

May 27th, 2020

Dear Client,

There is an unprecedented demand and urgent need for access to medications during the COVID-19 pandemic, which has led to reported drug shortages in Canada. Given the medical necessity of timolol maleate ophthalmic solution, Health Canada has added Timo-Stulln 0.5% to the List of Drugs for Exceptional Importation and Sale in Canada.

Pharma Stulln Inc. is pleased to provide **Timo-Stulln 0.5%** (Timolol Maleate Ophthalmic Solution) as an alternative to Canadian timolol maleate 0.5% ophthalmic solution during the current shortage. The product meets high quality standards, and is labelled in German. Translated copies of the label, carton and package leaflet in English and French are provided for your convenience; these materials are also available in English or French on our website at www.pharmastulln.ca.

Note that Timo-Stulln 0.5% should not be stored at a temperature above 25°C and should be kept in its original carton to protect it from light.

If you require any additional information, or to report any adverse reactions to this product, please contact:

Pharma STULLN Inc.

By email: info@phamastulln.ca

By tel.: **1-833-278-8556**

Or by fax: **1-855-631-0332**

Dr Chafic Chebli
CEO

P[®] Timo-Stulln

Timolol Maleate Ophthalmic Solution 0.5% w/v Timolol
Lowers the pressure in the eye for conditions such as glaucoma
and ocular hypertension

Special Note:

Timo-Stulln (Timolol Maleate Ophthalmic Solution) has been approved as an exception by Health Canada due to a product shortage in Canada during this COVID-19 period. The product label, box and leaflet are in German. Official English and French translations are available at www.pharmastulln.ca.



Timo-Stulln® 0,5%

5 ml Augentropfen
Wirkstoff: Timololmaleat

Zum Eintropfen in den Bindehautsack. Packungsbeilage beachten! Nach Anbruch 4 Wochen verwendbar. **Zusammensetzung:** Enthält Benzalkoniumchlorid. Packungsbeilage beachten. Enthält Phosphate. Packungsbeilage beachten.
Verschreibungspflichtig! | Zul.-Nr.: 30346.02.00 | Art.-Nr.: 23.0166.06
Pharma Stulln GmbH | Werksstraße 3 | 92551 Stulln pharma STULLN

Verw. bis: / Ch.-B.:

Timo-Stulln® 0,5%

5 ml Eye Drops
Active material: Timolol Maleate

To administer into the conjunctival sac. Note leaflet! Can be used for 4 weeks after opening. **Composition:** Contains benzalkonium chloride. Note leaflet. Contains phosphates. Note leaflet.
Prescription only! | Zul.-Nr.: 30346.02.00 | Art.-Nr.: 23.0166.06
Pharma Stulln GmbH | Werksstraße 3 | 92551 Stulln pharma STULLN

Exp. Date / Batch #:

English Translation

P[®] Timo-Stulln (Timolol Maleate)

Health Canada ID Code: 06879051
Ontario PIN: 09858120
Product Code: 1061 | 1061-3PK
McKesson Code: 152780 | 152781
Format: Box of 1 bottle | 3 bottles

Strength: 0.5% w/v (5mL Bottle)
Active Ingredient: Timolol Maleate
Preservative: Benzalkonium Chloride 0.1%
Shelf Life: 36 Months
Storage: 15-25°C.

All products available through McKesson.
To order by email: orders@pharmastulln.ca
To order by telephone: 1.833.278.8556 (Customer Service)

Package leaflet: Information for the user

Available on prescription only

Timo-Stulln® 0.5%

Timolol maleate ophthalmic solution

0.5% w/v timolol

For use in adults only.

Read all of this leaflet carefully before you start using this medicine.

- Keep this leaflet. You may need to read it again.
- If you have any further questions, ask your doctor or pharmacist.
- This medicine has been prescribed for you. Do not pass it on to others. It may harm them, even if their symptoms are the same as yours.
- If any of the side effects gets serious, or if you notice any side effects not listed in this leaflet, please tell your doctor or pharmacist.

In this leaflet:

1. What Timo-Stulln® 0.5% is and what it is used for
2. What you need to know before you use Timo-Stulln® 0.5%
3. How to use Timo-Stulln® 0.5%
4. Possible side effects
5. How to store Timo-Stulln® 0.5%
6. Further information

1. WHAT TIMO-STULLN® 0.5% IS AND WHAT IT IS USED FOR

Timo-Stulln® 0.5% is an ophthalmic solution which lowers the pressure in the eye for conditions such as glaucoma and ocular hypertension and is only available through a prescription by your physician.

Timolol maleate, the active substance of Timo-Stulln® 0.5%, belongs to a class of drugs known as beta-receptor blockers. These substances help lower the pressure in the eye.

Timo-Stulln® 0.5% is used to reduce intraocular pressure in:

- Patients with chronic open-angle glaucoma
- Patients with increased intraocular pressure (ocular hypertension)
- Patients with glaucoma who had their lens removed by cataract surgery, including those wearing contact lenses
- Patients with narrow angles and a history of narrow angle closure in the opposite eye in whom reduction of intraocular pressure is necessary

2. WHAT YOU NEED TO KNOW BEFORE YOU USE TIMO-STULLN® 0.5%

Do not use Timo-Stulln® 0.5%:

- if you are allergic (hypersensitive) to the active substance or any of the other ingredients of Timo-Stulln® 0.5%.
- if you have now, or have had in the past, one of the following disorders:
 - asthma, chronic obstructive lung disease or any other lung or breathing problems
 - heart failure, slow or irregular heartbeat, heart-related shock or any other heart problems
 - severe nasal allergy symptoms
 - genetic disease of your cornea
 - night-time pressure elevation, timolol eye drops are not suitable for such patients

Take special care with Timo-Stulln® 0.5%

- Beta-receptor blockers can affect the blood pressure and pulse, which is why care must be exercised in patients with reduced flow to the brain.
- An alternative treatment should be taken into consideration if signs of reduced blood flow to the brain are

- observed after commencing treatment with Timo-Stulln® 0.5%.
- In patients with muscle weakness: increased muscle weakness, and symptoms such as double vision, drooping upper eyelids and general weakness, has been reported in such patients when receiving beta-receptor blockers.
 - If you wear contact lenses. Remove contact lenses prior to application to the eye and wait at least 15 minutes before putting them back in.
 - If you have or have had in the past, diabetes or other blood sugar problems.
 - If you had past eye problems such as choroidal detachment.
 - If you are pregnant or intend to become pregnant.
 - If you are breast feeding or intend to breast feed. Timolol has been detected in human breast milk. Discuss with your physician.
 - NOT RECOMMENDED FOR CHILDREN.

Please make sure to attend your regular intraocular pressure and cornea check-ups.

Other medicines and Timo-Stulln® 0.5%

Please tell your doctor or pharmacist if you are using or have recently used any other medicines, including medicines obtained without a prescription.

The following medicines may affect the action of Timo-Stulln® 0.5%, or be affected by it:

Application of eye drops containing adrenalin at the same time as Timo-Stulln® 0.5% may lead to pupil dilation. The intraocular pressure reducing effect of timolol is enhanced by use of eye drops containing adrenalin or pilocarpine.

Use of two beta-receptor blocker eye drops at the same time is not recommended.

Taking other beta-receptor blockers (drugs for treating high blood pressure and coronary vessel diseases) at the same time as Timo-Stulln® 0.5% eye drops may result in increased effect both in the eye (pressure reduction) and on the cardiovascular system.

Please note that this information may also apply to recently used medication.

Use of Timo-Stulln® 0.5% at the same time as drugs for reducing high blood pressure or for treatment of heart diseases (e.g. calcium antagonists, reserpine or beta-receptor blockers) may lead to an increased occurrence of unwanted effects such as a drop in blood pressure and/or significant slowing down of the pulse (bradycardia).

Use in children

Safety and effectiveness of Timo-Stulln® 0.5% in children have not been established. Timo-Stulln® 0.5% is NOT RECOMMENDED FOR CHILDREN.

Pregnancy and breast-feeding

Ask your doctor or pharmacist for advice before using any medicine.

Since available experience concerning human use of timolol during pregnancy and breast-feeding is inadequate, Timo-Stulln® 0.5% should only be used if the doctor considers this to be specifically necessary.

Driving and using machines

Visual impairment, double vision, headache, feeling of weakness, nausea and dizziness may occur in very rare cases. You may then not be able to react fast enough to unexpected or sudden events. Do not drive a car or other vehicles. Do not operate any electrical tools or machines. Do not perform hazardous activities. Take into account that alcohol will reduce your fitness to drive even further.

Important information about some of the ingredients of Timo-Stulln® 0.5%

Benzalkonium chloride can cause eye irritations.

Avoid contact with soft contact lenses. Benzalkonium chloride can lead to discolouration of soft contact lenses.

Contact lenses must be removed before administration and may be replaced 15 minutes after administration at the

earliest.

3. HOW TO USE TIMO-STULLN® 0.5%

Always use Timo-Stulln 0.5% exactly as your doctor has told you. You should check with your doctor or pharmacist if you are not sure. Unless otherwise directed by the doctor, the usual dose is:

Administer 1 drop into the affected eye(s) twice a day at the beginning of the treatment.

Once regular check-ups confirm that the desired value of the intraocular pressure has been reached, the doctor may prescribe a dose of 1 drop a day.

If you take additional beta-receptor blockers, further reduction of ocular pressure must be expected. It should therefore be verified whether local administration of timolol-containing eye drops is still indicated under such circumstances.

Please consult your doctor if you take or plan to discontinue additional beta-receptor blockers.

Pressure reduction may be delayed or weaker in patients with a strongly pigmented iris.

Method of administration

For application to the eye.

Open the bottle by turning the cap clockwise. Hold the bottle slanted and press it slightly to dispense the eye drops. After use, immediately close the bottle by turning the cap anti-clockwise.

Place Timo-Stulln® 0.5% in the eye dropwise. Tilt your head slightly backwards, look upwards and gently pull the lower lid away from the eye to form a pouch. Slightly squeeze the lower part of the bottle to dispense one drop into the pouch. Do not allow the dropper tip to touch the eye. Close your eyes slowly after instilling the drops.

Possible systemic side effects may be reduced by exerting pressure on the tear duct with a finger for about 1 minute after the administration.

Duration of administration

Treatment with Timo-Stulln® 0.5% generally takes place over a longer period of time. Any interruptions or dosage changes must be according to instruction by the doctor.

Please consult your doctor or pharmacist if you think the action of Timo-Stulln® 0.5% might be too strong or too weak.

If you use more Timo-Stulln® 0.5% than you should

Please seek medical advice in case of overdosage. Your doctor will decide if treatment of overdosage is necessary.

If you forget to use Timo-Stulln® 0.5%

Do not take a double dose to make up for a missed dose.

Elevated intraocular pressure may persist over a long period of time without the patient experiencing any noticeable symptoms. The eye can nevertheless be damaged permanently if the pressure is not controlled. Regular treatment is therefore imperative for maintaining your sight.

If you stop using Timo-Stulln® 0.5%

Do not stop treatment without consulting the doctor.

If you have any further questions on the use of this product, ask your doctor or pharmacist.

4. POSSIBLE SIDE EFFECTS

Like all medicines, Timo-Stulln® 0.5% can cause side effects, although not everybody gets them.

Side effects are assessed on the basis of the following frequency ratings:

Very common:	More than 1 in 10 treated persons
Common:	Less than 1 in 10, but more than 1 in 100 treated persons
Uncommon:	Less than 1 in 100, but more than 1 in 1,000 treated persons
Rare:	Less than 1 in 1,000, but more than 1 in 10,000 treated persons
Very rare:	Less than 1 in 10,000 treated persons, or not known

Possible side effects:

Eyes

Eye irritations (which may become serious) as well as a sensation of dryness, reduced corneal sensitivity and visual disturbances may occur. Double vision and a drooping upper eyelid or complications after operations to reduce ocular pressure have been reported in few isolated cases.

Action may persist for several days after discontinuation. If timolol-containing eye drops are discontinued after a longer period of administration, a pressure reducing effect may continue for 2–4 weeks. Beta-blockers administered to one eye may also have a pressure reducing effect on the untreated eye.

Systemic side effects

When using eye medication, side effects may also occur in areas of the body other than the eye. Use of Timo-Stulln® 0.5% may be associated with the same side effects as for oral administration of timolol maleate.

The following side effects either occurred during use in the eye of this or other pharmaceutical forms of timolol-containing eye drops in clinical trials, or were reported since placement on the market:

Respiratory tract

Timo-Stulln® 0.5% may increase airway resistance. Shortness of breath (bronchial spasm) may occur in patients with a predisposition to spasms of the respiratory tract (e.g. bronchial asthma).

Skin

Hypersensitivity reactions such as e.g. rash or hives have been observed. Loss of hair (alopecia), psoriasis-like skin rash or worsening of psoriasis.

Cardiovascular system

Side effects such as slower or irregular pulse, drop in blood pressure, palpitations, heart block and heart failure may occur, particularly in patients with a heart condition. Further possible side effects include swelling (oedema), intermittent claudication, Raynaud syndrome, cold hands and feet as well as pain in the chest area. Reduced blood flow to the brain may also occur. Cardiac arrest and stroke have been reported in extremely rare cases.

Endocrine system

Masking of the symptoms of low blood sugar in insulin-dependent diabetics.

Nervous system and mental disorders

Headache, drowsiness, altered mood or other signs of limited brain performance, nightmares, memory loss, increase in symptoms in myasthenia gravis, abnormal sensations in the extremities (paraesthesia), reduced libido.

Functional disorders of the ear and inner ear

Sounds in the ear (tinnitus).

Gastrointestinal complaints

Nausea, diarrhoea, diarrhoea with vomiting (dyspepsia), dry mouth.

Functional disorders of the reproductive system and breast

Hardening of the spongy bodies of the penis.

General complaints

Dizziness, feeling of weakness

Other possible side effects

In very rare cases, patients with pronounced corneal defects who were treated with phosphate-containing eye drops developed corneal clouding due to formation of calcium phosphate.

If any of the side effects gets serious, or if you notice any side effects not listed in this leaflet, please tell your doctor or pharmacist.

Reporting of side effects

If you get any side effects, talk to your doctor or pharmacist or nurse. This includes any possible side effects not listed in this leaflet.

You can report any suspected side effects associated with the use of health products to Health Canada by:

- Visiting the Web page on Adverse Reaction Reporting (<https://www.canada.ca/en/health-canada/services/drugs-health-products/medeffect-canada/adverse-reaction-reporting.html>) for information on how to report online, by mail or by fax; or
- Calling toll-free at 1-866-234-2345.

NOTE: Contact your health professional if you need information about how to manage your side effects. The Canada Vigilance Program does not provide medical advice.

5. HOW TO STORE TIMO-STULLN® 0.5%

Keep out of the reach and sight of children.

Do not use the medicine after the expiry date which is stated on the outer carton (see “EXP”). The expiry date refers to the last day of the month shown.

Storage conditions

Store between 15°C and 25°C. Keep the bottle tightly closed and store bottle in outer carton in order to protect from light.

Information regarding shelf life after opening or preparation

The bottle can be used for four weeks after first opening.

6. FURTHER INFORMATION What Timo-Stulln® 0.5% contains

The active substance is: timolol maleate

1 ml eye drops contains 6.84 mg timolol maleate corresponding to 5.0 mg timolol.

The other ingredients are: benzalkonium chloride, sodium dihydrogen phosphate monohydrate, sodium monohydrogen phosphate dodecahydrate (Ph. Eur.), water for injections

What Timo-Stulln® 0.5% looks like and contents of the pack

Packs with eye drop bottles with screw cap and safety ring made of LDPE:

- 1 x 5 ml
- 3 x 5 ml

For more information contact:

Pharma STULLN Inc.

info@phamastulln.ca

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www.pharmastulln.ca

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