



СРПСКИ АРХИВ
ЗА ЦЕЛОКУПНО ЛЕКАРСТВО
SERBIAN ARCHIVES
OF MEDICINE

Address: 1 Kraljice Natalije Street, Belgrade 11000, Serbia

+381 11 4092 776, Fax: +381 11 3348 653

E-mail: office@srpskiarhiv.rs, Web address: www.srpskiarhiv.rs

Paper Accepted*

ISSN Online 2406-0895

Original Article / Оригинални рад

Miroslav Stamenković^{1,2,*}, Ivan Marjanović^{3,4}, Vesna Marić^{3,4}, Tanja Kalezić^{3,4},
Marija Božić^{3,4}

**Intraocular pressure and central corneal thickness
in a healthy student population**

Висина интраокуларног притиска и централна дебљина роњаче
код здраве студентске популације

¹Zvezdara University Medical Center, Clinic for Eye Diseases, Belgrade, Serbia;

²University of Belgrade, Faculty of Special Education and Rehabilitation, Belgrade, Serbia;

³University of Belgrade Faculty of Medicine, Belgrade, Serbia;

⁴University Clinical Center of Serbia, University Eye Hospital, Belgrade, Serbia

Received: October 8, 2023

Accepted: February 10, 2024

Online First: February 13, 2024

DOI: <https://doi.org/10.2298/SARH231008016S>

***Accepted papers** are articles in press that have gone through due peer review process and have been accepted for publication by the Editorial Board of the *Serbian Archives of Medicine*. They have not yet been copy-edited and/or formatted in the publication house style, and the text may be changed before the final publication.

Although accepted papers do not yet have all the accompanying bibliographic details available, they can already be cited using the year of online publication and the DOI, as follows: the author's last name and initial of the first name, article title, journal title, online first publication month and year, and the DOI; e.g.: Petrović P, Jovanović J. The title of the article. *Srp Arh Celok Lek*. Online First, February 2017.

When the final article is assigned to volumes/issues of the journal, the Article in Press version will be removed and the final version will appear in the associated published volumes/issues of the journal. The date the article was made available online first will be carried over.

***Correspondence to:**

Miroslav STAMENKOVIĆ

Zvezdara University Medical Center, Clinic for Eye Diseases, Dimitrija Tucovica 161, 11000 Belgrade, Serbia

E-mail: drmiroslavstamenkovic@gmail.com

Intraocular pressure and central corneal thickness in a healthy student population

Висина интраокуларног притиска и централна дебљина рожњаче
код здраве студентске популације

SUMMARY

Introduction/Objective Intraocular pressure is an important parameter of eye health, especially when glaucoma is suspected. So far, few studies have been published that aimed to determine the average value of intraocular pressure and central corneal thickness in a healthy population aged 20–30 years. The aim of this study was to determine the distribution of the values of intraocular pressure and central corneal thickness in healthy student population.

Methods In a cross-sectional study, intraocular pressure and central corneal thickness were measured on a sample of a healthy population, aged 22–37 years. Intraocular pressure was measured using the Goldmann applanation tonometry method, while central corneal thickness was measured using ultrasound pachymetry. The analysis of numerical values was done using the methods of descriptive statistics.

Results By measuring intraocular pressure and central corneal thickness in 641 subjects (1282 eyes), the average value of intraocular pressure was determined to be 14.79 ± 2.31 mmHg, and central corneal thickness was 553.92 ± 25.56 μ m. By comparing two groups of subjects, one group was male, and the other female, we determined that there was no statistically significant difference in the average value of intraocular pressure (t-test, $p > 0.05$), and the average value of central corneal thickness (t-test, $p > 0.05$) between the sexes.

Conclusion The determined average value of intraocular pressure and central corneal thickness is similar to those determined in other cross-sectional studies of this type. No statistically significant difference was found in the intraocular pressure values and the central thickness of the cornea by sex.

Keywords: intraocular pressure; central corneal thickness; students

САЖЕТАК

Увод/Циљ Интраокуларни притисак је значајан параметар здравља ока, а посебно када постоји сумња на глауком. Ретке су до сада објављене студије које су имале за циљ утврђивање просечне вредности интраокуларног притиска и централне дебљине рожњаче на здравој популацији старости 20–30 година. Циљ ове студије је утврђивање дистрибуције вредности интраокуларног притиска и централне дебљине рожњаче код особа здраве студентске популације.

Методе У студији пресека вршено је мерење вредности интраокуларног притиска и централне дебљине рожњаче на узорку здраве популације, старости 22–37 година. Мерење интраокуларног притиска вршено је методом Голдманове апланационе тонометрије, док је мерење централне дебљине рожњаче вршено ултразвучном пахиметријом. Анализа нумеричких вредности рађена је методама описне статистике.

Резултати Мерењем интраокуларног притиска и централне дебљине рожњаче на 641 испитанику (1282 ока), утврђена је просечна вредност интраокуларног притиска од 14.79 ± 2.31 mmHg, и централне дебљине рожњаче од 553.92 ± 25.56 μ m. Поређењем две групе испитаника, од којих је једна група била мушког пола, а друга женског, утврђено је да нема статистички значајне разлике у просечној вредности интраокуларног притиска (t-тест, $p > 0.05$), и просечној вредности централне дебљине рожњаче (t-тест, $p > 0.05$) између полова.

Закључак Утврђена просечна вредност интраокуларног притиска и централне дебљине рожњаче је слична онима утврђеним у осталим студијама пресека овог типа. Није утврђена статистички значајна разлика у висини притиска и централне дебљине рожњаче поређењем по полу.

Кључне речи: интраокуларни притисак; централна дебљина рожњаче; студенти

INTRODUCTION

Intraocular pressure (IOP) is one of the most important parameters of eye health. Its values represent the result of the dynamic balance of aqueous humor production and outflow. Elevated IOP is the most significant risk factor for glaucoma, and factor for the conversion of ocular hypertension to primary open-angle glaucoma [1, 2]. IOP is routinely measured for diagnosis and monitoring of glaucoma suspects and patients [3]. All this indicates the great

importance of determining the correct IOP values. Goldmann applanation tonometry (GAT) is the gold standard technique for measuring IOP. However, the accuracy of the results obtained by this procedure can be affected by several factors [4, 5], the most significant of which is central corneal thickness (CCT). In general, a thinner cornea leads to a lower IOP reading, while a thicker cornea leads to a higher IOP reading than their actual values [6,7].

Statistically, an IOP value of 21 mm Hg is widely accepted as the borderline between normal and elevated. When calibrating the Goldmann tonometer, Goldmann and Schmidt [8] assumed a CCT of 0.5 mm and emphasized that variations in corneal thickness could, in theory, affect the measurement. Information on differences in CCT values obtained through in vivo measurements subsequently became available [9].

CCT can be measured by different methods, but ultrasound pachymetry is considered more reliable [10]. High precision in measuring CCT values is desirable since it is as an indicator corneal endothelium health [11]. Finally, the association between decreased CCT values and readings of apparently decreased IOP values has prompted research into the role of CCT measurements in the early diagnosis of glaucoma [12, 13]. Most of the studies on central corneal thickness were performed on the population suffering from glaucoma or other ophthalmic diseases.

There are not many studies that have dealt with normal IOP values in healthy young population. The aim of the present study was to investigate IOP and CCT values in the healthy population of aged 20-40 years.

METHODS

In this cross-sectional, population-based, observational study, 641 students of the Faculty of Medicine, University of Belgrade of both sexes, aged 22–37, participated. This study was conducted according to the principles of the Helsinki declaration and we had the consent of the Ethics Committee of the Faculty of Medicine, Belgrade. All subjects were informed about the test methods before the measurement, and written informed consent was obtained. All subjects underwent a complete ophthalmic examination consisting of a medical history, best corrected visual acuity (BCVA), slit-lamp biomicroscopy (Haag-Streit, Switzerland), Goldmann applanation tonometry (GAT; Haag Streit AG, Bern, Switzerland), funduscopy, central corneal thickness. Exclusion criteria were as follows: any form of glaucoma or systemic disease that might influence IOP values, previous intraocular surgery or trauma, pregnancy, allergy to tetracaine.

Goldmann tonometer, slit lamp mounted (Haag-Streit, Switzerland) was used for Goldmann applanation tonometry (GAT). Tetracaine 1% and Fluorescein sodium 2% strips were used for the GAT measurements. All GAT measurements were done during morning hours (9-11h) of the day, in sitting position. The mean IOP and CCT value was obtained from three consecutive measurements. Palm Scan AP 2000, Ophthalmic Ultrasound, 2007, Micro Medical Devices Inc., Calabasas, CA, 91302 USA was used for central corneal thickness measurements after instillation of 1.0% tetracaine, and the mean of three readings was calculated for each tested eye.

The analysis of numerical values was done using classic methods of descriptive statistics, χ^2 test (for data analysis within groups) and t-test (for analysis between groups), arithmetic mean, median of mean values, and measures of variability with standard deviation, coefficient of variation and standard error, as well as the minimum and maximum value. A value of $p < 0.05$ was considered statistically significant.

RESULTS

The examination was performed on a sample of a healthy student population of 641 subjects (1282 eyes). The average age of the respondents was 24.41 ± 0.99 years.

The determined average values of IOP and CCT are listed in the Table 1.

By comparing two groups of subjects, one of which was male (227 subjects, 454 eyes) and the other female (414 subjects, 828 eyes), it was determined that there was no statistically significant difference in the average value of intraocular pressure (student's t test, $p > 0.05$), and the average value of central corneal thickness (student's t test, $p > 0.05$) between sexes (Table 2).

Analysis of the average values of IOP and CCT of the right and left eyes revealed no statistically significant differences ($p > 0.05$) (Table 3).

DISCUSSION

The role of IOP and its connection with glaucoma has been the focus of scientific research practically since the first definition of glaucoma as an eye disease. While this definition of glaucoma currently rests more on structural and functional damage [14], IOP measurement is still used as a mandatory, simple, accessible and economical method in approaching high-risk patients. Many studies have documented an association of increased incidence of glaucoma

with increasing IOP values [15, 16], and especially with values above 20-23 mmHg [17, 18], however, a study in a Latino population found this association with IOP values above 30 mmHg [19]. There are numerous data in the literature for the average statistical normal value of IOP, but few studies have addressed this question in different age groups, especially in the age group of 20-30 years [20]. In the study of Dane and al, which was done on 125 subjects, finding of higher intraocular pressure in women was explained by oestrogen effects. Some of published studies aimed to study IOP daily fluctuations in young people or the influence of sleeping position on IOP values, but all of them are characterized by a small number of subjects (10 or 20) [21, 22].

Normal IOP ranges between 10 and 22 mmHg, with an average of 16 mmHg. Values for normal IOP have been obtained by examining large population groups. One of the largest studies was conducted on the population in Serbia in 1970, when Prof. Cvetković et al examined 3,550 people of both sexes over 40 years of age in the municipality of Opovo [23]. Measurements were made with a Schiotz impression tonometer, and mean IOP values of 16.85 ± 3.0 mmHg were obtained. There was no statistically significant difference in the IOP level according to gender (although the IOP in women was slightly higher, 17.0 mmHg, compared to men, where it was 16.7 mmHg). As part of the above-mentioned project, part of the examination was performed using the applanation tonometry method, but on a smaller sample (512 subjects of both sexes), with very similar results - the average IOP value was 16.47 ± 3.0 mmHg. The mean IOP value measured in this study (15.11 ± 2.35 mmHg) corresponds to those recorded in earlier studies of this type involving healthy Caucasian subjects of approximately the same age [24]. One of the larger studies made in the territory of the Republic of Serbia was conducted in the period from 2007 to 2012 in the territory of the city of Novi Sad, but on the population of people who are being treated for glaucoma in the ophthalmology services of primary health care centers [25].

IOP is a dynamic parameter that changes depending on heart action (systole/diastole), inspiratory/expiratory pressure, extraocular muscle tone, hormonal status, daily rhythm of vagotonia and sympathicotonia, body position, and is even related to the seasons. Also, IOP is known to change with age. In newborn children and infants, and during the entire first decade of life, lower IOP values than those determined for the adult population are considered to be normal [26]. In children in the first years of life, the average normal IOP is below 15 mmHg, from the age of 6 to 12 years it is 11 ± 2.5 mmHg [27], and in the decades after the 50s, the average IOP value gradually increases, but without statistically significant differences.

The IOP in the right and left eye of the same person is practically the same, and 3 mmHg is accepted as a normal difference. When measuring, it is usual to measure the IOP first in the right eye, then in the left eye, and it is noted that repeated measurement in the right eye usually gives lower values. Probably one of the reasons is the relaxation of the extraocular muscles during the repeated measurement, or the discrete opening of the chamber angle due to the pressure of the prism on the cornea. In our study, it was found that there is no significant difference between the average IOP value between the right and left eyes, which agrees with the results of earlier studies [28].

As for gender differences, it was found that women have slightly higher IOP on average, but without statistical significance. In our study, the determined average value of IOP in female subjects is 15.23 ± 0.43 mmHg, while in male subjects it is 14.89 ± 0.52 mmHg, with a difference that is not statistically significant ($p > 0.05$).

CCT is routinely measured in clinical settings before corneal refractive procedures, but also because it can potentially significantly influence the reading of real IOP values and consequently the classification and therapy of glaucoma.

The average CCT value measured in this study (563.65 ± 27.74 μ m) confirms the values documented in earlier studies performed on a similar sample [11, 29, 30]. In earlier clinical studies, the average value of CCT varied from 520 μ m when CCT was determined by optical pachymetry [9, 29] to 540 μ m when determined by ultrasound [30]. By comparing the average CCT values between the sexes, we found that there is no statistically significant difference in the CCT value in the healthy population sample, which confirms previous studies [9], although the average value was slightly higher in female subjects.

In this study, the average value of CCT between the right and left eyes was determined, which was statistically not significantly different. Previous studies with optical pachymetry [30] have shown that there is a systematic difference between the right and left eyes. This may be due to measurement error in the optical method when the measurement is not positioned normal to the cornea. Such measurement errors do not occur when using an ultrasonic pachymeter because it reads a value only when the probe is directed normally to the cornea. Indeed, other studies using ultrasound pachymetry also found no statistically significant difference between the right and left eyes [30].

CONCLUSION

In this study, we determined the average values of IOP and CCT in a healthy student population, that is, the age group from 22 to 37 years old. So far, similar studies have not been done in our population. The average values of IOP and CCT in our sample did not differ significantly from the values obtained in similar previously published studies.

Conflict of interest: None declared.

Paper accepted

REFERENCES

1. Kontić Đ. Tonometry. In: Cvetkovic D, Kontić Đ, Hentova-Senčanić P, editors. *Glaucoma*. Belgrade: Zavod za udžbenike i nastavna sredstva; 1996. p. 43–71.
2. Why measure central corneal thickness to confirm a diagnosis of glaucoma? *Cont Lens Anterior Eye*. 2022 Dec;45(6):101777. doi: 10.1016/j.clae.2022.101777. Epub 2022 Nov 3.
3. Belovay GW, Goldberg I. The thick and thin of the central corneal thickness in glaucoma. *Eye (Lond)*. 2018 May;32(5):915-923. [DOI: 10.1038/s41433-018-0033-3] [PMID: 29445115]
4. Rüfer F. Aktueller Stellenwert der Druckmessung: Messverfahren und Fehlerquellen [Value of Pressure Measurements: Methods and Sources of Errors]. *Klin Monbl Augenheilkd*. 2016 Jul;233(7):847-55. German.[DOI: 10.1055/s-0042-101552] [PMID:27130978]
5. Zeppieri M, Gurnani B. Applanation Tonometry. 2023 Jun 11. In: StatPearls [Internet]. Treasure Island (FL): StatPearls Publishing; 2023 Jan.
6. Rewri P, Ali W. Erroneous assumption of ocular hypertension in patients with elevated intraocular pressure. *Indian J Ophthalmol*. 2022 Feb;70(2):564-568. [DOI:10.4103/ijo.IJO_938_21] [PMID: 35086238]
7. Badr M, Masis Solano M, Amoozgar B, Nguyen A, Porco T, Lin S. Central Corneal Thickness Variances Among Different Asian Ethnicities in Glaucoma and Nonglaucoma Patients. *J Glaucoma*. 2019 Mar;28(3):223-230. [DOI: 10.1097/IJG.0000000000001180] [PMID: 30624387]
8. De Bernardo M, Casaburi C, De Pascale I, Capasso L, Cione F, Rosa N. Comparison between dynamic contour tonometry and Goldmann applanation tonometry correcting equations. *Sci Rep*. 2022 Nov 23;12(1):20190. [DOI: 10.1038/s41598-022-24318-y] [PMID: 36418360]
9. Realini T, Gupta PK, Radcliffe NM, Garg S, Wiley WF, Yeu E, Berdahl JP, Kahook MY. The Effects of Glaucoma and Glaucoma Therapies on Corneal Endothelial Cell Density. *J Glaucoma*. 2021 Mar 1;30(3):209-218.[DOI: 10.1097/IJG.0000000000001722.] [PMID: 33105305]
10. Doğan M, Ertan E. Comparison of central corneal thickness measurements with standard ultrasonic pachymetry and optical devices. *Clin Exp Optom*. 2019 Mar;102(2):126-130. [DOI: 10.1111/cxo.12865][PMID: 30557910]
11. Pluháček F, Unzeitigová A, Marešová K, Rybář J. Influence of cornea on intraocular pressure measurement by ICARE PRO and ORA. *Cesk Slov Oftalmol*. 2019;75(3):111-118. [DOI: 10.31348/2019/3/1] [PMID: 31779459]
12. Özkan Aksoy N, Çakır B, Aksoy YE, Demir Boncuğu K, Özmen S, Çelik E, Alagöz G. Effects of glaucoma and central corneal thickness on optic nerve head biomechanics. *Int Ophthalmol*. 2021. 41(4):1283-1289. [DOI: 10.1007/s10792-020-01686-w] [PMID: 33387111]
13. Brandt JD, Beiser JA, Kass MA, Gordon MO; Ocular Hypertension Treatment Study (OHTS) Group. Central Corneal Thickness in the Ocular Hypertension Treatment Study (OHTS). *Ophthalmology*. 2020 Apr;127(4S):S72-S81. [DOI: 10.1016/j.opht.2020.01.028] [PMID: 32200829]
14. Foster PJ, Buhrmann R, Quigley HA, Johnson GJ. The definition and classification of glaucoma in prevalence surveys. *Br J Ophthalmol*. 2002 Feb;86(2):238-42. [DOI: 10.1136/bjo.86.2.238][PMID: 11815354]
15. Schuster AK, Wagner FM, Pfeiffer N, Hoffmann EM. Risk factors for open-angle glaucoma and recommendations for glaucoma screening. *Ophthalmologe*. 2021 Jul;118(Suppl 2):145-152. English. [DOI: 10.1007/s00347-021-01378-5][PMID: 33881589]
16. Bertaud S, Aragno V, Baudouin C, Labbé A. Le glaucome primitif à angle ouvert [Primary open-angle glaucoma]. *Rev Med Interne*. 2019 Jul;40(7):445-452. French. [DOI: 10.1016/j.revmed.2018.12.001] [PMID: 30594326]

17. Singh A, Gale J, Cheyne K, Ambler A, Poulton R, Wilson G. The prevalence of glaucoma among 45-year-old New Zealanders. *N Z Med J*. 2022 Apr 14;135(1553):35-42.
18. Selvan H, Gupta S, Wiggs JL, Gupta V. Juvenile-onset open-angle glaucoma - A clinical and genetic update. *Surv Ophthalmol*. 2022 Jul-Aug;67(4):1099-1117. [DOI: 10.1016/j.survophthal.2021.09.001][PMID: 34536459]
19. Tashtitova L, Aldasheva N. Study of the Prevalence of Glaucoma in Kazakhstan. *Klin Monbl Augenheilkd*. 2022 Feb;239(2):202-207. English. [DOI: 10.1055/a-1327-3999] [PMID: 33853192]
20. Kuo RN, Yang CC, Yen AM, Liu TY, Lin MW, Chen SL. Gender Difference in Intraocular Pressure and Incidence of Metabolic Syndrome: A Community-Based Cohort Study in Matsu, Taiwan. *Metab Syndr Relat Disord*. 2019 Jul/Aug;17(6):334-340. [DOI: 10.1089/met.2018.0131] [PMID: 31188053]
21. Vera J, Redondo B, Perez-Castilla A, Jiménez R, García-Ramos A. Intraocular pressure increases during dynamic resistance training exercises according to the exercise phase in healthy young adults. *Graefes Arch Clin Exp Ophthalmol*. 2020 Aug;258(8):1795-1801. [DOI: 10.1007/s00417-020-04736-2] [PMID: 32405701]
22. Nelson ES, Myers JG Jr, Lewandowski BE, Ethier CR, Samuels BC. Acute effects of posture on intraocular pressure. *PLoS One*. 2020 Feb 6;15(2):e0226915. [DOI: 10.1371/journal.pone.0226915] [PMID: 32027692]
23. Cvetkovic D, Blagojevic M, Stefanovic B, Brasic B, Kostadinovic D. Distribucija nalaza intraokularnog pritiska kod naseg stanovnistva. *Acta Ophthalmol Jug* 1971; 9:46-51
24. N.Babić, A.Miljković, S. Davidović, S. Barišić, V.Čanadanović. Prevalence of glaucoma in the city of Novi Sad *Srp Arh Celok Lek*. 2022 Sep-Oct;150(9-10):558-563. [DOI: <https://doi.org/10.2298/SARH211005078B>]
25. 25.Civcic-Drljaca N. Gonioskopske promjene komornog ugla sa starenjem organizma, magistarska teza. Medicinski fakultet Beograd 1990.
26. Samy E, El Sayed Y, Awadein A, Gamil M. Effect of general inhalational anesthesia on intraocular pressure measurements in normal and glaucomatous children. *Int Ophthalmol*. 2021 Jul;41(7):2455-2463. [DOI: 10.1007/s10792-021-01800-6][PMID: 33759070]
27. Melchior B, De Moraes CG, Paula JS, A Cioffi G, Girkin CA, Fazio MA, N Weinreb R, M Zangwill L, M Liebmann J. Relationship between mean follow-up intraocular pressure, rates of visual field progression and current target intraocular pressure guidelines. *Br J Ophthalmol*. 2022 Feb;106(2):229-233. [DOI: 10.1136/bjophthalmol-2020-317406][PMID: 33130556]
28. Han F, Li J, Zhao X, Li X, Wei P, Wang Y. Distribution and analysis of intraocular pressure and its possible association with glaucoma in children. *Int Ophthalmol*. 2021 Aug;41(8):2817-2825. [DOI: 10.1007/s10792-021-01838-6][PMID: 33842987]
29. Schuster AK, Fischer JE, Vossmerbaeumer U. Central Corneal Thickness in Spectral-Domain OCT and Associations with Ocular and Systemic Parameters. *J Ophthalmol*. 2016;2016:2596956. [DOI: 10.1155/2016/2596956][PMID: 27340561]
30. Lešták J, Pitrová Š, Nutterová E. Changes of Central Corneal Thickness in Normotensive and Hypertensive Glaucoma. *Cesk Slov Oftalmol*. 2019 Spring;74(5):186-189. English. [DOI: 10.31348/2018/5/3] [PMID: 31234631]

Table 1. Average intraocular pressure and central corneal thickness values in tested students

	IOP	CCT
\bar{x}	14.79 \pm 2.31 mmHg	553.92 \pm 25.56 μ m
range	10–24 mmHg	470–697 μ m

IOP – intraocular pressure; CCT – central corneal thickness

Table 2. Average intraocular pressure and central corneal thickness by sex

	female	male
Number of subjects/%	414 (828 eyes)/62.81	227 (454 eyes)/37.19
IOP (mmHg)	14.69 ± 0.41	14.932 ± 0.48
CCT (μm)	553.39 ± 4.13	554.99 ± 7.44

IOP – intraocular pressure; CCT – central corneal thickness

Table 3. average intraocular pressure and central corneal thickness by right/left eye

	Right eye	Left eye	p
IOP (mmHg)	15.13 ± 0.48	15.9 ± 0.45	0.9
CCT (μm)	563.64 ± 5.82	563.23 ± 5.23	0.99

IOP – intraocular pressure; CCT – central corneal thickness