doi 10.34172/aim.28831

Original Article

Open Access

Prevalence of Certain Corneal Conditions and their Demographic Risk Factors; Tehran Geriatric Eye Study



Alireza Hashemi¹⁰, Hassan Hashemi^{2*0}, Mohamadreza Aghamirsalim³, Alireza Jamali², Mehdi Khabazkhoob⁴

¹Noor Research Center for Ophthalmic Epidemiology, Noor Eye Hospital, Tehran, Iran

²Noor Ophthalmology Research Center, Noor Eye Hospital, Tehran, Iran

³Eye Research Center, Tehran University of Medical Sciences, Tehran, Iran

⁴Department of Medical Surgical Nursing, School of Nursing and Midwifery, Shahid Beheshti University of Medical Sciences, Tehran, Iran

Abstract

Background: Corneal abnormalities are one of the important reasons for visual impairment. There is little evidence of the prevalence of different types of corneal abnormalities. The aim of this study was to assess the prevalence of various corneal abnormalities and identify the key risk factors associated with these abnormalities in an elderly population residing in Tehran.

Methods: The Tehran Geriatric Eye Study (TGES) was conducted as a cross-sectional study, utilizing a population-based approach and employing stratified cluster random sampling. The study focused on individuals aged 60 years and above residing in Tehran. An ophthalmologist performed a slit lamp examination to evaluate the eyelid, cornea, and crystalline lens.

Results: The prevalence of posterior embryotoxon (PE), punctate epithelial defect (PED), pigment on endothelium (POE), corneal dystrophy (CDys), corneal vascularization (CV), and corneal degeneration (CDeg) were estimated to be 0.08% (95% confidence interval [CI]: 0.02 to 0.40), 8.77% (95% CI: 6.64 to 11.51), 0.57% (95% CI: 0.33 to 0.98), 0.53% (95% CI: 0.33 to 0.82), 0.95% (95% CI: 0.60 to 1.52), and 44.87% (95% CI: 41.80 to 47.98), respectively. Overall, approximately 49.08% of the participants exhibited some form of corneal abnormality in at least one eye. The multiple logistic regression model revealed that increasing age was significantly associated with PED, CV, and CD. Furthermore, illiterate participants had a significantly higher prevalence of PE. **Conclusion:** The findings of this study indicate that approximately half of the elderly population aged 60 years and above in Tehran have at least one corneal abnormality, with corneal degeneration being the most prevalent. Age was identified as the primary determinant of corneal abnormalities.

Keywords: Corneal abnormalities, Corneal degeneration, Epithelial defect, Geriatric, Population-based, Posterior embryotoxon, Prevalence

Cite this article as: Hashemi A, Hashemi H, Aghamirsalim M, Jamali A, Khabazkhoob M. Prevalence of certain corneal conditions and their demographic risk factors; Tehran Geriatric eye study. Arch Iran Med. 2024;27(8):414-420. doi: 10.34172/aim.28831

Received: January 20, 2024, Accepted: June 12, 2024, ePublished: August 1, 2024

Introduction

Corneal abnormalities are a significant contributor to visual impairment and rank as the fifth leading cause of blindness globally, following cataracts, refractive errors, glaucoma, and age-related macular degeneration.¹⁻³ Estimates suggest that corneal abnormalities are responsible for 8%-25% of the cases of blindness in developed countries; this is while the major burden of corneal abnormalities is related to developing countries.² Due to the effects of corneal abnormalities on vision⁴ and its high prevalence, 184576 corneal transplants were done across the world in 2012.5 The importance of corneal abnormalities is due to their outcomes; for example, corneal vascularization (CV) results in opacity in the vision path and causes significant visual impairment which sometimes requires corneal transplantation in advanced stages.^{6,7} Furthermore, some corneal abnormalities like posterior embryotoxon (PE) are associated with increased intraocular pressure and the risk of open-angle glaucoma.8,9

The epidemiology of corneal pathologies varies in different studies.¹⁰ Moreover, there is little evidence on the

prevalence of different types of corneal abnormalities; for example, studies have reported prevalence rates of 1.68%¹¹ and 27.2%¹² for corneal opacity and a prevalence of 0.3%¹³ and 0.2%14 for corneal dystrophy (CDys) in Iranian and Indian elderly populations, respectively. Also, 897 per one million people were observed to have CDys in the USA.15 This variety in the results of different studies is due to different factors including causative factors like age, gender, education level, availability and general standards of eye care, and study population.^{3,7,10} A literature search only found one population-based study addressing the prevalence of different corneal abnormalities in an Iranian elderly population,¹³ although other studies have been conducted to evaluate some corneal abnormalities individually.¹⁶ Therefore, more epidemiological studies are required in this regard in different age groups and in different societies for proper health policymaking.

According to projections, the world's elderly population has expanded significantly due to the improvement in life expectancy. Currently, 11% of the global population is aged 60 years or above, and this percentage is expected

to double by 2050, reaching 22%. Iran, as a developing country, exhibits a population aging trend that mirrors the global pattern.¹⁷ Projections indicate that by the year 2050, the elderly population in Iran is expected to reach 20 million, constituting approximately 22% of the total population of the country.¹⁸ Since most ocular diseases are attributable to poor knowledge of eye health and poor access to eye health care services in this age group, it is of great importance to determine the prevalence of vision-related abnormalities in this population. A few epidemiological studies have evaluated the prevalence of corneal abnormalities worldwide.15,19 The limitation of studies on the prevalence of corneal abnormalities in the elderly population of Iran and the need to examine this issue at different ages along with expectations for an increase in the prevalence of eye diseases due to population aging and lifestyle modification encouraged us to conduct this study to determine the prevalence of corneal abnormalities and their most important risk factors in an Iranian elderly population.

Materials and Methods

Design and Sampling

The data presented in this report is derived from the Tehran Geriatric Eye Study (TGES), a cross-sectional study carried out in 2019 on individuals aged 60 and above in Tehran, Iran. The study utilized a multi-stage stratified random cluster sampling method. The primary objective of the study was to examine the prevalence of visual impairment among the elderly population in Tehran. To achieve this goal, the sample size was determined based on a 5.2% prevalence rate of visual impairment, with a desired accuracy of 1% and a 95% confidence level. Initially, the sample size was calculated to be 1,894 individuals. However, after accounting for a 1.5 design effect and a 10% non-response rate, the final sample size was adjusted to 3,155 individuals, which was rounded up to 3200 participants.

In the TGES, a total of 160 clusters were chosen in a random manner, with the selection being proportional to the size of each cluster. These clusters were selected from 22 different strata in Tehran. Once each cluster was determined, a sampling team was dispatched to the corresponding address. The first house located on the southwest side of the chosen block was designated as the cluster head. Subsequently, the neighboring households were selected in a counterclockwise direction. All individuals aged 60 years and above were then invited to take part in the study after being provided with a clear explanation of the study's objectives and receiving assurance regarding the confidentiality of their data. If an individual expressed their willingness to participate, informed consent was obtained, an identification card was issued, and the person was subsequently transported to Noor Eye Hospital for further examinations. In cases where a household was found to be absent during the initial visit, a follow-up visit was scheduled, preferably in

the afternoon on the same day.

Trained research assistants in the hospital gathered comprehensive demographic, anthropometric, and socioeconomic (SES) information from the participants. Subsequently, the participants underwent optometric and ophthalmologic evaluations. The optometric assessments commenced with refraction utilizing the Nidek ARK-510A auto-refractometer/keratometers (Nidek Co. LTD, Aichi, Japan). Following this, uncorrected (UCVA) and best-corrected (BCVA) visual acuities were assessed using the Smart LC 13 LED visual chart (Medizs Inc., Korea) at a distance of 6 meters. Subsequent examinations of the anterior and posterior ocular segments were conducted utilizing the B900 slit-lamp (Haag-Streit AG, Bern, Switzerland) and a + 90 D lens. Individuals with a history of corneal injury due to trauma or corneal transplant were excluded from the study.

Definition of Corneal Abnormalities

To diagnose corneal abnormalities, the ophthalmologists of this study were first trained to define corneal conditions based on a uniform pattern, and all of the abnormalities were defined based on standard definitions.

Posterior embryotoxon (PE): In this abnormality, evaluation of the posterior peripheral cornea under direct light of the slit-lamp shows a peripheral corneal ring at the margin of Descemet's membrane displaced closer to the corneal center.²⁰

Punctate epithelial erosion: This corneal abnormality is a non-specific finding appearing clinically as tiny defects in the epithelium. It is an early sign indicating epithelial compromise that stains positively with fluorescein.²¹

Pigment on endothelium (POE): This abnormality is characterized by corneal endothelial deposits often appearing colored and may be found in the corneal center, periphery, or both.

Corneal dystrophy: A bilateral hereditary disorder affecting various corneal layers, such as the epithelium, basal membrane, stroma, and endothelium, presenting as opacities, vesicles, rings, and streaks.²²

Corneal vascularization: This abnormality is characterized by the presence of blood vessels in different corneal layers, particularly the stroma.²³

Corneal degeneration: It is defined as pathological (abnormal) corneal changes during aging.²⁴

Any condition in at least one eye: This abnormality refers to any condition in at least one eye.

Socioeconomic Status

To assess the socioeconomic status (SES), we collected information on 13 assets owned by households and utilized principle component analysis to create an asset index based on the weights assigned to the first component.

Statistical Analysis and Model Building

The prevalence of corneal anomalies in Tehran was estimated by standardizing the samples based on age and

sex according to the 2016 census. The percentage of corneal abnormalities, along with their 95% confidence intervals, was calculated using the exact method. Due to the low prevalence of some corneal abnormalities, Firth's logistic regression was employed for model building to address sparse data bias.25 The relationship between various determinants (such as age, sex, education, employment status, marital status, eye examination, insurance, smoking, alcohol consumption, SES, and outdoor activity) and corneal abnormalities was assessed through simple Firth's logistic regression analysis. Determinants with a P value of less than 0.05 were included in the multiple Firth's logistic regression analysis. Correction for the cluster effect was applied to standard error, and data analysis was conducted using Stata version 12.0 software, with statistical significance set at P < 0.05.

Results

Among the 3791 individuals who were invited to participate in the study, a total of 3310 individuals accepted, resulting in a response rate of 87.31%. The average age of the participants was 68.25 ± 6.55 years, ranging from 60 to 97 years, with 57.8% female (n = 1913) and the remaining male. Additionally, 43.56% (n = 1442) were retired, 73.99% (n = 2449) were married, and 12.48% (n = 413) had received a university education.

The data presented in Table 1 illustrates the age and sex standardized prevalence of various corneal abnormalities. The prevalence rates for PE, PED, POE, CDys, and CV were 0.08% (95% CI: 0.02 to 0.40), 8.77% (95% CI: 6.64 to 11.51), 0.57% (95% CI: 0.33 to 0.98), 0.53% (95% CI: 0.33 to 0.82), and 0.95% (95% CI: 0.60 to 1.52), respectively. Furthermore, the age- and sex-standardized prevalence of any corneal condition in at least one eye was 49.08% (95% CI: 45.90 to 52.26). The prevalence of any corneal condition in at least one eye was observed to increase with

age, ranging from 37.52% in the age group of 60 to 64 years to 64.85% in the age group over 85 years.

Figure 1 depicts the prevalence of any corneal condition in at least one eye based on age, categorized by gender.

The prevalence of corneal degeneration was 44.87%. Among different age groups, individuals aged 80 years and above exhibited the highest prevalence of PED, POE, CV, and corneal degeneration [13.75% (95% CI: 8.92 to 20.60), 1.21% (95% CI: 0.38 to 3.82), 2.20% (95% CI: 0.92 to 5.21), and 56.78% (95% CI: 47.77 to 65.35), respectively]. In terms of gender, women showed the highest prevalence of PED, POE, and CDys [8.88% (95% CI: 6.60 to 11.84), 0.70% (95% CI: 0.38 to 1.29), and 0.57% (95% CI: 0.30 to 1.07), respectively]. The prevalence of any eye condition in at least one eye is detailed based on education and economic status in Table 2. The findings of the multiple logistic regression analysis are presented in Table 3. Among the determinants considered, such as age, sex, education, employment status, marital status, eye examination, insurance, smoking, alcohol consumption, SES, and outdoor activity, only some met the criteria for inclusion in the multiple model. Notably, no models were developed for CDys and PE as none of the determinants



Figure 1. Prevalence of Certain Corneal Conditions in at Least One Eye According to Age and Gender

 Table 1. Prevalence of Corneal Abnormalities in Tehran Geriatric Eye Study (TGES), 2019

Variables		Prevalence% (95% CI)							
		PE	PED	POD	CDys	CV	CDeg	Any Condition in at Least One Eye	
Total*		0.08 (0.02 to 0.40)	8.77 (6.64 to 11.51)	0.57 (0.33 to 0.98)	0.53 (0.33 to 0.82)	0.95 (0.60 to 1.52)	44.87 (41.80 to 47.98)	49.08 (45.90 to 52.26)	
Age	60-64 (n=1166)	0	6.97 (4.74 to 10.13)	0.36 (0.12 to 1.04)	0.39 (0.17 to 0.87)	0.7 (0.25 to 1.94)	34.18 (30.62 to 37.94)	37.52 (34.94 to 40.18)	
	65-69 (n=953)	0.08 (0.01 to 0.61)	8.74 (6.23 to 12.14)	0.67 (0.30 to 1.49)	0.22 (0.05 to 0.92)	0.76 (0.36 to 1.59)	43.05 (38.50 to 47.73)	46.47 (42.55 to 50.43)	
	70-74 (n=632)	0	8.28 (6.03 to 11.28)	0.61 (0.23 to 1.61)	1.03 (0.50 to 2.13)	0.77 (0.32 to 1.85)	52.44 (47.89 to 56.95)	56.42 (50.99 to 61.7)	
	74-79 (n=317)	0	9.44 (5.75 to 15.10)	0.23 (0.03 to 1.62)	1.07 (0.39 to 2.88)	1.01 (0.32 to 3.15)	55.54 (49.77 to 61.17)	60.13 (53.89 to 66.06)	
	$\geq 80 \ (n = 242)$	0.05 (0.07 to 3.56)	13.75 (8.92 to 20.60)	1.21 (0.38 to 3.82)	0.28 (0.04 to 2.00)	2.20 (0.92 to 5.21)	56.78 (47.77 to 65.35)	64.85 (57.36 to 71.68)	
Gender	Male (n=1397)	0	8.67 (6.02 to 12.33)	0.43 (0.19 to 0.98)	0.48 (0.24 to 0.96)	1.22 (0.65 to 2.26)	48.51 (44.94 to 52.09)	53.43 (49.41 to 57.41)	
	Female (n = 1913)	0.17 (0.03 to 0.79)	8.88 (6.60 to 11.84)	0.70 (0.38 to 1.29)	0.57 (0.30 to 1.07)	0.69 (0.35 to 1.37)	41.29 (37.58 to 45.09)	43.81 (40.81 to 46.86)	

*Age and sex standardized

CI, confidence interval; PE, posterior embryotoxon; PED, Punctate epithelial defect; POD, Pigment on Endothelium; CDys, Corneal Dystrophy; CV, Corneal Vascularization; CDeg, Corneal degeneration.

 Table 2. Prevalence of Corneal Abnormalities in Tehran Geriatric Eye Study by

 Education and Socioeconomic Status

Variables		% (95% CI)			
	Illiterate	54.55 (48.18-60.92)			
	Primary School	47.78 (42.29-53.27)			
Education	Guide School	48.14 (43.73-52.55)			
	High School	46.73 (41.99-51.48)			
	College	51.55 (45.67-57.43)			
	Low	52.92 (48.09-57.75)			
Socioeconomic	Mid	49.39 (43.22-55.56)			
status	High	48.77 (44.44-53.10)			
	Highest	43.59 (38.58-48.60)			

met the criteria for inclusion in the multiple model.

Based on the findings presented in Table 2, PED exhibited a positive correlation with age (OR: 2.03 for individuals over 80 years old compared to those aged 60-64 years, P=0.001) and a negative relationship with SES (OR: 0.52 for the highest SES group compared to the lowest, P=0.004). Conversely, corneal degeneration showed a direct association with age (OR: 2.79 for individuals over 80 years compared to those aged 60-64 years; P<0.001) and an inverse correlation with female gender (OR: 0.57, P<0.001). Additionally, employment status was significantly linked to corneal degeneration (OR: 2.04 for retired individuals versus employed, P=0.001; OR: 2.72 for housekeepers versus employed, P<0.001; OR: 1.96 for others versus employed, P=0.016). Female gender was negatively associated with corneal degeneration (OR: 0.57, P<0.001).

In relation to the outcomes, CV demonstrated a positive connection with age (OR: 3.74 for individuals over 80 years compared to those aged 60-64 years, P=0.028) and negative relationships with SES (OR: 0.17 for high SES group compared to the lowest, P=0.017) and female gender (OR: 0.44, P=0.048).

The average best corrected visual acuity in eyes with at least one abnormality was 0.17 ± 0.51 , while eyes without any problems had a mean value of 0.16 ± 0.43 logMAR (*P*=0.748).

Discussion

The TGES was the first Iranian population-based study exclusively conducted on subjects aged 60 years and over living in Tehran, capital of Iran. This study, as part of the TGES, was performed to determine the prevalence of corneal abnormalities according to age, sex, and other demographic characteristics to shed light on their risk factor; therefore, its results can help to better understand the epidemiologic profile of corneal abnormalities.¹⁵

An extensive literature search showed only one population-based study on the prevalence of cornmeal abnormalities¹³ and the majority of the studies in this regard are clinic-based or medical records-based. According to the results, 50% of the study population had at least one of the corneal abnormalities, which was higher than the prevalence reported by Hashemi et al.¹³ Moreover,

the most common abnormality in the present study was corneal degeneration with a prevalence of 44.87%, while in the study by Hashemi et al, the most common abnormality was PE with a prevalence of 14.7%.¹³

Although few studies have investigated corneal abnormalities in the world and Iran, caution should be practiced when comparing their results due to differences in the age structure and design.^{20,26,27} The results of the present study showed that less than 0.1% of the participants had PE, which was much lower than previous studies; for example, Hashemi et al reported a prevalence of 14.7% for PE in the Shahroud Eye Cohort Study.¹³ Clinic-based studies have also reported a higher prevalence compared to the present study; for example, the prevalence of PE was 6.8% in a study by Rennie et al,²⁰ 15% in a study by Burian et al,²⁸ 24.3% in a study by Ozeki et al,²⁷ and 95% in subjects suffering from Alagille syndrome.²⁶ What is clear is that the difference in the prevalence of PE between this study and the study conducted by Hashemi et al¹³ is due to age structure since the age range of the participants was 40-65 years in the above study and 60-97 years in the present study. It becomes more difficult to observe and detect PE with an increase in age due to increase in other corneal degenerative diseases and peripheral corneal opacity including arcus senilis,²⁰ which may be a reason for the lower prevalence of this corneal opacity in the present population compared to another study.¹⁶ On the other hand, the prevalence of PE was lower in our study compared to other clinic-based studies, which could be due to diagnosis based on histology in those studies^{20,26,27} which is more accurate than diagnosis based on slit lamp examination.

Although CDys is one of the most important reasons for corneal transplant in the world, especially among children, 6,29 less than 1% of the participants had CDys and this abnormality had the lowest prevalence among all corneal abnormalities in the present study. In line with the results of this study, Musch et al evaluated all records available in the national managed-care network and reported a prevalence of less than 1% in the USA.15 Similarly, Hashemi et al also reported a prevalence of less than 1% in the Shahroud Eye Cohort Study.13 According to other studies, the prevalence of CDys is 11% in individuals aged 50 years and over,³⁰ 4.5%-9% in Europe,³¹ and 3.8-4.1% in Asia.^{19,32} It is clear that genetics play the most important role in CDys^{15,33}; however, previous studies found that personal factors also increased its occurrence and worsened its manifestations.³¹ Although some studies reported that the cumulative effect of environmental factors can result in increased prevalence of CDys in the elderly,³⁴ this relationship was not observed in the present study. To explain this finding, it should be noted that CDys can be usually detected with a slit lamp after 30 years of age, and the chance of detection increases with age. Therefore, its high prevalence in advanced ages is related to the higher chance of diagnosis. In other words, age is not a causative factor but is a proxy for detectability. Table 3. Association Between Corneal Abnormalities with Varies Determinants Based on Multiple Firth's Logistic Regression in Tehran Geriatric Eye Study (TGES), 2019

	Variables		OR (95% CI)	P Value
		65-69	1.27 (0.92 to 1.75)	0.147
		70-74	1.17 (0.81 to 1.69)	0.380
	Age (baseline: 60 years old)	75-79	1.34 (0.86 to 2.11)	0.180
Durantete en ite eliel defent		>80	2.03 (1.32 to 3.16)	0.001*
Punctate epitnellal delect		Low	0.71 (0.49 to 1.07)	0.082
	SEC (baseline) Lowest)	Mid	0.73 (0.48 to 1.10)	0.130
	SES (baseline: Lowest)	High	0.67 (0.42 to 1.02)	0.061
		highest	0.52 (0.33 to 0.81)	0.004*
		Primary school	0.23 (0.07 to 0.72)	0.012*
Digmont on Endothalium	Education (baselines illiterate)	Guide school	0.29 (0.08 to 1.03)	0.056
Pigment on Endothelium	Education (baseline: initerate)	High school	0.03 (0.01 to 0.54)	0.017*
		Collage	0.44 (0.12 to 1.52)	0.194
		65-69	1.27 (0.44 to 3.65)	0.663
	Age (baseline: 60 years ald)	70-74	1.35 (0.42 to 4.29)	0.615
	Age (baseline: 60 years old)	75-79	1.52 (0.40 to 5.85)	0.541
		>80	3.74 (1.15 to 12.11)	0.028*
Corneal Vascularization		Low	0.31 (0.11 to 0.91)	0.033*
	SEC (baseline) Lowest)	Mid	0.55 (0.19 to 1.59)	0.269
	SES (baseline: Lowest)	High	0.17 (0.04 to 0.72)	0.017*
		highest	0.43 (0.13 to 1.41)	0.165
	Sex (baseline: Male)		0.44 (0.20 to 0.99)	0.048*
		65-69	1.42 (1.19 to 1.71)	< 0.001*
	Age (baseline: 60 years ald)	70-74	2.13 (1.74 to 2.62)	< 0.001*
	Age (baseline: 60 years old)	75-79	2.24 (1.72 to 2.92)	< 0.001*
		>80	2.79 (2.06 to 3.78)	< 0.001*
		Low	1.02 (0.8 to 1.31)	0.859
	SES (baseline) Lowest)	Mid	1.05 (0.81 to 1.38)	0.701
	SES (baseline: Lowest)	High	1.02 (0.78 to 1.35)	0.878
		Highest	0.81 (0.6 to 1.09)	0.160
Corneal degeneration		Primary school	0.86 (0.68 to 1.09)	0.206
	Education (baselines illiterate)	Guide school	0.82 (0.63 to 1.07)	0.139
	Education (baseline: initerate)	High School	0.97 (0.75 to 1.27)	0.833
		Collage	1.29 (0.93 to 1.79)	0.134
		Retired	2.04 (1.32 to 3.16)	0.001*
	Employment (baseline: Employed)	housekeeper	2.72 (1.65 to 4.5)	< 0.001*
	(Section Employed)	other	1.96 (1.13 to 3.38)	0.016*
	Smoking (baseline: No)		1.05 (0.84 to 1.31)	0.681
	Sex (baseline: Male)		0.57 (0.44 to 0.76)	< 0.001*

SES, socioeconomic status; NN, No Need.

*Significance.

None of the determinants has any significant association with Corneal dystrophy and Posterior Embryotoxon.

Moreover, in the present study, the prevalence of CDys was higher in women but the difference was not significant, which could be due to differences in the study population (ethnicity), diagnostic tool, and diagnostic accuracy of the tool used for diagnosis.³⁵ Other studies have also reported a higher prevalence of CDys in women,^{13,15,31} which could be mostly due to genetic differences between men and women. However, this difference may be in part due to hormonal differences and environmental factors

that affect its occurrence and manifestations.³⁶ Zoega et al found that smoking increased the chance of CDys by more than twice.³¹ In the present study, the prevalence of CDys was higher in smokers versus non-smokers (data not shown) but the difference was not significant.

The prevalence of PED was 8.77% in the present study. According to some reports, about 200000 cases of PED occur in the USA every year,³⁷ comprising more than 10% of all eye-related emergency room visits.³⁸

The prevalence of CV was about 1% in the present study, which was higher than the value reported by Hashemi et al.¹³ The reason for this disparity may be the age difference between the two studies. However, the prevalence of CV was largely affected by age, such that it increased from 0.7% in individuals aged 60-64 years to 2.2% in participants aged 80 years and over in the present study. According to the results of the multiple logistic model, age had a direct relationship, and economic status had an indirect relationship with the prevalence of PED and CV, which is consistent with other studies.¹³ Considering the findings of other studies, an increase in age can predict an increase in corneal abnormalities including PED and CV.^{13,20,27} It should be borne in mind that different factors such as trauma, infectious diseases, UV exposure, MGD, dry eye, and malnutrition including vitamin A deficiency are effective in the occurrence of CV and PED and age and economic status may serve as proxies for these factors.

As mentioned earlier, corneal degeneration was the most common corneal abnormality, comprising 91% of all cases excluding corneal opacity. It should be noted that corneal degeneration includes a number of abnormalities like fatty degeneration and arcus senilis, all of which have a direct relationship with age³⁹ and can be a reason for the higher prevalence of this abnormality in this age range. The present study found a direct relationship between corneal degeneration. This is line with the literature that other ocular disorders have a strong association with age.³⁹

Another finding of the present study was the role of gender in corneal abnormalities. The results of the present study show that there is an inter-gender difference in the prevalence of CV and corneal degeneration in such a way that the chance of these abnormalities was lower in women. Studies have rarely discussed the reason for this difference; however, it has been shown that women are less susceptible to disorders affecting corneal changes (like dry eye) due to hormonal differences,³ which could be a reason for the lower prevalence of CV and degeneration in women.

Conclusion

In summary, it can be deduced that half of the individuals who are 60 years old and above possess at least a single corneal abnormality. PE was the rarest, and corneal degeneration was the most common corneal abnormality, comprising 91% of all cases. According to the results of model building, age, economic status, education level, and sex were the determinants of the occurrence of corneal abnormalities, among which age was the most important determinant. Age stands out as the primary factor influencing the development of corneal abnormalities. Consequently, special attention should be directed towards the elderly population when formulating strategies for diagnosis and treatment. It should be noted that the results of the present study are related to the city of Tehran, which has a heterogeneous population in terms of race and ethnicity, and its results can be generalized to all of Iran. Moreover, the present study was conducted

on people aged 60 years and above. There is a limitation of comparison with this age group. However, due to the lack of studies in this field, the results of this study are valuable. Selection bias and measurement bias may affect the results; however, these errors are minimized due to stratified cluster random sampling and the fact that the definition and protocol for determining corneal problems are the same for everyone.

Authors' Contribution

Conceptualization: Hassan Hashemi, Mehdi Khabazkhoob. Data curation: Mohammadreza Aghamirsalim, Alireza Jamali. Formal Analysis: Mehdi Khabazkhoob. Funding Acquisition: Hassan Hashemi. Investigation: Mehdi Khabazkhoob. Methodology: Hassan Hashemi, Mehdi Khabazkhoob. Project administration: Mehdi Khabazkhoob. Resources: Hassan Hashemi. Supervision: Mehdi Khabazkhoob. Writing-original draft: Alireza Hashemi, Alireza Jamali. Writing-review & editing: Hassan Hashemi, Mehdi Khabazkhoob, Mohamadreza Aghamirsalim, Alireza Jamali, Alireza Hashemi.

Competing Interests

No conflicting relationship exists for any author.

Ethical Approval

The study protocol was approved by the National Institute for Medical Research Development (NIMAD), which is affiliated with the Iranian Ministry of Health. In accordance with the principles outlined in the Helsinki Declaration, this study was conducted, ensuring that informed consent was obtained from all participants. The ethics code for this study is IR.NIMAD.REC.1397.292.

Funding

This project was supported by National Institute for Medical Research Development(NIMAD) affiliated with the Iranian Ministry of Health and Medical Education (grant code: 963660).

References

- Flaxman SR, Bourne RR, Resnikoff S, Ackland P, Braithwaite T, Cicinelli MV, et al. Global causes of blindness and distance vision impairment 1990-2020: a systematic review and metaanalysis. Lancet Glob Health. 2017;5(12):e1221-34. doi: 10.1016/s2214-109x(17)30393-5.
- 2. Garg P, Krishna PV, Stratis AK, Gopinathan U. The value of corneal transplantation in reducing blindness. Eye (Lond). 2005;19(10):1106-14. doi: 10.1038/sj.eye.6701968.
- Mathews PM, Lindsley K, Aldave AJ, Akpek EK. Etiology of global corneal blindness and current practices of corneal transplantation: a focused review. Cornea. 2018;37(9):1198-203. doi: 10.1097/ico.00000000001666.
- Hashemi H, Pakzad R, Khabazkhoob M, Yekta A, Emamian MH, Fotouhi A. Distribution of different corneal topography patterns in Iranian schoolchildren: the Shahroud Schoolchildren Eye Cohort Study. Eye Contact Lens. 2020;46(3):154-9. doi: 10.1097/icl.000000000000652.
- Gain P, Jullienne R, He Z, Aldossary M, Acquart S, Cognasse F, et al. Global survey of corneal transplantation and eye banking. JAMA Ophthalmol. 2016;134(2):167-73. doi: 10.1001/jamaophthalmol.2015.4776.
- Bajracharya L, Gurung R, Demarchis EH, Oliva M, Ruit S, Tabin G. Indications for keratoplasty in Nepal: 2005-2010. Nepal J Ophthalmol. 2013;5(2):207-14. doi: 10.3126/ nepjoph.v5i2.8730.
- Rezaei Kanavi M, Javadi MA, Motevasseli T, Chamani T, Rezaei Kanavi M, Kheiri B, et al. Trends in indications and techniques of corneal transplantation in Iran from 2006 to 2013; an

8-year review. J Ophthalmic Vis Res. 2016;11(2):146-52. doi: 10.4103/2008-322x.183930.

- Cockburn DM. Posterior embryotoxon: its epidemiology and significance as a sign of glaucoma. Clin Exp Optom. 1995;78(2):74-8. doi: 10.1111/j.1444-0938.1995.tb00791.x.
- Michels K, Bohnsack BL. Ophthalmological manifestations of Axenfeld-Rieger syndrome: current perspectives. Clin Ophthalmol. 2023;17:819-28. doi: 10.2147/opth.s379853.
- Merali FI, Schein OD. Epidemiology of corneal diseases. In: Colby K, Dana R, eds. Foundations of Corneal Disease: Past, Present and Future. Cham: Springer International Publishing; 2020. p. 307-30. doi: 10.1007/978-3-030-25335-6_26.
- 11. Hashemi H, Pakzad R, Yekta A, Khabazkhoob M. The prevalence of corneal opacity in rural areas in Iran: a populationbased study. Ophthalmic Epidemiol. 2018;25(1):21-6. doi: 10.1080/09286586.2017.1337912.
- Worku EM, Fekadu SA, Alemie BW, Lorato MM. Prevalence and associated factors of corneal opacity among adults in Kolladiba town, Northwest Ethiopia: a cross-sectional study. BMJ Open Ophthalmol. 2024;9(1):e001665. doi: 10.1136/ bmjophth-2024-001665.
- Hashemi H, Khabazkhoob M, Emamian MH, Shariati M, Yekta A, Fotouhi A. The frequency of occurrence of certain corneal conditions by age and sex in Iranian adults. Cont Lens Anterior Eye. 2015;38(6):451-5. doi: 10.1016/j.clae.2015.05.005.
- 14. Das AV, Chaurasia S. Clinical profile and demographic distribution of corneal dystrophies in India: a study of 4198 patients. Cornea. 2021;40(5):548-53. doi: 10.1097/ ico.00000000002450.
- 15. Musch DC, Niziol LM, Stein JD, Kamyar RM, Sugar A. Prevalence of corneal dystrophies in the United States: estimates from claims data. Invest Ophthalmol Vis Sci. 2011;52(9):6959-63. doi: 10.1167/iovs.11-7771.
- Hashemi H, Pakzad R, Aghamirsalim MR, Yekta A, Sadoughi MM, Norouzikhiabani N, et al. Age and sex standardized prevalence of corneal opacity and its determinants; Tehran Geriatric Eye Study (TGES). Iran J Public Health. 2022;51(3):643-51. doi: 10.18502/ijph.v51i3.8941.
- 17. Kanasi E, Ayilavarapu S, Jones J. The aging population: demographics and the biology of aging. Periodontol 2000. 2016;72(1):13-8. doi: 10.1111/prd.12126.
- Mirzaie M, Darabi S. Population aging in Iran and rising health care costs. Iran J Ageing. 2017;12(2):156-69. doi: 10.21859/ sija-1202156.
- 19. Higa A, Sakai H, Sawaguchi S, Iwase A, Tomidokoro A, Amano S, et al. Prevalence of and risk factors for cornea guttata in a population-based study in a southwestern island of Japan: the Kumejima study. Arch Ophthalmol. 2011;129(3):332-6. doi: 10.1001/archophthalmol.2010.372.
- Rennie CA, Chowdhury S, Khan J, Rajan F, Jordan K, Lamb RJ, et al. The prevalence and associated features of posterior embryotoxon in the general ophthalmic clinic. Eye (Lond). 2005;19(4):396-9. doi: 10.1038/sj.eye.6701508.
- Winkler S. Punctate epithelial defects/erosions. In: Schmidt-Erfurth U, Kohnen T, eds. Encyclopedia of Ophthalmology. Berlin, Heidelberg: Springer; 2016. p. 1-2. doi: 10.1007/978-3-642-35951-4_929-1.
- Soh YQ, Kocaba V, Weiss JS, Jurkunas UV, Kinoshita S, Aldave AJ, et al. Corneal dystrophies. Nat Rev Dis Primers. 2020;6(1):46. doi: 10.1038/s41572-020-0178-9.
- 23. Sharif Z, Sharif W. Corneal neovascularization: updates on pathophysiology, investigations & management. Rom J Ophthalmol. 2019;63(1):15-22. doi: 10.22336/rjo.2019.4.
- 24. Alkatan H, Alkheraiji N, Alzahem T. Corneal dystrophies and degenerations. In: Ziaei A. Frontiers in Ophthalmology and

Ocular Imaging. Rijeka: IntechOpen; 2019. doi: 10.5772/ intechopen.84426.

- 25. Greenland S, Mansournia MA, Altman DG. Sparse data bias: a problem hiding in plain sight. BMJ. 2016;352:i1981. doi: 10.1136/bmj.i1981.
- Hingorani M, Nischal KK, Davies A, Bentley C, Vivian A, Baker AJ, et al. Ocular abnormalities in Alagille syndrome. Ophthalmology. 1999;106(2):330-7. doi: 10.1016/s0161-6420(99)90072-6.
- Ozeki H, Shirai S, Majima A, Sano M, Ikeda K. Clinical evaluation of posterior embryotoxon in one institution. Jpn J Ophthalmol. 1997;41(6):422-5. doi: 10.1016/s0021-5155(97)00080-4.
- 28. Burian HM, Braley AE, Allen L. Visibility of the king of Schwalbe and the trabecular zone: an interpretation of the posterior corneal embryotoxon and the so-called congenital hyaline membranes on the posterior corneal surface. AMA Arch Ophthalmol. 1955;53(6):767-82. doi: 10.1001/ archopht.1955.00930010775001.
- 29. Limaiem R, Chebil A, Baba A, Ben Youssef N, Mghaieth F, El Matri L. Pediatric penetrating keratoplasty: indications and outcomes. Transplant Proc. 2011;43(2):649-51. doi: 10.1016/j.transproceed.2011.01.055.
- Eghrari AO, McGlumphy EJ, Iliff BW, Wang J, Emmert D, Riazuddin SA, et al. Prevalence and severity of Fuchs corneal dystrophy in Tangier Island. Am J Ophthalmol. 2012;153(6):1067-72. doi: 10.1016/j.ajo.2011.11.033.
- 31. Zoega GM, Fujisawa A, Sasaki H, Kubota A, Sasaki K, Kitagawa K, et al. Prevalence and risk factors for cornea guttata in the Reykjavik Eye Study. Ophthalmology. 2006;113(4):565-9. doi: 10.1016/j.ophtha.2005.12.014.
- 32. Kitagawa K, Kojima M, Sasaki H, Shui YB, Chew SJ, Cheng HM, et al. Prevalence of primary cornea guttata and morphology of corneal endothelium in aging Japanese and Singaporean subjects. Ophthalmic Res. 2002;34(3):135-8. doi: 10.1159/000063656.
- 33. Eghrari AO, Gottsch JD. Fuchs' corneal dystrophy. Expert Rev Ophthalmol. 2010;5(2):147-59. doi: 10.1586/eop.10.8.
- Han S, Mueller C, Wuebbolt C, Kilcullen S, Nayyar V, Calle Gonzalez B, et al. Selective effects of estradiol on human corneal endothelial cells. bioRxiv [Preprint]. April 28, 2023. Available from: https://biorxiv.org/ content/10.1101/2023.04.27.538629v1.
- 35. Hashemi H, Khabazkhoob M, Pakzad R, Bakhshi S, Ostadimoghaddam H, Asaharlous A, et al. Pentacam accuracy in discriminating keratoconus from normal corneas: a diagnostic evaluation study. Eye Contact Lens. 2019;45(1):46-50. doi: 10.1097/icl.00000000000531.
- Krachmer JH, Purcell JJ Jr, Young CW, Bucher KD. Corneal endothelial dystrophy. A study of 64 families. Arch Ophthalmol. 1978;96(11):2036-9. doi: 10.1001/ archopht.1978.03910060424004.
- 37. Wirostko B, Rafii M, Sullivan DA, Morelli J, Ding J. Novel therapy to treat corneal epithelial defects: a hypothesis with growth hormone. Ocul Surf. 2015;13(3):204-12.e1. doi: 10.1016/j.jtos.2014.12.005.
- Channa R, Zafar SN, Canner JK, Haring RS, Schneider EB, Friedman DS. Epidemiology of eye-related emergency department visits. JAMA Ophthalmol. 2016;134(3):312-9. doi: 10.1001/jamaophthalmol.2015.5778.
- Hashemi H, Pakzad R, Heydarian S, Yekta A, Aghamirsalim M, Shokrollahzadeh F, et al. Global and regional prevalence of strabismus: a comprehensive systematic review and meta-analysis. Strabismus. 2019;27(2):54-65. doi: 10.1080/09273972.2019.1604773.

2024 The Author(s). This is an open-access article distributed under the terms of the Creative Commons Attribution License (http://creativecommons. org/licenses/by/4.0), which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.