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Translation model for anterior segment tomographic data to corneal spherical aberration derived from a Monte-Carlo simulation based on raytracing

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ABSTRACT.

Background: Intraocular lenses with a negative aspherical design for correction of corneal spherical aberration (SA) have gained popularity in recent decades. In most cases, a 'one size fits all' concept is followed, where all eyes receive lenses with the same SA correction. The purpose of this study is to develop a strategy based on raytracing using anterior segment tomography data to extract corneal SA and to provide simple multivariable linear models for prediction of corneal SA.

Methods: The analysis was based on a large dataset of 8737 measurements of 8737 eyes from 1 clinical centre, using the Casia2 anterior segment tomographer. An optical model based on: corneal front and back surface radius Ra and Rp, asphericities Qa and Qp, corneal thickness CCT, anterior chamber depth ACD, and pupil centre position (X-Y position: PupX and PupY), was defined for each measurement. Corneal SA was derived using a 6-mm aperture perpendicular to the incident ray and centred on the chief ray, and linear prediction models were derived for SA using biometric data. Cross-validation was used for model performance evaluation.

Results: Using raytracing, the wavefront error within an aperture (6-mm diameter centred on the intersection of the chief ray with the cornea) was calculated and corneal SA was extracted. After identifying the relevant effect sizes (Ra, Qa, Rp, Qp, ACD, PupX, and PupY) using stepwise linear regression, linear mixed-effects models (model 1: all effect sizes, model 2: Ra, Qa, Rp, and Qp, model 3: Ra and Qa) were set up on the training data in terms of a Monte-Carlo simulation. On the test data (training data), model 1 with a mean absolute/root-mean-squared prediction error of 0.0095/0.0130 (0.0095/0.0127) performed similarly to model 2 with 0.0097/0.0131 (0.0096/0.0127), and both outperformed model 3 with 0.0152/0.0197 (0.0148/0.0190).

Conclusion: Based on the Casia2 anterior segment tomographer, corneal SA could be derived using shape data (curvature and asphericities) of both corneal surfaces (model 2). This information could easily be used for selection of the appropriate negative aspherical lens design in cataract surgery.

Key words: cataract surgery – cornea asphericity – linear mixed-effects model – Monte-Carlo simulation – raytracing – spherical aberration

Background

Over the last 2 decades aspherical intraocular lenses have gained popularity in cataract surgery as replacements for the opaque crystalline lens. Many different optical designs have been brought to market: in addition to spherical designs with spherical front and back lens surfaces, we have the category of aberration-free or aberration-neutral lenses that are intended not to change overall ocular spherical aberration (SA) (the SA of the pseudophakic eye equals the SA of the aphakic eye). In addition, we have the category of aberration correcting lenses that are intended to reduce or eliminate the SA of the eye by compensating for the SA of the cornea (which shows typically positive values) with a negative SA in the lens (Kohnen & Klaproth 2008; Wang et al. 2012).

The term ‘aberration-free’ or ‘aberration-neutral’ in the context of cataract surgery is not clearly defined in the literature: Whereas some lens designs neutralize the intrinsic SA for a collimated entrance beam and show positive SA for a convergent incident ray bundle (e.g. located behind a (model) cornea), others neutralize the intrinsic SA for a convergent entrance beam and show negative SA for a...
collimated incident ray bundle (Langenbucher et al. 2017). In contrast, aberration correcting lenses always induce some amount of negative SA if located behind a (model) cornea (with an incident convergent beam). Depending on the correction concept, which is mostly based on a schematic model eye, and the amount of SA to be corrected, there are lenses on the market with an SA correction in a range between 0.10 μm and 0.27 μm.

Unlike ocular spherical aberration, corneal spherical aberration cannot be measured directly with any device (Sicam et al. 2006; Calossi 2007; Nochez et al. 2010; Al-Sayyari et al. 2014; De Jong et al. 2016). In place of measurement, corneal SA can be calculated from corneal tomography data, which requires raytracing strategies (Langenbucher et al. 2021). In most tomographers, a conversion from corneal asphericity to corneal spherical aberration is implemented, but the methodology of the conversion, the reference plane, the radius and the centre of the unit circle for the Zernike decomposition, as well as the reference axis are not known/disclosed (Wang & Bao 2017). The amount of SA correction typically mentioned in the data specification sheet refers to the SA correction at corneal front surface plane referenced to a 6-mm zone. This means that the value provided by the lens manufacturer in terms of a Zernike coefficient (given in a μm scale) is related to a specific environment characterized by a specific location of the lens relative to the cornea and pupil position, where the pupil centre together with the angle of the incident ray bundle defines the visual axis. As this environment is not standardized, lens manufacturers use different lens models for specifying the lens SA, and therefore, the ‘real’ amount of SA correction in an individual eye may vary (Schrecker et al. 2019).

In current clinical practice, the selection of lens type (spherical or aspherical) according to the corneal SA of the individual eye is rarely considered. (Solomon 2010; Du et al. 2019; Schrecker et al. 2019). Wavefront aberrometers measure the optical aberrations of the entire eye (Atchison et al. 2016), but the result of ocular aberration measurement prior to cataract surgery is not relevant for lens selection, as the crystalline lens with its intrinsic optical aberrations is removed from the optical system. However, in the absence of corneal pathologies, with the new generation of anterior segment tomographers, many clinically relevant parameters, including the curvature of the corneal front and back surface and asphericity, and the central corneal thickness can be measured quite precisely. In cases of corneal pathologies such as tear film insufficiency, ectatic diseases such as keratoconus, keratoglobus or pellucid marginal degeneration, or with fixation instabilities, the variability of the measurement parameters may however increase significantly.

A Cartesian coordinate system is typically defined with its origin at the corneal front apex with axial direction Z towards the retina and lateral directions X (horizontal axis, positive values to the right) and Y (vertical direction, positive values up). In addition, there are measurement data for the axial and lateral position of the pupil centre and pupil diameter. From these data, the corneal SA value for a 6-mm zone at corneal front apex plane can be derived using raytracing techniques. With this value and the general definition of the amount of SA correction value from the lens data sheet, clinicians can select the appropriate lens model to compensate for the individual corneal SA (Schrecker et al. 2019).

The purpose of this study is (i) to provide a calculation strategy for extracting the corneal SA value referenced to a 6-mm zone at the corneal apex plane from measurement data of a clinically established high-resolution anterior segment OCT in a very large dataset based on raytracing techniques, (ii) to identify the relevant effect sizes and (iii) to model the results with a linear mixed-effects model in terms of a Monte-Carlo simulation. The resulting spherical aberration value from the model could be directly used together with the nominal SA correction from the lens data sheet to estimate the spherical aberration of the pseudophakic eye after cataract surgery.

Methods

Dataset for the Monte-Carlo simulation

In total, a dataset with 11 277 measurements from one clinical centre (Augenklinik Castrop-Rauxel, Germany) taken using the Casia2 anterior segment tomographer (Tomy, Nagoya, Japan) was considered for this retrospective study. Duplicate measurements of eyes and incomplete data in the dataset were excluded. Measurements from pseudophakic eyes or in mydriasis and data indexed with situation after refractive surgery, ectatic corneal diseases or other corneal pathologies were excluded from the dataset. The dataset was transferred to a .csv data table using the data export module of the Casia2 software. Data tables were reduced to the relevant parameters required for our raytracing and analysis, consisting of patient ID, laterality (left or right eye), central corneal curvature of the corneal front (Rf) and back (Rb) surface in mm, asphericity of the corneal front (Qf) and back (Qb) surface, central corneal thickness (CCT in mm), anterior chamber depth (ACD) measured from the corneal front apex to the lens front apex in mm, pupil size (Pup) in mm, as well as the location of the pupil centre (PupX and PupY in horizontal and PupZ in vertical direction), both in mm. The data were transferred to Matlab (Matlab 2019b, MathWorks, Natick, USA) for further processing. The study was registered with the local Ethics committee (Arztetekammer des Saarlandes, 157/21).

Preprocessing of the data and raytracing

Custom software was written in Matlab. By convention we defined a Cartesian coordinate system with its origin at the corneal front apex, X axis to the right, Y axis in the superior direction, and Z axis towards the retina. The optical model is represented by a structure consisting of a cornea with 2 coaxial aspheric surfaces defined by Rf, Qf, Rp, Qp and CCT. Accordingly, the coordinates of the elements in the optical model were X/Y/Z = 0/0/0 for the corneal front apex, 0/0/CCT for the corneal back apex and PupX/PupY/PupZ for the pupil centre (de-centration PupX in horizontal and PupY in vertical direction) respectively. For air/cornea/aqueous humour, we used refractive index values of 1.0/1.337/1.336, respectively, taken from the Liou-Brennan schematic model eye (Liou & Brennan 1997). Without loss of generality, in order to consider all samples as left eyes, the optical model was flipped horizontally for right eyes, which means that the sign for PupX
was changed keeping PupY unchanged. Positive values for X and Y refer to the temporal and superior direction.

As the incident ray angle with components Ix/Iy cannot be measured and therefore we do not have normative values, we used for the mean value the data from the Liou-Brennan schematic model eye (mean of Ix = 5° from nasally, mean of Iy = 0°). For right eyes, the sign of the Ix component was flipped. Additionally, we added a normal distribution with a standard deviation of 2° and finally discarded values outside the range [−9° to −1°/−4° to 4°] for Ix/Iy. Using this strategy, we considered the eccentric location of the fovea shifted in the temporal direction (positive values of X).

The entire dataset was split into training data (70%) and test data (30%) for a hold-out cross-validation strategy. The training data were used for extracting the relevant effect sizes and building up the model, and the performance of the prediction model was subsequently evaluated with the test data.

**Calculating the SA**

For initialization, a parametric ray with an incident ray angle Ix/Iy and variable offset in X and Y direction was projected onto the corneal front apex and traced through both corneal surfaces to the pupillary plane. The distance of the ray–pupil intersection was extracted, and using an iterative nonlinear optimisation strategy (Interior Point Method (IPM), Karmarkar 1984), the ray was shifted in X and Y to pass through the pupil centre (chief ray). The coordinates of the intersection of the chief ray with the corneal front surface plane were calculated as X0/Y0. In the next step, a circular aperture with a diameter of 6 mm was defined centred on X0/Y0 and perpendicular to the incident-collimated ray. This new aperture is tilted by Ix/Iy to the optical axis (Z direction of the Cartesian coordinate system) in order to be aligned to the incident ray (or the fixation axis of the patient). After removing the initial pupil from the optical system (located with its centre at X/Y/Z = ACD, Pupx, Pupy) a bundle of collimated rays (20 000 equidistant rays) with a diameter of 7 mm was traced through the optical system consisting of an aperture, the corneal front surface and the corneal back surface, and the best wavefront focus was identified by minimizing the root-mean-squared wavefront error. The optical path lengths were calculated from the source to the best wavefront focus. For all rays passing through the 6-mm aperture at X0/Y0, the optical path length differences were derived with respect to the aperture plane. For that purpose, all coordinates of the ray intersections with a plane Z = 0 were calculated, and the X/Y/Z coordinates were back-rotated by Euler angles −Ix/Iy to extract the wavefront with respect to the plane perpendicular to the collimated incident ray. This rotation was performed using quaternion transformation (St. Pierre 2021). Finally, a Zernike decomposition of all optical path length differences within the aperture (3-mm radius of the unit circle) was performed, and the Zernike coefficient Z0 0 was extracted representing the spherical aberration of the cornea independent from the pupil size measured with the Casia2. Figure 1 shows an example for 10 000 rays and model data derived from the Liou-Brennan schematic model eye (Ra, Qa, Rp, Qp, CCT, ACD, Pupx, Pupy/IX/IY = 7.77 mm/−0.18/6.4 mm/−0.6/0.5 mm/3.66 mm/−0.5 mm/0.0 mm/−5°/0°) the characteristics of the wavefront error (optical path length differences) in a plane perpendicular to the ray bundle together with the unit circle (6 mm in diameter, red circle) on which the Zernike coefficients were based. The resulting SA value was 0.27 μm.

**Identifying the relevant effect sizes for the multivariable regression model**

A stepwise linear regression (Draper & Smith 1998) was implemented to analyse the relevant effect sizes for a multilinear prediction model for the target parameter SA from the potential input parameters Ra, Qa, Rp, Qp, CCT, ACD, Pupx, Pupy, IX and Iy. This stepwise algorithm was applied to the training data, with the test data excluded. The stepwise strategy begins with an initial constant model and takes forward and backward steps to add or remove variables, until a stopping criterion is satisfied. As stopping criteria, we restricted the number of iterations to a maximum of 100, iteration steps smaller than 10e-9 or improvement of the root-mean-squared prediction error by less than 10e-12. The tolerance for adding terms was a significance value less than 0.05, and the tolerance for removing terms was a significance value larger equal 0.05.

**Setup of a multivariable linear prediction model**

With the effect sizes identified from this stepwise fit, a linear mixed-effects model (Herber et al. 2020) based on the training data was set up to predict the corneal SA. The biometric parameters were used as fixed effects, whereas the patient ID was considered as a grouping variable (random effects) in order to consider the effect of correlation between both eyes of an individual (Herber et al. 2020). In addition, a simplified linear mixed-effects model restricted to corneal front surface measurement data available from corneal topography was defined. These prediction models were applied to the test data for cross-validation in order to obtain a readout of the performance characteristics.

**Statistical analysis**

For exploratory data analysis of the input and output (SA) data, we provided the mean (MEAN) and standard deviation (SD), median value (MEDIAN), as well as 90% confidence interval (lower boundary: 5% quantile and upper boundary: 95% quantile), and 99% confidence interval (lower boundary: 0.5% quantile and upper boundary: 99.5% quantile). The performance of the prediction models derived from the training data were analysed using the test data with scatterplots displaying the modelled SA data versus the observed SA data from raytracing. The model prediction error PE was considered as the deviation of the multivariable linear model output minus the observed SA from raytracing. The performance of the model was expressed using the correlation coefficient R, significance level P (comparing the actual model against a constant model), the lower and upper boundaries of the 95% confidence interval for R (RLO and RUP), the mean prediction error (meanPE), the mean absolute prediction error (meanABSPE) and the root-mean-squared prediction error (rmsPE).
Fig. 1. Example of a wavefront (WF) error (optical pathlength differences) extracted from raytracing through model data derived from the Liou-Brennan schematic model eye. For this simulation with 10 000 equidistant rays we used (Ra, Qa, Rp, Qp, CCT, ACD, Pupx, Pupy, IX, IY = 7.77 mm/−0.18/6.4 mm/−0.6/0.5 mm/3.66 mm/−0.5 mm/0.0 mm/−5°/0°). After identifying the chief ray passing through the pupil centre, the pupil was removed and a new aperture with a 6-mm diameter was defined perpendicular to the incident ray and centred on the intersection of the chief ray with the corneal front surface. The reference unit circle for the Zernike decomposition with a 6-mm diameter is shown as a red line.

value took 24 733 seconds (6 hours 52 min) on a standard office PC. Table 1 shows the explorative data for Ra, Qa, Rp, Qp, CCT, ACD, Pup (not used for calculating corneal SA), the pupil centre position Pux and Pupy derived from the Casia2 anterior segment OCT in horizontal and vertical direction, the incident ray angles IX and IY in horizontal and vertical direction, as well as the corneal spherical aberration SA (Z4°). The wavefront error was considered within a 6-mm zone at a plane perpendicular to the incident-collimated beam centred on the intersection of the chief ray with the corneal front surface. The horizontal position of the pupil centre Pux and the horizontal incident ray angle IX were flipped in sign for right eyes to simulate left eyes only, meaning that positive values for X and Y refer to the temporal and superior directions respectively. The dataset was split randomly into training data (N = 6116, 70%) and test data (N = 2621, 30%).

Figure 2 shows the scatterplot matrix for the input parameters Ra, Qa, Rp, Qp, CCT, ACD, Pupx, Pupy, IX, IY, and the output parameter SA, separately for left eyes (in green) and right eyes (in red) before flipping the sign of horizontal pupil position and horizontal incident ray angle of left eye data. The respective histograms are plotted on the diagonal of the matrix. From the scatterplot matrix, it can be seen that left eyes and right eyes show similar behaviour in Ra, Qa, Rp, Qp, CCT, ACD, Pupx, IY and SA. The distributions of Pupx and IX are mostly symmetrical about the Y axis (flipped in sign). The most important effect size for corneal SA appears to be the asphericity value of the corneal front surface Qa.

Figure 3 displays the cumulative density function (CDF) of corneal SA extracted from the wavefront error considered in a 6-mm zone (unit circle) at a plane perpendicular to the incident rays. In the plot, we have added the benchmarks of the 90% and 99% confidence intervals shown as red and blue lines. From the graph, it can be seen that the SA value shows a mostly symmetrical distribution.

To investigate the relevant effect sizes, we used the following parameters as potential input values for our stepwise linear regression: Ra, Qa, Rp, Qp, CCT, ACD, Pupx and Pupy. In our final optical model for extracting corneal SA, we used a circular aperture with a 6-mm diameter centred on the intersection of the chief ray (passing through the pupil centre) with the corneal front surface oriented perpendicularly to the incident ray. Therefore, the pupil size measured with the Casia2 (Pup) was not considered for our model. As the incident ray angle was synthesized using a random generator, we decided to discard the parameters IX and IY from the modelling. The stepwise fit algorithm that qualifies the input parameters for our linear multivariable prediction model based on the training data showed convergence after 7 iterations. As relevant effect sizes, we identified the parameters Ra, Qa, Rp, Qp, ACD, Pupx and Pupy.

Model 1: For the parameters identified as relevant effect sizes, we set up a linear mixed-effects model based on the training data considering the patient ID as a grouping variable which reads:

\[
S.A = 0.9262 - 0.0935 \times R_d + 0.7154 \times Q_d + 0.0223 \times R_p - 0.1219 \times Q_p - 0.0026 \times ACD + 0.0062 \times Pup_x - 0.002 \times Pup_y
\]

The respective standard errors P values for the intercept and the 7 regression coefficients Ra, Qa, Rp, Qp, ACD, Pupx, Pupy mentioned above are 0.0048/<1e-100, 0.0008/<1e-100, 0.0013/<1e-100, 0.0010/<1e-100, 0.0014/<1e-100,
Table 1. Explorative data extracted from the dataset of the Casia2 anterior segment tomograph. Ra, Qa, Rp, Qp, CCT, ACD, Pup, PupX, PupY, IX, IY refer to the corneal front surface curvature and asphericity, corneal back surface curvature and asphericity, central corneal thickness, anterior chamber depth measured from the corneal front apex, pupil diameter, pupil lateral position of the pupil centre in X (positive values in the temporal direction) and Y (positive values in the superior direction), and simulated incident ray angle in X and Y. Right eyes were flipped in X. MEAN, SD, MEDIAN refer to mean value, standard deviation and median, and 5% quantile/95% quantile and 0.5% quantile/99.5% quantile to the lower and upper boundaries of the 90% and 99% confidence intervals respectively.

<table>
<thead>
<tr>
<th>N = 8737</th>
<th>Ra in mm</th>
<th>Qa</th>
<th>Rp in mm</th>
<th>Qp</th>
<th>CCT in mm</th>
<th>ACD in mm</th>
<th>Pup in mm</th>
<th>PupX in mm</th>
<th>PupY in mm</th>
<th>IX in °</th>
<th>IY in °</th>
<th>SA</th>
</tr>
</thead>
<tbody>
<tr>
<td>MEAN</td>
<td>7.76</td>
<td>−0.23</td>
<td>6.56</td>
<td>−0.11</td>
<td>0.55</td>
<td>3.36</td>
<td>3.32</td>
<td>−0.31</td>
<td>−0.10</td>
<td>−5.02</td>
<td>0.01</td>
<td>0.19</td>
</tr>
<tr>
<td>SD</td>
<td>0.28</td>
<td>0.13</td>
<td>0.25</td>
<td>0.11</td>
<td>0.04</td>
<td>0.40</td>
<td>0.81</td>
<td>0.21</td>
<td>0.18</td>
<td>1.73</td>
<td>1.46</td>
<td>0.09</td>
</tr>
<tr>
<td>MEDIAN</td>
<td>7.74</td>
<td>−0.22</td>
<td>6.56</td>
<td>−0.10</td>
<td>0.55</td>
<td>3.37</td>
<td>3.31</td>
<td>−0.30</td>
<td>−0.10</td>
<td>−5.01</td>
<td>0.01</td>
<td>0.19</td>
</tr>
<tr>
<td>5% quantile</td>
<td>7.33</td>
<td>−0.45</td>
<td>6.17</td>
<td>−0.34</td>
<td>0.49</td>
<td>2.67</td>
<td>2.45</td>
<td>−0.60</td>
<td>−0.40</td>
<td>−7.94</td>
<td>−2.44</td>
<td>0.03</td>
</tr>
<tr>
<td>95% quantile</td>
<td>8.27</td>
<td>−0.02</td>
<td>6.99</td>
<td>0.06</td>
<td>0.60</td>
<td>3.99</td>
<td>4.66</td>
<td>0.00</td>
<td>0.20</td>
<td>2.14</td>
<td>2.41</td>
<td>0.34</td>
</tr>
<tr>
<td>0.5% quantile</td>
<td>7.13</td>
<td>−0.49</td>
<td>5.99</td>
<td>−0.46</td>
<td>0.42</td>
<td>2.30</td>
<td>2.21</td>
<td>−0.90</td>
<td>−0.60</td>
<td>−8.80</td>
<td>−2.93</td>
<td>−0.01</td>
</tr>
<tr>
<td>99.5% quantile</td>
<td>8.79</td>
<td>0.08</td>
<td>7.26</td>
<td>0.10</td>
<td>0.64</td>
<td>4.37</td>
<td>4.95</td>
<td>0.30</td>
<td>0.40</td>
<td>−1.26</td>
<td>2.92</td>
<td>0.44</td>
</tr>
</tbody>
</table>

0.0004/1.20e-9, 0.0008/5.74e-14 and 0.0008/0.0203 respectively.

Applying this model to the training data/test data yields the performance data as shown in the upper part of Table 2. The respective performance plot where the modelled corneal SA is displayed versus the SA derived from raytracing is shown in the upper graph of Fig. 4.

Model 2: In addition, a simplified prediction model was set up based on the training data with the parameters Ra, Qa, Rp, Qp:

$$SA = 0.9180 - 0.0939 \times Ra + 0.7154 \times Qa + 0.0224 \times Rp - 0.1222 \times Qp$$

The respective standard errors $P$ values for the intercept and the 4 regression coefficients mentioned above are 0.0067/<1e-100, 0.0009/<1e-100 and 0.0019/<1e-100 respectively.

Applying this model to the training data/test data yields the performance data as shown in the lower part of Table 2. The respective performance plot where the modelled corneal SA is displayed versus the SA derived from raytracing is shown in the lower graph of Fig. 4.

Discussion

In recent decades, many different concepts for aspherical intraocular lenses have been developed (Kohnen & Klaaproth 2008; Wang et al. 2012). Currently, we have so called aberration-neutral or aberration-free lens designs having a slightly negative aspherical base curve, as well as aberration correcting lenses with some higher amount of negative asphericity at one or both optical surfaces (Langenbucher et al. 2017). However, to date, the philosophy behind aspherical lenses could simply add up the corneal SA (Langenbucher et al. 2017) as obtained from raytracing and the SA correction provided by the lens manufacturer to estimate the resulting SA (at corneal plane) in the pseudophakic eye after cataract surgery. However, we have to keep in mind that spherical aberration of the eye also enlarges the depth of focus, which could support near vision. Therefore, many intraocular lens manufacturers do not provide aspherical lenses, which are designed to fully correct the spherical aberration (of a schematic model eye).

The concept for extracting corneal SA from tomographic data is simple: we construct an optical model based on the data from an anterior segment tomograph using the shape of the corneal front and back surface, central corneal thickness, anterior chamber depth and the location of the pupil centre. Assuming that the axial location of the pupil is in the plane of the anterior lens surface, only the direction of collimated incident ray is required as a...
missing link. For simplification, we assume that all refractive surfaces in the eye are coaxially aligned, that only the pupil centre may be decentred and that the incident ray may be oblique. In this simplification, the ‘optical axis’ of the system is defined by all the coaxial refractive surfaces, and the incident ray describes the angle \(\alpha\), which in general cannot be measured with any instrumentation. In the present study, we used a random value for the incident ray angle with a mean of \(-5/5\) degree for left/right eyes and a standard deviation of \(\pm 2^\circ\) for both directions. As the angle \(\alpha\) cannot be measured, we decided to omit \(I_X\) and \(I_Y\) from the prediction models. However, as can be seen from our performance plots, the variation between the corneal SA derived from raytracing and the corneal SA derived from the 3 linear mixed-effects models is quite good, meaning that the incident ray angle may not only have a major impact on the tilt terms \(Z_1^1\) and \(Z_1^{-1}\) of the wavefront), but only a minor effect on the SA term \(Z_{40}\). Therefore, we feel that information on the visual axis is not mandatory for estimating the corneal SA from the tomographic data. As an alternative, we could use information from the Chang-Waring chord as offset between the Purkinje light reflex PI and the pupil centre to retrieve data on the visual axis (Chang & Waring 2014; Hollanday 2019; Jiang et al. 2020) in order to predict the incident ray angle from tomography.

Using a large dataset derived from the high-resolution anterior segment OCT Casia 2 from one clinical centre, we implemented a raytracing concept to calculate the optical path length differences in terms of a wavefront analysis. As the instrument axis is oriented coaxially with the incident beam, the optical path length differences had to be considered with respect to a plane perpendicular to the incident beam. Therefore, the reference plane considered for the wavefront analysis was tilted using a quaternion transformation with Euler angles as shown in the literature (St. Pierre 2021).

Fig. 2. Matrix of grouped scatterplots for the potential input parameters of the multivariable linear prediction model (Ra/Qa/Rp/Qp/CCT/ACD/Pup\(_X\)/Pup\(_Y\)/I\(_X\)/I\(_Y\)) and the spherical aberration SA calculated from the Zernike decomposition of the wavefront error considered in a plane perpendicular to the incident ray with a 6-mm diameter. Left eyes/right eyes are plotted in green/red. Please note that for this graph, left eyes were not flipped horizontally. The respective histograms are plotted on the diagonal of the matrix. From the graph, it is obvious that the corneal front surface asphericity is the most relevant effect size for SA.
Finally, the resulting wavefront error was decomposed into Zernike terms with a unit circle diameter of 6 mm to obtain the corneal SA term. The corneal SA was then modelled with a linear mixed-effects model (Herber et al. 2020). Linear mixed-effects models were used instead of standard linear models to consider the effect of intrapatient correlations in the dataset. To obtain realistic values for the model performance, we split our dataset into training and test data for cross-validation (hold-out strategy) using typical proportions of 70% for the training data and 30% for the test data. In this table, we provide the correlation coefficients R with the lower and upper limits of the confidence intervals RLO and RUP, the significance level comparing the model to a constant model (P), together with the mean prediction error (meanPE), the mean absolute prediction error (meanABSPE) and the root-mean-squared prediction error (rmsPE). There is no relevant difference in the performance data when applying the prediction model to the training data or to the test data, confirming that we do not have overfitting in the models.

<table>
<thead>
<tr>
<th>Model Description</th>
<th>Training Data</th>
<th>Test Data</th>
</tr>
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<tr>
<td>Prediction model based on Ra, Qa, Rp, Qp, CCT, PupX, PupY</td>
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<tr>
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<td>Simplified prediction model based on Ra, Qa, Rp, Qp</td>
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<td>Sparse prediction model based on Ra, Qa only</td>
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<td>rmsPE</td>
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</table>

Table 2. Performance of the 3 linear mixed-effects models (with patient ID as a grouping variable) derived from the training data. The models were applied to the training data and to the test data in a cross-validation process to check for potential overfitting. In this table, we provide the correlation coefficients R with the lower and upper limits of the confidence intervals RLO and RUP, the significance level comparing the model to a constant model (P), together with the mean prediction error (meanPE), the mean absolute prediction error (meanABSPE) and the root-mean-squared prediction error (rmsPE). There is no relevant difference in the performance data when applying the prediction model to the training data or to the test data, confirming that we do not have overfitting in the models.
model could also be used where measurement data are only available for the corneal front surface (e.g. from corneal topography), and not for the corneal back surface shape or central corneal thickness. Overall, we feel that all 3 linear prediction models perform well in predicting corneal SA from measurement data of the Casia2 anterior segment analyser. As can be seen from Table 2, there is not too much difference between Model 1 and Model 2, but both models slightly outperform the sparse model (Model 3) in terms of mean absolute prediction error (mean ABSPE) and root-mean-squared prediction error (rmsPE). The respective performance plots for the initial model, the simplified model and the sparse model are shown in Fig. 4. Here, the modelled corneal SA is displayed versus the corneal SA derived from raytracing together with the least-squares linear regression line.

There are some limitations in our study: we restricted the model to a simple optical model for the cornea with rotationally symmetric aspherical surfaces aligned on a common axis (which was considered as the ‘optical axis’). Additionally, we ignored the effect that the location of the pupil centre is affected by the pupil size as shown by Erdem et al. (2008); Yang et al. (2002). According to the convention of reporting the amount of SA correction with a 6-mm optical zone at corneal plane, we restricted our raytracing setup to this scenario. In the future, different optical zones could be used to investigate the effect of corneal spherical aberration as a function of pupil size. Further, there were no measurement data of the incident ray angle, and therefore, we had to use random data in our simulation instead based on mean values ($I_X = -5°$ / $I_Y = 0°$ for left eyes) from a modern schematic model eye (Liou & Brennan 1997), and last but not least, there are only limited data about the quality of the corneal asphericity measurements, about the characteristics in a cataractous population (De Sanctis et al. 2014) and especially about long-term changes (Amorim-de-Sousa et al. 2019) or iatrogenic changes due to cataract surgery (Li et al. 2012).

In conclusion, in this paper, we have developed simple linear prediction models to estimate corneal spherical aberration to be used for selecting the appropriate intraocular lens model in cataract surgery. A raytracing strategy was set up to derive corneal spherical aberration from a Zernike decomposition of the wavefront error considered at corneal plane perpendicularly to the incident ray. Our concept is based on a Monte-Carlo simulation with a large dataset from the anterior segment OCT. With cross-validation, the models showed a good performance with all relevant effect sizes identified by a stepwise regression, with a simplified prediction model considering the corneal front and back surface shape, but the performance was slightly lower with a sparse model considering only shape data of the corneal front surface.

### References


Li ZH, Jia LX & Huang YF (2012): Analysis of corneal spherical aberration derived from corneal topography, and not for the corneal back surface shape or central corneal thickness. Overall, we feel that all 3 linear prediction models perform well in predicting corneal SA from measurement data of the Casia2 anterior segment analyser. As can be seen from Table 2, there is not too much difference between Model 1 and Model 2, but both models slightly outperform the sparse model (Model 3) in terms of mean absolute prediction error (mean ABSPE) and root-mean-squared prediction error (rmsPE). The respective performance plots for the initial model, the simplified model and the sparse model are shown in Fig. 4. Here, the modelled corneal SA is displayed versus the corneal SA derived from raytracing together with the least-squares linear regression line.

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### References


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