Ophthalmic viscoelastic device injection for the treatment of flat anterior chamber after trabeculectomy: a case series study

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Purpose: Flat anterior chamber (FAC) in association with overfiltration is a complication after trabeculectomy. The aim of the current study was to investigate the efficacy of injection of an ophthalmic viscoelastic device (OVD) to treat FAC after trabeculectomy in a retrospective consecutive case series study.

Materials and methods: Twenty-five consecutive patients who underwent injection of OVD for the treatment of FAC after trabeculectomy were analyzed. The primary outcome variable was the qualified success of trabeculectomy, which was defined at three different levels by the achievement of one of three IOP-related criteria with or without glaucoma medication. Kaplan–Meier plots were generated to estimate survival functions for trabeculectomy with OVD injection.

Results: Twenty-one patients (21/25 = 84.0%) recovered from FAC by one OVD injection. The postoperative intraocular pressure (IOP) measured at 12 months, 24 months, 36 months, 48 months, and 60 months was 14.7 ± 3.6 mmHg, 15.6 ± 7.0 mmHg, 14.2 ± 2.8 mmHg, 14.2 ± 3.8 mmHg, and 14.6 ± 2.7 mmHg, respectively. All postoperative IOP measurements were significantly lower than the preoperative IOP ($P < 0.001$). The probabilities of qualified success for 5 years after trabeculectomy with OVD injection were as follows: 24.4% (IOP ≤ 21 mmHg and $\geq 20\%$ reduction of preoperative IOP), 16.5% (IOP ≤ 16 mmHg and $\geq 20\%$ reduction) and 6.3% (IOP ≤ 12 mmHg and $\geq 30\%$ reduction). One patient, who had undergone deep lamellar keratoplasty, experienced a complication associated with the injection of the viscoelastic material, Descemet's membrane detachment.

Conclusion: Injection of OVD may be effective for the treatment of FAC; however, the long-term success rate was relatively low.

Keywords: glaucoma, trabeculectomy, flat anterior chamber, ophthalmic viscoelastic device injection

Introduction

The incidence of flat anterior chamber (FAC), a complication associated with overfiltration after a trabeculectomy with mitomycin C, is reported to be 13%–24%.^{1,2} Persistent FAC causes peripheral anterior synechiae, cataracts, and endothelial dysfunction.^{3–6} FAC has been successfully treated with atropine eye drops, a firmly applied eye patch, balanced salt solution injections, sodium hyaluronate, air bubbles, and injection of perfluoropropane or sulfur hexafluoride gas into the anterior chamber.^{7–11} Fisher et al⁸ described using sodium hyaluronate to treat FAC in 1982, and Hoffman et al⁷ reported using Healon 5[®] (Abbott Medical Optics, Inc, Santa Ana, CA, USA) to treat FAC after trabeculectomy in 2002.

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This retrospective, consecutive case series study investigated the efficacy of injecting an ophthalmic viscoelastic device (OVD) to treat FAC after trabeculectomy.

Materials and methods

This study followed the tenets of the Declaration of Helsinki and was approved by the Ethics Committee of the Keio University School of Medicine (authorization number 2010214). This study included 25 consecutive patients who received OVD injections to treat FAC occurring after a trabeculectomy with mitomycin C between April 1, 2004 and March 31, 2008. In this retrospective study, we included all the subjects who underwent viscoelastic injection for the treatment of grade 2 FAC (entire iridocorneal touch without contact between the lens and the cornea) without conjunctival leakage.¹² There were no exclusion criteria. These patients had no leakage from the conjunctiva in the early postoperative period. All trabeculectomy surgeries were performed by one of three glaucoma specialists educated at the same institute. The trabeculectomies were performed using a limbal-based conjunctival flap, sub-Tenon dissection, and hemostasis. A half-thickness 4.0×4.0 mm rectangular scleral flap was fashioned in the superior area. A 4.0×4.0 mm sponge soaked in a 0.4 mg/mL solution of mitomycin C was applied for 3–5 minutes, and the area was irrigated thoroughly with 500 cc of physiological saline. The trabeculectomy was followed by a peripheral iridectomy. The scleral flap was sutured with 10-0 nylon sutures (Mani, Inc, Tochigi, Japan). All patients were given a postoperative regimen of topical 0.5% levofloxacin (Cravit® ophthalmic solution; Santen Pharmaceutical Co, Ltd, Osaka, Japan) three times per day, and betamethasone eye drops (Sanbetasone®; Santen Pharmaceutical Co, Ltd) four times per day.

OVD injections were performed after applying oxybutyrocaine hydrochloride (Benoxil ophthalmic solution; Santen Pharmaceutical Co, Ltd) topically as an anesthetic, and disinfected with 5% povidone-iodine (Isodine; Meiji Seika Co, Ltd, Tokyo, Japan). An Opegan Hi® (Santen Pharmaceutical Co, Ltd), Healon®, or Healon 5® (AMO Japan, Inc, Tokyo, Japan) OVD was injected into the anterior chamber through a paracentesis made during the trabeculectomy. Using a 23 gauge blunt cannula, the OVD was injected until the anterior chamber was deepened sufficiently. After the injection, the intraocular pressure (IOP) was checked using a Tono-Pen® XL (Reichert™ Technologies, Inc, Buffalo, NY, USA). If the IOP was less than 5 mmHg, additional OVDs were injected into the anterior chamber until the IOP was greater than 10 mmHg.

We considered the patients' age, sex, type of glaucoma, pre-operative lens status, surgical procedure (with or without cataract surgery), the type of OVD, pre- and post-trabeculectomy IOP, the IOP 24 hours after OVD injection, the number of laser suture lysis procedures, the prevalence of choroidal detachment, treatment prior to the OVD injection, the number of OVD injections required, the time from injection to FAC recovery, and any complications associated with the OVD injection. Goldmann applanation tonometry was used to measure the IOP each day during the admission period, 1 month after surgery, 3 months after surgery, and every 3 months thereafter up to 60 months after surgery.

The primary outcome variable was the qualified success of the trabeculectomy, defined by three levels of IOP-related criteria with or without glaucoma medication, as follows: (1) IOP ≤ 21 mmHg, reduced $\geq 20\%$ from the preoperative level (ie, the last IOP value before surgery); (2) IOP ≤ 16 mmHg, reduced $\geq 20\%$ from the preoperative level; or (3) IOP ≤ 12 mmHg, reduced $\geq 30\%$ from the preoperative level.^{13,17} The surgery was deemed to have failed if these criteria were not satisfied at two consecutive visits, or if a secondary surgery to lower the IOP was required. Kaplan–Meier plots were generated to estimate the survival functions for trabeculectomy with OVD injection. A one-way analysis of variance followed by Bonferroni's multiple comparison test was used for statistical analysis of the IOP before and after trabeculectomy. Significance was defined as $P < 0.05$. Statistical analysis was performed using SPSS software, version 21.0 (IBM Software, Las Vegas, NV, USA).

Results

Demographic characteristics are listed in Table 1. After OVD injection treatment, all subjects had sufficient anterior chamber depth to achieve maintained anterior chamber. None of the subjects had total apposition between the iris and cornea with lens–cornea contact.

The number of OVD injections, the IOP 24 hours after injection, time from the injection until the FAC improved, and the type of injected OVD are shown in Table 2. Of the 25 patients, 21 (84.0%) recovered from FAC with one OVD injection; three patients required two injections, and one patient required four injections. All subjects recovered from FAC after OVD treatment without any other surgical procedures.

The mean preoperative IOP was 28.8 ± 9.2 mmHg. The postoperative IOP measured at 12 months, 24 months, 36 months, 48 months, and 60 months was 14.7 ± 3.6 mmHg, 15.6 ± 7.0 mmHg, 14.2 ± 2.8 mmHg, 14.2 ± 3.8 mmHg,

Table 1 Demographic characteristics of 25 patients given OVD injections

Age (years)	57.5 ± 17.8 (24–85)
Sex	
Male/female	18/7
Eye laterality	
Right/left	16/9
Trabeculectomy/phacotrabeculectomy	17/8
	Number
Type of glaucoma	
POAG	11/25
Uveitis	5/25
Neovascular	1/25
Postvitrectomy	2/25
Other*	6/25
Preoperative lens status	
Phakia	18/25
IOL	6/25
Aphakia	1/25
Postoperative choroidal detachment (serous/hemorrhagic)	15/0
Treatment before OVD injection	
Firm eye patching	8/25
Autologous blood Injection into bleb	2/25

Notes: The data represents average ± standard deviation (range) or percentage.
 *Includes chronic angle-closure glaucoma, postpenetrating keratoplasty, postdeep lamellar keratoplasty, Axenfeld–Rieger syndrome, steroid-induced glaucoma, and exfoliation glaucoma.

Abbreviations: OVD, ophthalmic viscoelastic device; POAG, primary open-angle glaucoma; IOL, intraocular lens.

and 14.6 ± 2.7 mmHg, respectively. All postoperative IOP measurements were significantly lower than the preoperative IOP ($P < 0.001$; Figure 1).

The trabeculectomy postoperative success rate is shown in Figure 2. The probability of 5-year success after trabeculectomy with OVD-treated FAC was 24.4% (criterion A), 16.5% (criterion B), and 6.3% (criterion C) (Figure 2).

One patient, who had undergone deep lamellar keratoplasty, experienced a complication associated with the injection of the OVD – Descemet's membrane detachment. Healon 5® was injected through the paracentesis, and was

Table 2 Properties of OVD injection procedures

Number of viscoelastic injections until FAC improvement	1.3 ± 0.8 (1–4)
IOP 24 hours after injection (mmHg)	7.6 ± 8.7 (0–38)
Duration until FAC improvement after viscoelastic injection (days)	8.1 ± 5.3 (1–18)
Type of injected OVD (number of subjects)	
Healon®	16
Opegan Hi®	5
Healon 5®	4

Note: Data represent the mean ± standard deviation (range).

Abbreviations: OVD, ophthalmic viscoelastic device; FAC, flat anterior chamber; IOP, intraocular pressure.

Figure 1 IOP measured prior to trabeculectomy, 1 month after surgery, and at 3-month intervals from the time of surgery until 60 months after surgery.

Notes: All postoperative IOP levels measured were significantly lower than the preoperative IOP (** $P < 0.001$). Data are presented as the mean ± standard deviation and are censored after reoperation for glaucoma.

Abbreviation: IOP, intraocular pressure; pre, preoperative IOP.

accidentally injected between Descemet's membrane and the stroma where lamellar keratoplasty had been performed. The OVD could not be removed from the area between Descemet's membrane and the stroma by aspiration with the cannula or by air tamponade from the anterior chamber.

Figure 2 Survival curves of surgical failure of trabeculectomy with OVD-treated FAC.

Notes: Criterion A: IOP ≤21 mmHg and a ≥20% reduction of the preoperative IOP. Criterion B: IOP ≤16 mmHg and a ≥20% IOP reduction. Criterion C: IOP ≤12 mmHg and a ≥30% IOP reduction.

Abbreviations: OVD, ophthalmic viscoelastic device; FAC, flat anterior chamber; IOP, intraocular pressure.

Therefore, the donor cornea was removed, irrigated with balanced salt solution to wash out the OVD, and resutured on the recipient cornea.

In one patient, the IOP rose to 38 mmHg the day after a Healon® OVD injection, but fell to 12 mmHg after two laser suture lysis procedures. This was the only subject to experience an IOP higher than 20 mmHg after an OVD injection.

Discussion

To our knowledge, this is the largest case series study analyzing the results of OVD injection to treat FAC after trabeculectomy. In our study, a single OVD injection resolved FAC and achieved deep anterior chamber in 84% of the subjects. Case reports by Hoffman et al,⁷ as well as Gutierrez-Ortiz and Moreno-López¹³ on the use of OVD injection to treat postoperative FAC suggested that OVD injection is a simple and effective procedure for managing this complication. Our case series study showed that OVD injection is an effective treatment for FAC after trabeculectomy.

In our study, the long-term, post-trabeculectomy IOP in patients with FAC treated by OVD was significantly lower than the preoperative IOP. We observed an IOP of 14.6 ± 2.7 mmHg 5 years after the trabeculectomy. This result suggests that OVD injection is an effective treatment for FAC. However, previous studies have measured post-trabeculectomy IOPs of 10.0 mmHg, 10.3 mmHg, and 11.1 mmHg; our results were relatively high in comparison.^{14–17}

Our study found success rates of trabeculectomy after OVD injection of 24.4% (criterion A), 16.5% (criterion B), and 6.3% (criterion C). The qualified success rates of trabeculectomy reported previously are 71%¹⁸ (criterion A), 61%,¹⁵ 57%,¹⁷ 60%¹⁹ (criterion B), and 40%,¹⁹ 44%,¹⁷ and 52%¹⁵ (criterion C). Thus, our study, which included only patients who were treated with OVD for FAC after trabeculectomy, found a lower success rate for trabeculectomy along with the higher postoperative IOP.

In a randomized, controlled trial, de Barros et al¹² compared three groups treated for FAC: one with the anterior chamber reformed with OVD; one with the anterior chamber reformed with a balanced salt solution and concurrent drainage of choroidal effusion; and one treated with pharmacologic therapy. The authors concluded that the final IOP tended to be lower with reformation with OVD. The success rate of trabeculectomy in our study, which included only patients with FAC treated with OVD, was lower than that found in previous studies. However, in de Barros et al's¹² study, trabeculectomy complicated by FAC had a higher success rate when the FAC was treated by OVD injection than by

other procedures. Thus, OVD injection may be the best procedure for treating FAC after trabeculectomy.

One subject who had undergone deep lamellar keratoplasty showed Descemet's membrane detachment. A blunt cannula can be inserted in the gap between the stroma and Descemet's membrane, especially in lamellar surgery patients.²⁰ Therefore, extra care is required when injecting an OVD into patients who have undergone lamellar surgery, such as Descemet's stripping endothelial keratoplasty or deep lamellar keratoplasty.

The limitations of this study include its retrospective case series design and a lack of analysis according to OVD type; we did not evaluate the efficacy of the different types of OVD separately. Another limitation is that we did not evaluate the effect of peripheral anterior synechiae formation after OVD injection because of a lack of accurate information about peripheral anterior synechiae in the medical record. The third limitation is that, after the 30-month follow up, eleven subjects were lost to follow up, four subjects underwent additional glaucoma surgery, and one subject had retinal detachment. The success rate or the postoperative IOP may be affected by the relatively large number of participants lost to follow up.

In conclusion, our findings indicate that OVD injection is effective for treating FAC after trabeculectomy.

Disclosure

The authors report no conflicts of interest in this work.

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