Relative Contribution of Central and Peripheral Aberrations to Overall High Order Corneal Wavefront Aberration

Marco Lombardo, MD; Giuseppe Lombardo, Eng, PhD; Michele Manzulli, MD; Marino Palombi, MD; Sebastiano Serrao, MD, PhD

ABSTRACT

PURPOSE: To analyze the influence of specific combinations of corneal high order aberrations on the optical image quality of the cornea before and after photorefractive keratectomy (PRK) for low to high myopia and myopic astigmatism.

METHODS: Corneal topography was obtained for 80 eyes that underwent PRK using a scanning-spot excimer laser. The eyes were subdivided into three groups according to the preoperative refraction. The topographical data were imported into a custom software program that combined the Zernike high order terms having the same sign and angular frequency up to seventh order for 4-mm and 6-mm pupils, ie, coma and spherical aberrations, and midperipheral and peripheral high order aberrations.

RESULTS: Photorefractive keratectomy induced a significant amount of the root-mean-square (RMS) values of the combinations of midperipheral and peripheral high order aberrations over the smaller pupil size for deeper myopic ablations (P<.05). Over the larger pupil, spherical myopic ablations showed a significant increase (P<.05) of the RMS values of coma and spherical aberrations. In the astigmatism group, the combination of terms having higher angular frequency increased significantly (P<.05) after surgery both over 4-mm and 6-mm pupils.

CONCLUSIONS: After surface ablation, ablation depth and profile significantly influence the distribution and contribution of determined combinations of high order aberrations to the overall high order corneal wavefront aberration. Terms having high angular frequency were increased following large myopic correction and wide treatment zone. Quality of the whole corneal optics will be enhanced by designing future ablation profiles to compensate for peripheral high order optical aberrations.

Wavefront-guided customized ablation is an emerging technique in refractive surgery performed with the goal to minimize the whole eye optical aberrations. With the development of this technique, many authors are establishing aberration standards for normal individuals and the visual effects of each aberration to optimize the results of correction. Improvements in the technology and methodology of wavefront-guided laser ablation are expected to correct the variability in some individual factors and thereby further optimize the quality of vision. Besides the biophysical factors that cannot be predicted by a wavefront approach and need to be further investigated to minimize unwanted changes of the cornea after refractive procedures, the assessment of the preoperative eye should take into account the interaction between aberrations and the effect of their combinations on vision function. The wide variability of high order aberrations of the eye in the healthy population and the complex combination of individual terms in modulating visual performance make it difficult to correctly treat the individual high order wavefront aberration to achieve a super-normal vision.

Adaptive optics and the generation of aberrated acuity charts can reliably inform us on the combined effects of high order aberrations on the retinal image. Applegate et al stated the interaction between terms having the same sign and angular frequency (eg, 0, –2, or 2) may produce a combined effect on improving acuity if compared to the individual effect of each mode. Furthermore, it is known that for an equal amount of root-mean-square (RMS) error, each mode of Zernike expansion has a different impact on visual performance and that the Zernike modes concentrated near the cen

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The corneal epithelium was removed using the Amoils’ brush. The PRK procedure was performed using a Chiron Technolas Keracor 217C excimer laser (Bausch & Lomb, Dornach, Germany) with an ablation zone diameter of 6 mm and a transition zone up to 9 mm in diameter; an active eye tracker device was used. The astigmatism group was treated with a cross-cylinder technique \(^{23}\) (photorefractive astigmatic keratectomy) with a 6-mm ablation zone diameter and a transition zone up to 9 mm in diameter. As described previously, \(^{24}\) phototherapeutic keratectomy was performed (using a viscous-masking 0.25% sodium hyaluronate solution for a smoothing technique) at the end of the procedure. The smoothing technique proved to be more effective in inducing less high order aberrations than standard PRK. \(^{24}\)

All patients underwent a complete ocular evaluation, including corneal topography and pupillometry (Keratron Scout; Optikon 2000, Rome, Italy). The topographer is equipped with a special lamp board placed behind the mires cone that illuminates the mires with either visible light or infrared to obtain pupil measurements in both photopic and mesopic light conditions. The follow-up protocol of this study included examinations preoperatively and 1 year postoperatively. For each eye, measurements were repeated five times to assess the repeatability of the topography. \(^{25}\) Reproducibility and accuracy of videokeratoscopic measurements were tested using an artificial spherical cornea provided by the manufacturer and calculated from five independent measurements. The standard deviation for the high order RMS surface error of the sphere was ±0.007 µm and ±0.02 µm at 4 mm and 6 mm diameter, respectively.

The topographer software calculated the corneal wavefront aberration on the corneal elevation with respect to an ideal aspherical corneal shape with eccentricity 1/n (where n=1.3375); the wavefront aberration then was obtained from the derivatives using a least-squares best-fit \(^{26}\) procedure and described as a seventh order Zernike polynomial expansion. \(^{27}\) We then exported the aberration data output, computed with respect to the line of sight using the move axis function of the topographer and the photopic and mesopic pupil diameters from the topographer for analysis. Preoperative and 1-year postoperative corneal aberration data were computed for 4-mm and 6-mm pupils. \(^{28}\) The smaller pupil size ensured that we analyzed the high order corneal wavefront aberration changes exclusively within the ablation zone, whereas the larger pupil size allowed us to calculate the high order corneal wavefront aberration in the scotopic condition. First and second orders were not considered. The Zernike polynomials were normal-
ized such that over the pupil, the RMS for each Zernike mode was unity.29 The recommended Optical Society of America notation is used describing individual Zernike terms with a two-index scheme.30,31

Pre- and postoperative aberration data were processed using custom software written in MATLAB (software version 7.0) (MathWorks Inc, Natick, Mass). The parameters analyzed included: 1) the RMS of the spherical aberration (the square root of the sum of the squared coefficients of Z0 and Z2); 2) the RMS of coma (the square root of the sum of the squared coefficients of Z3, Z1, Z1, and Z2); 3) the RMS of midperipheral high order aberrations (the square root of the sum of the squared coefficients Z3, Z3, Z3, Z3, Z3, Z3, Z3, Z3, and Z2); and 4) the RMS of peripheral high order aberrations (the square root of the sum of the squared coefficients Z3, Z3, Z3, Z3, Z3, Z3, Z3, and Z2). The combinations of central and paracentral high order terms of the Zernike pyramid for analysis were chosen based on previous publications,15-17,20,32-34 and the combinations of terms having higher angular frequency represented a subset of the possible combinations of higher order Zernike modes.16 To assess the accuracy of this type of analysis, a vector approach has been applied to the Zernike representation of the wavefront error. Terms having single or higher angle symmetries were represented in the magnitude/axis method, as reported by Campbell35 and also in the American National Standards Institute (ANSI) standard for reporting wavefront error (ANSI Z80.28-2004). The alignment of vector axes for each determined combination of terms having the same azimuthal index has been examined. For example, the mean angle of the vector axes created by combining functions having a been examined. For example, the mean angle of the vector axes created by combining functions having a

RESULTS

Table 1 summarizes the pre- and postoperative mean spherical equivalent refraction for all of the study groups. The refractive target was emmetropia in all cases. All of the procedures were uneventful; after surgery, no eye lost one or more Snellen lines of spectacle-corrected visual acuity. The average photopic and mesopic pupil diameters were 3.40±0.55 mm and 5.77±1.03 mm, respectively, in the low myopia group; 3.77±0.78 mm and 5.69±1.04 mm, respectively, in the high myopia group; and 3.41±0.38 mm and 5.55±0.75 mm, respectively, in the astigmatism group.

The preoperative analysis of the high order aberrations showed absolute values that were highest for the coma terms in comparison with determined combinations of Zernike modes independent of spherical equivalent and cylinder. The level of significance was .05 for all of the tests performed.
TABLE 1

Mean (±SD) Preoperative and 1-year Postoperative Spherical Equivalent Refraction (D) for the Three Study Groups

<table>
<thead>
<tr>
<th>Group</th>
<th>Preoperative</th>
<th>Postoperative</th>
<th>P Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Low myopia</td>
<td>-3.04±0.73</td>
<td>+0.03±0.27</td>
<td>&lt;.05</td>
</tr>
<tr>
<td>(range: -1.50 to -4.50 D) (n=30)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>High myopia</td>
<td>-5.91±1.26</td>
<td>+0.07±0.38</td>
<td>&lt;.05</td>
</tr>
<tr>
<td>(range: -4.60 to -8.00 D) (n=30)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Astigmatism</td>
<td>-4.21±2.42</td>
<td>-0.75±0.83</td>
<td>&lt;.05</td>
</tr>
<tr>
<td>(range: -2.00 to -5.00 D cylinder) (range: -0.25 to -5.50 D sphere) (n=20)</td>
<td></td>
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</tr>
</tbody>
</table>

sis of the high order RMS wavefront variance showed no distinction between the study groups, as reported previously, except for slightly larger values in the astigmatism group in comparison with the simple myopia groups. Tables 2, 3, and 4 summarize the mean RMS of determined combinations of high order aberrations for all of the study groups.

TABLE 2

Mean (±SD) Preoperative and Postoperative Root-Mean-Square of High Order Aberrations (µm) in 4-mm and 6-mm Pupils for the Low Myopia Group

<table>
<thead>
<tr>
<th></th>
<th>4-mm Pupil</th>
<th>6-mm Pupil</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Preoperative</td>
<td>Postoperative</td>
</tr>
<tr>
<td>Coma</td>
<td>0.088±0.043</td>
<td>0.124±0.120</td>
</tr>
<tr>
<td>Spherical aberration</td>
<td>0.058±0.016</td>
<td>0.100±0.098*</td>
</tr>
<tr>
<td>Midperipheral high order aberrations</td>
<td>0.071±0.026</td>
<td>0.070±0.029</td>
</tr>
<tr>
<td>Peripheral high order aberrations</td>
<td>0.044±0.015</td>
<td>0.052±0.022</td>
</tr>
<tr>
<td>*Statistically significant P&lt;.05</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

TABLE 3

Mean (±SD) Preoperative and 1-year Postoperative Root-Mean-Square of High Order Aberrations (µm) in 4-mm and 6-mm Pupils for the High Myopia Group

<table>
<thead>
<tr>
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<th>6-mm Pupil</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Preoperative</td>
<td>Postoperative</td>
</tr>
<tr>
<td>Coma</td>
<td>0.081±0.039</td>
<td>0.206±0.192*</td>
</tr>
<tr>
<td>Spherical aberration</td>
<td>0.074±0.022</td>
<td>0.110±0.026*</td>
</tr>
<tr>
<td>Midperipheral high order aberrations</td>
<td>0.069±0.024</td>
<td>0.111±0.047*</td>
</tr>
<tr>
<td>Peripheral high order aberrations</td>
<td>0.057±0.027</td>
<td>0.076±0.034*</td>
</tr>
<tr>
<td>*Statistically significant P&lt;.05</td>
<td></td>
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RMS of the high order RMS wavefront variance showed no distinction between the study groups, as reported previously, except for slightly larger values in the astigmatism group in comparison with the simple myopia groups. Tables 2, 3, and 4 summarize the mean RMS of determined combinations of high order aberrations for all of the study groups.

At the end of follow-up, with the exception of the astigmatism group, the mean RMS of coma and spherical aberration increased over a 4-mm pupil. In detail,
The increase in the magnitude value of spherical aberration was significant in the low myopia group ($P<.001$), and there was a statistically significant increase both for coma ($P<.001$) and spherical aberration ($P<.001$) in the high myopia group. The surgically induced amount of both coma and spherical aberration was significantly related to the amount of refractive correction ($R=0.43$ and $P<.001$ for coma; $R=0.36$ and $P<.01$ for spherical aberration). In the astigmatism group, there was a decrease, although not statistically significant, in the mean RMS of coma and spherical aberrations ($P>.05$) over the smaller pupil size. The low myopia group exhibited no significant changes in the RMS values of midperipheral and peripheral high order aberrations over a 4-mm pupil after surgery ($P>.05$), whereas the RMS error values of the combinations of terms with higher angular frequency (ie, midperipheral and peripheral high order aberrations) were significantly increased in the high myopia ($P<.05$) and astigmatism groups ($P<.05$) over the smaller pupil diameter.

Coma-like aberration increased in all of the study groups over a 6-mm pupil size; however, the increase was statistically significant only in the two spherical ablation groups ($P<.05$) and it was significantly related to the amount of refractive correction ($R=0.42$ and $P<.01$) (Fig 1). With pupil dilation, a large and significant increase in spherical aberration ($P<.001$) was observed in both spherical myopia groups compared to the preoperative state, with a greater increase for deeper ablations (Fig 2). The postoperative spherical aberration was not statistically different ($P>.05$) compared with the preoperative state in the astigmatism group over the larger pupil size. The combinations of terms having high angular frequency were not significantly increased after spherical myopic ablations ($P>.05$). On the other hand, the surgically induced increase in mid-peripheral and peripheral high order aberrations were significantly increased in the low myopia ($P<.05$) and astigmatism groups ($P<.05$) over the smaller pupil diameter.

### TABLE 4

<table>
<thead>
<tr>
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<th>4-mm Pupil</th>
<th>6-mm Pupil</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Preoperative</td>
<td>Postoperative</td>
</tr>
<tr>
<td>Coma</td>
<td>0.123±0.042</td>
<td>0.112±0.050</td>
</tr>
<tr>
<td>Spherical aberration</td>
<td>0.073±0.016</td>
<td>0.057±0.034</td>
</tr>
<tr>
<td>Midperipheral high order aberrations</td>
<td>0.083±0.032</td>
<td>0.130±0.041*</td>
</tr>
<tr>
<td>Peripheral high order aberrations</td>
<td>0.040±0.009</td>
<td>0.058±0.020*</td>
</tr>
</tbody>
</table>

*Statistically significant $P<.05$
Peripheral (P<.05) and peripheral (P<.05) high order aberrations was statistically significant in the astigmatism group over a dilated pupil. Further, the increase in midperipheral and peripheral high order aberrations was significantly related to the amount of cylinder correction (R = −0.57 and P < .05 for midperipheral high order aberrations; R = −0.47 and P < .05 for peripheral high order aberrations) (Fig 3).

Corneal PSFs then were evaluated for each study group by averaging the effect of high order aberrations on the imaging performance of the first optical surface of the cornea, as specified in detail in a previous work.49 The postoperative increased contribution of spherical aberration to overall high order wavefront aberration decreased the optical quality of the anterior corneal optics following simple myopic PRK, especially for deeper ablations (Fig 4), whereas following astigmatic correction, the optical image quality of the anterior cornea was mainly affected by the increased contribution of peripheral asymmetrical high order aberrations to overall high order corneal wavefront aberration.

**DISCUSSION**

In an effort to optimize visual performance following procedures designed to reduce optical aberrations of the eye, a great deal of work has been carried out on how laser refractive surgery can modify both the distribution and contribution of specific high order terms to the overall high order wavefront aberration and the effect of their combinations on vision function.13,14,16,41,42 Studies have demonstrated that each high order Zernike mode influences visual performance differently and that high order aberrations can interact specifically to improve acuity despite the increase in total wavefront error.15-17 It has been suggested that minimizing the aberrations near the center of the Zernike pyramid may improve visual performance after refractive procedure. Besides the role of the eye optics on image formation,15-17 this phenomenon may be related to the Stiles-Crawford effect,43 which causes the retina to have a less effective response to the peripheral portions of the aberrant incident wavefront. The Stiles-Crawford effect may have an additional retinal basis in the optical behavior of cone photoreceptors with a differential sensitivity of the cones to light from different points in the pupil.44-46 However, less attention has been paid to the combination of Zernike terms having higher angular frequency, especially when the total RMS is relatively low.

To determine whether peripheral corneal high order aberrations may affect the optical image quality of the anterior cornea following PRK, we developed a power software tool that allowed us to separately analyze the weight and effect of specific combinations of Zernike high order terms having the same sign and angular frequency over two pupil diameters (4 mm and 6 mm) to characterize the optical image quality of the anterior cornea in different light conditions.

Previous works have shown that PRK changes the relative contribution of individual high order aberrations to the overall wavefront aberration and that the magnitude of the high order aberrations greatly increases with the attempted correction as well as pupil dilation.33,47-50 Our findings agreed with previous results on the increase of high order aberrations following standard myopic laser correction. Preoperatively, aberrations were relatively low and dominated by coma-like aberration in all of the study groups over the smaller pupil size; following surface ablation, the magnitude of the corneal first surface wavefront aberration was increased within the optical zone, and the distribution of determined combinations of high order aberrations varied according to the amount of spherical equivalent treated and the ablation profile.

We noted the positive spherical aberration value increased significantly following the spherical treatments even within the central 4-mm pupil diameter. In a previous work, Hersh et al51 hypothesized changes of corneal asphericity within the ablation zone could be related to the effect of corneal convexity on the ef-
effective fluence of the incoming laser beam and hence the resulting decrease in peripheral fluence may be the cause for unforeseen underablation of the peripheral treatment zone with a consequent oblate result. In addition, Marcos et al\textsuperscript{52} stated the increase in spherical aberration for the central 4 mm of the cornea after myopic ablation was not due to the ablation profile and parameters but possibly to biomechanical and healing effects; nevertheless, ablation algorithms are proprietary, and we cannot correctly simulate them.\textsuperscript{53} On the other hand, the cross-cylinder technique did not deeply modify spherical aberration, probably due to the flattening of the corneal periphery induced by photorefractive astigmatic keratectomy ablation.\textsuperscript{54}

Over a dilated pupil, spherical-like aberration dominated the high order aberration structure after simple myopic ablations, whereas peripheral asymmetrical aberrations were predominant following cylindrical treatment. The contribution of combinations of terms having higher angular frequency (ie, midperipheral and peripheral high order aberrations) to the overall high order wavefront aberration over the smaller pupil size was significantly increased by either the deeper spherical myopic and cylindrical ablations, whereas after pupil dilation, peripheral aberrations were found to significantly change only with cylindrical corrections. Low myopia treatment did not appear to influence the quality of peripheral corneal optics. These results may be related primarily to the alignment of ablation on the pupillary axis, which is decentered with respect to the corneal reference axis,\textsuperscript{55} and to the asymmetric biomechanical response between the nasal and temporal meridians of the cornea to photoablation, which has been shown to be positively related both to the amount of tissue removed as well as to larger treatment zones.\textsuperscript{56} Hence, the greater impact of peripheral high order aberrations on the overall high order corneal wavefront in comparison with paracentral high order aberrations following cylindrical ablation may result from the more peripheral ablation of photorefractive astigmatic keratectomy and then from the different biomechanical properties of the periphery of the corneal plane, as stated previously.\textsuperscript{56}

The PSF was more practical than the normalized...
Zernike expansion to show the quality of the corneal optics because of the complex interaction between modes at low levels of optical error. In this work, we confirmed that deeper myopic ablations have a greater impact on the image quality formation of the central corneal optics and showed that the more peripheral the ablation, the more the image quality of peripheral corneal optics worsened. Further work is required to understand the effect of the complex interactions of individual terms on visual performance in both natural and treated eyes. Nevertheless, weighting the effect of interactions between all of the possible combinations of Zernike terms on visual performance may be too complicated to realistically achieve.

Our data cannot be extrapolated for LASIK or other excimer laser systems in which the effect of the flap creation and the proprietary ablation profile and parameters may have different influences on the corneal remodeling and the related biomechanical and optical responses of the cornea. Limitations of our work include the fact that we did not consider internal wavefront aberrations. It is widely held that wavefront-guided procedures should always take the corneal front surface and the total wavefront aberrations into consideration. Hence, full knowledge of the preoperative corneal topography as well as of the ocular high order aberration pattern and its impact on visual performance would be necessary for designing an ideal compensating algorithm for the individual cornea to maximize the quality of the whole eye optics, especially in eyes with large pupils. An accurate spherocylindrical correction combined with an additional midperipheral ablation to optimize the asphericity of the postoperative corneal profile and minimize the regional asymmetries of the cornea may improve future advanced ablations.

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