

Accuracy of Biometric Formulae for Intraocular Lens Power Calculation in a Teaching Hospital

Running Title: Accuracy of Intraocular Lens Power Calculation

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Abstract

Aim To evaluate the accuracy of three commonly used biometric formulae across different axial lengths (ALs) at one United States Veterans Affairs teaching hospital.

Methods A retrospective chart review was conducted from November 2013 to May 2018. One eye of each patient who underwent cataract surgery with a monofocal intraocular lens (IOL) was included. The postoperative follow-up period was from 3 weeks to 4 months. The Holladay 2, Barrett Universal II, and Hill-Radial Basis Function (Hill-RBF) formulae were used to predict the postoperative refraction for all cataract surgeries. For each formula, we calculated the prediction errors (including Mean Absolute Prediction Error [MAE]) and the percentage of eyes within ± 0.25 diopter (D) and $\pm 0.5D$ of predicted refraction. We performed subgroup analyses for short (AL < 22.0 mm), medium (AL 22.0-25.0 mm), and long eyes (AL > 25.0 mm).

Results Of the 1131 patients screened, 909 met the study inclusion criteria. Resident ophthalmologists were the primary surgeons in 710 (78.1%) cases. We found no statistically significant difference in predictive accuracy among the three formulae over the entire AL range or in the short, medium, and long eye subgroups. Across the entire AL range, the Hill-RBF formula resulted in the lowest MAE (0.384D) and the highest percentage of eyes with postoperative refraction within $\pm 0.25D$ (42.7%) and $\pm 0.5D$ (75.5%) of predicted. All three formulae had the highest MAEs (>0.5D) and lowest percentage within $\pm 0.5D$ of predicted refraction (<55%) in short eyes.

Conclusion In cataract surgery patients at our teaching hospital, three commonly used biometric formulae demonstrated similar refractive accuracy across all ALs. Short eyes posed the greatest challenge to predicting postoperative refractive error.

Keywords: Cataract Surgery; Biometry; Intraocular Lens; Power Calculation

INTRODUCTION

Cataract surgery is one of the most frequently performed procedures in the Veterans Health Administration, the largest integrated health care system and the largest provider of health care training in the United States (US).¹ Advances in optical biometry and intraocular lens (IOL) power formulae have led to continued improvements in postoperative refractive outcomes^{2,3}: in 2017, 97.3% of cataract surgeries were within ± 1 diopter (D) of predicted postoperative refraction.⁴ Determining the postoperative effective lens position (ELP) and better accounting for the role of axial length (AL) remain challenges to further improvements in the accuracy of preoperative biometry.^{2,5}

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While studies done in the past five years have generally found the Barrett Universal II formula to be most accurate,^{4,6-9} the relative accuracy of different formulae is dependent on a multitude of factors, including AL,^{4,7,9,10} the type of biometry used (optical low-coherence reflectometry [OLCR] versus partial coherence interferometry [PCI]),¹¹ preoperative anterior chamber depth (ACD) values,¹² and interocular AL and corneal power differences.¹³

Evaluating biometric accuracy in a teaching hospital setting is important as this is where residents are learning their approach to patient care. In teaching hospitals, refractive outcomes may be impacted not only by resident surgeons with variable experience,^{14,15} but also different personnel who may perform biometry and refractions.¹⁶ However, the few studies published within the last five years in teaching hospitals are limited by size (<300 patients) or focus (eyes with AL>25.0 mm) (Table A, Appendix).^{2,12} The primary objective of this study was to identify the most accurate biometric formula at a single US Veterans Affairs teaching hospital. The secondary objective was to evaluate which biometric formula had the lowest prediction error in patients with short, medium, and long ALs.

SUBJECTS AND METHODS

The Providence Veterans Affairs Medical Center (PVAMC) Institutional Review Board approved this retrospective study. The Holladay 2, Barrett Universal II, and Hill-Radial Basis Function (Hill-RBF, first version) formulae were used to predict the postoperative refraction for all cataract surgeries. We did not include older formulae such as the SRK-T and Hoffer Q in our analysis as prior studies have demonstrated the superiority of current generation formulae.^{2,4,7,11} Optical biometry was performed using the Lenstar optical biometer (Haag-Streit USA, Mason, OH, USA). We included patients who received cataract surgery using monofocal spherical SN60WF IOLs at the PVAMC teaching hospital between November 2013 and May 2018. Only one eye was included from each patient to prevent compounding of data with the use of bilateral eyes; correlation between outcomes between a patient's two eyes would decrease the power of the study.¹⁷ Furthermore, as not all cataract patients at the PVAMC received bilateral surgery, including both eyes from eligible patients would have disproportionately weighted outcomes from these patients. If a patient had cataract surgery in both eyes, we included the eye with the better postoperative best corrected visual acuity (BCVA) in the study as refraction accuracy decreases with worsening BCVA.¹⁷ If both eyes had the same postoperative BCVA, we included the earlier cataract surgery.⁴ These inclusion criteria are based on recommendations by Hoffer et al¹⁷ for optimized study protocol in examining IOL formula accuracy. Patients were excluded if they had no postoperative refraction within 3 weeks to 4 months,^{4,11} AL or lens thickness (LT) not measurable by optical biometry, history of corneal disease, history of refractive surgery, posterior capsular rupture, sulcus IOL postoperative, or best corrected visual acuity (BCVA) worse than 20/40.

Information extracted from patient charts included patient age, race, ethnicity, gender, pupil size, prior cataract surgery, preoperative refraction, preoperative BCVA, postoperative refraction, postoperative BCVA, IOL type, and IOL power. Preoperative and postoperative refractive values were recorded in spherical equivalents. The preoperative biometry and the majority of the postoperative refractions were performed by experienced technicians certified by the Joint Commission on Allied Health Personnel in Ophthalmology.¹⁸

Information extracted from the Lenstar device included AL, ACD, preoperative flat corneal front power (K1), preoperative steep corneal front power (K2), LT, horizontal White-to-White (WTW) corneal diameter, and central corneal thickness (CCT). Predicted postoperative refractions from the Barrett Universal II and Hill-Radial Basis

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Function (Hill-RBF) formulae were extracted from the Haag-Streit EyeSuite software. Predictive measurements from the Holladay 2 formula were extracted from the Holladay IOL Consultant program.

We plotted overall refractive outcomes and calculated mean prediction error (ME), mean absolute prediction error (MAE), median absolute prediction error (MedAE), and the percentage of eyes with a prediction error of $\pm 0.25D$ and $\pm 0.5D$ for each formula. The MAE and MedAE provided a glimpse into the overall accuracy of each formula, while the ME showed whether each formula tends to produce more negative or positive refractive outcomes than predicted. These conventions follow those established by prior studies.^{4,7,11} Statistical comparisons of MAE among the three formulas were performed using one-way repeated measures analysis of variance (Friedman test). Subgroup analyses for short (AL<22.0 mm), medium (AL 22.0-25.0 mm), and long eyes (AL>25.0 mm) were also performed. STATA 11 (StataCorp LLC, College Station, Texas, USA) was used for statistical analysis.

RESULTS

A breakdown of patient demographics can be found in Table 1. Out of 1131 total charts reviewed in the study period, we included 909 eyes from 909 patients in the final study; 170 patients were excluded due to lack of postoperative refraction within the designated follow-up period, 33 for worse than 20/40 postoperative BCVA (27 had pre-existing ocular disease), 14 for complications, and five for missing data. Resident ophthalmologists were the primary surgeons in 78.1% (710/909) of the cases.

Overall refractive outcomes are displayed in Figure 1 and prediction error data for all AL subgroups are found in Table 2. While the Hill-RBF formula had the lowest MAE across the entire AL range, one-way analysis of variance showed no significant difference among the three formulae for monofocal IOL implantation ($F=0.37$, $p=0.69$). The Hill-RBF also predicted the highest percentage of eyes with postoperative refraction within $\pm 0.25D$ (42.5%) and $\pm 0.5D$ (75.5%) across the entire AL range.

The outcomes for short, medium, and long AL subgroups were similar: no statistically significant differences were found among the three formulae for all three subgroups ($p=0.97$, 0.75 , and 0.91 for short, medium, and long ALs, respectively). The Hill-RBF formula, however, consistently had the lowest MAE across all eye lengths. All three formulae produced their highest respective MAEs in the short AL subgroup: Holladay 2 had a MAE of 0.512D, Hill-RBF had one of 0.502D, and Barrett Universal II had one of 0.535D. The Hill-RBF and Barrett also produced the lowest percentage of eyes in the short AL subgroup with postoperative refraction within $\pm 0.25D$ and $\pm 0.5D$. Conversely, all three formulae produced their most accurate results in the medium AL subgroup: Holladay 2 had a MAE of 0.384D, Hill-RBF had one of 0.370D, and Barrett Universal II had one of 0.376D. All three formulae produced their highest percentage of eyes with postoperative refraction within $\pm 0.25D$ and $\pm 0.5D$ in the medium AL subgroup. Further graphical representations with trend lines of each formula's predictive accuracy are displayed in Figures A, B, and C in the Appendix.

Table 1. Demographics of Patients

Demographics	Percent of Total (count) N=909
Left eye	49% (444)
Female sex	2.3% (21)
Race	
Asian	0.2% (2)

Black	3.5% (32)
White	94% (855)
Other	2.2% (20)
Ethnicity	
Hispanic or Latino	0.4% (4)
Not Hispanic or Latino	99% (897)
Unknown	0.9% (8)
Axial length subgroups	
Short, <22.0 mm	1.8% (16)
Medium, 22.0-25.0 mm	84% (762)
Long, >25.0 mm	14% (125)
	Mean (SD)
Age, in years	74.5 (0.26)
Preoperative refraction [25%tile, median, 75%tile]	-2.125, -0.375, 1.125
Postoperative refraction [25%tile, median, 75%tile]	-0.5, -0.25, 0.0
IOL power	20.6 (2.8)
Anterior chamber depth, in mm	3.2 (0.43)
Lens thickness, in mm	4.61 (0.50)
Preoperative flat corneal front power, K1	43.1 (1.52)
Preoperative steep corneal front power, K2	43.8 (1.75)
Horizontal white-to-white corneal diameter, in mm	12.2 (0.51)
Central corneal thickness, in μ m	547 (38)

SD = standard deviation

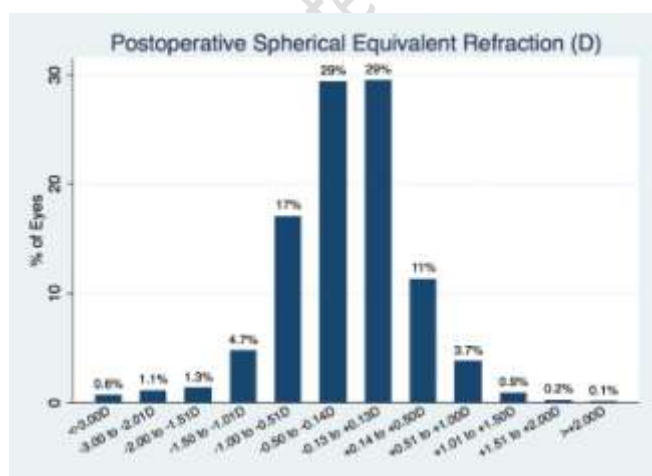


Figure 1. Bar graph of the distribution of refractive outcomes for the SN60WF model intraocular lens.

Table 2. Prediction errors in different AL groups (n=909)

Formula	MAE (D)	MedAE (D)	ME (D)	SD	Percentage of Eyes within Diopter Range Indicated	
					% ±0.25D	% ±0.5D
Entire AL range (n=909, F=0.37, p=0.69)						
Barrett 2	0.397	0.300	-0.0760	0.564	41.8%	74.1%
Holladay 2	0.399	0.310	0.0661	0.570	40.6%	71.2%
Hill-RBF	0.384	0.300	-0.0023	0.554	42.7%	75.5%
Short eyes (n=16, F=0.02, p=0.97)						
Barrett 2	0.535	0.470	0.137	0.669	26.7%	53.3%
Holladay 2	0.512	0.480	0.115	0.672	37.5%	50.0%
Hill-RBF	0.502	0.410	0.057	0.664	46.7%	53.3%
Medium eyes (n=762, F=0.28, p=0.75)						
Barrett 2	0.376	0.285	-0.060	0.517	42.8%	76.4%
Holladay 2	0.384	0.302	0.055	0.527	41.4%	72.7%
Hill-RBF	0.370	0.295	0.015	0.517	43.0%	77.6%
Long eyes (n=125, F=0.08, p=0.91)						
Barrett 2	0.507	0.355	-0.203	0.772	37.9%	62.9%
Holladay 2	0.483	0.368	0.130	0.785	36.4%	65.3%
Hill-RBF	0.474	0.335	-0.174	0.763	39.8%	62.5%

DISCUSSION

To our knowledge, this is the largest study to date comparing the accuracy of biometric formulae in cataract surgery in a teaching hospital setting. In our current sample size, we found no statistically significant difference between Holladay 2, Hill-RBF, and Barrett Universal II biometric formulae across multiple ALs. Kane et al found that the Hill-RBF formula had a significantly lower MAE than the Barrett Universal II in short eyes, and that the Hill-RBF performed better in long eyes than in medium eyes.⁸ In our analysis, the Hill-RBF retained the lowest MAE for the entire AL range, though this was not statistically significant, most likely due to the smaller size of our study relative to that of Kane et al (n=3241). In addition, we found that all three formulae produced their highest MAE in the short AL subgroup. This was consistent with previous studies that stratified predictive errors according to AL (Table A, Appendix).^{10,11}

Only two previous studies by Gokce et al^{2,12} were done in a teaching hospital, but they were both limited by their small sample sizes and focus on short eyes (Table A, Appendix). These two studies did not evaluate the accuracy of the Holladay 2 or Hill-RBF formulae, but our MAE for the Barrett Universal II was consistent with theirs.² Our MAEs for each of the formulae was also consistent with values demonstrated in previous non-teaching hospital studies (Table A, Appendix).^{4,8,11} While this is not a one-to-one comparison between resident and attending surgical outcomes, it is a realistic representation of the differences between teaching (where a percentage of cases will still be performed by attendings) and non-teaching hospital settings.

This study has several limitations. First, we targeted patients receiving care in the VHA; hence, our findings may not be generalizable to patients receiving cataract surgery elsewhere, including other teaching hospitals.¹⁹ Second,

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our sample size may have precluded achieving statistically significant differences among the three biometric formulae. However, our findings confirm that the overall accuracy of biometric formulae in predicting refractive outcomes are comparable between teaching and non-teaching hospital settings. Third, we excluded 18.3% of patients due to lack of postoperative refractive follow-up within the designated timeframe. Some patients may have followed up with providers outside of the PVAMC, but others may have neglected to come to follow-up appointments due to satisfactory postoperative visual outcomes; this may have resulted in selection bias toward patients with worse refractive outcomes.^{4,20}

Conclusions

This study found no difference in the accuracy of the Holladay 2, Hill-RBF, and Barrett Universal II formulae for cataract surgery in a US teaching hospital, although all three formulae were least accurate in short eyes.

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Conflicts of Interest: **Tang K:** none, **Tran E:** none, **Chen A:** none, **Rivera D:** none, **Rivera J:** none, **Greenberg P:** none.

Disclaimer: The views expressed in this article are those of the authors and do not necessarily reflect the position or policy of the United States Department of Veterans Affairs or the United States government.

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Appendix

Table 1. Previous studies comparing biometric formulae

Study	No. of eyes	Formulae	MAE (D)				Conclusions
			Overall	Short AL	Medium AL	Long AL	
Melles et al, 2017 ⁴	13,301	Barrett 2	0.311	-	-	-	Barrett was most consistently accurate in different AL groups
		Holladay 2	0.450	-	-	-	
Kane et al, 2017 ⁸	3,122	Barrett 2	0.381	0.451	0.383	0.375	Hill-RBF was more accurate than Barrett in short AL group; the Barrett was more accurate overall and in medium ALs
		Holladay 2	0.410	-	-	-	
		Hill-RBF	0.407	0.423	0.412	0.373	
Cooke et al, 2016 ¹¹	1,454	Barrett 2	0.306	0.338	-	0.274	All formulae were least accurate for short eyes
		Holladay	0.346	0.426	-	0.394	

		2					
Gokce et al, 2017 ² *	86	Barrett 2	-	0.39	-	-	Only short eyes were analyzed; no statistically significant difference in formula accuracy
		Holladay 2	-	0.40	-	-	
		Hill-RBF	-	0.36	-	-	
Gokce et al, 2018 ¹² *	270	Barrett 2	0.29	-	-	-	Compared formulae accuracy for patient groups with varying ACD; Barrett had lowest MAE for ACD<3.0 mm and ACD>3.5 mm
		Holladay 2	0.31	-	-	-	
		Hill-RBF	0.28	-	-	-	
Carifi et al, 2015 ²¹	28	Holladay 2	-	0.82	-	-	Only short eyes; no difference between formulae, but all with large MAE
Kane et al, 2016 ⁷	3,241	Barrett 2	0.385	0.469	0.386	0.435	All formulae were less accurate in short AL group; Barrett was most accurate for all other ALs
		Holladay 2	0.420	0.466	0.416	0.544	

MAE=Mean absolute prediction error; ACD=Anterior Chamber Depth; AL=Axial Length; D=Diopters

*Study performed in a teaching hospital

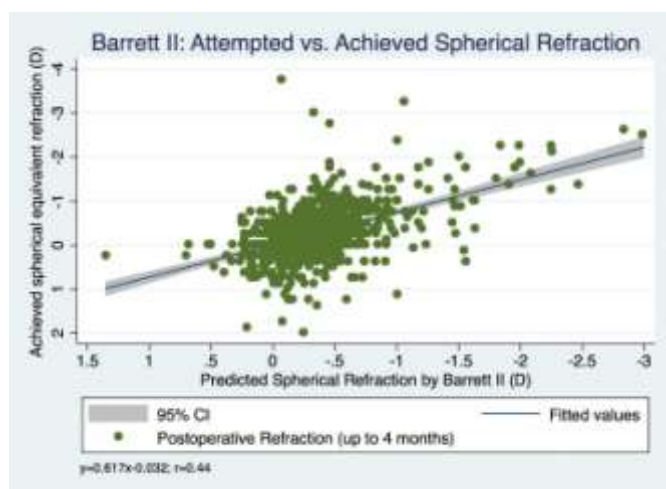


Figure A. Scatterplot with trend line of attempted vs. achieved spherical refraction for SN60WF model intraocular lens implantation using the Barrett II biometric formula.

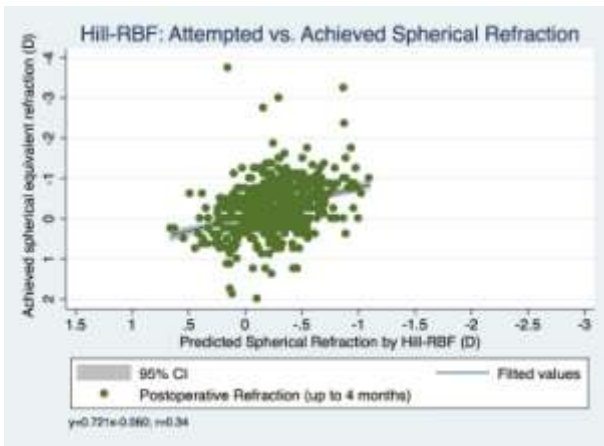


Figure B. Scatterplot with trend line of attempted vs. achieved spherical refraction for SN60WF model intraocular lens implantation using the Hill-RBF biometric formula.

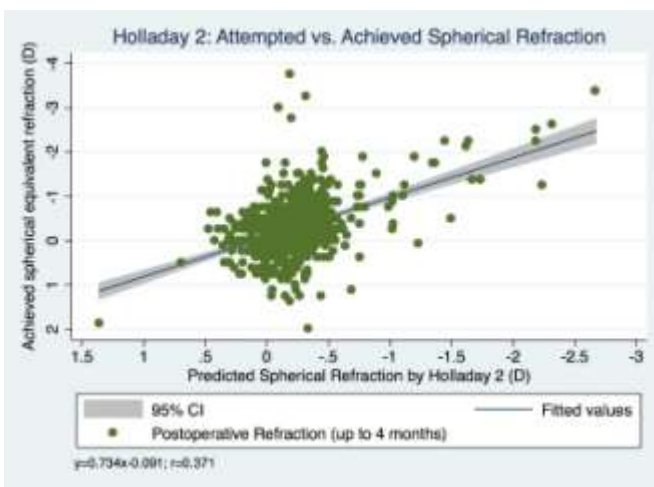


Figure C. Scatterplot with trend line of attempted vs. achieved spherical refraction for SN60WF model intraocular lens implantation using the Holladay 2 biometric formula.