

SULFADIAZINE- sulfadiazine tablet
EPIC PHARMA, LLC

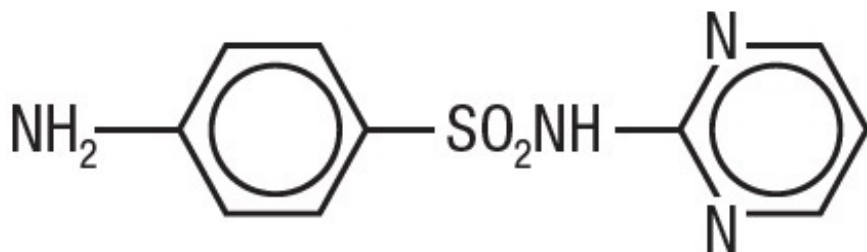
sulfADIAZINE Tablets, USP

DESCRIPTION

Sulfadiazine is an oral sulfonamide antibacterial agent.

Each tablet, for oral administration, contains 500 mg sulfadiazine. In addition, each tablet contains the following inactive ingredients: croscarmellose sodium, docusate sodium, microcrystalline cellulose, povidone, sodium benzoate, sodium starch glycolate and stearic acid.

Sulfadiazine occurs as a white or slightly yellow powder. It is odorless or nearly so and slowly darkens on exposure to light. It is practically insoluble in water and slightly soluble in alcohol. The chemical name of sulfadiazine is N¹-2-pyrimidinylsulfanilamide. The molecular formula is C₁₀H₁₀N₄O₂S. It has a molecular weight of 250.27. The structural formula is shown below:



Most sulfonamides slowly darken on exposure to light.

CLINICAL PHARMACOLOGY

The systemic sulfonamides are bacteriostatic agents having a similar spectrum of activity. Sulfonamides competitively inhibit bacterial synthesis of folic acid (pteroylglutamic acid) from aminobenzoic acid. Resistant strains are capable of utilizing folic acid precursors or preformed folic acid.

Sulfonamides exist in the blood in 3 forms – free, conjugated (acetylated and possibly others) and protein bound. The free form is considered to be the therapeutically active one.

Sulfadiazine given orally is readily absorbed from the gastrointestinal tract. After a single 2 g oral dose, a peak of 6.04 mg/100 mL is reached in 4 hours; of this, 4.65 mg/100 mL is free drug.

When a dose of 100 mg/kg of body weight is given initially and followed by 50 mg/kg every 6 hours, blood levels of free sulfadiazine are about 7 mg/100mL. Protein binding is 38% to 48%. Sulfadiazine diffuses into the cerebrospinal fluid; free drug reaches 32% to

65% of blood levels and total drug 40% to 60%.

Sulfadiazine is excreted largely in the urine, where concentrations are 10 to 25 times greater than serum levels. Approximately 10% of a single oral dose is excreted in the first 6 hours, 50% within 24 hours and 60% to 85% in 48 to 72 hours. Of the amount excreted in the urine, 15% to 40% is in the acetyl form.

INDICATIONS AND USAGE

sulfADIAZINE Tablets, USP are indicated in the following conditions:

Chancroid

Trachoma

Inclusion conjunctivitis

Nocardiosis

Urinary tract infections (primarily pyelonephritis, pyelitis and cystitis) in the absence of obstructive uropathy or foreign bodies, when these infections are caused by susceptible strains of the following organisms: *Escherichia coli*, *Klebsiella species*, *Enterobacter species*, *Staphylococcus aureus*, *Proteus mirabilis* and *P. vulgaris*. Sulfadiazine should be used for urinary tract infections only after use of more soluble sulfonamides has been unsuccessful.

Toxoplasmosis encephalitis in patients with and without acquired immunodeficiency syndrome, as adjunctive therapy with pyrimethamine.

Malaria due to chloroquine-resistant strains of *Plasmodium falciparum*, when used as adjunctive therapy.

Prophylaxis of meningococcal meningitis when sulfonamide-sensitive group A strains are known to prevail in family groups or larger closed populations (the prophylactic usefulness of sulfonamides when group B or C infections are prevalent is not proved and may be harmful in closed population groups).

Meningococcal meningitis, when the organism has been demonstrated to be susceptible.

Acute otitis media due to *Haemophilus influenzae*, when used concomitantly with adequate doses of penicillin.

Prophylaxis against recurrences of rheumatic fever, as an alternative to penicillin.

H. influenzae meningitis, as adjunctive therapy with parental streptomycin.

IMPORTANT NOTES

In vitro sulfonamide susceptibility tests are not always reliable. The test must be carefully coordinated with bacteriologic and clinical response. When the patient is already taking sulfonamides, follow-up cultures should have aminobenzoic acid added to the culture media.

Currently, the increasing frequency of resistant organisms limits the usefulness of antibacterial agents, including the sulfonamides, especially in the treatment of recurrent and complicated urinary tract infections.

Wide variation in blood levels may result with identical doses. Blood levels should be

measured in patients receiving sulfonamides for serious infections. Free sulfonamide blood levels of 5 mg to 15 mg per 100 mL may be considered therapeutically effective for most infections and blood levels of 12 mg to 15 mg per 100 mL may be considered optimal for serious infections. Twenty mg per 100 mL should be the maximum total sulfonamide level, since adverse reactions occur more frequently above this level.

CONTRAINDICATIONS

Sulfadiazine is contraindicated in the following circumstances: Hypersensitivity to sulfonamides.

In infants less than 2 months of age (except as adjunctive therapy with pyrimethamine in the treatment of congenital toxoplasmosis).

In pregnancy at term and during the nursing period, because sulfonamides cross the placenta and are excreted in breast milk and may cause kernicterus.

WARNINGS

The sulfonamides should not be used for the treatment of group A beta-hemolytic streptococcal infections. In an established infection, they will not eradicate the streptococcus and, therefore, will not prevent sequelae such as rheumatic fever and glomerulonephritis.

Deaths associated with the administration of sulfonamides have been reported from hypersensitivity reactions, agranulocytosis, aplastic anemia and other blood dyscrasias.

The presence of such clinical signs as sore throat, fever, pallor, purpura or jaundice may be early indications of serious blood disorders.

The frequency of renal complications is considerably lower in patients receiving the more soluble sulfonamides.

PRECAUTIONS

General

Sulfonamides should be given with caution to patients with impaired renal or hepatic function and to those with severe allergy or bronchial asthma.

Hemolysis may occur in individuals deficient in glucose-6-phosphate dehydrogenase. This reaction is dose related.

Adequate fluid intake must be maintained in order to prevent crystalluria and stone formation.

Information for Patients

Patients should be instructed to drink an eight ounce glass of water with each dose of medication and at frequent intervals throughout the day. Caution patients to report promptly the onset of sore throat, fever, pallor, purpura or jaundice when taking this drug, since these may be early indications of serious blood disorders.

Laboratory Tests

Complete blood counts and urinalyses with careful microscopic examinations should be done frequently in patients receiving sulfonamides.

Drug Interactions

Administration of a sulfonamide may increase the effect of oral anticoagulants and methotrexate, probably by displacement of these drugs from binding sites on plasma albumin. Potentiation of the action of sulfonylurea hypoglycemic agents, thiazide diuretics and uricosuric agents may also be noted. This may also be due to displacement of the drugs from albumin or a pharmacodynamic mechanism may play a role. Conversely, agents such as indomethacin, probenecid and salicylates may displace sulfonamides from plasma albumin and increase the concentrations of free drug in plasma.

Carcinogenesis, Mutagenesis, Impairment of Fertility

The sulfonamides bear certain chemical similarities to some goitrogens. Rats appear to be especially susceptible to the goitrogenic effects of sulfonamides and long-term administration has produced thyroid malignancies in rats.

Pregnancy

Teratogenic Effects

The safe use of sulfonamides in pregnancy has not been established. The teratogenic potential of most sulfonamides has not been thoroughly investigated in either animals or humans. However, a significant increase in the incidence of cleft palate and other bony abnormalities in offspring has been observed when certain sulfonamides of the short, intermediate and long acting types were given to pregnant rats and mice in high oral doses (7 to 25 times the human therapeutic dose).

Nursing Mothers

Sulfadiazine is contraindicated for use in nursing mothers because the sulfonamides cross the placenta, are excreted in breast milk and may cause kernicterus.

Because of the potential for serious adverse reactions in nursing infants from sulfadiazine, a decision should be made whether to discontinue nursing or to discontinue the drug, taking into account the importance of the drug to the mother. See CONTRAINDICATIONS.

Pediatric Use

Sulfadiazine is contraindicated in infants less than 2 months of age (except as adjunctive therapy with pyrimethamine in the treatment of congenital toxoplasmosis). See CONTRAINDICATIONS and DOSAGE AND ADMINISTRATION.

ADVERSE REACTIONS

Blood Dyscrasias

Agranulocytosis, aplastic anemia, thrombocytopenia, leukopenia, hemolytic anemia, purpura, hypoprothrombinemia and methemoglobinemia.

Allergic Reactions

Erythema multiforme (Stevens-Johnson syndrome), generalized skin eruptions, epidermal necrolysis, urticaria, serum sickness, pruritus, exfoliative dermatitis, anaphylactoid reactions, periorbital edema, conjunctival and scleral injection, photosensitization, arthralgia, allergic myocarditis, drug fever and chills.

Gastrointestinal Reactions

Nausea, emesis, abdominal pains, hepatitis, diarrhea, anorexia, pancreatitis and stomatitis.

C.N.S. Reactions

Headache, peripheral neuritis, mental depression, convulsions, ataxia, hallucinations, tinnitus, vertigo and insomnia.

Renal

Crystalluria, stone formation, toxic nephrosis with oliguria and anuria; periarteritis nodosa and lupus erythematosus phenomenon have been noted.

Miscellaneous Reactions

The sulfonamides bear certain chemical similarities to some goitrogens, diuretics (acetazolamide and the thiazides) and oral hypoglycemic agents. Goiter production, diuresis and hypoglycemia have occurred rarely in patients receiving sulfonamides. Cross-sensitivity may exist with these agents.

DOSAGE AND ADMINISTRATION

SYSTEMIC SULFONAMIDES ARE CONTRAINDICATED IN INFANTS UNDER 2 MONTHS OF AGE except as adjunctive therapy with pyrimethamine in the treatment of congenital toxoplasmosis.

Usual Dosage for Infants over 2 Months of Age and Children

Initially, one-half the 24-hour dose. Maintenance, 150 mg/kg or 4 g/m², divided into 4 to 6 doses, every 24 hours, with a maximum of 6 g every 24 hours. Rheumatic fever prophylaxis, under 30 kg (66 pounds), 500 mg every 24 hours; over 30 kg (66 pounds), 1 g every 24 hours.

Usual Adult Dosage

Initially, 2 g to 4 g. Maintenance, 2 g to 4 g, divided into 3 to 6 doses, every 24 hours.

HOW SUPPLIED

sulfADIAZINE Tablets, USP for oral administration are available as

500 mg

White, unscored, capsule-shaped tablets, debossed "E 757" on one face and supplied as:

NDC 42806-757-60 bottles of 60

Storage

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

Protect from moisture.

Dispense contents in a tight, light-resistant container as defined in the USP with a child-resistant closure, as required.

KEEP TIGHTLY CLOSED.

KEEP OUT OF THE REACH OF CHILDREN.

To report SUSPECTED ADVERSE REACTIONS, contact Epic Pharma, LLC at 1-888-374-2791 or FDA at 1-800-FDA-1088 or www.fda.gov/medwatch.

Distributed by:

Epic Pharma, LLC

Laurelton, NY 11413

Rev. 06-2023-00

MF757REV06/23

OS0006

PACKAGE/LABEL PRINCIPAL DISPLAY PANEL - 500 mg

No Varnish	<p>Each tablet contains: Sulfadiazine 500 mg.</p> <p>USUAL DOSAGE: See accompanying literature for complete prescribing information.</p> <p>Store at 20° to 25°C (68° to 77°F) [see USP Controlled Room Temperature].</p> <p>LE0029</p> <p>Rev. 07-2021-00</p>	<p>NDC 42806-757-60</p> <p>sulfADIAZINE Tablets, USP</p> <p>500 mg</p> <p>Rx Only 60 Tablets</p> 	<p>Dispense contents in a tight, light-resistant container as defined in the USP with a child-resistant closure, as required.</p> <p>Protect from moisture.</p> <p>KEEP TIGHTLY CLOSED.</p> <p>KEEP OUT OF THE REACH OF CHILDREN.</p>  <p>3 42806-757-60 4</p>
	<p>Distributed by: Epic Pharma, LLC Laurelton, NY 11413</p>		

SULFADIAZINE

sulfadiazine tablet

Product Information

Product Type	HUMAN PRESCRIPTION DRUG	Item Code (Source)	NDC:42806-757
Route of Administration	ORAL		

Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
SULFADIAZINE (UNII: 0N7609K889) (SULFADIAZINE - UNII:0N7609K889)	SULFADIAZINE	500 mg

Inactive Ingredients

Ingredient Name	Strength
CROSCARMELOSE SODIUM (UNII: M28OL1HH48)	
DOCUSATE SODIUM (UNII: F05Q2T2JA0)	
MICROCRYSTALLINE CELLULOSE (UNII: OP1R32D61U)	
POVIDONE, UNSPECIFIED (UNII: FZ989GH94E)	
SODIUM BENZOATE (UNII: OJ245FE5EU)	
SODIUM STARCH GLYCOLATE TYPE A POTATO (UNII: 5856J3G2A2)	
STEARIC ACID (UNII: 4ELV7Z65AP)	

Product Characteristics

Color	WHITE	Score	no score
Shape	CAPSULE	Size	17mm
Flavor		Imprint Code	E757
Contains			

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:42806-757-60	60 in 1 BOTTLE; Type 0: Not a Combination Product	12/15/2021	

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
ANDA	ANDA040091	12/15/2021	

Labeler - EPIC PHARMA, LLC (827915443)

Registrant - EPIC PHARMA, LLC (827915443)

Establishment

Name	Address	ID/FEI	Business Operations
EPIC PHARMA, LLC		827915443	MANUFACTURE(42806-757)

Revised: 11/2023

EPIC PHARMA, LLC