**Research Paper** 

## Global, regional, and national burden of blindness and vision loss due to common eye diseases along with its attributable risk factors from 1990 to 2019: a systematic analysis from the global burden of disease study 2019

Xiaorong Yang<sup>1,2</sup>, Hui Chen<sup>1,2</sup>, Tongchao Zhang<sup>3</sup>, Xiaolin Yin<sup>3</sup>, Jinyu Man<sup>3</sup>, Qiufeng He<sup>3</sup>, Ming Lu<sup>1,2,3</sup>

<sup>1</sup>Clinical Epidemiology Unit, Qilu Hospital of Shandong University, Jinan, China <sup>2</sup>Clinical Research Center of Shandong University, Qilu Hospital, Cheeloo College of Medicine, Shandong University, Jinan, China <sup>3</sup>Department of Epidemiology and Health Statistics, School of Public Health, Cheeloo College of Medicine,

Shandong University, Jinan, China

Correspondence to: Xiaorong Yang, Ming Lu; email: <a href="mailto:yangxiaorong@sdu.edu.cn">yangxiaorong@sdu.edu.cn</a>, <a href="mailto:lwming@sdu.edu.cn">lwming@sdu.edu.cn</a>, <a href="mailto:lwming@sdu.edu.edu">lwming@sdu.edu.edu</a>, <a href="mailto:lwming@sdu.edu">lwming@sdu.edu</a>, <a href="mailto:lwming@sdu">lwming@sdu</a>, <a href="mailto:lwming@sdu"/>lwming@sdu</a>, <a href="mailto:lwming@sdu"/>lwming@sdu</a>, <a href="mailto:lwming@sdu"/>lwming@sdu</a>, <a href="mailto:lwming@sdu"/>lwming@sdu</a>, <a href="mailto:lwming@sdu</a>, <a href="mailto:lwming@sdu</a>, <a href="mailto:lwming@sdu</a>, <a href="mailto:lwmi

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#### ABSTRACT

To map the magnitudes and temporal trends of blindness and vision loss (BVL) due to common eye diseases along with its attributable risk factors at the national, regional, and global levels. The annual burden of BVL in 204 countries and territories was extracted from the Global Burden of Disease Study 2019. The estimated annual percentage change (EAPC) and causes composition change were calculated to quantify the temporal trends of BVL-related disease burden by sex, region, and eye disease. The global disability-adjusted life years (DALYs) of BVL increased from 12.44 million in 1990 to 22.56 million in 2019, with a slightly decreased rate from 3.03 to 2.78 per 1000 population (EAPC = -0.30). About 29.6% of BVL-related DALYs worldwide were caused by cataract, followed by refraction disorders (29.1%), near vision loss (21.7%), other vision loss (13.7%), glaucoma (3.3%), and age-related macular degeneration (2.5%) in 2019. The age-standardized DALYs rates due to each eye disease type in most regions were decreased, especially in countries with high burden and high-middle socio-demographic index. Moreover, the contribution of smoking and air pollution from solid fuels to BVL burden decreased, however, the age-standardized burden of BVL attributed to high body-mass index and high fasting plasma glucose elevated gradually across almost all regions. The temporal trend of BVL burden due to specific eye diseases varies remarkably by region, sex and age. Understanding the real-time patterns of BVL burden is crucial for formulating more effective and targeted prevention and healthcare strategies to decrease the BVL burden.

#### **INTRODUCTION**

The latest global estimation and project on blindness and distance and near vision impairment among the population aged 50 years and older documents that about 43.3 million people worldwide suffer from blindness and

295 million people have moderate and severe vision impairment in 2020 [1]. Globally, an estimated 22.6 million disability-adjusted life years (DALYs) were caused by blindness and vision loss (BVL) in 2019, accounting for 0.88% of total DALYs due to all causes [2]. BVL brings severe personal educational and financial difficulties [3], reduces the quality of daily living [4], and increases mortality [5]. More than onethird of the patients with vision impairment could have not received effective prevention and treatment [6]. Understanding the global, regional and national characteristics as well as the temporal trends in specific causes of BVL is the essential basis for targeted public policy-making, such as medical resource allocation and health program planning.

The updated global and regional prevalence of BVL categorized by the severity of visual impairment along with the eye diseases has been assessed among the population aged over 50 years old [1, 7], which provides comprehensive knowledge for understanding the landscape of BVL burden. But the burden of the entire population and the impact of potential risk factors have not been assessed vet. The latest Global Burden of Disease (GBD) 2019 Study comprehensively estimated the annual burden of 369 diseases using extensive data sources and robust statistical methods in 204 countries and territories from 1990 to 2019 [2], which provided a unique opportunity to grasp the progress of the BVL burden. In this study, we aim to retrieve the global, regional, and national prevalence and DALYs of BVL caused by six major eye diseases by sex and age group from 1990 to 2019 based on the GBD 2019 Study, and further estimate the changes in the DALYs due to specific eye disease of BVL attributed to potential risk factors. Our results will serve as an important supplement and extension to previous studies [1, 7], optimize the process of universal eve health care, and facilitate the establishment of targeted eye health strategies tailoring to different characteristics in the future.

## **RESULTS**

#### The global burden due to overall BVL

Globally, the accumulated number of BVL patients increased from 353.2 (95% uncertainty interval [UI]: 298.7, 414.0) million in 1990 to 713.9 (95% UI: 593.2, 841.1) million in 2019, with a stable age-standardized prevalence rate (ASPR) of about 8.5 cases per 100 people (Supplementary Table 1). The estimated DALYs of BVL worldwide increased from 12.44 (95% UI: 8.64, 17.38) million in 1990 to 22.56 (95% UI: 15.58, 31.74) million in 2019, while the estimated annual percentage change (EAPC) in age-standardized DALYs rate (-0.30; 95% confidence interval [CI]: -0.34, -0.25) showed a slight decline from 3.03 to 2.78 per 1000 population during the period (Table 1). The ASPR and agestandardized DALYs rate of BVL among women were somewhat greater than that among men (9.10/100vs. 8.25/100 in 2019, 2.91/1000 vs. 2.64/1000 in 2019, respectively), and the gender disparity in BVL

burden further widened gradually over the past 30 years (Table 1 and Supplementary Table 1).

# Variation in BVL burden at the national and regional level

The global variety of ASPR and age-standardized DALYs rate of BVL were around 11 and 7 times, respectively, in 2019 (Figure 1), with the highest ASPR in Nepal (21.27/100) and age-standardized DALYs rate in Indonesia (5.59/1000), and the lowest ASPR in Sweden (1.94/100) and age-standardized DALYs rate in Canada (0.81/1000). Overall, the ASPR in 2019 was higher than 11/100 in 29 countries and territories, including Tanzania, Zimbabwe, Lesotho, India, etc., (Figure 1A and Supplementary Table 2), which also showed a severe burden in age-standardized DALYs rate (Figure 1B and Supplementary Table 3). Conversely, Japan, Canada, Greenland and other 8 countries presented an ASPR of less than 2.5/100 (Figure 1A). In Equatorial Guinea, both ASPR and agestandardized DALYs rate had the largest decline (EAPC = -0.99; EAPC = -2.45, respectively) (Figure 1C, 1D). The top declines in age-standardized DALYs rate were also observed in Saudi Arabia, Cambodia, Qatar and Tunisia (Supplementary Table 4). Only 6 countries and territories (Burkina Faso, Cote d'Ivoire, Benin, Central African Republic, Chad, and Somalia) reported a significantly increasing age-standardized DALYs rate from 1990 and 2019 (Figure 1D and Supplementary Table 5).

For five socio-demographic index (SDI) regions, the number of DALYs of BVL increased across all SDI regions (Table 1 and Supplementary Figure 1), while the corresponding age-standardized DALYs rates decreased (Table 1). The greatest decrease observed in low-middle SDI region (EAPC = -0.82; 95% CI: -0.86, -0.77). As for GBD regions, absolute numbers in DALYs of BVL also increased in all regions (Table 1), but all 21 GBD regions showed a significant reduction in age-standardized DALYs rate of BVL, with the largest reduction found in Southeast Asia (EAPC = -1.02; 95% CI: -1.07, -0.98).

#### **BVL** burden due to specific eye diseases

The distribution of BVL-related DALYs due to specific eye diseases at the global and regional levels between 1990 and 2019 are presented in Supplementary Figure 1 and Figure 2. The BVL-related DALYs number due to all six eye diseases at global and regional levels increased during the monitoring period, especially for the near vision loss (Table 1 and Supplementary Figure 1). Globally, about 29.6% of BVL-related DALYs were caused by cataract, followed by refraction disorders

	199	0	2019		1990-2019	
Characteristics	DALYs No.×10 <sup>5</sup> (95% UI)	Age-standardized DALYs rate per 1000 No. (95% UI)	DALYs No.×10 <sup>5</sup> (95% UI)	Age-standardized DALYs rate per 1000 No. (95% UI)	EAPC in age- standardized DALYs Rate No. (95% CI)	
Overall	124.4 (86.4, 173.8)	3.03 (2.12, 4.21)	225.6 (155.7, 317.4)	2.78 (1.92, 3.92)	-0.30 (-0.34, -0.25)*	
Sex						
Males	56.4 (38.81, 79.01)	2.96 (2.08, 4.11)	100.7 (69.1, 142.4)	2.64 (1.82, 3.72)	-0.40 (-0.45, -0.36)*	
Females	68.02 (47.26, 94.69)	3.11 (2.18, 4.31)	124.9 (86.4, 175.0)	2.91 (2.01, 4.09)	-0.22 (-0.26, -0.17)*	
SDI						
High SDI	10.76 (7.42, 14.92)	1.13 (0.77, 1.56)	16.75 (11.59, 23.17)	1.08 (0.73, 1.52)	-0.16 (-0.20, -0.12)*	
High-middle SDI	25.96 (17.85, 36.81)	2.48 (1.72, 3.51)	45.15 (30.91, 64.58)	2.35 (1.61, 3.40)	-0.17 (-0.22, -0.11)*	
Middle SDI	41.33 (28.77, 57.19)	3.95 (2.79, 5.42)	80.07 (55.00, 112.7)	3.32 (2.29, 4.63)	-0.58 (-0.64, -0.52)*	
Low-middle SDI	34.17 (23.87, 47.50)	5.53 (3.94, 7.57)	59.55 (41.07, 83.20)	4.35 (3.02, 6.05)	-0.82 (-0.86, -0.77)*	
Low SDI	12.14 (8.39, 16.90)	4.91 (3.48, 6.72)	23.96 (16.49, 33.37)	4.35 (3.05, 5.98)	-0.42 (-0.48, -0.36)*	
GBD region						
High-income Asia Pacific	1.94 (1.36, 2.66)	1.05 (0.74, 1.44)	3.35 (2.37, 4.56)	0.97 (0.67, 1.34)	-0.28 (-0.33, -0.23)*	
High-income North America	2.82 (1.95, 3.91)	0.87 (0.59, 1.21)	4.36 (3.03, 6.02)	0.84 (0.58, 1.17)	-0.12 (-0.16, -0.08)*	
Western Europe	6.90 (4.85, 9.37)	1.38 (0.97, 1.89)	9.28 (6.52, 12.58)	1.27 (0.88, 1.74)	-0.27 (-0.29, -0.24)*	
Australasia	0.23 (0.16, 0.32)	1.08 (0.74, 1.51)	0.41 (0.29, 0.56)	1.02 (0.69, 1.42)	-0.17 (-0.21, -0.14)*	
Tropical Latin America	3.73 (2.63, 5.09)	3.67 (2.60, 4.97)	7.18 (5.03, 9.76)	3.06 (2.14, 4.17)	-0.28 (-0.42, -0.14)*	
Andean Latin America	0.92 (0.65, 1.26)	4.06 (2.90, 5.48)	1.82 (1.26, 2.54)	3.20 (2.23, 4.45)	-0.93 (-0.99, -0.88)*	
Central Latin America	3.28 (2.28, 4.52)	3.50 (2.48, 4.78)	6.74 (4.65, 9.40)	2.85 (1.97, 3.95)	-0.69 (-0.73, -0.66)*	
Southern Latin America	0.74 (0.52, 1.02)	1.61 (1.13, 2.20)	1.11 (0.77, 1.51)	1.43 (0.99, 1.97)	-0.36 (-0.38, -0.34)*	
Caribbean	0.77 (0.53, 1.07)	2.82 (1.96, 3.90)	1.22 (0.84, 1.72)	2.39 (1.65, 3.38)	-0.56 (-0.58, -0.55)*	
Eastern Europe	6.85 (4.63, 9.90)	2.61 (1.77, 3.76)	8.02 (5.3, 11.91)	2.50 (1.66, 3.69)	-0.26 (-0.30, -0.21)*	
Central Europe	2.40 (1.58, 3.62)	1.72 (1.13, 2.56)	3.09 (1.99, 4.69)	1.58 (1.03, 2.40)	-0.29 (-0.30, -0.27)*	
Central Asia	1.49 (1.01, 2.11)	3.12 (2.15, 4.40)	2.00 (1.36, 2.87)	2.76 (1.88, 3.95)	-0.46 (-0.48, -0.43)*	
North Africa and Middle East	7.84 (5.56, 10.77)	4.15 (2.96, 5.57)	13.93 (9.74, 19.11)	3.15 (2.22, 4.28)	-0.95 (-0.96, -0.94)*	
South Asia	38.94 (26.86, 54.07)	6.71 (4.78, 9.15)	72.99 (50.55, 101.9)	5.18 (3.62, 7.19)	-0.90 (-0.95, -0.85)*	
Southeast Asia	13.71 (9.73, 18.64)	5.30 (3.82, 7.11)	23.31 (16.44, 31.85)	4.00 (2.85, 5.43)	-1.02 (-1.07, -0.98)*	
East Asia	22.63 (15.43, 32.70)	2.63 (1.81, 3.77)	48.09 (32.41, 71.39)	2.43 (1.65, 3.56)	-0.19 (-0.36, -0.03)*	
Oceania	0.11 (0.08, 0.16)	3.59 (2.50, 4.93)	0.24 (0.17, 0.34)	3.33 (2.31, 4.65)	-0.26 (-0.34, -0.18)*	
Western Sub-Saharan Africa	3.89 (2.74, 5.41)	4.27 (3.01, 5.86)	8.48 (5.86, 11.93)	4.18 (2.91, 5.78)	-0.18 (-0.28, -0.08)*	
Eastern Sub-Saharan Africa	3.42 (2.38, 4.75)	4.32 (3.03, 5.90)	6.55 (4.54, 9.16)	3.73 (2.62, 5.13)	-0.51 (-0.54, -0.48)*	
Central Sub-Saharan Africa	0.56 (0.37, 0.85)	2.35 (1.57, 3.46)	1.25 (0.81, 1.90)	2.19 (1.44, 3.28)	-0.20 (-0.27, -0.14)*	
Southern Sub-Saharan Africa	1.24 (0.84, 1.80)	4.23 (2.92, 6.09)	2.14 (1.42, 3.25)	3.66 (2.46, 5.48)	-0.57 (-0.64, -0.50)*	
Specific eye diseases						
Glaucoma	4.42 (3.02, 6.26)	0.13 (0.09, 0.18)	7.48 (5.16, 10.45)	0.10 (0.07, 0.13)	-1.00 (-1.04, -0.95)*	
Cataract	34.93 (24.82, 47.2)	0.93 (0.66, 1.25)	66.76 (47.61, 90.06)	0.83 (0.59, 1.12)	-0.23 (-0.31, -0.14)*	
Age-related macular degeneration	2.97 (2.05, 4.19)	0.08 (0.06, 0.12)	5.64 (3.93, 7.89)	0.07 (0.05, 0.10)	-0.71 (-0.78, -0.65)*	
Refraction disorders	40.81 (27.56, 57.87)	0.89 (0.61, 1.24)	65.69 (44.34, 92.91)	0.82 (0.55, 1.15)	-0.31 (-0.35, -0.27)*	
Near vision loss	22.56 (10.35, 44.41)	0.56 (0.26, 1.09)	49.01 (22.56, 96.69)	0.59 (0.27, 1.17)	0.05 (0.00, 0.11)	
Other vision loss	18.73 (13.37, 25.56)	0.45 (0.32, 0.61)	31.00 (22.06, 42.43)	0.38 (0.27, 0.52)	-0.65 (-0.70, -0.60)*	

Table 1. DALYs and age-standardized DALYs rate per 1000 people for all blindness and vision loss in 1990 and 2019, and its estimated annual percentage change from 1990 to 2019.

DALYs, disability-adjusted life years; No., number; UI, uncertainty interval; EAPC, estimated annual percentage change; CI, confidential interval; SDI, socio-demographic index; GBD, global burden of disease. \* *P* value<0.05.

(29.1%), near vision loss (21.7%), other vision loss (13.7%), glaucoma (3.3%) and age-related macular degeneration (2.5%) in 2019 (Figure 2). Compared with the proportions in 1990, the decrease in refraction disorders (-3.7%) and the increase in near vision loss

(3.6%) in 2019 were notable. Besides, the proportions presented a great temporal and spatial variation. For instance, in Southern Latin America, the proportion of refraction disorders dropped from 47.9% in 1990 to 44.0% in 2019, and the proportion of age-related



**Figure 1. The global disease burden of BVL for both sexes in 204 countries and territories.** (A) The ASPR of BVL in 2019; (B) The age-standardized DALYs rate of BVL in 2019; (C) The EAPC in ASPR of BVL from 1990 to 2019. (D) The EAPC in age-standardized DALYs rate of BVL from 1990 to 2019. BVL, blindness and vision loss; ASPR, age-standardized prevalence rate; DALYs, disability-adjusted life years; EAPC, estimated annual percentage change.

macular degeneration in West Europe (10.5% in 2019) was much higher than that in other regions.

The age-standardized BVL burden rate along with the temporal trend due to specific eye diseases was significantly heterogeneous across the world (Supplementary Figures 2, 3). Overall, the agestandardized DALYs rates due to each categorized eye disease in most regions and countries were decreased. However, the age-standardized DALYs rate due to agerelated macular degeneration increased in some countries located in Sub-Saharan Africa; the near vision loss burden increased in Eastern Europe; and the refraction disorders burden increased in Western Sub-Saharan Africa, Tropical Latin America, and Australasia.

#### The BVL burden and age structure

Figure 3 presents the age distribution of BVL-related burden by specific eye diseases in 2019. The near vision loss was the predominant subtype in the prevalence of BVL across almost all age groups for both sexes. The proportion of refraction disorders in DALYs became pronounced across all age groups and the cataract was changed to an overwhelming subtype for DALYs due to BVL among the elders aged 60 years and above. The



Figure 2. Contribution of glaucoma, cataract, age-related macular degeneration, refraction disorders, near vision loss, and, other vision loss to absolute DALYs of BVL, both sexes, globally and by region, in 1990 and 2019. DALYs, disability-adjusted life years; BVL, blindness and vision loss.

burden rate of overall BVL kept pace with age increasing, especially after 55 years of age. The proportion of patients over 50 years old steadily increased among the DALYs number due to BVL for all six eye diseases (Figure 4A). Although the EAPCs in DALYs due to total BVL among all age groups were closed to -0.5, the remarkable dropped rate of burden due to cataract and other vision loss could be observed among the people under 50 years old (Figure 4B, 4C). Furthermore, the DALYs rate due to near vision loss among aged 5-49 years presented a slight upward trend.

#### The influential factors for EAPC

As shown in Figure 5, a significant negative association was detected between EAPC and age-standardized

overall BVL burden rates in 1990 at the national level 2.2e-16 for DALYs), which reflected that the BVL burden might be given priority intervention in countries with a high burden rate. Although the positive relationship between EAPC in age-standardized burden rate and SDI in 2019 was significant, the obvious U shape relationship could be observed. Moreover, we investigated the correlation between SDI and agestandardized DALYs rate by specific eye diseases from 1990 to 2019 in 21 GBD regions around the world (Supplementary Figure 4 and Figure 6). The results indicated that the age-standardized DALYs rate was markedly negatively associated with SDI in GBD regions with SDI ranging from 0.4 to 0.7, especially for the burden in glaucoma, cataract, age-related macular degeneration and other vision loss.





#### The BVL-related DALYs attributable to risk factors

We searched the identified potential risk factors for specific BVL disorders from the GBD database. Eventually, we found four risk factors involved in cataract: smoking, high body-mass index, household air pollution from solid fuels, and high fasting plasma glucose. Of these risk factors, household air pollution from solid fuels remained the greatest contributor to cataract-related DALYs, particularly for females in lower SDI regions, although the corresponding agestandardized DALYs rate gradually decreased from 1990 to 2019 (Figure 7). Moreover, we could observe the incremental influence on cataract from the high body-mass index and high fasting plasma glucose across almost all SDI regions.

The age-standardized DALYs rate of glaucoma attributable to high fasting plasma glucose in 2019 was negatively associated with SDI in 2019 at the national



Figure 4. The change of DALYs of BVL by different eye diseases and age groups, both sexes, from 1990 to 2019. (A) Number of DALYs; (B) DALYs rate; (C) EAPC in DALYs rate. The figures of six eye diseases are further enlarged in its top-right panel. BVL, blindness and vision loss; DALYs, disability-adjusted life years; EAPC, estimated annual percentage change.



Figure 5. The factors affected the EAPCs in age-standardized burden rate of BVL from 1990 to 2019, both sexes, at the national level. (A) ASPR of BVL in 1990 and EAPC in ASPR (B) SDI in 2019 and EAPC in ASPR; (C) age-standardized DALYs rate of BVL in 1990

and EAPC in age-standardized DALYs rate; (**D**) SDI in 2019 and EAPC in age-standardized DALYs rate. The circles represent countries and the size of circle is increased with the number of DALYs. The  $\rho$  indices and *P* values presented were derived from Spearman rank analysis. The blue line and its shade was fitted by LOESS. ASPR, age-standardized prevalence rate; EAPC, estimated annual percentage change; DALYs, disability-adjusted life years; BVL, blindness and vision loss; SDI, socio-demographic index.

levels ( $\rho = -0.548$ , P < 2.2e-16) (Figure 8). Besides, the proportion of glaucoma-related DALYs due to high fasting plasma glucose increased in almost all countries and territories between 1990 to 2019, except for Ethiopia. (Figure 8 and Supplementary Table 6). We could detect a significant association between the agestandardized DALYs rate of age-related macular degeneration attributable to smoking in 2019 and SDI in 2019 at the national levels ( $\rho = 0.200$ , P = 0.004) (Figure 9). The proportion of DALYs of age-related macular degeneration due to smoking decreased among most countries and territories during the monitoring period, except for 33 countries and territories, including Bosnia and Herzegovina, Russian and Lebanon, etc., (Figure 9 and Supplementary Tables 7, 8).

#### **DISCUSSION**

We comprehensively summarized the global, regional, and national burden in prevalence and DALYs of BVL by specific eye diseases along with the temporal trend over the past three decades in 204 countries and





territories. From a global perspective, the prevalence and DALYs cases of BVL were continuously increasing, with an estimated 0.71 billion people suffering from BVL in 2019, while the age-standardized rates only presented a mild downward trend during the period. However, the heterogeneous patterns in specific eye diseases, regions, sexes and age groups across the world make the prevention and healthcare of BVL complex [1, 7]. For example, the age-standardized DALYs rate caused by age-related macular degeneration increased in Sub-Saharan Africa, but the age-standardized DALYs rate caused by near vision loss increased in Eastern Europe. Therefore, understanding the exact change pattern of the BVL burden is essential for policy-makers to allocate rationally limited medical resources and formulate adapted prevention and treatment strategies.

The top three eye diseases for DALYs due to BVL remained cataract, refractive disorders, and near vision

loss over the past three decades. The DALYs due to cataract tended to be more prevalent in low or lowmiddle SDI regions, while it is promising that the agestandardized DALYs rates due to cataract in these regions have been decreasing over time, except for Burkina Faso, Chad, Afghanistan and Guinea, etc. The identified main contribution to cataract, namely, air pollution from solid fuels and smoking, dropped shapely across most countries, because the clean fuel for cooking has been popularized gradually with economic development and environmental requirement [8–10], and the prevalence of tobacco smoking further fell in more than 125 countries since 2003 under the WHO Framework Convention on Tobacco Control (FCTC) mobilization across the world [11, 12]. However, the age-standardized DALYs rates due to household air pollution from solid fuels in low SDI and low-middle regions still exceeded 74/100 000 over the past 30 years, especially in women, which was more than 300 times the





burden in the high SDI region in 2019. A recent study from India reported that the application of clean cooking fuels such as kerosene or liquefied petroleum gas didn't significantly increase with income growth, because the housework of cooking and fuel collection remains primarily women's responsibility and gender inequality within the family determines low household investment in clean fuel in developing areas [13]. Promoting healthy lifestyles and eliminating gender inequality can alleviate the corresponding burden. Moreover, we noted that the age-standardized DALYs rate of cataract attributed to high body-mass index and high fasting plasma glucose elevated gradually across almost all SDI regions and even surpassed the contributor of smoking and air pollution from solid fuels in the high SDI region. Despite the improvement in prevention and treatment, cataract remains a major threat to eye health and accounts for one-third blindness [7, 14].



**Figure 8. High fasting plasma glucose contributing to glaucoma-related DALYs, both sexes, at the national level.** (A) The association between age-standardized DALYs rates of glaucoma attributable to high fasting plasma glucose in 2019 and SDI in 2019. (B) The proportion of DALYs of glaucoma attributable to high fasting plasma glucose in 1990 and 2019. DALYs, disability-adjusted life years; SDI, socio-demographic index.

Unlike other BVL eye diseases, the distribution in DALYs due to refractive disorders did not exhibit an obvious association with SDI levels, and the corresponding geographical variation was lowest, except for a high burden in South Asia. The age-standardized DALYs rates due to refractive disorders in almost high-burden areas have been declining, which is consistent with the previous studies [15, 16]. Although the rate of refractive disorders slightly increased with age, the

number of DALYs due to refractive disorders remained relatively stable across age groups among the population under 80 years old, which indicates that eye health promotion among young people may decrease the burden [17]. No clear risk factors for refractive disorders were evaluated in the GBD study, but some studies reported that education years, unreasonable use of eyes, fine particulate matter and ozone in the environment are associated with refractive disorders [18–20].



Figure 9. Smoking contributing to DALYs of age-related macular degeneration, both sexes, at the national level. (A) The association between age-standardized DALYs rates of age-related macular degeneration attributable to smoking in 2019 and SDI in 2019. (B) The proportion of DALYs of age-related macular degeneration attributable to smoking in 1990 and 2019. DALYs, disability-adjusted life years; SDI, socio-demographic index.

The DALYs due to near vision loss tended to be pronounced in Sub-Sahara Africa and India. Of all the identified BVL eye diseases, only the age-standardized DALYs rate due to near vision loss in most countries and regions remained relatively stable over the last couple of decades, which means the number of near vision loss patients would still shapely increase under the population growth in Sub-Sahara Africa and India. The relatively stable rate might be related to the negligence of the control of near vision loss in public health policy, because of the safe and effective treatment with optical correction [21], which is also illustrated by the difference between prevalence and DALYs proportion. Albeit, no potential environmental risk factor was well-identified for near vision loss in the GBD study, we observed that the burden due to near vision loss in high-risk areas was highly related to the low socio-economic status, which was in agreement with the previous studies [22, 23]. Low socioeconomic status usually indicates poor awareness of vision loss, less access to vision correction, lack of ophthalmologists and optometrists, and inability to afford suitable glasses. Reasonable eye use, eye care exercises, and the availability of glasses could reduce the burden of near vision loss [24, 25].

The age-standardized DALYs rates due to glaucoma and age-related macular degeneration were significantly lower than the above-mentioned eye diseases, but they are responsible for about one-sixth of blindness [16, 21]. The geographical distributions for glaucoma and age-related macular degeneration were highly similar. and the disease burden was pronounced in low SDI areas, except the high burden of age-related macular degeneration was found in Western Europe [26-28]. The obvious decline in DALYs due to glaucoma was detected in 194 countries and territories, especially in high-burden areas, but the proportion of glaucomarelated DALYs due to high fasting plasma glucose increased in almost all countries and territories, which further emphasizes the importance of the prevention and control of high fasting plasma glucose in public policy [29, 30]. By contrast, 181 countries and territories presented a downward trend in DALYs due to agerelated macular degeneration, and the contribution of smoking fell in most countries and territories, with the implementation of various tobacco control measures [12]. Some studies have indicated that the high smoking prevalence, Western diet pattern, increased high-density lipoprotein, low 25-hydroxyvitamin D, and decreased amino acids are the risk factors for age-related macular degeneration [12, 31–36], which may contribute to the high burden in high SDI areas.

Apart from glaucoma, cataract, age-related macular degeneration, refraction disorders and near vision loss,

other eve diseases such as diabetic retinopathy, retinopathy of prematurity, trachoma, and onchocerciasis, were integrated into "other vision loss" in this study; these have also been recognized as causations of BVL [21, 37, 38]. In the current study, we observed that agestandardized DALYs rates due to other vision loss decreased in 193 regions, especially in North Africa and Middle East, and Southeast Asia. However, the highburden areas, including most Western Sub-Saharan Africa countries and Afghanistan, presented a minor increased trend. Our results may indicate that the various risk factors responsible for other vision loss, such as the lack of access to water and crowded living conditions that cause trachoma [39], need to be improved, and the available treatment, such as the use of azithromycin for trachoma [40], need to be strengthened, especially in high-burden areas.

In addition, we found that the temporal trend in agestandardized DALYs rate due to BVL from 1990 to 2019, namely EAPC, was significantly negatively associated with baseline age-standardized rate. For those heavily burdened countries in 1990, the age-standardized DALYs rate of BVL was more likely to decline, which could be explained by the fact that countries with high burden are likely to take into account BVL and identified risk factors as a high priority in disease prevention and treatment schemes [41]. The annualized declined trend of age-standardized DALYs rates due to BVL was associated with the increasing SDI, except for the SDI areas exceeding 0.8 with a low burden in BVL. Possible reasons involve the highly public attention to BVL, intensive eye health education, the availability of clean cooking fuel, effective tobacco control intervention, access to visual examination, and improved vision correction and treatment [22, 42].

Some deficiencies should be taken into account in the interpretation of our results. First, although the collaborators in GBD 2019 study have done enormous work every year to assess the disease burden worldwide, bias could not be avoided in fitting the unavailable data, as described previously [2, 29]. Second, the heterogeneity in quality and quantity of data sources, diagnosis of BVL and its disease subtypes, assessment for risk factors of BVL in different regions might hamper the conclusion.

BVL is still one of the major global public health concerns. Albeit minor achievements have been attained in the age-standardized burden rate in most countries and territories, the absolute BVL burden continues increasing rapidly in the context of population growth and aging, especially in low SDI areas. We need to further strengthen tobacco control and replace solid fuels with clean energy to minimize household air pollution. Meanwhile, we should control the body-mass index and fasting plasma glucose to reduce the related BVL burden. Fortunately, WHO united with Member States and their partners have initiated and implemented several related programs, such as Vision 2020 the right to sight, Universal eye health: a global action plan 2014-2019, and World report on vision. These programs will help to reduce the BVL burden by providing guidance and technical support for comprehensive vision protection systems and promoting the available ophthalmic care. The information addressed in the current study would clarify the global disease burden of BVL by specific eye diseases and formulate more effective and targeted BVL prevention and healthcare strategies.

#### **MATERIALS AND METHODS**

#### Data source

The detailed resources search, inclusion and exclusion criteria, data processing steps and modeling methods of GBD 2019 have been delineated in previous GBD studies [2, 29], and the repeatable analytical process and statistical codes for estimating BVL burden could be obtained from the accessible supporting website (http://ghdx.healthdata.org/gbd-2019/code). Here, we briefly introduced the methods specific to the estimation of the BVL burden. Each step performed in this study to analyze and report the GBD database complied with the Guidelines for Accurate and Transparent Health Estimates Reporting (GATHER) statements [43]. The following search terms were used to collect BVL burden information on MEDLINE, Embase, WHOLIS, SciELO, Open Grey and other gray literature searches: ("glaucoma" OR "cataract" OR "macular" OR "refractive error" OR "presbyopia" OR "blindness" OR "vision, low") AND ("prevalence" OR "incidence" OR "epidemiology"). A total of 387 original BVL-related data sources were extracted for this cause-specific assessment, covering representative population-based studies, peer-reviewed publications, grey literature and recorded surveys. All ICD-9 and ICD-10 codes pertaining to BVL (360.8-362, 362.1-363.9, 365-369.9, 377-378.9 and H25-H28.8, H31-H36.8, H40-H40.9, H42-H42.8, H46-H54.9, respectively) were claimed in the GBD estimates. The Disease Modelling Meta-Regression (DisMod-MR) 2.1 modeling was developed by the GBD Collaborators to estimate nonfatal diseases burden based on Bayesian mixed-effects meta-regression [2]. In the DisMod model, all original data from all available regions was input into a mixedeffects nonlinear model to produce a global estimation of BVL burden as well as fixed and random effects. The model outputs were delivered to the regional level as a prior to fit the regional estimations, which were further used as the prior values to fit the national estimations.

The 95% uncertainty interval (UI) for all estimations was generated using the 2.5th and 97.5th centiles of 1000 random draws of the posterior distribution.

We exacted annual data on BVL burden by sex, 5-year age group and eye diseases (glaucoma, cataract, agerelated macular degeneration, refraction disorders, near vision loss, and, other vision loss) in 204 countries and regions from 1990 to 2019 from the Institute for Health Metrics and Evaluation (http://ghdx.healthdata.org/gbdresults-tool). In order to delineate the disease burden of BVL across the world, the 204 available countries and territories were divided into five regions according to the corresponding SDI, a comprehensive indicator combining per capita income, years of education, and fertility, namely, low, low-middle, middle, high-middle, and high SDI regions. Besides, the world was geographically categorized into 21 GBD areas, e.g., high-income Asia Pacific, Tropical Latin America, and West Europe (Table 1), which were further simplified into seven super GBD regions.

Among 87 behavioral, environmental and occupational. and metabolic risk factors quantified in GBD 2019, the attributable risks of smoking, high body-mass index, household air pollution from solid fuels, and high fasting plasma glucose for BVL burden were well estimated, based on the established risk-outcome inclusion criterion. The comparative risk assessment framework was applied to calculate the population attributable fraction of potential risk factors for disease-specific BVL burden [29]: identify convincing risk-outcome pairs with the summary relative risk based on systematic review and meta-regression, estimate the exposure level and distribution by spatio-temporal Gaussian process regression and Bayesian mixed-effects meta-regression, define the theoretical minimum risk exposure level, and calculate the population attributable fraction and attributable burden.

#### Statistical analysis

The ASPR and age-standardized DALYs rate were used to quantify the variation of BVL burden by calendar year, sex, and region, to avoid the differences in the age composition of the population, based on the WHO World Standard Population Distribution 2001. We calculated the EAPC to describe the temporal trend in various agestandardized rates of BVL burden. We performed a regression model to fit the natural logarithm of the agestandardized rate with the calendar year, namely, ln (agestandardized rate) =  $\alpha + \beta^*$  calendar year +  $\epsilon$ , as our previous articles [44, 45]. The EAPC with its 95%CI were estimated according to the formula of 100 × (exp ( $\beta$ ) – 1). In our study, the 95% lower boundary of EAPC greater than 0 indicated that the changing trend was increasing, whereas the 95% upper boundary of EAPC less than 0 indicated that the changing trend was decreasing. We applied the Spearman rank test to calculate the relationship correlation between the EAPCs in BVL-related burden and the baseline burden in 1990 and the SDI in 2019 at the national level. The age-standardized rate of BVL burden in 1990 could be used as a proxy for the baseline disease reservoir, and the SDI in 2019 could represent the level and availability of health care in each country. All statistical analyses in the current study were conducted using R program version 4.0.3 (https://www.R-project.org/), and the two-sided *P* value less than 0.05 was considered statistically significant, which was presented in scientific notation format when it was less than 0.001.

#### Ethics approval and consent to participate

The GBD 2019 study is a publicly available database and all data were anonymous. Our study protocol was approved by the Institutional Review Boards of Qilu Hospital of Shandong University with approval number KYLL-202011(KS)-134.

#### Data availability statement

All data could be extracted from the online database (<u>http://ghdx.healthdata.org/gbd-results-tool</u>).

## Abbreviations

ASPR: age-standardized prevalence rate; BVL: blindness and vision loss; CI: confidence interval; DALYs: disability-adjusted life years; EAPC: estimated annual percentage change; GBD: global burden of disease; SDI: socio-demographic index; UI: uncertainty interval.

## **AUTHOR CONTRIBUTIONS**

Study design: XRY, ML; Data collection: XRY, HC, TCZ, XLY, JYM, QFH; Data analyses: XRY, HC, TCZ; Results visualization: XRY, XLY, JYM, QFH; Results interpretations: All authors; Manuscript writing: XRY, ML.

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## **CONFLICTS OF INTEREST**

The authors declare no potential conflicts of interest.

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## REFERENCES

- GBD 2019 Blindness and Vision Impairment Collaborators, and Vision Loss Expert Group of the Global Burden of Disease Study. Trends in prevalence of blindness and distance and near vision impairment over 30 years: an analysis for the Global Burden of Disease Study. Lancet Glob Health. 2021; 9:e130–43. <u>https://doi.org/10.1016/S2214-109X(20)30425-3</u> PMID:<u>33275950</u>
- GBD 2019 Diseases and Injuries Collaborators. Global burden of 369 diseases and injuries in 204 countries and territories, 1990-2019: a systematic analysis for the Global Burden of Disease Study 2019. Lancet. 2020; 396:1204–22. https://doi.org/10.1016/S0140-6736(20)30925-9

https://doi.org/10.1016/S0140-6736(20)30925-9 PMID:<u>33069326</u>

- Chakravarthy U, Biundo E, Saka RO, Fasser C, Bourne R, Little JA. The Economic Impact of Blindness in Europe. Ophthalmic Epidemiol. 2017; 24:239–47. <u>https://doi.org/10.1080/09286586.2017.1281426</u> PMID:28665742
- Cupples ME, Hart PM, Johnston A, Jackson AJ. Improving healthcare access for people with visual impairment and blindness. BMJ. 2012; 344:e542. <u>https://doi.org/10.1136/bmj.e542</u> PMID:<u>22290101</u>
- Zhang T, Jiang W, Song X, Zhang D. The association between visual impairment and the risk of mortality: a meta-analysis of prospective studies. J Epidemiol Community Health. 2016; 70:836–42. <u>https://doi.org/10.1136/jech-2016-207331</u> PMID:<u>27095181</u>
- World Health Organization. World report on vision. (Geneva, Switzerland: World Health Organization). 2019. <u>https://www.who.int/publications/i/item/97892</u> <u>41516570</u>
- 7. GBD 2019 Blindness and Vision Impairment Collaborators, and Vision Loss Expert Group of the Global Burden of Disease Study. Causes of blindness and vision impairment in 2020 and trends over 30 years, and prevalence of avoidable blindness in relation to VISION 2020: the Right to Sight: an analysis

for the Global Burden of Disease Study. Lancet Glob Health. 2021; 9:e144–60. https://doi.org/10.1016/S2214-109X(20)30489-7 PMID:33275949

- Puzzolo E, Zerriffi H, Carter E, Clemens H, Stokes H, Jagger P, Rosenthal J, Petach H. Supply Considerations for Scaling Up Clean Cooking Fuels for Household Energy in Low- and Middle-Income Countries. Geohealth. 2019; 3:370–90. <u>https://doi.org/10.1029/2019GH000208</u> PMID:<u>32159025</u>
- Balmes JR. Household air pollution from domestic combustion of solid fuels and health. J Allergy Clin Immunol. 2019; 143:1979–87. <u>https://doi.org/10.1016/j.jaci.2019.04.016</u> PMID:<u>31176380</u>
- Puzzolo E, Pope D, Stanistreet D, Rehfuess EA, Bruce NG. Clean fuels for resource-poor settings: A systematic review of barriers and enablers to adoption and sustained use. Environ Res. 2016; 146:218–34. <u>https://doi.org/10.1016/j.envres.2016.01.002</u> PMID:<u>26775003</u>
- Gravely S, Giovino GA, Craig L, Commar A, D'Espaignet ET, Schotte K, Fong GT. Implementation of key demand-reduction measures of the WHO Framework Convention on Tobacco Control and change in smoking prevalence in 126 countries: an association study. Lancet Public Health. 2017; 2:e166–74. <u>https://doi.org/10.1016/S2468-2667(17)30045-2</u> PMID:<u>29253448</u>
- Bilano V, Gilmour S, Moffiet T, d'Espaignet ET, Stevens GA, Commar A, Tuyl F, Hudson I, Shibuya K. Global trends and projections for tobacco use, 1990-2025: an analysis of smoking indicators from the WHO Comprehensive Information Systems for Tobacco Control. Lancet. 2015; 385:966–76. <u>https://doi.org/10.1016/S0140-6736(15)60264-1</u> PMID:<u>25784347</u>
- Choudhuri P, Desai S. Gender inequalities and household fuel choice in India. J Clean Prod. 2020; 265:121487. <u>https://doi.org/10.1016/j.jclepro.2020.121487</u> PMID:32831484
- Khairallah M, Kahloun R, Bourne R, Limburg H, Flaxman SR, Jonas JB, Keeffe J, Leasher J, Naidoo K, Pesudovs K, Price H, White RA, Wong TY, et al, and Vision Loss Expert Group of the Global Burden of Disease Study. Number of People Blind or Visually Impaired by Cataract Worldwide and in World Regions, 1990 to 2010. Invest Ophthalmol Vis Sci. 2015; 56:6762–69. https://doi.org/10.1167/joys.15-17201

https://doi.org/10.1167/iovs.15-17201 PMID:<u>26567788</u>.

- Fricke TR, Tahhan N, Resnikoff S, Papas E, Burnett A, Ho SM, Naduvilath T, Naidoo KS. Global Prevalence of Presbyopia and Vision Impairment from Uncorrected Presbyopia: Systematic Review, Meta-analysis, and Modelling. Ophthalmology. 2018; 125:1492–99. <u>https://doi.org/10.1016/j.ophtha.2018.04.013</u> PMID:<u>29753495</u>
- 16. Flaxman SR, Bourne RR, Resnikoff S, Ackland P, Braithwaite T, Cicinelli MV, Das A, Jonas JB, Keeffe J, Kempen JH, Leasher J, Limburg H, Naidoo K, et al, and Vision Loss Expert Group of the Global Burden of Disease Study. Global causes of blindness and distance vision impairment 1990-2020: a systematic review and meta-analysis. Lancet Glob Health. 2017; 5:e1221–34. <u>https://doi.org/10.1016/S2214-109X(17)30393-5</u> PMID:<u>29032195</u>
- Hu Y, Ding X, Guo X, Chen Y, Zhang J, He M. Association of Age at Myopia Onset With Risk of High Myopia in Adulthood in a 12-Year Follow-up of a Chinese Cohort. JAMA Ophthalmol. 2020; 138:1129–34. <u>https://doi.org/10.1001/jamaophthalmol.2020.3451</u> PMID:<u>32940622</u>
- Lin H, Guo Y, Ruan Z, Yang Y, Chen Y, Zheng Y, Cummings-Vaughn LA, Rigdon SE, Vaughn MG, Sun S, Zhang L, Wang X, Qian ZM, Wu F. Ambient PM<sub>2.5</sub> and O<sub>3</sub> and their combined effects on prevalence of presbyopia among the elderly: A cross-sectional study in six low- and middle-income countries. Sci Total Environ. 2019; 655:168–73.

https://doi.org/10.1016/j.scitotenv.2018.11.239 PMID:<u>30469062</u>

 Mountjoy E, Davies NM, Plotnikov D, Smith GD, Rodriguez S, Williams CE, Guggenheim JA, Atan D. Education and myopia: assessing the direction of causality by mendelian randomisation. BMJ. 2018; 361:k2022.

https://doi.org/10.1136/bmj.k2022 PMID:29875094

- Plotnikov D, Williams C, Atan D, Davies NM, Ghorbani Mojarrad N, Guggenheim JA, and UK Biobank Eye and Vision Consortium. Effect of Education on Myopia: Evidence from the United Kingdom ROSLA 1972 Reform. Invest Ophthalmol Vis Sci. 2020; 61:7. <u>https://doi.org/10.1167/iovs.61.11.7</u> PMID:<u>32886096</u>
- Resnikoff S, Pascolini D, Mariotti SP, Pokharel GP. Global magnitude of visual impairment caused by uncorrected refractive errors in 2004. Bull World Health Organ. 2008; 86:63–70. <u>https://doi.org/10.2471/blt.07.041210</u> PMID:18235892
- 22. Wang Y, Lou L, Cao J, Shao J, Ye J. Socio-economic disparity in global burden of near vision loss: an analysis for 2017 with time trends since 1990. Acta Ophthalmol. 2020; 98:e138–43.

#### https://doi.org/10.1111/aos.14198 PMID:<u>31429212</u>

- 23. Zebardast N, Friedman DS, Vitale S. The Prevalence and Demographic Associations of Presenting Near-Vision Impairment Among Adults Living in the United States. Am J Ophthalmol. 2017; 174:134–44. <u>https://doi.org/10.1016/j.ajo.2016.11.004</u> PMID:<u>27865728</u>
- 24. Han X, Ellwein LB, Guo X, Hu Y, Yan W, He M. Progression of Near Vision Loss and Incidence of Near Vision Impairment in an Adult Chinese Population. Ophthalmology. 2017; 124:734–42. <u>https://doi.org/10.1016/j.ophtha.2017.01.020</u> PMID:<u>28336059</u>
- Wolffsohn JS, Davies LN. Presbyopia: Effectiveness of correction strategies. Prog Retin Eye Res. 2019; 68:124–43. <u>https://doi.org/10.1016/j.preteyeres.2018.09.004</u> PMID:<u>30244049</u>
- Tham YC, Li X, Wong TY, Quigley HA, Aung T, Cheng CY. Global prevalence of glaucoma and projections of glaucoma burden through 2040: a systematic review and meta-analysis. Ophthalmology. 2014; 121:2081– 90.

https://doi.org/10.1016/j.ophtha.2014.05.013 PMID:24974815

- Ye X, She X, Shen L. Association of sex with the global burden of glaucoma: an analysis from the global burden of disease study 2017. Acta Ophthalmol. 2020. [Epub ahead of print]. https://doi.org/10.1111/aos.14330 PMID:31912656
- Wong WL, Su X, Li X, Cheung CM, Klein R, Cheng CY, Wong TY. Global prevalence of age-related macular degeneration and disease burden projection for 2020 and 2040: a systematic review and meta-analysis. Lancet Glob Health. 2014; 2:e106–16. <u>https://doi.org/10.1016/S2214-109X(13)70145-1</u> PMID:25104651
- 29. GBD 2019 Risk Factors Collaborators. Global burden of 87 risk factors in 204 countries and territories, 1990-2019: a systematic analysis for the Global Burden of Disease Study 2019. Lancet. 2020; 396:1223–49. <u>https://doi.org/10.1016/S0140-6736(20)30752-2</u> PMID:<u>33069327</u>
- 30. Ye L, Xu J, Zhang T, Lin X, Pan X, Zeng W, Ren Y, Shan P. Global burden of noncommunicable diseases attributable to high fasting plasma glucose. J Diabetes. 2020; 12:807–18. <u>https://doi.org/10.1111/1753-0407.13072</u> PMID:32472661
- 31. Dighe S, Zhao J, Steffen L, Mares JA, Meuer SM, Klein BE, Klein R, Millen AE. Diet patterns and the incidence

of age-related macular degeneration in the Atherosclerosis Risk in Communities (ARIC) study. Br J Ophthalmol. 2020; 104:1070–76. https://doi.org/10.1136/bjophthalmol-2019-314813 PMID:31810976

- 32. Millen AE, Nie J, Mares JA, Lutsey PL, LaMonte MJ, Meuer SM, Sahli MW, Andrews CA, Klein BE, Klein R. Serum 25-Hydroxyvitamin D Concentrations and Incidence of Age-Related Macular Degeneration: The Atherosclerosis Risk in Communities Study. Invest Ophthalmol Vis Sci. 2019; 60:1362–71. <u>https://doi.org/10.1167/iovs.18-25945</u> PMID:30934055
- 33. Acar İE, Lores-Motta L, Colijn JM, Meester-Smoor MA, Verzijden T, Cougnard-Gregoire A, Ajana S, Merle BM, de Breuk A, Heesterbeek TJ, van den Akker E, Daha MR, Claes B, et al, and EYE-RISK Consortium. Integrating Metabolomics, Genomics, and Disease Pathways in Age-Related Macular Degeneration: The EYE-RISK Consortium. Ophthalmology. 2020; 127:1693–709. <u>https://doi.org/10.1016/j.ophtha.2020.06.020</u> PMID:32553749
- Myers CE, Klein BE, Gangnon R, Sivakumaran TA, Iyengar SK, Klein R. Cigarette smoking and the natural history of age-related macular degeneration: the Beaver Dam Eye Study. Ophthalmology. 2014; 121:1949–55. https://doi.org/10.1016/j.ophtha.2014.04.040

PMID:24953792

 Detaram HD, Joachim N, Liew G, Vu KV, Burlutsky G, Mitchell P, Gopinath B. Smoking and treatment outcomes of neovascular age-related macular degeneration over 12 months. Br J Ophthalmol. 2020; 104:893–98.

https://doi.org/10.1136/bjophthalmol-2019-314849 PMID:<u>31558491</u>

- 36. Casey C, Woodside JV, McGinty A, Young IS, McPeake J, Chakravarthy U, Rahu M, Seland J, Soubrane G, Tomazzoli L, Topouzis F, Vioque J, Fletcher AE. Factors associated with serum 25-hydroxyvitamin D concentrations in older people in Europe: the EUREYE study. Eur J Clin Nutr. 2019; 73:319–28. <u>https://doi.org/10.1038/s41430-018-0353-1</u> PMID:<u>30367156</u>
- 37. Leasher JL, Bourne RR, Flaxman SR, Jonas JB, Keeffe J, Naidoo K, Pesudovs K, Price H, White RA, Wong TY, Resnikoff S, Taylor HR, and Vision Loss Expert Group of the Global Burden of Disease Study. Global Estimates on the Number of People Blind or Visually Impaired by Diabetic Retinopathy: A Meta-analysis From 1990 to 2010. Diabetes Care. 2016; 39:1643–49. Erratum in: Diabetes Care. 2016; 39:2096.

https://doi.org/10.2337/dc15-2171. PMID:27555623

- 38. Courtright P, Rotondo LA, MacArthur C, Jones I, Weaver A, Negash BK, Olobio N, Binnawi K, Bush S, Abdala M, Haddad D, Bonfield A, Emerson P, et al. Strengthening the links between mapping, planning and global engagement for disease elimination: lessons learnt from trachoma. Br J Ophthalmol. 2018; 102:1324–27. <u>https://doi.org/10.1136/bjophthalmol-2018-312476</u> PMID:<u>29907634</u>
- 39. West SK. Milestones in the fight to eliminate trachoma. Ophthalmic Physiol Opt. 2020; 40:66–74. <u>https://doi.org/10.1111/opo.12666</u> PMID:<u>32017172</u>
- 40. Lietman TM, Oldenburg CE, Keenan JD. Trachoma: Time to Talk Eradication. Ophthalmology. 2020; 127:11–13. <u>https://doi.org/10.1016/j.ophtha.2019.11.001</u> PMID:31864470
- 41. Yang X, Man J, Chen H, Zhang T, Yin X, He Q, Lu M. Temporal trends of the lung cancer mortality attributable to smoking from 1990 to 2017: A global, regional and national analysis. Lung Cancer. 2021; 152:49–57. https://doi.org/10.1016/j.lungcan.2020.12.007

PMID:<u>33348250</u>

42. Mansouri A, Emamian MH, Zeraati H, Hashemi H, Fotouhi A. Economic Inequality in Presenting Vision in Shahroud, Iran: Two Decomposition Methods. Int J Health Policy Manag. 2018; 7:59–69. <u>https://doi.org/10.15171/ijhpm.2017.48</u> PMID:29325403

- Stevens GA, Alkema L, Black RE, Boerma JT, Collins GS, Ezzati M, Grove JT, Hogan DR, Hogan MC, Horton R, Lawn JE, Marušić A, Mathers CD, et al, and The GATHER Working Group. Guidelines for Accurate and Transparent Health Estimates Reporting: the GATHER statement. Lancet. 2016; 388:e19–23. <u>https://doi.org/10.1016/S0140-6736(16)30388-9</u> PMID:<u>27371184</u>
- 44. Yu J, Yang X, He W, Ye W. Burden of pancreatic cancer along with attributable risk factors in Europe between 1990 and 2019, and projections until 2039. Int J Cancer. 2021; 149:993–1001. <u>https://doi.org/10.1002/ijc.33617</u> PMID:33937984
- Yang X, Fang Y, Chen H, Zhang T, Yin X, Man J, Yang L, Lu M. Global, regional and national burden of anxiety disorders from 1990 to 2019: results from the Global Burden of Disease Study 2019. Epidemiol Psychiatr Sci. 2021; 30:e36.

https://doi.org/10.1017/S2045796021000275 PMID:<u>33955350</u>

## SUPPLEMENTARY MATERIALS



## **Supplementary Figures**

Supplementary Figure 1. The DALYs number of BVL by different eye diseases, by SDI regions, from 1990 to 2019. The data from five SDI regions are further enlarged in its top-right panel. DALYs, disability-adjusted life years; BVL, blindness and vision loss; SDI, socio-demographic index.



Supplementary Figure 2. The age-standardized DALYs rate and the EAPC in age-standardized DALYs rate of BVL by six eye diseases for both sexes in 204 countries and territories. (A, B) glaucoma; (C, D) cataract; (E, F) age-related macular degeneration; (G, H) refraction disorders; (I, J) near vision loss; (K, L) other vision loss. DALYs, disability-adjusted life years; EAPC, estimated annual percentage change.



Supplementary Figure 3. The EAPCs of BVL burden from 1990 to 2019, both sexes, by region and eye diseases. (A) The EAPCs in ASPR of BVL; (B) The EAPCs in age-standardized DALYs rate of BVL. EAPC, estimated annual percentage change; BVL, blindness and vision loss; ASPR, age-standardized prevalence rate; DALYs, disability-adjusted life years.



Supplementary Figure 4. The changing trend in age-standardized burden rate of overall BVL across 21 GBD regions with SDI, both sexes, from 1990 to 2019. (A) ASPR; (B) age-standardized DALYs rate. BVL, blindness and vision loss; GBD, global burden of disease; SDI, socio-demographic index; ASPR, age-standardized prevalence rate; DALYs, disability-adjusted life years.

## **Supplementary Tables**

Supplementary Table 1. Prevalence cases and age-standardized prevalence rate per 100 people for all blindness and vision loss in 1990 and 2019, and its estimated annual percentage change from 1990 to 2019.

	19	90	2019		EAPC in ASPR,	
Characteristics	Prevalence cases No.×10 <sup>6</sup> (95% UI)	ASPR per 100	Prevalence cases	ASPR per 100	1990-2019 No. (95% CI)	
Overall	<b>No.×10 (95% U1)</b> 353.2 (298.7, 414.0)	No. (95% UI) 8.43 (7.07, 9.86)	<b>No.×10<sup>6</sup> (95% UI)</b> 713.9 (593.2, 841.1)	No. (95% UI) 8.69 (7.27, 10.22)	0.02 (-0.02, 0.06)	
Sex	555.2 (298.7, 414.0)	0.45 (7.07, 9.80)	/13.9 (395.2, 641.1)	8.09 (7.27, 10.22)	0.02 (-0.02, 0.00)	
Males	162.2 (136.6, 191.2)	8.18 (6.88, 9.61)	323.6 (268.9, 383.8)	8.25 (6.91, 9.73)	-0.06 (-0.10, -0.02)*	
Females	190.9 (161.4, 222.6)	8.67 (7.29, 10.11)	323.0 (208.9, 383.8) 390.4 (325.0, 458.9)	9.10 (7.61, 10.68)	0.09 (0.05, 0.12)*	
SDI	190.9 (101.4, 222.0)	8.07 (7.29, 10.11)	590.4 (525.0, 458.9)	9.10 (7.01, 10.08)	0.09 (0.03, 0.12)	
High SDI	28 20 (24 02 22 02)	207(262,224)	47 11 (41 14 52 92)	2.09 (2.72, 2.49)	0.12(0.00.0.15)*	
c	28.30 (24.92, 32.03) 85.69 (70.93, 102.1)	2.97 (2.63, 3.34)	47.11 (41.14, 53.83)	3.08 (2.72, 3.48) 8.31 (6.90, 9.88)	0.12 (0.09, 0.15)* 0.04 (0.00, 0.09)*	
High-middle SDI Middle SDI		8.00 (6.67, 9.50)	162.5 (133.2, 194.6)			
	111.8 (93.78, 132.0)	10.12 (8.44, 11.92)	247.0 (204.3, 293.6)	9.82 (8.22, 11.61)	-0.19 (-0.23, -0.14)*	
Low-middle SDI	90.84 (76.88, 106.5)	13.57 (11.47, 15.79)	179.8 (149.2, 213.2)	12.5 (10.45, 14.73)	-0.35 (-0.39, -0.31)*	
Low SDI	36.40 (30.48, 42.71)	13.63 (11.43, 15.95)	77.22 (65.04, 90.76)	13.13 (10.98, 15.37)	-0.19 (-0.25, -0.14)*	
GBD region						
High-income Asia Pacific	4.14 (3.71, 4.59)	2.25 (2.03, 2.48)	7.26 (6.47, 8.11)	2.21 (1.99, 2.44)	-0.06 (-0.08, -0.04)*	
High-income North America	7.54 (6.65, 8.54)	2.32 (2.05, 2.62)	11.91 (10.39, 13.61)	2.31 (2.04, 2.61)	-0.01 (-0.03, 0.02)	
Western Europe	14.40 (13.03, 15.89)	2.98 (2.70, 3.26)	19.60 (17.66, 21.68)	2.87 (2.59, 3.15)	-0.11 (-0.12, -0.10)*	
Australasia	0.62 (0.57, 0.68)	2.86 (2.61, 3.12)	1.04 (0.92, 1.17)	2.64 (2.35, 2.94)	-0.25 (-0.32, -0.19)*	
Tropical Latin America	8.68 (7.66, 9.77)	8.16 (7.15, 9.32)	17.90 (15.61, 20.43)	7.55 (6.60, 8.62)	-0.03 (-0.10, 0.04)	
Andean Latin America	2.17 (1.90, 2.46)	9.15 (7.95, 10.50)	4.93 (4.25, 5.66)	8.56 (7.38, 9.87)	-0.27 (-0.30, -0.24)*	
Central Latin America	8.61 (7.43, 9.96)	8.98 (7.62, 10.56)	20.40 (17.20, 24.06)	8.55 (7.22, 10.11)	-0.18 (-0.19, -0.17)*	
*Southern Latin America	1.63 (1.48, 1.80)	3.48 (3.16, 3.82)	2.52 (2.28, 2.78)	3.32 (3.01, 3.65)	-0.14 (-0.15, -0.13)*	
Caribbean	2.27 (1.91, 2.69)	8.26 (6.89, 9.86)	3.95 (3.27, 4.72)	7.72 (6.43, 9.21)	-0.25 (-0.26, -0.23)*	
Eastern Europe	25.92 (21.28, 31.08)	9.61 (7.95, 11.45)	33.05 (26.85, 40.07)	10.1 (8.27, 12.13)	0.03 (-0.03, 0.09)	
Central Europe	11.17 (8.88, 13.89)	7.79 (6.25, 9.64)	14.84 (11.75, 18.68)	7.37 (5.92, 9.13)	-0.19 (-0.20, -0.17)*	
Central Asia	4.65 (3.87, 5.51)	9.64 (7.97, 11.45)	6.93 (5.71, 8.3)	9.25 (7.67, 11.11)	-0.16 (-0.17, -0.15)*	
North Africa and Middle East	15.98 (14.19, 17.84)	7.92 (6.99, 8.95)	32.49 (28.62, 36.58)	7.04 (6.19, 8.00)	-0.39 (-0.40, -0.38)*	
South Asia	104.7 (89.03, 122.1)	16.14 (13.75, 18.72)	224.1 (187.5, 265.6)	15.00 (12.58, 17.61)	-0.34 (-0.39, -0.29)*	
Southeast Asia	26.88 (23.47, 30.48)	9.64 (8.41, 10.96)	52.77 (45.5, 60.45)	8.67 (7.54, 9.95)	-0.38 (-0.39, -0.36)*	
East Asia	84.75 (67.36, 104.5)	9.44 (7.56, 11.57)	195.9 (155.6, 240.3)	9.47 (7.66, 11.49)	-0.12 (-0.19, -0.05)*	
Oceania	0.32 (0.27, 0.37)	9.72 (8.28, 11.36)	0.73 (0.62, 0.85)	9.55 (8.14, 11.15)	-0.07 (-0.11, -0.04)*	
Western Sub-Saharan Africa	10.99 (9.06, 13.19)	11.38 (9.35, 13.64)	26.13 (21.62, 31.21)	12.00 (9.92, 14.34)	-0.03 (-0.14, 0.08)*	
Eastern Sub-Saharan Africa	10.35 (8.45, 12.3)	12.49 (10.22, 14.89)	21.47 (17.69, 25.53)	11.62 (9.57, 13.76)	-0.24 (-0.26, -0.23)*	
Central Sub-Saharan Africa	2.55 (2.01, 3.17)	10.38 (8.13, 12.84)	5.80 (4.60, 7.24)	9.89 (7.85, 12.30)	-0.16 (-0.21, -0.11)*	
Southern Sub-Saharan Africa	4.88 (3.88, 5.96)	16.18 (12.77, 19.68)	10.20 (8.00, 12.49)	16.48 (13.05, 20.19)	-0.08 (-0.15, -0.01)*	
Specific subtypes						
Glaucoma	3.88 (3.30, 4.54)	0.11 (0.09, 0.13)	7.47 (6.35, 8.77)	0.09 (0.08, 0.11)	-0.55 (-0.61, -0.50)*	
Cataract	42.34 (37.73, 47.62)	1.15 (1.03, 1.29)	97.02 (85.37, 109.7)	1.21 (1.07, 1.36)	0.41 (0.32, 0.50)*	
Age-related macular degeneration	3.58 (3.03, 4.19)	0.10 (0.08, 0.11)	7.79 (6.53, 9.16)	0.10 (0.08, 0.11)	-0.15 (-0.20, -0.11)*	
Refraction disorders	97.61 (87.51, 108.5)	2.08 (1.87, 2.31)	157.4 (140.9, 174.9)	1.96 (1.75, 2.18)	-0.25 (-0.28, -0.22)*	
Near vision loss	227.2 (164.2, 298.4)	5.61 (4.08, 7.34)	493.2 (358.9, 645.6)	5.94 (4.34, 7.77)	0.04 (-0.01, 0.10)	
Other vision loss	21.10 (19.06, 23.56)	0.52 (0.47, 0.58)	38.40 (34.35, 43.08)	0.47 (0.43, 0.53)	-0.38 (-0.42, -0.34)*	

No., number; ASPR, age-standardized prevalence rate; UI, uncertainty interval; EAPC, estimated annual percentage change; CI, confidential interval; SDI, Socio-demographic Index; GBD, global burden of disease.

\* *P* value<0.05.

Number	Location	ASPR per 100 No. (95% UI)
1	Nepal	21.27 (18.16, 24.24)
2	Tanzania	18.01 (14.90, 21.20)
3	Zimbabwe	16.97 (13.31, 20.93)
4	Lesotho	16.75 (13.16, 20.80)
5	South Africa	16.45 (13.06, 20.13)
6	Eswatini	16.27 (12.58, 20.19)
7	Namibia	15.88 (12.24, 19.87)
8	Botswana	15.78 (12.25, 19.56)
9	Niger	15.41 (13.08, 17.92)
10	India	15.29 (12.79, 18.05)
11	Bangladesh	13.78 (11.39, 16.55)
12	Nigeria	12.87 (10.63, 15.31)
13	Pakistan	12.42 (10.87, 14.02)
14	Bhutan	12.39 (9.74, 15.45)
15	Somalia	12.39 (9.57, 15.43)
16	Mozambique	11.96 (9.62, 14.61)
17	Mali	11.95 (9.82, 14.52)
18	Malawi	11.81 (9.49, 14.41)
19	South Sudan	11.59 (9.33, 14.18)
20	Burkina Faso	11.57 (9.36, 14.01)
21	Eritrea	11.54 (9.29, 13.97)
22	Guinea	11.45 (9.31, 14.02)
23	Moldova	11.40 (9.29, 13.74)
24	Guinea-Bissau	11.39 (9.31, 13.83)
25	Chad	11.38 (9.23, 13.87)
26	Madagascar	11.14 (8.80, 13.74)
27	Benin	11.07 (8.98, 13.36)
28	Comoros	11.07 (8.81, 13.53)
29	Gambia	11.05 (8.99, 13.56)

Supplementary Table 2. Countries/territories with ASPR per 100 greater than 11 in 2019.

ASPR, age-standardized prevalence rate; No., number; UI, uncertainty interval.

Number	Location	Age-standardized DALYs rate per 1000 No. (95% UI)
1	Indonesia	5.59 (4.04, 7.48)
2	Pakistan	5.39 (3.85, 7.30)
3	India	5.25 (3.66, 7.27)
4	Mali	5.13 (3.62, 7.02)
5	Timor-Leste	4.99 (3.52, 6.81)
6	Nepal	4.78 (3.14, 7.20)
7	Tanzania	4.75 (3.21, 6.73)
8	Afghanistan	4.70 (3.33, 6.33)
9	Nigeria	4.68 (3.30, 6.47)
10	Saudi Arabia	4.65 (3.32, 6.22)
11	Oman	4.55 (3.26, 6.15)
12	Bangladesh	4.48 (3.06, 6.33)
13	Cambodia	4.47 (3.17, 6.12)
14	Niger	4.34 (2.99, 6.05)
15	Lesotho	4.23 (2.88, 6.11)
16	Myanmar	4.21 (2.96, 5.75)
17	Ethiopia	4.21 (3.03, 5.69)
18	Zimbabwe	4.20 (2.87, 6.20)
19	Guinea-Bissau	4.18 (2.94, 5.82)
20	Guinea	4.04 (2.80, 5.60)
21	Senegal	3.96 (2.79, 5.47)
22	Burkina Faso	3.94 (2.76, 5.45)
23	Eswatini	3.91 (2.66, 5.76)
24	South Sudan	3.87 (2.70, 5.35)
25	Eritrea	3.86 (2.68, 5.35)
26	Botswana	3.83 (2.63, 5.65)
27	Mozambique	3.81 (2.66, 5.28)
28	Guatemala	3.80 (2.65, 5.21)

Supplementary Table 3. Countries/territories with age-standardized DALYs rate per 1000 greater than 3.8 in 2019.

DALYs, disability-adjusted life years; No., number; UI, uncertainty interval.

Number	Location	Age-standardized DALYs rate per 1000 in 1990 No. (95% UI)	Age-standardized DALYs rate per 1000 in 2019 No. (95% UI)	EAPC in age-standardized DALYs rate No. (95% CI)
1	Equatorial Guinea	6.25 (4.44, 8.41)	3.34 (2.33, 4.63)	-2.45 (-2.59, -2.31)
2	Saudi Arabia	8.04 (5.73, 10.77)	4.65 (3.32, 6.22)	-1.85 (-1.91, -1.78)
3	Cambodia	6.45 (4.54, 8.72)	4.47 (3.17, 6.12)	-1.44 (-1.52, -1.36)
4	Qatar	4.10 (2.92, 5.49)	2.87 (2.02, 3.94)	-1.29 (-1.35, -1.23)
5	Tunisia	3.97 (2.81, 5.39)	2.79 (1.97, 3.78)	-1.26 (-1.33, -1.20)
6	Malaysia	4.35 (3.07, 5.90)	3.14 (2.20, 4.35)	-1.14 (-1.17, -1.10)
7	Vietnam	4.18 (2.96, 5.71)	3.09 (2.15, 4.34)	-1.11 (-1.16, -1.07)
8	Bhutan	3.91 (2.68, 5.69)	2.90 (1.91, 4.31)	-1.10 (-1.15, -1.05)
9	Thailand	3.77 (2.59, 5.26)	2.83 (1.92, 4.10)	-1.10 (-1.16, -1.05)
10	Peru	4.56 (3.24, 6.19)	3.54 (2.47, 4.91)	-1.07 (-1.17, -0.98)
11	Lebanon	4.15 (2.94, 5.61)	3.08 (2.16, 4.19)	-1.06 (-1.09, -1.03)
12	Syria	4.45 (3.15, 5.95)	3.31 (2.34, 4.51)	-1.05 (-1.12, -0.98)
13	Bahrain	3.84 (2.74, 5.17)	2.87 (2.01, 3.92)	-1.04 (-1.10, -0.99)
14	Myanmar	5.40 (3.87, 7.34)	4.21 (2.96, 5.75)	-1.01 (-1.13, -0.88)
15	Sudan	5.10 (3.66, 6.87)	3.73 (2.67, 5.03)	-1.01 (-1.08, -0.95)
16	Iraq	4.16 (2.96, 5.58)	3.16 (2.24, 4.30)	-1.01 (-1.06, -0.97)
17	India	6.95 (4.97, 9.48)	5.25 (3.66, 7.27)	-0.98 (-1.02, -0.93)
18	Botswana	5.06 (3.47, 7.03)	3.83 (2.63, 5.65)	-0.97 (-0.99, -0.94)
19	Palestine	4.07 (2.91, 5.52)	2.99 (2.11, 4.10)	-0.97 (-1.04, -0.90)
20	Jordan	3.11 (2.21, 4.23)	2.41 (1.68, 3.34)	-0.97 (-1.00, -0.93)

Supplementary Table 4. Top 20 countries/territories with the most rapid decrease in agestandardized DALYs rate from 1990 to 2019.

DALYs, disability-adjusted life years; No., number; CI, confidence interval.

# Supplementary Table 5. Countries/territories with the increased age-standardized DALYs rate from 1990 to 2019.

Number	Location	Age-standardized DALYs rate per 1000 in 1990 No. (95% UI)	Age-standardized DALYs rate per 1000 in 2019 No. (95% UI)	EAPC in age-standardized DALYs rate No. (95% CI)
1	Burkina Faso	2.91 (2.00, 4.19)	3.94 (2.76, 5.45)	0.63 (0.35, 0.91)
2	Cote d'Ivoire	2.43 (1.65, 3.57)	3.74 (2.60, 5.21)	0.54 (0.06, 1.02)
3	Benin	3.04 (2.09, 4.40)	3.66 (2.52, 5.10)	0.52 (0.37, 0.67)
4	Central African Republic	2.20 (1.45, 3.34)	2.31 (1.53, 3.39)	0.22 (0.19, 0.26)
5	Chad	3.69 (2.56, 5.18)	3.75 (2.60, 5.18)	0.10 (0.01, 0.20)
6	Somalia	3.54 (2.46, 5.01)	3.45 (2.38, 4.87)	0.09 (0.03, 0.15)

DALYs, disability-adjusted life years; No., number; CI, confidence interval.

Number	Location	Proportion in 1990	Proportion in 2019	Proportion change
1	Luxembourg	4.40%	13.66%	9.26%
2	Seychelles	9.48%	18.17%	8.69%
3	Qatar	16.94%	25.21%	8.28%
4	Bahrain	15.83%	24.09%	8.26%
5	Mauritius	10.86%	18.97%	8.10%
6	Sri Lanka	10.54%	18.30%	7.76%
7	Micronesia	8.76%	15.69%	6.93%
8	Libya	9.63%	16.33%	6.70%
9	Fiji	14.78%	21.38%	6.60%
10	Cambodia	5.23%	11.65%	6.42%
11	Palestine	9.37%	15.75%	6.38%
12	Niue	13.22%	19.54%	6.32%
13	Marshall Islands	15.00%	21.16%	6.16%
14	Ireland	3.63%	9.70%	6.07%
15	Solomon Islands	7.84%	13.86%	6.02%
16	Afghanistan	8.37%	14.36%	5.98%
17	Sudan	7.61%	13.56%	5.95%
18	Czech	10.20%	16.13%	5.93%
19	Tunisia	9.44%	15.36%	5.92%
20	Bosnia and Herzegovina	7.61%	13.51%	5.90%

Supplementary Table 6. Top 20 countries/territories where the proportion of glaucomarelated DALYs attributable to high fasting plasma glucose increased between 1990 to 2019.

DALYs, disability-adjusted life years.

Number	Location	Proportion in 1990	Proportion in 2019	<b>Proportion change</b>
1	Bosnia and Herzegovina	16.42%	21.38%	4.96%
2	Russian	8.43%	12.05%	3.62%
3	Lebanon	19.13%	22.65%	3.52%
4	Austria	12.01%	14.86%	2.85%
5	Serbia	16.03%	18.69%	2.66%
6	Egypt	10.18%	12.83%	2.64%
7	Montenegro	20.68%	23.28%	2.60%
8	North Korea	6.94%	9.40%	2.47%
9	Kiribati	17.12%	19.23%	2.11%
10	Azerbaijan	7.21%	8.97%	1.77%
11	Mali	2.39%	4.16%	1.76%
12	Afghanistan	3.52%	5.11%	1.59%
13	Rwanda	7.91%	9.46%	1.55%
14	Qatar	8.55%	10.08%	1.53%
15	China	13.13%	14.59%	1.46%
16	Guinea	4.94%	6.33%	1.39%
17	Albania	15.70%	16.96%	1.26%
18	Estonia	11.66%	12.91%	1.25%
19	Moldova	11.70%	12.63%	0.94%
20	Antigua and Barbuda	4.58%	5.40%	0.82%
21	Cote d'Ivoire	4.56%	5.38%	0.82%
22	United Arab Emirates	8.35%	9.14%	0.79%
23	Georgia	8.10%	8.77%	0.67%
24	Indonesia	9.43%	10.09%	0.66%
25	Saudi Arabia	7.55%	8.20%	0.65%
26	Niger	1.21%	1.78%	0.57%
27	Sao Tome and Principe	2.48%	2.93%	0.45%
28	Kyrgyzstan	8.71%	8.95%	0.24%
29	Uzbekistan	4.48%	4.71%	0.22%
30	Bermuda	6.98%	7.19%	0.21%
31	Gabon	2.65%	2.77%	0.12%
32	Vietnam	10.44%	10.49%	0.05%
33	Malawi	5.14%	5.17%	0.02%

Supplementary Table 7. Countries/regions where the proportion of DALYs in age-related macular degeneration attributable to smoking increased between 1990 to 2019.

DALYs, disability-adjusted life years.

Number	Location	Proportion in 1990	Proportion in 2019	Proportion change
1	Myanmar	18.73%	8.40%	-10.33%
2	Canada	21.32%	12.00%	-9.32%
3	Netherlands	21.73%	12.46%	-9.27%
4	Ireland	22.09%	12.92%	-9.17%
5	New Zealand	16.86%	8.25%	-8.61%
6	Brazil	18.33%	9.86%	-8.47%
7	South Africa	13.68%	6.21%	-7.47%
8	United Kingdom	19.47%	12.48%	-7.00%
9	USA	19.00%	12.14%	-6.86%
10	Venezuela	12.82%	6.21%	-6.60%
11	Iceland	19.18%	12.71%	-6.47%
12	Thailand	15.17%	8.78%	-6.39%
13	Mexico	11.67%	5.40%	-6.27%
14	Australia	14.57%	8.69%	-5.88%
15	Brunei Darussalam	14.49%	8.75%	-5.74%
16	Japan	18.08%	12.50%	-5.58%
17	Croatia	22.47%	16.95%	-5.52%
18	Colombia	10.70%	5.29%	-5.41%
19	Denmark	23.31%	17.99%	-5.32%
20	Sri Lanka	9.59%	4.41%	-5.18%

Supplementary Table 8. Top 20 countries/regions where the proportion of DALYs in agerelated macular degeneration attributable to smoking decreased between 1990 to 2019.

DALYs, disability-adjusted life years.