# Clinical study of coparision of pterygium surgery with and without Mitomycin-C on bare sclera technique

Gupta ML<sup>1</sup>, Meena RK<sup>2</sup>, Bhardwaj V<sup>3</sup>

<sup>1</sup>Dr. Mohan Lal Gupta, Associate Professor, Department. Of Ophthalmology, Jhalawar Medical College, Jhalawar, Rajasthan, Dr. Ravindra Kumar Meena, Assistant Professor, Department Of Ophthalmology, Jhalawar Medical College, Jhalawar, Rajasthan, Dr Veena Bhardwaj, Professor Department Of Ophthalmology, Jhalawar Medical College, Jhalawar, Rajasthan, India

Address for correspondence: Dr. Mohan Lal Gupta, Email: dr.mohanlalgupta14@gmail.com

.....

# **Abstract**

**Introduction:** A pterygium is a triangular wedge of fibro-vascular conjunctival tissue that appears on the epibulbar conjunctiva, which can be removed by various methods. Recurrence of Pterygium after exicision is a very common problem encountered by ophthalmologist. Several methods have been suggested to avoid these recurrences. We studied the recurrence rate of pterygium after application of intraoperative mitomycin C (0.04%). **Method:** This is a retrospective study of fifty eyes in fifty patients who underwent pterygium excision by the same surgeon using intraoperative topical mitomycin C (25 patients) and without using mitomycin c(25 patients) during September 20014—September 2015 in the ophthalmology department at Jhalawar medical college, Jhalawar, Rajasthan. 0.04% Mitomycin applied to bare sclera after excision for two minutes by swab sticks. Postoperative follow up period was 6 months. Outcomes measured in the form of recurrence and complications were analyzed. **Results:** In Group A with use of mitomycin C there was no recurrence after 6 months follow-up while in Group B recurrence was seen in 5 patients within 3-6 months, however in group A 1 patient had scleral thinning. **Conclusion:** Intraoperative administration of mitomycin C 0.04% is safe and effective to prevent pterygium recurrences.

**Keywords:** Bare Sclera, Mitomycin C, Pterygium, Recurrence.

## Introduction

A pterygium is a wing-shaped growth of fibrovascular subconjunctival tissue onto the cornea. Several hypotheses have been ascribed to its etiology. Prevalence rates range from 0.7-31% in various populations around the world and the condition is more common in warm, dry climates. Ultraviolet radiation exposure is a major risk factor for its development [1, 2].

Some findings suggest that an immunologic dysfunction plays a role in the pathogenesis of pterygium, and recent studies have shown that pterygia have increased levels of proangiogenic growth factors such as basic fibroblast growth factor (FGF) and vascular endothelial growth factor (VEGF) [3,4].

Manuscript received: 15<sup>th</sup> Nov 2015 Reviewed: 1<sup>st</sup> Dec 2015 Author Corrected: 4<sup>th</sup> Dec 2015 Accepted for Publication: 10<sup>th</sup> Dec 2015 Pterygium is a worldwide condition with a "pterygium belt" between the latitudes 30°north and south of the equator [5,6] it is more common in sub-tropical and tropical areas [7]. Pterygium is prevalent in Hong Kong, situated 22°north of the equator [2]. Ultraviolet radiation exposure is a major risk factor for its development [1,3].

The first line of treatment for primary pterygium is surgical excision [1]. In pterygium surgery a variety of surgical procedures are in use. The bare sclera technique is still common because of its simplicity. There are overwhelming evidences that the sole use of the bare sclera technique is associated with a high risk of local recurrence. Adjunctive therapies, as betairradiation and antimetabolic drugs, like mitomycin C (MMC) are used to decrease the recurrence rate [8].

Simple excision carries a high recurrence rate ranging from 24%–89% [2, 7]. Recurrent pterygia are more difficult to treat and various other treatment modalities are usually indicated i.e. Conjunctival autograft transplantation and triethylene-thiophosphor- amide (Thiotepa), an antimitotic, Mitomycin C [7].

To prevent recurrence, two major adjunctive therapies are usually performed: (1) the application of antimetabolites, and (2) conjunctival or limbal autograft. Although many other therapeutic modalities have been proposed, it requires extensive studies of efficiency and safety before a new procedure can be

considered as ideal [6]. Addition of MMC at various concentrations has been reported to be effective in preventing recurrence [2]. Mitomycin C is an antibiotic-antineoplastic agent. It is a metabolic inhibitor extracted from Streptomyces caespitosus that inhibits DNA synthesis [1] resulting in an inhibition of the cellular proliferation for a long time [8] which selectively inhibits the synthesis of DNA, cellular RNA, and protein [7]. The mechanism of action seems to be inhibition of fibroblast proliferation at the level of the episclera. However, MMC may cause devastating complications such as scleral necrosis and microbial infections [2].

## **Material and Method**

A prospective, comparative clinical study was conducted from September 2014 until September 2015 in the department. Of ophthalmology at Jhalawar medical college, Jhalawar. Fifty eyes of 50 patients with primary pterygium were included in the study. Informed consent was obtained from all patients before recruitment. All patients underwent full ophthalmologic examination before and after surgery, including visual acuity, slit-lamp examination, fundoscopy On slit lamp examination grading of Pterygium was done based on extent of corneal involvement: Grade I – crossing limbus, Grade II – midway between limbus and pupil, Grade III – reaching up to pupillary margin, Grade IV – crossing pupillary margin.









Figure 1: Pterygium grading severity (1-4); a) grade 1- at the conjunctiva (to the limbus); b) grade 2-1-2mm across the cornes; c) grade 3-te the pupil: d) grade 4- half way across the pupil

# **Grades of Pterygium**

# **Exclusion criteria**

- 1. Collagen vascular disease or other
- 2. Autoimmune disease,
- 3. Pregnancy,
- 4. Ocular surface pathology or infection, and
- 5. Previous limbal surgery,
- 6. Diabetes mellitus,
- 7. Recurrent pterygium, Pseudo Pterygium,
- 8. Lime injury associated symblephron,
- 9. History of ocular injuries,
- 10. Corneal opacities and degeneration were excluded.

All 50 patients divided in two groups 25 in each: Group A in which 0.04% mitomycin c was used. Group B: without mitomycin c use.

All surgeries were done by single surgeon. Steps of surgery (1) eye was anesthesised by instillation of proparacaine hydrochloride 0.5% eyedrops in conjunctival sac two-to-three times every one-to-two minute's (2) eye was paint by butadiene and draped

after that wire speculum was applied.(3) Pterygium was excised by 11 number bard - parker blade and separated from underlines sclera. The remaining sub-conjunctival degenerative part of Pterygium was excised with wescott scissor. The conjuctival and corneal surfaces were smoothened by scraping with a bared – parker blade and bleeding vessels were cauterized by electric cautery. (4) In patients of Group A, 0.04% percent of mitomycin C applied on bare sclera by cotton swab for two minutes and after that sclera is irrigated by 50 ml of normal saline.(5) In group B, sclera was left as such without mitomycin C application. (6) Eye was patched with antibiotic ointment for 24 hours.

On Postoperatively 0.3% tobramycin sulphate eye drop applied six times a day for 15 days, 1% predinsolone acetate eye drop four times a day for one week, followed by tapering dose for subsequent three weeks and 0.5% carboxymethylcelloose eye drop four times a day for one month were given. Patients were asked to follow up on 1-day, 7-day, 1-month, 3-month and every

6-monthly regarded complications and recurrence of Pterygium.

The main postoperative outcomes were recurrence of Pterygium and complication of mitomycin C. Recurrence was defined as fibro-vascular proliferation invading the cornea more than 1.5 mm at the site of previously excised Pterygium. The complications were in terms of postoperative corneal complication and complications of mitomycin C i.e. puncate keratopathy, corneal melting, scleral necrosis, corneal perforation, cataract and secondary glaucoma.

#### **Results**

Fifty eyes of fifty patients were operated for Pterygium surgery in which 25 patients were included in Group A, in which mitomycin C was applied and rest 25 patients were included in Group B and they left with bare sclera. All patients were followed up for 6 months.

**In Group A:** Mean Age of surgically managed patients was 46.08± 9.11 (range 30 - 60 yrs) out of which 60% were males (15) and 40% were females (10), 20 Patients have nasal Pterygium and 5 patients have bilateral Pterygium. 5 patients were having Grade IV Pterygium, 15 patients were having Grade III Pterygium, 3 patients were having Grade I Pterygium and 2 patients were having Grade I Pterygium.

**In Group B:** Mean Age of surgically managed patients was  $45.52\pm 9.90$  (range 30 - 60 yrs) out of which 60% were males (15) and 40% were females (10), 15 Patients have nasal Pterygium and 10 patients have bilateral Pterygium. 7 patients were having Grade IV Pterygium, 10 patients were having Grade III Pterygium, 5 patients were having Grade II Pterygium and 3 patients were having Grade I Pterygium. In Group A, 1 patient developed thinning of sclera after six months and there is no recurrence after follow up. In Group B, 5 patients developed recurrence after six month follow up.

Table No 1: Distribution of cases according to site of pterygium

Gender	Nasal	Nasal/temporal	Total
Male	22	08	30
Female	13	07	20
Total	35	15	50

Chi sq=0.397,p value=0.529(ns)

Table No 2: Comparison of visual acuity before surgery

Visual Acuity	Group A before Surgery	Group B before surgery
6/6	11	11
6/9	02	02
6/12	12	12
Total	25	25

Chi sq=0.00,pvalue=1.00(ns)

Table No. 3: Comparison of visual acuity after surgery

Visual acuity	Group A after surgery	Group B after surgery	
6/6	14	14	
6/9	11	11	
6/12	00	00	
Total	25	25	

Chi sq=0.00,p value=1.00(ns)

Table No 4: Distribution of cases according to recurrence rate

Group A	Group B	Z value	P value	significance
00/25	05/25	2.2704	0.0232	significant

Table 5: Distribution of cases according to complication

Group A	Group B	Z value	P value	significance
01/25	01/25	0.9947	0.3199	Non significant

#### Discussion

There were no statistically significant differences observed in age, gender, laterality and visual acuity among the groups. All patients completely recovered, and no other abnormal ocular or systemic complications were observed during the six-month follow-up period except scleral thinning in one case in group A. our main outcome measure was recurrence rate Generally, pterygium recurrences occur during the first 6 months after surgery. A number of factors such as the type of pterygium, age of the patient, environmental agents, and surgical technique may be responsible [9] but No recurrence was observed up to one month after surgery in group A However, at three months, recurrence was observed in three eyes (12%) and at four months in two eyes (8%) in the group B. In Group A MMC were used intraoperatively at a concentration of 0.4 mg/ml over bare sclera for 3 min. The rate of recurrence was 00% in comparison with 38% reported by Chen et al.,[10] and 10.5% by Manning et al.,[11] with the application of 0.4 mg/ml for 3 min. This concurs with previous studies on intraoperative application of MMC with a rate of recurrence of 25% [12]. Various concentrations of MMC with different durations of application have been used, but the minimal safe and effective dosage and application time are still not certain [13].

Because recurrence rates differed significantly at three months (p = 0.02372) these observation had enough support by various study A L Young et al[5] Allan BDS et al[14].

One and only complication which was seen in group A was scleral thinning which was supported by Rubinfeld et al[15], Safianik B et al[16] Zhivov A et al[17], Peponis V et al[18].

The limitations of the study were the short follow-up period and the small size of the study groups. With a longer follow-up period, recurrence rates and side effects related to the adjunctive drugs could be analyzed more accurately. Larger sample sizes could make the statistical analyses stronger. Overall we can say that 0.04% mitomycin c is safe and effective to prevent recurrence after pterygium excision.[19]

#### Conclusion

0.04% mitomycin c is effective to prevent recurrence after perygium surgery.

Funding: Nil, Conflict of interest: None.

Permission of IRB: Yes

#### Reference

- 1. Hwang S, Choi S<sup>1</sup>. A Comparative Study of Topical Mitomycin C, Cyclosporine, and Bevacizumab after Primary Pterygium Surgery. Korean J Ophthalmol. 2015 Dec;29(6):375-81. doi: 10.3341/kjo.2015.29.6.375. Epub 2015 Nov 25.
- 2. Ozsutcu M, Ayintap E, Akkan JC, Koytak A, Aras C. Repeated bevacizumab injections versus mitomycin C in rotational conjunctival flap for prevention of pterygium recurrence. Indian J Ophthalmol. 2014 Apr;62(4):407-11. doi: 10.4103/0301-4738.120220.
- 3. Kria L, Ohira A, Amemiya T. Immunohistochemical localization of basic fibroblast growth factor, platelet derived growth factor, transforming growth factor-beta and tumor necrosis factor-alpha in pterygium. Acta Histochem. 1996;98:195–201.
- 4. Aspiotis M, Tsanou E, Gorezis S, Ioachim E, Skyrlas A, Stefaniotou M, Malamou-Mitsi V. Angiogenesis in pterygium: study of microvessel density, vascular endothelial growth factor, and thrombospondin-1. Eye (Lond). 2007 Aug;21(8):1095-101. Epub 2006 Jun 23.
- 5. Young AL, Leung GY, Wong AK, Cheng LL, Lam DS. A randomised trial comparing 0.02% mitomycin C and limbal conjunctival autograft after excision of

- primary pterygium. Br J Ophthalmol. 2004 Aug;88(8):995-7.
- 6. Ma DH, See LC, Liau SB, Tsai RJ. Amniotic membrane graft for primary pterygium: comparison with conjunctival autograft and topical mitomycin C treatment. Br J Ophthalmol. 2000 Sep;84(9):973-8.
- 7. Oguz H, Basar E, Gurler B. Intraoperative application versus postoperative mitomycin C eye drops in pterygium surgery. Acta Ophthalmol Scand. 1999 Apr;77(2):147-50.
- 8.Gabor Koranyi,1 Ditte Artze´ n,2 Stefan Seregard2 and Eva Dafgard Kopp2 Intraoperative mitomycin C versus autologous conjunctival autograft in surgery of primary pterygium with four-year follow-up. Acta Ophthalmol. 2012: 90: 266–270 a 2010 The Authors Journal compilation a 2010 Acta Ophthalmol doi: 10.1111/j.1755-3768.2010.01936.x.
- 9. Mutlu FM, Sobaci G, Tatar T, Yildirim E. A comparative study of recurrent pterygium surgery: Limbal conjunctival autograft transplantation versus mitomycin C with conjunctival flap. Ophthalmology. 1999;106:817–21. [PubMed]
- 10. Chen PP, Ariyasu RG, Kaza V, LaBree LD, McDonnell PJ. A randomized trial comparing mitomycin C and conjunctival autograft after excision of primary p terygium. Am J Ophthalmol. 1995 Aug;120(2):151-60.
- 11. Manning CA, Kloess PM, Diaz MD, Yee RW. Intraoperative mitomycin in primary pterygium excisio n. A prospective, randomized trial. Ophthalmology. 1997 May;104(5):844-8.

- 12. Alpay A, Uğurbaş SH, Erdoğan B. Comparing techniques for pterygium surgery. Clin Ophthalmol. 2009;3:69-74. Epub 2009 Jun 2.
- 13. Lam DS, Wong AK, Fan DS, Chew S, Kwok PS, Tso MO. Intraoperative mitomycin C to prevent recurrence of pterygium after excision: a 30-month follow-up study. Ophthalmology. 1998 May;105(5):901-4; discussion 904-5.
- 14. Allan BD, Short P, Crawford GJ, Barrett GD, Constable IJ. Pterygium excision with conjunctival autografting: an effective and safe technique. Br J Ophthalmol. 1993 Nov;77(11):698-701.
- 15. Rubinfeld RS, Pfister RR, Stein RM, Foster CS, Martin NF, Stoleru S, Talley AR, Speaker MG. Serious complications of topical mitomycin-C after pterygium surgery. Ophthalmology. 1992 Nov;99(11):1647-54.
- 16. Safianik B, Ben-Zion I, Garzozi HJ. Serious corneoscleral complications after pterygium excision with mitomycin C. Br J Ophthalmol. 2002 Mar;86(3):357-8.
- 17. Zhivov A, Beck R, Guthoff RF. Corneal and conjunctival findings after mitomycin C application in pterygium surgery: an in-vivo confocal microscopy study. Acta Ophthalmol. 2009 Mar;87(2):166-72. doi: 10.1111/j.1755-3768.2008.01198.x. Epub 2008 Jun 3.
- 18. Peponis V, Rosenberg P, Chalkiadakis SE, Insler M, Amariotakis A . Fungal sclera keratitis and endophthalmitis following pterygium exci sion. Eur J Ophthalmol. 2009 May-Jun;19(3):478-80.
- 19. Rodriguez JA, Ferrari C, Hernández GA. Intraoperative application of topical mitomycin C 0.05% for pterygium surgery. Bol Asoc Med P R. 2004 Mar-Apr;96(2):100-2.

# How to cite this article?

Gupta ML, Meena RK, Bhardwaj V. Clinical study of coparision of pterygium surgery with and without Mitomycin-C on bare sclera technique. *Int J Med Res Rev* 2016;4(1):33-37. doi: 10.17511/ijmrr.2016.i01.005.

.....