

Data Sheet

CANESTEN™

Canesten Clotrimazole Thrush Treatment Once Cream, vaginal cream 100 mg/g (10%)

Canesten Clotrimazole Thrush Treatment 3 Day Cream, vaginal cream 20 mg/g (2%)

Canesten Clotrimazole Thrush Treatment 6 Day Cream, vaginal cream 10 mg/g (1%)

Canesten Clotrimazole Thrush Treatment 6 Day Pessary, vaginal pessary 100 mg

Canesten Clotrimazole Thrush Treatment Once Pessary, vaginal pessary 500 mg

Canesten Clotrimazole Thrush Treatment Once Pessary + Cream, vaginal pessary 500 mg + topical cream 10 mg/g (1%)

Canesten Clotrimazole Thrush Treatment Once Soft Vaginal Pessary, soft vaginal pessary 500 mg (not marketed)

Canesten Clotrimazole Thrush Treatment Once Soft Vaginal Pessary + Cream, soft vaginal pessary 500 mg + topical cream 10 mg/g (1%) (not marketed)

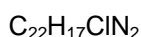
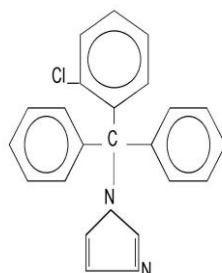
Canesten Topical Cream 10 mg/g (1%)

Canesten Topical Solution 10 mg/mL (1%)

Name of Medicine

Clotrimazole

1-(o-chloro- α, α - diphenylbenzyl) imidazole



Molecular Weight 344.84

Clotrimazole is a colourless, crystalline, weakly alkaline substance, melting point 141°- 145°C, soluble in acetone, chloroform and ethanol and practically insoluble in water. It forms stable salts with both organic and inorganic acids. It is not photosensitive but is slightly hygroscopic, and may be hydrolysed in acid media.

Qualitative and Quantitative Composition

Canesten Clotrimazole Thrush Treatment Once Cream

Canesten Clotrimazole Thrush Treatment Once Cream contains 100 mg/g (10% w/w) of clotrimazole in a vanishing cream base. The cream also contains sorbitan stearate, polysorbate 60, cetyl palmitate, cetostearyl alcohol, isopropyl myristate, benzyl alcohol (1% w/w) and purified water.

Canesten Clotrimazole Thrush Treatment 3 Day Cream

Canesten Clotrimazole Thrush Treatment 3 Day Cream contains 20 mg/g (2% w/w) of clotrimazole in a vanishing cream base. The cream also contains sorbitan stearate, polysorbate 60, cetyl palmitate, cetostearyl alcohol, octyldodecanol, benzyl alcohol (2% w/w) and purified water.

Canesten Clotrimazole Thrush Treatment 6 Day Cream

Canesten Clotrimazole Thrush Treatment 6 Day Cream contains 10 mg/g (1% w/w) of clotrimazole in a vanishing cream base. The cream also contains sorbitan stearate, polysorbate 60, cetyl palmitate, cetostearyl alcohol, octyldodecanol, benzyl alcohol (2% w/w) and purified water.

Canesten Clotrimazole Thrush Treatment 6 Day Pessary

Canesten Clotrimazole Thrush Treatment 6 Day Pessary contains 100 mg of clotrimazole in each pessary. The pessaries also contain lactose, maize starch, magnesium stearate, silicon dioxide, calcium lactate pentahydrate, crospovidone, lactic acid, hypromellose, microcrystalline cellulose.

Canesten Clotrimazole Thrush Treatment Once Pessary

Canesten Clotrimazole Thrush Treatment Once Pessary contains 500 mg of clotrimazole in each pessary. The pessaries also contain lactose, microcrystalline cellulose, lactic acid, maize starch, crospovidone, calcium lactate; magnesium stearate, silicon dioxide, hypromellose.

Canesten Clotrimazole Thrush Treatment Once Pessary + Cream

Canesten Clotrimazole Thrush Treatment Once Pessary + Cream contains 500 mg of clotrimazole in each pessary and 10 mg/g clotrimazole in the cream. The pessary also contains lactose, microcrystalline cellulose, lactic acid, maize starch, crospovidone, calcium lactate; magnesium stearate, silicon dioxide, hypromellose. The cream also contains sorbitan stearate, polysorbate 60, cetyl palmitate, cetostearyl alcohol, octyldodecanol, benzyl alcohol (2% w/w) and purified water.

Canesten Clotrimazole Thrush Treatment Once Soft Vaginal Pessary (not marketed)

Canesten Clotrimazole Thrush Treatment Once Soft Vaginal Pessary contains 500 mg of clotrimazole in each soft vaginal pessary. The soft pessary also contains white, soft paraffin, liquid paraffin, gelatin, glycerol, titanium dioxide, quinoline yellow, sunset yellow, and traces of purified water, lecithin and fractionated coconut oil.

Canesten Clotrimazole Thrush Treatment Once Soft Vaginal Pessary + Cream (not marketed)

Canesten Clotrimazole Thrush Treatment Once Soft Vaginal Pessary + Cream contains 500 mg of clotrimazole in each soft vaginal pessary and 10 mg/g clotrimazole in the cream. The soft pessary also contains white, soft paraffin, liquid paraffin, gelatin, glycerol, titanium dioxide, quinoline yellow, sunset yellow, and traces of purified water, lecithin and fractionated coconut oil. The cream also contains sorbitan stearate, polysorbate 60, cetyl palmitate, cetostearyl alcohol, octyldodecanol, benzyl alcohol (2% w/w) and purified water.

Canesten Topical Cream

Canesten topical cream contains 10 mg/g (1%) of clotrimazole in a vanishing cream base. The cream also contains sorbitan stearate, polysorbate 60, cetyl palmitate, cetostearyl alcohol, octyldodecanol, benzyl alcohol (2% w/w) and purified water.

Canesten Topical Solution

Canesten topical solution contains 10 mg/mL (1% w/v) of clotrimazole in a solution of propylene glycol, isopropyl alcohol and polyethylene glycol.

Pharmaceutical Form

Canesten Clotrimazole Thrush Treatment Once Cream

Vaginal cream.

Canesten Clotrimazole Thrush Treatment 3 Day Cream

Vaginal cream.

Canesten Clotrimazole Thrush Treatment 6 Day Cream

Vaginal cream.

Canesten Clotrimazole Thrush Treatment 6 Day Pessary

Vaginal pessary.

Canesten Clotrimazole Thrush Treatment Once Pessary

Vaginal pessary.

Canesten Clotrimazole Thrush Treatment Once Pessary + Cream

Combination product – vaginal pessary and topical cream.

Canesten Clotrimazole Thrush Treatment Once Soft Vaginal Pessary (not marketed)

Vaginal soft gel pessary.

Canesten Clotrimazole Thrush Treatment Once Soft Vaginal Pessary + Cream (not marketed)

Combination product – vaginal soft gel pessary and topical cream.

Canesten Topical Cream

Topical cream.

Canesten Topical Solution

Topical solution.

Clinical Particulars

Therapeutic Indications

Canesten Clotrimazole Thrush Treatment Once, 3 Day and 6 Day creams are indicated for the topical treatment of vulvovaginal candidiasis. Canesten Clotrimazole cream may also be used in conjunction with Canesten Clotrimazole vaginal pessaries in the management of *Candida* vulvovaginitis or infection of the peri-anal area while application of the cream to the glans penis of the partner may help prevent re-infection of the female.

Canesten Clotrimazole Thrush Treatment Once and 6 Day vaginal pessaries are indicated for the topical treatment of vaginal candidiasis.

Canesten Clotrimazole Thrush Treatment Once Pessary + Cream is indicated for the topical treatment of vulvovaginal candidiasis. The cream can also be used in the management of *Candida* vulvovaginitis or infection of the peri-anal area while application of the cream to the glans penis of the partner may help prevent re-infection of the female.

Canesten topical cream and topical solution are indicated for the topical treatment of the following dermal infections:

1. Tinea pedis (athlete's foot), tinea cruris (jock itch) and tinea corporis (ringworm of the trunk and limbs) due to *Trichophyton rubrum*, *Trichophyton mentagrophytes*, *Epidermophyton floccosum* and *Microsporum canis*.
2. Candidiasis due to *Candida albicans* including cutaneous candidiasis, onychia (infection of the nail or nail bed), paronychia (infection of the nail fold), external genital candidiasis, candida balanitis (*Candida* in the male genital area), nappy rash.
3. Pityriasis versicolor due to *Malassezia furfur*.
4. Erythrasma (a chronic skin infection due to the bacterium *Corynebacterium minutissimum*).

Posology and Method of Administration

Canesten Clotrimazole Thrush Treatment 6 Day Cream

Once daily, preferably in the evening for six successive days, one applicator should be filled with cream (approx. 5 g) and inserted as deeply as possible into the vagina with the patient lying on her back. The 35 g tube of cream for vaginal use provides for six such doses.

Canesten Clotrimazole Thrush Treatment 3 Day Cream

Once daily, preferably in the evening for three successive days, one applicator should be filled with cream (approx. 5 g) and inserted as deeply as possible into the vagina with the patient lying on her back. The 20 g tube of cream for vaginal use provides for three such doses.

Canesten Clotrimazole cream may also be used in conjunction with Canesten Clotrimazole vaginal pessaries in the management of *Candida* vulvovaginitis or infection of the peri-anal area while application of the cream to the glans penis of the partner may help prevent re-infection of the female.

Canesten Clotrimazole Thrush Treatment Once Cream

The disposable applicator should be filled with Canesten cream, ensuring the entire contents of the tube are used (approx. 5 g). The cream is then inserted as gently and deeply as possible into the vagina with the patient lying on her back at bedtime as a single dose of treatment.

Canesten Clotrimazole Thrush Treatment Vaginal Pessaries

The pessaries should be inserted as deeply as possible into the vagina once daily, preferably in the evening. This is best achieved using the plastic applicator provided and following the directions on the patient instruction sheet. In pregnancy digital insertion may be preferable to use of the applicator.

A course of treatment normally consists of either a single 500 mg pessary (Canesten Clotrimazole Thrush Treatment Once Pessary) or of six 100 mg pessaries (Canesten Clotrimazole Thrush Treatment 6 Day Pessary). The latter may be given either as two pessaries, inserted one after the other, daily for three days or as one pessary daily for six days. Clinical investigations have shown comparable efficacy from either dosage scheme. Where a first course proved unsuccessful, a second course produced success in 8 of 12 women treated.

It is suggested that treatment be timed so as to avoid the menstrual period and to be complete before the onset of menstruation.

Canesten Topical Cream and Solution

Apply sparingly to the affected areas and rub in gently, two or three times daily.

Duration of Treatment

The following recommendations are made:

Cutaneous candidiasis	2 weeks
Dermatomycoses	2 - 4 weeks
Onychia and Paronychia due to <i>C. albicans</i>	4 - 8 weeks or more

Regular application of Canesten topical cream or solution is essential for successful treatment and, whether or not a cure is confirmed mycologically, treatment should be continued for two weeks after all clinical signs have disappeared.

Special Remarks

Attention to hygiene is important in the management of fungal diseases of the feet. After washing, the feet - especially between the toes - should be dried thoroughly. Canesten Clotrimazole cream may be useful in mycotic paronychia or onychia following removal of the nail.

Contraindications

Hypersensitivity to clotrimazole or to any other component of the product.

Hypersensitivity to cetostearyl alcohol (a component of all cream presentations).

Special Warnings and Precautions for Use

Clotrimazole creams may reduce the effectiveness and safety of latex products such as condoms and diaphragms when applied to the genital area (women: labia and adjacent area of the vulva; men: prepuce and glans of the penis). The effect is temporary and occurs only during treatment.

If the patient has a fever (temperature of 38°C or above), lower abdominal pain, back pain, foul smelling vaginal discharge, nausea, vaginal haemorrhage and/or associated shoulder pain the patient should consult a doctor.

Vaginal Treatments

If symptoms persist for more than 7 days, or do not improve within 4 days, since starting treatment the patient may have a medical condition that requires treatment by a doctor.

The treatment can be repeated if necessary, however recurrent infections may indicate an underlying medical cause, including diabetes or HIV infection. Patients should seek medical advice if they have had 3 or more infections within 6 months.

Treatment during the menstrual period should not be performed. The treatment should be finished before the onset of menstruation.

Do not use tampons, intravaginal douches, spermicides or other vaginal products while using this product.

Avoidance of vaginal intercourse is recommended while using this product because the infection could be transferred to your partner, and the effectiveness and safety of latex products such as condoms and diaphragms may be reduced.

For All Presentations

Keep the medicine out of reach of children.

Avoid contact with eyes.

Do not swallow.

For products containing cetostearyl alcohol i.e. the cream presentations:

Cetostearyl alcohol may cause local skin reactions (e.g. contact dermatitis).

Interactions with Other Medicinal Products and Other Forms of Interaction

Concomitant medication with vaginal clotrimazole and oral tacrolimus (FK-506 immunosuppressant) might lead to increased tacrolimus plasma levels. Patients should thus be thoroughly monitored for symptoms of tacrolimus overdose, if necessary by determination of the respective plasma levels.

Pregnancy and Lactation

Pregnancy (Category A)

While controlled clinical studies in pregnant women do not exist, epidemiological investigations give no indications that harmful effects on the mother and child should be anticipated when Canesten Clotrimazole is used during pregnancy. However, as with all medicines in the first three months of pregnancy, Canesten Clotrimazole should only be used after first consulting a doctor.

Sanitation of the birth canal should be ensured particularly during the last 4 – 6 weeks of pregnancy.

During pregnancy, the treatment should be carried out with Canesten Clotrimazole vaginal pessaries since these can be inserted without using an applicator.

Lactation

Because systemic absorption of clotrimazole following topical or vaginal application is marginal, there should be no risk for the infant during lactation.

Effects on Ability to Drive and Use Machines

No effects on ability to drive and use machines have been observed.

Undesirable Effects

The listed adverse reactions are based on spontaneous reports, thus an organisation according to CIOMS III categories of frequency is not possible.

Immune System Disorders

Allergic reactions (syncope, hypotension, dyspnoea, urticaria).

Skin and Subcutaneous Tissue Disorders

Blisters, discomfort/pain, oedema, irritation, peeling/exfoliation, pruritis, rash, stinging/burning.

Reproductive System and Breast Disorders

Genital peeling, pruritis, rash, oedema, discomfort, burning, irritation, pelvic pain.

Gastrointestinal Disorders

Abdominal pain.

Overdose

Not applicable.

Pharmacological Properties

Pharmacotherapeutic Group

Clotrimazole is an imidazole derivative with a broad spectrum antimycotic activity.

Mechanism of Action

Clotrimazole acts against fungi by inhibiting ergosterol synthesis. Inhibition of ergosterol synthesis leads to structural and functional impairment of the cytoplasmic membrane.

Pharmacodynamic Properties

Clotrimazole has a broad antimycotic spectrum of action in vitro and in vivo, which includes dermatophytes, yeasts, moulds, etc.

Under appropriate test conditions, the MIC values for these types of fungi are in the region of less than 0.062 – 8.0 µg/mL substrate.

The mode of action of clotrimazole is primarily fungistatic or fungicidal depending on the concentration of clotrimazole at the site of infection. In vitro activity is limited to proliferating elements; fungal spores are only slightly sensitive.

In addition to its antimycotic action, clotrimazole also acts on *Trichomonas vaginalis*, gram-positive micro-organisms (*Streptococci* / *Staphylococci*) and gram-negative micro-organisms (*Bacteroides* / *Gardnerella vaginalis*).

In vitro clotrimazole inhibits the multiplication of *Corynebacteria* and gram-positive cocci (with the exception of *Enterococci*) in concentrations of 0.5 – 10 µg/mL substrate and exerts a trichomonacidal action at 100 µg/mL.

Primarily resistant variants of sensitive fungal species are very rare; the development of secondary resistance by sensitive fungi has so far only been observed in very isolated cases under therapeutic conditions.

Pharmacokinetic Properties

Pharmacokinetic investigations after vaginal application have shown that only a small amount of clotrimazole (3 – 10%) is absorbed. Due to the rapid hepatic metabolism of absorbed clotrimazole into pharmacologically inactive metabolites the resulting peak plasma concentrations of clotrimazole after vaginal application of a 500 mg dose were less than 10 ng/mL, suggesting that clotrimazole applied intravaginally is unlikely to lead to measurable systemic effects or side effects.

Pharmacokinetic investigations after dermal application have shown that clotrimazole is minimally absorbed from the intact or inflamed skin into the human blood circulation. The resulting peak serum concentrations of clotrimazole were below the detection limit of 0.001 µg/mL, suggesting that clotrimazole applied topically on the skin is unlikely to lead to measurable systemic effects or side effects.

Preclinical Safety Data

Toxicological studies in different animals with intravaginal or local application showed good vaginal and local tolerability.

Preclinical data reveal no special hazard for humans based on conventional studies of single and repeated dose toxicity, genotoxicity and toxicity to reproduction.

Pharmaceutical Particulars

Incompatibilities

None known.

Special Precautions for Storage

Canesten Clotrimazole Thrush Treatment Once Soft Vaginal Pessaries (not marketed): Store at or below 30°C.

All other presentations: Store at or below 25°C.

Nature and Contents of Container

Canesten Clotrimazole Thrush Treatment Once Cream

One tube containing 5 g of vaginal cream, 100 mg clotrimazole per gram (10% w/w) packed with a single-use applicator and patient instruction sheet.

Canesten Clotrimazole Thrush Treatment 3 Day Cream

One tube containing 20 g of vaginal cream, 20 mg clotrimazole per gram (2% w/w) packed with three single-use disposable applicators and patient instruction sheet.

Canesten Clotrimazole Thrush Treatment 6 Day Cream

One tube containing 35 g of vaginal cream, 10 mg clotrimazole per gram (1% w/w) packed with six single-use disposable applicators and patient instruction sheet.

Canesten Clotrimazole Thrush Treatment 6 Day Pessary

Packs of six vaginal pessaries each sealed in a blister with plastic applicator and patient instruction sheet. Each pessary contains 100 mg clotrimazole per tablet.

Canesten Clotrimazole Thrush Treatment Once Pessary

One vaginal pessary sealed in a blister with plastic applicator and patient instruction sheet. Each pessary contains 500 mg clotrimazole.

Canesten Clotrimazole Thrush Treatment Once Pessary + Cream

One vaginal pessary sealed in a blister with plastic applicator, one tube containing 10 g topical cream and patient instruction sheet. Each pessary contains 500 mg clotrimazole. Each gram of cream contains 10 mg clotrimazole.

Canesten Clotrimazole Thrush Treatment Once Soft Vaginal Pessary (not marketed)

One soft vaginal pessary sealed in a blister with plastic applicator and patient instruction sheet. Each pessary contains 500 mg clotrimazole.

Canesten Clotrimazole Thrush Treatment Once Soft Vaginal Pessary + Cream (not marketed)

One soft vaginal pessary sealed in a blister with plastic applicator, one tube containing 10 g topical cream and patient instruction sheet. Each pessary contains 500 mg clotrimazole. Each gram of cream contains 10 mg clotrimazole.

Canesten Topical Cream

Packs of one tube containing 20 g or 50 g of cream, 10 mg clotrimazole per gram (1% w/w).

Canesten Topical Solution

Packs of one plastic dropper-type bottle containing 20 mL of solution, 10 mg of clotrimazole per mL (1% w/v).

Instructions for Use/Handling

Medicines should not be disposed of via wastewater or household waste. Ask a pharmacist how to dispose of medicines no longer required. These measures will help to protect the environment.

Medicine Classification

Pharmacist Only Medicines

Canesten Clotrimazole Thrush Treatment

- Once Cream
- 3 Day Cream
- 6 Day Cream
- Once Pessary
- 6 Day Pessaries
- Once Pessary + Cream
- Once Soft Vaginal Pessary (not marketed)
- Once Soft Vaginal Pessary + Cream (not marketed)

Pharmacy Medicines

Canesten

- Topical Cream
- Topical Solution

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Date of Preparation

6 January 2012

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