Photorefractive keratectomy in the management of postradial keratotomy hyperopia and astigmatism

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In clinical practice, there may be situations, in which an ophthalmologist visits a patient with severe post-RK refractive errors; herein comes the question of whether further surgical procedures bring acceptable refractive efficacy and safety.

Before 2000, to correct patients’ residual refractive errors, RK and laser refractive procedures were applied interchangeably.[6,7] In the literature, post-RK laser in situ keratomileusis (LASIK) has been acknowledged as a reasonable choice to correct the residual refractive errors;[8‑13] however, LASIK flap complications, epithelial ingrowths, and keratectasia are not uncommon.[14‑16]

Regarding the use of PRK, existing literatures lack consensus and none of the therapeutic procedures

INTRODUCTION

Radial keratotomy (RK) is now named as an obsolete procedure though by its advent in 1980’s RK had been considered to be a reasonable treatment for low to moderate degrees of myopia.[1‑2] As regards, the procedure of RK, 90%–95% radial incisions on the cornea were made to produce peripheral steepening and central flattening. The main complication of RK over the ensuing postoperative years has been refractive instability and progressive hyperopia.[3‑4] Since 1991, photorefractive keratectomy (PRK) has been used to correct residual refractive errors of RK.[5]

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put forward therein are completely convincing.[17] One of the protocols employs off-label use of intraoperative mitomycin-C (MMC) which can effectively control cornea haze as a major complication of post-RK PRK.[18-20] As far as we have been able to ascertain, the only recent large report on this topic comes from Brazil that enrolled 61 eyes,[21] since then few large series of patients has yet to be studied.[22,23] The present study was planned to address this topic and evaluate the refractive efficacy and safety of PRK with MMC in the management of post-RK hyperopia and astigmatism in a large Iranian cohort.

MATERIALS AND METHODS

Study design and ethics
This prospective nonrandomized, noncomparative, interventional case series enrolled consecutive eyes treated with PRK after RK in Iran between April 2012 and December 2014. All procedures were performed by 1 surgeon (M. G.) using a uniform technique with one protocol.

Patients had a positive defocus equivalent (consecutive hyperopia after primary RK surgery).

To minimize the difference of corneal parameters in the morning and evening, all subjects had a concise preoperative examination at about 10 AM. Examinations were uncorrected distant visual acuity (UDVA), corrected distant visual acuity (CDVA), subjective/cycloplegic refraction, wavefront (WF) analysis with Hartman-Shack based Zywave aberrometer, and Pentacam topography (Pentacam HR, Oculus, Wetzlar, Germany).

The study protocol adhered to the tenets of the Declaration of Helsinki and was approved by the Ethics Committee of Isfahan University of Medical Sciences. Data collection was in conformity with country laws, and all of the patients were informed about every possible side effects of each procedure. Signed informed consent was obtained from each patient before inclusion. All of patients’ data were recorded, documented, and reported confidentially.

Patients and the participation criteria
Eighty eyes from 58 patients were enrolled at baseline. Subjects were eligible for participation if they were otherwise healthy, literate with history of RK surgery with refraction of + 8.0 diopters of hyperopia, and −7.0 diopters of astigmatism. Corneal thickness should have been 480 μm or more.

The exclusion criteria of the study were cataract, keratoconus and corneal pathology, glaucoma and/or ocular hypertension, endocrine or collagen vascular diseases, diabetic retinopathy, pregnancy/lactation. Patients were also excluded if they did not undergo follow-up examinations for 6 months [Flow Chart 1].

Surgical method and postoperative care
Following a 5% a povidone-iodine scrub of eyelids and face, topical tetracaine 1% was instilled into each eye preoperatively. After primary preparations, the 9 mm central cornea was deepithelized by a blunt spatula. The flying spot Technolas 217-z excimer system (Bausch and Lomb) was employed for laser ablation.

Patients were treated with two protocols based on the following criteria:
1. In cases, in which (1) WF scan was undetectable during primary examinations; and/or, (2) WF data were not transferable to the excimer laser device, patients were treated with the tissue-saving (TS) mode
2. Patients with detectable/transferable WF were assigned to WF-guided advanced personalized treatment (APT).

The amount of ablation performed was 90% of cycloplegic refraction. MMC 0.02% was applied to the stromal bed for up to 60 s; the surface was irrigated with a balanced salt solution. Then, topical ciprofloxacin drop was instilled, and soft bandage contact lenses (Acuvue; Johnson and Johnson Vision Care, Jacksonville, FL, USA) were administered.

Postoperatively, topical betamethasone 0.1% was used every 3 h and topical ciprofloxacin every 6 h. On the day 6 of follow-up, patients were assessed for complete corneal epithelial healing; and consequently, contact lenses were removed.[24]

Up to 1-month after surgery, Fluorometholone and Artelac drops were used every 6 h. At this time, patients were evaluated for their intraocular pressure. UDVA, CDVA, cycloplegic refraction, WF, and grade of haze formation were assessed at 6 months.

Using a questionnaire, patients were asked for the following objective items: overall satisfaction (as more than 80%, 60%-80%, 40%-60%, and <40%), excellence of day/night vision (as excellent, good, moderate, and weak), and severity of postoperative symptoms, i.e. halo, photophobia, pain, burn, drying, and burning (as nil, mild, moderate, and severe).

Statistical measures
All data were collected in an Excel database (Microsoft Office 2003, Microsoft Corp.). Decimal UDVA and CDVA were converted from Snellen values to the logarithm of the minimum angle of resolution for statistical calculations.

Statistical analysis was performed using SPSS software version 13.0 (SPSS Inc., Chicago, IL, USA). Data were
expressed as means ± standard deviation and \( P < 0.05 \) was considered statistically significant. Paired \( t \)-test was used to assess the difference between preoperative and postoperative data. Linear regression analysis was used for the evaluation of the correlation between intended and achieved correction. Chi-square test was used for the analysis of qualitative data.

**RESULTS**

Of 58 patients included in the study, one missed the 6-months follow-up examinations and was excluded from our study. Of the rest of the patients, 18 were male, and 39 were female. Mean age was 43.40 ± 4.60 with a range of 31–56 years. The majority of the eyes (73%) had 8 RK incisions; 21.6% and 5.4% had 6 and 4 RK incisions, respectively.

Thirty-two and 47 eyes were managed by APT and TS modes, respectively. Demographics, visual and refractive results of the patients are summarized in Tables 1 and 2.

Pooled analysis of both APT and TS groups showed improvement in UDVA and CDVA. The amount of sphere, cylinder, corneal cylinder, spherical equivalent, defocus equivalent, and total aberration showed improvement as well [Tables 1 and 2].

Changes in higher order and spherical aberration were not statistically significant (\( P = 0.670 \) and 0.405, respectively). Eighty-four percent of our patients achieved UDVA of 20/40 or better. Figure 1 shows that attempted spherical equivalent was strongly correlated with achieved one (\( R^2 = 0.8671 \)). Figure 2 depicts changes in pre- and post-operative astigmatism. Figures 3-5 show excellence of day/night vision, severity of postoperative symptoms, and overall satisfaction, respectively.

![Figure 1: Spherical equivalent attempted versus achieved](image)

**Table 1: Visual and refractive parameters pre- and post-operative**

| Group (n) | APT (32) | Tissue-saving (47) | Pooled analysis (79 eyes) |
|-----------|----------|--------------------|--------------------------|
| Male/female | 11/21 | 16/31 | 27/52 |
| Age (years), mean±SD (range) | 43.33±6.4 (31-56) | 43.43±3.8 (36-50) | 43.4±4.6 (31-56) |
| UDVA (logMAR) | | | |
| Preoperative | 0.49±0.35 (0.15-1.3) | 0.45±0.37 (0.05-2) | 0.46±0.36 (0.05-2) |
| Postoperative | 0.12±0.13 (0-0.4) | 0.23±0.18 (0-0.7) | 0.2±0.17 (0-0.7) |
| \( P \) | 0.002 | 0.001 | 0.000 |
| CDVA (logMAR) | | | |
| Preoperative | 0.16±0.16 (0-0.52) | 0.18±0.13 (0-0.52) | 0.18±0.13 (0-0.52) |
| Postoperative | 0.07±0.12 (0-0.04-0.4) | 0.11±0.12 (0-0.52) | 0.10±0.12 (0-0.04-0.52) |
| \( P \) | 0.006 | 0.009 | 0.000 |
| Sphere (D) | | | |
| Preoperative | 1.56±1.33 (0.0-4.0) | 3.13±1.97 (0-0.7) | 2.68±1.93 (0-0.7) |
| Postoperative | 0.58±0.81 (0.0-1.75) | 0.11±0.9 (-1.75-2.75) | 0.24±0.89 (-1.75-2.75) |
| \( P \) | 0.061 | 0.000 | 0.000 |
| Cylinder (D) | | | |
| Preoperative | -2.77±1.46 (-5.25-0.75) | -2.8±2.04 (-8.0-0.0) | -2.8±1.87 (-8.0-0.0) |
| Postoperative | -0.62±0.69 (-1.75-0.0) | -1.37±1.0 (-3.5-0.0) | -1.15±0.98 (-3.5-0.0) |
| \( P \) | 0.001 | 0.000 | 0.000 |
| SE (D) | | | |
| Preoperative | 0.18±1.81 (-2.37-3.12) | 1.73±2.1 (-2.0-6.0) | 1.29±2.12 (-2.37-6.0) |
| Postoperative | 0.27±0.84 (-0.62-2.75) | -0.58±1.02 (-2.75-2.75) | -0.33±1.04 (-2.75-2.75) |
| \( P \) | 0.387 | 0.000 | 0.000 |
| DE (D) | | | |
| Preoperative | 2.95±1.15 (1.37±5.5) | 4.53±2.32 (0.62-9.37) | 4.08±2.17 (0.62 to 9.37) |
| Postoperative | 0.89±0.93 (0.0-2.75) | 1.16±0.87 (0.0-3.62) | 1.11±0.89 (0.0 to 3.62) |
| \( P \) | 0.001 | 0.000 | 0.000 |

CDVA = Corrected distance visual acuity; DE = Defocus equivalent; logMAR = Logarithm of the minimum angle of resolution; SD = Standard deviation; SE = Spherical equivalent; UDVA = Uncorrected distance visual acuity; APT = Advanced personalized treatment
DISCUSSION

For refractive surgeons, encountering RK patients with consecutive hyperopia is a challenging situation. This may come either from a primary overcorrection or from a progressive postoperative process. To manage such a complicated condition, LASIK and PRK have been proposed to be promising.\textsuperscript{[18-20]} Although either mechanical microkeratome or the femtosecond LASIK surgeries have shown favorable initial outcomes, such methods are not without complications.\textsuperscript{[18,19]} Flap-related issues, especially epithelial in growth and splitting RK incisions can cause flap fragmentation along with the possibility corneal ectasia. Due to such possibilities, some authors believe that LASIK should be avoided in eyes with more than 8 RK incisions.\textsuperscript{[18,19]} About the advantages of PRK over LASIK, the

### Table 2: Aberrometry and keratometry parameters pre- and post-operative

| Group (n) | APT (32) | Tissue-saving (47) | Pooled analysis (79 eyes) |
|----------|----------|--------------------|---------------------------|
| **Total aberration (6 mm)** | | | |
| Preoperative | 4.16±1.23 (1.75-5.75) | 4.67±1.94 (1.92-8.32) | 4.47±1.69 (1.92-8.32) |
| Postoperative | 2.39±0.64 (1.6-3.54) | 2.97±1.09 (1.56-5.32) | 2.74±0.97 (1.56-5.32) |
| \(P\) | 0.000 | 0.000 | 0.000 |
| **Higher-order aberration (6 mm)** | | | |
| Preoperative | 1.28±0.47 (0.56-2.3) | 1.98±0.91 (0.41-4.04) | 1.67±0.83 (0.41-4.04) |
| Postoperative | 1.29±0.54 (0.63-2.38) | 1.63±0.61 (0.78-2.73) | 1.59±0.59 (0.63-2.73) |
| \(P\) | 0.901 | 0.031 | 0.670 |
| **Spherical aberration (6 mm)** | | | |
| Preoperative | 1.12±0.44 (0.54-2.22) | 1.82±0.94 (0.24-3.89) | 1.53±0.82 (0.24-3.89) |
| Postoperative | 1.25±0.52 (0.62-2.28) | 1.55±0.62 (0.57-2.73) | 1.44±0.60 (0.57-2.73) |
| \(P\) | 0.371 | 0.149 | 0.405 |
| \(K_{\text{min}}\) (D) | | | |
| Preoperative | 37.69±2.63 (32.8-42.1) | 35.09±4.42 (29.5-46.5) | 35.83±4.13 (29.5-46.5) |
| Postoperative | 38.4±3.31 (32-43.1) | 37.95±3.77 (32-46.5) | 38.08±3.61 (32-46.5) |
| \(P\) | 0.259 | 0.000 | 0.000 |
| \(K_{\text{max}}\) (D) | | | |
| Preoperative | 40.43±3.15 (36.8-46.3) | 37.93±4.21 (32.4-48) | 38.65±3.98 (32.4-48) |
| Postoperative | 40.06±3.28 (32.9-44.9) | 39.54±3.11 (32.9-48) | 36.69±3.44 (32.9-48) |
| \(P\) | 0.659 | 0.001 | 0.016 |
| **Average K (D)** | | | |
| Preoperative | 39.05±2.73 (34.7-44.0) | 36.46±3.98 (31.5-47.3) | 37.18±3.67 (31.5-47.3) |
| Postoperative | 38.95±3.23 (32.4-43.7) | 38.77±3.66 (32.4-47.3) | 38.82±3.50 (32.4-47.3) |
| \(P\) | 0.920 | 0.000 | 0.000 |
| **Corneal cylinder** | | | |
| Preoperative | 2.82±1.38 (0.6-5.0) | 3.18±2.66 (0.3-11.5) | 3.08±2.35 (0.3-11.5) |
| Postoperative | 1.66±1.42 (0.2-5.0) | 1.59±1.04 (0.2-4.1) | 1.61±1.12 (0.2-5.0) |
| \(P\) | 0.008 | 0.001 | 0.000 |

\(K = \text{Keratometry}; \text{APT} = \text{Advanced personalized treatment}\)
following points are noteworthy: first: PRK avoids additive weakening of the cornea, as a risk factor of ectasia; and second: PRK does not interrupt with RK incisions, saving the remaining corneal structural integrity.\cite{19} Corneal ectasia has occurred neither in the present study nor in our daily practice over recent years. Taken together, based on the current literature on the management of post-RK hyperopia, PRK seems to be a reasonable method with a favorable safety profile. However, in this method, increased risk of haze cannot be overlooked as a limitation in handling high hyperopia.\cite{19} With administration of MMC in PRK, corneal haze has become less important.\cite{20} In some reports, there had been a matter of uncertainty about the safety of using MMC in RK because increased permeability of the cornea for MMC due to deep RK incisions could cause excessive toxicity to the eye.\cite{18} Notwithstanding, use of MMC for 60 s in a study and up to 40 s in another caused no endothelial cell loss after 1 year.\cite{18,21}

In our study, 23% of patients achieved 20/20 UDVA, and 84% achieved 20/40. Such findings are in correspondence to other reports, in which a UDVA of 20/40 or better was observed in 70%,\cite{19} 65%,\cite{11} 82%,\cite{9} 96%,\cite{13} and 91%.\cite{12}

In our study, over 61% of patients gained 1 line or more of CDVA, 29% had no change, and about 10% lost 1 line or more at 6 months of follow-up. This is in line with previous studies in which 68% of eyes gained at least 1 line after 1 year.\cite{19} In studies on hyperopic LASIK, this value was 16%, 20%, 24%, 30%, and 56%.\cite{8,11-13} Regarding the above-mentioned reports, the relative favorable safety profile PRK against LASIK may be due to the absence of flap-related complications.\cite{19}

It is crucial for practitioners to warn their patients about the fact that refractive instability may still progress regardless of the use of the laser method of vision correction. Indeed, patients and practitioners should be aware that refractive correction is not in deal with the cornea structural stabilization.\cite{20} In spite of an early reduction in hyperopia with corrective methods, hyperopic changes are shown to persist in most of the cases; and in this regard, factors that might play role are RK-induced progressive central corneal flattening, alterations in the lens, and post-PRK corneal healing.\cite{21}

**CONCLUSION**

This was the largest clinical study to have been undertaken on the management of post-RK refractive errors. PRK
seems to bring favorable outcome and safety profile in the management of post-RK hyperopia and astigmatism.

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Conflicts of interest
There are no conflicts of interest.

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