<u>Title:</u>

Corneal topography with an aberrometry-topography system

Authors:

Michael Mülhaupt,¹ Sven Dietzko,¹ James Wolffsohn,² Stefan Bandlitz,^{1,2}

¹Höhere Fachschule für Augenoptik Köln (Cologne School of Optometry), Cologne,

Germany

²Ophthalmic Research Group, Life and Health Sciences, Aston University, Birmingham, UK

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Corresponding Author

Stefan Bandlitz

Höhere Fachschule für Augenoptik Köln

(Cologne School of Optometry)

Bayenthalgürtel 6-8

D-50968 Köln, Germany

e-mail: bandlitz@hfak.de

Telephone: 0049-221-348080

ABSTRACT

Purpose: To investigate the agreement between the central corneal radii and corneal eccentricity measurements generated by the new Wave Analyzer 700 Medica (WAV) compared to the Keratograph 4 (KER) and to test the repeatability of the instruments.

Methods: 20 subjects (10 male, mean age 29.1 years, range 21-50 years) were recruited from the students and staff of the Cologne School of Optometry. Central corneal radii for the flat (r_{c/fl}) and steep (r_{c/st}) meridian as well as corneal eccentricity for the nasal (e_{nas}), temporal (e_{temp}), inferior (e_{inf}) and superior (e_{sup}) directions were measured using WAV and KER by one examiner in a randomized order.

Results: Central radii of the flat (rc/fl) and steep (rc/st) meridian measured with both instruments were statically significantly correlated (r=0.945 and r=0.951; p<0.001). Comparison between the WAV and KER showed that $r_{c/fl}$ and $r_{c/st}$ measured with WAV were significantly steeper than those measured with KER (p<0.001). Corneal eccentricities were statistically significantly correlated in all meridians (p<0.05). Compared to KER, e_{temp} and e_{sup} measured with WAV were greater (p<0.05), while there were no statistically significant differences for e_{nas} and e_{inf} (p=0.350 and p=0.083). For the central radii, repeated measurements were not significantly different for the KER or WAV (p>0.05). Limits of agreement (LoA) indicate a better repeatability for the KER compared to WAV.

Conclusions: Corneal topography measurements captured with the WAV were strongly correlated with the KER. However, due to the differences in measured corneal

radii and eccentricities, the devices cannot be used interchangeably. For corneal topography the KER demonstrated better repeatability.

Key words: Corneal topography, placido-based, corneal radius, corneal eccentricity, aberrometry-topography.

The measurement of the shape, refractive power and thickness of the cornea is essential for the planning of corneal refractive surgery, for diagnosis of corneal diseases and for fitting contact lenses, in particular speciality lenses. Various diagnostic procedures have been developed for the analysis of the corneal surface. Corneal topographical measurements can be performed by classic Placido-based topographers as well as by tomography systems that produce three-dimensional corneal models from cross-sectional images [1].

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9 Placido-based computerized videokeratoscopy, proposed first by Klyce in 1984 [2], are 10 the most frequently used corneal topography systems in clinical practice [3]. This 11 method of imaging of the anterior corneal surface analyses tear film reflected images 12 of multiple concentric rings projected on the cornea. In contrast, corneal tomography 13 provides an analysis of the shape of anterior and posterior corneal surfaces, as well 14 as the thickness distribution of the cornea [4]. Corneal tomography can be performed 15 by a scanned slit, rotating Scheimpflug cameras or by optical coherence tomography 16 [5].

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Recently, a new corneal topography with an integrated aberrometry-topography system named the Wave Analyzer 700 Medica (Essilor, Freiburg, Germany) has been introduced to the market. The Wave Analyzer is a multifunctional device for performing objective refraction, aberrometry, pupillometry, pachymetry, non-contact tonometry, measurement of anterior chamber depth and angle as well as corneal topography. The instrument combines a Hartmann-Shack aberrometer, an air tonometer, a Scheimpflug camera and a Placido-based topographer. However, the data for the corneal radii and

corneal eccentricity is only generated from the Placido-disc measurement without any
 contribution of the Scheimpflug camera.

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28 Consequently, the aims of this study were (i) to investigate the agreement in the 29 measurement of central corneal radii and corneal eccentricity between the new Wave 30 Analyzer 700 Medica (WAV) and the Placido-based Keratograph 4 (KER) (Oculus 31 Optikgeräte GmbH, Wetzlar, Germany) and (ii) to test the repeatability of the 32 instruments.

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35 MATERIALS AND METHODS

36 Instruments

37 To measure central corneal radii as well as corneal eccentricity, two placido based 38 corneal topographers were used in this study. The Keratograph 4 (Oculus Optikgeräte 39 GmbH, Wetzlar, Germany) uses a placido cone consisting of 22 red illuminated rings 40 (650nm) at 80mm from the eye to generate 22 000 measuring points. The Wave 41 Analyzer 700 Medica (Essilor, Freiburg, Germany) is a diagnostic device that performs 42 objective refraction, aberrometry, pupillometry, crystalline lens opacity, pachymetry, 43 tonometry and topography. For corneal topography it uses a placido cone off 24 rings to generate 6144 measuring points. Instruments had been calibrated following the 44

45 manufacturer's recommendations. The room temperature was maintained at 18 to46 22°C.

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48 In Vitro Study

Four precision glass balls (radii: 6.00, 7.00, 8.00 and 9.00 mm; CA 100-Caldev, Topcon, Tokyo, Japan) were used as a model of the cornea. The mean of three consecutive measurements of the four glass balls was compared between the KER and the WAV at two different sessions at the same time of day (day 1 and day 2).

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56 In Vivo Study

57 Twenty healthy subjects (mean age 29.1 ± 9.2 (SD) years, range 21 to 50 years, even 58 male to female split) were recruited from the students and staff of the Höhere 59 Fachschule für Augenoptik Köln (Cologne School of Optometry), Cologne, Germany. All subjects underwent a medical history and a slit lamp examination. Subjects were 60 61 excluded if: they had a current or previous condition known to affect the cornea, 62 conjunctiva or the sclera such as pterygium and pinguecula; had a history of previous 63 ocular surgery, including refractive or strabismus surgery, eyelid surgery, or corneal 64 surgery; had any previous ocular trauma; were diabetic; were taking medication known to affect the ocular surface or sclera; and/or had worn rigid contact lenses or soft 65 66 contact lenses during the preceding 24 hours prior to the study.

The study was approved by the Research Ethics Committee and all subjects gave written informed consent before participating in the study. The procedures were conducted in accordance with the requirements of the Declaration of Helsinki (1983) and patient data were used only in anonymized form.

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Central corneal radii for the flat ($r_{c/fl}$) and steep ($r_{c/st}$) meridian as well as corneal eccentricity for the nasal (e_{nas}), temporal (e_{temp}), inferior (e_{inf}) and superior (e_{sup}) direction were measured by one examiner using the WAV and the KER in a randomized order. Corneal eccentricities were taken from the data given for an angle of 30°. The mean of three consecutive measurements of the right eye was recorded for both instruments at two different sessions at the same time of day (day 1 and day 2).

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82 Statistical Analyses

Normal distribution of data was analyzed by Shapiro-Wilk test. As the data was normally distributed, differences between sessions (day 1 and day 2) and instruments were analyzed using Bland-Altman plots, coefficient of repeatability (CR), and paired t-tests. The relationship between the WAV and KER measurements was analyzed by Pearson product-moment correlation. Data were analyzed using SigmaPlot 12 (Systat Software Inc., Chicago, USA).

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90 **RESULTS**

91 In Vitro Study

92 The measured radii of the four glass balls were 6.01, 6.97, 7.99, and 8.99 mm for the 93 WAV and 6.02, 7.01, 8.00, and 9.00 mm for the KER. The mean difference between the measurements of the two devices was 0.018 mm (95% confidence interval [CI], -94 95 0.015 to + 0.050 mm; p = 0.125) (Figure 5). Repeated measurements from day 1 and day 2 were not significantly different for the KER (paired t-test: p = 0.391), but they 96 were different for the WAV (p = 0.034). The mean difference and the limits of 97 98 agreement (LoA) indicate a better in vitro repeatability for the KER (0.005 mm; LoA -99 0.013 to 0.008 mm) compared to the WAV (0.030 mm; LoA -0.003 to +0.118 mm).

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101 In Vivo Study

Table 1 summarizes the mean values ± standard deviations of central corneal radii and corneal eccentricities, mean difference and limits of agreement (LoA) of the two measuring sessions (day 1 to day 2) and the mean differences and 95% confidence interval between the two instruments.

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107 Central corneal radii of the flat ($r_{c/fl}$) and steep ($r_{c/st}$) meridian measured with both 108 instruments were statically significantly correlated (r=0.945 and r=0.951; both 109 p<0.001). On average the mean central radii measured with the WAV were significantly 110 steeper than those measured with the KER (-0.05mm; CI -0.08 to -0.02; paired t-test; 111 p<0.001) (Figure 6).

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The measured corneal eccentricities were statistically significantly correlated in all meridians (e_{nas} ;r=0.747, e_{temp} ;r=0.541, e_{inf} ;r=0.783 and superior e_{sup} ;r=0.661; all p<0.05). On average the mean corneal eccentricities measured with the WAV were significantly greater than those measured with the KER (+0.06; CI 0.0126 to 0.105; paired t-test; p=0.009) (Figure 7). Compared to the KER, e_{temp} and e_{sup} measured with

- the WAV were greater (p<0.05), while there were no statistically significant differences for e_{nas} and e_{inf} (p=0.350 and p=0.083) (Table 1).
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For the central radii, repeated measurements from day 1 to day 2 were not significantly different for the KER and WAV (paired t-test; rc/fl: p=0.523 and p=0.860; rc/st: p=0.783 and p=0.154). The mean difference and the limits of agreement (LoA) indicate a better repeatability for the KER compared to the WAV (Table 1).

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For the overall corneal eccentricity, repeated measurements from day 1 to day 2 were not significantly different for the KER and the WAV (paired t-test; p > 0.05). The mean difference and the limits of agreement (LoA) indicate a better repeatability for the KER compared to the WAV (Table 1).

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132 **DISCUSSION**

The Wave Analyzer is a multifunctional device for performing objective refraction, aberrometry, pupillometry, pachymetry, non-contact tonometry and corneal topography. Comparing the values obtained for corneal topography with those of a placido-based Keratograph 4 showed a high correlation. However, radii measured with the Wave Analyzer were, on average, 0.06 mm and 0.09 mm (flat or steep meridian) steeper than those of the Keratograph 4.

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Shneor et al. [6] compared the L80 (Visionix Luneau, Chartes, France), a multi-function
device similar to the Wave Analyzer, with a manual Bausch & Lomb ophthalmometer.
As in the present study, they report statistically significantly steeper central radii
measurements (by 0.05 mm and 0.07 mm in the flat or steep meridians respectively)

144 compared to the manual ophthalmometer. For the Keratograph 4 (Oculus, Germany),
145 Best et al. reported flatter central corneal radii compared to Tonoref II (Nidek, Japan)
146 [7].

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148 Likewise, a comparison of the Placido-based Allegro Topolyzer system (Alcon 149 Research, Ltd., Fort Worth, TX, USA) with a Scheimpflug camera-based Galilei G4 150 system (Ziemer Ophthalmic Systems AG, Port, Switzerland) showed statistically 151 significant differences in the central corneal radii [8]. The Scheimpflug camera-based 152 system showed steeper radii than the Placido-based system; the differences in 153 patients with keratoconus were even greater [8, 9]. Comparing the Orbscan II (Orbtek), 154 a combination of a slit scanning technique and Placido disc image, with the Palcido-155 based EyeSys (Houston, TX, USA), Douthwaite and Mallen [10] found that the 156 Orbscan appears to under-read slightly for both apical radius and p-value.

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In contrast, Laursen et al. [11] reported no significant differences in the measurement
of mean corneal power between different devices: Keratograph 4, Pentacam (Oculus,
Germany), Tonoref II (Nidek, Japan), IOLMaster 500 and Lensstar LS 900 (Haag-Streit,
Switzerland). A comparison of three Scheimpflug camera-based systems (Pentacam,
Galilei G2 and Sirus 3D) in a study by Hernández-Camarena et al. [12] also did not
show any statistically significant differences in the measurement of the central corneal
radii.

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For corneal eccentricities, significant differences (mean differences from 0.08 to 0.26)
were found comparing four topographers (Humphrey, Atlas 991 (Zeiss), Dicon CT200
(Dicon, US), Orbscan II (Orbtek) and Medmont E300 (Medmont, Australia)) [13], which

is in concordance to the mean differences of 0.07 and 0.08 reported for the temporaland superior eccentricities in the present study.

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Furthermore, in the present study, a better in vivo repeatability of the measurements was obtained for the Keratograph 4 compared to the WaveAnalyzer. The values for the Keratograph 4 described in this study are in good agreement with repeatability described by Riede-Pult et al. [14] for the Keratograph 2. Device-specific differences in the repeatability of the measurement of central corneal radii as well as corneal eccentricities have already been reported in several studies [11-13, 15, 16].

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179 For the differences in measurement and in repeatability described in the various 180 studies, several causes can be considered: differences in the measuring principle 181 (manual keratometry, Placido-based systems, Scheimpflug camera-based systems); 182 differences in the measured area of the cornea (e.g. number of Placido-rings); different 183 calculation algorithms of the devices; as well as differences between the subjects (eg. 184 keratokonus or dry eye). Hamer et al. suggested, that the Placido-based systems seem 185 to be more susceptible to changes in the tear film than the Scheimpflug camera-based 186 systems [16]. Corneal topographers such as those utilising a Placido disc, analyse the 187 pattern of light rays reflected off the cornea and tear film-air interface and therefore any 188 disruption of the tear film may influence the measurement [16]. Since the reflection 189 quality of the placido mires indicates the quality of the tear film over time, topographers 190 can also be used to assess tear film stability [7].

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A limitation of the present study results from the eye models used for the *in vitro* study.
The glass balls had spherical surfaces which does not ideally reflects the aspherical
shape of most corneas. Therefore, Douthwaite [17] proposed the use of conicoidal

195 surface convex polymethylmethacrylate buttons to produce surfaces similar to the 196 normal healthy human cornea. However, both instruments in the present study where 197 calibrated using the manufactures spherical glass probes which corresponds to the 198 normal procedure in clinical practice. Furthermore, it should be noted that in vitro 199 models are never able to accurately reproduce the complexity of *in vivo* conditions [18, 200 19]. As a further limitation it should be noted, that in this study only healthy eyes were 201 included. McMahon et al. [20, 21] reported a loss in repeatability and reliability of 202 corneal topography measurements when corneal irregularity was present.

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204 Although corneal topography has improved over time, it appears that even two devices, 205 which are based on the same measuring principle as in this study, do not necessarily 206 lead to the same measurement result and equivalent repeatability. Some devices have 207 better repeatability than others, and therefore not all devices can be used 208 interchangeable. It has been suggested that mathematicals models should be 209 constructed to adjust the data of one instrument to be comparable to another [20], but 210 this presumes instruments are repeatable and differences are systematic across all 211 subjects.

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Practitioners should be aware of the measuring accuracy and the repeatability of the topography instrument used. This is important for the appropriate selection of the first contact lens to be trialled, as well as for the diagnosis and monitoring of corneal changes, especially when different topography systems are in use.

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219 CONCLUSIONS

220	Comparing the corneal topography determined by the Wave Analyzer with that of the
221	Keratograph 4 showed a high correlation. However, due to the differences in measured
222	corneal radii and eccentricities, the devices cannot be used interchangeably. For
223	corneal topography the KER demonstrated better repeatability.
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226	Conflict of interest
227	None
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318	Figures
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320	Figure 1. Wave Analyzer 700 Medica (Essilor, Freiburg, Germany).
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322	Figure 2. Keratograph 4 (Oculus GmbH, Wetzlar, Germany).
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324	Figure 3. Output of the Wave Analyzer 700 (Essilor, Freiburg, Germany).
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326	Figure 4. Output of the Keratograph 4 (Oculus GmbH, Wetzlar, Germany).
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328	Figure 5. In vitro difference in mean radius (mm) between the Keratograph 4 and the
329	Wave Analyzer.
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331	Figure 6. In vivo difference in mean radius (mm) between the Keratograph 4 and the
332	Wave Analyzer (solid line: mean; dashed line: 95% confidence interval).
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334	Figure 7. In vivo difference in mean eccentricity between the Keratograph 4 and the
335	Wave Analyzer (solid line: mean; dashed line: 95% confidence interval).
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337	Tables
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339	Table 1. Mean values ± standard deviations of three repeated measurements of
340	central corneal radii and corneal eccentricities, mean difference and limits of
341	agreement (LoA) of two measuring sessions (day 1 to day 2) and the mean differences
342	and 95% confidence interval between both instruments (n=20 eyes). *Indicates

343 statistically significant differences.

Table 1

	Wave Analyzer	Mean Difference (95% LoA) Day1 to Day 2	p value	Keratograph	Mean Difference (95% LoA) Day 1 to Day 2	p value	Mean Difference (95% CI) KER - WAV	p value
Central corneal radii								
Flat meridian (rc/fl)	7.82 ± 0.26	-0.01 (-0.26 to 0.25)	p=0.860	7.88 ± 0.27	+0.01 (-0.08 to 0.09)	p=0.594	-0.06 (-0.10 to -0.02)	p = 0.006*
Steep meridian (r _{c/st})	7.62 ± 0.30	+0.02 (-0.15 to 0.20)	p=0.308	7.71 ± 0.26	0.00 (-0.06 to 0.06)	p=0.783	-0.09 (-0.17 to -0.01)	p < 0.001*
Corneal eccentricity								
Nasal (enas)	0.71 ± 0.24	+0.01 (-0.36 to 0.38)	p=0.810	0.68 ± 0.11	-0.02 (-0.11 to 0.14)	p=0.469	+0.04 (-0.04 to +0.12)	p = 0.350
Temporal (et _{emp})	0.50 ± 0.39	+0.01 (-0.78 to 0.79)	p=0.340	0.43 ± 0.08	-0.01 (-0.12 to 0.11)	p=0.615	+0.07 (-0.10 to +0.23)	p = 0.014*
Inferior (e _{inf})	0.56 ± 0.19	-0.02 (-0.29 to 0.25)	p=0.496	0.51 ± 0.15	0.00 (-0.12 to 0.11)	p=0.823	+0.05 (-0.01 to +0.11)	p = 0.083
Superior (e _{sup})	0.61 ± 0.14	+0.03 (-0.77 to 0.82)	p=0.090	0.53 ± 0.13	+0.01 (-0.18 to 0.21)	p=0.402	+0.08 (+0.03 to +0.13)	p = 0.004*
Overall	0.60 ± 0.26	+0.04 (-0.50 to 0.49)	p=0.592	0.53 ± 0.15	0.00 (-0.13 to 0.12)	p=0.780	+0.06 (+0.01 to +0.11)	p = 0.009*











Average of KER and WAV







