LASIK and PRK in hyperopic astigmatic eyes: is early retreatment advisable?

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Purpose: To analyze the refractive and keratometric stability in hyperopic astigmatic laser in situ keratomileusis (LASIK) or photorefractive keratectomy (PRK) during the first 6 months after surgery.

Patients and methods: This retrospective cross-sectional study included 97 hyperopic eyes; 55 were treated with LASIK and 42 with PRK. Excimer ablation for all eyes was performed using the ALLEGRETTO excimer laser platform using a mitomycin C for PRK and a mechanical microkeratome for LASIK. Keratometric and refractive data were analyzed during three consecutive follow-up intervals (6 weeks, 3 months, and 6 months). The corneal topography was obtained using Scheimpflug topography, and subjective refractions were acquired by expert optometrists according to a standardized protocol.

Results: After 3 months, mean keratometry and spherical equivalent were stable after LASIK, whereas PRK-treated eyes presented statistically significant \((P<0.001)\) regression of hyperopia. In eleven cases, hyperopic regression of \(>1\) D occurred. The optical zone diameter did not correlate with the development of regression.

Conclusion: After corneal laser refractive surgery, keratometric changes are followed by refractive changes and they occur up to 6 months after LASIK and for at least 6 months after PRK, and therefore, caution should be applied when retreatment is planned during the 1st year after surgery because hyperopic refractive regression can lead to suboptimal visual outcome. Keratometric and refractive stability is earlier achieved after LASIK, and therefore, retreatment may be independent of late regression.

Keywords: hyperopia, astigmatism, regression, keratometry

Introduction
The safety and efficacy of laser in situ keratomileusis (LASIK) and photorefractive keratectomy (PRK) in correcting hyperopic astigmatism have been reported.1 However, the efficacy of any corneal refractive surgery relies on refractive and keratometric stability.

Refractive hyperopic regression is common after hyperopic treatment and, therefore, limits the hyperopic correction range to 3–4 D of hyperopia compared with the more than double dioptic range for myopic correction.1–4 In our experience, refractive hyperopic regression of \(>0.25\) D after 3 months caused by hyperopic refractive shift associated with changes in the crystalline lens (due to age) represents a considerable refractive problem, demanding a retreatment in a significant proportion of cases. Since refractive patients often request the earliest possible retreatment, this study analyses the refractive and keratometric stability during the first 6 months after LASIK or PRK in hyperopic astigmatic eyes.
Materials and methods

Patients and examinations

This retrospective study included 97 eyes from 97 hyperopic patients consecutively treated between August 1, 2013, and January 5, 2014, and was based on the Hamburg Refractive Database (data retrieved from Care Vision Refractive Centers in Germany). For each patient, one randomly selected eye was analyzed. The inclusion criteria were LASIK and PRK for hyperopia with >0.5 D, no previous refractive surgery, and follow-up (FU) examinations at 6 weeks, 3 months, and 6 months after surgery with full refractive data available.

The written informed consent for retrospective data analysis was obtained from refractive surgery candidates during their recruiting process. The study and consent procedure were approved by the University of Hamburg Ethics Committee (no 2882) and adhered to the tenets of the Declaration of Helsinki.

The hyperopic correction was based on the manifest refraction and presented here as spherical equivalent (SE). Whether a PRK or a LASIK was performed depends on the preoperative pachymetry, topography, and subsequent assumed risk of ectasia as well as on individual risk profile for flap-associated complications. PRK was preferred in patients with hobbies or jobs with a higher risk of eye trauma or in eyes with mild ectasia risk. Moreover, previous studies reported that epithelial remodeling following PRK may be substantially more in comparison to LASIK. These unavoidable inherent differences in pachymetry and refractions between LASIK and PRK eyes may be statistically disadvantageous but clinically speaking it reflects the actual population of LASIK and PRK patients and is useful for drawing clinical conclusions for these populations.

Manifest spherical and cylindrical refraction, cycloplegic refraction, as well as visual acuity with and without correction were assessed pre- and postoperatively (except cycloplegic refraction). FU included examinations at 1 day (LASIK) to 5 days (PRK) after surgery, and during three consecutive FU examinations (6 weeks, 3 months, and 6 months after surgery), and were recorded electronically.

Refractive stability was defined as a maximum of ±0.25 D change in SE or mean keratometry (Km) between 3-month and 6-month examination. The corneal topography was obtained using Scheimpflug topography (Pentacam HR; Oculus, Wetzlar, Germany). All refractions were acquired by expert optometrists using the same refractometers, visual acuity tables, and documentation protocol. Each patient was examined pre- and postoperatively by the same optometrist. Examinations were carried out according to a standardized protocol. To minimize the potential bias, latent hyperopia eyes with preoperative cycloplegic refraction of >1 D difference to manifest refraction were excluded from this study.

Laser treatment

Excimer ablation for all eyes was performed using the ALLEGRO GRETTO excimer laser platform (Eye-Q 400 Hz; WaveLight GmbH, Erlangen, Germany) under constant eye tracking using a wavefront-optimized ablation protocol. The surgical procedures using mitomycin C for PRK and a mechanical microkeratome (SBK 90 µm; Moria SA, Antony, France) for LASIK have been described previously.6,7

The ablation was centered on the visual axis (first Purkinje image), which is required in hyperopic eyes with relatively large-angle kappa. The patient was asked to focus on a target light offered by the excimer platform. The first Purkinje image of this light and its relation to the pupil center were documented, and the eye tracker used this reference point as ablation center (“off-set”). All eyes were treated with an optical zone (OZ) of either 6 mm (n=16), 6.5 mm (n=65), or 7 mm (n=16) according to scotopic pupil size. The between-group difference was not statistically significant (tested with chi-square, P=0.180).

All three refractive surgeons were experienced consultants and followed a standard protocol of indications, and the pre-, intra-, and postoperative treatment plan was written and implemented by the last author in all three refractive centers involved in this study.

Pre- and postoperative Km readings were measured in the central 3 mm as simulated K (simK) in millimeter radius and converted to D using the air to stroma refraction index of 1:1.367. The central 3 mm includes the visual axis and reflects the main refractive change in the whole cornea. The difference between the true anterior corneal Km preoperatively and postoperatively follows precisely the change in manifest refraction and hence is an appropriate parameter for the corneal and refractive development over time. The targeted Km was calculated using the preoperative Km and adding the SE that was used by the laser platform with compensation for the vertex distance of the manifest refraction (12 mm) to corneal plane (0 mm). For example, correcting a manifest SE of +4.0 D should increase the central Km by 4/(1–0.012×4) =4.2 D. In an eye with preoperative Km of 39 D and SE of +4 D, the treatment should result in Km of 43.2 D and SE of 0 D. The expected change is in Km of +4.2 D and in SE of +4.0 D. We calculated the postoperative actual Km and SE at each FU and analyzed changes between the FU examinations.
Statistical analysis
To analyze the differences in biometrical data between LASIK and PRK, either a parametrical independent t-test or a non-parametric Mann–Whitney test was used, depending on the normality of the data. To analyze the predictability of the treatment, linear regression was applied. Changes in SE and Km during FU were evaluated using repeated measure analysis of variance. The differences between achieved SE change and achieved Km change were tested with a paired t-test.

Results
Table 1 gives the preoperative refractive data for both LASIK and PRK groups and demonstrates that there were no statistically significant differences except for preoperative pachymetry. In 55 eyes, LASIK was performed and another 42 patients were treated with PRK.

The postoperative data originate from the 6-month FU examination. There were no statistically significant differences in any refractive parameter after surgery (Table 1). Figure 1 shows the development of Km and SE during FU. In the LASIK group (Figure 1A; Table 2), postoperative changes in Km and SE were significant (P<0.001) until 3 months. In other words, there was statistically significant change in refractive and keratometric stability (P<0.001) (Table 2).

In the 42 eyes that had PRK, postoperative changes in Km and refraction were significant (P<0.001) up to 6 months after surgery (Figure 1B, Table 2). Km and SE progressed in terms of hyperopic regression (= undercorrection) (Figure 1C). In eleven cases, refractive regression was >1 D (three LASIK and eight PRK eyes), and this difference was statistically significant between LASIK and PRK (chi-square, P=0.036).

All laser treatments applied an OZ of either 6 mm, 6.5 mm, or 7 mm based on scotopic pupil size; there was no statistically significant difference in OZ between LASIK and PRK. The OZ diameter did not correlate with keratometric or refractive regression.

Discussion
This study analyses the refractive and keratometric stability in hyperopic astigmatic LASIK or PRK during the first 6 months after surgery.

Our results show that stability of refraction and Km was earlier presented in eyes after LASIK; PRK-treated eyes showed statistically significant higher hyperopic regression displayed by changes in postoperative SE and Km up to 6 months after surgery. In 11 (11%) cases, regression of >1 D occurred. Among them were three LASIK-(5.45% of all LASIK cases) and eight PRK-treated eyes (19.05% of all PRK cases). As refractive and topographic stability in LASIK is present at the 6-month FU examination, retreatment could be performed earlier than in PRK eyes. After PRK, we do not recommend any retreatment within the first 6 months after surgery due to hyperopic regression and recommend reevaluation of refractive and topographic stability after 1 year. In general, Km and refraction developed in a parallel manner that implies that regression is Km driven and not due to latent hyperopia; moreover, it was independent of OZ diameter.

Refractive stability after hyperopic corneal refractive surgery is still controversial. After LASIK, stability has been reported after 1 month, 9 at 3 months, 9 at 6 months, 10 and at 12 months.11 Nevertheless, caution should be applied as refractive stability is often not clearly defined or hard to compare between studies. In our opinion, clinically relevant refractive stability should be defined as a maximum of ±0.25 D change in SE or Km after at least 3 months.

We are aware of the fact that the hyperopic shaping of the cornea is three-dimensional and includes multiple changes; however, we chose the simK parameter because it is the only single parameter that presents the total corneal refractive power as close as possible and was measured in all FU examinations and its change corresponds to the change in refraction.12

Both PRK and LASIK are effective and safe in the correction of hyperopia. However, PRK tends to lead to initial temporary myopic overshoot, which can be followed by a hyperopic regression over months to years.1 The results of our study concur with that finding (Figure 1; Table 2). de Ortueta and Arba Mosquera showed that topography can be used as an objective method to analyze regression after LASIK for hyperopia.2 They reported that virtually no topographic regression between 3-month and 36-month FU could be observed; between 12 months and 36 months a regression of 0.001 D/month was calculated and this was not statistically significant.2 Others have reported that regression can occur for years; Jaycock et al found that regression could not be explained by age-related changes in refraction that still occurred after 5 years.3 By contrast, Kezirian et al, observing a 4.5-year period after surgery, demonstrated that the selection of the 6-month examination as the time point of stability was justified.4 This conclusion is supported by our data showing stability in LASIK-treated eyes after 3 months.

Other factors that have been related to the impact of LASIK on spherical aberration include surface smoothing as a result of corneal wound healing and epithelial
| Variable                              | LASIK n=55                      | PRK n=42                      | P-value* |
|---------------------------------------|---------------------------------|-------------------------------|----------|
|                                       | Min | Max | Median (Q25/Q75) | Mean (SD) | Min | Max | Median (Q25/Q75) | Mean (SD) |          |
| Preoperative                          |     |     |                 |           |     |     |                 |           |          |
| Male                                  | –    | –    | –                | 0.42 (±0.50) | –    | –    | –                | 0.60 (±0.50) | 0.084*** |
| Age (years)                           | 18   | 60   | 41 (29/49)       | 40 (±12)   | 22   | 62   | 34 (28/47)       | 37 (±12)   | 0.281*** |
| Refractive sphere (D)                 | 0.75 | 3.5  | 1.75 (1.25/2.25) | 1.89 (±0.69) | 0.75 | 3.5  | 1.75 (1.25/2.50) | 1.84 (±0.73) | 0.747**  |
| Refractive cylinder (D)               | –3.75 | 0    | –1.75 (–2.50/–0.75) | –1.69 (±1.07) | –4   | –0.25 | –1.38 (–2.25/–1.00) | –1.61 (±1.00) | 0.701*** |
| Refractive cylinder axis (°)          | 4    | 180  | –                | 99         | 3    | 175  | –                | 86         | 0.844*** |
| Spherical equivalent (SE in D)        | 0.13 | 1.88 | 1.00 (0.50/1.63) | 1.04 (±0.58) | 0.13 | 1.88 | 1.06 (0.63/1.63) | 1.04 (±0.58) | 0.43   |
| UDVA (logMAR)                         | 0.05 | 1    | 0.34 (0.22/0.52) | 0.38 (±0.21) | –0.04 | 1    | 0.30 (0.18/0.52) | 0.37 (±0.28) | 0.300*** |
| CDVA (logMAR)                         | –0.1 | 0.3  | 0.00 (–0.02/0.05) | 0.01 (±0.07) | –0.11 | 0.55 | 0.00 (–0.04/0.03) | 0.02 (±0.12) | 0.494*** |
| Mean keratometry (Km in D)            | 40   | 45.8 | 43.10 (42.20/44.00) | 43.03 (±1.33) | 38.8 | 46.45 | 43.45 (42.20/44.75) | 43.33 (±1.89) | 0.353*   |
| Km cylinder (D)                       | –4.5 | –0.1 | –1.60 (–2.60/–0.80) | –1.75 (±1.03) | –3.6  | –0.2 | –1.60 (–2.40/–1.10) | –1.70 (±0.90) | 0.801*   |
| Km axis (°)                           | 1    | 180  | –                | 94         | 2    | 180  | –                | 85         | –       |
| Pachymetry (µm)                       | 484  | 632  | 550 (522/580)    | 552 (±35)  | 384  | 639  | 530 (496/560)    | 531 (±52)  | 0.023*   |
| Postoperative (6-month follow-up)     |     |     |                 |           |     |     |                 |           |          |
| Refractive sphere (D)                 | –0.75 | 2.25 | 0.25 (0.00/0.50) | 0.25 (±0.55) | –0.75 | 0.75 | 0.00 (0.00/0.50) | 0.11 (±0.34) | 0.178*** |
| Refractive cylinder (D)               | –1.5  | 0    | –0.50 (–0.75/0.00) | –0.49 (±0.43) | –1   | 0    | –0.50 (–0.50/0.00) | –0.37 (±0.29) | 0.135*   |
| Refractive cylinder axis (°)          | 1    | 180  | –                | 98         | 5    | 180  | –                | 99         | 0.196*** |
| SE (D)                                | –0.88 | 1.5  | 0.00 (–0.25/0.25) | 0.01 (±0.50) | –1   | 0.5  | 0.00 (–0.38/0.25) | –0.08 (±0.38) | 0.045*** |
| UDVA (logMAR)                         | –0.1  | 0.47 | 0.05 (0.00/0.12) | 0.07 (±0.13) | –0.1 | 0.42 | 0.00 (–0.06/0.03) | 0.03 (±0.11) | 0.352*   |
| CDVA (logMAR)                         | –0.12 | 0.27 | 0.00 (–0.04/0.03) | 0.00 (±0.07) | –0.1 | 0.42 | 0.00 (–0.06/0.03) | 0.00 (±0.10) | 0.352*   |
| Km (D)                                | 40.25 | 46.1 | 43.90 (43.30/44.80) | 43.84 (±1.33) | 39.4 | 47.25 | 44.13 (43.10/45.60) | 44.22 (±1.82) | 0.242*   |
| Km cylinder (D)                       | –2.8  | –0.3 | –1.10 (–1.50/–0.70) | –1.11 (±0.58) | –3.2 | 0    | –0.80 (–1.20/–0.50) | –1.00 (±0.71) | 0.168*** |
| Km axis (°)                           | 1    | 180  | –                | 90         | 2    | 180  | –                | 86         | –       |
| EI                                    | 0.36 | 1.2  | 0.96 (0.76/1.05) | 0.90 (±0.20) | 0.62 | 1.36 | 0.98 (0.93/1.04) | 0.98 (±0.14) | 0.157*** |
| SI                                    | 0.67 | 1.33 | 1.04 (0.98/1.11) | 1.03 (±0.13) | 0.9  | 1.36 | 1.00 (0.98/1.06) | 1.04 (±0.10) | 0.551*** |

Notes: Depending on the assumption of normality either Student's t-test or Mann–Whitney test was applied. *Student's t-test. **The counted data were compared with chi-square test. ***Mann–Whitney test. "Mean direction defined as (ArcTan(S/C))/2, where S = SUM sin axis and C = SUM cos axis and axis is in range from 0° to 360°."

Abbreviations: LASIK, laser in situ keratomileusis; PRK, photorefractive keratectomy; min, minimum; max, maximum; SD, standard deviation; UDVA, uncorrected distance visual acuity; CDVA, corrected distance visual acuity; EI, efficacy index; SI, safety index; Km, mean keratometry; logMAR, logarithm of the minimum angle of resolution.
remodulation,\textsuperscript{13} biomechanical changes,\textsuperscript{14} and variation in the laser ablation depth per pulse across the treatment area in relation to corneal curvature.\textsuperscript{15} This first factor is minimized in our study by using a wavefront-optimized ablation protocol.

In eyes after PRK, regression could be explained by a prolonged epithelial healing process. As previously described, after hyperopic PRK, epithelial remodeling aims to compensate for the ablated corneal tissue.\textsuperscript{1} On the other hand, central epithelial hyperplasia would manifest as refraction overshoots over the first week to month, and after complete healing causes regression after 3–4 months.

Regression has also been strongly associated with the magnitude of hyperopia and some argue that less regression goes along with the use of larger OZs;\textsuperscript{16} it is considerably greater in high hyperopia and can continue up to 12 months.\textsuperscript{17} In our study, there was no statistically significant correlation of OZ diameter or magnitude of preoperative hyperopia with regression during FU. Moreover, there was no statistically significant difference in OZ between LASIK and PRK groups. Since simK is taken from the central 3 mm, the OZ should not affect the expected central Km.

In contrast to previous literature, advantages of our study are that we analyzed only one eye per patient who were of relatively young age, we compared two homogeneous groups of LASIK and PRK, and the ablation profiles took into account intraoperative off-set. On the other hand, we acknowledge limitations to our study. Especially the role of cycloplegic refraction in treatment planning is important when analyzing hyperopic treatment results. Although we focused on patients with minimal latent hyperopia, we did not analyze the influence of postoperative cycloplegic refraction. However, Km and refraction developed in a parallel manner that implies that regression is Km driven and not due to latent hyperopia.

**Conclusion**

In conclusion, after corneal laser refractive surgery, keratometric changes are followed by refractive changes and they occur up to 6 months after LASIK and for at least 6 months after PRK, and therefore, caution should be applied when retreatment is planned during the 1st year after surgery as
hyperopic refractive regression can lead to suboptimal visual outcome. Keratometric and refractive stability is earlier achieved after LASIK, and therefore, retreatment may be independent of late regression.

Disclosure

The authors report no conflicts of interest in this work.

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