COLCIGEL- colchicinum 4x gel
Gensco Laboratories, LLC

Disclaimer: This homeopathic product has not been evaluated by the Food and Drug Administration for safety or efficacy. FDA is not aware of scientific evidence to support homeopathy as effective.

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HIGHLIGHTS OF PRESCRIBING INFORMATION

These highlights do not include all the information needed to use colchicine safely and effectively. See full prescribing information for ColciGel®.

Initial U.S. Approval

INDICATIONS AND USAGE

ColciGel® (colchicinum 4X) transdermal gel is an alkaloid indicated for:
Treatment and Prophylaxis of Gout Flares in adults. (1.1)

ColciGel® (colchicinum 4X) is not an analgesic medication and should not be used to treat pain from other causes.

DOSAGE FORMS AND STRENGTHS

ColciGel® is Colchicinum 4X in a transdermal gel base. The gel is viscous and opaque in appearance. ColciGel® is available in 15 mL sealed dispensing containers that produce 0.25 mL of gel per each manual depression of the plunger top (pump). (3)

CONTRAINDICATIONS

Traumatized skin, secondary bacterial infection of the area of proposed application and known hypersensitivity to any of the components. (4)

Revised: 12/2018

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* Sections or subsections omitted from the full prescribing information are not listed.
1.1 Gout Flares
ColciGel® transdermal gel is indicated for prophylaxis and the treatment of acute gout flares.

Treatment of Gout Flares:
ColciGel® gel is indicated for treatment of acute gout flares when used at the first sign of a flare.

Prophylaxis of Gout Flares:
ColciGel® is indicated for prophylaxis of gout flares.

2 DOSAGE AND ADMINISTRATION

Apply 1 – 4 pumps up to four times per day.
1 application = 1 pump (0.25ml) covers an area 4” in diameter.

Children under 16 years of age do not use unless directed by a physician.

DISCARD 6 MONTHS AFTER FIRST USE

3 DOSAGE FORMS AND STRENGTHS
ColciGel® is Colchicinum 4X in a transdermal gel base. The gel is viscous and opaque in appearance. ColciGel® is available in 15 mL sealed dispensing containers that produce 0.25 mL of gel per each manual depression of the plunger top (pump).

4 CONTRAINDICATIONS
Patients with severe renal or severe hepatic impairment should not be given ColciGel® in conjunction with P-gp or strong CYP3A4 inhibitors (this includes all protease inhibitors, except fosamprenavir). In these patients, life-threatening and fatal colchicine toxicity has been reported with ORAL colchicine taken in therapeutic doses.

5 WARNINGS
For External Use Only • Avoid Contact With Eyes and Mouth • Keep Out of Reach of Children
Store at 25º C (77º F); exclusions permitted to 15º to 30ºC (59º to 86º F). See USP Controlled Room Temperature

PRECAUTIONS
Use in Pregnancy: No human or animal studies on the effect of ColciGel® (colchicinum 4X) in pregnancy have been conducted.

Nursing Mothers: It is not known whether ColciGel® (colchicinum 4X) is excreted in breast milk.

6 ADVERSE REACTIONS
Prophylaxis and Treatment of Gout Flares: The most commonly reported adverse reaction in clinical trials for the prophylaxis and treatment of gout was mild skin irritation at the site of application.
7 DRUG INTERACTIONS
ColciGel® (Colchicinum 4X) is a substrate of the efflux transporter P-glycoprotein (P-gp). Of the cytochrome P450 enzymes tested, CYP3A4 was mainly involved in the metabolism of colchicinum. If ORAL colchicine is administered with drugs that inhibit P-gp, most of which also inhibit CYP3A4, increased concentrations of colchicinum are likely. Fatal drug interactions have been reported. Topical application of ColciGel® has demonstrated insignificant systemic absorption in animal testing and confirmed in limited human pharmacokinetic evaluations and, therefore, poses a limited risk of clinically significant drug interactions. Physicians should, however, ensure that patients are suitable candidates for treatment with ColciGel® and remain alert for signs and symptoms of toxicities related to increased colchicinum exposure as a result of a drug interaction. Signs and symptoms of ColciGel® toxicity should be evaluated promptly and, if toxicity is suspected, ColciGel® should be discontinued immediately.

12 CLINICAL PHARMACOLOGY
The mechanism by which colchicinum exerts its beneficial effect in patients has not been fully elucidated; however, evidence suggests that colchicinum may interfere with the intracellular assembly of the inflammasome complex present in neutrophils and monocytes that mediates activation of interleukin-1alpha. Additionally, colchicine disrupts cytoskeletal functions through inhibition of alpha-tubulin polymerization into microtubules and consequently prevents the activation, degranulation, and migration of neutrophils thought to mediate some gout symptoms.

13 NONCLINICAL TOXICOLOGY
Carcinogenesis
Carcinogenicity studies of colchicinum have not been conducted. Due to the potential for colchicinum to produce aneuploid cells (cells with an unequal number of chromosomes), there is theoretically an increased risk of malignancy.

Mutagenesis
Colchicine was negative for mutagenicity in the bacterial reverse mutation assay. In a chromosomal aberration assay in cultured human white blood cells, colchicine treatment resulted in the formation of micronuclei. Since published studies demonstrated that colchicine induces aneuploidy from the process of mitotic non-disjunction without structural DNA changes, colchicine is not considered clastogenic, although micronuclei are formed.

Impairment of Fertility
No studies of colchicinum effects on fertility were conducted with ColciGel®. However, published nonclinical studies demonstrated that colchicine-induced disruption of microtubule formation affects meiosis and mitosis. Reproductive studies also reported abnormal sperm morphology and reduced sperm counts in males, and interference with sperm penetration, second meiotic division, and normal cleavage in females when exposed to colchicine. Colchicine administered to pregnant animals resulted in fetal death and teratogenicity. These effects were dose dependent, with the timing of exposure critical for the effects on embryofetal development. The nonclinical doses evaluated were generally higher than an equivalent human ORAL therapeutic dose, but safety margins for reproductive and developmental toxicity could not be determined. Case reports and epidemiology studies in human male subjects on colchicine therapy indicated that infertility from colchicine is rare. A case report indicated that azoospermia was reversed when therapy was stopped. Case reports and epidemiology studies in female subjects on colchicine therapy have not established a clear relationship between colchicinum use and female infertility. The use of colchicinum needs to be weighed against the potential risks.
14 HOW SUPPLIED/STORAGE AND HANDLING

14.1 How Supplied
ColciGel® is Colchicum 4X in a transdermal gel base. The gel is viscous and opaque in appearance. ColciGel® is available in 2x15mL sealed dispensing containers that produce 0.25 mL of gel per each manual depression of the plunger top (pump).
2x15mL (1.01 fl oz) Bottles NDC 35781-0400-4

14.2 Storage
Store at 20° to 25°C (68° to 77°F).
[See USP Controlled Room Temperature]
DISPENSE IN ORIGINAL CONTAINER.
US Patent Pending

Manufactured for:
Gensco Pharma, LLC
8550 NW 33rd Street, Suite 200.
Miami, FL 33122
(855) 743-6726
www.genscopharma.com
COLCIGEL
colchicinum 4x gel

Product Information

Product Type: HUMAN PRESCRIPTION DRUG
Item Code (Source): NDC:35781-0400

Discard 6 months after first use.

DOSAGE AND ADMINISTRATION
Apply 1 - 4 pumps up to four times per day.
1 application = 1 pump (0.25ml) covers an area 4" in diameter.
Children under 16 years of age do not use unless directed by a physician.

Caution: Federal law prohibits dispensing without prescription.
US Patent Pending

Manufactured for:
Gensco Pharma, LLC | Miami, FL 33122
855-743-6726 | www.genscopharma.com

For Oint Flare
For Oral Use
For External Use Only
Avoid Contact With Eyes or Mouth
Keep Out of Reach of Children
Store at 25°C (77°F); excursions permitted to 15°C to 30°C (59°F to 86°F).
See USP Controlled Room Temperature
## Route of Administration

**TRANSDERMAL**

### Active Ingredient/Active Moiety

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<th>Ingredient Name</th>
<th>Basis of Strength</th>
<th>Strength</th>
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<tr>
<td>COLCHICINE (UNII: SML2Y3J35T) (COLCHICINE - UNII:SML2Y3J35T)</td>
<td>COLCHICINE</td>
<td>4 [hp_X] in 15 mL</td>
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### Inactive Ingredients

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<th>Ingredient Name</th>
<th>Strength</th>
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<td>ISOPROPYL MYRISTATE (UNII: 0RE8K4LNJS)</td>
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<td>LECITHIN, SOYBEAN (UNII: 1DI56QDM62)</td>
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<td>UREA (UNII: 8W8T17847W)</td>
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<td>DOCUSATE SODIUM (UNII: F05Q2T2JA0)</td>
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### Packaging

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<th>Marketing End Date</th>
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<td>1</td>
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<td>2 in 1 PACKAGE</td>
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<tr>
<td>1</td>
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### Marketing Information

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**Labeler** - Gensco Laboratories, LLC (831042325)

**Registrant** - Gensco Laboratories, LLC (831042325)

**Establishment**

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<th>Name</th>
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<td>Gensco Laboratories, LLC</td>
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<td>manufacture(35781-0400)</td>
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Revised: 12/2018