Single-step transepithelial photorefractive keratectomy in high myopia: qualitative and quantitative visual functions

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Abstract

• AIM: To investigate quantitative and qualitative optical outcomes of single-step transepithelial photorefractive keratectomy (TransPRK) in high myopia.

• METHODS: In a prospective interventional case-series, 30 eyes with high myopia (-6.00 to -8.75 D) with (up to -3.00 D) or without astigmatism were enrolled from Bina Eye Hospital, Tehran, Iran. One-step TransPRK was performed with aberration-free aspherical optimized profile and SCHWIND AMARIS 500 laser. One-year follow-up results for refraction, visual acuities, vector analysis, ocular wave-front (OWF) and corneal wave-front (CWF) higher order aberrations (HOA), contrast sensitivity (CS), and post-operative haze were assessed.

• RESULTS: After the surgery, both photopic and mesopic CSs significantly improved (both \( P < 0.001 \)). We detected significant induction of OWF coma and trefoil (\( P < 0.001 \) for both) HOAs; CWF coma (\( P = 0.002 \)), spherical (\( P < 0.001 \)), and tetrafoil (\( P = 0.003 \)) HOAs in 6 mm analysis diameter; and CWF trefoil (\( P = 0.04 \)) HOA in 4 mm analysis diameter. The range of mean induction observed for various HOAs was 0.005-0.11 \( \mu \)m. The 86.7% of eyes reached an uncorrected distance visual acuity of 20/20 or better; 96.7% of eyes were within ±0.5 D of targeted spherical refraction. In vector analysis, mean correction index value was 1.03 and mean index of success was 0.22. By 12mo after the operation, no eye lost any number of corrected distance visual acuity lines. We detected no corneal haze greater than 1+ throughout the follow-up.

• CONCLUSION: Our findings show promising effects of single-step TransPRK on quality of vision in high myopic eyes. It also improves refraction and visual acuity.

• KEYWORDS: single-step transepithelial photorefractive keratectomy; transepithelial photorefractive keratectomy; high myopia; contrast sensitivity; higher order aberrations; vector analysis

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INTRODUCTION

Transepithelial photorefractive keratectomy (TransPRK) was introduced in late 1990s to avoid mechanical debriment complications associated with PRK and flap related complications and ectasia after laser in situ keratomileusis (LASIK)[1-4]. It has been used to correct primary refractive errors as well as secondary refractive errors subsequent to other corneal surgeries[5].

There are two main platforms of TransPRK. The older two-step platform removes epithelium by phototherapeutic keratectomy (PTK) followed by conventional PRK for stromal ablation. Clinical results for efficacy of this platform in correction of primary refractive errors are inconclusive[6-7]. The next is one-step platform which exploits an aspheric nomogram with Amaris 500 excimer laser to simultaneously ablate surface and stroma[8-9]. It utilizes a population-based profile of corneal thickness to provide even energy on entire surface of the cornea. It also could prevent stromal dehydration[9]. Studies have reported promising efficacy of one-step TransPRK in correction of different types of refractive errors[8-13].

Correction of high myopia remains a big challenge for refractive surgeons. Results in LASIK and PRK confirmed better correction, more stability and predictability in low myopia compared to high myopia[14-15]. Higher corrections attempted in high myopic eyes induce larger intra-ablation...
transitional zones and spherical aberrations and result in further transformation of corneal asphericity from prolate to oblate. This diminishes central effective optical zone (EOZ) of the cornea and compromise contrast sensitivity (CS)\textsuperscript{[16-17]}. In case of TransPRK, we hypothesized that delivery of even energy to different parts of cornea by one-step platform is expected to attenuate harmful effects of higher corrections on quality of vision in high myopia. To the best of our knowledge, no study has addressed the topic. We aimed to investigate the effect of single-step TransPRK on qualitative as well as quantitative visual functions in high myopic eyes.

**SUBJECTS AND METHODS**

Patients with high myopia (subjective spherical refraction above -6 D) and astigmatism were included in this prospective interventional case-series at Bina Eye Hospital, Tehran, Iran from July 2013 to December 2014. The study was ethically approved by the Ethics Committees of Local Institutional Review Board and was in accordance with Declaration of Helsinki. A written informed consent was taken from all of the patients. The participants did not receive any stipend. Concurrent ocular diseases, systemic diseases (with ocular involvement), previous corneal or ocular surgery, keratoconus, high aberration values (corneal abberations above 0.35 μm for a 6 mm pupil diameter analyzed with Optikon Scout corneal analyzer), and night vision disturbances (e.g. patients with retinitis pigmentosa and chorioretinal atrophic changes) were considered as our exclusion criteria. Four weeks prior to surgery, patients discontinued wearing hard or soft contact lenses.

We assessed uncorrected distance visual acuity (UDVA); corrected distance visual acuity (CDVA); refraction; keratometry and topography with Scout (Optikon 2000 SPA, Rome, Italy) and Orbscan (Bausch and Lomb, Rochester, NY, USA); CS in photopic and mesopic conditions (M&S Smart System 20/20, M&S Technologies Inc., Niles, IL, USA) using a myopia-correcting spectacles; anterior corneal wave-front (CFW) aberrometry in 4 mm and 6 mm analysis diameters (Keratron Scout Corneal Analyser; Optikon 2000 SPA, Rome, Italy); and ocular wave-front (OWF) aberrometry (ORK Wavefront Analyzer; SCHWIND eye-tech-solutions GmbH, Kleinostheim, Germany). In our logarithmic scale for CS, lower values indicated better CSs. Follow-up data were collected for 1, 3, 6, and 12mo after the operation.

For laser ablation, first we used local anesthetic drops, and irrigated the eyes with balanced salt solution (BSS). We used no alcohol. The ablations were performed by Amaris 500 Hz excimer laser (SCHWIND eye-tech-solutions, GmbH, Kleinostheim, Germany) in aberration-free mode. In our refined approach, we considered comprehensive demographic and optical characteristic of each patient to set the ablation profile. Age, keratometry, central corneal thickness, radius of corneal curvature were amongst the main parameters considered to adjust the target refraction and optical zone. After primary assessment with Orbscan, Scout Analyser was used to get detailed topographic data of the patients’ eye. Each single eye underwent 8-10 consecutive scans. The surgeon chose the one with the highest precision to use for adjusting the ablation profile. Ablation profile was centered on the corneal vertex for the eyes with a pupillary offset (distance between pupil center and corneal vertex) greater than 0.35 mm. For all other eyes, an ablation centered on pupil center was used. Optical zone (OZ) ranged from 6.1 to 7.0 mm. Administration of the laser occurred in a single continuous session to ablate both the epithelium and stroma in a single step using an aberration free and aspheric profile. According to a population-based epithelium-thickness profile, the ablation plan utilized 55 µm centrally, and 65 µm peripherally, with further adjustment of the differential ablation rate in the epithelium compared to the stroma. It provides an even laser energy on the entire corneal surface. Eye movements throughout the ablation were compensated by static and dynamic cyclotorsion corrections (SCC and DCC). All treatments were performed by the same surgeon (Adib-Moghaddam S). Just after the ablation, a soaked and then squeezed sponge with 0.02% mitomycin-C (MMC) was placed over the stromal bed for 25s\textsuperscript{[18]}.

Then, we irrigated the eye by copious amounts of chilled BSS. A soft bandage contact lens with high diffusion constant of oxygen permeability and base curve of 8.6 (Bausch & Lomb, New York, USA) was placed over cornea for two or three days. We used our refined protocol for postoperative medication (Iran regimen)\textsuperscript{[12]}: chloramphenicol eye drop 0.5% (Sina Daru, Tehran, Iran) were administered every 4h up to 2-3d. Oral ibuprofen capsule 400 mg (DaanaPharmo Co, Tabriz, Iran) twice a day and alprazolam tablet 1 mg (Pursina Pharmaceutical Co, Tehran, Iran) daily were prescribed as well. Neither topical steroid nor topical non-steroidal anti-inflammatory drug was used during epithelial healing phase. Contact lens was removed 2-3d after the surgery when the corneal re-epithelialization was complete. If no corneal epithelial defect was detected after three days, patients were prescribed loteprednol eye drops 0.5% (Lotemax, Bausch & Lomb, New York, USA) every 6h for 2wk, then tapering every 2wk to once daily and continued up to 6mo. Chloramphenicol 0.5% every 8h for a week, and preservative-free artificial tear (Artelac Advanced, Bausch & Lomb, New York, USA) for six months, were also prescribed after the complete re-epithelialization. All patients were instructed to protect eyes from ultra-violet light by wearing sunglasses. Corneal haze was assessed by slit lamp and graded as previously described\textsuperscript{[19]}.

**Vector Analysis** We used vector analysis according to the Alpins method\textsuperscript{[20]} to assess the accuracy of astigmatism correction. Astigmatism was vertexed to corneal plane considering a back
vertex distance of 12 mm. To address the mirror symmetric effects, cylinder axes of left eyes were transformed to a new axis by the formula: 180-original axis. For each individual eye, X and Y coordinates were calculated from astigmatism power (D) and axis (degree). They were used to calculate components and indices of vector analysis (Figure 1)\[^{20}\]. The optimal values for magnitude of error (ME), index of success (IOS), angle of error (AE) and the magnitude of difference vector (DV) are zero and for the correction index (CI) and flattening index (FI) are one. These values indicate a successful correction. CI values above one and positive values for ME indicate overcorrection and CI values below one and negative values for ME indicate under correction. We considered emmetropia as the target for our correction.

Statistical Analysis

The continuous variables were shown as mean±standard deviation (SD). The categorical variables were presented as frequency and percentage. We used parametric t-tests (independent or paired) or non-parametric Mann-Whitney or Wilcoxon signed-rank (WSR) tests to compare the mean values. Chi-squared or Fischer-exact tests were used to compare categorical variables. No missing data was encountered in main study parameters. Cases with loss to follow-up were excluded from the analysis. All the statistical analyses were conducted using IBM SPSS statistics for Windows, version 23.0 (Armonk, NY: IBM Corp.).

RESULTS

From 40 eyes screened for eligibility, 33 eyes were enrolled in the study (Figure 2). Finally, three eyes were lost to follow-up and a total number of 30 high myopic eyes from 16 patients were included in the analysis. In two patients, only one eye met the eligibility criteria. All of the patients completed one-year follow-up. The mean±SD age of patients was 30.8±6.4y with a range of 23-48y and 20 (66.7%) eyes belonged to female and 10 (33.3%) eyes belonged to male patients (Table 1). All of the eyes were primary high-myopic (-6.00 to -8.75 D), with or without astigmatism (-0.25 to -3.0 D).

Contrast Sensitivity and Higher Order Aberrations

Both photopic and mesopic CSs improved after the operation. Mean photopic CS improved from preoperative value of 1.42±1.48 to 0.96±0.49 three months after the operation (\(P=0.003\)), and further improved to 0.94±0.37 and 0.83±0.36 by 6 and 12mo after the operation, respectively. The mean±SD age of patients was 30.8±6.4y with a range of 23-48y and 20 (66.7%) eyes belonged to female and 10 (33.3%) eyes belonged to male patients (Table 1). All of the eyes were primary high-myopic (-6.00 to -8.75 D), with or without astigmatism (-0.25 to -3.0 D).

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coma ($P=0.002$), spherical ($P=0.001$), and tetrafoil ($P=0.003$) HOA in 6 mm analysis diameter; and CWF trefoil ($P=0.04$) HOA in 4 mm analysis diameter (Table 2). For other HOA, the numerical values increased; however, their changes did not reach statistical significance ($P=0.31$ for coma, $P=0.09$ for spherical, $P=0.25$ for secondary astigmatism, and $P=0.82$ for tetrafoil HOA at 4 mm diameter; and $P=0.3$ for trefoil and $P=0.08$ for secondary astigmatism HOA at 6 mm diameter). The range of mean induction observed for various HOAs was $0.005$ to $0.11 \mu m$.

### Visual Acuity

**Efficacy** Preoperative mean UDVA logMAR of $0.87\pm0.49$ significantly improved to $-0.05\pm0.11$, three months after the operation ($P<0.001$) and then further improved to $-0.09\pm0.12$ and $-0.08\pm0.10$ by month 6 and 12, respectively. Number of eyes with UDVA equal or better than 20/20 was $28$ ($93.3\%$), $27$ ($90.0\%$) and $26$ ($86.7\%$) by 3, 6 and 12 months, respectively (Figure 3). All of the eyes achieved UDVA equal or better than 20/25 by 12 months after the operation. Mean logMAR UDVA achieved 12 months after the operation was better than mean logMAR of preoperative CDVA (mean difference $=0.07\pm0.08$, $P<0.001$) and $93.3\%$ of the treated eyes achieved postoperative UDVA equal or better than their preoperative CDVA (Figure 3).

**Safety** By 12 months after the operation, no eye lost any number of CDVA lines (Figure 3). At this time point, $13$ ($43.3\%$) eyes gained one line of postoperative CDVA and the others preserved their preoperative CDVA.

**Refraction** After the operation, mean ($\pm SD$) spherical equivalent (SE) of the eyes improved from preoperative value of $-7.12\pm1.00$ to $-0.11\pm0.30$ D ($P<0.001$) three months after the operation and then slightly regressed to $-0.15\pm0.43$ and $-0.19\pm0.29$ D by 6 and 12 months of follow-up, respectively (Figure 3). By this time, $96.6\%$ of the preoperative SE was corrected. Linear regression analysis of achieved versus attempted corrections revealed a slope of $1.05$ (slight over correction) with a predictability coefficient ($R^2$) of $0.88$ (Figure 3). Arithmetic mean ($\pm SD$) of astigmatism improved from preoperative value of $0.80\pm0.64$ to $0.19\pm0.31$ D ($P<0.001$). No eye suffered from astigmatism induction. Respectively, $90\%$ and $100\%$ of the eyes achieved $\pm0.5$ D and $\pm1.0$ D of astigmatism targeted (Figure 3).

**Predictability** Six months after the operation, $24$ ($80.0\%$) eyes achieved $\pm0.5$ D of target refraction. It was $96.7\%$ for $\pm1.0$ predictability. One-year after the operation, $\pm0.5$ D and $\pm1.0$ predictabilities improved to $96.7\%$ and $100\%$, respectively (Figure 3).

**Stability** Mean rate of SE change between third and sixth months of follow-up was $-0.01\pm0.11$ D/mo (Figure 3). During the next six months, mean SE regressed by a rate of $-0.01\pm0.06$ D/mo. Six months after the operation, mean SE of treated eyes was comparable to their mean SE at third month of follow-up ($P=0.4$). Similarly, mean SE obtained twelve months after the operation was comparable to mean SE at the sixth month of follow-up ($P=0.2$). In other words, treated eyes achieved a stable refraction by third month of follow-up. The number of eyes with more than $0.5$ D of change in SE between third and sixth months of follow-up was five ($16.7\%$). No eye had more than one dioptr change in SE during this period. Between 6th and 12th months of follow-up, number of eyes with more than $0.5$ and $1$ D change in SE were $3$ ($10\%$) and $0$, respectively.

**Vector Analysis** We achieved almost optimal correction of astigmatism according to vector analysis. Mean CI, DV, and ME were $1.03\pm0.26$, $0.19\pm0.31$, and $0.02\pm0.19$, respectively (Table 3). Mean CI was close to one and mean DV and ME were close to zero and indicated that magnitudes of targeted
and achieved corrections were almost comparable. Mean AE was 2.43 degree, which indicated that the axis of correction achieved was slightly counter-clockwise to the axis of correction intended (Figure 3).

Complications We detected no delayed re-epithelialization after the laser ablation. Complete re-epithelialization was observed in 90% and 100% of treated eyes 48 and 72h after the operation, respectively. One eye with trace and another eye with grade 1 + haze were detected three months after the operation which resolved till the next visit at sixth month of follow-up. We detected no corneal haze greater than 1 + throughout the follow-up.

Figure 3 Visual outcomes of transepithelial photorefractive keratectomy in high myopia The figure demonstrates results of uncorrected distance visual acuity (A), uncorrected distance visual acuity vs corrected distance visual acuity in terms of numbers of decimal line change (B), change in corrected distance visual acuity (C), spherical equivalent attempted vs achieved (D), spherical equivalent refractive accuracy (E), stability of spherical equivalent refraction (F), refractive astigmatism (G), target induced astigmatism vs surgically induced astigmatism (H), and refractive astigmatism angle of error (I).
Index of success; FI: Flattening index. of error; ME: Magnitude of error; CI: Correction index; IOS: Standard deviation; SIA: Surgically induced astigmatism; TIA: TransPRK: Transepithelial photorefractive keratectomy; SD: TransPRK in high myopia deteriorates CS also reduce central EOZ. This induces HOAs and as a result transform corneal asphericity from prolate to oblate. They due to delivery of uneven energy to different parts of cornea, moderate myopic eyes induced HOAs and deteriorated CS or did not affect it in high surgeries. Studies reported that PRK, LASIK, and LASIK myopia is promising compared to other laser-assisted refractive Our results for effects of TransPRK on visual quality in high myopic eyes undergoing LASIK and LASEK, respectively. It was reported to be 77% and 82%-97% in previous studies of single-step TransPRK in high myopic eyes in higher corrections attempted, which is the case in high myopic eyes. Single-step platform of TransPRK uses a population-based profile of corneal thickness to calculate amount of energy delivered to different parts of cornea. This prevents delivery of uneven amounts of energy to central cornea compared to peripheral cornea. We believe that this difference could account for better quality of vision outcomes of TransPRK in high myopia. We found promising efficacy of TransPRK in improvement of visual acuity and refraction of high myopic eyes. Difference between preoperative CDVA and postoperative UDVA is amongst the main parameters contributing to patient satisfaction. In this study, patients achieved a postoperative UDVA that was better than their preoperative CDVA by a mean difference of 0.07 which was comparable with a mean difference of 0.07 reported by Ghadhfan et al[32] and 0.03 reported by Aslanides et al[10] for single-step TransPRK in high myopia. In contrast, high myopic eyes treated by two-step TransPRK, mechanical PRK, or LASIK could not achieve a postoperative UDVA comparable with their preoperative CDVA. In our study, percentage of eyes achieving one-year postoperative UDVA equal to or higher than 20/20 and 20/25 were 87% and 100%, respectively. It was reported to be 77% and 82%-97% in previous studies of single-step TransPRK in high myopic eyes[8-10]. In other modalities, percentage of high myopic eyes achieving UDVA equal or above 20/25 were 65.5% in two-step TransPRK, 50%-87% in LASIK, 76% in LASEK, and 45%-75% in mechanical PRK[32-34].

DISCUSSION

Our results indicated that single-step TransPRK with aberration-free mode improves CS, visual acuity and refraction in high myopic eyes. It may not induce a clinically significant change in HOAs. Vector analysis demonstrated accuracy of astigmatism correction.

A recent study reported one-year refraction and visual acuity results of one-step TransPRK in high myopic eyes[10]. The study did not report CS and HOAs results. Our study is the first to report effect of single-step TransPRK on photopic and mesopic CSs as well as ocular and corneal HOAs in high myopic eyes. As hypothesized, TransPRK improved CSs in high myopic eyes. In addition, although most of HOAs were induced, the mean change observed in different HOAs ranged from 0.005 to 0.11 µm. This range of change is not expected to result in a clinically significant impairment of visual function[21]. Our results indicated promising results of single-step TransPRK in quality of vision in this high risk subgroup of refractive errors.

Our results for effects of TransPRK on visual quality in high myopia is promising compared to other laser-assisted refractive surgeries. Studies reported that PRK, LASIK, and LASIK induced HOAs and deteriorated CS or did not affect it in high myopic eyes[22-25]. They also worsened CS and HOAs in low to moderate myopic eyes[26-29]. It is postulated that laser ablations, due to delivery of uneven energy to different parts of cornea, transform corneal asphericity from prolate to oblate. They also reduce central EOZ. This induces HOAs and as a result deteriorates CS[18,29]. These changes are further emphasized in higher corrections attempted, which is the case in high myopic eyes[30-31]. Single-step platform of TransPRK uses a population-based profile of corneal thickness to calculate amount of energy delivered to different parts of cornea[6-9].

### Table 3: Results of vector analysis in eyes with high myopic astigmatism after TransPRK

<table>
<thead>
<tr>
<th>Parameters</th>
<th>Mean</th>
<th>SD</th>
<th>Min</th>
<th>Max</th>
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</thead>
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<td>SIA Magnitude (D)</td>
<td>0.78</td>
<td>0.56</td>
<td>0.25</td>
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<td>SIA Axis (degree)</td>
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<td>41.11</td>
<td>12.00</td>
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<tr>
<td>TIA Magnitude (D)</td>
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<td>0.64</td>
<td>0.25</td>
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<tr>
<td>TIA Axis (degree)</td>
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<td>38.30</td>
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</tr>
<tr>
<td>DV Magnitude (D)</td>
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<td>0.31</td>
<td>0.00</td>
<td>0.99</td>
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<td>DV Axis (degree)</td>
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<td>135.00</td>
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<tr>
<td>ME (D)</td>
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<td>0.19</td>
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<td>0.69</td>
</tr>
<tr>
<td>AE (degree)</td>
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<td>-75.14</td>
<td>86.04</td>
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<tr>
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<td>0.53</td>
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<td>IOS</td>
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<tr>
<td>FI</td>
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<td>0.27</td>
<td>-0.09</td>
<td>1.00</td>
</tr>
</tbody>
</table>

TransPRK: Transepithelial photorefractive keratectomy; SD: Standard deviation; SIA: Surgically induced astigmatism; TIA: Target induced astigmatism; DV: Difference vector; AE: Angle of error; ME: Magnitude of error; CI: Correction index; IOS: Index of success; FI: Flattening index.

Our findings showed that high myopic eyes achieved almost emmetropia by three months after TransPRK and then underwent slight regression till the twelfth month of follow-up. Similar pattern was seen in previous studies of single- and two-step TransPRK in correction of high myopia[10,32]. In our study, percentage of eyes achieving one-year postoperative UDVA equal or above 20/20 and 20/25 were 87% and 100%, respectively. It was reported to be 77% and 82%-97% in previous studies of single-step TransPRK in high myopic eyes[10,32]. In other modalities, percentage of high myopic eyes achieving UDVA equal or above 20/25 were 65.5% in two-step TransPRK, 50%-87% in LASIK, 76% in LASEK, and 45%-75% in mechanical PRK[32-34].

Almost 75% of preoperative arithmetic mean astigmatism of treated eyes were corrected by 12mo after the operation in our study. We did not detect induction of astigmatism in any treated eye. In support, Aslanides et al[10] reported high efficacy of single-step TransPRK in correction of high myopic astigmatism. Our study was the first to report results of vector analysis in correction of high myopic astigmatism by TransPRK. Promising values of CI, DV and ME in vector analysis further established the accuracy of astigmatism correction in our study. However, AE showed that there was a slight misalignment of correction axis achieved against correction axis attempted.
No eye lost any number of preoperative CDVA in our study. Similarly, previous studies reported that no high myopic eye treated with single-step TransPRK lost 2 or more lines of CDVA. In contrast, loss of two or more lines of CDVA have been reported in 15%, 2.7%, and 0.7% of high myopic eyes treated by mechanical PRK, LASEK, and LASIK, respectively. We also did not detect delayed re-epithelialization or notable degrees of haze after the operation. These findings imply acceptable safety of single-step TransPRK in correction of high myopia.

We postulated contribution of some parameters to promising findings obtained in our study. We scrutinized the demographic data and optical parameters (e.g. topographic map of the cornea) to best tailor the ablation plan for each individual eye. Some considerations in surgical technique such as individualized calculation of target refraction, large optical zones, immediate irrigation with chilled BSS, and appropriate regimen of MMC application should also be noted. In addition, aberration-free mode of our ablation profile prevented further alterations of physiologic corneal curvature and removal of corneal tissue that happens in aberration-guided ablations. We took advantage of a high-tech laser ablative system equipped with Intelligent Thermal Effect Control. Strong ultra violet protection and our modified post-operative regimen might contribute to the healing process of the cornea as well.

Almost low number of eyes included in this study and lack of a control group are limitations and should be considered in interpretation of our findings. Future studies on larger samples of high myopic eyes and controlled with parallel groups would yield more robust findings in the topic.

In conclusion, single-step TransPRK could be considered effective and safe in correction of high myopia. It improves both qualitative and quantitative visual functions in terms of CS, refraction and visual acuity. It may not result in clinically significant induction of HOAs. Future studies with larger sample size and longer follow-up seem necessary to draw more confident conclusions in the topic.

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Author Contribution: Adib-Moghaddam S: concept and design, writing the manuscript, critical revision of the manuscript, administrative, technical, or material support; Soleyman-Jahi S: concept and design, analysis and interpretation of data, writing the manuscript, critical revision of the manuscript, statistical expertise; Arba Mosquera S: concept and design, writing the manuscript, critical revision of the manuscript, administrative, technical, or material support; Hooshad N: data collection, writing the manuscript, critical revision of the manuscript; Tofighi S: data collection, writing the manuscript, critical revision of the manuscript.

Conflicts of Interest: Adib-Moghaddam S, None; Soleyman-Jahi S, None; Adili-Aghdam F, None; Arba Mosquera S, is an employee of SCHWIND eye-tech-solutions and is paid for employment. Additionally he owns rights for a patent pertaining to SCHWIND eye-tech-solutions with no related payment. Hooshad N, None; Tofighi S, None.

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