The Effect of Optical Zone Decentration on Lower- and Higher-Order Aberrations after Photorefractive Keratectomy in a Cat Model

Jens Bühren,1 Geunyoung Yoon,1,2 Shawn Kenner,1,3 Scott MacRae,1,2 and Krystel Huxlin1,2

PURPOSE. To simulate the effects of decentration on lower- and higher-order aberrations (LOAs and HOAs) and optical quality, by using measured wavefront error (WFE) data from a cat photorefractive keratectomy (PRK) model.

METHODS. WFE differences were obtained from five cats’ eyes 19 ±7 weeks after spherical myopic PRK for −6 D (three eyes) and −10 D (two eyes). Ablation-centered WFEs were computed for a 9.0 mm pupil. A computer model was used to simulate decentration of a 6-mm subaperture in 100-μm steps over a circular area of 3000 μm diameter, relative to the measured WFE difference. Changes in LOA, HOA, and image quality (visual Strehl ratio based on the optical transfer function; VSOTF) were computed for simulated decentrations over 3.5 and 6.0 mm.

RESULTS. Decentration resulted in undercorrection of sphere and induction of astigmatism; among the HOAs, decentration mainly induced coma. Decentration effects were distributed asymmetrically. Decentration >1000 μm led to an undercorrection of sphere and cylinder of >0.5 D. Computational simulation of LOA/HOA interaction did not alter threshold values. For image quality (decrease of best-corrected VSOTF by >0.2 log units), the corresponding thresholds were lower. The amount of spherical aberration induced by the centered treatment significantly influenced the decentration tolerance of LOAs and log best corrected VSOTF.

CONCLUSIONS. Modeling decentration with real WFE changes showed irregularities of decentration effects for rotationally symmetric treatments. The main aberrations induced by decentration were defocus, astigmatism, and coma. Treatments that induced more spherical aberration were less tolerant of decentration. (Invest Ophthalmol Vis Sci. 2007;48:5806–5814) DOI:10.1167/iovs.07-0661

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Correct alignment of the ablation to the visual axis of the eye is an essential requirement for optimal outcome in laser refractive surgery (LRS). Decentration of the ablation zone leads to incomplete refractive correction and induction of higher-order aberrations (HOAs), especially coma.1-4 The expected benefit of less HOA induction in eye tracker–controlled treatments has been demonstrated.5 However, decentration still occurs as a result of misalignment of the tracking system,6 static registration errors due to surgeon offsets,7 and pupil center shifts as a function of dilation.8 In most cases, the magnitude of such misalignments is <500 μm.1,7,9,10 A recent study showed that in uneventful wavefront-guided LASIK, coma induction occurred in a random fashion, independent of factors such as attempted correction and optical zone (OZ) diameter.11 Thus, microdecentrations can be considered ubiquitous, random errors; however, their impact on optical quality is poorly understood.10 In contrast, gross decentrations of >500 μm are one of the most visually disturbing complications after LRS. Besides causing severe deterioration of visual quality, such complications are difficult to treat, and success is often limited.12-18

Although several studies on decentration-induced aberrations after conventional1-2,7 and wavefront-guided LRS3-4,8,19 have been published, all assumed a perfect ablation and did not consider the inherent induction of HOA which occurs in real corneas as a result of wound healing and biomechanical effects.20,21 The present study was conducted to investigate the effects of decentration of the laser ablation relative to the entrance pupil of the eye on LOA, HOA, and optical quality, in a cat photorefractive keratectomy (PRK) model. Although the optical effects of PRK for myopia, such as reduction of defocus, induction of coma, and positive spherical aberration are similar in cats and humans,22,23 the greater corneal surface area and the naturally large scotopic pupil diameter (PD) of ~12 mm in cats allowed us to measure wavefront changes well beyond the ablation OZ. A simplified computational model was used to simulate decentration effects over a circular area of 5000 μm in diameter by calculating wavefront errors (WFEs) for systematically offset subapertures of 3.5 and 6.0 mm. Using this paradigm, we assessed (1) the nature and magnitude and spatial distribution of optical aberrations induced by different amounts of decentration, (2) the impact of such aberrations on theoretical optical quality, (3) whether residual refractive errors could be partially attributed to microdecentrations (<500 μm), and (4) the impact of optical aberrations induced by laser refractive surgery on tolerance of decentration.

MATERIALS AND METHODS

Subjects

Data were obtained from five eyes of five normal, male domestic short hair cats (Felis catus), who underwent myopic PRK with an uncomplicated follow-up of at least 3 months and in which wavefront aberrations could be measured over a PD of 9 mm. Procedures were

5806
Table 1. Treatment Characteristics and WFE Changes for the Centered Treatment (ΔW(x, y)) over 6- and 9-mm PDs

<table>
<thead>
<tr>
<th>Eye</th>
<th>Treatment (D)</th>
<th>OZ (mm)</th>
<th>TTZ (mm)</th>
<th>PD (mm)</th>
<th>Sphere (D)</th>
<th>Cylinder (D)</th>
<th>Axis (°)</th>
<th>Total HOA RMS (μm)</th>
<th>Coma RMS (μm)</th>
<th>SA RMS (μm)</th>
<th>rHOA RMS (μm)</th>
</tr>
</thead>
<tbody>
<tr>
<td>c1-005 OD</td>
<td>−6</td>
<td>8</td>
<td>11.1</td>
<td>9</td>
<td>+3.33</td>
<td>−0.58</td>
<td>172</td>
<td>2.412</td>
<td>1.019</td>
<td>1.838</td>
<td>1.184</td>
</tr>
<tr>
<td>c2-001 OS</td>
<td>−6</td>
<td>6</td>
<td>9.1</td>
<td>9</td>
<td>+4.71</td>
<td>−0.61</td>
<td>12</td>
<td>0.541</td>
<td>0.197</td>
<td>0.333</td>
<td>0.388</td>
</tr>
<tr>
<td>c2-006 OS</td>
<td>−6</td>
<td>6</td>
<td>9.1</td>
<td>9</td>
<td>+2.17</td>
<td>−0.65</td>
<td>88</td>
<td>0.618</td>
<td>0.285</td>
<td>0.527</td>
<td>0.440</td>
</tr>
<tr>
<td>c5-005 OD</td>
<td>−10</td>
<td>6</td>
<td>9.1</td>
<td>9</td>
<td>+2.49</td>
<td>−0.28</td>
<td>29</td>
<td>2.182</td>
<td>0.414</td>
<td>2.108</td>
<td>0.381</td>
</tr>
<tr>
<td>c5-026 OD</td>
<td>−10</td>
<td>6</td>
<td>9.1</td>
<td>9</td>
<td>+4.11</td>
<td>−0.28</td>
<td>37</td>
<td>0.574</td>
<td>0.296</td>
<td>0.426</td>
<td>0.246</td>
</tr>
</tbody>
</table>

OZ, diameter of the programmed optical zone; TTZ, diameter of the total treatment zone; total HOA RMS, root mean square value of 3rd- to 10th-order aberrations; coma RMS, RMS value of 3rd- to 9th-order coma; SA RMS, RMS value of 4th- to 10th-order spherical aberration; rHOA RMS, residual RMS of all non coma, nonspherical HOA.

Conducted according to the guidelines of the University of Rochester Committee on Animal Research (UCAR), the ARVO Statement for the Use of Animals in Ophthalmic and Vision Research, and the NIH Guide for the Care and Use of Laboratory Animals.

Photorefractive Keratectomy

Three cats' eyes underwent PRK for −6 D, two with a programmed OZ of 6 mm and one with an 8 mm OZ; two eyes received a PRK for −10 D (6 mm OZ). The procedure has been described in detail elsewhere.

Briefly, all eyes received a conventional spherical ablation (Planoscan 4.14; Bausch & Lomb, Inc., Rochester, NY) after surgery, the cats received 2 drops of 0.3% tobramycin 0.1% dexamethasone 0.1% (Tobradex; Alcon, Fort Worth, TX) per eye, once a day, until the surface epithelium healed.

Wavefront Sensing

As described previously, the cats were trained to fixate single spots of light presented on a computer monitor. Wavefront measurements were performed before surgery and 19 ± 7 (12–24) weeks after surgery, with a custom-built Hartmann-S普遍 shock wavefront sensor. The wavefront sensor was aligned to the visual axis of one eye, while the other eye fixated a spot on the computer monitor.

Calculation of Centered WFE Differences

From each single drop array pattern, WFEs were calculated with a 2nd–10th-order Zernike polynomial expansion according to Vision Science and Its Applications (VISA) standards for reporting aberation data of the eye. WFE changes were simulated in a multistep process. The first step included the determination of the center of the OZ. Because PRK was performed with the cat under general anesthesia and the ablation was registered to the pupil center, an alignment to the visual axis of the cat's eye during surgery could not be ensured, and possible decentration effects had to be compensated for. Therefore, the centroiding area (analysis pupil) of 6-mm diameter was shifted manually in steps of 300 μm according to the distance between the lenslet centers to find the wavefront that yielded the most negative Z 2 0 value (i.e., the maximum treatment effect. This was then averaged from single measurements over a PD of 9 mm. In the second step, the horizontal and vertical offsets between the center of the OZ and the center of the original pupil were used to calculate preoperative WFEs (W pre(x, y)), for the position that equaled the later treatment center. Like W pre(x, y), W pre(x, y) was computed for a 9-mm PD. In a third step, the change in WFEs, ΔW(x, y), was obtained by subtracting the pre- from the postoperative Zernike coefficients. Thus, ΔW(x, y) reflected the treatment effect over a 9-mm PD, for a perfectly centered OZ, minimizing the potential influence of internal aberrations. The Zernike coefficient spectrum of each ΔW(x, y) (Table 1) was consistent with data obtained in humans after PRK.

Computer Modeling of Treatment Decentration

For each eye, decentration of a 6-mm subpupil relative to ΔW(x, y) was simulated by using custom software (MatLab 7.2; The MathWorks Inc., Natick, MA). Decentered WFE differences ΔW(x’, y’) were calculated for the size of the 6-mm subaperture along Cartesian decentrations Δx and Δy, where Δx and Δy were changed in steps of 100 μm, covering the entire 9-mm centroiding area and resulting in a maximum decetration range of 5000 μm over a circular region. Zernike polynomials for the 2nd to the 6th order were fitted to the data of each decentered wavefront ΔW(x’, y’) by using a singular value decomposition algorithm to calculate the pseudoinverse of the Zernike data to get the decentered subpupil Zernike coefficients. As a refinement of the manual determination of the centered position, the algorithm assigned the center coordinates (Δx = 0, Δy = 0) to the ΔW(x’, y’) with the lowest Z 2 0 value. For each eye, 709 WFEs, 1 centered and 708 decelerated were calculated over a 6-mm PD.

Simulation of Decentered Treatment Effects and VSOtF Calculation

Theoretical optical quality was investigated by calculating the VSOtF metric (visual Strehl ratio based on the optical transfer function [OTF]). The VSOtF is the ratio of the contrast sensitivity-weighted OTF to the contrast sensitivity-weighted OTF of the diffraction-limited eye.

Because the preoperative WFEs W pre(x, y) were decentered, calculating the VSOtF from preoperative HOA could lead to misinterpretation of optical quality due to over- or underestimation of HOA. Thus, we calculated a standard preoperative WFE, W pre(x, y), from all eyes included in this study. For the calculation of W pre(x, y), all preoperative, pupil-centered WFEs were averaged, resulting in a WFE representing the typical preoperative range of HOA (Table 2). Simulated postoperative WFEs, W post(x, y), were calculated by subtracting the W pre(x, y) from each ΔW(x’, y’). This treatment simulation relates to a standard preoperative WFE allowed us to eliminate interindividual differences in preoperative optical quality and internal optics. Therefore, the independent variables in this experiment were the five different centered treatment effects ΔW(x, y) and...
their corresponding $\Delta W(x', y')$. A computer program (Visual Optics Laboratory, VOL-Pro 7.14; Sarver and Associates, Carbondale, IL) was used to calculate the VSOTF over an analysis PD of 3.5 and 6.0 mm. The VSOTF for a given WFE was calculated for the combination of LOA terms that provided the highest VSOTF simulating the optical quality with best spherocylindrical correction (BCVSOTF). Thus, for each simulated $W_{\text{post}}(x', y')$, an LOA-derived refractive error based on 2nd-order terms and an “effective” refractive error based on the BCVSOTF were obtained. Differences between refractive errors were expressed as dioptric power vectors $(M, J0, J45)$, where $M$ corresponds to the spherical equivalent and $J0$ to the 0°/90° and $J45$ to the 45°/135° astigmatic components. The difference between the VSOTF- and 2nd-order-based power vectors could be considered a function of the interaction between HOA and LOA. Since “sphere” and “cylinder” are most commonly used in clinical settings, we displayed most of the results in terms of sphere and cylinder magnitude. To visualize decentration effects for single eyes, color maps plotting $\Delta LOA$, $\Delta HOA$, and $\Delta \log$ BCVSOTF against horizontal and vertical decentration were created. For further statistical analysis, data for decentration along the 0°, 90°, 180°, and 270° meridians were averaged for each eye.

Calculating Decentration Tolerance

Analysis of tolerance was performed by calculating the maximum permissible decentration that yielded a critical refraction or BCVSOTF difference. For sphere and cylinder, this threshold value was defined as $-0.5$ D. For the optical quality metric BCVSOTF, we chose a critical decrease of 0.2 log units, which roughly equals a decrease of 2 logMAR steps. For each parameter investigated, vectors $r$ between the centered position $(x, y)$ and each outmost coordinate below the criterion (threshold coordinates $x', y'$) were calculated. The mean value, $\bar{r}$, reflects the average maximum permissible decentration (in micrometers) that allows one to remain below the threshold criterion and equals the radius of a circle around the centered position. The standard deviation ($SD$) of $\bar{r}$ and the coefficient of variation ($CV$) of $\bar{r}$ served as metrics for regularity of decentration effects, where $SD$ of $\bar{r}$ reflects the absolute and $CV$ of $\bar{r}$ the relative irregularity. The smaller the $SD$ and $CV$, the less variable were the decentration effects along different meridians (i.e., the more circle-shaped was the decentration pattern).

### Statistical Analysis

All analyses were based on the difference values $\Delta W$ and $\Delta \log$ BCVSOTF, which reflected the treatment effects. Main outcome measures were the change of log BCVSOTF, the change of LOA, expressed in dioptries, and the change of HOA as a function of decentration. All differences for the center position $(x, y)$ were normalized to zero. Thus, values for decentered coordinates $x'$ and $y'$ reflect the deviation from the centered treatment effect. The difference between wavefront- and VSOTF-based refraction was considered an effect of interaction between LOA and HOA. Tolerance metrics were calculated as de-

<table>
<thead>
<tr>
<th>PD (mm)</th>
<th>Log BCVSOTF</th>
<th>Total HOA RMS ($\mu$m)</th>
<th>Coma RMS ($\mu$m)</th>
<th>SA RMS ($\mu$m)</th>
<th>rHOA RMS ($\mu$m)</th>
</tr>
</thead>
<tbody>
<tr>
<td>3.5</td>
<td>$-0.05$</td>
<td>0.036</td>
<td>0.031</td>
<td>0.012</td>
<td>0.014</td>
</tr>
<tr>
<td>6.0</td>
<td>$-0.38$</td>
<td>0.185</td>
<td>0.145</td>
<td>0.078</td>
<td>0.083</td>
</tr>
</tbody>
</table>

For all calculations LOA were set to zero. BCVSOTF (visual Strehl ratio based on the optical transfer function, simulated for best correction); total HOA RMS, root mean square value of 3rd- to 6th-order aberrations; coma RMS, RMS of 3rd to 5th order coma; SA RMS, RMS of $Z_4^0$ and $Z_6^0$; rHOA RMS, residual RMS of all noncoma, nonspherical HOA.

### Table 2. The Averaged Preoperative Mean WFE $W_{\text{measured}}(x', y')$, Computed for 3.5- and 6-mm PDs

**FIGURE 1.** Second-order refraction change decentration maps for the right eye of cat 5-005. (A) Change in sphere, 3.5-mm PD. (B) Change in cylinder magnitude, 3.5-mm PD. (C) Change in sphere, 6-mm PD. (D) Change in cylinder magnitude, 6-mm PD. The center (crosshair) is set to zero; dotted lines: 0.25-D steps.
scribed earlier. HOAs were broken down into coma root mean square (RMS) (the RMS of all coma terms $Z_6^{(1)}$), spherical aberration RMS (SA RMS, the RMS value of all coefficients $Z_6^{(0)}$), and the RMS of the residual noncoma, nonspherical aberrations (rHOA, the RMS value of all remaining HOA $Z_6^{(n)}$).

The influence of the magnitude of HOA induction on decentration tolerance was assessed with linear regression analysis. The dependent variables were the mean vectors $\bar{r}$ and their SD. To investigate the impact of HOAs on log BCVSOTF, we applied a multiple-regression model using HOAs as predictors and log BCVSOTF as dependent variables. The role of interaction on decentration tolerance was investigated by comparing $\bar{r}$ and SD for 2nd-order sphere and cylinder with their VSOTF-based equivalents using a nonparametric test for matched pairs (Wilcoxon test). The same test was also applied to compare decentration tolerance for PDs of 3.5 and 6.0 mm. All statistical tests were performed with a commercial program (SPSS 11.0; SPSS Inc., Chicago, IL), assuming a significance level of $P < 0.05$ and using the Bonferroni adjustment for multiple tests.

**RESULTS**

**Change in Second-Order Aberrations**

For all eyes examined, increasing decentration caused increasing undercorrection of 2nd-order sphere and induction of 2nd-order astigmatism. However, the pattern of decentration effects was triangular in shape rather than rotationally symmetric, as might have been predicted from the intended refractive correction (Fig. 1). These irregularities were more pronounced for 3.5-mm PDs which also showed reduction of cylinder magnitude with decentration (Figs. 1A, 1B). When averaging over all five eyes, decentrations of $\pm 1000$ μm had a limited effect on sphere and cylinder magnitude, since the average undercorrection and cylinder induction were $> -0.5$ D (Fig. 2). In contrast, decentrations $\pm 1000$ μm resulted in larger deviation from the central treatment effect. The mean induction of astigmatism was higher for 6- than for 3.5-mm PDs; however, the differences between the two PDs for decentrations $\pm 900$ μm reached only local significance of $P < 0.05$, which was nonsignificant with the Bonferroni correction.

**Decentration Effects and the Interaction between HOAs and LOAs**

VSOTF-based refraction data included interaction effects of LOA with HOA. Apart from a tendency of VSOTF-based $M$ values to be more hyperopic at the centered position, there were no significant differences between 2nd-order and VSOTF-based power vectors (Table 3). Decentration effects were more irregular for VSOTF-based refraction data than for the corresponding wavefront-derived data, particularly for sphere measured over 6-mm PDs (local $P < 0.05$; Table 5). The effects of decentration on the VSOTF cylinder magnitude also showed high interindividual variability among the eyes.

**TABLE 3.** Effects of Interaction between LOAs and HOAs on Treatment Effects as a Function of Decentration

<table>
<thead>
<tr>
<th>PD (mm)</th>
<th>Decentration (μm)</th>
<th>$M$</th>
<th>$J0$</th>
<th>$J45$</th>
</tr>
</thead>
<tbody>
<tr>
<td>3.5</td>
<td>0</td>
<td>0.05 ± 0.11</td>
<td>-0.05 ± 0.07</td>
<td>-0.02 ± 0.06</td>
</tr>
<tr>
<td></td>
<td>200</td>
<td>0.03 ± 0.26</td>
<td>-0.04 ± 0.09</td>
<td>-0.01 ± 0.07</td>
</tr>
<tr>
<td></td>
<td>500</td>
<td>0.01 ± 0.37</td>
<td>-0.02 ± 0.08</td>
<td>0.05 ± 0.09</td>
</tr>
<tr>
<td></td>
<td>1000</td>
<td>0.03 ± 0.39</td>
<td>-0.02 ± 0.13</td>
<td>0.04 ± 0.10</td>
</tr>
<tr>
<td></td>
<td>1500</td>
<td>0.13 ± 0.43</td>
<td>-0.02 ± 0.11</td>
<td>0.02 ± 0.05</td>
</tr>
<tr>
<td>6.0</td>
<td>0</td>
<td>0.68 ± 0.31</td>
<td>-0.18 ± 0.14</td>
<td>0.05 ± 0.06</td>
</tr>
<tr>
<td></td>
<td>200</td>
<td>0.61 ± 0.45</td>
<td>-0.16 ± 0.17</td>
<td>0.04 ± 0.08</td>
</tr>
<tr>
<td></td>
<td>500</td>
<td>0.63 ± 0.28</td>
<td>-0.10 ± 0.25</td>
<td>0.00 ± 0.09</td>
</tr>
<tr>
<td></td>
<td>1000</td>
<td>0.58 ± 0.53</td>
<td>-0.06 ± 0.13</td>
<td>-0.01 ± 0.12</td>
</tr>
<tr>
<td></td>
<td>1500</td>
<td>0.56 ± 0.65</td>
<td>-0.16 ± 0.37</td>
<td>0.01 ± 0.15</td>
</tr>
</tbody>
</table>

The data are averaged from the 0°, 90°, 180°, and 270° meridian and expressed as mean and SD of the difference between second-order and VSOTF-based dioptic power vectors $M$, $J0$, and $J45$. Differences were not statistically significant. VSOTF refraction, simulated endpoint of the subjective refraction based on the BCVSOTF; $M$, spherical equivalent; $J0$, 0°/90° astigmatic component; $J45$, 45°/135° astigmatic component.
Table 4. Change of VSOTF and Induction of Higher-Order Aberrations as a Function of Decentration

<table>
<thead>
<tr>
<th>PD (mm)</th>
<th>Decentration (µm)</th>
<th>Δlog BCVSOTF</th>
<th>Δ Coma RMS</th>
<th>Δ SA RMS</th>
<th>Δ rHOA RMS</th>
</tr>
</thead>
<tbody>
<tr>
<td>3.5</td>
<td>0</td>
<td>0 ± 0</td>
<td>0 ± 0</td>
<td>0 ± 0</td>
<td>0 ± 0</td>
</tr>
<tr>
<td></td>
<td>200</td>
<td>-0.04 ± 0.04</td>
<td>0.004 ± 0.041</td>
<td>-0.001 ± 0.015</td>
<td>-0.001 ± 0.031</td>
</tr>
<tr>
<td></td>
<td>500</td>
<td>-0.03 ± 0.09</td>
<td>0.059 ± 0.069</td>
<td>0.011 ± 0.025</td>
<td>0.002 ± 0.044</td>
</tr>
<tr>
<td></td>
<td>1000</td>
<td>-0.12 ± 0.18</td>
<td>0.112 ± 0.079</td>
<td>0.015 ± 0.034</td>
<td>0.003 ± 0.058</td>
</tr>
<tr>
<td>6.0</td>
<td>0</td>
<td>-0.14 ± 0.19</td>
<td>0.058 ± 0.039</td>
<td>0.270 ± 0.149</td>
<td>0.186 ± 0.042</td>
</tr>
<tr>
<td></td>
<td>200</td>
<td>-0.18 ± 0.17</td>
<td>0.111 ± 0.080</td>
<td>0.275 ± 0.138</td>
<td>0.187 ± 0.041</td>
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<tr>
<td></td>
<td>500</td>
<td>-0.24 ± 0.18</td>
<td>0.500 ± 0.147</td>
<td>0.295 ± 0.147</td>
<td>0.195 ± 0.047</td>
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<tr>
<td></td>
<td>1000</td>
<td>-0.29 ± 0.20</td>
<td>0.660 ± 0.289</td>
<td>0.358 ± 0.184</td>
<td>0.223 ± 0.048</td>
</tr>
<tr>
<td></td>
<td>1500</td>
<td>-0.29 ± 0.20</td>
<td>0.932 ± 0.403</td>
<td>0.317 ± 0.188</td>
<td>0.252 ± 0.042</td>
</tr>
</tbody>
</table>

The data are averaged from the 0°, 90°, 180°, and 270° meridian and expressed as mean and standard deviations. The values are normalized to the values for the centered position for a 3.5 mm pupil diameter (PD), i.e. each value reflects the difference to the value obtained from centered position over a 3.5 mm PD. Δx, horizontal decentration; Δy, vertical decentration. log BCVSOTF, visual Strehl ratio based on the optical transfer function, simulated for best correction; coma RMS, RMS of 3rd and 5th order coma terms; SA RMS, RMS of Z₄ and Z₆; rHOA RMS, residual RMS of all noncoma, nonspherical HOA.

Changes of HOAs and BCVSOTF

HOAs induced by decentration were dominated by coma (Table 4). As for 2nd-order aberrations, the decentration patterns for coma RMS were not rotationally symmetric, and displayed flatter slopes but higher irregularity for 3.5- than for 6-mm PDs (Fig. 3). We found a significant influence of the amount of decentration on the induction of coma RMS at a PD of 6 mm (adjusted $R^2 = 0.51$; B = $0.7 \times 10^{-3}$, $P < 0.001$). At 3.5 mm, although much less pronounced (adjusted $R^2 = 0.23$, B = $0.08 \times 10^{-3}$, $P < 0.001$), the same tendency was observed (Fig. 4). The induction of SA RMS and rHOA RMS was less influenced by decentration (no significant correlation), with irregular decentration patterns and high variability between individual eyes at the two PDs.

In all eyes, theoretical best-corrected optical quality expressed as BCVSOTF decreased by $-0.41 \pm 0.13$ log units for 3.5-mm pupils and by $-0.55 \pm 0.19$ log units for 6.0 mm pupils after a centered treatment. Decentration resulted in even higher decrease in log BCVSOTF (Figs. 5, 6). Furthermore, if Δlog BCVSOTF was computed for a 3.5-mm PD, the position that yielded the minimum decrease of BCVSOTF was located paracentrally in all eyes (Fig. 5A). The regression model revealed a significant influence of the HOAs on log BCVSOTF at both PDs (adjusted $R^2 = 0.84$ for 6-mm PD, $R^2 = 0.81$ for 3.5-mm PD) with the highest impact of coma RMS in both models.

Analysis of Decentration Tolerance and Irregularity

Table 5 shows the mean vectors $\bar{r}$ and their SDs. Both for wavefront-derived and for VSOTF-based sphere, the critical $\bar{r}$ for an undercorrection of 0.5 D was greater than 1000 µm in all cases. The mean change of decentration tolerance due to interaction was $82 \pm 232$ µm for 6-mm PD and $-92 \pm 73$ µm for 3.5-mm PD (both $P < 0.05$). For the 6-mm PD, $\bar{r}$ of cylinder magnitude decreased by $-160 \pm 42$ µm when interaction was simulated ($P > 0.05$). At the 3.5-mm PD, values remained almost constant ($14 \pm 153$ µm; $P > 0.05$). While the $\bar{r}$ of sphere and cylinder was similar at the two PDs, the data (Figs. 5, 6) suggested a higher decentration tolerance at the 3.5-mm PD with regard to log BCVSOTF. This was confirmed by analysis of $\bar{r}$ (Table 5; local $P < 0.05$). Analysis of the SD and CV of $\bar{r}$ showed that the 2nd-order sphere (6.0-mm PD) had more regular decentration patterns than did the other parameters (Table 5).

Linear regression analysis revealed that decentration tolerance $\bar{r}$ was influenced significantly by spherical aberrations induced by the centered treatment. At 6-mm PD, 2nd-order sphere ($R^2 = 0.87$, B = $-181$; $P < 0.05$), 2nd-order cylinder (adjusted $R^2 = 0.80$, B = $-278$; $P < 0.05$), and the VSOTF sphere ($R^2 = 0.80$, B = $-407$; $P < 0.05$) were significantly influenced by ΔSA RMS but not by the amount of defocus or coma and rHOA RMS changes. Likewise, sphere and cylinder obtained over a 3.5-mm PD appeared not to be influenced by defocus change or HOA induction of the treatment. Steeper
FIGURE 4. Mean effects of decentration on the change of coma RMS (averaged data from 0°, 90°, 180°, and 270° meridians for the five eyes). The center is set to zero.

decentration-induced LOAs

As observed by others,1,2,14,51 there was undercorrection of the spherical refractive error and induction of astigmatism as a function of decentration in all eyes examined. However, to our knowledge, the present study is the first decentration model study that is based on real wavefront data. Because model studies in the literature have always assumed the Munnerlyn algorithm1,2 or a perfect wavefront-guided ablation,3,4,19 they probably underestimated the effects of HOA induced by the primary treatment.21,32 Unlike spherical aberrations that dominated ∆W(x, y), the amount of coma RMS and rHOA RMS induced by the treatment did not significantly influence decentration tolerance of sphere and cylinder. With calculation of a simulated endpoint of the subjective refraction based on the metric VSOTF, an investigation of interaction effects between LOA and HOA was possible. In particular, we asked whether induced HOA affected the endpoint of the subjective refraction and caused “residual refractive error.” All eyes showed a tendency toward hyperopic VSOTF sphere values over a 6-mm PD which could be explained by interaction with spherical aberration.53 Furthermore, interindividual variability of interaction effects increased with decentration (Table 3, higher SDs for larger decentrations). Although cautious because of our small sample size, we believe that contrary to its effects on LOA induction, decentration did not consistently affect LOA/HOA interactions. However, VSOTF-based refraction results27 may differ from subjective refraction, particularly with HOA-related image distortion. Our finding that only decentrations ≥1000 μm caused spherical and cylindrical undercorrection ≥0.5 D and larger suggests that ubiquitous microdecentrations7,9,10 ≤500 μm are not a significant source of postoperative residual refractive errors.

HOA and Best-Corrected Optical Quality

Induction of HOAs, especially coma, is a key contributor to symptoms after LRS in humans.1,12–14,17,34,35 We noted that the induction of HOAs by decentration occurred in an irregular pattern that may have resulted from treatment-induced, non-rotationally symmetric aberrations. There was also a large difference in the induction of coma at 3.5- and 6-mm PDs. Although on average, the amount of spherical aberrations decrease of ∆log BCVSOTF with decentration was also associated with higher amounts of SA RMS induction by the treatment (Fig. 7), but this association did not reach statistical significance. In this series of eyes, we could not establish any correlation between the induced defocus or HOA and the irregularity index SD of r̄, neither for sphere and cylinder, nor for log BCVSOTF.

DISCUSSION

Decentration Effects Followed an Irregular Pattern

The present experiments revealed that decentration effects were distributed asymmetrically, although the treatment involved only rotationally symmetric ablation patterns. This behavior affected all parameters investigated and was more pronounced at the smaller (3.5-mm) pupil size. The fact that only decentrations that may have resulted from treatment-induced, non-rotationally symmetric ablation patterns. This behavior affected all parameters investigated and was more pronounced at the smaller (3.5-mm) pupil size. The fact that only decentrations...
induced was not affected by decentration, the SDs increased with decentration, reflecting high interindividual differences. In all eyes examined, log BCVSOTF decreased asymmetrically as a function of decentration, displaying asymmetric decentration patterns. The obvious relationship between coma and log BCVSOTF in the decentration maps (Figs. 3, 4) was confirmed by regression analysis that revealed a highly significant, numerical impact of coma on log BCVSOTF at both 3.5- and 6-mm PDs. The large discrepancy between decentration tolerance at 3.5- and 6-mm PDs suggests that microdecentrations could be one cause of night vision disturbances in eyes that are asymptomatic under photopic conditions, particularly if center shifts between constricted and dilated pupil are involved. Nevertheless, the computational model described in the present study is a powerful and versatile tool for the analysis of decentration effects on refractive outcome. Although based on a small sample of experimental eyes, it allowed us to reach conclusions of potential interest for refractive surgical practice: (1) Decentration of myopic treatments leads to consistent undercorrection of the defocus term and the induction of astigmatism. However, the decentration tolerance of sphere and cylinder (including simulated interaction effects) makes it unlikely that microdecentrations ≤500 μm are a significant cause of residual refractive errors in otherwise asymptomatic eyes.

**CONCLUSIONS**

Anatomy, optics, function, and subjective perception are key levels in the concept of quality of vision after refractive surgical procedures. The model described herein allowed us to investigate decentration tolerance as a novel dimension of the "optics" level in the quality of vision concept. Our calculations reduced possible biases resulting from aberrometer misalignments or internal optics so that "pure" WFE changes could be investigated. Although these computations are laborious, an evaluation of decentration effects on novel treatment modalities (e.g., presbyopia-correcting laser profiles or new multifocal intraocular lenses) is now possible. As demonstrated in the context of image quality, it appears logical that different aberrations should interact, affecting decentration tolerance. A limitation of our computational model, however, is that it simulates decentration by pupil shifts rather than by shifts of the treatment zone. Given that some of our treatments were centered themselves, this could be a problem, especially if different portions of the central cornea yield significantly different biological responses.

Nevertheless, the computational model described in the present study is a powerful and versatile tool for the analysis of decentration effects on refractive outcome. Although based on a small sample of experimental eyes, it allowed us to reach conclusions of potential interest for refractive surgical practice: (1) Decentration of myopic treatments leads to consistent undercorrection of the defocus term and the induction of astigmatism. However, the decentration tolerance of sphere and cylinder (including simulated interaction effects) makes it unlikely that microdecentrations ≤500 μm are a significant cause of residual refractive errors in otherwise asymptomatic eyes.

**TABLE 5. Analysis of Decentration Tolerance (Maximum Permissible Decentration to Maintain a Threshold Value)**

<table>
<thead>
<tr>
<th>Parameter</th>
<th>Threshold Value</th>
<th>3.5-mm PD</th>
<th>6-mm PD</th>
<th>3.5-mm PD</th>
<th>6-mm PD</th>
</tr>
</thead>
<tbody>
<tr>
<td>Δ2nd-order sphere</td>
<td>−0.5 D</td>
<td>1255 ± 160</td>
<td>1313 ± 136</td>
<td>228 ± 60</td>
<td>111 ± 47</td>
</tr>
<tr>
<td>Δ2nd-order cylinder</td>
<td>−0.5 D</td>
<td>1304 ± 150</td>
<td>1008 ± 214</td>
<td>208 ± 64</td>
<td>173 ± 102</td>
</tr>
<tr>
<td>ΔVSOTF sphere</td>
<td>−0.5 D</td>
<td>1348 ± 104</td>
<td>1232 ± 314</td>
<td>204 ± 63</td>
<td>246 ± 55</td>
</tr>
<tr>
<td>ΔVSOTF cylinder</td>
<td>−0.5 D</td>
<td>1289 ± 201</td>
<td>1167 ± 271</td>
<td>226 ± 100</td>
<td>207 ± 63</td>
</tr>
<tr>
<td>Δlog BCVSOTF</td>
<td>−0.2</td>
<td>1219 ± 210</td>
<td>800 ± 512</td>
<td>248 ± 82</td>
<td>143 ± 40</td>
</tr>
</tbody>
</table>

The radius $r$ is the mean length of the vectors between the center and the locations with threshold values. The SD and CV of $r$ reflect the irregularity of the decentration behavior. All data are expressed as the mean and SD. PD, analysis pupil diameter. VSOTF sphere/cylinder, simulated endpoint of the subjective refraction based on the BCVSOTF.
eyes. (2) In contrast to effects on LOAs, microdecenterations appear to be a source of HOA-related visual symptoms under mesopic conditions in a proportion of eyes that would be asymptomatic under photopic conditions. (3) Given the intra- and interindividual variability of effects in our model, it appears that only some eyes will experience symptoms in clinical practice. (4) Finally, our results suggest that minimizing the induction of spherical aberration by maximizing the functional optical zone of the cornea40 using aspheric ablation profiles41,42 or large OZ diameters43 could significantly increase decentration tolerance and by doing so, optimize refractive outcome.

References


