

**MUPIROCIN - mupirocin ointment**  
**Keltman Pharmaceuticals Inc.**

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**MUPIROCIN OINTMENT USP, 2%**

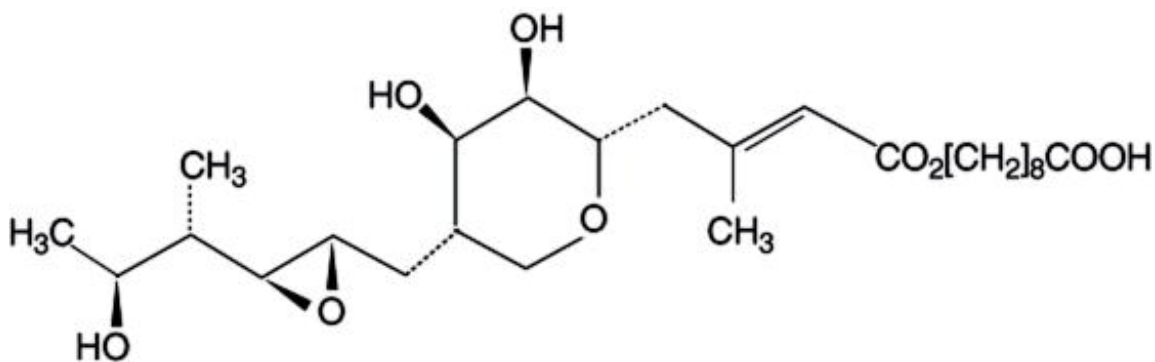
Rx only  
For Dermatologic Use

**DESCRIPTION**

Each gram of mupirocin ointment USP, 2% contains 20 mg mupirocin in a bland water miscible ointment base (polyethylene glycol ointment, N.F.) consisting of polyethylene glycol 400 and polyethylene glycol 3350.

Mupirocin is a naturally occurring antibiotic. The chemical name is (*E*)-(2*S*,3*R*,4*R*,5*S*)-5-[(2*S*,3*S*,4*S*,5*S*)-2,3-Epoxy-5-hydroxy-4-methylhexyl]tetrahydro-3,4-dihydroxy- $\beta$ -methyl-2*H*-pyran-2-crotonic acid, ester with 9-hydroxynonanoic acid.

The molecular formula of mupirocin is C<sub>26</sub>H<sub>44</sub>O<sub>9</sub> and the molecular weight is 500.63. The chemical structure is:



**CLINICAL PHARMACOLOGY**

Application of <sup>14</sup>C-labeled mupirocin ointment to the lower arm of normal male subjects followed by occlusion for 24 hours showed no measurable systemic absorption (<1.1 nanogram mupirocin per milliliter of whole blood). Measurable radioactivity was present in the stratum corneum of these subjects 72 hours after application.

Following intravenous or oral administration, mupirocin is rapidly metabolized. The principal metabolite, monic acid, is eliminated by renal excretion, and demonstrates no antibacterial activity. In a study conducted in seven healthy adult male subjects, the elimination half-life after intravenous administration of mupirocin was 20 to 40 minutes for mupirocin and 30 to 80 minutes for monic acid. The pharmacokinetics of mupirocin has not been studied in individuals with renal insufficiency.

**Microbiology:** Mupirocin is an antibacterial agent produced by fermentation using the organism *Pseudomonas fluorescens*. It is active against a wide range of gram-positive bacteria including methicillin-resistant *Staphylococcus aureus* (MRSA). It is also active against certain gram-negative bacteria. Mupirocin inhibits bacterial protein synthesis by reversibly and specifically binding to bacterial isoleucyl transfer-RNA synthetase. Due to this unique mode of action, mupirocin demonstrates no *in vitro* cross-resistance with other classes of antimicrobial agents.

Resistance occurs rarely. However, when mupirocin resistance does occur, it appears to result from

the production of a modified isoleucyl-tRNA synthetase. High-level plasmid-mediated resistance (MIC >1024 mcg/mL) has been reported in some strains of *S. aureus* and coagulase-negative staphylococci.

Mupirocin is bactericidal at concentrations achieved by topical administration. However, the minimum bactericidal concentration (MBC) against relevant pathogens is generally eight-fold to thirty-fold higher than the minimum inhibitory concentration (MIC). In addition, mupirocin is highly protein bound (>97%), and the effect of wound secretions on the MICs of mupirocin has not been determined.

Mupirocin has been shown to be active against most strains of *Staphylococcus aureus* and *Streptococcus pyogenes*, both *in vitro* and in clinical studies. (See **INDICATIONS AND USAGE**.) The following *in vitro* data are available, BUT THEIR CLINICAL SIGNIFICANCE IS UNKNOWN. Mupirocin is active against most strains of *Staphylococcus epidermidis* and *Staphylococcus saprophyticus*.

## INDICATIONS AND USAGE

Mupirocin ointment, 2% is indicated for the topical treatment of impetigo due to: *Staphylococcus aureus* and *Streptococcus pyogenes*.

## CONTRAINDICATIONS

This drug is contraindicated in individuals with a history of sensitivity reactions to any of its components.

## WARNINGS

Mupirocin ointment is not for ophthalmic use.

## PRECAUTIONS

If a reaction suggesting sensitivity or chemical irritation should occur with the use of mupirocin ointment 2%, treatment should be discontinued and appropriate alternative therapy for the infection instituted. As with other antibacterial products, prolonged use may result in overgrowth of nonsusceptible organisms, including fungi. Mupirocin ointment is not formulated for use on mucosal surfaces. Intranasal use has been associated with isolated reports of stinging and drying.

Polyethylene glycol can be absorbed from open wounds and damaged skin and is excreted by the kidneys. In common with other polyethylene glycol-based ointments, mupirocin ointment should not be used in conditions where absorption of large quantities of polyethylene glycol is possible, especially if there is evidence of moderate or severe renal impairment. A paraffin-based formulation- \*Bactroban<sup>®</sup> Nasal (mupirocin calcium ointment) - is available for intranasal use.

**Information for Patients:** Use this medication only as directed by your healthcare practitioner. It is for external use only. Avoid contact with the eyes. The medication should be stopped and your healthcare practitioner contacted if irritation, severe itching, or rash occurs. If impetigo has not improved in 3 to 5 days, contact your healthcare practitioner.

**Drug Interactions:** The effect of the concurrent application of mupirocin ointment and other drug products has not been studied.

**Carcinogenesis, Mutagenesis, Impairment of Fertility:** Long-term studies in animals to evaluate carcinogenic potential of mupirocin have not been conducted. Results of the following studies performed with mupirocin calcium or mupirocin sodium *in vitro* and *in vivo* did not indicate a potential for genotoxicity: rat primary hepatocyte unscheduled DNA synthesis, sediment analysis for DNA strand breaks, *Salmonella* reversion test (Ames), *Escherichia coli* mutation assay, metaphase analysis of human lymphocytes, mouse lymphoma assay, and bone marrow micronuclei assay in mice. Reproduction

studies were performed in male and female rats with mupirocin administered subcutaneously at doses up to 14 times a human topical dose (approximately 60 mg mupirocin per day) on a mg/m<sup>2</sup> basis and revealed no evidence of impaired fertility and reproductive performance from mupirocin.

**Pregnancy Teratogenic Effects. Pregnancy Category B:** Reproduction studies have been performed in rats and rabbits with mupirocin administered subcutaneously at doses up to 22 and 43 times, respectively, the human topical dose (approximately 60 mg mupirocin per day) on a mg/m<sup>2</sup> basis and revealed no evidence of harm to the fetus due to mupirocin. There are, however, no adequate and well-controlled studies in pregnant women. Because animal studies are not always predictive of human response, this drug should be used during pregnancy only if clearly needed.

**Nursing Mothers:** It is not known whether this drug is excreted in human milk. Because many drugs are excreted in human milk, caution should be exercised when *mupirocin ointment* is administered to a nursing woman.

**Pediatric Use:** The safety and effectiveness of *mupirocin ointment* have been established in the age range of 2 months to 16 years. Use of *mupirocin ointment* in these age groups is supported by evidence from adequate and well-controlled studies of *mupirocin ointment* in impetigo in pediatric patients studied as a part of the pivotal clinical trials. (See **CLINICAL STUDIES**.)

## ADVERSE REACTIONS

The following local adverse reactions have been reported in connection with the use of *mupirocin ointment*: burning, stinging, or pain in 1.5% of patients; itching in 1% of patients; rash, nausea, erythema, dry skin, tenderness, swelling, contact dermatitis, and increased exudate in less than 1% of patients.

## DOSAGE AND ADMINISTRATION

A small amount of mupirocin ointment should be applied to the affected area three times daily. The area treated may be covered with a gauze dressing if desired. Patients not showing a clinical response within 3 to 5 days should be re-evaluated.

## CLINICAL STUDIES

The efficacy of topical *mupirocin ointment* in impetigo was tested in two studies. In the first, patients with impetigo were randomized to receive either *mupirocin ointment* or vehicle placebo t.i.d. for 8 to 12 days. Clinical efficacy rates at end of therapy in the evaluable populations (adults and pediatric patients included) were 71% for *mupirocin ointment* (n=49) and 35% for vehicle placebo (n=51). Pathogen eradication rates in the evaluable populations were 94% for *mupirocin ointment* and 62% for vehicle placebo. There were no side effects reported in the group receiving *mupirocin ointment*. In the second study, patients with impetigo were randomized to receive either *mupirocin ointment* t.i.d. or 30 to 40 mg/kg oral erythromycin ethylsuccinate per day (this was an unblinded study) for 8 days. There was a follow-up visit 1 week after treatment ended.

Clinical efficacy rates at the follow-up visit in the evaluable populations (adults and pediatric patients included) were 93% for *mupirocin ointment* (n=29) and 78.5% for erythromycin (n=28). Pathogen eradication rates in the evaluable patient populations were 100% for both test groups. There were no side effects reported in the *mupirocin ointment* group.

## Pediatrics

There were 91 pediatric patients aged 2 months to 15 years in the first study described above. Clinical efficacy rates at end of therapy in the evaluable populations were 78% for *mupirocin ointment* (n=42) and 36% for vehicle placebo (n=49). In the second study described above, all patients were pediatric except two adults in the group receiving *mupirocin ointment*. The age range of the pediatric patients was

7 months to 13 years.

The clinical efficacy rate for *mupirocin ointment* (n=27) was 96%, and for erythromycin it was unchanged (78.5%).

### HOW SUPPLIED

Mupirocin Ointment USP, 2% is supplied:  
NDC 68387-550-01 22 gram tube

Store at 20° to 25°C (68° to 77°F)[see USP Controlled Room Temperature].

\*Bactroban® Nasal is a registered trademark of SmithKline Pharmaceuticals.

### E. FOUGERA & CO.

A division of Nycomed US Inc.  
MELVILLE, NEW YORK 11747

This Product was Repackaged By Sandhills Packaging For:

### Keltman Pharmaceuticals Inc.

1 Lakeland Square, Suite A  
Flowood, MS 39232  
United States

### PACKAGE LABEL – PRINCIPAL DISPLAY PANEL – 22 G CONTAINER

Fougera®

### MUPIROCIN

### OINTMENT USP, 2%

Rx only

For dermatologic use only.

Each gram contains 20 mg  
mupirocin in polyethylene  
glycol ointment, NF base.

**NET WT 22 grams**

The image shows a product label for Mupirocin Ointment 2%. The label features a blue header with the Keltman Pharmaceuticals, Inc. logo. Below the logo, the NDC number 68387-550-01 and the product name 'Mupirocin Ointment 2%' are prominently displayed. The label also indicates '1 Unit' and 'Mupirocin 20mg/gram'. A barcode is located on the left side, with the number 68387-0550-01++586C1 printed vertically next to it. The text 'Apply to affected area one or more times daily.' is written vertically to the right of the barcode. The label includes storage instructions: 'Store at controlled room temperature 20-25 C (68-77 F). Keep out of reach of children. Dosage: See package insert. Caution: Federal law prohibits transfer of this drug to any person other than the patient for whom prescribed.' It also lists the manufacturer 'E. Fougera & Co. a division of Nycomed US Inc. Melville, NY 11747' and the distributor 'Keltman Pharmaceuticals, Inc. Flowood, MS 39232'. A 'Rx Only' label is placed on the right side of the product information. On the far right, there are three identical blocks of text, each containing the NDC number, Rx number (84473), product name, lot number (586C 1), and expiration date (6/2010). These blocks are labeled 'Chart', 'Patient', and 'Bill' from top to bottom, and a 'Log' label is at the bottom right.

**Keltman Pharmaceuticals, Inc.**  
NDC:68387-550-01  
**Mupirocin Ointment 2%**  
1 Unit  
Mupirocin 20mg/gram  
Rx Only  
Lot: 586C 1 Exp: 6/2010 Rx #: 84473  
Store at controlled room temperature 20-25 C (68-77 F).  
Keep out of reach of children. Dosage: See package insert  
Caution: Federal law prohibits transfer of this drug to any person other than the patient for whom prescribed.  
Manufactured by: E. Fougera & Co. a division of Nycomed US Inc. Melville, NY 11747  
Distributed by: Keltman Pharmaceuticals, Inc. Flowood, MS 39232  
Call your doctor for medical advice about side effects.  
You may report side effects to FDA at 1-800-FDA-1088.

68387-0550-01++586C1  
Apply to affected area one or more times daily.

NDC: 68387-550-01 Rx #: 84473  
Mupirocin Ointment 2% (mupirocin)  
Lot: 586C 1 Exp: 6/2010  
Chart

NDC: 68387-550-01 Rx #: 84473  
Mupirocin Ointment 2% (mupirocin)  
Lot: 586C 1 Exp: 6/2010  
Patient

NDC: 68387-550-01 Rx #: 84473  
Mupirocin Ointment 2% (mupirocin)  
Lot: 586C 1 Exp: 6/2010  
Bill

NDC: 68387-550-01 Rx #: 84473  
Mupirocin Ointment 2% (mupirocin)  
Lot: 586C 1 Exp: 6/2010  
Log

# MUPIROCIN

mupirocin ointment

## Product Information

<b>Product Type</b>	HUMAN PRESCRIPTION DRUG	<b>Item Code (Source)</b>	NDC:68387-550(NDC:0168-0352)
<b>Route of Administration</b>	TOPICAL		

## Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
MUPIROCIN (UNII: D0GX8630A5) (mupirocin - UNII:D0GX8630A5)	MUPIROCIN	20 mg in 1 g

## Inactive Ingredients

Ingredient Name	Strength
POLYETHYLENE GLYCOL 3350 (UNII: G2M7P15E5P)	
POLYETHYLENE GLYCOL 400 (UNII: B697894SGQ)	

## Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:68387-550-01	1 in 1 BAG		
1		22 g in 1 TUBE		

## Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
ANDA	ANDA065192	06/22/2009	

**Labeler** - Keltman Pharmaceuticals Inc. (362861077)

## Establishment

Name	Address	ID/FEI	Business Operations
Keltman Pharmaceuticals Inc.		362861077	repack, relabel

Revised: 6/2011

Keltman Pharmaceuticals Inc.