

Glaucoma after open globe injury at a tertiary care university hospital in Central Saudi Arabia

Cumulative incidence and risk factors

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ABSTRACT

الأهداف: دراسة عوامل حدوث الخطر ومعدل الإصابة التراكمي بالزرق بعد جروح العين المفتوحة.

الطريقة: أجريت دراسة استرجاعية على السجلات الطبية لجميع المرضى الذين تم تشخيصهم كحالات جروح عين مفتوحة ومن ثم خضعوا لعملية إصلاح أولية خلال الفترة من يناير 1996م وحتى ديسمبر 2011م في مستشفى الملك عبد العزيز الجامعي، الرياض، المملكة العربية السعودية.

النتائج: خلال فترة الـ 15 عاماً التي استعرضتها الدراسة، تم تحديد 775 مريضاً من الذين خضعوا لعملية إصلاح لإصابات العين المفتوحة. وكانت نسبة الإصابة الكلية بينهم بالزرق 5.3% (41/775) بعد متوسط متابعة قدره (12±6.5) شهراً. وقد كشف تحليل المتغيرات الفردي عن عوامل الخطر الفعلية للإصابة بالزرق وهي: إصابة المنطقة الثانية $p=0.027$ ، والإصابات التي تخترق العين $p=0.0008$ ، وإصابات العدسة $p=0.011$ ، ونزيف الجسم الزجاجي $p=0.002$ ، ووجود جسم غريب داخل العين $p<0.0001$. بينما وقع المرضى في الفئة العمرية من 18 عاماً وأكثر في المنطقة الحرجة ($p=0.054$). وبتطبيق الإنحدار اللوجستي، وجد أن الإصابات التي تخترق العين $p=0.019$ ، وإصابة العدسة $p=0.002$ ، ونزف الجسم الزجاجي $p=0.037$ هي عوامل الخطر المعنوية.

خاتمة: ليس من المستبعد إصابة العين بالزرق بعد جروح العين المفتوحة. كما تعتبر إصابة المنطقة الثانية، والإصابات التي تخترق العين، وإصابة العدسة، ونزف الجسم الزجاجي، ووجود جسم غريب داخل العين أهم عوامل الخطر الفعلية للإصابة بالزرق بعد جروح العين المفتوحة.

Objectives: To investigate the incidence and risk factors of glaucoma after open globe injury (OGI).

Methods: The medical records of all patients admitted with the diagnosis of OGI at the Department of Ophthalmology, King Abdulaziz University Hospital, Riyadh, Kingdom of Saudi Arabia and had undergone primary repair from January 1996 to December 2011 were retrospectively reviewed.

Results: For the 15-year study period, 775 patients who underwent repair of an OGI were identified. The overall risk of post-traumatic glaucoma was 5.3% (41/775) with a mean \pm SD follow-up period of 12 \pm 6.5 months. Univariate analysis revealed that Zone II injury ($p=0.027$), penetrating ocular injury ($p=0.0008$), lens injury ($p=0.011$), vitreous hemorrhage ($p=0.002$), and presence of intraocular foreign body ($p<0.0001$) were significantly associated with glaucoma. Age of more than 18 years was critical ($p=0.054$). Following logistic regression, penetrating ocular injury ($p=0.019$), lens injury ($p=0.002$), and vitreous hemorrhage were significant ($p=0.037$).

Conclusion: Glaucoma after OGI is not uncommon. Zone II injury, penetrating ocular injury, lens injury, presence of vitreous hemorrhage, and presence of an intraocular foreign body were significant risk factors for developing post-traumatic glaucoma.

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Ocular trauma remains an important cause of preventable blindness globally.¹⁻³ Several risk factors are associated with visual loss caused by ocular trauma. Such factors may include; glaucoma, corneal scarring, lens injury, vitreous hemorrhage, retinal and choroidal detachments, and endophthalmitis. Increased intraocular pressure after repair is one of the well known complications, which may persist and require further management. The mechanism of developing glaucoma either goes through meshwork injuries, or inflammation, which may block the outflow of the aqueous humor, leading to secondary glaucoma. Consequently, the function of the trabecular meshwork may get recovered or decompensated.^{4,5} Little is known regarding the risk factors of glaucoma after open ocular trauma in the literature.^{6,7} In the current study, we aimed to evaluate the incidence and potential risk factors for developing glaucoma after repair of open globe injuries (OGIs) at a tertiary care university hospital in Riyadh, Kingdom of Saudi Arabia (KSA).

Methods. The search strategy of the literature was carried out using identified relative key words and targeting PubMed, EBSCO, OVID, and Google scholar. Search findings were retrieved, then filtered to select the more relative ones according to our objectives. The medical records of all patients admitted with the diagnosis of OGI, and had undergone primary repair from January 1996 to December 2011 at the Department of Ophthalmology, King Abdulaziz University Hospital, Riyadh, KSA were retrospectively reviewed. In the current study, an OGI was defined as any traumatic, full-thickness wound of the globe. Patients were excluded for the following reasons: undergoing primary repair elsewhere before presentation; presenting with a diagnosis of endophthalmitis; and undergoing primary enucleation for an irreparably traumatized eye. The location and extent of the OGI (cornea only) was identified as Zone I, corneoscleral within 5 mm of the limbus (Zone II), and corneoscleral extended more than 5 mm beyond the limbus (Zone III). Medical records were reviewed to determine patients' demographics such as: visual acuity at presentation; type, extent, and location of ocular injury; mechanism of the injury (blunt, sharp, missile); presence of an intraocular

foreign body (IOFB); lens injury and vitreous hemorrhage and prolapse; during the follow-up period. Patients included in this study were classified into one of 3 categories according to the time of onset of post-traumatic glaucoma: 1) early stage (\leq one month after ocular trauma); 2) intermediate stage (2-6 months); and 3) late stage ($>$ 6 months).⁸ This study was registered with the institutional review board and adheres to the tenets of the Declaration of Helsinki. Approval was obtained from the Research and Ethics Board of the College of Medicine, King Saud University (approval #E-11-501).

Statistical analysis. Data were collected and stored using a specifically designed Microsoft Excel 2007[®] sheet. Data were then reviewed, managed, and cleaned for analysis with Statistical Package for Social Sciences version 19 (IBM, Chicago, IL, USA) and MedCalc version 11.6 (MedCalc Software, Mariakerke, Belgium). The chi-square test was performed to investigate the association between different potential risk factors and the occurrence of glaucoma as a complication of interest (Fisher exact test whenever indicated). The student's t-test was used to compare means between glaucoma and non-glaucoma cases. A $p < 0.05$ indicated statistical significance. Moreover, a binary logistic regression was conducted to adjust for interaction and other confounding variables where all significant variables were introduced into the model.

Results. The records of 818 patients who had undergone repair of an OGI were reviewed. Of these, 43 records were excluded because of missing or incomplete data. Of the 775 patients included in the current study, 675 (87.1%) were males and 100 (12.9%) were females with a mean \pm standard deviation (SD) age of 25.7 ± 15 years (range: 1-84 years; median - 27 years). We estimated the overall cumulative rate of post-traumatic secondary glaucoma at 5.3% (41/775) over a mean follow-up period of 12 ± 6.5 months (range: 3 months to 13.8 years). In the univariate analysis, the development of post-traumatic glaucoma was significantly associated with Zone II injury ($p=0.027$), penetrating ocular injury ($p=0.0008$), lens injury ($p=0.011$), presence of an intraocular foreign body ($p < 0.0001$), and presence of vitreous hemorrhage ($p=0.002$), whereas age greater than 18 years was critical ($p=0.052$). Logistic regression analysis revealed that the following risk factors were significantly associated with the development of post-traumatic glaucoma: penetrating ocular injury (odds ratio [OR] - 3.5; 95% confidence interval [CI]: 1.41-8.9; $p=0.019$), vitreous hemorrhage (OR - 2.4; 95% CI: 1.53-6.2; $p=0.037$), and lens injury (OR - 2.1; 95% CI: 1.56-7.7; $p=0.002$) (Table 1). During

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Table 1 - Univariate analysis to assess different factors associated with post-traumatic glaucoma (N=775).

| Variable | n (n=41) | (%) | No glaucoma (n=734) | Total | P-value | 95% confidence intervals (LCL - UCL) |
|---|-------------|--------|------------------------|-------|---------|---|
| <i>Age, years</i> | | | | | | |
| ≤18* | 8 | (3.1) | 251 | 259 | 0.052 | 0.952 - 5.449 |
| >18 | 33 | (6.4) | 483 | 516 | | |
| <i>Gender</i> | | | | | | |
| Male* | 37 | (5.5) | 638 | 675 | 0.537 | 0.182 - 2.067 |
| Female | 4 | (4.0) | 96 | 100 | | |
| <i>Vision</i> | | | | | | |
| ≤HM* | 18 | (6.9) | 243 | 261 | - | |
| CF | 8 | (4.3) | 176 | 184 | 0.614 | 0.226 - 1.525 |
| ≥20/200 | 15 | (4.5) | 315 | 330 | 0.216 | 0.295 - 1.383 |
| <i>Mechanism of injury</i> | | | | | | |
| Blunt* | 3 | (5.6) | 51 | 54 | - | |
| Sharp | 24 | (4.5) | 505 | 529 | 0.730 | 0.233 - 4.336 |
| Missile | 14 | (7.3) | 178 | 192 | 0.998 | 0.353 - 7.526 |
| <i>Type of trauma</i> | | | | | | |
| Penetrating | 35 | (4.7) | 717 | 752 | 0.0008 | 0.048 - 0.458 |
| Perforating* | 6 | (26.1) | 17 | 23 | | |
| <i>Extent of trauma</i> | | | | | | |
| 10 mm* | 34 | (5.0) | 652 | 686 | - | |
| 10-20 mm | 6 | (7.7) | 72 | 78 | 0.285 | 0.529 - 4.027 |
| >20 mm | 1 | (9.1) | 10 | 11 | 0.435 | 0.0429 - 4.027 |
| <i>Location of trauma</i> | | | | | | |
| Limited to cornea (Zone I)* | 21 | (4.1) | 495 | 516 | - | |
| Corneoscleral not extended (Zone II) | 19 | (7.9) | 220 | 239 | 0.027 | 1.012 - 4.063 |
| Corneoscleral extended (Zone III) | 1 | (5.0) | 19 | 20 | 0.574 | 0.0285 - 8.594 |
| <i>Lens injury</i> | | | | | | |
| Yes | 19 | (8.5) | 204 | 223 | 0.011 | 1.121 - 4.441 |
| No* | 22 | (4.0) | 530 | 552 | | |
| <i>IOFB</i> | | | | | | |
| Yes | 5 | (0.8) | 634 | 639 | <0.0001 | 0.008 - 0.058 |
| No* | 36 | (26.5) | 100 | 136 | | |
| <i>Vitreous hemorrhage</i> | | | | | | |
| Yes | 7 | (15.6) | 38 | 45 | 0.002 | 1.320 - 9.382 |
| No* | 34 | (4.7) | 696 | 730 | | |
| <i>Vitreous prolapse</i> | | | | | | |
| Yes | 1 | (5.6) | 17 | 18 | 0.998 | 0.025 - 7.707 |
| No* | 40 | (5.3) | 717 | 757 | | |

*reference group in the analysis. HM - hand movement, CF - counting fingers, IOFB - intraocular foreign body, LCL - lower confidence interval limits, UCL - upper confidence interval limits

the early stage (≤one month), high intraocular pressure (IOP) developed in 11 patients (26.8%) because of unremoved lens particles in 6 patients (14.6%) due to inflammation, and in 3 patients (7.3%) due to hyphema. During the intermediate phase (2-6 months), synechial angle closure was the recognized cause in 9 patients (22%), ghost cell in 3 patients (7.3%), and unremoved lens particles in 2 patients (4.9%) of those who developed post-traumatic glaucoma. During the late stage (>6 months), the IOP gradually rose in 4 patients (9.8%) because of angle recession, and in 3 patients (7.3%) because of synechial angle closure

(Table 2). At the last follow-up visit, all 41 patients who had developed post-traumatic secondary glaucoma underwent gonioscopy. This procedure revealed that 23 patients (56.1%) had open-angle glaucoma, 14 patients (34.1%) had closed-angle glaucoma, and 4 patients (9.8%) had angle recessions.

Discussion. In this retrospective cohort study, we determined the overall incidence of glaucoma after open ocular injury to be 5.3%. The risk factors significantly associated with the development of glaucoma in the present study were: Zone II injury; penetrating ocular

Table 2 - Causes and time of onset of post-traumatic glaucoma.

| Stage | Causes of glaucoma |
|-------------------------------------|---|
| Early (\leq one month) (n=20) | Unremoved lens particles (n=11) Inflammatory (n=6) Hyphema (n=3) |
| Intermediate (2-6 months) (n=14) | Synechial angle closure (n=9) Ghost cell (n=3) Unremoved lens particles (n=2) |
| Late ($>$ 6 months) (n=7) | Angle recession (n=4) Synechial angle closure (n=3) |

injury; lens injury; presence of vitreous hemorrhage; and presence of an intraocular foreign body during the follow-up period.

The incidence of post-traumatic glaucoma is unpredictable in closed and open eye injuries. Several studies have estimated the incidence rate after a closed eye injury to be between 3.4 and 19%. Apparently, very few studies have estimated the incidence of glaucoma after a penetrating ocular injury. Based on data from the United States Eye Injury Registry (USEIR), Girkin et al⁶ determined the risk of developing post-traumatic glaucoma after a penetrating eye injury to be 2.67% over a 6-month follow-up period. In our study, the incidence of post-traumatic glaucoma was 5.3%. Recently, Turalba et al⁷ estimated the prevalence at 17%, however according to their criteria, among their series, those with ocular hypertension ≥ 22 mm Hg at one visit and treated were included, which may have overestimated their prevalence. Additionally, although our incidence is higher than Girkin's,⁶ this may be explained by our longer mean follow-up duration (12 ± 6.5 months).

Glaucoma that occurs after open eye injuries tends not to follow any predictable patterns and is most probably multifactorial in origin.⁹ For example, it may develop following inflammation, which can lead to synechial angle closure, particularly in patients with a shallow or flat anterior chamber. Other risk factors for developing glaucoma following an open eye injury include the alteration of the trabecular meshwork caused by blood cells inflammatory debris, lens particles, and ghost cells associated with long-lasting vitreous hemorrhage.⁹ In agreement with what has been reported in the USEIR study,⁶ we found a significant association between the development of glaucoma, lens injury, and presence of vitreous hemorrhage during the follow-up period. Although age was not explicitly significant, its probability value was quite critical ($p=0.054$), which reflects a high probability of reaching the significance level if the sample size were slightly increased. This finding is highly supported by the literature,^{6,7} where

an association between traumatic glaucoma and an increase in age is expected because the trabecular meshwork might be more compromised in this age group. In addition, it is well known that the incidence of primary open-angle glaucoma (POAG) is higher in the healthy elderly population.^{10,11} Girkin et al⁶ detected an association between glaucoma and other risk factors reported here (poor visual acuity at presentation, lens injury, presence of vitreous hemorrhage) might be explained by increased severity of trauma because these clinical signs are commonly observed in more serious ocular injuries. Moreover, Turalba et al⁷ found that the increased age, hyphema, lens injury, and Zone II injury are potential risk factors for developing ocular hypertension after OGI.

Additionally, Girkin et al⁶ found no significant association between the presence of retained IOFB and glaucoma. This disagreement with our study findings may be explained by the differences in the removal time of IOFB. Our clinical protocol involved the removal of the IOFB one to 2 weeks after primary repair, whereas the protocol followed by Girkin et al⁶ involved the removal of most IOFB during the initial examination, if detected. Turalba et al⁷ did not look for IOFB as a risk factor due to exclusion of a significant number of patients with IOFB from their study. One possible reason for such a positive association in our study is the longer duration (one to 2 weeks) of ocular retention of foreign bodies containing ferrous metals, which may lead to siderosis and iron staining of the trabecular structures.⁶ However, this explanation seems unlikely because the period of time (one to 2 weeks) is rather short for siderosis to develop.

The current study have faced a number of limitations as per its retrospective nature, which hindered the collection of all required information, and the relatively small number of cases. In the future, a prospective study with trauma registry program might enable better in depth understanding of the risk factors for glaucoma and other post-traumatic ocular disease

In conclusion, we determined the incidence of glaucoma after open ocular injury to be 5.3%. Several baseline risk factors were identified that were significantly and independently associated with the development of glaucoma, including Zone II injury, penetrating ocular injury, lens injury, presence of vitreous hemorrhage, and the presence of an intraocular foreign body during the follow-up period. Patients with OGI must be carefully observed, and special attention should be paid to any early signs of glaucomatous changes, particularly in patients who are considered at high risk.

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References

1. McCall BP, Horwitz IB, Taylor OA. Occupational eye injury and risk reduction: Kentucky workers' compensation claim analysis 1994-2003. *Inj Prev* 2009; 15: 176-182.
2. Négrel AD, Thylefors B. The global impact of eye injuries. *Ophthalmic Epidemiol* 1998; 5: 143-169.
3. Thylefors B. Epidemiological patterns of ocular trauma. *Aust N Z J Ophthalmol* 1992; 20: 95-98.
4. De Leon-Ortega JE, Girkin CA. Ocular trauma-related glaucoma. *Ophthalmol Clin North Am* 2002; 15: 215-223.
5. Sihota R, Kumar S, Gupta V, Dada T, Kashyap S, Insan R, et al. Early predictors of traumatic glaucoma after closed globe injury: trabecular pigmentation, widened angle recess, and higher baseline intraocular pressure. *Arch Ophthalmol* 2008; 126: 921-926.
6. Girkin CA, McGwin G Jr, Morris R, Kuhn F. Glaucoma following penetrating ocular trauma: a cohort study of the United States Eye Injury Registry. *Am J Ophthalmol* 2005; 139: 100-105.
7. Turalba AV, Shah AS, Andreoli MT, Andreoli CM, Rhee DJ. Predictors and Outcomes of Ocular Hypertension After Open-globe Injury. *J Glaucoma* 2012 Aug 29. [Epub ahead of print]
8. Bai HQ, Yao L, Wang DB, Jin R, Wang YX. Causes and treatments of traumatic secondary glaucoma. *Eur J Ophthalmol* 2009; 19: 201-206.
9. Milder E, Davis K. Ocular trauma and glaucoma. *Int Ophthalmol Clin* 2008; 48: 47-64.
10. Al-Mansouri FA, Kanaan A, Gamra H, Khandekar R, Hashim SP, Al Qahtani O, et al. Prevalence and determinants of glaucoma in citizens of Qatar aged 40 years or older: a community-based survey. *Middle East Afr J Ophthalmol* 2011; 18: 141-149.
11. Kim M, Kim TW, Park KH, Kim JM. Risk factors for primary open-angle glaucoma in South Korea: the Namil study. *Jpn J Ophthalmol* 2012; 56: 324-329.

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Al-Obeidan SA, Mousa A, Naseem A, Abu-Amero KK, Osman EA. Efficacy and safety of non-penetrating deep sclerectomy surgery in Saudi patients with uncontrolled open angle glaucoma. *Saudi Med J* 2013; 34: 54-61.

Khandekar RB, Jaffer MA, Al-Harthy H. Should the health information glaucoma be hospital based or community based? *Saudi Med J* 2008; 29: 1156-1158.

Rismanchian A, Eslami F, Moeini H, Attarzade H, Naderibeni A. Efficacy of the latanoprost versus timolol/dorzolamide combination therapy in patients with primary open angle glaucoma. *Saudi Med J* 2008; 29: 384-387.