

1 **Analysis of age-dependence of the anterior and posterior cornea with Scheimpflug**
2 **imaging**

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21 **Abstract**

22

23 **Purpose:** The assessment of keratometric and higher order aberrations (HOA) of the anterior
24 and posterior cornea and their age-related changes.

25 **Patient and methods:** Our study investigated one healthy eye of 227 subjects (age:
26 55.15 ± 21.2 years, range: 16-90 years, right eye 135, left eye 92). Images were captured from
27 each eye with Pentacam HR using automatic mode. Keratometric, astigmatism data and
28 corneal HOA were analyzed.

29 **Results:** With respect to laterality, no deviance was found in any of the parameters ($p > 0.05$).
30 Mean refractive error was 0.52 ± 0.23 D. Regarding both the anterior and posterior corneal
31 surfaces, the level of astigmatism decreased significantly with advancing age ($p < 0.05$). The
32 overall root mean square (RMS) of the HOA is continuously increasing with age ($r = 0.517$;
33 $p < 0.01$), which can be explained by the combined effect of the increase in both the anterior
34 and posterior corneal RMS HOA. Of the HOAs, the constant increase of the primary and the
35 secondary spherical aberration (SA) with aging ($p < 0.01$) is caused by the SA growth of the
36 anterior surface. Apart from these, only the vertical coma aberration of the posterior surface
37 and the vertical trefoil aberrations of both anterior and posterior surfaces showed a
38 significantly positive correlation with aging ($p < 0.05$). Horizontal components of coma and
39 trefoil aberrations showed no deviation with advancing age.

40 **Conclusions:** Corneal astigmatism shows a significant decrease with aging. Of the higher
41 order aberrations, primary and secondary spherical aberrations, vertical coma and vertical
42 trefoil significantly increase with age, while other HOAs show no correlation with aging.

43

44 **Keywords:** higher order aberration, corneal spherical aberration, coma aberration, Pentacam
45 HR

46

47 **Introduction**

48 The cornea represents about 80% of the refraction of the eye. In addition to lower
49 order aberrations (prism, defocus), higher order aberrations (HOA) of the eye and the cornea
50 are also known. Significant deviations in HOA can affect visual quality and contrast
51 sensitivity negatively. Subsequent to ophthalmic surgeries, subjects often complain about
52 dysphotopsia and low contrast sensitivity, especially under low-light conditions. Higher order
53 aberrations of the eye can account for some of such symptoms.¹ One of the HOAs being most
54 particularly investigated is spherical aberration (SA). Modern, aspheric intraocular lenses are
55 designed for its correction in different ways.

56 In order to develop appropriate aspheric intraocular lenses it is highly important to
57 investigate corneal SA measurements and to understand what changes they may undergo with
58 aging. Intraocular lenses are to compensate such HOAs of the eye as strongly as possible
59 bringing it close to the optimal state free of aberrations.

60 Both the anterior and the posterior surfaces contribute to the lower and higher order
61 aberrations of the cornea, however earlier, posterior surface examinations could be carried out
62 only with Orbscan. At present, posterior corneal analysis based on Scheimpflug imaging has
63 become well known with its good repeatability.²

64 A human eye changes with advancing age: in addition to several anatomic parameters,
65 higher-order aberration of the cornea will also undergo alterations.³ Our goal was to analyze
66 corneal aberrations of both the anterior and posterior surfaces changing with age using the
67 high resolution version of Pentacam (Pentacam HR) based on Scheimpflug imaging, the

68 already widely used device for the assessment of the anterior segment. According to our
69 knowledge this is the first study, involving a larger population, to assess anterior and posterior
70 corneal aberrations and to investigate their correlations with aging using Pentacam HR device.

71

72 **Patients and methods**

73

74 Our study comprised healthy volunteers with good distance vision (minimum of 20/25
75 Snellen equivalent) who had no clinically documented ocular deviation other than low
76 refraction error (lower than 0.75 D) and cataract. None of them was contact lens wearer.

77 Subsequent to the visual acuity test and prior to any other measurements and slit-lamp
78 examination 3 images were captured of each eye with the high resolution version of Pentacam
79 (Pentacam HR, software version 1.17r139) using Scheimpflug imaging. The device was set to
80 a 25 images/second mode and images were taken in auto mode at perfect eye-set. In case of
81 image distortion (e.g. blinking) or lack of data, the snapshot was repeated. Images were
82 captured of only one randomly chosen eye of each patient, which underwent further
83 evaluation.

84 The anterior segment rotating Scheimpflug camera rotates along the optical axis of the
85 eye. This device uses 475 nm monochromatic blue light for imaging, the camera captures 25,
86 50 or 100 scans in two seconds with 2760 measuring points. The software allows for
87 automatic analysis of the anterior segment, anterior and posterior topography of the cornea,
88 pachymetry, calculation of the chamber angle, volume, chamber height (anterior chamber
89 depth) and analysis of the lens. Finally, the instrument creates a 3D model of the anterior
90 segment. The software of the device corrects distortions in the Scheimpflug images based on
91 the geometry of the Scheimpflug principle.

92 Pentacam also allows us to assess and analyze the lower and higher order aberrations
93 of the eye. It calculates Zernike coefficients and also the values of the root mean square
94 (RMS) of the coma (square root of the sum of the squared coefficients of $Z(3,-1)$, $Z(3,1)$,
95 $Z(5,-1)$, and $Z(5,1)$) and the RMS of the spherical aberration (square root of the sum of the
96 squared coefficients of $Z(4,0)$ and $Z(6,0)$). Aberrations were evaluated at 8.0 mm pupil
97 setting. The research protocol adhered to the tenets of the Declaration of Helsinki.

98 Statistical analysis was performed with MedCalc 10.0. Descriptive statistical results
99 were described as mean, standard deviation (SD) and 95% confidence interval (95% CI) for
100 the mean. Normality of data was tested by Kolmogorov-Smirnov test. If the normality was
101 rejected ($p < 0.05$), nonparametric test was used. Mann-Whitney U test was carried out for
102 comparison between groups or variables, and Spearman rank test for correlation. Besides, we
103 performed linear regression analysis adjusting aberrometric data for age. A P value below
104 0.05 was considered statistically significant, and Bonferroni correction was applied for
105 multiple tests.

106

107 **Results**

108 Our study examined one eye of each of the 227 subjects (age: 55.15 ± 21.2 years, range:
109 16-90 years, 95% CI: 52.37-57.9 years). In 135 cases the right eyes, in 92 cases the left eyes
110 were investigated. With respect to laterality, no deviance was found in any of the parameters
111 ($p > 0.05$). Mean refractive error was 0.52 ± 0.23 D (range: -0.75- +0.75 D). Normality was
112 rejected for all parameters examined ($p < 0.001$). Detailed keratometric results of the anterior
113 and posterior corneal surfaces are displayed in Table 1. No significant correlations were found
114 between keratometric data and age ($p > 0.05$), however the level of keratometric astigmatism in
115 both the anterior and posterior surfaces decreased weakly, but significantly with advancing

116 age ($p < 0.05$) (Table 1). During the assessment of lower order aberrations of the eye, at Z2
117 level, the anterior and posterior corneal surface measurements demonstrated that the level of
118 aberrations increase significantly with aging ($p < 0.01$) (Table 2). Intraclass correlation was
119 good for all HOA data ($ICC = 0.86-0.91$). The examination of corneal aberration RMS shows
120 that the overall RMS and the total RMS HOA are continuously increasing with age ($p < 0.01$).
121 This process is due to the increase of the RMS and RMS HOA of the anterior surface
122 ($p < 0.01$); while the values of RMS and RMS HOA of the posterior surface do not show any
123 deviations with increasing age ($p > 0.05$) (Table 3). Of the HOAs, the constant increase of the
124 primary and secondary spherical aberration with age is striking ($p < 0.01$), which is due to the
125 increase of the SA of the anterior surface (Figure 1). The investigation of coma aberration
126 showed a significant and positive correlation between vertical coma aberration of the
127 posterior surface and age ($r = 0.273$; $p < 0.05$). (Figure 2.) However, vertical coma values of the
128 anterior and the entire corneal surface did not change significantly with advancing age.
129 Regarding trefoil aberration the vertical trefoil values of the posterior corneal surface
130 increased with age ($r = 0.154$; $p = 0.02$), while vertical trefoil values of the anterior and the
131 entire corneal surface slightly decreased with age ($r = -0.134$; $p = 0.04$ and $r = -0.15$; $p = 0.02$)
132 (Figure 3.). The horizontal components of coma- and trefoil aberrations showed no deviation
133 with aging ($p > 0.05$). The correlation between tetrafoil aberration and age was not significant
134 ($p > 0.05$). Detailed HOA data presented in Table 4.

135

136 **Discussion**

137 The investigations of the higher order aberration of the cornea and its correlation with
138 age have had controversial results so far, highly due to the differences in the investigation
139 methods. Present study, the first in literature, aimed to investigate the keratometric, lower and

140 higher order aberration of the anterior and posterior surfaces and their changes with age using
141 the high resolution version of Pentacam based on Scheimpflug imaging.

142 The keratometric parameters and the assessment of the anterior corneal surface have
143 already been well defined; however the level of the astigmatism of the posterior cornea was
144 investigated by only few studies.^{4,5} The mean posterior power in healthy eyes was -6.2 D,⁶
145 corneal curvature ranged between 5.8-6.78 mm.^{5,7} Dunne found that the spherical component
146 was -6.25--6.45 D and the cylindrical component was 0.34-0.38 D in the posterior corneal
147 surface.⁴ Our results, verifying a -6.15 D/-6.48 D mean keratometry in the posterior surface
148 comparable well with Dunne's findings.

149 The correlation between biometric data and age has already been observed and
150 published in cross-sectional examinations,⁸⁻¹² and was partly explained with the reduction in
151 the length of the constituent collagen fibers in tissues.⁸ According to the findings of Ho and
152 his coworkers using Pentacam,¹³ there were age-related shifts toward against-the-rule and
153 with-the-rule astigmatisms for the anterior and posterior corneal surfaces, respectively. Our
154 data demonstrate that keratometric astigmatism decreases slightly, however significantly in
155 both the anterior and posterior surfaces of the cornea with age. Contradictory, at Z2 level,
156 aberration data increase significantly with aging. Keratometric and Zernike astigmatism are
157 not calculated in the same way, and we think that the reason for this apparent contradiction is
158 that the latter one was calculated at large pupil diameter.

159 Certain higher order aberrations of the entire eye have already been investigated for at
160 least 50 years ago¹⁴⁻¹⁶ The reducing effect of HOA on image quality are also widely known.¹⁷
161 The posterior surface of the cornea has a role in these errors, but has been by far the least
162 investigated up till now. OrbScan¹⁸ and later Pentacam provided the opportunity for
163 assessment of the posterior surface, however, the limitations of Orbscan are well known.¹⁹

164 HOA measurements can be obtained with several devices.^{16,19,20-22} The dissimilarity of the
165 methods may give explanation for the differences in the results in the studies observing the
166 changes of total and corneal HOA with aging.

167 Eye aberrations belonging to various groups of refraction errors can be of different
168 levels;^{23,24} however, corneal topography¹⁰ and the total eye aberrometry²⁵ did not verify this
169 difference. In present study, eyes with high refractive errors were excluded, so the possible
170 correlation between the refractive state and the total corneal aberration was not analyzed.

171 The total corneal HOA, SA and coma are higher than the total ocular aberrations,²⁶
172 although Artal's study suggests that this applies only for a young age, at an older age it is just
173 the opposite.²⁷ Some of the total ocular aberrometric deviations change with advancing
174 age.^{27,28} Previous publications supported the increase of the corneal^{9,26,29} and the total
175 ocular^{16,27,28} coma aberration with aging. Other researches involving smaller population and
176 using spatially resolved refractometer observed that total corneal SA increases with aging, but
177 coma aberration shows no deviation.¹⁶ Our results are similar to the these findings, although it
178 obtained only the change of the vertical component of the coma aberration on the posterior
179 corneal surface. McLellan et al.¹⁶ described that the overall RMS wavefront error (excluding
180 tilt, astigmatism and defocus) significantly increases with advancing age especially with
181 respect to the fifth and seventh order. However, our data prove that the sixth order secondary
182 spherical aberration shows a significant change with age, while such deviation was not
183 detected in the fifth and seventh order.

184 The change of corneal SA with age is also controversial: according to several
185 researchers, corneal SA shows no change with aging,^{26,29} although topographic and
186 videokeratographic investigations⁹ reported changes and increase with aging. Using
187 Scheimpflug photography, they observed an increase with advancing age³⁰ because

188 the spherical aberration of the posterior corneal surface is negative at a young age and
189 becomes positive at an older age.³⁰ Another study using Pentacam suggests, that the anterior
190 corneal surface accounts for most of the corneal HOAs, and coma and spherical aberration
191 also increase with age.³¹ Our investigations based on Scheimpflug analysis demonstrated that
192 the total HOA of the cornea significantly increases with advancing age due to the increase of
193 the SA, though the coma and the vertical components of the trefoil aberration also play a role
194 in the increase of total HOA. Glasser et al. found that the overall SA of the examined eyes
195 was 0.25 μm .³² According to our measurements the corneal SA is much higher this value.

196 The videokeratographic and wavefront aberration results of Amano³ suggest that the
197 increase of the coma aberration of the total eye is mainly due to corneal changes. The age-
198 related increase of the ocular spherical aberration is not to be attributed to corneal changes,
199 but to the increase of the spherical aberration of the internal optics and the deviations in the
200 compensating effects of the crystalline lens radius^{33,34} and the refractive index³⁵ may also
201 contribute to its development. Systems compensating for refraction errors are known both in
202 young^{27,36} and in older eyes.³⁷ On one hand, the astigmatism of the posterior corneal surface
203 partially compensates for the astigmatism of the anterior corneal surface.⁵ The age-related
204 increase of SA can be of a lens origin,^{27,28} moreover, with the development of cataract the
205 total HOA also changes.³⁸ In a young eye the negative SA of the lens compensates for the
206 positive SA of the cornea,³⁵ however, at an older age, the SA of the lens increases, thus the
207 total ocular SA also increases.^{27,35} This can be the root cause of the development of a
208 decreasing image quality with an advancing age.⁹ It is known, that it is mainly the vertical
209 coma that can help the reading ability of a presbyopic eye because of its increasing effect on
210 the depth of focus, although we could only verify the increase of the vertical component of the
211 trefoil aberration with similar effect. Eventually, the degradation of the optics of an older eye

212 is due to the overbalance between the corneal and other surfaces of the eye.²⁷ Besides, it is
213 known that the HOA is pupil dependent,²⁹ and the pupil is becoming more miotic with
214 age,^{11,39} so the decreasing effect of corneal aberrations on vision quality impair decrease,
215 moreover, the Stiles-Crawford effect⁴⁰ decreases the effect of aberrations on visual acuity.

216 In conclusion, our data prove that the assessment of the anterior and posterior corneal
217 surfaces shows that the level of astigmatism continuously, weakly decreases. We proved that
218 primary and secondary spherical aberration and vertical trefoil aberration change significantly
219 with age. Along with other changes of the eye, these changes are probably part of a complex
220 system compensating for presbyopia, which may serve as a base for further studies.

221

222 **References**

223

224 1. Fan-Paul NI, Li J, Miller JS, Florakis GJ. Night vision disturbances after corneal refractive
225 surgery. *Surv Ophthalmol.* 2002;47:533-546.

226 2. Chen D, Lam AKC. Intrasession and intersession repeatability of the Pentacam system on
227 posterior corneal assessment in the normal human eye. *J Cataract Refract Surg.* 2007;33:448-
228 454.

229 3. Amano S, Amano Y, Yamagami S, Miyai T, Miyata K, Samejima T, Oshika T. Age-related
230 changes in corneal and ocular higher-order wavefront aberrations. *Am J Ophthalmol.*
231 2004;137:988-992.

232 4. Dunne MC, Royston JM, Barnes DA. Normal variations of the posterior corneal surface.
233 *Acta Ophthalmol (Copenh).* 1992;70:255-261.

234 5. Dubbelman M, Sicam VA, Van der Heijde GL. The shape of the anterior and posterior
235 surface of the aging human cornea. *Vision Res.* 2006;46:993-1001.

236 6. Seitz B, Langenbacher A, Hofmann B, Behrens A, Kus MM. Refractive power of the
237 human posterior corneal surface in vivo in relation to gender and age. *Ophthalmologe.*
238 1998;95:S50.

239 7. Seitz B, Torres F, Langenbacher A, Behrens A, Suárez E. Posterior corneal curvature
240 changes after myopic laser in situ keratomileusis. *Ophthalmology.* 2001;108:666-672.

241 8. Leighton DA, Tomlinson A. Changes in axial length and other dimensions of the eyeball
242 with increasing age. *Acta Ophthalmol.* 1972;50:815-826.

243 9. Guirao A, Redondo M, Artal P. Optical aberrations of the human cornea as a function of
244 age. *J Opt Soc Am A Opt Image Sci Vis.* 2000;17:1697-1702.

- 245 10. Philip K, Martinez A, Ho A, Conrad F, Ale J, Mitchell P, Sankaridurg P. Total ocular,
246 anterior corneal and lenticular higher order aberrations in hyperopic, myopic and emmetropic
247 eyes. *Vision Res.* 2012;52:31-37.
- 248 11. O'Donnell C, Hartwig A, Radhakrishnan H. Correlations between refractive error and
249 biometric parameters in human eyes using the LenStar 900. *Cont Lens Anterior*
250 *Eye.* 2011;34:26-31.
- 251 12. Xu L, Cao WF, Wang YX, Chen CX, Jonas JB. Anterior chamber depth and chamber
252 angle and their associations with ocular and general parameters: the Beijing Eye Study. *Am J*
253 *Ophthalmol.* 2008;145:929-936.
- 254 13. Ho JD, Liou SW, Tsai RJ, Tsai CY. Effects of aging on anterior and posterior corneal
255 astigmatism. *Cornea.* 2010;29:632-637.
- 256 14. Ivanof A. About the spherical aberration of the eye. *J Opt Soc Am.* 1953;46:901-903.
- 257 15. Castejón-Mochón JF, López-Gil N, Benito A, Artal P. Ocular wave-front aberration
258 statistics in a normal young population. *Vision Res.* 2002;42:1611-1617.
- 259 16. McLellan JS, Marcos S, Burns SA. Age-related changes in monochromatic wave
260 aberrations of the human eye. *Invest Ophthalmol Vis Sci.* 2001;42:1390-1395.
- 261 17. Chalita MR, Krueger RR. Correlation of aberrations with visual acuity and symptoms.
262 *Ophthalmol Clin North Am.* 2004;17:135-142.
- 263 18. Wang Z, Chen J, Yang B. Posterior corneal surface topographic changes after laser in situ
264 keratomileusis are related to residual corneal bed thickness. *Ophthalmology.* 1999;106:406-
265 409.
- 266 19. Cairns G, McGhee CNJ. Orbscan computerized topography: attributes, applications, and
267 limitations. *J Cataract Refract Surg.* 2005;31:205-220.

- 268 20. Webb RH, Penney CM, Thompson KP. Measurement of ocular wavefront distortion with
269 a spatially resolved refractometer. *Appl Optics*. 1992;31:3678-3686.
- 270 21. Liang J, Grimm B, Goelz S, Bille JF. Objective measurement of wave aberrations of the
271 human eye with the use of a Hartmann-Shack wave-front sensor. *J Opt Soc Am A*.
272 1994;11:1949-1957.
- 273 22. Guirao A, Artal P. Corneal wave aberration from videokeratography: accuracy and
274 limitations of the procedure. *J Opt Soc Am A Opt Image Sci Vis*. 2000;17:955-965.
- 275 23. Paquin MP, Hamam H, Simonet P. Objective measurements of optical aberrations in
276 myopic eyes. *Optom Vis Sci*. 2002;79:285-291.
- 277 24. He JC, Sun P, Held R, Thorn F, Sun X, Gwiazda JE. Wavefront aberrations in eyes of
278 emmetropic and moderately myopic school children and young adults. *Vis Res*. 2002;42:1063-
279 1070.
- 280 25. Cheng X, Bradley A, Hong X, Thibos LN. Relationship between refractive error and
281 monochromatic aberrations of the eye. *Optom Vis Sci*. 2003;80:43-49.
- 282 26. Wang L, Dai E, Koch DD. Optical aberrations of the human anterior cornea. *J Cataract*
283 *Refract Surg*. 2003;29:1514-1521.
- 284 27. Artal P, Berrio E, Guirao A, Piers P. Contribution of the cornea and internal surfaces to
285 the change of ocular aberrations with age. *J Opt Soc Am A Opt Image Sci Vis*. 2002;19:137-
286 143.
- 287 28. Wang L, Koch DD. Ocular higher-order aberrations in individuals screened for refractive
288 surgery. *J Cataract Refract Surg*. 2003;29:1896-1903.
- 289 29. Oshika T, Klyce SD, Applegate RA, Howland HC. Changes in corneal wavefront
290 aberrations with aging. *Invest Ophthalmol Vis Sci*. 1999;40:1351-1355.

- 291 30. Sicam VA, Dubbelman M, van der Heijde RG. Spherical aberration of the anterior and
292 posterior surfaces of the human cornea. *J Opt Soc Am A Opt Image Sci Vis.* 2006;23:544-549.
- 293 31. Cermáková S, Skorkovská S. Corneal higher order aberrations and their changes with
294 aging. *Cesk Slov Oftalmol.* 2010;66:254-257.
- 295 32. Glasser A, Campbell MCW. Presbyopia and the optical changes in the human crystalline
296 lens with age. *Vision Res.* 1998;38:209-229.
- 297 33. Brown N. The changes in lens curvature with age. *Exp Eye Res.* 1974;19:175-183.
- 298 34. Dubbelman M, Van der Heijde GL. The shape of the aging human lens: curvature,
299 equivalent refractive index and the lens paradox. *Vis Res.* 2001;41:1867-1877.
- 300 35. Smith G, Atchison DA, Pierscionek BK. Modeling the power of the aging human eye. *J*
301 *Opt Soc Am A.* 1992;9:2111-2117.
- 302 36. Artal P, Guirao A, Berrio E, Williams DR. Compensation of corneal aberration by the
303 internal optics in the human eye. *J Vis.* 2001;1:1-8.
- 304 37. He JC, Gwiazda J, Thorn F, Held R. Wave-front aberrations in the anterior corneal surface
305 and the whole eye. *J Opt Soc Am A Opt Image Sci Vis.* 2003;20:1155-1163.
- 306 38. Kuroda T, Fujikado T, Maeda N, Oshika T, Hirohara Y, Mihashi T. Wavefront analysis of
307 higher-order aberrations in patients with cataract. *J Cataract Refract Surg.* 2002;28:438-444.
- 308 39. Buckley C, Curtin DM, Docherty J, Eustace P. Ageing and alpha adrenoreceptors in the iris.
309 *Eye.* 1987;1:211-216.
- 310 40. Stiles WS, Crawford BH. The luminous efficiency of rays entering the eye pupil at
311 different points. *Proc R Soc Lond B.* 1933;112:428-450.

312 **Legends of figures**

313

314 Figure 1.

315 Corneal primary spherical aberration (SA) as a function of age. It can be seen, that the total
316 SA increase of the cornea ($r=0.273$; $p<0.001$) is due to the SA increase of the anterior corneal
317 surface ($r=0.255$; $p=0.001$).

318

319 Figure 2.

320 Graph showing the correlation between vertical coma aberration and age. The vertical coma
321 aberration of the posterior surface showed a slight regression line with age ($r=0.291$; $p<0.01$).
322 Values of total corneal vertical coma and that of the anterior surface showed no significant
323 change with advancing age.

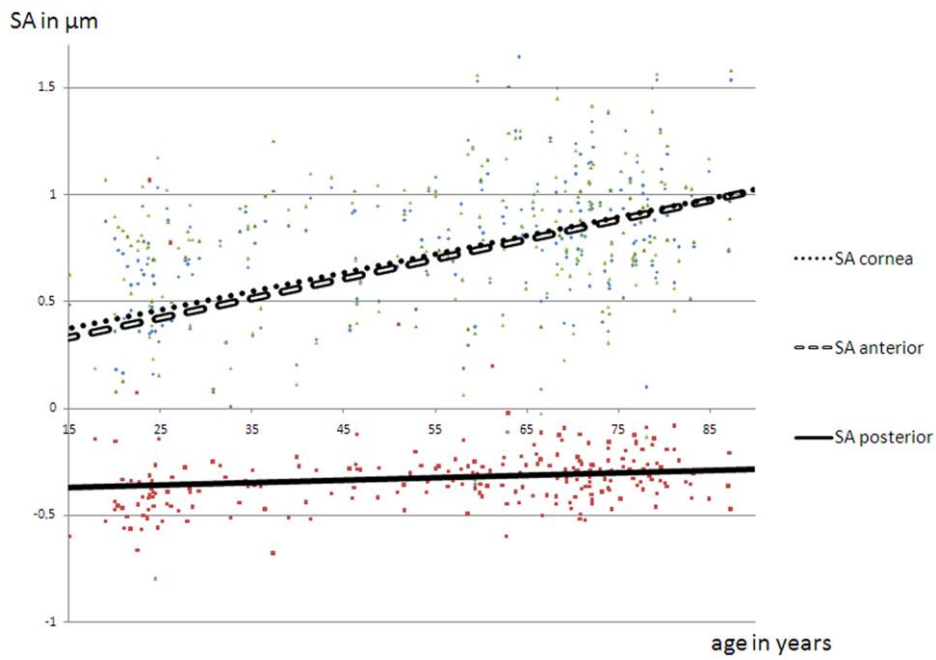
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325 Figure 3.

326 Vertical trefoil aberration changes of the total cornea, the anterior corneal surface and the
327 posterior corneal surface with age. The vertical trefoil value of the posterior corneal surface
328 increases moderately with age ($r=0.162$; $p=0.018$), while the vertical trefoil values of the
329 anterior surface and the entire cornea moderately decrease ($r=-0.146$; $p=0.048$ and $r=-0.151$;
330 $p=0.031$).

331

Age dependency of anterior and posterior corneal aberration

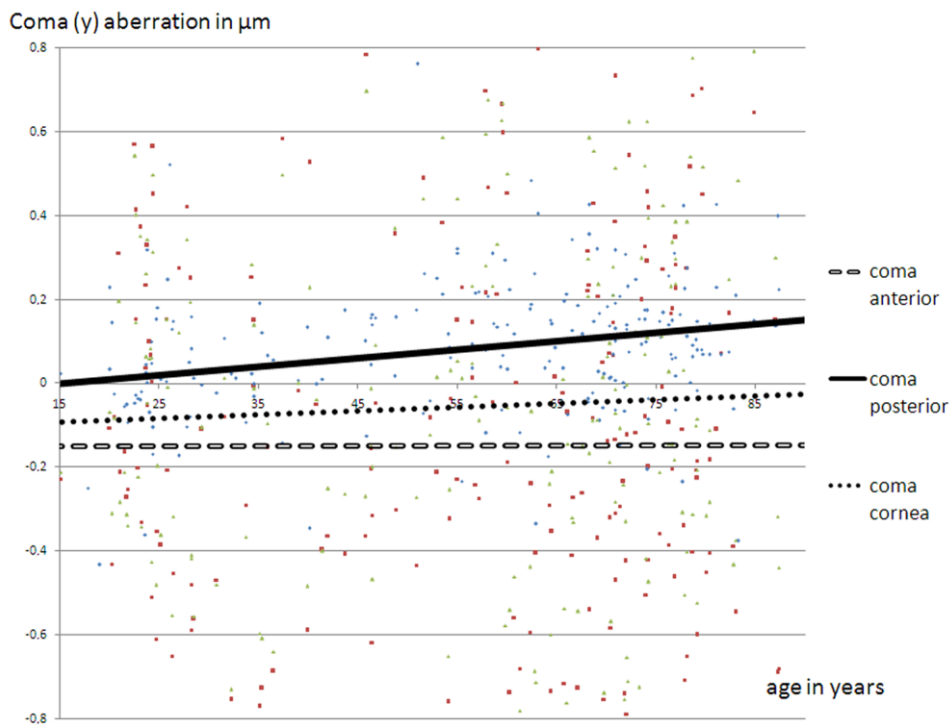


332

333 Figure 1

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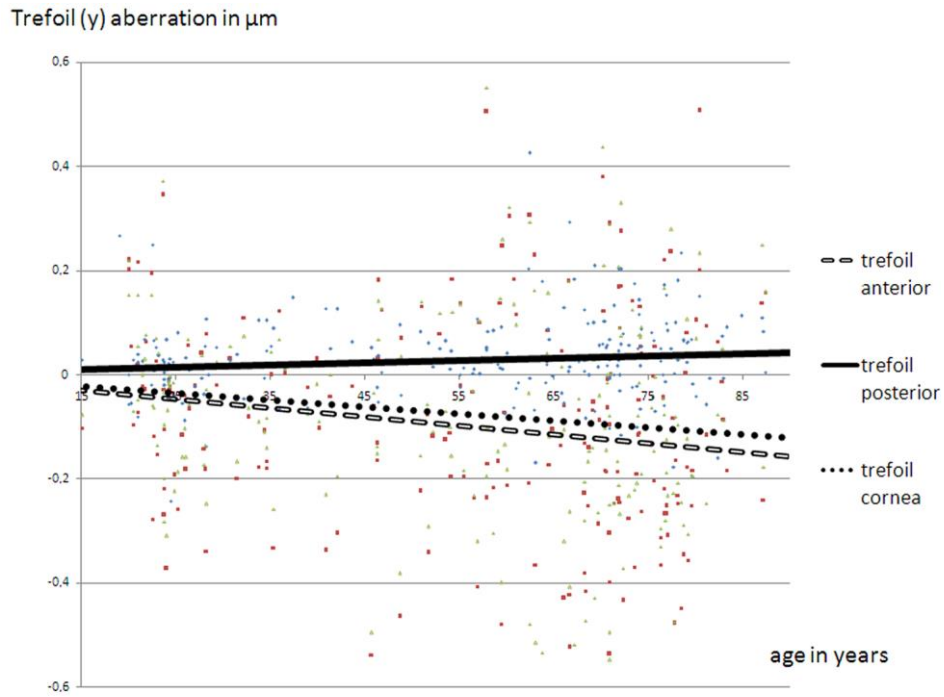
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337 Figure 2

Age dependency of anterior and posterior corneal aberration



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339 Figure 3

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Age dependency of anterior and posterior corneal aberration

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	Mean	SD	95% CI	Minimum	Maximum	b	p
K1 B (D):	-6.15	0.28	-6.19 - -6.11	-7.73	-4.93	0.008	0.508
K1 F (D):	43.24	1.95	42.98 - 43.45	38.66	48.76	0.031	0.381
K2 B (D):	-6.47	0.32	-6.52 - -6.43	-9.23	-5.70	0.134	0.114
K2 F (D):	44.21	1.88	43.96 - 44.46	40.03	57.36	0.091	0.147
Astigmatism B (D):	0.32	0.19	0.29 - 0.35	0.03	1.56	-0.170	0.010
Astigmatism F (D):	0.96	1.14	0.81 - 1.11	0.03	9.33	-0.259	<0.001

342

343

344 Table 1.: Statistical data of keratometric values in a healthy population measured with
 345 Pentacam HR (N=227). K1: keratometric value in diopter (D) on anterior corneal surface (F)
 346 and posterior corneal surface (B), in flat axis, K2: keratometric value in diopter (D) in the
 347 steep axis on anterior corneal surface (F) and posterior corneal surface (B). SD: standard error
 348 of mean, 95% CI: 95% confidence interval for mean, b: regression coefficients between age
 349 and the dependents, p: significance level of regression

350

351

352

	name of Zernike order	Mean	SD	95% CI	Minimum	Maximum	b	p
Z 2 -2 in μm (CB):	astigmatism (y)	0.0443	0.231	0.014 - 0.075	-0.604	0.772	-0.072	0.265
Z 2 -2 in μm (CF):	astigmatism (y)	-0.037	1.088	-0.182 - 0.106	-4.900	4.527	0.243	<0.001
Z 2 -2 in μm (Cornea):	astigmatism (y)	-0.033	1.235	-0.197 - 0.130	-6.281	3.932	0.215	<0.001
Z 2 0 in μm (CB):	defocus	-1.443	0.974	-1.572 - -1.314	-4.426	6.020	0.493	<0.001
Z 2 0 in μm (CF):	defocus	3.167	1.721	2.625 - 3.711	-4.062	8.132	0.312	<0.001
Z 2 0 in μm (Cornea):	defocus	3.397	1.690	2.947 - 3.391	-8.280	7.512	0.478	<0.001
Z 2 2 in μm (CB):	astigmatism (x)	0.371	0.319	0.328 - 0.413	-1.305	1.555	-0.365	<0.001
Z 2 2 in μm (CF):	astigmatism (x)	-0.905	1.807	-1.144 - -0.666	-7.641	11.713	0.436	<0.001
Z 2 2 in μm (Cornea):	astigmatism (x)	-0.640	1.276	-0.893 - -0.386	-13.12	10.503	0.497	<0.001

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355 Table 2.: Astigmatism as a lower order aberration of the entire cornea and the anterior corneal

356 surface (CF) and posterior corneal surface (CB) respectively, measured with Pentacam HR.

357 SD: standard error of mean, 95% CI: 95% confidence interval for mean, b: regression

358 coefficients between age and the dependents, p: significance level of regression

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Age dependency of anterior and posterior corneal aberration

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	Mean	SD	95% CI	Minimum	Maximum	b	p
RMS in μm (CB):	1.175	0.541	1.108 - 1.252	0.393	3.905	0.042	0.531
RMS in μm (CF):	3.119	1.631	2.954 - 3.446	0.589	10.929	0.383	<0.001
RMS in μm (Cornea):	2.905	1.661	2.683 - 3.127	0.603	9.895	0.512	<0.001
RMS HOA in μm (CB):	0.274	0.165	0.252 - 0.291	0.085	0.945	0.304	<0.001
RMS HOA in μm (CF):	0.858	0.698	0.766 - 0.950	0.206	3.882	0.496	<0.001
RMS HOA in μm (Cornea):	0.834	0.731	0.737 - 0.931	0.170	5.695	0.517	<0.001
RMS LOA in μm (CB):	1.138	0.524	1.069 - 1.207	0.380	3.882	0.027	0.681
RMS LOA in μm (CF):	2.973	1.445	2.779 - 3.167	0.524	7.121	0.476	<0.001
RMS LOA in μm (Cornea):	2.783	1.565	2.573 - 2.992	0.531	9.710	0.532	<0.001

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363 Table 3.: Root mean square (RMS) values with respect to lower and higher order aberration
 364 (LOA and HOA), regarding the entire cornea and the anterior corneal surface (CF) and
 365 posterior corneal surface (CB) with Pentacam HR, in a healthy population (N=227). SD:
 366 standard error of mean, 95% CI: 95% confidence interval for mean, b: regression coefficients
 367 between age and the dependents, p: significance level of regression

Age dependency of anterior and posterior corneal aberration

	name of Zernike order	Mean	SD	95% CI	Minimum	Maximum	b	p
Z 3 -3 in μm (CB):	trefoil (y)	0.028	0.109	0.013 - 0.042	-0.637	0.630	0.162	0.018
Z 3 -3 in μm (CF):	trefoil (y)	-0.123	0.577	-0.198 - -0.046	-4.34	4.585	-0.146	0.048
Z 3 -3 in μm (Cornea):	trefoil (y)	-0.049	0.521	-0.157 - 0.058	-3.795	4.035	-0.151	0.031
Z 3 -1 in μm (CB):	coma (y)	0.078	0.164	0.056 - 0.099	-0.45	0.762	0.291	<0.001
Z 3 -1 in μm (CF):	coma (y)	-0.136	0.703	-0.229 - -0.042	-3.652	2.134	-0.027	0.678
Z 3 -1 in μm (Cornea):	coma (y)	-0.045	0.683	-0.136 - 0.045	-3.378	3.757	0.039	0.535
Z 3 1 in μm (CB):	coma (x)	0.017	0.162	-0.004 - 0.038	-0.665	0.806	0.047	0.395
Z 3 1 in μm (CF):	coma (x)	-0.118	0.677	-0.203 - -0.028	-4.785	2.638	0.055	0.354
Z 3 1 in μm (Cornea):	coma (x)	-0.114	0.604	-0.194 - -0.034	-4.387	2.076	0.145	0.092
Z 3 3 in μm (CB):	trefoil (x)	0.011	0.098	-0.002 - 0.024	-0.343	0.978	-0.118	0.124
Z 3 3 in μm (CF):	trefoil (x)	-0.011	0.325	-0.055 - 0.031	-1.922	2.052	0.041	0.522
Z 3 3 in μm (Cornea):	trefoil (x)	-0.015	0.384	-0.065 - 0.036	-3.349	1.759	0.037	0.651
Z 4 -4 in μm (CB):	tetrafoil (y)	0.008	0.085	-0.002 - 0.020	-0.252	0.884	0.031	0.665
Z 4 -4 in μm (CF):	tetrafoil (y)	-0.082	0.479	-0.141 - -0.018	-4.311	0.458	-0.07	0.256
Z 4 -4 in μm (Cornea):	tetrafoil (y)	-0.062	0.478	-0.123 - 0.002	-3.803	2.527	-0.076	0.224
Z 4 -2 in μm (CB):	secondary astigmatism (y)	0.002	0.056	-0.004 - 0.010	-0.268	0.404	-0.122	0.092
Z 4 -2 in μm (CF):	secondary astigmatism (y)	0.025	0.289	-0.012 - 0.063	-1.243	2.009	0.041	0.522
Z 4 -2 in μm (Cornea):	secondary astigmatism (y)	0.021	0.279	-0.015 - 0.058	-1.131	1.941	0.039	0.488
Z 4 0 in μm (CB):	primary spherical	-0.323	0.175	-0.346 - -0.300	-0.682	1.063	0.139	0.014
Z 4 0 in μm (CF):	primary spherical	0.711	0.753	0.612 - 0.811	-5.291	2.877	0.255	0.001
Z 4 0 in μm (Cornea):	primary spherical	0.729	0.667	0.641 - 0.817	-3.992	3.682	0.273	<0.001
Z 4 2 in μm (CB):	secondary astigmatism (x)	-0.035	0.076	-0.045 - -0.025	-0.589	0.359	-0.345	<0.001
Z 4 2 in μm (CF):	secondary astigmatism (x)	-0.042	0.373	-0.091 - 0.006	-1.415	3.115	0.123	0.084
Z 4 2 in μm (Cornea):	secondary astigmatism (x)	-0.061	0.413	-0.115 - -0.005	-1.342	3.266	0.034	0.654
Z 4 4 in μm (CB):	tetrafoil (x)	-0.032	0.087	-0.043 - -0.020	-0.891	0.213	-0.032	0.567
Z 4 4 in μm (CF):	tetrafoil (x)	-0.058	0.422	-0.115 - -0.002	-2.933	4.094	0.041	0.462
Z 4 4 in μm (Cornea):	tetrafoil (x)	-0.114	0.533	-0.184 - -0.043	-5.337	3.303	0.065	0.342
Z 6 0 in μm (CB):	secondary spherical	0.011	0.043	0.005 - 0.016	-0.322	0.158	-0.377	<0.001
Z 6 0 in μm (CF):	secondary spherical	-0.075	0.184	-0.099 - -0.050	-0.608	1.509	-0.286	<0.001
Z 6 0 in μm (Cornea):	secondary spherical	-0.067	0.234	-0.098 - -0.036	-2.648	1.286	-0.378	<0.001

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369 Table 4.: Higher order aberrations labeled on the basis of Zernike pyramid for the entire cornea, and the anterior corneal surface (CF) and
370 posterior corneal surface (CB). SD: standard error of mean, 95% CI: 95% confidence interval for mean, b: regression coefficients between age
371 and the dependents, p: significance level of regression