ask your healthcare provider or pharmacist for information about ERTACZO cream that is written for health professionals. Do not use ERTACZO cream for a condition for which it was not prescribed. Do not give ERTACZO cream to other people, even if they have the same symptoms you have. It may harm them.

What are the ingredients in ERTACZO cream?

Active ingredient: sertaconazole nitrate

Inactive ingredients: ethylene glycol, glyceryl isostearate, glycolized saturated glycerides, light mineral oil, methylparaben, polyethylene glycol palmitostearate, polyoxyethylened saturated glycerides, sorbic acid and purified water

Distributed by: Valeant Pharmaceuticals North America LLC, Bridgewater, NJ 08807

Manufactured by: DPT Laboratories, Ltd., San Antonio, TX 78215

Product of Spain

For more information call 1-800-321-4567.

This Patient Information has been approved by the U.S. Food and Drug Administration.

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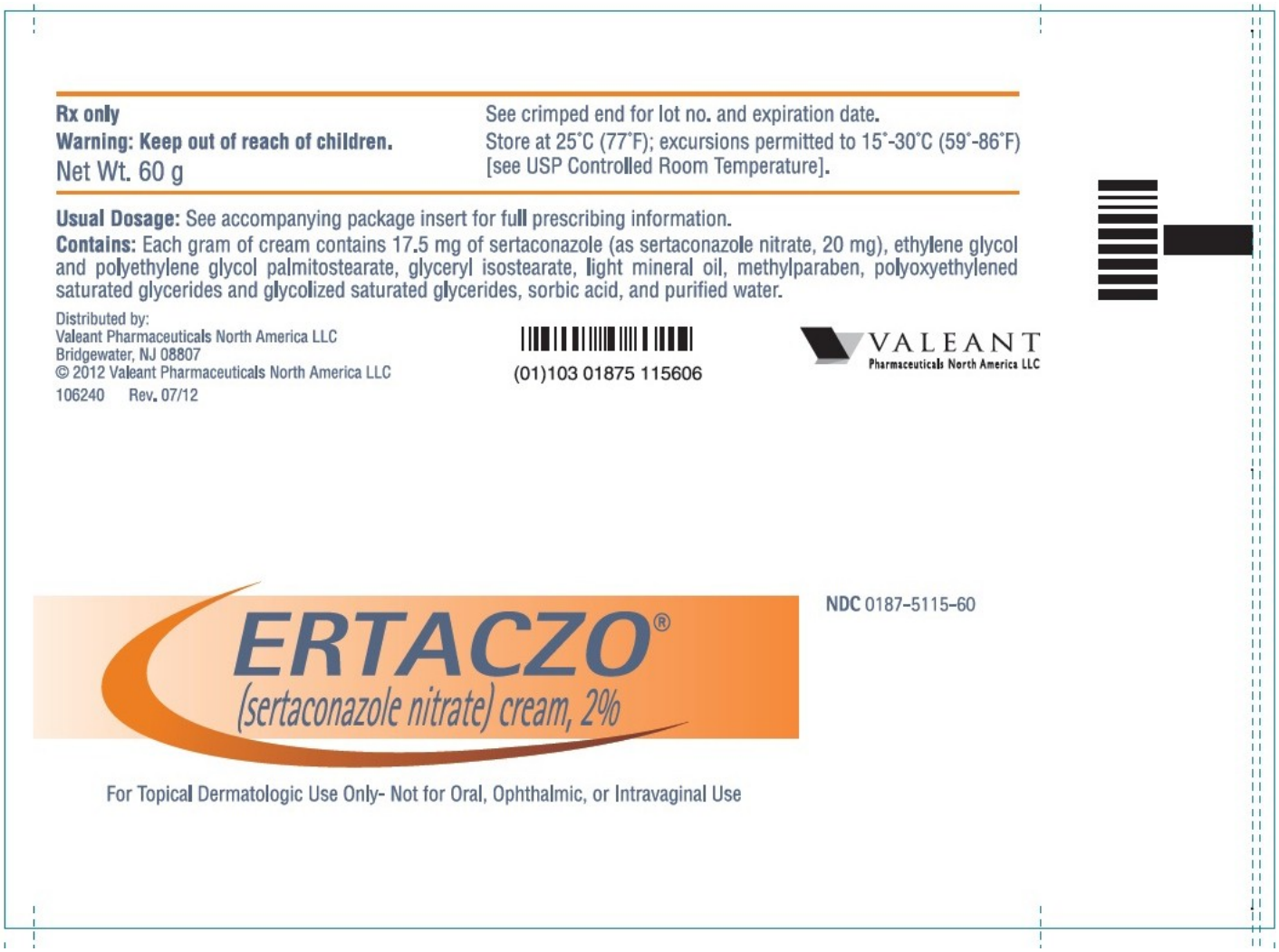
Rev. 01/14

PRINCIPAL DISPLAY PANEL - 60 g Tube

NDC 0187-5115-60

ERTACZO®
(sertaconazole nitrate) cream, 2%

For Topical Use Only – Not for Oral, Ophthalmic, or Intravaginal Use



ERTACZO

sertaconazole nitrate cream

Product Information			
Product Type	HUMAN PRESCRIPTION DRUG	**Item Code (Source)**	NDC:0187-5115
Route of Administration	TOPICAL		

Active Ingredient/Active Moiety		
Ingredient Name	**Basis of Strength**	**Strength**
Sertaconazole Nitrate (UNII: 1DV05410M5) (Sertaconazole - UNII:72W71I16EG)	Sertaconazole Nitrate	20 mg in 1 g

Inactive Ingredients	
Ingredient Name	**Strength**
Ethylene Glycol (UNII: FC72KVT52F)	
Glyceryl Isostearate (UNII: HYE7O27HAO)	
Light Mineral Oil (UNII: N6K5787QVP)	
Methylparaben (UNII: A2I8C7HI9T)	
Sorbic Acid (UNII: X045WJ989B)	

Water (UNII: 059QF0KO0R)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0187-5115-60	1 in 1 CARTON		
1		60 g in 1 TUBE		
2	NDC:0187-5115-02	18 in 1 CARTON		
2		2 g in 1 TUBE		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
NDA	NDA021385	12/10/2003	

Labeler - Valeant Pharmaceuticals North America LLC (042230623)

Establishment

Name	Address	ID/FEI	Business Operations
DPT Laboratories, Ltd.		832224690	MANUFACTURE(0187-5115)

Revised: 1/2014 Valeant Pharmaceuticals North America LLC