Long-term Improvement after the Athens Protocol for Advanced Keratoconus with Significant Ectasia Progression in the Fellow Eye

Luis Claudio Pereira et al

ABSTRACT
To describe the long-term result of the Athens protocol (custom topography-guided advanced surface ablation followed by riboflavin-UVA collagen cross-linking in the same day) for progressive advanced keratoconus in the left eye, while the fellow right eye first presented with relatively mild keratoconus that was treated with a more conservative therapeutic approach of continuous contact lenses use with the goal for ectasia stabilization. Five years after the Athens protocol, a regression of over 10D (diopters) was observed on the sagittal curvature with significant regularization of corneal shape and improvement of distance corrected visual acuity (DCVA) in the left eye. The right eye, which was treated more conservatively, had an over 12D increase on sagittal curvature along with significant distortion of the corneal shape and reduction of DCVA.

Keywords: Keratoconus, Athens protocol, Contact lens.


Source of support: Nil
Conflict of interest: None declared

INTRODUCTION
Keratoconus is a progressive, bilateral ectatic disorder in which the cornea becomes irregular and cone-shaped. The ectatic condition is related to a progressive thinning and bulging due to chronic biomechanical failure of the corneal stroma. Corneal collagen cross-linking was developed in 1998 by Prof. Seiler and coworkers for augmenting corneal rigidity through the application of riboflavin (vitamin-B2) eye drops followed by ultraviolet A (UV-A) radiation. The application of customized topography-guided surface ablation has been reported in patients with stable or subclinical keratoconus. The Athens Protocol consists of the combination of same-day advanced custom surface ablation and riboflavin-UVA collagen cross-linking. The main goal of the procedure is to regularize the cornea, not to provide primary refractive correction, so that this approach is considered as therapeutic.

Special contact lens fitting, including rigid gas permeable (RGP) contact lenses, represents the main conservative therapeutic approach for visual rehabilitation of ectatic corneal diseases and is responsible for over 50% of cases. Contact lenses may provide the best possible visual acuity because it regulates the corneal surface along with the tear film. However, there is no evidence that contact lens stabilize or prevent ectasia progression, while there is some anecdotal evidence that it may be associated with a higher risk of ectasia progression due to continuous microtrauma from chronic RGP contact lens use.

CASE REPORT
A Caucasian 26-year-old presented with a history of decreased vision in his left eye since 2007. On examination, uncorrected distance visual acuity (UDVA) was 20/60 of the right eye and <20/400 of the left eye. Distance best corrected vision (DCVA) with spectacles was 20/20 in the right eye and 20/80 in the left eye. Fundus examination was within normal limits in both eyes. Goldmann applanation intraocular pressure (IOP) was 10 mm Hg in oculus dextrus (OD) and 7 mm Hg in oculus sinister (OS). Goldmann-correlated IOP (IOPg), corneal-compensated intraocular pressure (IOPcc), corneal hysteresis (CH), and corneal resistance factor (CRF) from the ocular response analyzer (ORA, Reichert, Buffalo, NY) were, respectively, 14.1, 11.4, 7.8 and 8.8 mm Hg in the right eye and 11.7, 7.6, 6.1 and 8.1 mm Hg of the left eye. Keratoconus was confirmed by placido disk-based corneal tomography (iTrace) and rotating Scheimpflug corneal tomography (Pentacam, Oculus, Wetzlar, Germany). Wavefront optimized manifest refraction with data from iTrace (Tracey Technologies, Houston, TX) was −1.25 − 2.75 × 56° = 20/20 of the right eye and −9.75 −3.75 × 1° = 20/60.
Long-term Improvement after the Athens Protocol for Advanced Keratoconus with Significant Ectasia Progression

of the left eye. The patient was fitted with RGP contact lens, providing a best corrected visual acuity of 20/15 of the right eye and 20/30 of the left eye.

Six months later, the patient returned complaining of decreased vision of the left, which failed to improve with contact lens wear. Wavefront (WF) optimized manifest refraction was $-1.50 - 2.75 \times 49^\circ = 20/20$ of the right eye, and $-11.75 - 6.00 \times 162^\circ = 20/80$ of the left. Significant progression of ectasia was demonstrated of the left eye with an increase of the maximal keratometric value ($K_{max}$) of 2.77 D, while the right eye remained stable.

Considering the progressive nature of the ectasia, the intolerance to further contact lens wear, the amount of irregularity, and the thinnest corneal thickness value of 456 µm of the left eye, the Athens Protocol was performed accordingly to the description by Kanellopoulos as an alternative for keratoplasty. Advanced surface ablation was performed with the Allegretto (Alcon-Wavelight, Earlagen, Germany). Epithelial removal was accomplished using the (phototherapeutic keratectomy) (PTK) mode set for 50 microns with a 6.5 mm optical zone and an 8.8 mm blend zone followed by subsequent same day custom topography-guided ablation (TCAT), based on oculyzer data, which removed 47 microns of tissue in the deepest ablation zone. Mitomycin C (0.005%) was applied for 45 seconds followed by copious irrigation with 50 cc of cold (4°C) BSS. Dextran-free 0.1% riboflavin solution was instilled every 2 minutes for 30 minutes. The yellow flare was verified before UVA irradiation at 3 mW/cm$^2$ for 30 minutes was competed. A bandage soft contact lens (SofLens 66, Bausch and Lomb Inc, Rochester, New York) was placed until the cornea epithelialized on postoperative day 6 with no adverse event besides moderate pain in the first 3 days after the procedure. One month after the procedure, UCVA was 20/100. The patient had a second opinion with another corneal subspecialist, who advised against corneal collagen cross-linking OD and who also indicated that continuous RGP contact lens wear in both eyes was necessary.

The patient returns in 2012 (4 years after the procedure) complaining of bad vision in both eyes and contact lens intolerance. UCVA was 20/100 of the right eye and 20/60 of the left eye. The wavefront (WF) optimized manifest refraction was $-13.50 - 7.00 \times 8^\circ = 20/70$ of the right eye and $-11.50 - 1.00 \times 169^\circ = 20/25$ of the left eye. Comparative subtraction front sagittal curvature maps from 2008 and 2012 are presented in Figure 1. Figures 2 and 3 include the elevation differential maps for OD and OS, respectively. 12.6D of corneal steepening is observed OD (Figs 1A and B) and 10.6D of flattening is observed OS (Figs 1C and D).

Figure 2 provides the front elevation maps for OD in 2008 and 2012. The same reference was chosen of 7.8 mm as best fit sphere (BFS) for the first evaluation. Over 80 microns of increase in elevation is observed as well as a decrease in central corneal thickness from 511 to 455 µm (at pupil center) and an increase in ACD (anterior chamber depth) (ACD) from 3.54 to 3.87 mm.

Figure 3 provides the front elevation maps for OS in 2008 and 2012. The same reference was chosen of 7.04 mm as best fit sphere (BFS) for the first evaluation. 36 microns of decrease in elevation was observed and a decrease in central keratometric astigmatism from 5.9D to 0.3D. A mild decrease in central corneal thickness from 478 to 443 µm (at pupil center) and an decrease in anterior chamber depth (ACD) from 3.87 to 3.77 mm.

The patient was advised about the need for therapeutic surgery for OD and surgical options, including the possibility of keratoplasty, and phakic intraocular lens (IOL) as a refractive option for OS.

![Fig. 1: Sagittal curvature differential maps](image-url)
DISCUSSION

The patient presented with advanced progressive keratoconus of the left eye, which was treated by the Athens protocol, and the right eye had a mild and relatively stable keratoconus presentation with good distance corrected visual acuity with spectacles. The patient had a second opinion, who advised against cross-linking of the right eye and to continuously wear RGP contact lens OD. This case illustrates that the use of contact lens does not stop the progression of keratoconus. Moreover, continuous microtrauma related to continuous contact lens wear may be an important factor to accelerate ectasia progression in some patients. After 4 years, a regression of more than 10D of astigmatism of the left eye was associated with regularization of the corneal surface and improvement in the BCVA after the Athens protocol. This result is consistent with the literature.

CONCLUSION

The Athens protocol is a therapeutic alternative for patients with progressive keratoconus to prevent or even delay keratoplasty. This is not a refractive procedure. Refractive correction with a phakic IOL becomes a possibility, if best spectacle corrected vision is good. However, the progressive nature of ectatic corneal diseases should be considered and patient education is fundamental. Surgery for keratoconus should be indicated when visual rehabilitation is not possible with glasses or contact lenses and when there is documented or a significant risk of ectasia progression. While contact lenses provide the best possible treatment for the irregular astigmatism, there is no evidence to support the benefit of contact lens for decreasing the risk of ectasia progression. In fact, contact lens fitting should be considered with caution, if spectacles provide adequate visual acuity.
REFERENCES