

# **TIOPRONIN- tiopronin tablet, delayed release**

## **BioComp Pharma, Inc.**

-----

### **HIGHLIGHTS OF PRESCRIBING INFORMATION**

#### **Tiopronin Delayed-Release Tablets delayed-release tablets, for oral use**

**These highlights do not include all the information needed to use Tiopronin Delayed-Release Tablets safely and effectively. See full prescribing information for Tiopronin Delayed-Release Tablets.**

#### **Initial U.S. Approval: 1988**

#### **INDICATIONS AND USAGE**

Tiopronin Delayed-Release Tablets are a reducing and complexing thiol indicated, in combination with high fluid intake, alkali, and diet modification, for the prevention of cystine stone formation in adults and pediatric patients 20 kg and greater with severe homozygous cystinuria, who are not responsive to these measures alone. (1) (1)

#### **DOSAGE AND ADMINISTRATION**

- The recommended initial dosage in adult patients is 800 mg/day. In clinical studies, the average dosage was

about 1,000 mg/day. (2.1) (2)

- The recommended initial dosage in pediatric patients 20 kg and greater is 15 mg/kg/day. Avoid dosages greater

than 50 mg/kg per day in pediatric patients. (5.1, 8.4) (2)

- Measure urinary cystine 1 month after initiation of Tiopronin Delayed-Release Tablets and every 3 months

thereafter (2.3) (2)

- Administer Tiopronin Delayed-Release Tablets in 3 divided doses at the same times each day, with or without

food. Maintain a routine pattern with regard to meals. (2.1) (2)

- Tiopronin Delayed-Release Tablets can be crushed and mixed with applesauce. For preparation and administration instructions, see the full prescribing information. (2.2) (2)

#### **DOSAGE FORMS AND STRENGTHS**

Tablets: 100 mg and 300 mg (3) (3)

#### **CONTRAINDICATIONS**

- Hypersensitivity to tiopronin or any component of Tiopronin Delayed-Release Tablets (4)

#### **WARNINGS AND PRECAUTIONS**

- Proteinuria, including nephrotic syndrome, and membranous nephropathy, has been reported with tiopronin use.

Pediatric patients receiving greater than 50 mg/kg of tiopronin per day may be at increased risk for proteinuria.

(2.1, 5.1, 8.4) (5)

- Hypersensitivity reactions have been reported during tiopronin treatment. (4, 5.2)

#### **ADVERSE REACTIONS**

Most common adverse reactions ( $\geq 10\%$ ) are nausea, diarrhea or soft stools, oral ulcers, rash, fatigue, fever, arthralgia, proteinuria, and emesis. (6)

**To report SUSPECTED ADVERSE REACTIONS, contact BioComp Pharma, Inc. at toll-free**

phone # 1-866-

762-2365 or FDA at 1-800-FDA-1088 or [www.fda.gov/medwatch](http://www.fda.gov/medwatch). (6)

----- **USE IN SPECIFIC POPULATIONS** -----

- Lactation: Breastfeeding is not recommended. (8.2)
- Geriatric: Choose dose carefully and monitor renal function in the elderly. (8.5)

Revised: 12/2024

---

**FULL PRESCRIBING INFORMATION: CONTENTS\***

**1 INDICATIONS AND USAGE**

**2 DOSAGE AND ADMINISTRATION**

**2.1 Recommended Dosage**

**2.2 Preparation and Administration Instructions**

**2.3 Monitoring**

**3 DOSAGE FORMS AND STRENGTHS**

**4 CONTRAINDICATIONS**

**5 WARNINGS AND PRECAUTIONS**

**5.1 Proteinuria**

**5.2 Hypersensitivity Reactions**

**6 ADVERSE REACTIONS**

**6.1 Clinical Trials Experience**

**6.2 Postmarketing Experience**

**7 DRUG INTERACTIONS**

**7.1 Alcohol**

**8 USE IN SPECIFIC POPULATIONS**

**8.1 Pregnancy**

**8.2 Lactation**

**8.4 Pediatric Use**

**8.5 Geriatric Use**

**10 OVERDOSAGE**

**11 DESCRIPTION**

**12 CLINICAL PHARMACOLOGY**

**12.1 Mechanism of Action**

**12.2 Pharmacodynamics**

**12.3 Pharmacokinetics**

**13 NONCLINICAL TOXICOLOGY**

**13.1 Carcinogenesis, Mutagenesis, Impairment of Fertility**

**16 HOW SUPPLIED/STORAGE AND HANDLING**

\* Sections or subsections omitted from the full prescribing information are not listed.

---

**FULL PRESCRIBING INFORMATION**

**1 INDICATIONS AND USAGE**

Tiopronin Delayed-Release Tablets are indicated, in combination with high fluid intake, alkali, and diet modification, for the prevention of cystine stone formation in adults and pediatric patients 20 kg and greater with severe homozygous cystinuria, who are not responsive to these measures alone.

## **2 DOSAGE AND ADMINISTRATION**

### **2.1 Recommended Dosage**

Adults: The recommended initial dosage in adult patients is 800 mg/day. In clinical studies, the average dosage was about 1,000 mg/day.

Pediatrics: The recommended initial dosage in pediatric patients weighing 20 kg and greater is 15 mg/kg/day. Avoid dosages greater than 50 mg/kg per day in pediatric patients [see *Warnings and Precautions (5.1)*, *Use in Specific Populations (8.4)*].

Administer Tiopronin Delayed-Release Tablets in 3 divided doses at the same times each day, with or without food. Maintain a routine pattern with regard to meals.

Consider starting Tiopronin Delayed-Release Tablets at a lower dosage in patients with history of severe toxicity to d-penicillamine.

### **2.2 Preparation and Administration Instructions**

For patients who cannot swallow the tablet whole, Tiopronin Delayed-Release Tablets can be crushed and mixed with applesauce. Administration of Tiopronin Delayed-Release Tablets with other liquids or foods has not been studied and is not recommended.

#### Preparation and Administration of Tiopronin Delayed-Release Tablets Mixed in Applesauce

For patients who can swallow semi-solid food, Tiopronin Delayed-Release Tablets can be crushed and mixed with applesauce:

1. Crush the Tiopronin Delayed-Release Tablet in a clean pill crusher or mortar and pestle. Always crush one tablet at a time.
2. Measure approximately one tablespoon of applesauce and transfer it into a container with the crushed Tiopronin Delayed-Release Tablet.
3. Mix the crushed Tiopronin Delayed-Release Tablet in the applesauce until the powder is well dispersed.
4. Administer the entire Tiopronin Delayed-Release Tablets-applesauce mixture to the patient's mouth immediately. (However, if this is not possible, the mixture can be stored in a refrigerator for up to 2 hours after adding the crushed tablet to the applesauce. Discard any mixture that has not been given within 2 hours.)
5. To assure that any leftover applesauce mixture from the container is recovered, add

tap water to the same container, mix, and have the patient drink the water.

### **2.3 Monitoring**

Measure urinary cystine 1 month after starting Tiopronin Delayed-Release Tablets and every 3 months thereafter. Adjust Tiopronin Delayed-Release Tablets dosage to maintain urinary cystine concentration less than 250 mg/L.

Assess for proteinuria before treatment and every 3 to 6 months during treatment [*see Warnings and Precautions (5.1)*].

Discontinue Tiopronin Delayed-Release Tablets in patients who develop proteinuria, and monitor urinary protein and renal function. Consider restarting Tiopronin Delayed-Release Tablets treatment at a lower dosage after resolution of proteinuria.

### **2.1 Recommended Dosage**

Adults: The recommended initial dosage in adult patients is 800 mg/day. In clinical studies, the average dosage was about 1,000 mg/day.

Pediatrics: The recommended initial dosage in pediatric patients weighing 20 kg and greater is 15 mg/kg/day.

Avoid dosages greater than 50 mg/kg per day in pediatric patients [*see Warnings and Precautions (5.1), Use in Specific Populations (8.4)*].

Administer Tiopronin Delayed-Release Tablets in 3 divided doses at the same times each day, with or without food. Maintain a routine pattern with regard to meals.

Consider starting Tiopronin Delayed-Release Tablets at a lower dosage in patients with history of severe toxicity to d-penicillamine.

### **2.2 Preparation and Administration Instructions**

For patients who cannot swallow the tablet whole, Tiopronin Delayed-Release Tablets can be crushed and mixed with applesauce. Administration of Tiopronin Delayed-Release Tablets with other liquids or foods has not been studied and is not recommended.

#### Preparation and Administration of Tiopronin Delayed-Release Tablets Mixed in Applesauce

For patients who can swallow semi-solid food, Tiopronin Delayed-Release Tablets can be crushed and mixed with applesauce:

1. Crush the Tiopronin Delayed-Release Tablet in a clean pill crusher or mortar and pestle. Always crush one tablet at a time.
2. Measure approximately one tablespoon of applesauce and transfer it into a container with the crushed

Tiopronin Delayed-Release Tablet.

3. Mix the crushed Tiopronin Delayed-Release Tablet in the applesauce until the powder is well dispersed.
4. Administer the entire Tiopronin Delayed-Release Tablets-applesauce mixture to the patient's mouth immediately. (However, if this is not possible, the mixture can be stored in a refrigerator for up to 2 hours after adding the crushed tablet to the applesauce. Discard any mixture that has not been given within 2 hours.)
5. To assure that any leftover applesauce mixture from the container is recovered, add tap water to the same container, mix, and have the patient drink the water.

### **2.3 Monitoring**

Measure urinary cystine 1 month after starting Tiopronin Delayed-Release Tablets and every 3 months thereafter. Adjust Tiopronin Delayed-Release Tablets dosage to maintain urinary cystine concentration less than 250 mg/L.

Assess for proteinuria before treatment and every 3 to 6 months during treatment [see *Warnings and Precautions (5.1)*].

Discontinue Tiopronin Delayed-Release Tablets in patients who develop proteinuria, and monitor urinary protein and renal function. Consider restarting Tiopronin Delayed-Release Tablets treatment at a lower dosage after resolution of proteinuria.

## **3 DOSAGE FORMS AND STRENGTHS**

Tablets for oral use:

100 mg tablets: round, white to off-white and imprinted in red with "T1" on one side

300 mg tablets: round, white to off-white and imprinted in red with "T3" on one side

## **4 CONTRAINDICATIONS**

Tiopronin Delayed-Release Tablets are contraindicated in patients with hypersensitivity to tiopronin or any other components of Tiopronin Delayed-Release Tablets [see *Warnings and Precautions (5.2)*].

## **5 WARNINGS AND PRECAUTIONS**

### **5.1 Proteinuria**

Proteinuria, including nephrotic syndrome, and membranous nephropathy, have been reported with tiopronin use. Pediatric patients receiving greater than 50 mg/kg of tiopronin per day may be at

increased risk for proteinuria [see *Dosage and Administration (2.3)*, *Adverse Reactions (6.1, 6.2)*, *Use in Specific Populations (8.4)*]. Monitor patients for the development of proteinuria and discontinue therapy in patients who develop proteinuria [see *Dosage and Administration (2.3)*].

## **5.2 Hypersensitivity Reactions**

Hypersensitivity reactions (drug fever, rash, fever, arthralgia and lymphadenopathy) have been reported [see *Contraindications (4)*].

## **5.1 Proteinuria**

Proteinuria, including nephrotic syndrome, and membranous nephropathy, have been reported with tiopronin use. Pediatric patients receiving greater than 50 mg/kg of tiopronin per day may be at increased risk for proteinuria [see *Dosage and Administration (2.3)*, *Adverse Reactions (6.1, 6.2)*, *Use in Specific Populations (8.4)*]. Monitor patients for the development of proteinuria and discontinue therapy in patients who develop proteinuria [see *Dosage and Administration (2.3)*].

## **5.2 Hypersensitivity Reactions**

Hypersensitivity reactions (drug fever, rash, fever, arthralgia and lymphadenopathy) have been reported [see *Contraindications (4)*].

## **6 ADVERSE REACTIONS**

The following adverse reactions are discussed in greater detail in other sections of the labeling:

- Proteinuria [see *Warnings and Precautions (5.1)*]
- Hypersensitivity [see *Warnings and Precautions (5.2)*]

### **6.1 Clinical Trials Experience**

Because clinical trials are conducted under widely varying conditions, the adverse reaction rates observed in the clinical trials of the drug cannot be directly compared to rates in the clinical trials of another drug and may not reflect the rates observed in practice.

Adverse reactions occurring at an incidence of  $\geq 5\%$  in an uncontrolled trial in 66 patients with cystinuria age 9 to 68 years are shown in the table below. Patients in group 1 had previously been treated with d-penicillamine; those in group 2 had not. Of those patients who had stopped taking d-penicillamine due to toxicity (34 out of 49 patients in group 1), 22 were able to continue treatment with Tiopronin Tablets. In those without prior history of d-penicillamine treatment, 6% developed reactions of sufficient severity to require Tiopronin Tablets withdrawal.

**Table 1** presents adverse reactions  $\geq 5\%$  in either treatment group occurring in this trial.

**Table 1: Adverse Reactions Occurring in One or More Patients**

System Organ Class	Adverse Reaction	Group 1 Previously treated with d-penicillamine (N = 49)	Group 2 Naïve to d-penicillamine (N = 17)
Blood and Lymphatic System Disorders	anemia	1 (2%)	1 (6%)
Gastrointestinal Disorders	nausea	12 (25%)	2 (12%)
	emesis	5 (10%)	–
	diarrhea/soft stools	9 (18%)	1 (6%)
	abdominal pain	–	1 (6%)
	oral ulcers	6 (12%)	3 (18%)
General Disorders and Administration Site Conditions	fever	4 (8%)	–
	weakness	2 (4%)	2 (12%)
	fatigue	7 (14%)	–
	peripheral (edema)	3 (6%)	1 (6%)
Metabolism and Nutrition Disorders	chest pain	–	1 (6%)
	anorexia	4 (8%)	–
Musculoskeletal and Connective Tissue Disorders	arthralgia	–	2 (12%)
Renal and Urinary Disorders	proteinuria	5 (10%)	1 (6%)
	impotence	–	1 (6%)
Respiratory, Thoracic and Mediastinal Disorders	cough	–	1 (6%)
Skin and Subcutaneous Tissue Disorders	rash	7 (14%)	2 (12%)
	ecchymosis	3 (6%)	–
	pruritus	2 (4%)	1 (6%)
	urticaria	4 (8%)	–
	skin wrinkling	3 (6%)	1 (6%)

### Taste Disturbance

A reduction in taste perception may develop. It is believed to be the result of chelation of trace metals by tiopronin. Hypogeusia is often self-limited.

## **6.2 Postmarketing Experience**

Adverse reactions have been reported from the literature, as well as during post-

approval use of Tiopronin

Tablets. Because the post-approval 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 Tiopronin Tablets exposure.

Adverse reactions reported during the postmarketing use of Tiopronin Tablets are listed by body system in

**Table 2.**

**Table 2: Adverse Reactions Reported for Tiopronin Tablets Pharmacovigilance by System Organ Class and Preferred Term**

System Organ Class	Preferred Term
Cardiac Disorders	congestive heart failure
Ear and Labyrinth Disorder	vertigo
Gastrointestinal Disorders	abdominal discomfort; abdominal distension; abdominal pain; chapped lips; diarrhea; dry mouth; dyspepsia; eructation; flatulence; gastrointestinal disorder; gastroesophageal reflux disease; nausea; vomiting; jaundice; liver transaminitis
General Disorders and Administration Site Conditions	asthenia; chest pain; fatigue; malaise; pain; peripheral swelling; pyrexia; swelling
Investigations	glomerular filtration rate decreased; weight increased
Metabolism and Nutrition Disorders	decreased appetite; dehydration; hypophagia
Musculoskeletal and Connective Tissue Disorders	arthralgia; back pain; flank pain; joint swelling; limb discomfort; musculoskeletal discomfort; myalgia; neck pain; pain in extremity
Nervous System Disorders	ageusia; burning sensation; dizziness; dysgeusia; headache; hypoesthesia
Renal and Urinary Disorders	nephrotic syndrome; proteinuria; renal failure
Skin and Subcutaneous Tissue Disorders	dry skin; hyperhidrosis; pemphigus foliaceus; pruritus; rash; rash pruritic; skin irritation; skin texture abnormal; skin wrinkling; urticaria

## 7 DRUG INTERACTIONS

### 7.1 Alcohol

Tiopronin is released faster from Tiopronin Delayed-Release Tablets in the presence of

alcohol and the risk for adverse events associated with Tiopronin Delayed-Release Tablets when taken with alcohol is unknown. Avoid alcohol consumption 2 hours before and 3 hours after taking Tiopronin Delayed-Release Tablets [see *Clinical Pharmacology* (12.3)].

## **8 USE IN SPECIFIC POPULATIONS**

- Lactation: Breastfeeding is not recommended. (8.2)
- Geriatric: Choose dose carefully and monitor renal function in the elderly. (8.5)

**See 17 for PATIENT COUNSELING INFORMATION.**

**Revised: 07/2021**

### **8.1 Pregnancy**

#### Risk Summary

Available published case report data with tiopronin have not identified a drug-associated risk for major birth defects, miscarriage, or adverse maternal or fetal outcomes. Renal stones in pregnancy may result in adverse pregnancy outcomes (see Clinical Considerations). In animal reproduction studies, there were no adverse developmental outcomes with oral administration of tiopronin to pregnant mice and rats at doses up to 2 times a 2 grams/day human dose (based on mg/m<sup>2</sup>). The estimated background risk of major birth defects and miscarriage for the indicated population is unknown. All pregnancies have a background risk of birth defect, loss, or other adverse outcomes. In the U.S. general population, the estimated background risk of major birth defects and miscarriage in clinically recognized pregnancies are 2% to 4% and 15% to 20%, respectively.

#### Clinical Considerations

##### *Disease-associated maternal and/or embryo/fetal risk*

Renal stones in pregnancy may increase the risk of adverse pregnancy outcomes, such as preterm birth and low birth weight.

#### Data

##### *Animal Data*

No findings of fetal malformations could be attributed to the drug in reproduction studies in mice and rats at doses up to 2 times the highest recommended human dose of 2 grams/day (based on mg/m<sup>2</sup>).

## **8.2 Lactation**

### Risk Summary

There are no data on the presence of tiopronin in either human or animal milk or on the effects of the breastfed child. A published study suggests that tiopronin may suppress milk production. Because of the potential for serious adverse reactions, including nephrotic syndrome, advise patients that breastfeeding is not recommended during treatment with Tiopronin Delayed-Release Tablets.

## **8.4 Pediatric Use**

Tiopronin Delayed-Release Tablets are indicated in pediatric patients weighing 20 kg or more with severe homozygous cystinuria, in combination with high fluid intake, alkali, and diet modification, for the prevention of cystine stone formation who are not responsive to these measures alone. This indication is based on safety and efficacy data from a trial in patients 9 years to 68 years of age and clinical experience. Proteinuria, including nephrotic syndrome, has been reported in pediatric patients. Pediatric patients receiving greater than 50 mg/kg tiopronin per day may be at greater risk [see *Dosage and Administration (2.1, 2.3), Warnings and Precautions (5.1) and Adverse Reactions (6.1)*].

Tiopronin Delayed-Release Tablets are not approved for use in pediatric patients weighing less than 20 kg [see *Dosage and Administration (2.1)*].

## **8.5 Geriatric Use**

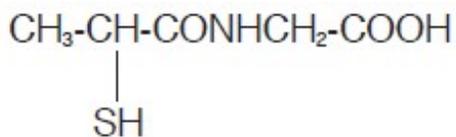
This drug is known to be substantially excreted by the kidney, and the risk of adverse reactions to this drug may be greater in patients with impaired renal function. Because elderly patients are more likely to have decreased renal function, care should be taken in dose selection, and it may be useful to monitor renal function.

## **10 OVERDOSAGE**

There is no information on overdosage with tiopronin.

## **11 DESCRIPTION**

Tiopronin Delayed-Release Tablets are a reducing and cystine-binding thiol drug (CBTD) for oral use. Tiopronin is N-(2-Mercaptopropionyl) glycine and has the following structure:



Tiopronin has the empirical formula  $\text{C}_5\text{H}_9\text{NO}_3\text{S}$  and a molecular weight of 163.20. In this drug product tiopronin exists as a dl racemic mixture.

Tiopronin is a white crystalline powder, which is freely soluble in water.

Each Tiopronin Delayed-Release Tablet contains 100 or 300 mg of tiopronin. The inactive ingredients in

Tiopronin Delayed-Release Tablets include lactose monohydrate, hydroxypropyl cellulose, hydroxypropyl cellulose (low substitute), magnesium stearate, hydroxypropyl methylcellulose E5, methacrylic acid: ethyl acrylate copolymer (Eudragit L 100-55), talc, triethyl citrate.

## 12 CLINICAL PHARMACOLOGY

### 12.1 Mechanism of Action

The goal of therapy is to reduce urinary cystine concentration below its solubility limit.

Tiopronin is an active

reducing agent which undergoes thiol-disulfide exchange with cystine to form a mixed disulfide of tiopronincysteine.

From this reaction, a water-soluble mixed disulfide is formed and the amount of sparingly soluble cystine is reduced.

### 12.2 Pharmacodynamics

The decrement in urinary cystine produced by tiopronin is generally proportional to the dose. A reduction in

urinary cystine of 250-350 mg/day at tiopronin dosage of 1 g/day, and a decline of approximately 500 mg/day

at a dosage of 2 g/day, might be expected. Tiopronin has a rapid onset and offset of action, showing a fall in

cystine excretion on the first day of administration and a rise on the first day of drug withdrawal.

### 12.3 Pharmacokinetics

#### Absorption

#### *Tiopronin Delayed-Release Tablets*

When Tiopronin Tablets and Tiopronin Delayed-Release Tablets single doses were given to fasted healthy

subjects, the median time to peak plasma levels ( $T_{\text{max}}$ ) was 1 (range: 0.5 to 2.1) and 3 (range: 1.0 to 6.0)

hours, respectively. The peak exposure ( $C_{\text{max}}$ ) and total exposure ( $\text{AUC}_{0-t}$ ) of tiopronin from Tiopronin Delayed-

Release Tablets were decreased by 22% and 7% respectively compared to Tiopronin Tablets.

When Tiopronin Delayed-Release Tablets were administered crushed in applesauce, the median time to peak plasma levels of tiopronin ( $T_{max}$ ) was 1 hour (range: 0.5 to 2.0) compared to 3.1 hours (range: 1.5 to 4.0) when administered as intact Tiopronin Delayed-Release Tablets.

When Tiopronin Delayed-Release Tablets were administered crushed in applesauce, the maximum concentration ( $C_{max}$ ) and exposure ( $AUC_{0-t}$ ) to tiopronin were increased by 38% and 14%, respectively, compared to Tiopronin Delayed-Release Tablets administered intact.

#### Food Effects

Administration of the Tiopronin Delayed-Release Tablet with food decreases  $C_{max}$  of tiopronin by 13% and  $AUC_{0-t}$  by 25% compared to Tiopronin Delayed-Release Tablets administered in a fasted state.

Since the drug is dosed to effect, the study results support administration of Tiopronin Delayed-Release Tablets with or without food; administer at the same time each day with a routine pattern with regard to meals.

#### Elimination

##### *Excretion*

When tiopronin is given orally, up to 48% of dose appears in urine during the first 4 hours and up to 78% by 72 hours.

#### Drug Interactions

##### *Alcohol*

An *in vitro* dissolution study was conducted to evaluate the impact of alcohol (5, 10, 20, and 40%) on the dose dumping of Tiopronin Delayed-Release Tablets. The study results showed that the addition of alcohol to the dissolution media increases the dissolution rate of Tiopronin Delayed-Release Tablets in the acidic media of 0.1N HCl [see *Drug Interactions (7.1)*].

## **13 NONCLINICAL TOXICOLOGY**

### **13.1 Carcinogenesis, Mutagenesis, Impairment of Fertility**

#### Carcinogenesis

Long-term carcinogenicity studies in animals have not been performed.

#### Mutagenesis

Tiopronin was not genotoxic in the chromosomal aberration, sister chromatid exchange, and *in vivo* micronucleus assays.

### Impairment of Fertility

High doses of tiopronin in experimental animals have been shown to interfere with maintenance of pregnancy and viability of the fetus. In 2 published male fertility studies in rats, tiopronin at 20 mg/kg/day intramuscular (IM) for 60 days induced reductions in testis, epididymis, vas deferens, and accessory sex glands weights and in the count and motility of cauda epididymal sperm.

## **16 HOW SUPPLIED/STORAGE AND HANDLING**

100 mg delayed-release, round, white to off-white tablet imprinted with "T1" on one side with red ink and blank on the other side: Bottles of 300 **NDC** 44523-054-01.

300 mg delayed-release, round, white to off-white tablet imprinted with "T3" on one side with red ink and blank on the other side: Bottles of 90 **NDC** 44523-055-01.

Store at 25°C (77°F); excursions permitted to 15-30°C (59-86°F) [see *USP Controlled Room Temperature*].

## **17 PATIENT COUNSELING INFORMATION**

### Administration Instructions

For patients who cannot swallow the tablet whole, the Tiopronin Delayed-Release Tablets can be crushed and mixed with applesauce. See *Dosage and Administration (2.2)* for preparation and administration instructions.

### Lactation

Advise women that breastfeeding is not recommended during treatment with Tiopronin Delayed-Release Tablets [see *Use in Specific Populations (8.2)*].



Manufactured for:  
BioComp Pharma, Inc.  
San Antonio, TX 78230 1355

L45020R0822

BCP013R0423

NDC 44523-054-01

Rx only. Usual dosage: See prescribing information. Store at 25°C (77°F); excursions permitted to 15-30°C (59-86°F) [see USP Controlled Room Temperature]. L05401R0721

Manufactured for: BioComp Pharma, Inc.  
by Mission Pharmacal Company  
San Antonio, TX 78230 1355

NDC 44523-054-01

**Tiopronin**  
Delayed-Release Tablets

100 mg  
Rx only  
300 TABLETS

 BioComp Pharma.

3 44523 05401 1

Tiopronin Delayed-Release Tablets 300mg

NDC 44523-055-01

Rx only. Usual dosage: See prescribing information. Store at 25°C (77°F); excursions permitted to 15-30°C (59-86°F) [see USP Controlled Room Temperature]. L05501R0721

Manufactured for: BioComp Pharma, Inc.  
by Mission Pharmacal Company  
San Antonio, TX 78230 1355

NDC 44523-055-01

**Tiopronin**  
Delayed-Release Tablets

300 mg  
Rx only  
90 TABLETS

 BioComp Pharma.

3 44523 05501 8

## TIOPRONIN

tiopronin tablet, delayed release

### Product Information

Product Type	HUMAN PRESCRIPTION DRUG	Item Code (Source)	NDC:44523-054
Route of Administration	ORAL		

### Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
TIOPRONIN (UNII: C5W04GO61S) (TIOPRONIN - UNII:C5W04GO61S)	TIOPRONIN	100 mg

### Inactive Ingredients

Ingredient Name	Strength
METHACRYLIC ACID-ETHYL ACRYLATE COPOLYMER (1:1) TYPE A (UNII: NX76LV5T8J)	
LACTOSE MONOHYDRATE (UNII: EWQ57Q8I5X)	
HYPROMELLOSE, UNSPECIFIED (UNII: 3NXW29V3WO)	
TALC (UNII: 7SEV7J4R1U)	
TRIETHYL CITRATE (UNII: 8Z96QXD6UM)	
MAGNESIUM STEARATE (UNII: 70097M6I30)	
HYDROXYPROPYL CELLULOSE, UNSPECIFIED (UNII: 9XZ8H6N6OH)	
HYDROXYPROPYL CELLULOSE, LOW SUBSTITUTED (UNII: 2165RE0K14)	

### Product Characteristics

Color	white	Score	no score
Shape	ROUND	Size	8mm
Flavor		Imprint Code	T1
Contains			

### Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:44523-054-01	300 in 1 BOTTLE; Type 0: Not a Combination Product	12/05/2024	

### Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
NDA authorized generic	NDA211843	12/05/2024	

## TIOPRONIN

tiopronin tablet, delayed release

### Product Information

Product Type	HUMAN PRESCRIPTION DRUG	Item Code (Source)	NDC:44523-055
Route of Administration	ORAL		

### Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
TIOPRONIN (UNII: C5W04GO61S) (TIOPRONIN - UNII:C5W04GO61S)	TIOPRONIN	300 mg

### Inactive Ingredients

Ingredient Name	Strength
-----------------	----------

<b>METHACRYLIC ACID-ETHYL ACRYLATE COPOLYMER (1:1) TYPE A</b> (UNII: NX76LV5T8J)	
<b>LACTOSE MONOHYDRATE</b> (UNII: EWQ57Q8I5X)	
<b>HYPROMELLOSE, UNSPECIFIED</b> (UNII: 3NXW29V3WO)	
<b>TRIETHYL CITRATE</b> (UNII: 8Z96QXD6UM)	
<b>MAGNESIUM STEARATE</b> (UNII: 70097M6I30)	
<b>HYDROXYPROPYL CELLULOSE, UNSPECIFIED</b> (UNII: 9XZ8H6N6OH)	
<b>HYDROXYPROPYL CELLULOSE, LOW SUBSTITUTED</b> (UNII: 2165RE0K14)	
<b>TALC</b> (UNII: 7SEV7J4R1U)	

### Product Characteristics

<b>Color</b>	white	<b>Score</b>	no score
<b>Shape</b>	ROUND	<b>Size</b>	8mm
<b>Flavor</b>		<b>Imprint Code</b>	T3
<b>Contains</b>			

### Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:44523-055-01	90 in 1 BOTTLE; Type 0: Not a Combination Product	12/05/2024	

### Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
NDA authorized generic	NDA211843	12/05/2024	

**Labeler** - BioComp Pharma, Inc. (829249718)

**Registrant** - Mission Pharmacal Company (927726893)

### Establishment

Name	Address	ID/FEI	Business Operations
Mission Pharmacal Company		927726893	manufacture(44523-054, 44523-055) , label(44523-054, 44523-055) , analysis(44523-054, 44523-055) , pack(44523-054, 44523-055)

Revised: 6/2025

BioComp Pharma, Inc.