

Each rectal suppository contains:

Mesalamine, USP 1000 mg

Inactive Ingredient: Hard Fat

Adult Dosage: One (1000 mg) rectal suppository daily. See accompanying prescribing information.

Store below 25°C (77°F). May be refrigerated.

Keep away from direct heat, light or humidity.

Keep this and all medication out of the reach of children.

Keep container tightly closed.

AFFIX PHARMACY LABEL HERE

NDC 0378-9230-93

Mesalamine Rectal Suppositories

1000 mg

PHARMACIST: Dispense the accompanying Patient Information Leaflet to each patient.



For Rectal Use Only

Rx only

30 Rectal Suppositories (5 Strips of 6)

NDC 0378-9230-93

**Mesalamine
Rectal Suppositories**

1000 mg

PHARMACIST: Dispense the accompanying Patient Information Leaflet to each patient.



For Rectal Use Only

Rx only

30 Rectal Suppositories (5 Strips of 6)



0732-00

Manufactured by:

Halo Pharmaceuticals Incorporation
Whippany, NJ 07891 U.S.A.

Manufactured for:

Mylan Pharmaceuticals Inc.
Morgantown, WV 26505 U.S.A.

HL:9230:30C:R1



NDC 0378-9230-93

Mesalamine Rectal Suppositories

1000 mg

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For Rectal Use Only

Rx only

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NDC 0378-9230-23

Mesalamine Rectal Suppositories

1000 mg

PHARMACIST: Dispense the accompanying Patient Information Leaflet to each patient.

For Rectal Use Only



Rx only

**42 Rectal Suppositories (7 Strips of 6)
Full Course of Therapy**



Manufactured by:
Halo Pharmaceuticals Incorporation
Whippany, NJ 07891 U.S.A.
Manufactured for:
Mylan Pharmaceuticals Inc.
Morgantown, WV 26505 U.S.A.
Mylan.com



NDC 0378-9230-23

Mesalamine Rectal Suppositories

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Rx only

**42 Rectal Suppositories (7 Strips of 6)
Full Course of Therapy**

NDC 0378-9230-23

Mesalamine Rectal Suppositories

1000 mg

PHARMACIST: Dispense the accompanying Patient Information Leaflet to each patient.

For Rectal Use Only



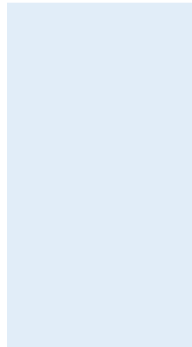
Rx only

**42 Rectal Suppositories (7 Strips of 6)
Full Course of Therapy**

Each rectal suppository contains:
Mesalamine, USP 1000 mg
Inactive Ingredient: Hard Fat
Adult Dosage: One (1000 mg) rectal suppository daily. See accompanying prescribing information.
Store below 25°C (77°F). May be refrigerated.
Keep away from direct heat, light or humidity.
Keep this and all medication out of the reach of children.
Keep container tightly closed.



LOT
EXP



MESALAMINE

RECTAL SUPPOSITORY

1000 mg

Store below 25°C (77°F). May be refrigerated.

Mfd. for: Mylan Pharmaceuticals Inc.

Morgantown, WV 26505 U.S.A.

HL:9230:1S:R1



Mesalamine

Rectal Suppositories

1000 mg



HIGHLIGHTS OF PRESCRIBING INFORMATION

These highlights do not include all the information needed to use mesalamine rectal suppositories safely and effectively. See full prescribing information for mesalamine rectal suppositories.

MESALAMINE rectal suppository, for rectal administration

Initial U.S. Approval: 1987

INDICATIONS AND USAGE

Mesalamine rectal suppositories are an aminosalicilate indicated for the treatment of mild to moderately active ulcerative proctitis. Safety and effectiveness of mesalamine rectal suppositories beyond 6 weeks have not been established. (1)

DOSAGE AND ADMINISTRATION

The dosage is one 1000 mg rectal suppository once daily at bedtime. (2)

DOSAGE FORMS AND STRENGTHS

- 1000 mg rectal suppository (3)

CONTRAINDICATIONS

Hypersensitivity to mesalamine or to any components of the formulation (4)

WARNINGS AND PRECAUTIONS

- Renal impairment may occur. Assess renal function at the beginning of treatment and periodically during treatment. (5.1)
- Mesalamine-induced acute intolerance syndrome has been reported. Observe patients closely for worsening of these symptoms while on treatment. (5.2)
- Use caution when treating patients who are hypersensitive to sulfasalazine. (5.3)

- Mesalamine-induced cardiac hypersensitivity reactions (myocarditis and pericarditis) have been reported. (5.3)
- Hepatic failure has been reported in patients with pre-existing liver disease. Use caution when treating patients with liver disease. (5.4)

ADVERSE REACTIONS

The most common adverse reactions occurring in more than 1% of mesalamine suppository treated patients are: dizziness, rectal pain, fever, rash, acne and colitis. (6.1)

To report SUSPECTED ADVERSE REACTIONS, contact Mylan Pharmaceuticals Inc. at 1-877-446-3679 (1-877-4-INFO-RX) or FDA at 1-800-FDA-1088 or www.fda.gov/medwatch.

DRUG INTERACTIONS

- Nephrotoxic agents including NSAIDs: renal reactions have been reported. (7.1)
- Azathioprine or 6-mercaptopurine: blood disorders have been reported. (7.2)

USE IN SPECIFIC POPULATIONS

- Renal impairment: Use mesalamine with caution in patients with a history of renal disease. (5.1, 7.1, 8.5, 13.2)
- Nursing Women: Caution should be exercised when administered to a nursing woman. (8.3)
- Geriatric Patients: Monitor blood cell counts in geriatric patients. (8.5)

See 17 for PATIENT COUNSELING INFORMATION and FDA-approved patient labeling.

JANUARY 2015

HL: MESASUP-R1

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FULL PRESCRIBING INFORMATION

1 INDICATIONS AND USAGE

Mesalamine 1000 mg rectal suppositories are indicated for the treatment of mild to moderately active ulcerative proctitis. Safety and effectiveness of mesalamine rectal suppositories beyond 6 weeks have not been established.

2 DOSAGE AND ADMINISTRATION

The dosage of mesalamine 1000 mg suppositories is one rectal suppository once daily at bedtime.

The suppository should be retained for one to three hours or longer, if possible. The usual course of therapy is from 3 to 6 weeks depending on symptoms and sigmoidoscopic findings.

If a patient misses a dose of mesalamine rectal suppositories, it should be administered as soon as possible, unless it is almost time for next dose. Patients should not use two mesalamine rectal suppositories at the same time to make up for a missed dose.

3 DOSAGE FORMS AND STRENGTHS

Mesalamine 1000 mg suppositories for rectal administration are available as white to off-white torpedo shaped suppositories containing 1000 mg mesalamine.

4 CONTRAINDICATIONS

Mesalamine rectal suppositories are contraindicated in patients who have demonstrated hypersensitivity to mesalamine (5-aminosalicylic acid) or to the suppository vehicle [saturated vegetable fatty acid esters (Hard Fat, NF)], or to salicylates (including aspirin) [see Warnings and Precautions (5.3), Adverse Reactions (6.2), and Description (11)].

5 WARNINGS AND PRECAUTIONS

5.1 Renal Impairment

Renal impairment, including minimal change nephropathy, acute and chronic interstitial nephritis, and, rarely, renal failure, has been reported in patients given products such as mesalamine rectal suppositories that contain mesalamine or are converted to mesalamine. It is recommended that patients have an evaluation of renal function prior to initiation of mesalamine therapy and periodically while on therapy. Exercise caution when using mesalamine in patients with known renal dysfunction or a history of renal disease. In animal studies, the kidney was the principal organ for toxicity [see Drug Interactions (7.1) and Nonclinical Toxicology (13.2)].

5.2 Mesalamine-Induced Acute Intolerance Syndrome

Mesalamine has been associated with an acute intolerance syndrome that may be difficult to distinguish from an exacerbation of ulcerative colitis. Although the exact frequency of occurrence has not been determined, it has occurred in 3% of patients in controlled clinical trials of mesalamine or sulfasalazine. Symptoms include cramping, acute abdominal pain and bloody diarrhea, and sometimes fever, headache, and rash. Observe patients closely for worsening of these symptoms while on treatment. If acute intolerance syndrome is suspected, promptly discontinue treatment with mesalamine.

5.3 Hypersensitivity Reactions

Some patients who have experienced a hypersensitivity reaction to sulfasalazine may have a similar reaction to mesalamine tablets or to other compounds that contain or are converted to mesalamine.

Mesalamine-induced cardiac hypersensitivity reactions (myocarditis and pericarditis) have been reported with mesalamine rectal suppositories and other mesalamine medications. Caution should be taken in prescribing mesalamine rectal suppositories to patients with hypersensitivity to 5-ASA products.

5.4 Hepatic Impairment

There have been reports of hepatic failure in patients with pre-existing liver disease who have been administered other products containing mesalamine. Caution should be exercised when administering mesalamine to patients with liver disease.

5.5 Drug-Laboratories Test Interactions

There have been several reports of possible interference with measurements, by liquid chromatography, of urinary normetanephrine in patients exposed to sulfasalazine or its metabolite, mesalamine/mesalazine.

6 ADVERSE REACTIONS

The most serious adverse reactions seen in mesalamine rectal suppository clinical trials or with other products that contain or are metabolized to mesalamine are:

- Renal impairment, including renal failure [see Warnings and Precautions (5.1)]
- Mesalamine-induced acute intolerance syndrome [see Warnings and Precautions (5.2)]
- Hypersensitivity reactions [see Warnings and Precautions (5.3)]
- Hepatic impairment, including hepatic failure [see Warnings and Precautions (5.4)]

6.1 Clinical 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.

The most frequent adverse reactions observed in the double-blind, placebo-controlled trials are summarized in the Table 1 below.

Table 1: ADVERSE REACTIONS OCCURRING IN MORE THAN 1% OF MESALAMINE SUPPOSITORY TREATED PATIENTS (COMPARISON TO PLACEBO)

Symptom	Mesalamine (n = 177)		Placebo (n = 84)	
	N	%	N	%
Dizziness	5	3	2	2.4
Rectal Pain	3	1.8	0	0
Fever	2	1.2	0	0
Rash	2	1.2	0	0
Acne	2	1.2	0	0
Colitis	2	1.2	0	0

In a multicenter, open-label, randomized, parallel group study comparing the mesalamine 1000 mg suppository administered nightly to that of the mesalamine 500 mg suppository twice daily, the two treatment groups had similar adverse event profiles. The most frequent AEs were headache (14.4%), flatulence (5.2%), abdominal pain (5.2%), diarrhea (3.1%), and nausea (3.1%). Three (3) patients had to discontinue medication because of an adverse reaction; one of these adverse reactions (headache) was deemed possibly related to study medication.

6.2 Postmarketing Experience

In addition to the adverse reactions reported above in clinical trials involving mesalamine rectal suppositories, the adverse reactions listed below have been identified during post-approval use of mesalamine rectal suppositories and other mesalamine-containing products. 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.

- Body as a Whole: drug fever, fatigue, lupus-like syndrome, medication residue
- Cardiac Disorders: myocarditis, pericarditis, pericardial effusion
- Eye Disorders: eye swelling
- Gastrointestinal Disorders: abdominal cramps, abdominal distension, anal pruritus, anorectal discomfort, constipation, feces discolored, flatulence, frequent bowel movements, gastrointestinal bleeding, mucus stools, nausea,

painful defecation, pancreatitis, proctalgia, rectal discharge, rectal tenesmus, stomach discomfort, vomiting

- Hepatic Disorders: cholestatic jaundice, hepatitis, jaundice, Kawasaki-like syndrome including changes in liver enzymes, liver necrosis, liver failure
- Hematologic Disorders: agranulocytosis, aplastic anemia, thrombocytopenia
- Neurological/Psychiatric Disorders: Guillain-Barre syndrome, peripheral neuropathy, transverse myelitis
- Renal Disorders: interstitial nephritis
- Respiratory, Thoracic and Mediastinal Disorders: hypersensitivity pneumonitis (including allergic alveolitis, eosinophilic pneumonitis, interstitial pneumonitis)
- Skin and Subcutaneous Tissue Disorders: alopecia, erythema, erythema nodosum, pruritus, psoriasis, pyoderma gangrenosum, urticaria
- Urogenital: reversible oligospermia

7 DRUG INTERACTIONS

No investigations of interaction between mesalamine rectal suppositories and other drugs have been performed. However, the following interactions between mesalamine medications and other drugs have been reported.

7.1 Nephrotoxic Agents, Including Non-Steroidal Anti-Inflammatory Drugs

The concurrent use of mesalamine with known nephrotoxic agents, including non-steroidal anti-inflammatory drugs (NSAIDs) may increase the risk of renal reactions.

7.2 Azathioprine or 6-mercaptopurine

The concurrent use of mesalamine with azathioprine or 6-mercaptopurine may increase the risk for blood disorders.

8 USE IN SPECIFIC POPULATIONS

8.1 Pregnancy

Teratogenic Effects: Pregnancy Category B: Reproduction studies have been performed in rats at oral doses up to 320 mg/kg/day (about 1.7 times the recommended human intra-rectal dose, based on body surface area) and in rabbits at oral doses up to 495 mg/kg/day (about 5.4 times the recommended human intra-rectal dose, based on body surface area) and have revealed no evidence of impaired fertility or harm to the fetus due to mesalamine. There are, however, no adequate and well controlled studies in pregnant women. Because animal reproduction studies are not always predictive of human response, this drug should be used in pregnancy only if clearly needed.

8.3 Nursing Mothers

Mesalamine and its N-acetyl metabolite have been detected in human breast milk. The clinical significance of this has not been determined. Caution should be exercised when mesalamine is administered to a nursing woman.

8.4 Pediatric Use

Safety and effectiveness in pediatric patients have not been established.

8.5 Geriatric Use

Reports from uncontrolled clinical studies and postmarketing reporting systems suggested a higher incidence of blood dyscrasias (i.e., neutropenia and pancytopenia) in patients who were 65 years or older who were taking mesalamine-containing products such as mesalamine rectal suppositories. Caution should be taken to closely monitor blood cell counts during mesalamine therapy.

Clinical trials of mesalamine rectal suppositories did not include sufficient numbers of patients aged 65 and over to determine whether they respond differently from younger patients. Other reported clinical experience has not identified differences in responses between the elderly and younger patients. Systemic exposures are increased in elderly subjects [see Clinical Pharmacology (12.3)]. In general, dose selection for an elderly patient should be cautious, usually starting at the low end of the dosing range, reflecting the greater frequency of decreased hepatic, renal, or cardiac function, and of concurrent disease or other drug therapy in elderly patients.

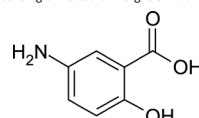
10 OVERDOSAGE

There have been no documented reports of serious toxicity in man resulting from massive overdosing with mesalamine suppository. Under ordinary circumstances, mesalamine absorption from the colon is limited.

11 DESCRIPTION

The active ingredient in mesalamine 1000 mg rectal suppositories is mesalamine, also known as mesalazine or 5-aminosalicylic acid (5-ASA). Chemically, mesalamine is 5-amino-2-hydroxybenzoic acid, and is classified as an anti-inflammatory drug. Each mesalamine rectal suppository contains 1000 mg of mesalamine, USP in a base of Hard Fat.

The molecular formula is C₇H₇NO₃, representing a molecular weight of 153.14. The structural formula is:



12 CLINICAL PHARMACOLOGY

12.1 Mechanism of Action

The mechanism of action of mesalamine is not fully understood, but appears to be topical rather than systemic. Although the pathology of inflammatory bowel disease is uncertain, both prostaglandins and leukotrienes have been implicated as mediators of mucosal injury and inflammation.

12.3 Pharmacokinetics

Absorption: Mesalamine (5-ASA) administered as a rectal suppository is variably absorbed. In patients with ulcerative colitis treated with mesalamine 500 mg rectal suppositories, administered once every 8 hours for 6 days, the mean mesalamine peak plasma concentration (C_{max}) was 353 ng/mL (CV = 55%) following the initial dose and 361 ng/mL (CV = 67%) at steady state. The mean minimum steady state plasma concentration (C_{min}) was 89 ng/mL (CV = 89%). Absorbed mesalamine does not accumulate in the plasma.

Distribution: Mesalamine administered as rectal suppositories distributes in rectal tissue to some extent. In patients with ulcerative proctitis treated with mesalamine 1000 mg rectal suppositories, rectal tissue concentrations for 5-ASA and N-acetyl-5-ASA have not been rigorously quantified.

Metabolism: Mesalamine is extensively metabolized, mainly to N-acetyl-5-ASA. The site of metabolism has not been elucidated. In patients with ulcerative colitis treated with one 500 mg mesalamine rectal suppository every eight hours for six days, peak concentration (C_{max}) of N-acetyl-5-ASA ranged from 467 ng/mL to 1399 ng/mL following the initial dose and from 193 ng/mL to 1304 ng/mL at steady state.

Elimination: Mesalamine is eliminated from plasma mainly by urinary excretion, predominantly as N-acetyl-5-ASA. In patients with ulcerative proctitis treated with one mesalamine 500 mg rectal suppository every 8 hours for 6 days, ≤ 12% of the dose was eliminated in urine as unchanged 5-ASA and 8% to 77% as N-acetyl-5-ASA following the initial dose. At steady state, ≤ 11% of the dose was eliminated as unchanged 5-ASA and 3-35% as N-acetyl-5-ASA. The mean elimination half-life was five hours (CV = 73%) for 5-ASA and six hours (CV = 63%) for N-acetyl-5-ASA following the initial dose. At steady state, the mean elimination half-life was seven hours for both 5-ASA and N-acetyl-5-ASA (CV = 102% for 5-ASA and 82% for N-acetyl-5-ASA).

13 NONCLINICAL TOXICOLOGY

13.1 Carcinogenesis, Mutagenesis, Impairment of Fertility

Mesalamine caused no increase in the incidence of neoplastic lesions over controls in a 2-year study of Wistar rats fed up to 320 mg/kg/day of mesalamine admixed with diet (about 1.7 times the recommended human intra-rectal dose, based on body surface area).

Mesalamine was not mutagenic in the Ames test, the mouse lymphoma cell (TK⁺/-) forward mutation test, or the mouse micronucleus test.

No effects on fertility or reproductive performance of the male and female rats were observed at oral mesalamine doses up to 320 mg/kg/day (about 1.7 times the recommended human intra-rectal dose, based on body surface area).

13.2 Animal Toxicology and/or Pharmacology

Toxicology studies of mesalamine were conducted in rats, mice, rabbits and dogs, and the kidney was the main target organ of toxicity. In rats, adverse renal effects were observed at a single oral dose of 600 mg/kg (about 3.2 times the recommended human intra-rectal dose, based on body surface area) and at IV doses of > 214 mg/kg (about 1.2 times the recommended human intra-rectal dose, based on body surface area). In a 13-week oral gavage toxicity study in rats, papillary necrosis and/or multifocal tubular injury were observed in males receiving 160 mg/kg (about 0.86 times the recommended human intra-rectal dose, based on body surface area) and in both males and females at 640 mg/kg (about 3.5 times the recommended human intra-rectal dose, based on body surface area). In a combined 52-week toxicity and 127-week carcinogenicity study in rats, degeneration of the kidneys and hyalinization of basement membranes and Bowman's capsule were observed at oral doses of 100 mg/kg/day (about 0.54 times the recommended human intra-rectal dose, based on body surface area) and above. In a 14-day rectal toxicity study of mesalamine suppositories in rabbits, intra-rectal doses up to 800 mg/kg (about 8.6 times the recommended human intra-rectal dose, based on body surface area) was not associated with any adverse effects. In a 6-month oral toxicity study in dogs, doses of 80 mg/kg (about 1.4 times the recommended human intra-rectal dose, based on body surface area) and higher caused renal pathology similar to that described for the rat. In a rectal toxicity study of mesalamine suppositories in dogs, a dose of 166.6 mg/kg (about 3 times the recommended human intra-rectal dose, based on body surface area) produced chronic nephritis and pyelitis. In the 12-month eye toxicity study in dogs, keratoconjunctivitis sicca (KCS) occurred at oral doses of 40 mg/kg (about 0.72 times the recommended human intra-rectal dose, based on body surface area) and above.

14 CLINICAL STUDIES

Two double-blind, placebo-controlled, multicenter studies were conducted in North America in patients with mild to moderate active ulcerative proctitis. The primary measures of efficacy were the same in both trials (clinical disease activity index (DAI) and histologic evaluations). The DAI is a composite index reflecting rectal bleeding, stool frequency, mucosal appearance at endoscopy, and a physician's global assessment of disease. The main difference between the studies was dosage regimen: 500 mg three times daily (1.5 g/d) in Study 1; and 500 mg twice daily (1 g/d) in Study 2. A total of 173 patients were studied (Study 1, N = 79; Study 2, N = 94). Eighty-nine (89) patients received mesalamine suppositories, and eighty-four (84) patients received placebo suppositories. Patients were evaluated clinically and sigmoidoscopically after 3 and 6 weeks of suppository treatment. In Study 1, patients were 17 to 73 years of age (mean = 39 years), 57% were female, and 97% were white. Patients had an average extent of proctitis (upper disease boundary) of 10.8 cm. Eighty-four percent (84%) of the study patients had multiple prior episodes of proctitis. In Study 2, patients were 21 to 72 years of age (mean = 39 years), 62% were female, and 96% were white. Patients had an average extent of proctitis (upper disease boundary) of 10.3 cm. Seventy-eight percent (78%) of the study patients had multiple prior episodes of proctitis.

Compared to placebo, mesalamine suppository treatment was statistically ($p < 0.01$) superior to placebo in both trials with respect to improvement in stool frequency, rectal bleeding, mucosal appearance, disease severity, and overall disease activity at 3 and 6 weeks of treatment. The effectiveness of mesalamine suppositories was statistically significant irrespective of sex, extent of proctitis, duration of current episode, or duration of disease.

An additional multicenter, open-label, randomized, parallel group study in ninety-nine (99) patients diagnosed with mild

to moderate ulcerative proctitis compared the clinical efficacy of the mesalamine 1000 mg suppository to that of the mesalamine 500 mg suppository. The primary measures of efficacy included the clinical disease activity index (DAI) and histologic evaluations. Patients were randomized to one of two treatment groups, with a dosage regimen of one 500 mg mesalamine suppository twice daily, morning and at bedtime, or one 1000 mg mesalamine suppository at bedtime for 6 weeks. Patients were evaluated clinically and sigmoidoscopically at 3 and 6 weeks of suppository treatment. Of the eighty-one (81) patients in the Per Protocol population, forty-six (46) patients received mesalamine 500 mg suppositories twice daily, and thirty-five (35) patients received mesalamine 1000 mg suppositories at bedtime.

The efficacy of the 1000 mg at bedtime treatment was not different at 6 weeks from the 500 mg twice daily treatment, and both were effective in the treatment of ulcerative proctitis. Both treatments resulted in a significant decrease at 6 weeks in DAI. In the 500 mg twice daily group, the mean DAI value decreased from 6.6 to 1.6, and in the 1000 mg at bedtime group, the mean DAI value decreased from 6.2 to 1.3 over 6 weeks of treatment, representing a decrease of greater than 75% in both groups. Seventy-eight percent (78%; 36/46) of patients in the 500 mg twice daily group and 86% (30/35) of the patients in the 1000 mg at bedtime group achieved a DAI score of less than 3 after 6 weeks of treatment.

16 HOW SUPPLIED/STORAGE AND HANDLING

Mesalamine Rectal Suppositories for rectal administration are available containing 1000 mg mesalamine, USP.

The 1000 mg rectal suppositories are white to off-white torpedo shaped suppositories. They are available as follows:

NDC 0378-9230-93
Carton of 30 Rectal Suppositories (5 strips of 6 suppositories)
NDC 0378-9230-23
Carton of 42 Rectal Suppositories (7 strips of 6 suppositories)

Store below 25°C (77°F). May be refrigerated.

Keep away from direct heat, light or humidity.

PHARMACIST: Dispense a Patient Information Leaflet with each prescription.

Mesalamine rectal suppositories will cause staining of direct contact surfaces, including but not limited to fabrics, flooring, painted surfaces, marble, granite, vinyl, and enamel.

17 PATIENT COUNSELING INFORMATION

See FDA-approved patient labeling (Patient Information)

Instruct patients not to take mesalamine rectal suppositories if they have hypersensitivity to salicylates (e.g., aspirin) or other mesalamines.

Inform patients to let their physicians know all medications they are taking and if they:

- are allergic to sulfasalazine, salicylates or mesalamine;
- are taking non-steroidal anti-inflammatory drugs (NSAIDs) or other nephrotoxic agents;
- are taking azathioprine or 6-mercaptopurine;
- experience cramping, abdominal pain, bloody diarrhea, fever, headache or rash;
- have a history of myocarditis or pericarditis;
- have kidney or liver disease;
- have a history of stomach blockage;
- are pregnant, intend to become pregnant or are breast-feeding.

FDA-Approved Patient Labeling

PATIENT INFORMATION

MESALAMINE RECTAL SUPPOSITORIES

(me sal' a meen)

1000 mg

Read the Patient Information leaflet that comes with mesalamine before you start taking it and each time you get a refill. There may be new information. This leaflet does not take the place of talking with your doctor about your medical condition or treatment. If you have any questions about mesalamine rectal suppositories, ask your doctor or pharmacist.

What are mesalamine rectal suppositories?

Mesalamine rectal suppositories are a prescription medicine used to treat active ulcerative proctitis (ulcerative rectal colitis).

It is not known if mesalamine rectal suppositories are safe and effective for use for longer than 6 weeks.

It is not known if mesalamine rectal suppositories are safe and effective in children.

Who should not use mesalamine rectal suppositories?

Do not use mesalamine rectal suppositories if you are:

- allergic to medicines that contain salicylates, including aspirin.
- allergic to mesalamine or any of the ingredients in mesalamine rectal suppositories. See the end of this leaflet for a complete list of ingredients in mesalamine rectal suppositories.

Ask your doctor if you are not sure if your medicine is listed above.

What should I tell my doctor before using mesalamine rectal suppositories?

Before using mesalamine rectal suppositories, tell your doctor if you:

- have a history of allergic reaction to the medicine sulfasalazine (Azulfidine)
- have kidney problems
- have ever had inflammation of the sac around your heart (pericarditis).
- have any other medical condition
- are pregnant or plan to become pregnant. It is not known if mesalamine can harm your unborn baby.
- are breastfeeding or plan to breastfeed. Mesalamine can pass into your milk. Talk to your doctor about the best way to feed your baby if you use mesalamine rectal suppositories.

Tell your doctor about all the medicines you take, including prescription and non-prescription medicines, vitamins and herbal supplements.

Know the medicines you take. Keep a list of them to show your doctor and pharmacist when you get a new medicine.

How should I use mesalamine rectal suppositories?

- Use mesalamine rectal suppositories exactly as prescribed by your doctor. Your doctor will tell you how long to continue using mesalamine rectal suppositories.
- Mesalamine comes as a suppository that you insert into your rectum.
- Mesalamine rectal suppositories are used one time each day, at bedtime.
- After a mesalamine rectal suppository is inserted in your rectum, you should try to keep (retain) the suppository in your rectum for 1 to 3 hours, or longer if possible.
- Mesalamine rectal suppositories can cause staining of surfaces including, clothing and other fabrics, flooring, painted surfaces, marble, granite, vinyl and enamel. Keep mesalamine rectal suppositories away from these surfaces to prevent staining.

What are the possible side effects of mesalamine rectal suppositories?

Mesalamine rectal suppositories may cause serious side effects, including:

- **Allergic type reactions.** This can include sudden symptoms called "Acute intolerance syndrome." When this happens, it is usually in people who have had an allergic reaction to medicines containing sulfasalazine. Stop using mesalamine rectal suppositories and tell your doctor right away if you get any of these symptoms:
 - cramps
 - stomach (abdominal) pain
 - bloody diarrhea

- fever
- headache
- rash

- **Inflammation of the sac around the heart (pericarditis).** Tell your doctor right away if you get chest pain or shortness of breath. Your doctor may tell you to stop using mesalamine rectal suppositories if you get pericarditis.

The most common side effects of mesalamine rectal suppositories include:

- dizziness
- acne
- colitis (inflammation of the colon)
- rectal pain (pain in the final portion of the large intestine)
- fever
- rash

Tell your doctor if you have any side effect that bothers you or that does not go away.

These are not all of the possible side effects of mesalamine rectal suppositories. For more information ask your doctor or pharmacist.

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 mesalamine rectal suppositories?

- Store mesalamine rectal suppositories below 25°C (77°F).
- Mesalamine rectal suppositories may be refrigerated.
- Keep mesalamine rectal suppositories away from direct heat, light, or humidity.

Keep mesalamine rectal suppositories and all medicines out of the reach of children.

General information about mesalamine rectal suppositories

Medicines are sometimes prescribed for purposes other than those listed in a patient information leaflet. Do not use mesalamine for a condition for which it was not prescribed. Do not give mesalamine rectal suppositories to other people, even if they have the same symptoms you have. It may harm them.

This leaflet summarizes the most important information about mesalamine rectal suppositories. If you would like more information, talk with your doctor. You can ask your doctor or pharmacist for information about mesalamine rectal suppositories that is written for health professionals.

For more information, call Mylan Pharmaceuticals Inc. at 1-877-446-3679 (1-877-4-INFO-RX).

What are the ingredients in mesalamine rectal suppositories?

Active ingredients: Mesalamine, USP

Inactive ingredients: Hard Fat base

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



Manufactured for:
Mylan Pharmaceuticals Inc.
Morgantown, WV 26505 U.S.A.

Manufactured by:
Halo Pharmaceuticals Incorporation
Whippany, NJ U.S.A.

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