

FLISK[®] 150mg Capsules & 50mg/5mL Dry Suspension

[Fluconazole]

DESCRIPTION:

FLISK (Fluconazole) is a triazole antifungal agent.

QUALITATIVE AND QUANTITATIVE COMPOSITION:

FLISK (Fluconazole) is available for oral administration as:

1. FLISK Capsules 150mg

Each capsule contains:

Fluconazole USP 150mg

(Product Specs.: Sigma)

2. FLISK 50mg/5mL Dry Suspension

Each reconstituted 5mL contains:

Fluconazole USP 50mg

(Product Specs.: USP)

CLINICAL PHARMACOLOGY:

Mechanism of Action:

FLISK (Fluconazole) inhibits cytochrome P-450 dependent enzymes resulting in blockage of ergosterol synthesis and ultimately responsible for antifungal activity of fluconazole. Fluconazole has been shown to be more selective for fungal cytochrome P-450 enzymes than for various mammalian cytochrome P-450 enzyme systems.

Microbiology:

Fluconazole is active against:

- Blastomycosis dermatitidis.
- Candida species (except C. krusei).
- Cryptococcus neoformans.
- Coccidioides immitis.
- Epidermophyton Species.
- Histoplasma capsulatum.
- Microsporum Species.
- Trichophyton Species.

Pharmacokinetics:

Absorption:

FLISK is well absorbed following oral administration. Peak plasma concentrations reached in 1 to 2 hours of oral dose. Steady state levels are reached within 6 to 10 days following oral dose of 50mg to 400mg once daily. Oral absorption is not affected by concomitant food intake.

Distribution:

The protein binding is low (11-12%). Following either single or multiple oral doses for upto 14 days, Fluconazole is widely distributed in all body fluids and the apparent volume of distribution is close to that of the total body water. Concentration in joint fluid, saliva, sputum, vaginal fluid and peritoneal fluid are similar to those in plasma. Concentration in CSF ranges from 50% to 90% of plasma concentration. Even in the absence of meningeal inflammation.

Metabolism:

Fluconazole is metabolised only to a minor extent. Fluconazole is a selective inhibitor of the isozymes CYP2C9 and CYP3A4.

Elimination:

Plasma elimination half-life of fluconazole is approximately 30 hours. The major route of excretion is renal, with approximately 80% of the administered dose appearing in the urine as unchanged drug and remainder as metabolites. Fluconazole clearance is proportional to creatinine clearance.

Special Population:

Renal Impairment:

In patients with severe renal insufficiency half life increased and reduction of the dose is needed. Fluconazole is removed by dialysis.

THERAPEUTIC INDICATIONS:

FLISK (Fluconazole) is indicated for the treatment of:

1. **Mucosal Candidiasis:** These include oropharyngeal, oesophageal, non-invasive bronchopulmonary infections, candiduria, mucocutaneous & atrophic candidiasis (denture sore mouth).

2. **Systemic Candidiasis:** including candidemia, disseminated candidiasis and other forms of invasive candidal infection.

3. **Vaginal Candidiasis:** Acute or recurrent.

4. **For Prevention of Fungal Infections:** in patients with malignancy who are predisposed to such infections as a result of cytotoxic chemotherapy or radiotherapy.

5. **Dermatomycosis:** Tinea pedis, tinea corporis, tinea cruris, tinea versicolor and dermal candida infection.

6. **Cryptococcosis:** including cryptococcal meningitis and infections of other sites (pulmonary and cutaneous).

Normal hosts & patients with AIDS, organ transplant or other causes of immunosuppression may be treated.

FLISK can be used as maintenance therapy to prevent relapse of cryptococcal disease in patients with AIDS.

DOSAGE AND ADMINISTRATION: The usual adult oral dose of FLISK is given mentioned in the following table:

ADULT DOSE:

Indications	Dosage	Duration (Days)
Oropharyngeal candidiasis	50mg - 100mg daily	7 - 14 (Duration depend on severity)
Oesophageal candidiasis	200mg day 1 followed by 100 mg	14 - 30
Atrophic oral candidiasis	50mg daily	14
Systemic candidiasis and cryptococcal infections (including meningitis)	400mg day 1 followed by 200mg daily	28 or longer based on response
Acute and recurrent vaginal candidiasis	150mg single dose	In few cases duration should be individualized and ranges from 4-12 months 1 capsule per month
Prevention of fungal infections	50mg-100mg daily adjusted individually	

فلسک
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150mg Capsules & 50mg/5mL Dry Suspension

Dermal infections including Tinea pedis, tinea corporis, tinea cruris, dermal candida infection	150mg weekly	2-4 weeks (For tinea pedis 6 weeks)
Tinea versicolor	300mg once weekly	2 weeks
Prevention of relapse of cryptococcal meningitis	100mg-200mg daily	

Paediatric Population:

A maximum dosage of 400mg daily should not be exceeded in children.

Infants, children and adolescents (from 28 days to 17 years):

Indications	Dosage	Duration (Days)
Mucosal candidiasis	1-2mg/kg daily	-
Systemic candidiasis, Cryptococcal infections	3-6 mg/kg daily	Dosage dependant on the severity of the disease

Term newborn infants (0 to 27 days): The same mg/kg dosing as in older children administer every 72 hours in neonate up to 2 weeks old, every 48 hours in neonate 2-4 weeks old.

Directions for Preparing Oral Suspension:

1. Tap the bottle to release the powder.
2. Add a small quantity of previously boiled and cooled water and shake it vigorously.
3. Add water up to the mark on the bottle.
4. Shake well for 1 to 2 minutes to obtain a homogenous suspension.
5. The reconstituted suspension can be used for upto 7days, when stored at room temperature and 14days when stored in refrigerator.

Special Population:

Renal Impairment:

In patients with renal insufficiency dosage adjusted according to creatinine clearance.

CONTRAINDICATIONS:

FLISK is contraindicated in following conditions:

- Patients with known hypersensitivity to fluconazole or to related azole compounds.
- With cisapride and terfenadine.

PRECAUTIONS:

FLISK should be used with care in the following conditions:

- Fluconazole should be administered with caution to patients with liver dysfunction.
- Fluconazole should be administered with caution to patients with proarrhythmic conditions.

Pregnancy:

FLISK is used in pregnancy only if potential benefits justifies possible risk to the fetus.

Lactation:

FLISK is contraindicated during lactation.

DRUG INTERACTIONS:

- **Hydrochlorothiazide** increases plasma concentrations of fluconazole by 40%.

- Fluconazole increases plasma levels of **oral sulphonylureas**.

- In post-marketing experience, as with other azole antifungals, bleeding events (bruising, epistaxis, gastrointestinal bleeding, hematuria, and melena) have been reported, in association with increases in prothrombin time in patients receiving fluconazole concurrently with **warfarin**. Prothrombin time should be carefully monitored. Dose adjustment of the anticoagulant may be necessary.

- Concomitant administration with **phenytoin** may increase the level of phenytoin.

- Fluconazole increases **cyclosporine** level in renal transplant patient with or without renal impairment.

- **Rifampicin** increases metabolism and decreases availability of fluconazole.

- Fluconazole decreases clearance increases **theophylline** levels.

- Fluconazole increases plasma concentration of **cetacecab** dose should be reduced to half

- Fluconazole increases plasma concentration of **midazolam** and prolonged sedative effect.

- With **drugs metabolized by cytochrome P450** serum levels of these drugs increases.

ADVERSE REACTIONS:

Fluconazole is generally well tolerated.

Very common: nausea, abdominal discomfort, diarrhoea and flatulence and increased liver enzymes.

Uncommon: rash are rarely encountered. Less frequently dyspepsia, vomiting, taste disturbance, hepatic disorders, angioedema, dizziness, pruritis may occur.

OVERDOSEAGE:

In the event of overdose, supportive measure and gastric lavage should be instituted according to the patient's clinical conditions. As fluconazole is excreted largely in urine, forced volume diuresis would probably increase elimination rate. A three hour session of haemodialysis decreases plasma levels approximately 50%.

INSTRUCTIONS:

- Store below 30°C.

- Protect from light & moisture.

- Keep out of reach of children.

To be sold on prescription of a registered medical practitioner.

How Supplied:

FLISK (Fluconazole) capsules 150mg are available in pack of 1's.

FLISK (Fluconazole) 50mg/5mL Powder for Oral Suspension is available in pack size of 35mL Bottle.

غیراک ڈاکٹر کی ہدایت کے بغیر استعمال نہ کریں۔
ہدایت دہندہ اک ڈاکٹر کی ہدایت کے بغیر استعمال نہ کریں۔
دقیقہ انداز سے چاہیں۔
مرکز میں ڈاکٹر کے حکم کے مطابق استعمال کریں۔

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