### Prognostic factors for a change in eye health or vision: A rapid review

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### Abstract:

The general public are advised to have regular routine eye examinations to check their vision and ocular health; however current UK guidance on how often to have eve examinations is not evidence-based and was issued in 2002. This Rapid Review aims to provide an evidence base that stakeholders can use to form updated guidance for Wales by asking the question 'What are the prognostic factors for a change in ocular status in the general population attending routine eye examinations?'

The review included evidence available from January 2009 up until August 2023. Evidence was included from 2011 up until 2023. 19 studies were included: two systematic reviews; nine prospective cohort studies; three retrospective cohort studies; two longitudinal studies; two case-control studies; and one cross-sectional study were included.

### **Research Implications and Evidence Gaps:**

Future research to inform appropriate eye examination intervals should be narrower in focus to ensure as much relevant and useful evidence as possible is gathered. There are large amounts of evidence on prevalence and prognostic factors for prevalent conditions, which did not meet the inclusion criteria of this rapid review which looks at incident or changing conditions.

### **Policy and Practice Implications:**

Caution should be taken if using this review for decision making on appropriate eye examination intervals due to low certainty and generalisability. This review should be used to identify key prognostic factors and suggesting these for further targeted research and evidence synthesis.

### **Economic considerations:**

Sight loss costs the UK economy 25 billion pounds per annum, with more than 2 million people in the UK currently living with sight loss. The economic implications of appropriate or inappropriate testing intervals for different causes of vision loss will be different. When captured at a population wide scale, the earlier detection of conditions through examination can result in significant economic savings.

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### Prognostic factors for a change in eye health or vision: A rapid review

### Report number RR0010 (January 2024)

# EXECUTIVE SUMMARY

### What is a Rapid Review?

Our rapid reviews (RR) use a variation of the systematic review approach, abbreviating or omitting some components to generate the evidence to inform stakeholders promptly whilst maintaining attention to bias.

### Who is this summary for?

This Rapid Review is intended for use by clinical leaders and decision makers in Wales' primary eye care services. The evidence in this Review is intended to be used to examine the risk of a person experiencing a change in their ocular health, vision, or systemic health that affects their eves so that guidance can be produced on how often people should attend for routine eve examinations based on their individual risk factors.

It is also intended to identify gaps in the evidence to determine where further research is required for certain risk factors or patient groups.

### **Background / Aim of Rapid Review**

The general public are advised to have regular routine eye examinations to check their vision and ocular health; however current UK guidance on how often to have eye examinations is not evidence-based and was issued in 2002.

This Rapid Review aims to provide an evidence base that stakeholders can use to form updated guidance for Wales by asking the question "What are the prognostic factors for a change in ocular status in the general population attending routine eye examinations?"

### Results

### Recency of the evidence base

The review included evidence available from January 2009 up until August 2023. Evidence was included from 2011 up until 2023.

### Extent of the evidence base

19 studies were included: two systematic reviews; nine prospective cohort studies; three retrospective cohort studies; two longitudinal studies; two case-control studies; and one cross-sectional study were included.

### Key findings and certainty of the evidence

- Demographic prognostic factors: age, sex, ethnicity, and household net worth are potential prognostic factors for a change in ocular health or vision.
- Ocular prognostic factors: intraocular pressure, family history of glaucoma, visual acuity, visual field mean deviation, spherical equivalent refraction, high myopia, age-related macular degeneration, glaucoma, and cataract are potential prognostic factors for a change in ocular health or vision.

- Lifestyle/behaviour prognostic factors: diet, alcohol intake, smoking, time spent outdoors, and time spent reading are potential prognostic factors for a change in ocular health or vision.
- Systemic health prognostic factors: hypertension, heart disease, cholesterol, diabetes, peripheral arterial disease, hypercoagulable state, stroke, pregnancy, age at menarche, oral contraceptive use, and atopy are potential prognostic factors for a change in ocular health or vision.
- Increasing length of time between eye examinations is a potential prognostic factor for a change in ocular health or vision.
- The level of certainty for all prognostic factors is low as there was generally only one study reporting for each individual outcome.
- Studies were often performed in specific populations, meaning the results cannot be applied to the general population, particularly due to low study numbers per outcome.

### **Research Implications and Evidence Gaps**

- Future research to inform appropriate eye examination intervals should be narrower in focus to ensure as much relevant and useful evidence as possible is gathered. Prognostic factors or specific ocular conditions of interest potentially need to be investigated individually for their effect on a change in ocular status.
- There are large amounts of evidence on prevalence and prognostic factors for prevalent conditions, which did not meet the inclusion criteria of this rapid review which looks at incident or changing conditions. Further evidence generation could be conducted in this area.
- Very little evidence was identified in a UK setting, more primary evidence generation may be required.
- There is a notable lack of evidence in younger adults aged under 40 years.

### **Policy and Practice Implications**

- Caution should be taken if using this review for decision making on appropriate eye examination intervals due to low certainty and generalisability.
- This review should be used to identify key prognostic factors and suggesting these for further targeted research and evidence synthesis.

### **Economic considerations**

- Sight loss costs the UK economy £25 billion per annum, with more than 2 million people in the UK currently living with sight loss.
- The economic implications of appropriate or inappropriate testing intervals for different causes of vision loss will be different.
- A new case of age-related macular degeneration (AMD) in an adult aged 50 or over, costs the UK economy £73,350 over the person's lifetime. Lifetime costs to the UK economy for a person diagnosed with glaucoma are approximately £49,800 per person. Reducing the prevalence of these conditions by just 14 or 20 cases respectively could save the UK economy £1 million in lifetime costs.
- On economic grounds, early detection of AMD in eye care services and the eye care pathway may be of benefit due to the high level of prevalence and associated long term costs to the NHS as the condition causes irreversible, life limiting damage.
- When captured at a population wide scale, the earlier detection of conditions through examination can result in significant economic savings.

**Disclaimer:** The views expressed in this publication are those of the authors, not necessarily Health and Care Research Wales. The Health and Care Research Wales Evidence Centre and authors of this work declare that they have no conflict of interest.

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### Abbreviations:

Acronym	Full Description	
AMD	Age-related macular degeneration	
CI	Confidence interval	
CRVO	Central retinal vein occlusion	
HR	Hazard ratio	
IOP	Intraocular pressure	
OR	Odds ratio	
QUIPS	Quality in Prognostic factor Studies	
RCT	Randomised controlled trial	
ROBIS	Risk of Bias in Systematic reviews	
RR	Risk ratio / relative risk	
SER	Spherical equivalent refraction	
VA	Visual acuity	

# 1. BACKGROUND

### 1.1 Who is this review for?

This Rapid Review was conducted as part of the Health and Care Research Wales Evidence Centre Work Programme. The original question was suggested by the National Clinical Leads for Wales General Ophthalmic Services and the Optometry and Audiology Policy Branch, Welsh Government. Working with these stakeholders, the question was then amended to the one mentioned above.

This Rapid Review is intended for use by clinical leaders and decision makers in Wales' primary eye care services. The evidence in this Review is intended to be used to examine the risk of a person experiencing a change in their ocular health, vision, or systemic health that affects their eves so that guidance can be produced on how often people should attend for routine eye examinations based on their individual risk factors.

It is also intended to identify gaps in the evidence to determine where further research is required for certain risk factors or patient groups.

### **1.2** Background and purpose of this review

The general public are advised to attend routine eye examinations to regularly check their visual acuity, provide any necessary vision correction, and identify ocular health problems. Current guidance in the UK was issued in 2002 and is based on a consensus decision regarding minimum re-examination intervals, with no evidence base. The currently recommended minimum intervals are:

- Under 16 years in the absence of any binocular vision anomaly 1 year
- Under 7 years with binocular vision anomaly or corrected refractive error 6 months •
- 7 years and over and under 16 with binocular vision anomaly or rapidly progressing • myopia - 6 months
- 16 years and over and under 70 years 2 years •
- 70 years and over 1 year •
- 40 years and over with a family history of glaucoma or with ocular hypertension and not in a monitoring scheme - 1 year
- Diabetic patients 1 year •

With significant reform of Wales General Ophthalmic Services underway, it is pertinent to review the evidence that is available that may be able to inform recommendations on the frequency of routine eye examinations in Wales.

The evidence identified in this review will be used by stakeholders to help answer questions similar to the below:

- What is the risk of an asymptomatic person attending for a routine eye examination having experienced a change in ocular status?
- Is there evidence to suggest that this risk may vary between different groups?
- Can the evidence regarding this risk be used to inform appropriate time intervals between routine eye examinations?

# 2. RESULTS

### 2.1 Overview of the Evidence Base

Nineteen studies were identified that met the inclusion criteria of this rapid review. Two systematic reviews and 17 primary studies were included, all of which are observational studies. The study designs varied, with nine prospective cohort studies; three retrospective cohort studies; two longitudinal studies; two case-control studies; and one cross-sectional study included. Sample sizes were also very different across studies, with some having only a few hundred participants whilst others had more than 400,000. Full details of the eligibility criteria are presented in Section 5.1, Table 7. Full details of the included studies and the extracted data can be found in Section 6.2, Tables 8 and 9.

The results of this rapid review have been categorised into common prognostic factors. These factors are demographic, ocular, lifestyle/behaviour, and systemic health related. The factors are then further categorised into specific prognostic factors or similar categories of prognostic factors.

As the scope of this prognostic factor review is broad, and the review conducted with exploratory aims, essential restrictions were put in place to make sure the review remained tenable within the limits of rapid review methodology. Included studies were therefore limited to a pre-determined list of countries (Section 5.1, Table 7) that were determined by the review team to have similar demographics and eye care systems to the UK. Studies were also included only if they had presented their findings as odds ratios, risk ratios/relative risk, or hazard ratios – an approach that is in line with other prognostic factor reviews (Riley et al. 2019) – and had used multivariate analyses to determine these. The use of multivariate analyses means, where an association has been identified, the reported prognostic factors have an effect on the outcome that is independent of the other factors controlled for. Factors controlled for in each study are included in Table 8; there was considerable variation in the types of factors and the number of factors controlled for in each study. Owing to this being a rapid review, it was not feasible to convert other types of outcomes into ratios as is sometimes done in prognostic factor reviews. This is discussed in the limitations of this review.

Using the Quality in Prognostic factor Studies (QUIPS) tool, all but one of the primary evidence sources were determined to be of low or moderate risk of bias across all six domains of the tool. Ten of the 17 studies were assessed as moderate risk of bias for prognostic factor or outcome measurement, with eight of these relying on self-reporting, leading to increased risk of recall bias. Similarly, there were concerns regarding loss to follow-up or study attrition in seven studies and it was unclear whether the strategy for model building was appropriate and based on a conceptual framework or model in four studies. One study (Barsam et al. 2017) was determined to be at high risk of bias due to its case-control design and only including a small number of the cases in multivariable modelling.

Using the Risk of Bias in Systematic reviews (ROBIS) tool, both systematic reviews included in this rapid review were deemed to have either low risk of bias (Kessel et al. 2015) or an unclear risk of bias (Dinu et al. 2019). For the systematic review deemed unclear, issues were centred around the failure to address heterogeneity, and a lack of clarity on whether subgroup analyses were pre-specified. Both studies included meta-analyses and the results of these were extracted for this rapid review. None of the identified primary evidence sources were included in either of the systematic reviews.

### 2.2 Demographic prognostic factors

Results for this section are summarised in Table 1 with comprehensive details available in Section 6.2, Tables 8 and 9.

### 2.2.1 Age

Nine identified studies examined the association of age with various ocular/vision conditions, including five prospective cohort analyses, one retrospective cohort analysis, one population-based longitudinal study, one cross-sectional study, and one case-control study. All of the included observational studies were of low to moderate risk of bias, with three studies rated as moderate due to some measures being self-reported and lack of clarity whether the strategy for multivariate model building was appropriate.

Only three studies reported on the same outcome, and found that aging is associated with an increased risk of developing open-angle glaucoma in adults aged 65 to 74 years, 55 to 84 years, and 55 years or over, respectively (Ekström 2012, Ekström & Hårleman 2023, Marcus et al. 2012). The two studies by Ekström were, however, identified by the review team as having a high risk of double reporting. Kang et al. (2012) found that increasing age is also a risk factor for exfoliation glaucoma or being an exfoliation glaucoma suspect. When compared to 40 to 55 year olds, the rate ratio increased for every 5-year bracket, with 55 to 60 year olds having approximately four times (rate ratio 4.33) the risk of developing exfoliation glaucoma or being a suspect case and those over 75 years old having approximately 46 times the risk (rate ratio 46.22).

Aging is also associated with increased odds of needing an eye care referral in adults over 50 years of age (Keel et al. 2017) or experiencing any kind of change in ocular status (Irving et al. 2016). For every one-year increase in age, the odds of having a change in ocular status increased by 3%.

Aging is associated with a reduced risk of experiencing a myopic change in refractive error at five years in adults aged 35 to 74 years, by 48% per year (Stingl et al. 2023). At the same time, it is associated with increased risk of experiencing a hyperopic change in refractive error over five years in adults aged 35 to 74 years, with 62% increased risk per year (Stingl et al. 2023).

Age was found not to be associated with the risk of progression of myopic maculopathy in high myopes (people with extreme or severe near-sightedness) aged between 35 and 74 (Hopf et al. 2022) or with developing visual field damage in glaucoma suspects of African or European descent (Khachatryan et al. 2015).

### 2.2.2 Sex

Eight studies were identified that examined the effect of sex as a prognostic factor for a change in ocular status. These included four prospective cohort studies, one retrospective cohort analysis, one longitudinal study, one cross-sectional study and one case-control study. All included studies were identified as having low to moderate risk of bias, with concerns around the use of self-reported measures in four studies.

Male sex was found to be a risk factor for requiring an eye care referral in adults over 50 years of age, with 24% higher odds than females (Keel et al. 2017). Males were also found to be at higher risk of developing open-angle glaucoma in adults aged 55 years and older, with 37% higher risk (Marcus et al. 2012). However, females are more likely to be diagnosed with exfoliation glaucoma or a suspect case of this, with males having less (32%) chance than females do (Kang et al. 2012). Females are also nearly 50% more likely than males to

experience a myopic change in refractive error over five years in adults aged 35 to 74 years (Stingl et al. 2023).

Sex was not found to be associated with the risk of developing myopia in children (Guggenheim et al. 2012), or open-angle glaucoma in adults aged 55 to 84 years (Ekström & Hårleman 2023), progression of myopic maculopathy in high myopes aged 35 to 74 years (Hopf et al. 2022), or experiencing a change in ocular status (Irving et al. 2016).

### 2.2.3 Ethnicity/race

Two identified studies looked into the effect of ethnicity/race in relation to different eye conditions. This included a prospective cohort analysis and a retrospective cohort analysis. Both observational studies were judged to be at low risk of bias.

Glaucoma suspects, defined as eyes with a history of elevated intraocular pressure (IOP) and/or an optic disc appearance suspicious of glaucoma but normal visual fields at baseline in this study, of African descent were at higher risk of developing visual field damage than those of European descent if their mean IOP was 22 mmHg or higher with the hazard ratio increasing as IOP increased (Khachatryan et al. 2015). The study found that those with a mean IOP of 22 mmHg had double the risk of their European counterparts, whilst the risk was more than 3.5 times greater with a mean IOP of 26 mmHg. The study found that there was no significant association with race at IOPs of 10 to 20 mmHg.

Black ethnicity was also associated with increased risk of developing central retinal vein occlusion (CRVO) compared to White ethnicity in adults aged 55 years and over (hazard ratio 1.58 [95% confidence interval (CI) 1.25 to 1.99]) (Stem et al. 2013). Asian-American ethnicity was not deemed to be a risk factor according to this study.

### 2.2.4 Socioeconomic characteristics

Various socioeconomic factors were examined as potential prognostic factors in three identified studies. These were a prospective cohort study, a retrospective cohort study, and a cross-sectional study. All three studies were rated as low or low to moderate risk of bias.

Geographical remoteness and years of education were not found to be associated with the risk of eye care referral in adults over 50 years of age by Keel et al. (2017). This study was conducted in Australia and, thus there are concerns about the generalisability of this evidence to Wales due to much greater remoteness and distances to major urban settlements in Australia.

Education was found to not be associated with change in refractive error at five years in adults aged 35 to 74 years (Stingl et al. 2023). This same study also found that occupation is not associated with change in refractive error.

Lower household net worth was found to be associated with increased risk of developing CRVO in adults 55 years and older in an American study (Stem et al. 2013). Those with a household net worth of greater than US\$500,000 had 27% lower risk of developing CRVO than those with a net worth less than US\$25,000 (hazard ratio 0.73 [95% CI 0.56 to 0.96]).

### 2.2.5 Bottom line results for demographic prognostic factors

The evidence identifies suggests that age, sex, ethnicity, and household net worth are potential risk factors for changes in vision or ocular health. Aging and increasing age is associated with a general increased risk of change in ocular status, while sex, ethnicity and household net worth are dependent on the outcome examined. No studies were identified that examined the use of index of deprivation as a prognostic factor.

Across all studies in this review the certainty of the evidence is low due to the paucity of evidence for each outcome - with only one study identified in many cases. Though the evidence was deemed to be at low or moderate risk of bias, further research is necessary to inform any decision making in this area.

	Index	Ohic prognostic 1	
Citation (Country)	prognostic factor	Cultonio	Adjusted prognostic effect (95% confidence intervals) and interpretation
Ekström (2012)	Age	Incident OAG in adults aged 65-	Age (per year) HR 1.15 (1.05 to 1.26)
(Sweden)		74 years	Increasing age is associated with increased risk of incident OAG in adults aged 65-74 years.
n = 976			
Ekström &	Age	Incident OAG in	75-84 years OR 3.02 (1.13 to 8.08)
Hårleman (2023)		adults aged 55- 84 years	65-74 years OR 1.15 (0.44 to 3.00) 55-64 years (ref)
(Sweden)			Increasing age is associated with increased risk
n = 481			of incident OAG in adults aged 55-84 years.
Hopf et al. (2022)	Age	Progression of myopic maculopathy at 5	Age (per year) OR 0.94 (0.88 to 1.02, p = 0.134)
(Germany)		years in adults aged 35-74 years	Age is not associated with increased risk of myopic maculopathy progression at 5 years in
n = 350	A	Oʻrusifin su t	adults aged 35-74 years.
Irving et al. (2016)	Age	Significant change in optical	Age (per year) OR 1.03 (1.03 to 1.04)
(Canada)		status (see Table 9 for full description) in all	Increasing age is associated with increased risk of experiencing a significant change in ocular status.
n = 2656		ages	
Kang et al. (2012)	Age	Incident exfoliation	Rate ratio (RR) of age: 40 to 55 years (ref)
(USA)		glaucoma or exfoliation	55 to 60 years RR 4.33 (2.19 to 8.56) 60 to 65 years RR 10.43 (5.50 to 19.78)
n = 120,146		glaucoma suspect in adults	65 to 70 years RR 19.88 (10.41 to 37.96) 70 to 75 years RR 33.54 (17.23 to 65.29) Over 75 years RR 46.22 (22.77 to 93.80)
			Increasing age is associated with higher risk for incident exfoliation glaucoma or exfoliation glaucoma suspect.
Keel et al. (2017)	Age	Rates of eye care referral in adults	Age OR 1.02 (1.01 to 1.02, p < 0.001)
(Australia)		aged 50 years and over	Increasing age is associated with higher eye care referral rates in adults aged 50 years and
n = 3098			older.
Khachatryan et al. (2015)	Age	Incident visual field damage in	Age (per year) HR 1.02 (0.99 to 1.04)
(USA)		glaucoma suspects of African or	Age is not associated with increased risk of visual field damage.
n = 357		European descent	
Marcus et al. (2012)	Age	Incident OAG in adults aged 55	Age (per year) OR 1.06 (1.04 to 1.09, p < 0.001)
(The Netherlands)		year and over	Age is associated with increased risk of incident OAG in adults aged 55 years and older.
n = 3939			

Table 1: Summary of demographic prognostic factors

Stingl et al.	Age	Change in	Myopic change at 5 years:
(2023)		refractive error at	Age (per year) OR 0.52 (0.49 to 0.55, p <0.001)
		5 years in adults	
(Germany)		aged 35-74 years	Hyperopic change at 5 years:
10.175			Age (per year) OR 1.62 (1.52 to 1.72, p <0.001)
n = 10,175			
			Decreasing age is associated with increased
			risk of having a myopic shift in refractive error at
			5 years in adults aged 35 to 74 years.
			Increasing age is associated with an increased
			risk of a hyperopic shift in refractive error at 5
Ekström &	Sex	Incident OAG in	years in adults aged 35 to 74 years. Sex (male) OR 1.77 (0.91 to 3.43)
Hårleman (2023)	Sex	adults aged 55-	Sex (male) OK 1.77 (0.91 (0.3.43)
naneman (2023)		84 years	Sex is not associated with increased risk of
(Sweden)		04 years	OAG in adults aged 55-84 years.
(Oweden)			
n = 481			
Guggenheim et	Sex	Incident myopia	OR 1.058 (0.810 to 1.382, p = 0.679)
al. (2012)	UCX	after age 11	on 1.000 (0.010 to 1.002; p = 0.010)
un (2012)		alter age i i	Sex is not associated with incident myopia after
(UK)			age 11.
(0.1)			~g~
n = 2005			
Hopf et al. (2022)	Sex	Progression of	Sex (female) OR 5.54 (0.93 to 32.92, p = 0.060)
		myopic	
(Germany)		maculopathy at 5	Sex is not associated with increased risk of
		years in adults	myopic maculopathy progression at 5 years in
n = 350		aged 35-74 years	adults aged 35-74 years.
Irving et al.	Sex	Significant	Sex (female) OR 1.07 (0.90 to 1.29)
(2016)		change in optical	
		status (see Table	Sex is not associated with increased risk of
(Canada)		9 for full	significant change in ocular status.
		description) in all	
n = 2656	-	ages	
Kang et al. (2012)	Sex	Incident	Rate ratio (RR) of gender:
(110.4.)		exfoliation	Male RR 0.32 (0.23 to 0.46)
(USA)		glaucoma or	Female (ref)
n = 120,146		exfoliation	Eamole pay is appaciated with higher risk for
11 = 120,140		glaucoma	Female sex is associated with higher risk for incident exfoliation glaucoma or exfoliation
		suspect in adults	-
Keel et al. (2017)	Sex	Rates of eye care	glaucoma suspect. Sex (male) OR 1.24 (1.06 to 1.46, p = 0.007)
	UEX .	referral in adults	Sex (male) Six 1.24 (1.00 to 1.40, p = 0.007)
(Australia)		aged 50 years	Male sex is associated with higher eye care
(/ tubti alia)		and over	referral rates in adults aged 50 years and older.
n = 3098			
Marcus et al.	Sex	Incident OAG in	Sex (female) OR 0.63 (0.43 to 0.93, p = 0.022)
(2012)		adults aged 55	
x = 7		year and over	Male sex is associated with increased risk of
(The		-	incident OAG in adults aged 55 years and older.
Netherlands)			
n = 3939			
Stingl et al.	Sex	Change in	Sex (female) OR 1.49 (1.28 to 1.73, p < 0.001)
(2023)		refractive error at	
		5 years in adults	Female sex is associated with increased risk of
(Germany)		aged 35-74 years	having a myopic shift in refractive error at 5
			years in adults aged 35 to 74 years.
n = 10,175			-
Khachatryan et	Ethnicity/	Incident visual	African descent vs. European descent by IOP:
al. (2015)	race	field damage in	No significant association at IOP = 10 mmHg to
		glaucoma	20 mmHg
(USA)		suspects of	IOP 22 mmHg HR 2.03 (1.15 to 3.57)
n = 357		African or	IOP 24 mmHg HR 2.71 (1.39 to 5.29)
	1	1	IOP 26 mmHg HR 3.61 (1.61 to 8.08)

		European descent	Mean IOP of cohort (17.8 mmHg) HR 1.12 (0.66 to 1.90)
			Glaucoma suspects of African descent with higher mean IOP are associated with increased risk of visual field damage compared to European glaucoma suspects.
Stem et al. (2013)	Ethnicity/	Incident CRVO in	Ethnicity:
(USA)	race	adults aged 55 years and over	Black HR 1.58 (1.25 to 1.99, p < 0.0001) Asian-American HR 0.75 (0.43 to 1.30, p = 0.31)
n = 494,165			White (ref)
			Black ethnicity is associated with higher risk for incident CRVO in adults aged 55 years and over.
Keel et al. (2017)	Education	Rates of eye care	Years of education OR 0.98 (0.96 to 1.00, $p =$
(Australia)		referral in adults aged 50 years	0.11)
n = 3098		and over	Years of education is not associated with eye care referral rates in adults aged 50 years and older.
Stingl et al.	Education	Change in	Myopic change at 5 years:
(2023)		refractive error at	Secondary general school (ref)
( /		5 years in adults	Intermediate school OR 0.96 (0.81 to 1.14, p =
(Germany)		aged 35-74 years	0.64)
			High school OR 0.96 (0.81 to 1.12, p = 0.58)
n = 10,175			Others OR 0.52 (0.17 to 1.57, p = 0.25)
			None OR 0.91 (0.28 to 2.94, p = 0.88)
			Hyperopic change at 5 years: Secondary general school (ref) Intermediate school OR 1.11 (0.98 to 1.26, p = 0.10) High school OR 1.03 (0.91 to 1.16, p = 0.67) Others OR 0.64 (0.27 to 1.52, p = 0.31) None OR 1.20 (0.44 to 3.27, p = 0.72) Education is not associated with increased risk
			of either myopic or hyperopic shift at 5 years in adults aged 35 to 74 years.
Keel et al. (2017)	Geographical remoteness	Rates of eye care referral in adults	Geographical remoteness OR 1.04 (0.979 to $1.10, p = 0.27$ )
(Australia)		aged 50 years and over	Geographical remoteness is not associated with
n = 3098			eye care referral rates in adults aged 50 years and older.
Stem et al. (2013)	Net worth / wealth	Incident CRVO in adults aged 55	Household net worth: > \$500,000 HR 0.73 (0.56 to 0.96, p = 0.02)
(USA)		years and over	< \$25,000 (ref)
n = 494,165			Lower household net worth is associated with higher risk for incident CRVO in adults aged 55 years and over.
Stingl et al.	Occupation	Change in	ORs ranged from 0.81 to 1.28, all 95% CI
(2023)		refractive error at 5 years in adults	included 1.00, p ≥ 0.05 for all.
(Germany)		aged 35-74 years	Occupation is not associated with increased risk of either myopic or hyperopic shift at 5 years in
n = 10,175			adults aged 35 to 74 years.

### 2.3 Ocular prognostic factors

Results for this section are summarised in Table 2 with comprehensive details available in Section 6.2, Tables 8 and 9.

### 2.3.1 Vision-related

Five identified studies looked at vision-related characteristics as prognostic factors for changes in ocular status. Four were prospective cohort studies and one was a case-control study. Three of the studies were rated as low to moderate risk of bias, with one (Barsam et al. 2017) deemed high risk of bias, due to the case-control study design meaning that the prognostic data was collected after the outcome was known and the low number of cases (21%) included in the multivariate model.

In a case-control study comparing people with keratoconus, a progressive condition causing thinning and irregular curvature of the cornea, who developed acute corneal hydrops (a sight-threatening complication of keratoconus that can leaving scarring) to those who did not, Barsam et al. (2017) found that worse visual acuity (VA) was associated with increased odds of developing corneal hydrops.

A prospective cohort study by Khachatryan et al. (2015) of glaucoma suspects found that a worse result on visual field assessment (mean deviation) at baseline was a risk factor for developing visual field damage (hazard ratio 1.04 (95% CI 1.02 to 1.06) per 0.1 dB decrease).

In a study by Stingl et al. (2023), baseline spherical equivalent refraction (SER) was found to be a risk factor for having a myopic shift in refractive error at five years in adults aged 35 to 74 years, with increasing myopia being a greater risk factor for a myopic shift (11% per dioptre more myopic). However, SER was not found to be associated with the risk of developing visual field damage in glaucoma suspects (Khachatryan et al. 2015) or for the progression of myopic maculopathy in high myopes aged 35 to 74 years (Hopf et al. 2022). High myopia was found to be associated with increased risk of incident open-angle glaucoma in adults aged 55 years and over (Marcus et al. 2012).

### 2.3.2 Ocular pathology

Four studies identified looked at various ocular pathologies as prognostic factors for other ocular pathology or changes in refractive error. This included one systematic review and meta-analysis, one prospective cohort analysis, one retrospective cohort analysis, and one case-control study. The meta-analysis was judged to have low concerns for risk of bias and the three observational studies were judged to be of low to moderate risk of bias, with two studies unclear whether the strategy for model building was appropriate.

Stem et al. (2013) investigated whether age-related macular degeneration (AMD), openangle glaucoma, and cataract were prognostic factors for developing CRVO in adults of 55 years of age and older. They found that all three were associated with increased risk of developing CRVO with increased risks of 50%, 50% and 24% respectively compared to those without these conditions.

It was found that the presence of cataract is not associated with increased risk of a change in refractive error in adults aged 35 to 74 years (Stingl et al. 2023), and a meta-analysis of four studies found that undergoing cataract surgery was not associated with increased risk of progression to wet AMD in people with dry AMD 6 to 12 months after surgery (Kessel et al. 2015). Stingl et al. note that their findings are contrary to other cohort studies which report an association between nuclear cataract and a myopic shift in refractive error. They suggest this may be explained by the lack of differentiation of nuclear and cortical cataract in their study.

The presence of pseudoexfoliation was found not to be a risk factor for developing openangle glaucoma in adults aged 55 to 84 years; odds ratio 1.27 (95% CI 0.63 to 2.57) (Ekström & Hårleman 2023). This is thought to be due to the strong interaction between increased IOP and the presence of pseudoexfoliation (see Section 2.3.3) and, therefore, pseudoexfoliation is not independently associated with increased risk of open-angle glaucoma.

### 2.3.3 Intraocular pressure

Five identified studies investigated IOP as a prognostic factor for ocular pathology or a change in refractive error. There were three prospective cohort analyses, one longitudinal study, and a case-control study. All studies were assessed as low to moderate risk of bias.

Ekström (2012) and Ekström & Hårleman (2023) confirmed that increasing IOP is associated with increased risk of open-angle glaucoma, something which is well established as stated in the studies. While pseudoexfoliation is not independently associated with glaucoma, mean IOP greater than or equal to 25 mmHg concurrent with pseudoexfoliation is associated with greater risk of incident open-angle glaucoma in 65 to 74 year olds (Ekström 2012).

A study of glaucoma suspects by Khachatryan et al. (2015) found that increasing IOP was not independently associated with increased risk of developing visual field damage in suspects of African or European descent; HR 0.97 per 1 mmHg increase (95% CI 0.92 to 1.03). However, as stated in Section 2.2, the risk of visual field damage in glaucoma suspects of African descent compared to European descent did show a positive correlation with mean IOP increase.

Hopf et al. (2022) found that increasing IOP increased the risk of progression of myopic maculopathy in high myopes aged 35 to 74 years by 62% per mmHg at 5 years. Whilst another study found that IOP is not associated with increased risk of a change in refractive error at 5 years in adults aged 35 to 74 years (Stingl et al. 2023).

### 2.3.4 Family history

Four studies were identified that investigated family history as a potential prognostic factor. These include two prospective cohort studies, a longitudinal study, and a case-control study. All three studies were judged as low to moderate risk of bias, with self-reporting of measures being a factor.

Three of the studies investigated positive family history of glaucoma as a risk factor for different types of glaucoma. Positive family history of glaucoma was found to be associated with over double (rate ratio 2.29) the risk of developing exfoliation glaucoma or becoming an exfoliation glaucoma suspect in adults (Kang et al. 2012). Positive history was associated with more than double (OR 2.24) the risk of developing open-angle glaucoma in adults aged 55 years and over in a study from The Netherlands (Marcus et al. 2012) and more than three times (OR 3.21) increased risk in adults aged 55 to 84 years (Ekström & Hårleman 2023).

Positive parental history of myopia was not associated with increased risk of children becoming myopic after age 11 years (Guggenheim et al. 2012). This was true for children with only one myopic parent or both parents.

### 2.3.5 Ocular parameters

Two prospective cohort studies were identified that examined other ocular parameters as potential prognostic factors. One study was rated as low risk of bias, whilst Kang et al. (2012) was rated low to moderate risk of bias due to self-reporting of both outcomes and prognostic factors.

It was found that central corneal thickness is not associated with risk of developing visual field damage in glaucoma suspects of African or European descent (Khachatryan et al. 2015). Eve colour was also found not to be associated with the risk of developing exfoliation glaucoma or becoming a exfoliation glaucoma suspect in adult men (Kang et al. 2012).

### Bottom line results for ocular prognostic factors 2.3.6

The evidence identified shows that various ocular prognostic factors, which could be identified during a routine eye examination, were identified as risk factors for the development other ocular pathology or changes in vision or refractive error. This includes VA, visual field mean deviation, SER, various ocular pathologies, IOP, and family history of glaucoma.

As with section 2.2, it is pertinent to note that the evidence summarised in this section is specific to the outcomes mentioned above, and cannot be extrapolated to cover all eve conditions. In many cases, a lack of evidence means that there is often only one study per prognostic factor / outcome pairing, and therefore the certainty of the evidence is low. There were two studies that reported positive family history of glaucoma as a risk factor for incident open-angle glaucoma and the confidence in this finding is also higher. Most included primary studies were of low or moderate risk of bias, with one at high risk of bias, whilst the included meta-analysis states that the studies it included were of moderate or very low quality.

Citation	Index prognostic	Outcome	Adjusted prognostic effect (95% confidence
(Country)	factor		intervals) and interpretation
Barsam et al. (2017)	Visual acuity	Incident acute corneal hydrops in people with	VA in worse eye OR 4.11 (1.18 to 14.32, p = 0.026) Having worse visual acuity is associated with
(UK)		keratoconus	higher odds of developing acute corneal hydrops in people with keratoconus.
n = 159			
Khachatryan et al. (2015)	Visual field mean deviation	Incident visual field damage in glaucoma	Baseline visual field mean deviation (per 0.1 dB decrease) HR 1.04 (1.02 to 1.06)
(USA)		suspects of African or	Lower baseline visual field mean deviation is associated with increased risk of visual field
n = 357	5 ( )	European descent	damage in glaucoma suspects.
Hopf et al. (2022)	Refractive error	Progression of myopic maculopathy at 5	SER (per dioptre) OR 1.21 (0.99 to 1.49, $p = 0.063$ ) SER is not associated with increased risk of myopic
(Germany)		years in adults aged 35-74 years	maculopathy progression at 5 years in adults aged 35-74 years.
n = 350			
Khachatryan et al. (2015)	Refractive error	Incident visual field damage in	Lower SER (per D greater) HR 1.11 (0.84 to 1.34)
(USA)		glaucoma suspects of African or	SER is not associated with increased risk of visual field damage in glaucoma suspects.
n = 357		European descent	
Marcus et al. (2012)	Refractive error	Incident OAG in adults aged 55	High myopia OR 2.22 (1.13 to 4.38, $p = 0.021$ )
(The Netherlands)		year and over	High myopia is associated with increased risk of incident OAG in adults aged 55 years and older.
n = 3939			

Table 2: Summary of ocular prognostic factors

	1	r	
Stingl et al. (2023)	Refractive error	Change in refractive error at 5 years in adults	Baseline SER (per dioptre) OR 0.89 (0.87 to 0.91, p < 0.001)
(Germany)		aged 35-74 years	Baseline myopic SER is associated with increased risk of having a myopic shift in refractive error at 5
n = 10,175 Stem et al.	AMD	Incident CRVO in	years in adults aged 35 to 74 years. AMD HR 1.50 (1.31 to 1.72, p < 0.0001)
(2013)	AIVID	adults aged 55	AND TIC 1.50 (1.51 to $1.72$ , $p < 0.0001$ )
(USA)		years and over	AMD is associated with higher risk for incident CRVO in adults aged 55 years and over.
n = 404.165			
n = 494,165 Stem et al.	Cataract	Incident CRVO in	Cataract HR 1.24 (1.08 to 1.42, p = 0.003)
(2013)	Outuruot	adults aged 55	
(USA)		years and over	Cataract is associated with higher risk for incident CRVO in adults aged 55 years and over.
n = 494,165			
Stingl et al.	Cataract	Change in	Myopic change at 5 years:
(2023)		refractive error at 5 years in adults	Lens opacity OR 1.09 (0.91 to 1.30, p = 0.36)
(Germany)		aged 35-74 years	Hyperopic change at 5 years: Lens opacity OR 1.02 (0.91 to 1.16, p = 0.70)
n = 10,175			Lens opacity is not associated with increased risk of either myopic or hyperopic shift at 5 years in
			adults aged 35 to 74 years.
Kessel et al.	Cataract surgery	Progression of non-exudative	Meta-analysis results for progression of non-
(2015)		AMD to exudative	exudative AMD to exudative AMD after cataract surgery (follow-up 6 to 12 months):
(UK,		AMD 6-12 months	RR 1.33 (0.60-2.94) [Total], RR 3.21 (0.14-75.68)
Australia,		after undergoing	[RCTs], RR 1.25 (0.55-2.85) [case-control]
Germany,		cataract surgery in	
Austria)		adults	Cataract surgery is not associated with increased
4 studies			risk of progression to exudative AMD 6-12 months after surgery.
(only 3			anel surgery.
included in			
meta-			
analysis), n			
= 1574	On en en ele		
Stem et al. (2013)	Open-angle glaucoma	Incident CRVO in adults aged 55	OAG HR 1.50 (1.30 to 1.72, p < 0.0001)
(2010)	giadoonia	years and over	Open-angle glaucoma is associated with higher risk
(USA)		, , , , , , , , , , , , , , , , , , ,	for incident CRVO in adults aged 55 years and
			over.
n = 494,165 Ekström &	Dooudoovfaliation	Incident OAG in	Decude ovtaliation OD 4.27 (0.62 to 0.57)
Ekstrom & Hårleman	Pseudoexfoliation	adults aged 55-84	Pseudoexfoliation OR 1.27 (0.63 to 2.57)
(2023)		years	Pseudoexfoliation is not associated with increased
			risk of OAG in adults aged 55-84 years. The effect
(Sweden)			of pseudoexfoliation on glaucoma risk is mediated
n = 494			by elevated IOP.
n = 481 Ekström	IOP	Incident OAG in	Mean IOP ≥ 25 mmHg and pseudoexfoliation HR
(2012)		adults aged 65-74 years	2.38 (1.87 to 3.03)
(Sweden)		, 50.0	Time-dependent (per 10 years):
			Mean IOP ≥ 25 mmHg HR 15.4 (4.52 to 52.1)
n = 679			Mean IOP 20-24.99 mmHg HR 3.92 (2.13 to 7.22) Mean IOP < 20 mmHg (ref)
			Increasing IOP is associated with increased risk of
			incident OAG in adults aged 65-74 years.
			Mean IOP ≥ 25 mmHg concurrent with
			pseudoexfoliation is associated with an increased
			risk of incident OAG in adults aged 65-74 years.

Ekotröm 8	IOP	Incident OAC in	IOP (per 5 mmHg) OR 4.04 (2.91 to 5.62)
Ekström & Hårleman	IOP	Incident OAG in adults aged 55-84	10P (per 5 mmHg) OK 4.04 (2.91 to 5.62)
(2023)		years	Increasing IOP is associated with increased risk of
(2020)		years	incident OAG in adults aged 55-84 years.
(Sweden)			
(0			
n = 481			
	IOP	Progression of	IOP (per mmHg) OR 1.62 (1.51 to 1.59, p = 0.035)
(2022)		myopic	
,		maculopathy at 5	Increasing IOP is associated with increased risk of
(Germany)		years in adults	progression of myopic maculopathy at 5 years in
		aged 35-74 years	adults aged 35-74 years.
n = 350			
	IOP	Incident visual	Mean IOP (per 1 mmHg increase) HR 0.97 (0.92 to
et al. (2015)		field damage in	1.03)
		glaucoma	
(USA)		suspects of	Increasing IOP is not associated with increased risk
		African or	of visual field damage in glaucoma suspects of
n = 357		European descent	African or European descent.
Ų	IOP	Change in	Myopic change at 5 years: IOP per mmHg OR 1.01 (0.99 to 1.03, p = 0.54)
(2023)		refractive error at 5 years in adults	10P per mm g OR 1.01 (0.99 to 1.03, $p = 0.54$ )
(Germany)		aged 35-74 years	Hyperopic change at 5 years:
(Cermany)		aged 55-7 + years	IOP OR 0.99 (0.97 to 1.00, $p = 0.06$ )
n = 10,175			101 OK 0.00 (0.07 to 1.00, p = 0.00)
11 - 10,110			Increasing IOP is not associated with increased risk
			of either myopic or hyperopic shift at 5 years in
			adults aged 35 to 74 years.
Ekström &	Family history of	Incident OAG in	Positive family history of OAG OR 3.21 (1.38 to
	glaucoma	adults aged 55-84	7.45)
(2023)	-	years	
			Positive family history of OAG is associated with
(Sweden)			increased risk of incident OAG in adults aged 55-84
101			years.
n = 481			
	Family history of	Incident exfoliation	Rate ratio (RR) of family history of glaucoma:
(2012)	glaucoma	glaucoma or exfoliation	Positive history RR 2.29 (1.39 to 3.78) Negative history (ref)
(USA)		glaucoma suspect	Negative history (lef)
(00Å)		in adults	Positive family history of glaucoma is associated
n = 120,146		in dduito	with higher risk for incident exfoliation glaucoma or
			exfoliation glaucoma suspect for adults.
Marcus et al.	Family history of	Incident OAG in	Positive family history of glaucoma OR 2.24 (1.31
	glaucoma	adults aged 55	to 3.84, p = 0.003)
	0	year and over	
(The		-	Positive family history of glaucoma is associated
Netherlands)			with increased risk of incident OAG in adults aged
			55 years and older.
n = 3939	_		
	Parental myopia	Incident myopia	1 myopic parent OR 1.175 (0.900 to 1.533, p =
et al. (2012)		after age 11	0.236)
(1112)			2 myopic parent OR 1.143 (0.718 to 1.818, p =
(UK)			0.574)
n = 2005			No myopic parents (ref)
11 - 2000			Parental myopia is not associated with incident
			myopia after age 11.
Khachatryan	Corneal thickness	Incident visual	Central corneal thickness (per 40 microns thinner)
et al. (2015)		field damage in	HR 1.18 (0.86 to 1.60)
		glaucoma	- (
(USA)		suspects of	Decreasing central corneal thickness is not
			associated with increased risk of developing visual
		African or	
n = 357		African or European descent	field damage in glaucoma suspects of African or European descent.

Kang et al. (2012) (USA)	Eye colour	Incident exfoliation glaucoma or exfoliation glaucoma suspect in adults	Rate ratio (RR) of eye colour (males only): Hazel/green/medium RR 0.87 (0.43 to 1.74) Brown/dark RR 0.84 (0.42 to 1.68) Blue/light (ref)
n = 120,146			Eye colour is not associated with increased risk of incident exfoliation glaucoma or exfoliation glaucoma suspect for adult men.

### 2.4 Interval between eye examinations

Results for this section are summarised in Table 3 with comprehensive details available in Section 6.2, Tables 8 and 9.

Three of the identified studies investigated whether the length of time between eye examinations is a prognostic factor for changes in ocular health. This included two retrospective cohort analyses and a cross-sectional study. All three studies were of low or low to moderate risk of bias.

Keel et al. (2017) and Wright et al. (2020) examined whether increased time between eye examinations affected eye care referral rates and rates of referral to a general practitioner (GP), respectively. The odds of requiring an eye care referral increased 15% per year since last examination in the study by Keel et al. (2017) in adults aged 50 years and older. Delayed attendance at eye examinations was associated with 30% increased odds of requiring referral to a GP for 60 to 69 year olds and a 7% increase for those aged 70 years or over (Wright et al. 2020). Early attendance for an eye examination was associated with nearly three times increased odds of requiring GP referral in both age groups. However, this was believed to be due to early attendance usually being in response to the patient noticing symptomatic problems or because the optometrist recommended early assessment at the last examination.

The relationship between the time elapsed since previous eye examination and the odds of experiencing a change in ocular status (such as a change in vision or glasses prescription, emergence of new pathology, requiring a referral) was investigated by Irving et al. (2016). The study found that increasing the length of time elapsed between examinations is associated with an increased risk of experiencing a significant change in ocular status, with a 6% increase in risk per year.

### 2.4.1 Bottom line results for intervals between eye examinations

Increasing length of time between eye examinations is associated with increased risk of experiencing a change in ocular status, including requiring onward referral. The confidence in this finding is high as three studies reported this as a prognostic factor for similar outcomes. Due to the study designs of Keel et al. (2017) and Wright et al. (2020), it is possible that some of the participants in these studies were experiencing symptomatic eye issues.

Table 3: Summary of studies examining interval between eye examinations as a prognostic factor

Citation (Country)	Index prognostic factor	Outcome	Adjusted prognostic effect (95% confidence intervals) and interpretation
Keel et al.	Interval between	Rates of eye care	Time since last eye examination OR 1.15 per year
(2017)	eye examinations	referral in adults aged 50 years and	(1.12 to 1.19, p < 0.001)
(Australia)		over	Longer time period since last eye examination is associated with higher eye care referral rates in
n = 3098			adults aged 50 years and older.

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Irving et al. (2016)	Interval between eye examinations	Significant change in optical status (see Table 9 for	Interval between eye examinations (per year) OR 1.06 (1.02 to 1.11)
(Canada) n = 2656		full description) in all ages	Increasing length of time between eye examinations is associated with increased risk of experiencing a significant change in ocular status.
Wright et al. (2020) (UK)	Interval between eye examinations	Referral to a GP in adults aged 60 years and over	Aged 60-69: Delayed eye exam attendance OR 1.30 (1.04 to 1.61) Early eye exam attendance OR 2.86 (2.36 to 3.46)
n = 132,046			Aged ≥ 70: Delayed eye exam attendance OR 1.07 (1.01 to 1.13) Early eye exam attendance OR 2.72 (2.58 to 2.87)
			Delayed attendance for eye examinations is associated with increased risk of requiring a GP referral for adults aged 60 years and older. Early attendance is also associated with increased risk of referral for adults aged 60 years or older, though this is driven by early attendance usually being due to symptomatic problems or early recall
			suggested by the optometrist.

### 2.5 Lifestyle/behaviour prognostic factors

Results for this section are summarised in Table 4 with comprehensive details available in Section 6.2, Tables 8 and 9.

### 2.5.1 Diet

Three of the identified studies investigated diet or alcohol intake as prognostic factors for changes in ocular status. This included one systematic review and meta-analysis, one prospective cohort study, and one longitudinal population-based cohort study. The two observational studies were rated as moderate risk of bias due to uncertainties around study attrition and some self-reporting of measures. The meta-analysis was rated as having unclear concerns of risk of bias due to heterogeneity in the studies not being addressed and a lack of clarity as to whether subgroup analyses were pre-specified.

In a meta-analysis of 26 studies, Dinu et al. (2019) found that higher meat intake is associated with higher risk of the occurrence or progression of early AMD, increasing the risk by 17%. Increasing alcohol intake was found to be associated with increased risk of all AMD and early AMD (20% and 29% increased risk respectively) but not with late AMD. Increased dietary intake of fish was found to have a protective effect against AMD with the risk of early AMD being reduced by 16%, late AMD by 21%, and all AMD by 18%. An increased intake of dairy products, plant products, and fats was not found to be associated with risk of AMD.

In a study on post-menopausal women, Elmore et al. (2022) found no association between dietary intake of fish or fatty acids and incident AMD. The study also investigated red blood cell fatty acid levels as a longer-term biomarker of fatty acid intake and still found no association between levels of any red blood cell polyunsaturated fatty acid levels and incident AMD.

Gopinath et al. (2014) investigated the effect of diet on the 5-year incidence of dual sensory impairment (concurrent visual and hearing impairment). The study found no association between having a higher total diet score (healthier diet) and the incidence of dual sensory impairment in adults aged over 49 years.

### 2.5.2 Smoking

One identified study investigated smoking as a prognostic factor. This was a prospective cohort study. The study was rated as low to moderate risk of bias due to issues with study attrition and lack of clarity on whether the model building strategy was appropriate.

Smoking is associated with increased risk of experiencing a hyperopic change in refractive error at five years in adults aged 35 to 74 years (Stingl et al. 2023), with the risk increasing by 31%. However, the study found that being an occasional smoker or former smoker does not increase the risk of having a hyperopic change in prescription compared to non-smokers.

### 2.5.3 Activity-related

Two studies were identified that investigated whether activity levels, or the types of activities done, are prognostic factors for refractive changes. One was a prospective cohort analysis and the other was a longitudinal study. Both observational studies were of low to moderate risk of bias, with study attrition being a common issue.

Guggenheim et al. (2012) examined the role of time spent outdoors and physical activity on myopia prevalence and progression. The study found that increased time spent outdoors was independently associated with lower risk of developing myopia after the age of 11 years, with the risk reducing by 35%. It also found that increased time spent reading was associated with a 32% increased risk of developing myopia after age 11, but that the amount of physical activity done/amount of sedentary time was not independently associated with the risk of developing myopia.

Stingl et al. (2023) also found that the amount of physical activity was not associated with increased risk of a change in refractive error at five years in adults aged 35 to 74 years.

### 2.5.4 Bottom line results for lifestyle/behaviour prognostic factors

Dietary intake of meat, fish, and alcohol are prognostic factors for incident AMD. However, there is discrepancy in the literature on the protective effect of fish intake in the sub-group of post-menopausal women.

Smoking is potentially a risk factor for hyperopic changes in refractive error. Increased time spent outdoors may protect against children developing myopia, whilst reading may be a risk factor for it. However, the amount of time being physically active does not appear to be associated with changes in refractive error.

As before, though all the studies included in this section of the rapid review are of moderate or low risk of bias, the findings are of low certainty due to the fact that there is generally only one study for each prognostic factor/outcome pairing. The findings therefore cannot be generalised beyond the outcome or specific populations that are presented in this section.

	minaly of mesty	yic/bellaviour p	
Citation (Country)	Index prognostic factor	Outcome	Adjusted prognostic effect (95% confidence intervals) and interpretation
Dinu et al. (2019)	Alcohol intake	Incident AMD or progression of AMD in adults	Total AMD RR 1.20 (1.04-1.39, p = 0.01), Early AMD RR 1.29 (1.16-1.43, p < 0.001), Late AMD RR 0.98 (0.76-1.27)
(USA, Australia, The Netherlands,			Increasing alcohol intake is associated with increased risk of all AMD and early AMD, but not late AMD.

Table 4: Summary of lifestyle/behaviour prognostic factors

Denmark, Iceland, Japan, South Korea)			
12 studies, n = 120,440			
Dinu et al. (2019) (USA, Australia, The Netherlands, Denmark, Iceland, Japan, South Korea) Meat: 6 studies, n = 101,011 Dairy	Diet	Incident AMD or progression of AMD in adults	Meta-analysis results for incidence or progressionof AMD:Animal products:Meat Total AMD RR 1.11 (0.96-1.27, $p = 0.16$ ), EarlyAMD RR 1.17 (1.02-1.34, $p = 0.03$ ), Late AMD RR 0.99(0.70-1.39)Dairy products Total AMD RR 1.07 (0.68-1.70, $p = 0.77$ ), Early AMD RR 1.18 (0.93-1.50), Late AMD RR0.97 (0.27-3.48)Fish Total AMD RR 0.82 (0.75-0.90, $p < 0.001$ ), EarlyAMD RR 0.82 (0.75-0.90, $p < 0.001$ ), EarlyAMD RR 0.84 (0.73-0.97, $p = 0.02$ ), Late AMD RR 0.79(0.70-0.90, $p < 0.001$ )Plant products:Vegetables Total AMD RR 0.92 (0.82-1.03, $p = 0.33$ ),Early AMD RR 0.92 (0.67-1.25), Late AMD RR 0.80(0.76-1.00)Fruits Total AMD RR 0.91 (0.82-1.01, $p = 0.08$ ), Early
products: 3 studies, n = 73,772 Fish: 8 studies, n = 237,464			AMD RR 0.92 (0.82-1.03), Late AMD RR 0.83 (0.62- 1.12) Nuts Total AMD RR 0.81 (0.64-1.02, $p = 0.08$ ), Early AMD RR 0.73 (0.51-1.04), Late AMD RR 0.83 (0.62- 1.10) Grains Total AMD RR 0.84 (0.62-1.13, $p = 0.25$ )
Vegetables: 4 studies, n = 133,904 Fruits: 3 studies, n = 132,525 Nuts: 3 studies, n = 4711 Grains: 2 studies, n = 4335			Fats: Oils Total AMD RR 1.10 (0.98-1.23, p = 0.12), Early AMD RR 1.13 (0.93-1.37), Late AMD RR 1.05 (0.53- 2.07) Butter Total AMD RR 1.04 (0.93-1.16, p = 0.49), Early AMD RR 0.99 (0.75-1.30), Late AMD RR 0.85 (0.49- 1.47) Margarine Total AMD RR 1.05 (0.91-1.21, p = 0.54), Early AMD RR 1.07 (0.85-1.35), Late AMD RR 0.98 (0.56-1.70)
Oils: 2 studies, n = 77,078 Butter: 2 studies, n = 7862 Margarine: 3 studies, n = 79,336			Increasing dietary meat intake is associated with increased risk of early AMD. Increasing dietary fish intake is associated with decreased risk of all AMD, early AMD, and late AMD. Increasing dietary dairy product, plant product is not associated with risk of AMD.
Elmore et al. (2022) (USA) n = 1076	Diet	Incident AMD in post- menopausal women	RBC polyunsaturated fatty acid levels: No significant association between any RBC polyunsaturated fatty acid levels and incident AMD Dietary intake of fatty acids: No significant association between dietary intake of
			any fatty acids and incident AMD Dietary intake of fish: ≥ 1 serving per week HR 0.91 (0.53 to 1.58) ≥ 1 serving per month and < 1 serving per week HR 0.86 (0.48 to 1.54)

			None or < 1 serving per month (ref)
			Dietary intake of dark fish: ≥ 1 serving per week HR 1.20 (0.79 to 1.81) ≥ 1 serving per month and < 1 serving per week HR 0.60 (0.34 to 1.04) None or < 1 serving per month (ref) There is no association between fatty acid intake or
			fish intake and incident AMD in post-menopausal women.
Gopinath et al. (2014) (Australia) n = 2443	Diet	Incidence of dual sensory impairment at 5 years in adults aged over 49 years	Total diet score 5 <sup>th</sup> quintile vs. 1 <sup>st</sup> quintile and dual sensory impairment OR 1.03 (0.30 to 3.50) Diet is not associated with 5-year incidence of dual sensory impairment in adults aged over 49 years.
Stingl et al. (2023) (Germany) n = 10,175	Smoking	Change in refractive error at 5 years in adults aged 35- 74 years	Smoker OR 1.31 (1.14 to 1.50, p < 0.001) Occasional smoker OR 1.11 (0.69 to 1.42, p = 0.95) Former smoker OR 1.03 (0.98 to 1.21, p = 0.13) Non-smoker (ref) Being a smoker is associated with an increased risk of a hyperopic shift in refractive error at 5 years in adults
Guggenheim	Time spent	Incident myopia	aged 35 to 74 years. OR 0.65 (0.45 to 0.96, p = 0.029)
et al. (2012) (UK)	outdoors	after age 11	Increased time spent outdoors is associated with lower risk of developing myopia after age 11.
n = 2005			
Guggenheim et al. (2012) (UK)	Time spent reading	Incident myopia after age 11	OR 1.323 (1.023 to 1.712, p = 0.033) Increased time spent reading is associated with increased risk of developing myopia after age 11.
n = 2005			
Guggenheim et al. (2012) (UK)	Amount of physical activity/sedentary time	Incident myopia after age 11	Physical activity/sedentary behaviour: Mean counts per minute for whole week: OR 0.887 (0.773 to 1.017, $p = 0.086$ ) Time with moderate to vigorous activity per day:
n = 2005			OR 0.877 (0.764 to 1.006, $p = 0.062$ ) Time with sedentary counts: OR 1.095 (0.959 to 1.251, $p = 0.180$ ) Physical activity/sedentary behaviour are not associated with incident myopia after age 11.
Stingl et al. (2023) (Germany) n = 10,175	Physical activity	Change in refractive error at 5 years in adults aged 35- 74 years	Myopic change at 5 years: Physical activity OR 1.00 (1.00 to 1.00, p = 0.28) Hyperopic change at 5 years: Physical activity OR 1.00 (1.00 to 1.00, p = 0.73)
			Physical activity is not associated with increased risk of either myopic or hyperopic shift at 5 years in adults aged 35 to 74 years.

### Systemic health prognostic factors 2.6

Results for this section are summarised in Table 5 with comprehensive details available in Section 6.2, Tables 8 and 9.

### 2.6.1 Cardiovascular/vascular issues

Five of the identified studies examined cardiovascular or vascular issues as prognostic factors for ocular changes. This included two prospective cohort studies, one retrospective cohort study, one cross-sectional study, and one case-control study. All the studies were of low or low to moderate risk of bias.

Treated hypertension was found not to be associated with increased risk of incident openangle glaucoma in adults aged 55 to 84 years, however the effect of untreated hypertension was not investigated (Ekström & Hårleman 2023). However, increasing mean arterial pressure was associated with increased risk of visual field damage in glaucoma suspects of African or European descent, with a 3% increased risk per 1 mmHg (Khachatryan et al. 2015). Stem et al. (2013) found that hypertension is associated with increased risk of incident CRVO in adults aged 55 years and over. The risk increased by two-thirds compared to those without hypertension. Those with hypertension and high cholesterol still had increased risk of CRVO compared to those without these conditions but lower increased risk than hypertension alone (46% compared with 66%). Having hypertension, diabetes, and high cholesterol lead to CRVO risk increasing by 58%.

Peripheral arterial disease and hypercoagulable state were both found to be associated with higher risk of CRVO, increasing the risk 1.15 and 2.45 times respectively (Stem et al. 2013). Previous stroke was also associated with higher risk of CRVO (HR 1.44 [95% CI 1.23 to 1.68, p < 0.0001). However, self-reported previous stroke was not associated with higher risk of requiring eye care referral in adults aged 50 years or over (Keel et al. 2017).

Previous myocardial infarction was associated with a lower risk of incident CRVO in adults aged 50 years and over (Stem et al. 2013). The chance of developing CRVO was reduced by 28%; this may be due to treatments given after the myocardial infarction has happened. Ischaemic heart disease was also found to be associated with higher risk of incident openangle glaucoma in adults aged 55 to 84 years, more than doubling the risk (Ekström & Hårleman 2023).

Stingl et al. (2023) found that a range of cardiovascular parameters were not associated with change in refractive error at five years in adults aged 35 to 74 years.

### 2.6.2 Diabetes

Two of the identified studies examined diabetes as a prognostic factor. These were a retrospective cohort analysis and a cross-sectional study. The retrospective cohort analysis was rated as low risk of bias, whilst the cross-sectional study was rated low to moderate risk due to self-reporting of some prognostic factors and only including covariates that were significant in univariate analysis in the multivariable model.

Having hypertension and diabetes was associated with higher risk of developing CRVO in adults aged 55 years and over, increasing the risk by 82% (Stem et al. 2013). As mentioned in Section 2.5.1, diabetes alongside hypertension and high cholesterol increased the risk of incident CRVO in this population by 58%.

Self-reported history of diabetes was not found to be associated with eye care referral rates in adults aged 50 years and over (Keel et al. 2017).

### 2.6.3 Women's health

Two studies were identified that examined various aspects of women's health/reproductive health as prognostic factors for eye conditions. Both were prospective cohort analyses. Fernández-Montero et al. (2017) was rated as moderate risk of bias due to self-reporting of outcomes and prognostic factors and due to the loss to follow-up rate. Pasquale & Kang (2011) was also rated as moderate risk for similar reasons.

Fernández-Montero et al. (2017) found that pregnancy was associated with lower risk of developing myopia or progression of existing myopia in women aged 20 to 50 years. This risk was decreased by 42% compared to women who were not pregnant and was proposed to be due to increased time spent outdoors during periods of maternity leave.

For women aged 40 and over, age at menarche was associated with increased risk of incident normal tension glaucoma (Pasquale & Kang 2011). Those with age at menarche older than 13 had 47% increased risk compared to those with age at menarche under 12 years. The length of time between menarche and menopause (reproductive duration) was not associated with risk of incident open-angle glaucoma.

Pasquale & Kang (2011) also found that oral contraceptive use was associated with the risk of incident open-angle glaucoma in women 40 years of age and over. Five years or greater use of oral contraceptives and time since discontinuing use of oral contraceptives less than 10 years were associated with increased risk of open-angle glaucoma. The risk was increased by 25% and 39% respectively compared to those who had never used oral contraceptives. Whether a woman had had children or not, and the number of children they have had, was not associated with incident open-angle glaucoma.

### 2.6.4 Other systemic health issues

In a case-control study examining risk factors for acute corneal hydrops in keratoconics, two atopic conditions were found to be associated with increased risk of developing corneal hydrops (Barsam et al. 2017). History of vernal conjunctivitis increased the risk by 15 times and asthma increased the risk by nearly five times. However, both had very wide 95% CI, suggesting these results are not very precise, and the multivariable model only included 15 cases and 144 controls out of the samples of 64 and 1794 respectively. This study was rated as high risk of bias due to the case-control design and the prognostic data being collected after the outcome was known and the low number of cases included in the multivariable model.

When investigating adults over 55 years of age, Marcus et al. (2012) found that use of any type of corticosteroid medications was not associated with increased risk of incident openangle glaucoma. They acknowledged that this is contradictory to many other studies' findings, but also noted studies with consistent findings that often found that corticosteroid use is associated with increased IOP but not necessarily with a diagnosis of glaucoma. This study was rated as moderate risk of bias due to a fairly low number of participants having follow-up data and some measures being self-reported.

### 2.6.5 Bottom line results for systemic health prognostic factors

There are many systemic health conditions that can be risk factors or protective factors for ocular pathology, refractive error, or the need for eye care referral. These include hypertension, high cholesterol, diabetes, heart disease, peripheral artery disease, stroke, hypercoagulable state, and atopy. However, this list is not exhaustive. Many of these were associated with CRVO.

Factors related to reproductive health, such as current pregnancy, use of oral contraceptives, and age at menarche, are also linked to glaucoma and myopia. Use of corticosteroids was not identified as a risk factor for glaucoma.

All findings for this section are of low certainty due to the lack of evidence available. Further research is necessary to understand the relationship between systemic health factors and visual health. These findings cannot be extrapolated beyond the outcomes or populations explored in this section.

Citation (Country)	Index prognostic factor	Outcome	Adjusted prognostic effect (95% confidence intervals) and interpretation
Ekström & Hårleman (2023)	Hypertension	Incident OAG in adults aged 55-84 years	Treated hypertension OR 0.58 (0.29 to 1.15) Treated hypertension is not associated with risk of incident OAG in adults aged 55 to 84 years.
(Sweden)			
n = 481			
Khachatryan et al. (2015)	Hypertension	Incident visual field damage in glaucoma	Mean arterial pressure (per 1 mmHg increase) HR 1.03 (1.00 to 1.06)
(USA) n = 357		suspects of African or European descent	Increasing mean arterial pressure is associated with increased risk of visual field damage in glaucoma suspects of African or European descent.
Stem et al. (2013) (USA)	Hypertension / high cholesterol / diabetes	Incident CRVO in adults aged 55 years and over	Hypertension HR 1.66 (1.14 to 2.42, $p = 0.01$ ) Hypertension and diabetes HR 1.82 (1.15 to 2.89, $p = 0.01$ ) Hypertension and hyperlipidaemia HR 1.46 (1.04 to 2.05,
n = 494,165			p = 0.03) Hypertension, hyperlipidaemia, and diabetes HR 1.58 (1.11 to 2.23, $p = 0.01$ ) No diabetes, hypertension, or hyperlipidaemia (ref)
			Hypertension (alone or in combination with diabetes, hyperlipidaemia, or both) is associated with higher risk for incident CRVO in adults aged 55 years and over.
Stem et al. (2013) (USA)	Peripheral artery disease	Incident CRVO in adults aged 55 years and over	HR 1.15 (1.00 to 1.33, p = 0.05) Peripheral artery disease is associated with higher risk for incident CRVO in adults aged 55 years and over.
n = 494,165 Stem et al.	Hypercoagulable	Incident CRVO	HR 2.45 (1.40 to 4.28, p = 0.002)
(2013) (USA)	state	in adults aged 55 years and over	Hypercoagulable state is associated with higher risk for incident CRVO in adults aged 55 years and over.
n = 494,165			
Stem et al. (2013)	Stroke	Incident CRVO in adults aged	HR 1.44 (1.23 to 1.68, p < 0.0001)
(USA)		55 years and over	Stroke is associated with higher risk for incident CRVO in adults aged 55 years and over.
n = 494,165			
Keel et al. (2017) (Australia)	Stroke	Rates of eye care referral in adults aged 50 years and over	OR 1.00 (0.99 to 1.00, p = 0.64) Previous stroke is not associated with eye care referral rates in adults aged 50 years and older.
n = 3098			
Stem et al. (2013)	Myocardial infarction	Incident CRVO in adults aged	HR 0.72 (0.57 to 0.92, p = 0.01)
(USA)		55 years and over	Previous myocardial infarction is associated with lower risk of incident CRVO in adults aged 55 years and over.

 Table 5: Summary of systemic health prognostic factors

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n = 494,165			
Ekström & Hårleman (2023)	Ischaemic heart disease	Incident OAG in adults aged 55-84 years	OR 2.41 (1.15 to 5.06) Ischaemic heart disease is associated with increased risk of incident OAG in adults aged 55-84 years.
(Sweden)			
n = 481	O and in the second and	Oh e a se in	
Stingl et al. (2023)	Cardiovascular parameters	Change in refractive error at 5 years in	ORs ranged from 0.98 to 1.07, all 95% CI included 1.00, $p > 0.05$ for all.
(Germany) n = 10,175		adults aged 35-74 years	Cardiovascular parameters are not associated with change in refractive error at 5 years in adults aged 35 to 74 years.
Keel et al.	Diabetes	Rates of eye	OR 0.83 (0.67 to 1.04, $p = 0.11$ )
(2017)		care referral in adults aged 50	History of diabetes is not associated with eye care
(Australia) n = 3098		years and over	referral rates in adults aged 50 years and older.
Fernández- Montero et	Pregnancy	Incident myopia or	HR 0.58 (0.49 to 0.69, p < 0.001)
al. (2017) (Spain)		progression of myopia in women aged	Pregnancy is associated with a decreased risk of developing myopia or progression of existing myopia in women aged 20-50 years.
n = 10,401		20-50 years	
Pasquale &		Incident OAG	> 13 years and normal tension glaucoma RR 1.47 (1.01
(USA)	Age at menarche	in women aged 40 years and over	to 2.13) < 12 years (ref)
(USA) n = 79,440			Age at menarche older than 13 years is associated with increased risk of normal tension glaucoma in women aged 40 years or older.
Pasquale &	Reproductive	Incident OAG	< 36 years RR 0.93 (0.71 to 1.22)
Kang (2011)	duration	in women aged 40 years and over	36-38 years RR 0.94 (0.73 to 1.21) 39-40 years (ref) ≥ 41 years RR 0.96 (0.73 to 1.27)
(USA)		and over	241 years RR 0.90 (0.75 to 1.27)
n = 79,440			Reproductive duration is not associated with OAG in women aged 40 years or older.
Pasquale &	Oral	Incident OAG	Ever used RR 1.14 (0.98 to 1.34) < 2 years RR 1.10 (0.89 to 1.36)
Kang (2011)	contraceptives,	in women aged 40 years	2-4 years RR 1.04 (0.89 to 1.36)
(USA)	duration of use	and over	5+ years RR 1.25 (1.02 to 1.53)
n = 79,440			Never used (ref)
			Five years or greater use of oral contraceptives is associated with increased risk of OAG in women aged 40 years or older.
Pasquale &	Oral	Incident OAG	≥ 25 years RR 1.13 (0.91 to 1.40)
Kang (2011)	contraceptives,	in women	20-24 years RR 1.06 (0.82 to 1.38)
(USA)	time since discontinuing use	aged 40 years and over	15-19 years RR 1.20 (0.91 to 1.59) < 10 years RR 1.39 (1.01 to 1.91) Never used (ref)
n = 79,440			Time since discontinuing use of oral contraceptives less than 10 years is associated with increased risk of OAG
Pasquale &	Parity		in women aged 40 years or older.
Pasquale & Kang (2011)	Parity	Incident OAG in women aged 40 years	No children RR 0.85 (0.60 to 1.21) 1-2 children (ref) 3 children RR 1.08 (0.90 to 1.29)
(USA)		and over	4+ children RR 1.00 (0.84 to 1.19)

n = 79,440			Parity is not associated with OAG in women aged 40 years or older.
Barsam et al. (2017)	Atopy	Incident acute corneal hydrops in	Vernal conjunctivitis OR 15.00 (1.30 to 173.70, p = 0.026)
(UK)		people with keratoconus	Asthma OR 4.92 (1.22 to 19.78, p = 0.025)
n = 159			History of vernal conjunctivitis and asthma are associated with higher odds of developing acute corneal hydrops in people with keratoconus.
Marcus et al. (2012)	Corticosteroid use	Incident OAG in adults aged	Ophthalmic steroid use OR 1.04 (0.66 to 1.65, p = 0.86)
(The		55 year and over	Inhaled steroid use OR 0.79 (0.42 to 1.48, p = 0.46)
Netherlands)			Nasal steroid use OR 1.26 (0.74 to 2.13, p = 0.40)
n = 3939			Oral steroid use OR 1.03 (0.65 to 1.64, p = 0.89)
			Ointment steroid use OR 0.70 (0.47 to 1.05, p = 0.086)
			Corticosteroid use is not associated with OAG incidence in adults aged 55 years and older.

 Table 6: Summary of findings and numbers of studies reporting each finding

		Dei	nogra	ographic factors Ocular factors Ocular															Life	style	/beha	viou	r facto	stors Systemic health factors																					
				Soc	cioec fact	onom ors	nic	V	ision- fact		ed	(	Dcula	r patł	nolog	y			nily tory	Oc par ete	am			Di	et				ctivit <u>y</u> elated		Cai	diova	iscula	ar/va	scula	r issu	es		V	Vome	en's h	nealth	1	Oth	ier
	Aging	Sex (male)	Ethnicity (African/Black)	Geographical Remoteness	Years of education	Occupation	Lower household net worth	Worse visual acuity	Worse visual field mean deviation	SER	High myopia	AMD	Glaucoma	Cataract	Cataract surgery	Pseudoexfoliation	Higher IOP	Positive family history of glaucoma	Parental myopia	Central corneal thickness	Eye colour	Increasing time between eye examinations	Higher meat intake	Higher fish intake	Overall diet quality	Higher alcohol intake	Smoking	Increasing time outdoors	Increasing time reading	Amount of physical activity	HBP	High cholesterol	Peripheral arterial disease	Hypercoagulable state	Previous stroke	Previous myocardial infarction	Ischaemic heart disease	Diabetes	Pregnancy	Older age at menarche	Reproductive duration	Oral contraceptives	Parity	Atopy	Corticosteroids
Incident OAG	3	2									1					1	2	2													1 ^						1			1	1	1	1		1
Incident EG/EGS	1	1																1			1																								
Referral (to eye care or GP)	1	1		1	1																	2													1			1							
Hyperopi c change in refractive error	1				1	1								1			1										1			1		1													
Myopic change in refractive error	1	1			1	1				1 *				1			1													1		1													
Change in ocular status	1	1																				1																							
Myopic maculop athy progressi on	1	1								1							1																												

Incident visual field damage	1 Aging	Sex (male)	L Ethnicity (African/Black)	Geographical Remoteness	Years of education	Occupation	Lower household net worth	Worse visual acuity	Worse visual field mean deviation	SER.	High myopia	AMD	Glaucoma	Cataract	Cataract surgery	Pseudoexfoliation	Higher IOP	Positive family history of glaucoma	Parental myopia	Central corneal thickness	Eye colour	Increasing time between eye examinations	Higher meat intake	Higher fish intake	Overall diet quality	Higher alcohol intake	Smoking	Increasing time outdoors	Increasing time reading	Amount of physical activity	d8H	High cholesterol	Peripheral arterial disease	Hypercoagulable state	Previous stroke	Previous myocardial infarction	Ischaemic heart disease	Diabetes	Pregnancy	Older age at menarche	Reproductive duration	Oral contraceptives	Parity	Atopy	Corticosteroids
Incident myopia		1																	1									1	1	1									1 ‡						
Incident CRVO			1				1					1	1	1																	1	1	1	1	1	1		1							
Incident acute corneal hydrops								1																																				1	
AMD progressi on															1								1	2		1																			
Incident dual sensory impairme nt				o my		0.5														f					1																				

\* More myopic SER. ^ Treated hypertension. <sup>‡</sup> Incident myopia or progression of existing myopia. **Key:** Red = risk factor/increased risk; green = protective factor/lower risk; orange = mixed results, 1 study risk factor, 1 study no association; light green = mixed results, 1 study protective factor, 1 study no association; yellow = no association; grey = no studies investigated this prognostic factor/outcome pairing. **Abbreviations:** AMD: age-related macular degeneration; CRVO: central retinal vein occlusion; EG/EGS: exfoliation glaucoma/exfoliation glaucoma suspect; HBP: high blood

pressure; IOP: intraocular pressure; SER: spherical equivalent refraction.

# 3. DISCUSSION

### 3.1 Summary of the findings

The evidence included in this rapid review suggests that increasing age, sex, Black/African ethnicity, increasing IOP, positive family history of glaucoma, increasing length of time between eye examinations, hypertension, and heart disease are potential prognostic factors for a change in ocular health or vision. These were prognostic factors that were investigated in multiple studies; however, certainty in the evidence is low due to the majority of outcomes only being evidenced by one study. Similarly, the majority of studies were undertaken in specific populations, meaning that the association between these prognostic factors and the individual outcomes remains unclear in the general population. Single studies suggest that lower household net worth, worse VA, worse visual field mean deviation, SER, high myopia, AMD, glaucoma, cataract, diet, increasing alcohol intake, smoking, time spent outdoors, time spent reading, cholesterol, diabetes, peripheral arterial disease, hypercoagulable state, stroke, pregnancy, age at menarche, oral contraceptive use, and atopy are potential prognostic factors for a change in ocular status. This is summarised in Table 6.

Limited confidence in the results of this rapid review mean that these prognostic effects have limited applicability to the general population, owing to the specificity of the studies. Caution should therefore be taken when drawing from this rapid review, and further research is necessary to inform policy and practice.

It should also be noted that these findings are not specific to risk factors, with some studies also identifying protective factors. Each prognostic factor should be considered in relation to specific outcomes, rather than in relation to the overall category of a change in ocular status.

### 3.2 Strengths and limitations of the available evidence

One of the strengths of the available evidence is that all results were derived from multivariate analyses, presenting adjusting odds / risk / hazard ratios. This demonstrates that the prognostic factor of interest has an independent effect on the outcome. However, this is limited to only being independent of the other covariates included in the multivariate model, and these are not exhaustive. There is notable variation between studies in how many covariates they included in models, with some studies including as few as three covariates and others more than a dozen. Some studies did not clearly state what their adjustment factors were, or how they had decided which factors to include.

Limitations to the available evidence include only identifying two relevant studies that were carried out in the UK. Though search limits and eligibility criteria were applied to ensure only evidence from countries sufficiently similar to the UK were included in the review, the generalisability of any of these studies to Wales remains uncertain. Sample sizes of the studies also varied considerably with some being quite small, only several hundred participants. The sample populations were also often quite specific, such as post-menopausal women, female graduates, or glaucoma suspects, meaning that it is difficult or not possible to apply these findings to the broader population. There was comparatively less evidence identified that examined prognostic factors for conditions that would not require onward referral and could be managed by an optometrist, potentially influencing their decision on frequency of eye examinations, than conditions that would be referred to be managed by other health professionals.

There was a lack of relevant systematic reviews and meta-analyses identified for this review, which would have helped increase the certainty of the evidence by collating more studies. There were also no RCTs identified. However, the types of primary evidence included in the

review were of appropriate design for a prognostic factor review. Observational studies, such as cohort studies and case-control studies, can be useful to these types of review and variation in study designs is to be expected (Riley et al. 2019). The included studies were of reasonable quality, with risk of bias ranging from low to moderate as judged using the QUIPS and ROBIS tools. Only one primary study was rated as high risk of bias. However, the data in a number of studies were subject to bias as it was self-reported from participants or their carers. Questionnaires were used in several studies to collect both prognostic and outcome data, and this may lead to high risk of recall bias.

Finally, there was a distinct lack of evidence identified in children and younger adults. Only three studies included children in their study population, and only one of these reported results specifically for children. Similarly, only a small number of studies included adults aged 18-39 years and no studies reported specifically on younger adults. Further research is needed in these populations.

### Strengths and limitations of this Rapid Review 3.3

This rapid review was strictly limited to the included studies that were deemed to align with the research question and protocol, the scope of which was broad owing to the exploratory nature of this review. Controls intended to manage the amount of retrieved evidence were used, such as strict exclusion criteria and date limits and as such the methods used in this rapid review have been robust and pragmatic. However, it is crucial to note that due to the nature of rapid review methodology there remains the possibility that additional relevant studies were not identified. Additionally, some identified studies that reported findings relevant to the review question were excluded due to not reporting multivariate results or because the findings were not presented as odds/hazard/risk ratios. Sometimes these results can be converted to ratios, however, due to the nature of rapid review, it was not feasible to conduct this and include these studies. As such, the review team were reliant on interpreting the results of studies that have differing levels of quality and their own limitations. This therefore impacts the confidence of this review's conclusions.

### 3.4 Implications for policy and practice

The low certainty of the evidence in this review means caution should be taken should this review be used for decision making on appropriate eye examination intervals. Additionally, there is very little data from the UK and, thus, the generalisability of the findings to the Welsh population is uncertain.

This review should be used to identify what are thought to be the key prognostic factors and patient characteristics that could be used when an optometrist is determining an individual patient's risk of a change in ocular status, and therefore the appropriate interval until their next eye examination and suggesting these for further targeted research and evidence synthesis. The chosen factors or characteristics should be specific and narrow in scope, so that the limitations discussed above are mitigated. Alternatively, further research could be conducted looking at prognostic factors for specific ocular outcomes instead. The implications for future research are discussed in more detail below.

### 3.5 Implications for future research

Any further research undertaken to inform guidance on appropriate eye examination reassessment intervals should be much narrower in focus to ensure as much relevant and useful evidence as possible is gathered. Prognostic factors or specific ocular conditions of interest potentially need to be investigated individually for their effect on a change in ocular status.

This rapid review focused on incident conditions or progression of existing conditions. Prevalence data and prognostic factors for prevalent eye conditions or vision problems may also be useful for decision makers producing guidance on eye examination intervals and further evidence review could be performed in this area.

It has been noted in previous evidence-based guidelines on eye examination frequency that there is a lack of data in younger adults aged under 40 years (Robinson et al. 2012). This was also found to be the case during this review. Therefore, further research is required in this demographic. This is a demographic that is assumed to be at lower risk of ocular issues, but published evidence is lacking to support this claim and research should be conducted to determine if this is the case.

This review has identified a significant lack of evidence that would be needed to make confident conclusions to the research question and represents the findings of evidence that may not be generalisable to Wales, limiting the validity of this review's conclusions. More high-quality research must be undertaken in the populations of interest in order to inform and guide policy.

### 3.6 Economic considerations\*

- Sight loss costs the UK economy £25 billion per annum (RNIB 2021).
- Over 2 million people in the UK are currently living with sight loss (NHS 2021).
- The economic implications of appropriate or inappropriate testing intervals for different causes of vision loss will be different.
- A new case of age-related macular degeneration (AMD) in an adult aged 50 or over, costs the UK economy £73,350 over the person's lifetime. Lifetime costs to the UK economy for a person diagnosed with glaucoma are approximately £49.800 per person. Reducing the prevalence of these conditions by just 14 or 20 cases respectively could save the UK economy £1 million in lifetime costs (Fight for Sight 2020).
- On economic grounds, early detection of AMD in eye care services and the eye care pathway may be of benefit due to the level of prevalence and associated long term costs to the NHS as the condition causes irreversible, life limiting damage (Stahl 2020, Pezzullo et al. 2018).
- Draft National Institute for Health and Care Excellence (NICE) Guidance for Diabetic Retinopathy (DR) examinations suggest the use of ultrawide-field imaging for diagnosing and monitoring progression. The new guidance publishes in early 2024. DR costs the UK economy £80 million per annum when adjusted to October 2023 prices\*\* (Hex et al. 2012).
- The earlier detection of eve conditions through regular screening can identify conditions before severely impactful symptoms manifest. When captured at a population wide scale, this can result in significant economic savings (Fight for Sight 2020).

\*This section has been completed by the Centre for Health Economics & Medicines Evaluation (CHEME), Bangor University.

\*\* Prices adjusted using Bank of England Inflation Calculator.

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# 5. RAPID REVIEW METHODS

#### 5.1 Eligibility criteria

Table 7: Eligibility criteria

	Inclusion criteria	Exclusion criteria
Population	Asymptomatic adults and children attending for routine eye examinations, including those without refractive error or pre-existing ocular pathology, as well as those with pre-existing, managed ocular conditions.	People attending for eye examinations due to new symptoms.
Index prognostic factor	Prognostic factors that are available to optometrists during a routine eye examination will be examined in this review, with appropriate sub-group analyses performed. Specific prognostic factors of interest will include, but not be restricted to, age, sex, ethnicity, systemic health conditions (such as diabetes, hypertension), family history of eye disease, health behaviours (such as smoking, display screen use). Other prognostic factors/sub-groups that are identified during the evidence sift and data extraction stages will also be analysed.	
Comparison	Not applicable.	
Outcomes	<ul> <li>Primary outcomes:</li> <li>A change in ocular status: <ul> <li>change in refractive error</li> <li>change in visual acuity</li> <li>emergence of new ocular pathology, e.g. glaucoma, cataracts, macular degeneration</li> <li>ocular signs of new systemic pathology, e.g. Diabetes, hypertension</li> <li>change in existing ocular pathology</li> <li>ocular signs suggesting a change in existing systemic pathology</li> <li>new referral to general practitioner or secondary care</li> </ul> </li> <li>Secondary outcomes: <ul> <li>Prevalence of ocular pathology or refractive errors</li> <li>Rates of disease/condition progression</li> </ul> </li> </ul>	

Timing	Prognostic factor and outcomes measured at baseline and outcomes measured at any follow-up period up to 5 years. This time horizon was chosen as it was considered an appropriate length of time beyond the current most used interval between eye examinations in Wales of two years.				
Setting	To be used during primary care eye examinations to determine an individual patient's risk of experiencing a change in ocular status (as detailed in 'Outcome measures') with this risk being used to create dynamic re-examination intervals.				
Study design	Evidence-based clinical guidelines, systematic and rapid reviews, controlled trials, cohort analyses or population- based studies.				
Countries	We will prioritise studies from the UK and will not look at evidence from other countries where there is thought to be sufficient evidence from the UK. Where more evidence is required, studies from other countries, where optometry services are similarly comparable to Wales, will be prioritised in the following order: Ireland, Australia, New Zealand, Norway, Canada, Sweden, USA, Malta, Austria, Finland, Germany, Spain, The Netherlands, Switzerland. <sup>a</sup>				
Language of publication	English.				
Publication date	January 2009 to present.				
Publication type	Published and preprint.				
Other factors	We will include evidence on both prevalence and progression rates of ocular conditions where this is an outcome reported in relevant studies. We will report relevant recommendations for the frequency of eye examinations made by any evidence-based guidelines identified.				

<sup>a</sup>Prioritisation is based on similarity to UK optometry services and inclusion of countries in clinical guidelines by Robinson et al. (2012), and on data from the European Council of Optometry and Optics Blue Book (2020)

**Definitions: Refractive error** – A common eye disorder when the eye does not clearly focus images, which can usually be corrected by spectacles or contact lenses. The most common types of refractive error are myopia (shortsightedness), hypermetropia/hyperopia (longsightedness), astigmatism and presbyopia (reduced ability to focus on near objects); Visual acuity – A person's ability to recognise small details with precision, also referred to as clarity of vision.

#### 5.2 Literature search

Prior to planning this review, a preliminary search for existing reviews was undertaken of Cochrane Database of Systematic Reviews, NIHR Journals Library, Trip database, KSR Evidence, Canadian Agency for Drugs and Technologies in Health, Prospero, PubMed, NICE, SIGN, Epistemonikos, Google Advanced Search, and Google Scholar using the keywords sight test, eye examination, eye test, sight examination, routine, frequency, interval, recall, and time. The findings were presented to the stakeholders and used to refine the scope of the present rapid review, and to inform the methods.

A comprehensive search was conducted to identify any additional English-language reviews from 2009 onwards. An analysis of the text words contained in the title and abstracts, and of the index terms used to describe any relevant reviews already identified were used to inform the search. The full search strategy was designed and run using Ovid Medline and then translated to all other databases:

- CINAHL via the EBSCO platform •
- Embase via the Ovid platform
- Cochrane Library database
- Epistemonikos •

This was followed by a thorough search for relevant English-language primary studies from 2009 onwards on the following databases:

- CINAHL via the EBSCO platform •
- Medline and Embase via the Ovid platform •

The full searches for English-language reviews and primary studies can be found in Appendix 1.

Grey literature sources, including websites of key third sector and government organisations, identified by the review team, or provided by Stakeholders were also searched (see Appendix 2).

#### 5.3 Reference management

All citations retrieved from the database searches were imported or entered manually into EndNote<sup>™</sup> (Thomson Reuters, CA, USA) and duplicates removed by a single reviewer. The citations that remained were exported as a TXT file and imported to Rayyan<sup>™</sup> for study selection. Grey literature search results were added to an Excel spreadsheet and crosschecked against the database search results.

#### 5.4 Study selection process

Two reviewers screened 20% of titles and abstracts independently. If 20% is equal to less than 200 total records, then the two reviewers will screen 200 records. After this, the level of agreement was assessed with disagreements settled by discussion and consensus. Both reviewers had to achieve at least 80% agreement on screened records before progressing to the next stage. The remaining titles and abstracts were screened by the primary reviewer alone. 20% of full texts were screened by both reviewers, with the same agreement threshold (80%) as before necessary before the remaining records could be screened by the primary reviewer alone. During independent screening, the primary reviewer consulted with the secondary reviewer in the case of any uncertainties.

#### 5.5 Data extraction

Data extraction was based on the outlined eligibility criteria. We extracted details/characteristics on study country, study design, number of participants, relevant

outcomes (see eligibility criteria) and study settings. The <u>Checklist for Critical Appraisal and</u> <u>Data Extraction for Systematic Reviews of Prediction Modelling Studies (CHARMS-PF)</u> (Riley et al. 2019) was used to guide data extraction.

Data extraction was completed by individual reviewers and checked by a second reviewer (see Section 6.2 for completed data extraction forms for all included studies). In line with other prognostic factor reviews, data were only extracted from studies that reported prognostic factors as hazard ratios, odds ratios, or risk ratios/relative risk. Only multivariate or adjusted ratios were extracted so that only factors that were independently associated with outcomes were included in the report. Studies were excluded if they only reported unadjusted/univariate results. Relevant prevalence or condition progression rates were also extracted from studies that had reported the ratios listed above.

#### 5.6 Quality appraisal

Study quality was assessed using the Risk of Bias in Systematic Reviews (ROBIS) tool (Whiting et al. 2016) for included systematic reviews and using the Quality in Prognostic Factor Studies (QUIPS) tool for included primary studies (Hayden et al. 2013). Critical appraisal was completed by individual reviewers and checked by a second reviewer. Studies of all quality were included.

#### 5.7 Synthesis

We undertook narrative synthesis of the evidence identified based on the selection criteria outlined above.

#### 5.8 Assessment of body of evidence

All evidence selected after the sift stage was deemed fit for inclusion in the final review. Due to the scope of this review and the methodological constraints of rapid review, formal assessment of the body of evidence using GRADE was not feasible in this case. An informal assessment of the evidence has been conducted.

# 6. EVIDENCE

#### Search results and study selection 6.1

**Secondary Studies** 







## 6.2 Data extraction

All of the studies included in the rapid review are listed here. This includes two systematic reviews (Table 8) and 17 primary studies (Table 9).

Citation	Review details	Included studies	Quality	Key findings	Interpretation and observations
Dinu et al.	Review period: 1966 to January	26 studies:	Methodological	Animal products, RR (95% CI):	Increasing dietary meat
(2019)	2018	Meat: 6 studies, n = 101,011	quality of the	Meat Total AMD RR 1.11 (0.96-	intake is associated with
		Dairy products: 3 studies, n = 73,772	included studies	1.27, p = 0.16), Early AMD RR 1.17	increased risk of early
	Review purpose: To evaluate the	Fish: 8 studies, n = 237,464	was appraised	(1.02-1.34, p = 0.03), Late AMD RR	AMD.
	consumption of different food	Vegetables: 4 studies, n = 133,904	using the	0.99 (0.70-1.39, p NR)	Increasing alcohol intake
	groups and alcohol in relation to	Fruits: 3 studies, n = 132,525	Newcastle-	Dairy products Total AMD RR 1.07	is associated with
	occurrence and progression of AMD	Nuts: 3 studies, n = 4711	Ottawa Scale.	(0.68-1.70, p = 0.77), Early AMD	increased risk of all AMD
		Grains: 2 studies, n = 4335		RR 1.18 (0.93-1.50, p NR), Late	and early AMD, but not
	Study designs: Prospective cohort	Oils: 2 studies, n = 77,078	All but 3 studies	AMD RR 0.97 (0.27-3.48, p NR)	late AMD.
	studies	Butter: 2 studies, n = 7862	were ranked as	Fish Total AMD RR 0.82 (0.75-0.90,	Increasing dietary fish
		Margarine: 3 studies, n = 79,336	high quality.	p < 0.001), Early AMD RR 0.84	intake is associated with
	Outcome measures: Occurrence of	Alcohol: 12 studies, $n = 120,440$		(0.73-0.97, p = 0.02), Late AMD RR	decreased risk of all
	AMD		ROBIS RoB	0.79 (0.70-0.90, p < 0.001)	AMD, early AMD, and
		Country: 10 USA, 10 Australia, 2 The	assessment:		late AMD.
		Netherlands, 1 Denmark, 1 Iceland, 1	1. Study eligibility	Alcohol:	
		Japan, 1 South Korea	criteria: Low risk	Total AMD RR 1.20 (1.04-1.39, p =	Studies from countries
			of bias. Sufficient	0.01), Early AMD RR 1.29 (1.16-	included in the rapid
			information on	1.43, p < 0.001), Late AMD RR 0.98	review protocol
			eligibility criteria &	(0.76-1.27, p NR)	contributed more than
			their justification		85% of the weighting to
			2. Identification	Plant products:	all included meta-
			and selection of	Vegetables Total AMD RR 0.92	analyses.
			studies: Low risk	(0.82-1.03, p = 0.33), Early AMD	
			of bias. Search	RR 0.92 (0.67-1.25, p NR), Late	
			strategy appears	AMD RR 0.80 (0.76-1.00, p NR) Fruits Total AMD RR 0.91 (0.82-	
			appropriate. Dual sifting used	1.01, p = 0.08), Early AMD RR 0.92	
			3. Data collection	(0.82-1.03, p NR), Late AMD RR	
			and study	0.83 (0.62-1.12, p NR)	
			appraisal: Low	Nuts Total AMD RR 0.81 (0.64-1.02,	
			risk of bias. Dual	p = 0.08), Early AMD RR 0.73	
			independent data	(0.51-1.04, p NR), Late AMD RR	
			extraction.	0.83 (0.62-1.10, p NR)	
			Sufficient data	Grains Total AMD RR 0.84 (0.62-	
			extracted. RoB	1.13, p = 0.25)	
			analysed		

### Table 8: Summary of included systematic reviews

Citation	Review details	Included studies	Quality	Key findings	Interpretation and observations
			<ul> <li>4. Synthesis and findings: Unclear risk of bias.</li> <li>Serious heterogeneity (I2 &gt; 50%) not addressed in the synthesis. RoB not addressed for each pooled result. Unclear whether food groups or early vs late AMD subgroups were pre-specified in protocol.</li> <li>5. Risk of bias in the review: Unclear risk of bias.</li> <li>Heterogeneity not addressed in review. Unclear whether subgroup analyses were prespecified</li> </ul>	Fats:           Oils Total AMD RR 1.10 (0.98-1.23, p = 0.12), Early AMD RR 1.13 (0.93-1.37, p NR), Late AMD RR 1.05 (0.53-2.07, p NR)           Butter Total AMD RR 1.04 (0.93-1.16, p = 0.49), Early AMD RR 0.99 (0.75-1.30, p NR), Late AMD RR 0.85 (0.49-1.47, p NR)           Margarine Total AMD RR 1.05 (0.91-1.21, p = 0.54), Early AMD RR 1.05 (0.91-1.21, p = 0.54), Early AMD RR 1.07 (0.85-1.35, p NR), Late AMD RR 0.98 (0.56-1.70, p NR)	
Kessel et al. (2015)	<ul> <li>Review period: 1996 to August 2014</li> <li>Review purpose: To examine whether cataract surgery increases the risk of progression of dry AMD</li> <li>Study designs: RCTs, case-control studies</li> <li>Outcome measures: Best corrected distance visual acuity, funduscopic signs of AMD progression at least three months after surgery</li> </ul>	4 studies: 2 RCTs, 2 case-control studies. n = 1679. Country: UK, Australia, Germany, Austria	Quality was assessed using GRADE. The 2 RCTs were downgraded to 'moderate' quality due to imprecision and the 2 case- control studies were rated as 'very low' due to risk of bias.	Progression of non-exudative AMD to exudative AMD after cataract surgery (follow-up 6 to 12 months), RR (95% CI): RR 1.33 (0.60-2.94) [Total], RR 3.21 (0.14-75.68) [RCTs], RR 1.25 (0.55- 2.85) [case-control]	Cataract surgery is not associated with increased risk of progression to exudative AMD 6-12 months after surgery. The event rate was low (around 2%), however, and there is uncertainty about the effect estimate. Uncertain whether RRs have been adjusted for confounders.

Abbreviations: AMD: age-related macular degeneration; CI: confidence interval; NR: not reported; RCT: randomised controlled trial; RoB: risk of bias; RR: risk ratio/relative risk

#### Table 9: Summary of included primary studies

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
Barsam et al. (2017)	Study Design: Case- control study	Sample size: 64 cases, 1794 controls	Outcome: Acute corneal hydrops Method of	Prognostic factor: Vernal keratoconjunctivitis	Adjusted prognostic effect (95% confidence interval): Vernal conjunctivitis OR 15.00	History of vernal conjunctivitis, asthma, and having worse visual
(UK)		Participants: Cases:	measurement:	Method of measurement:	(1.30 to 173.70, p = 0.026)	acuity were associated with higher odds of

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
	Eligibility criteria/recruitment methods: The British Ophthalmological Surveillance Unit was used to identify new cases of acute corneal hydrops that occurred between November 2009 and December 2010. Clinicians who reported a case were sent an initial questionnaire that requested information on patient demographics, the best-corrected visual acuity before the onset of hydrops, previous keratometry values, if available, and prior ophthalmic and medical history. Patients with a completed questionnaire were defined as cases and included for further analysis. Controls with keratoconus who did not have a prior history of an acute corneal hydrops were identified from the public care hospital system from nine ophthalmic centres in the UK selected by a clustered, stratified random sampling procedure. The UK was divided into nine regions and selected hospitals within each region using	Mean (±SD) age 33.3 ± 12.9 years 75% male Ethnicity: 65.1% White, 22.2% South Asian, 11.1% Black, 1.6% other Controls: Mean (±SD) age 36.4 ± 12.1 years 66.1% male Ethnicity: 74.4% White, 17.1% South Asian, 4.3% Black, 4.3% other Dates of data collection: November 2009 to December 2010	Acute corneal hydrops was defined as the acute onset of bullous corneal oedema with an identifiable break in the Descemet's layer in the presence of keratoconus.	Clinician- completed questionnaire, based on clinical assessment. Prognostic factor: Asthma Method of measurement: Clinician- completed questionnaire, based on self- reported symptoms. Prognostic factor: VA in worse eye Method of measurement: Clinician- completed questionnaire.	Asthma OR 4.92 (1.22 to 19.78, p = 0.025) VA in worse eye OR 4.11 (1.18 to 14.32, p = 0.026) <b>Modelling method of</b> <b>analysis:</b> Backward stepwise multiple variable logistic regression <b>Adjustment factors used:</b> Not stated	developing acute corneal hydrops in people with keratoconus. The multivariable model only included 15 cases and 144 controls.

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
	computer-generated					
	random numbers with the					
	probability of selection					
	proportional to the					
	number of ophthalmic					
	consultants who worked					
	in each hospital. A local					
	investigator at each					
	centre then retrieved the					
	case notes of 20					
	consecutive patients with					
	keratoconus who had not					
	had an acute corneal					
	hydrops in the order they					
	attended clinic. The same					
	demographic and clinical					
	data were collected for					
	both cases and controls.					
	Quality rating: QUIPS					
	RoB assessment					
	1. Study participation:					
	High risk of bias. Case					
	control study design.					
	Prognostic data was					
	collected after the					
	outcome was known.					
	2. Study attrition: High					
	risk of bias. 88% of					
	eligible cases returned					
	questionnaire. 21% of					
	cases included in					
	analysis					
	3. Prognostic factor					
	measurement: Low risk of					
	bias.					
	4. Outcome					
	measurement: Low risk of					
	bias.					
	5. Adjustment for other					
	prognostic factors: Low					

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
	risk of bias. Relevant factors identified from the literature 6. Statistical analysis and reporting: Moderate risk of bias. Unclear whether strategy for model building is appropriate and is based on a conceptual framework or model	Sample size: 679	Outcome: Incident open-	Prognostic	Adjusted prognostic effect	Increasing age and
Ekström (2012) (Sweden)	<b>Study Design:</b> Population-based, longitudinal study	participants:	angle glaucoma (OAG) Method of measurement: Patients	factor: Age Method of measurement:	Adjusted prognostic effect (95% confidence interval): Age (per year) HR 1.15 (1.05 to 1.26)	increasing IOP are associated with increased risk of incident OAG in
	Eligibility criteria/recruitment methods: In 1984–1986, a population survey was conducted in the municipality of Tierp, south central Sweden. Its target population comprised 2429 residents 65–74 years of age. The size of the sample was limited to about one-third of the target population. Participants with normal and reliable visual fields, who completed the population survey, were invited to the follow-up study. To increase the cohort, 14 patients diagnosed with ocular hypertension at the Eye Department in Tierp in 1984–1986 were included. A further	Participants: 61.9% age 65-69 years 38.1% age 70-74 years 59.6% female Mean (± SD) follow-up time 9.0 ± 4.3 years Dates of data collection: 1984 to 2006	measurement: Patients diagnosed with OAG via supra-threshold visual field testing underwent manual Goldmann perimetry and repeated visual field testing using Competer threshold test logic. Threshold fields were sent for grading by an ophthalmologist otherwise unconnected with the study. Patients deemed to have progressive disease were classed as definite OAG cases. The glaucoma case records of patients with non-progressive disease, or missing threshold fields, were reviewed by an ophthalmologist, including optic disc characteristics. Patients were then classified as either definite OAG or	measurement:Self-reportedPrognosticfactor:IOPMethod ofmeasurement:GoldmannapplanationtonometryPrognosticfactor:PseudoexfoliationMethod ofmeasurement:Not stated	Mean IOP $\geq$ 25 mmHg and pseudoexfoliation HR 2.38 (1.87 to 3.03) Time-dependent (per 10 years): Mean IOP $\geq$ 25 mmHg HR 15.4 (4.52 to 52.1) Mean IOP 20-24.99 mmHg HR 3.92 (2.13 to 7.22) Mean IOP < 20 mmHg (ref) <b>Modelling method of</b> <b>analysis:</b> Cox proportional hazard models <b>Adjustment factors used:</b> Not clearly stated	Adults aged 65-74 years. Mean IOP ≥ 25 mmHg concurrent with pseudoexfoliation is associated with an increased risk of incident OAG in adults aged 65- 74 years. There is a risk of some double reporting between this study and Ekström & Hårleman (2023).

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
	259 people, participating		not.			
	in a case-control study in					
	1988–1995, were also					
	recruited. Those enrolled					
	were in the age range of					
	65–74 years and					
	underwent the same					
	baseline examination as					
	those in the population					
	survey. Exclusion criteria:					
	previous treatment for					
	OAG, previous cataract					
	surgery					
	Quality rating: QUIPS					
	RoB assessment					
	1. Study participation:					
	Low risk of bias.					
	Prospective cohort.					
	2. Study attrition: Low risk					
	of bias.					
	3. Prognostic factor					
	measurement: Moderate risk of bias. Unclear					
	whether prognostic factor					
	measures were updated					
	during follow-up.					
	4. Outcome					
	measurement: Low risk of					
	bias.					
	5. Adjustment for other					
	prognostic factors:					
	Moderate risk of bias.					
	Unclear how factors were					
	chosen. It appears family					
	history of glaucoma was					
	not included in the					
	multivariable model.					

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
	6. Statistical analysis and reporting: Low risk of bias.					
Ekström & Hårleman (2023) (Sweden)	Study Design: Nested case-control study Eligibility criteria/recruitment methods: Eligibility criteria for entry into the study included being a resident in one of the two rural districts of Tierp or Älvkarleby in the north of Uppsala County, south central Sweden, and being 55–84 years of age at the first consultation for eye-related problems at the Tierp Health Centre during the two recruitment periods, January 1988 to December 1995, or June 2003 to December 2003. In addition, the participants had to fulfil the IOP criteria for the study. The vast majority were referred from opticians or general practitioners. People who had seen an eye care provider in the last 3 years, were using pressure-reducing therapy, or had a history of intraocular surgery were not eligible.	Sample size: 481 participants Participants: Cases: n = 99 12 cases age 55-64 years, 38 cases age 65- 74 years, 49 cases age 75-84 years 50 cases male Controls: n =382 80 controls age 55-64 years, 183 controls age 65-74 years, 119 controls age 75-84 years 139 controls male Dates of data collection: 1988 to 2003	Outcome: Incident OAG Method of measurement: OAG was classified by a repeatable visual field defect in either eye, consistent with glaucoma and not explained by other causes. Participants with normal screening fields, who developed an abnormal test point within 2 years, were counted as incident OAG. Patients with a totally excavated optic disc and visual acuity < 0.05, unable to undergo automated perimetry, were also included in the OAG cases.	Prognostic factor: Age, sex, family history of glaucoma Method of measurement: Self-reported Prognostic factor: IOP Method of measurement: Goldmann applanation tonometry Prognostic factor: Hypertension, ischaemic heart disease Method of measurement: Self-reported, obtained from medical records (medical records (medical records prioritised if a discrepancy)	Adjusted prognostic effect (95% confidence interval): Age: 75-84 years OR 3.02 (1.13 to 8.08) 65-74 years OR 1.15 (0.44 to 3.00) 55-64 years (ref) Sex: Male OR 1.77 (0.91 to 3.43) Female (ref) Positive family history of OAG OR 3.21 (1.38 to 7.45) IOP (per 5 mmHg) OR 4.04 (2.91 to 5.62) Pseudoexfoliation OR 1.27 (0.63 to 2.57) Treated hypertension OR 0.58 (0.29 to 1.15) Ischaemic heart disease OR 2.41 (1.15 to 5.06) Modelling method of analysis: Multiple logistic regression analyses Adjustment factors used: Not stated	Increasing age, increasing IOP, positive family history of OAG, and ischaemic heart disease are associated with increased risk of incident OAG in adults aged 55-84 years. There is no evidence that sex and pseudoexfoliation are associated with increased risk of OAG in adults aged 55-84 years. The effect of pseudoexfoliation on glaucoma risk is mediated by elevated IOP. There is a risk of some double reporting between this study and Ekström (2012).

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
	With the intention of					
	reaching an equal					
	distribution of IOP					
	readings around 22					
	mmHg, the pressure limit					
	for inclusion in the study					
	was changed once or					
	twice every year, with a					
	lower level of 18 mmHg,					
	depending on the results					
	of those already included.					
	From January 1995 to					
	November 1995, people					
	with an IOP < 18 mmHg					
	were recruited, while all					
	pressures were accepted					
	in December 1995. From					
	June to December 2003,					
	the IOP had to be < 17 or					
	≥ 35 mmHg.					
	Quality rating: QUIPS					
	RoB assessment					
	1. Study participation:					
	Low risk of bias. Nested					
	case control study design					
	2. Study attrition: Low risk					
	of bias.					
	3. Prognostic factor					
	measurement: Moderate					
	risk of bias. Unclear					
	whether prognostic factor					
	measures were updated					
	during follow-up.					
	4. Outcome					
	measurement: Low risk of					
	bias.					
	5. Adjustment for other					
	prognostic factors: Low					
	risk of bias. Unclear how					
	factors were chosen.					

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
Elmore et al.	<ul> <li>6. Statistical analysis and reporting: Moderate risk of bias. Unclear whether strategy for model building is appropriate and is based on a conceptual framework or model</li> <li>Study Design:</li> </ul>	Sample size: 1076	Outcome: Incident AMD Method of	Prognostic	Adjusted prognostic effect	There is no association
(2022)	Prospective cohort analysis	participants Participants: No AMD,	measurement: Self-	factor: RBC fatty acid Method of	(95% confidence interval): RBC polyunsaturated fatty acid levels:	between fatty acid intake or fish intake and incident AMD in post-menopausal
(USA)	Eligibility criteria/recruitment methods: Post- menopausal women. Participants had to be enrolled in both the Women's Health Initiative Memory Study and the Women's Health Initiative Sight Exam Study (WHI-SE); two ancillary studies conducted in the Women's Health Initiative Hormone Therapy (WHI HT) trial. Exclusion criteria: missing or ungradable fundus photos, unreliable red blood cell fatty acid measures, self-reported energy intake above 5000 kcals or below 600 kcals, missing covariate data.	Participants: No AMD, n = 938 (follow-up): 59.0% < 70 years, 32.5% 70-74 years, 8.5% 75+ years Ethnicity 89.8% White, 6.5% Black, 1.8% Hispanic, 0.9% Asian or Pacific Islander, 0.1% American Indian or Alaskan Native, 1.0% other Incident AMD, $n = 138$ (follow-up): 48.6% < 70 years, 39.1% 70-74 years, 12.3% 75+ years Ethnicity 92.0% White, 4.3% Black, 0.0% Hispanic, 0.7% Asian or Pacific Islander, 0.0% American Indian or Alaskan Native, 2.9% other Dates of data collection: 2000 to 2015	Prevalent AMD status was determined from stereoscopic 30° colour fundus photographs taken as part of the WHI-SE Study (2000–2002) and graded using the Wisconsin Age-Related Maculopathy Grading Scheme. This identified 240 prevalent AMD cases and 1216 cases without AMD. Participants enrolled in the WHI Extension Study 1 (2005–2010) and Extension Study 2 (2010– 2015) received a mailed Medical History Update survey annually. This survey asked, "since the date on the front of this form, has a doctor told you for the first time that you have macular degeneration?"	measurement: Fasting blood samples were collected at WHI HT baseline (1993–1998). RBC fatty acid composition was analysed using gas chromatography with flame ionization detection and then expressed as percent weight of total fatty acids. Prognostic factor: Dietary intake of fatty acids and fish Method of measurement: A subset of the study sample were additionally enrolled in the WHI	Acid levels.         No significant association between any RBC polyunsaturated fatty acid levels and incident AMD         Dietary intake of fatty acids: No significant association between dietary intake of any fatty acids and incident AMD         Dietary intake of fish: ≥ 1 serving per week HR 0.91 (0.53 to 1.58)         ≥ 1 serving per month and < 1 serving per week HR 0.86 (0.48 to 1.54) None or < 1 serving per month (ref)         Dietary intake of dark fish: ≥ 1 serving per week HR 1.20 (0.79 to 1.81)         ≥ 1 serving per month and < 1 serving per week HR 0.60 (0.34 to 1.04) None or < 1 serving per month (ref)	RBC polyunsaturated fatty acid levels are a longer-term biomarker of fatty acid intake.

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
	<ol> <li>Study participation: Low risk of bias.</li> <li>Prospective cohort</li> <li>Study attrition:</li> <li>Moderate risk of bias.</li> <li>Only participants enrolled in the WHI extension study (1076/1216) provided incidence data.</li> <li>Prognostic factor measurement: Moderate risk of bias. Self-reported via questionnaire</li> <li>Outcome measurement: Moderate risk of bias. Self-reported AMD</li> <li>Adjustment for other prognostic factors: Low risk of bias.</li> <li>Statistical analysis and reporting: Low risk of bias</li> </ol>		Self-reported AMD was categorized as "yes" or "no." Participants not followed up into the extension studies were excluded. 1076 of the 1216 women without prevalent AMD at WHI-SE baseline were followed up, and 138 women were identified as developing incident AMD.	Dietary Modification Trial. Dietary variables were collected via a modified Block food frequency questionnaire assessing usual dietary intake during the previous three months.	Modelling method of analysis: Cox proportional hazards regression modelling Adjustment factors used: Age, race/ethnicity, pack- years of smoking, assignment to clinical trial (the hormone therapy trial, the dietary modification trial, or the calcium and vitamin D trial), hypertension, BMI, recreational physical activity, diabetes status	
Fernández- Montero et al. (2017) (Spain)	Study Design: Longitudinal, prospective cohort study Eligibility criteria/recruitment methods: The SUN project is a multipurpose, prospective, dynamic cohort of young adult university graduates conducted in Spain. The recruitment of participants started in 1999 and is permanently open. Mailed questionnaires are used to gather baseline	Sample size: 10,401 participants Participants: 3180 reported pregnancies Pregnancy mean (± SD) age 28.4 ± 4.2 years No pregnancy mean (± SD) age 35.3 ± 8.5 years Dates of data collection: 1999 to 2013	Outcome: Incident myopia or progression of myopia Method of measurement: All follow- up questionnaires included the following question: "Have you been diagnosed by a medical doctor of new-onset myopia or a progression of 0.5 or more dioptres in myopia, since the last questionnaire you filled in?" Participants who responded yes to this question were considered incident cases.	Prognostic factor: Pregnancy Method of measurement: Pregnancies were assessed in each biennial follow-up questionnaire. All questionnaires, except for the first 2-year follow-up, included the following question: "Have you been diagnosed by a doctor of a pregnancy since the last questionnaire? If	Adjusted prognostic effect (95% confidence interval): Pregnancy HR 0.58 (0.49 to 0.69, p < 0.001) Modelling method of analysis: Multivariable Cox regression analysis Adjustment factors used: Age, BMI, total energy intake, Mediterranean Diet, smoking habits, computer use, educational level, sleeping behaviour, time of television watching, physical activity	Pregnancy is associated with a decreased risk of developing myopia or progression of existing myopia in women aged 20-50 years.

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
	characteristics and information on diet, lifestyles and new medical diagnoses of disease every 2 years. This study only included women with a minimum of 2-years follow-up between the ages of 20 to 50 years. <b>Quality rating:</b> QUIPS RoB assessment 1. Study participation: Low risk of bias. Prospective cohort 2. Study attrition: Moderate risk of bias. 14% lost to follow-up 3. Prognostic factor measurement: Moderate risk of bias. Self-reported via questionnaire. Time outdoors was estimated & only at baseline. 4. Outcome measurement: Moderate risk of bias. Self-reported myopia 5. Adjustment for other prognostic factors: Low risk of bias. 6. Statistical analysis and reporting: Low risk of		Participants were considered a case of myopia progression when they reported a new diagnosis of myopia or an increase in myopia of at least -0.50 D in one eye.	so, please report estimated due data (month/year)".		
Gopinath et al. (2014) (Australia)	bias. Study Design: Longitudinal, population- based cohort study	Sample size: 2443 participants Participants: 1 <sup>st</sup> quintile of total diet score:	Outcome: 5-year incidence of dual sensory impairment Method of measurement:	Prognostic factor: Diet Method of measurement: At baseline, dietary data were	Adjusted prognostic effect (95% confidence interval): Total diet score 5 <sup>th</sup> quintile vs. 1 <sup>st</sup> quintile and dual sensory impairment OR 1.03 (0.30 to 3.50)	Diet is not associated with 5-year incidence of dual sensory impairment in adults aged over 49 years.

Citation	Study Details	Participants &	Outcomes	Prognostic	Key findings	Interpretation and
(Country)	-	setting		factors	itey indiage	observations
	Eligibility	Mean ( $\pm$ SD) age 67.3 $\pm$	Pure-tone audiometry	collected using a		
	criteria/recruitment	9.5 years, 53.9% male	was performed by	145-item self-	Modelling method of	
	methods: Part of the	5 <sup>th</sup> quintile of total diet	audiologists in sound-	administered food	analysis: Discrete linear	
	Blue Mountains Eye	score:	treated booths, using	frequency	logistic models	
	Study (BMES).	Mean ( $\pm$ SD) age 66.8 $\pm$	TDH-39 earphones and	questionnaire.		
	Following a door-to-door	8.4 years, 31.9% male	Madsen OB822	A total diet score	Adjustment factors used:	
	census of the region,		audiometers. Sound-proof	was established by	age, sex, education, current	
	baseline examinations of	Dates of data	rooms were set-up	allocating scores	smoking, noise exposure, type	
	3654 residents aged > 49	collection: 1992 to	according to International	for intakes of	2 diabetes	
	years were conducted	2004	Standards Organization	selected food		
	during 1992-4 (BMES-1).		protocol 8253-2. Bilateral	groups and		
	Surviving baseline		hearing impairment was	nutrients for each		
	participants were invited		determined as the pure-	participant as		
	to attend 5-year follow-up		tone average of	described in the		
	examinations (1997-9,		audiometric hearing	Dietary Guidelines		
	BMES-2), at which 2334		thresholds at 500,1000,	for Australian		
	(75.1% of survivors) and		2000, and 4000 Hz	Adults. The total		
	an additional 1174 newly		(PTA0.5 to 4 kHz) in the	diet score is		
	eligible residents were		better ear, defining any	divided into ten		
	examined. At BMES-2,		hearing loss as	components, and		
	2956 participants had		PTA0.5 to 4kHz > 25 dB	each component		
	audiometric testing		HL; mild hearing loss as	has a possible		
	performed. At BMES-3		PTA0.5 to 4kHz	score ranging from		
	(2002-4), 1952		> 25 to 40 dB HL; and	0 to 2. A maximum		
	participants were re-		moderate to severe	score of 2 was		
	examined.		hearing loss as PTA0.5 to	given to subjects		
	Visual acuity data were		4kHz > 40 dB HL.	who met the		
	collected at all three		Monocular distance	recommendations		
	BMES examinations.		logMAR VA was	with pro-rated		
			measured with forced	scores for lower		
	Quality rating: QUIPS		choice procedures using	intakes. These		
	RoB assessment		a retro-illuminated chart	were then		
	1. Study participation:		according to the early	summated		
	Moderate risk of bias.		treatment diabetic	providing a final		
	Prospective cohort.		retinopathy Study	score ranging		
	513/2956 (17%) were		protocol. Both presenting	between 0 and 20		
	ineligible due to missing		VA (with current	with higher scores		
	data at baseline.		eyeglasses, if worn) and	indicating closer		
	2. Study attrition:		after subjective refraction	adherence to the		
	Moderate risk of bias.		(best-corrected VA) were	dietary guidelines.		
	Unclear how many		measured.			

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
	participants completed the 5 year follow-up. 3. Prognostic factor measurement: Low risk of bias. Some factors based on self-report others measured. 4. Outcome measurement: Low risk of bias. 5. Adjustment for other prognostic factors: Moderate risk of bias. Unclear how factors were chosen. 6. Statistical analysis and reporting: Low risk of bias.		Any visual impairment was defined as presenting VA of the better eye < 20/40). Dual sensory impairment was defined as concurrent visual (either presenting or best- corrected) and hearing impairment, as determined using the above definitions.			
Guggenheim et al. (2012) (UK)	Study Design: Opportunistic, longitudinal studyEligibility criteria/recruitment methods: Avon Longitudinal Study of Parents and Children (ALSPAC) cohort: Pregnant women with an expected date of delivery between April 1, 1991 and December 31, 1992, resident in the former Avon health authority area in Southwest England, were eligible to participate in the study. A cohort of 14,541 pregnant women was established, resulting in 13,988	Sample size: 2005 participants with complete information on predictor variables and either were seen at the age 15-year clinic or who were already known to have become myopic when they attended the 12-year clinic. Participants: Mean (± SD) age 11.7 ± 0.2 years 49.1% male Dates of data collection: April 1991 to c.2008	Outcome: Incident myopia after age 11 years Method of measurement: Non- cycloplegic autorefraction. Participants were classified as myopic if the average of the SERs in their right and left eyes was ≤ -1.00 D. Subjects were classified as emmetropic or hyperopic if the averaged SER in their right and left eyes was ≥ -0.25 D.	Prognostic factor: Parental myopia Method of measurement: Participants' parents completed a questionnaire that included the question "How would you rate your sight without glasses?" and were classed as myopic if they answered "can't see clearly at distance" for both eyes. Prognostic factor: Time spent reading	Adjusted prognostic effect (95% confidence interval): Parental myopia: 1 myopic parent OR 1.175 (0.900 to 1.533, $p = 0.236$ ) 2 myopic parent OR 1.143 (0.718 to 1.818, $p = 0.574$ ) No myopic parents (ref) Time spent reading (low vs. high): OR 1.323 (1.023 to 1.712, $p = 0.033$ ) Time spent outdoors: OR 0.65 (0.45 to 0.96, $p = 0.029$ ) Sex (male vs. female): OR 1.058 (0.810 to 1.382, $p = 0.679$ )	Increased time spent reading is associated with increased risk of developing myopia after age 11. Increased time spent outdoors is associated with lower risk of developing myopia after age 11. Parental myopia, sex and physical activity/sedentary behaviour are not associated with incident myopia after age 11.

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
	children who were alive	3		Method of	Physical activity/sedentary	
	at 12 months of age.			measurement:	behaviour:	
	Data collection has been			When the	Mean counts per minute for	
	via various methods,			participants were	whole week:	
	including self-completion			aged 8 to 9 years,	OR 0.887 (0.773 to 1.017, p =	
	questionnaires sent to the			mothers completed	0.086)	
	mother and her partner,			a questionnaire		
	and after age 5 to the			including the	Time with moderate to	
	child, as well as direct			question "On	vigorous activity per day:	
	assessments and			normal days in	OR 0.877 (0.764 to 1.006, p =	
	interviews in a research			school holidays,	0.062)	
	clinic, biological samples,			how much time on		
	and linkage to school and			average does your	Time with sedentary counts:	
	hospital records.			child spend each	OR 1.095 (0.959 to 1.251, p =	
	All children still			day reading books	0.180)	
	participating in ALSPAC			for pleasure?"		
	were invited			Children were	Modelling method of	
	approximately yearly			classified as	analysis: Multivariate logistic	
	(starting at age 7 years)			spending a "high"	regression analyses	
	to sessions where a			amount of time	A diversion on the state was de	
	number of assessments and interviews, tailored to			reading for pleasure if the	Adjustment factors used: Parental myopia,	
	their age, took place.			response was "1–2	time reading, sex,	
	Vision-related data were			hours" or "3 or	physical activity/	
	included in the			more hours," and	sedentary behaviour, time	
	assessments carried out			as "low" otherwise.	spent outdoors.	
	at the 7-, 10-, 11-, 12-,				spent outdoors.	
	and 15-year clinics.			Prognostic	Three different measures for	
				factor: Time spent	physical activity/sedentary	
	Quality rating: QUIPS			outdoors	behaviour were used and	
	RoB assessment			Method of	modelled separately. The ORs	
	1. Study participation:			measurement: At	were similar regardless of	
	Low risk of bias.			8 to 9 years of age,	which measure was adjusted	
	Prospective cohort.			a questionnaire	for.	
	2. Study attrition:			was completed by		
	Moderate risk of bias.			participants'		
	Refractive error data was			mothers, asking		
	available for 9109/13988			"On a (weekend		
	children.			day)/(school week		
	3. Prognostic factor			day), how much		
	measurement: Moderate			time on average		

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
(ooundy)	risk of bias. Time	Setting		does your child		
	outdoors, time spent			spend each day		
	reading and parental			out of doors in		
	refractive error were all			(summer)/(winter)."		
	self-reported (by mother).			The response		
	4. Outcome			options were		
	measurement: Low risk of			"None at all," "1		
	bias.			hour," "1–2		
	5. Adjustment for other			hours," and "3 or		
	prognostic factors: Low			more hours."		
	risk of bias.			Children were		
	6. Statistical analysis and			classified as		
	reporting: Low risk of			spending a "high"		
	bias.			amount of time		
				outdoors in		
				summer if the		
				response was "3		
				or more hours,"		
				and as "low"		
				otherwise. For time		
				spent outdoors in		
				winter, children		
				were classified as		
				spending a "low"		
				amount of time		
				outdoors if the		
				response was		
				"None at all" or "1		
				hour," and as		
				"high" otherwise.		
				g. ettermeet		
				Prognostic		
				factor: Sex		
				Method of		
				measurement:		
				Self-reported		
				Prognostic		
				factor: Physical		
				activity/sedentary		
				behaviour		

Citation	Study Details	Participants &	Outcomes	Prognostic	Key findings	Interpretation and
(Country)	citaly zotalie	setting	• • • • • • • • • • • • • • • • • • • •	factors	ney mange	observations
				Method of		
				measurement:		
				Children attending		
				the research clinic		
				at age 11 years		
				were asked to		
				wear an Actigraph		
				accelerometer for		
				the following 7		
				days. Data from		
				the returned		
				accelerometers		
				were downloaded		
				and imported into a		
				database. Children		
				who did not		
				provide at least 10		
				hours of valid data		
				on at least 3		
				separate days		
				were omitted from		
				the analyses. Two		
				physical activity		
				variables were		
				derived from the		
				data: Mean counts		
				per min for the		
				whole week, and		
				minutes of		
				moderate to		
				vigorous activity		
				per day.		
				A variable		
				representing		
				sedentary		
				behaviour was		
				derived from count		
				per min by defining		
				sedentary time as		
				less than 200		
				counts per min.		

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
Hopf et al. (2022) (Germany)	<ul> <li>Study Design: Prospective, population- based cohort study</li> <li>Eligibility criteria/recruitment methods: Part of the Gutenberg Health Study. Random sampling of residents of the State of Rhine-Palatine by the regional registration office, stratified by gender, decade of age, residence, baseline age of 35 to 74 years. Inclusion criteria: Phakic eyes with SER ≤ -6.00 D at baseline examination, gradable fundus photographs at baseline and 5-year follow-up.</li> <li>Quality rating: QUIPS RoB assessment 1. Study participation: Low risk of bias. Prospective cohort 2. Study attrition: Low risk of bias. 3. Prognostic factor measurement: Low risk of bias. 4. Outcome measurement: Low risk of bias. 5. Adjustment for other prognostic factors: Low risk of bias. 6. Statistical analysis and</li> </ul>	Sample size: 350 participants (528 eyes) Participants (528 eyes) Participants (at baseline): Without baseline myopic maculopathy: Mean age 50.23 ± 9.17 years 50.8% female Median SER (RE) -7.19 D (IQR -8.62 to -6.25) Median SER (LE) -7.25 (-8.75 to -6.50) With baseline myopic maculopathy: Mean (± SD) age 56.70 ± 9.08 years 44.4% female Median SER (RE) -9.81 D (IQR -11.47 to -7.47) Median SER (LE) -8.75 (-11.25 to -7.25) Dates of data collection: 2007 to not stated	Outcome: Progression of myopic maculopathy at 5 years Method of measurement: Fundus photographs graded by two masked graders following the international photographic grading system for myopic maculopathy. A retinal specialist made the decision in a consensus meeting if the two graders disagreed. Progression was defined as an increase in stage of myopic maculopathy, enlargement(s) of existing lesion, or new lesion(s) at a different spot at the posterior pole.	Prognostic factor: Sex, age Method of measurement: Not explicitly stated Prognostic factor: IOP Method of measurement: Non-contact tonometry (Nidek NT-2000) Prognostic factor: SER Method of measurement: Non-cycloplegic autorefraction	Adjusted prognostic effect (95% confidence interval): Sex (female) OR 5.54 (0.93 to 32.92, p = 0.060) Age (per year) OR 0.94 (0.88 to 1.02, p = 0.134) IOP (per mmHg) OR 1.62 (1.51 to 1.59, p = 0.035) SER (per dioptre) OR 1.21 (0.99 to 1.49, p = 0.063) Modelling method of analysis: Multivariable logistic regression analyses Adjustment factors used: Sex, age, IOP, baseline SER	Increasing IOP is associated with increased risk of progression of myopic maculopathy at 5 years in adults aged 35- 74 years. Sex, age, and SER are not associated with increased risk of myopic maculopathy progression at 5 years in adults aged 35-74 years.

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
Irving et al.	reporting: Moderate risk of bias. Unclear whether strategy for model building is appropriate and is based on a conceptual framework or model Study Design:	Sample size: 2656 participants	Outcome: Significant change in ocular status	Prognostic factor: Age, sex,	Adjusted prognostic effect (95% confidence interval):	Increasing age and increasing length of time
(2016)	Retrospective cohort analysis		Method of	interval between	Åge (per year) OR 1.03 (1.03	between eye
(Canada)	analysis Eligibility criteria/recruitment methods: Retrospective cross-sectional database of patients who presented at the University of Waterloo Optometry Clinic during a 1-year period from January 2007 to January 2008. Data were extracted for all patients whose reason for presenting was to have a routine eye examination as reported in the case history (including those presenting for employment purposes, to obtain contact lenses, or to replace spectacles). There were some patients who initially presented for a routine eye examination but reported symptoms when specifically questioned. These patients were	Participants: Median age 38.5 years (range 0.4 to 93.9 years) 48% male Dates of data collection: January 2007 to January 2008	<b>measurement:</b> Defined as one or more of spectacle prescription change, new critical diagnosis, or new management of an existing condition. A spectacle prescription change was considered to be significant if in at least one eye, the sphere, cylinder, or any reading addition changed by > 0.50 D from the entering to the exiting spectacle prescription, or if the cylinder axis changed as follows: > 15 degrees if the absolute value of the final cylinder value was < 1.00 D, > 10 degrees if the cylinder was $\ge 1.00$ D but < $2.00$ D, or > 5 degrees if the cylinder was $\ge 2.00$ D. A critical diagnosis was considered new if it was not reported in the clinic file case history or at previous examinations.	eye examinations Method of measurement: Identified from case notes	to 1.04) Sex (female) OR 1.07 (0.90 to 1.29) Interval between eye examinations (per year) OR 1.06 (1.02 to 1.11) <b>Modelling method of</b> <b>analysis:</b> Multivariable logistic regression <b>Adjustment factors used:</b> Assessment interval, sex, age	examinations are associated with increased risk of experiencing a significant change in ocular status. Sex was not associated with increased risk of significant change in ocular status.

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
	excluded from the main analysis, but their overall percentage of significant change is reported for comparison. Quality rating: QUIPS RoB assessment 1. Study participation: Low risk of bias. Retrospective cross- sectional study 2. Study attrition: Low risk of bias. 3. Prognostic factor measurement: Low risk of bias. 4. Outcome measurement: Moderate risk of bias. Composite outcome of any critical ocular disorders or abnormal findings 5. Adjustment for other prognostic factors: Low risk of bias. 6. Statistical analysis and reporting: Low risk of bias.		A management (not including prescription change) was considered new if it was not initiated at a previous visit or if there was a change compared to the last available information. New managements included referrals, new treatment, or changes in monitoring schedule.			
Kang et al. (2012)	Study Design: Prospective cohort study	Sample size: 120,146 participants Participants:	Outcome: Incident exfoliation glaucoma or exfoliation glaucoma suspect	Prognostic factor: Age, gender, eye colour Method of	Adjusted prognostic effect (95% confidence interval): Rate ratio (RR) of age: 40 to 55 years (ref)	Increasing age, positive family history of glaucoma and female gender are associated
(USA)	Eligibility criteria/recruitment methods: The Nurses' Health Study is an ongoing population- based cohort of registered female nurses.	Women: 78,955 participants Ancestry: 7% Scandinavian, 16.4% Southern European, 74.3% other white, 1.4% black, 0.7% Asian, 0.2%	Method of measurement: In all biennial questionnaires from 1986, participants were asked if they had physician-diagnosed glaucoma. From among	measurement: Self-reported in questionnaires Prognostic factor: Family	55 to 60 years RR 4.33 (2.19 to 8.56) 60 to 65 years RR 10.43 (5.50 to 19.78) 65 to 70 years RR 19.88 (10.41 to 37.96)	with higher risk for incident EG or EGS. Eye colour is not associated with increased risk of incident EG or EGS for adult men.

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
	The Nurses' Health Study was established in 1976 when 121,700 United States women were invited to complete a questionnaire regarding lifestyle, health behaviour, and chronic diseases. The Health Professionals Follow-up Study (HPFS) is an ongoing cohort created in 1986 when 51,529 male health professionals (dentists, veterinarians, pharmacists, optometrists, osteopaths, and podiatrists) completed a similar health survey. The participants in both cohorts have been followed up biennially with mailed	setting Native American or Hawaiian, 0.8% Hispanic 288 incident cases of EG or EGS Mean (± SD) age at diagnosis 68.1 ± 6.6 years Men: 41,191 participants Ancestry: 11.2% Scandinavian, 23% Southern European, 61.3% other white, 0.9% black, 1.6% Asian, 2% Native American or Hawaiian 60 incident cases of EG or EGS Mean (± SD) age at diagnosis 70.8 ± 6.9 years	participants who gave a positive response to this question, permission was obtained to retrieve their medical information. The diagnosing eye care provider of record was sent a request to complete a glaucoma questionnaire, which asked about the presence of exfoliation material or other secondary causes for elevated IOP, maximum IOP, optic nerve features, and status of the filtration apparatus and was asked to send all available visual field (VF) reports. In lieu of completing the questionnaire, eye care providers could send the complete medical records		70 to 75 years RR 33.54 (17.23 to 65.29) Over 75 years RR 46.22 (22.77 to 93.80) Rate ratio (RR) of family history of glaucoma: Positive history RR 2.29 (1.39 to 3.78) Negative history (ref) Rate ratio (RR) of eye colour (males only): Hazel/green/medium RR 0.87 (0.43 to 1.74) Brown/dark RR 0.84 (0.42 to 1.68) Blue/light (ref) Rate ratio (RR) of gender: Male RR 0.32 (0.23 to 0.46) Female (ref) <b>Modelling method of</b>	
	with mailed questionnaires that have updated health and lifestyle information. The study period was 1980 through 2008 for the Nurses' Health Study and 1986 through 2008 for the HPFS. Data were collected from those who were prospectively followed for 20 years or more and who provided lifetime residence information as well as other lifestyle and health information were used to examine the	Dates of data collection: 1980 to 2008	complete medical records and all VF reports related to the glaucoma diagnosis. A glaucoma specialist (LRP) evaluated the questionnaire or medical record information as well as the VF data in a standardized manner for confirmation and classification. Cases of either EG or EGS were analysed. Specifically, EG was defined as the presence of exfoliation material in combination with 2 or		Modelling method of analysis: Cox proportional hazard analysis Adjustment factors used: Age, race, family history of glaucoma, BMI, self-reported hypertension, diabetes mellitus, high cholesterol, myocardial infarction, geographical tier	

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
	<ul> <li>descriptive epidemiologic features of exfoliation glaucoma (EG) or exfoliation glaucoma suspect (EGS).</li> <li>Participants contributed person-time until confirmed EG or EGS, self-report of glaucoma, death, loss to follow-up, diagnosis of cancer other than nonmelanoma skin cancer, self-report of cataract extraction, or the end of the study (2008).</li> <li>Quality rating: QUIPS RoB assessment 1. Study participation: Moderate risk of bias. Prospective cohort. Confirmation received from diagnosing eye-care provider of 6870/10737 of those self-reporting glaucoma.</li> <li>Study attrition: Low risk of bias.</li> <li>Prognostic factor measurement: Moderate risk of bias. Self reported via questionnaire</li> <li>Outcome measurement: Moderate risk of bias. Self-reported glaucoma - but verified by review of medical records.</li> <li>Adjustment for other prognostic factors: Low risk of bias.</li> </ul>		more reliable tests showing reproducible VF loss consistent with glaucoma, and EGS was defined as the presence of exfoliation material in combination with (1) a history of IOP of more than 21 mmHg; or (2) a cup-to-disc ratio of 0.6 or more or the inter-eye difference in a cup-to-disc ratio of 0.2 or more; or (3) only 1 reliable test showing VF loss consistent with glaucoma. Those with a presence of exfoliation material only without any VF loss or elevation in IOP or abnormal cup-to-disc ratios (as defined above) were not considered as cases of EG or EGS.			

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
	<ol> <li>Statistical analysis and reporting: Low risk of bias.</li> </ol>					
Keel et al. (2017) (Australia)	<ul> <li>Study Design: Population-based, cross-sectional study</li> <li>Eligibility criteria/recruitment methods: Thirty sites, across five Remoteness Areas (Major City, Inner Regional, Outer Regional, Remote and Very Remote), were selected using a multi- stage, random cluster sampling methodology. To obtain a nationally representative sample of the population, 100 non- Indigenous Australians aged 50 years and older and 50 Indigenous Australians aged 40 years and older were to be recruited at each site. Recruiters went door-to- door to determine the eligibility of the residents. All eligible residents were invited to participate.</li> <li>Quality rating: QUIPS RoB assessment 1. Study participation: Low risk of bias. Cross sectional study 2. Study attrition: Low risk</li> </ul>	Sample size: 3098 participants Participants: Referred (n = 994): Mean (± SD) age 67 ± 10 years 51.8% male Not referred (n = 2104): Mean (± SD) age 66.4 ± 9.5 years 43.8% male Dates of data collection: c.2015	Outcome: Rates of eye care referral Method of measurement: A referral protocol was developed by study investigators in conjunction with ophthalmologists. Participants were provided with a referral letter to be taken to their optometrist or local doctor if they met any of the following referral criteria: (1) evidence of eye disease or visual impairment detected during the NEHS eye examination; (2) participants with diabetes who had not undergone a screening eye examination within the timeframe recommended by the National Health and Medical Research Council diabetic retinopathy guidelines, or (3) individuals without diabetes who had undergone an eye examination in the past 5 years. Participants who were already under ophthalmological care were not provided with a	Prognostic factor: Sex, age, time since previous eye examination, geographical remoteness, years of education, diabetes, stroke Method of measurement: Each participant underwent an interviewer- administered questionnaire to collect information on socio- demographic factors, history of ocular problems, stroke and diabetes.	Adjusted prognostic effect (95% confidence interval): Age (per year) OR 1.02 (1.01 to 1.02, p < 0.001) Sex (male) OR 1.24 (1.06 to 1.46, p = 0.007) Years of education (per year) OR 0.98 (0.96 to 1.00, p = 0.11) Diabetes (self-reported) OR 0.83 (0.67 to 1.04, p = 0.11) Stroke (self-reported) OR 1.00 (0.99 to 1.00, p = 0.64) Geographical remoteness OR 1.04 (0.979 to 1.10, p = 0.27) Time since last eye examination OR 1.15 (1.12 to 1.19, p < 0.001) Modelling method of analysis: Multivariate logistic regression analyses Adjustment factors used: Sex, age, time since previous eye examination, geographical remoteness, years of education, diabetes, stroke	Increasing age, male sex, and longer time period since last eye examination are all associated with higher eye care referral rates in adults aged 50 years and older. Years of education, history of diabetes or stroke, and geographical remoteness are not associated with eye care referral rates in adults aged 50 years and older. Only the results of non- Indigenous participants have been extracted.

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
	of bias. 3. Prognostic factor measurement: Moderate risk of bias. Diabetes and stroke history self- reported 4. Outcome measurement: Low risk of bias. 5. Adjustment for other prognostic factors: Low risk of bias. 6. Statistical analysis and reporting: Moderate risk of bias. Only covariates that were significant in univariate analysis were included in the multivariable model.		referral unless new pathology was suspected.			
Khachatryan et al. (2015) (USA)	Study Design: Prospective cohort study Eligibility criteria/recruitment methods: Participants included in this study were selected from the African Descent and Glaucoma Evaluation Study (ADAGES) and Diagnostic Innovations in Glaucoma Study (DIGS). Suspect glaucoma defined as a history of elevated IOP and/or an optic disc appearance suspicious of glaucoma but normal visual fields at study entry. Elevated IOP defined as IOP > 21	Sample size: 357 participants (636 eyes) Participants: Mean (± SD) age at entry 58.1 ± 12.3 years 65% female 67% European descent 33% African descent Mean (± SD) follow-up time 7.1 ± 2.4 years Dates of data collection: January 2003 to not stated	Outcome: Incident visual field damage Method of measurement: Standard automated perimetry with 24-2 Swedish Interactive Threshold Algorithm. Visual fields were defined as abnormal if pattern standard deviation was ≤ 5% and/or glaucoma hemifield test was "outside normal limits." Eyes that developed a repeatable visual defect, defined as 3 consecutive abnormal tests, were defined as developed visual field damage. The development of damage was reviewed by an ophthalmologist to	Prognostic factor: Race Method of measurement: Self-reported Prognostic factor: Age Method of measurement: Self-reported Prognostic factor: IOP Method of measurement: Not stated Prognostic factor: Central corneal thickness	Adjusted prognostic effect (95% confidence interval): African descent vs. European descent by IOP: No significant association at IOP = 10 mmHg to 20 mmHg IOP 22 mmHg HR 2.03 (1.15 to 3.57) IOP 24 mmHg HR 2.71 (1.39 to 5.29) IOP 26 mmHg HR 3.61 (1.61 to 8.08) Mean IOP of cohort (17.8 mmHg) HR 1.12 (0.66 to 1.90) Age (per year) HR 1.02 (0.99 to 1.04) Central corneal thickness (per 40 microns thinner) HR 1.18 (0.86 to 1.60)	Glaucoma suspects of African descent with higher mean IOP are associated with increased risk of visual field damage compared to European glaucoma suspects. Lower baseline visual field mean deviation and increasing mean arterial pressure are associated with increased risk of visual field damage.

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
	mmHg or a history of		confirm that the damage	Method of	Lower SER (per D greater)	
	ocular hypotensive		was glaucomatous, and	measurement:	HR 1.11 (0.84 to 1.34)	
	treatment.		the location of damage	Not stated		
	Inclusion criteria: best-		was consistent on all 3		Disc area (per 0.4 mm <sup>2</sup>	
	corrected VA of 20/40 or		visual fields.	Prognostic	increase) HR 1.06 (0.84 to	
	better, spherical			factor: SER	1.34)	
	refraction less than 5.00			Method of		
	D, cylinder correction less			measurement:	Vertical cup-disc ratio (per 0.1	
	than 3.00 D, open angles			Not stated	increase) HR 1.25 (0.99 to	
	by gonioscopy, African or				1.57)	
	European descent,			Prognostic		
	classed as glaucoma			factor: Disc area,	Baseline visual field mean	
	suspect at baseline, at			vertical cup-disc	deviation (per 0.1 dB	
	least 2 years follow-up, at			ratio	decrease) HR 1.04 (1.02 to	
	least 4 good quality visual			Method of	1.06)	
	field results.			measurement:		
	Exclusion criteria:			Clinician	Mean IOP (per 1 mmHg	
	Coexisting ocular trauma,			assessment of	increase) HR 0.97 (0.92 to	
	retinal disease, uveitis,			stereo-	1.03)	
	non-glaucomatous optic			photographs		
	disc neuropathy, or other				Mean arterial pressure (per 1	
	diseases possibly			Prognostic	mmHg increase) HR 1.03	
	affecting the visual field,			factor: Baseline	(1.00 to 1.06)	
	evidence of consecutive			visual field mean		
	repeatable visual field			deviation	Modelling method of	
	damage at baseline.			Method of	analysis: Multivariable Cox	
				measurement:	proportional hazards model	
	Quality rating: QUIPS			Standard		
	RoB assessment			automated	Adjustment factors used:	
	1. Study participation:			perimetry with 24-2	Race, age, central corneal	
	Low risk of bias.			Swedish	thickness, SER, disc area,	
	Prospective cohort			Interactive	baseline stereophotograph-	
	2. Study attrition: Low risk			Threshold	based vertical cup-disc ratio,	
	of bias.			Algorithm.	baseline visual field mean	
	3. Prognostic factor			Deserves	deviation, mean IOP during	
	measurement: Low risk of			Prognostic	follow-up, mean arterial	
	bias.			factor: Arterial	pressure, and a mean	
	4. Outcome			pressure Mathad of	IOP*mean IOP interaction	
	measurement: Low risk of			Method of	term and a race*mean IOP	
	bias.			measurement:	interaction term	
	5. Adjustment for other			Blood pressure		

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
	prognostic factors: Low risk of bias. 6. Statistical analysis and reporting: Low risk of bias.	Sample size: 3939	Outcome: Incident OAG	measured using DINAMAP@ PRO Monitor Model 100. Mean arterial pressure = (2/3) diastolic pressure + (1/3) systolic pressure. <b>Prognostic</b>	Adjusted prognostic effect	Corticosteroid use is not
Marcus et al. (2012) (The Netherlands)	Study Design: Prospective, population- based cohort study Eligibility criteria/recruitment methods: Part of the Rotterdam Study examining age-related disorders of individuals aged 55 year and older from a district in Rotterdam. Data were used from participants who completed the baseline ophthalmic examination, did not have glaucoma at baseline and completed at least one follow-up examination. Cases with	participants Participants: Mean follow-up 9.8 years Incident glaucoma: 108 participants Mean (± SD) age 68.4 ± 7.1 years 49.1% female No glaucoma: 3831 participants Mean (± SD) age 65.7 ± 6.8 years 58.7% female Dates of data collection: 1991 to	Method of measurement: An incident OAG case was defined as a participant with no glaucomatous visual field loss in both eyes at baseline and glaucomatous visual field loss in at least one eye at follow-up. At each examination, three IOP measurements were taken on each eye and the median value of these three measurements was recorded. The visual field of each eye was screened using a 52-point supra- threshold test that	factor: Corticosteroid use Method of measurement: Data on corticosteroid prescriptions for all participants were obtained from seven fully automated pharmacies using a centralized computer network from 1 January 1991 onward. This included the product name, Anatomical Therapeutic	(95% confidence interval): Ophthalmic steroid use OR 1.04 (0.66 to 1.65, p = 0.86) Inhaled steroid use OR 0.79 (0.42 to 1.48, p = 0.46) Nasal steroid use OR 1.26 (0.74 to 2.13, p = 0.40) Oral steroid use OR 1.03 (0.65 to 1.64, p = 0.89) Ointment steroid use OR 0.70 (0.47 to 1.05, p = 0.086) Age (per year) OR 1.06 (1.04 to 1.09, p < 0.001) Sex (female) OR 0.63 (0.43 to	associated with OAG incidence in adults aged 55 years and older. Age, female sex, positive family history of glaucoma, and high myopia are associated with increased risk of incident OAG in adults aged 55 years and older.
	a history or signs of angle closure (gonioscopy was performed in all identified cases) or secondary glaucoma (except for steroid-induced glaucoma) were excluded. <b>Quality rating:</b> QUIPS RoB assessment	2006	covered the central visual field with a radius of 24° (Humphrey Field Analyser). Visual field loss was defined as non- response to a light stimulus of 6 dB above a threshold-related estimate of the hill of vision in at least three contiguous test points, or four including the blind spot. In	Chemical (ATC) code, number of prescriptions and the date of first prescription. Corticosteroids were classified as ophthalmic steroids, inhaled steroids, nasal steroids, oral steroids and	0.93, p = 0.022) Positive family history of glaucoma OR 2.24 (1.31 to 3.84, p = 0.003) High myopia OR 2.22 (1.13 to 4.38, p = 0.021) Modelling method of analysis: Multivariate logistic regression analyses	

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
	<ol> <li>Study participation: Low risk of bias.</li> <li>Prospective cohort</li> <li>Study attrition: Moderate risk of bias.</li> <li>3939/6630 (59%) eligible participants had follow-up data.</li> <li>Prognostic factor measurement: Moderate risk of bias. Family history self-reported</li> <li>Outcome measurement: Low risk of bias.</li> <li>Adjustment for other prognostic factors: Low risk of bias.</li> <li>Statistical analysis and reporting: Low risk of bias.</li> </ol>		participants with reproducible abnormalities on supra-threshold testing, Goldmann perimetry (baseline and first follow-up) or full- threshold Humphrey Field Analyser 24-2 testing (second follow-up) was performed on both eyes. Visual field loss was considered to be glaucomatous visual field loss only if reproducible and after excluding all other possible causes.	steroid ointments. The number of prescriptions during follow-up was used as a proxy for cumulative dose. Usage before baseline was not considered because the onset of the automated collection of medication data started on 1 January 1991. <b>Prognostic</b> factor: Age, sex, family history of glaucoma Method of measurement: Not explicitly stated <b>Prognostic</b> factor: High myopia Method of measurement: Refraction during eye examination, no further details given. High myopia defined as SER < - 4.00 D.	Adjustment factors used: Age, sex, positive family history of glaucoma, high myopia	
Pasquale & Kang (2011)	Study Design: Prospective cohort study	Sample size: 79,440 participants	Outcome: Incident OAG Method of measurement:	Prognostic factor: Age at menarche	Adjusted prognostic effect (95% confidence interval): Age at menarche:	Age at menarche older than 13 years is associated with increased risk of normal tension

Citation	Study Details	Participants &	Outcomes	Prognostic	Key findings	Interpretation and observations
(Country) (USA)	Eligibility	setting Participants: 100%	Self-reported in	factors Method of	> 13 years and normal tension	glaucoma in women aged
(03A)	criteria/recruitment	female	questionnaires, then	measurement:	glaucoma RR 1.47 (1.01 to	40 years or older.
	methods: The Nurses'	All were aged 40 or	followed up by	Self-reported in	2.13)	
	Health Study is an	more years	investigator review of	biennial	< 12 years (ref)	Five years or greater use
	ongoing population-	more years	medical records to	questionnaires.		of oral contraceptives and
	based cohort of	Dates of data	confirm diagnosis.	queettermaneet	Reproductive duration:	time since discontinuing
	registered female nurses.	collection: 1980 to	communication and generation	Prognostic	< 36 years RR 0.93 (0.71 to	use of oral contraceptives
	The Nurses' Health Study	2006		factor:	1.22)	less than 10 years is
	was established in 1976			Reproductive	36-38 years RR 0.94 (0.73 to	associated with increased
	when 121,700 United			duration	1.21)	risk of OAG in women
	States women were			Method of	39-40 years (ref)	aged 40 years or older.
	invited to complete a			measurement:	≥ 41 years RR 0.96 (0.73 to	5 ,
	questionnaire regarding			Self-reported in	1.27)	Reproductive duration
	lifestyle, health			biennial		and parity are not
	behaviour, and chronic			questionnaires.	Oral contraceptives, duration	associated with OAG in
	diseases. Follow-up			Taken as age at	of use:	women aged 40 years or
	biennial questionnaires			natural menopause	Ever used RR 1.14 (0.98 to	older.
	were used to update this			minus age at	1.34)	
	data and report newly			menarche.	< 2 years RR 1.10 (0.89 to	
	diagnosed medical				1.36)	
	conditions including			Prognostic	2-4 years RR 1.04 (0.81 to	
	glaucoma.			factor: Oral	1.35)	
	For this study, the follow-			contraceptive use	5+ years RR 1.25 (1.02 to	
	up was from 1980 to			Method of	1.53)	
	2006.			measurement:	Never used (ref)	
	Eligible participants			Self-reported in		
	contributed to the study if			biennial	Oral contraceptives, time	
	they reached age 40			questionnaires.	since discontinuing use:	
	years and if they reported			After 1984, oral	≥ 25 years RR 1.13 (0.91 to	
	having had an eye exam			contraceptive use	1.40)	
	in the period at risk. Eligible			was defined as ever / never.	20-24 years RR 1.06 (0.82 to	
				ever / never.	1.38) 15-19 years RR 1.20 (0.91 to	
	participants contributed person-time in 2-year			Prognostic	1.59)	
	units from the return date			factor: Parity	1.59) < 10 years RR 1.39 (1.01 to)	
	of the first biennial			Method of	1.91)	
	questionnaire until the			measurement:	Never used (ref)	
	occurrence of a report of			Self-reported in		
	glaucoma, cancer, death,			biennial	Parity:	
	or loss to follow-up, or			questionnaires.	No children RR 0.85 (0.60 to	
				94004011141100.	1.21)	

	until 2006, whichever came first. At each 2-year follow-up cycle, only women who indicated they received	setting			1-2 children (ref)	
	<ul> <li>an eye exam in the previous 2 years were eligible to contribute person-time to the study.</li> <li>Quality rating: QUIPS RoB assessment <ol> <li>Study participation:</li> <li>Moderate risk of bias.</li> <li>Prospective cohort.</li> <li>79440 / 121700 had sufficient data to be included</li> <li>Study attrition: Low risk of bias.</li> <li>Prognostic factor measurement: Moderate risk of bias.</li> <li>Prognostic factor measurement: Moderate risk of bias. Self reported via questionnaire</li> <li>Outcome measurement: Moderate risk of bias. Self-reported glaucoma - but verified by review of medical records.</li> <li>Adjustment for other prognostic factors: Low risk of bias.</li> </ol> </li> </ul>				3 children RR 1.08 (0.90 to 1.29) 4+ children RR 1.00 (0.84 to 1.19) Modelling method of analysis: Cox proportional hazard analyses Adjustment factors used: Age, time-interval at risk, family history of glaucoma, African ancestry, hypertension, diabetes, smoking status, alcohol intake, caffeine intake, BMI, physical activity, post- menopausal hormone use, oral contraceptive use, parity, age at menopause	
Stem et al. (2013) (USA)	bias. Study Design: Retrospective, longitudinal cohort analysis	Sample size: 494,165 participants	Outcome: Incident CRVO Method of measurement: Identified by reported ICD code. To	Prognostic factor: sex, ethnicity, household net	Adjusted prognostic effect (95% confidence interval): Sex (female) HR 0.75 (0.66 to 0.85, p < 0.0001)	Black ethnicity, lower household net worth, having hypertension (alone or in combination
Citation (Country)	Study Details	Participants &	Outcomes	Prognostic factors	Key findings	Interpretation and
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Citation (Country)	Eligibility criteria/recruitment methods: Investigators used the i3 InVision Data Mart database (Ingenix, Eden Prairie, MN), which contains detailed records of all beneficiaries in a managed care network throughout the United States. The dataset contains all individuals with ≥ 1 International Classification of Diseases, 9th Revision, Clinical Modification (ICD-9-CM) codes for eye-related diagnoses; ≥ 1 Current Procedural Terminology codes for any eye-related visits, diagnostic, or therapeutic procedures; or any other claim submitted by an ophthalmologist or optometrist from January 1 2001 to December 31 2009. The investigators had access to all medical claims for ocular and non-ocular conditions and sociodemographic information. Individuals were included	Participants & setting Participants: 1302 (0.26%) with newly diagnosed CRVO Mean (± SD) age 65.7 ± 8.1 years (those without CRVO diagnosis), 69.9 ± 8.4 (those with CRVO diagnosis) 41.7% male Ethnicity: 79.7% White, 4.9% Black, 3.4% Latino, 1.6% Asian- American, 0.7% other, 9.7% unknown Dates of data collection: January 2001 to December 2009	Outcomes case of CRVO, individuals must have had at least 1 eye care visit during their first 2 years in the plan (with no documented diagnosis of CRVO) and then must have been diagnosed with CRVO at a subsequent visit after the index date (2 years after entry into the plan). Beneficiaries were identified with a CRVO if they had ≥ 1 billing records with the ICD-9- CM code 362.35.	Prognostic factors syndrome components, vascular disease, ophthalmic disease Method of measurement: Data retrieved from i3 InVision Data Mart database.	Ethnicity: Black HR 1.58 (1.25 to 1.99, p < 0.0001) Asian-American HR 0.75 (0.43 to 1.30, p = 0.31) White (ref) Household net worth: > \$500,000 HR 0.73 (0.56 to 0.96, p = 0.02) < \$25,000 (ref) Metabolic syndrome: Hypertension HR 1.66 (1.14 to 2.42, p = 0.01) Hypertension and diabetes HR 1.82 (1.15 to 2.89, p = 0.01) Hypertension and hyperlipidaemia HR 1.46 (1.04 to 2.05, p = 0.03) Hypertension, hyperlipidaemia and diabetes HR 1.58 (1.11 to 2.23, p = 0.01) No diabetes, hypertension or hyperlipidaemia (ref) Vascular disease: Cerebrovascular accident HR 1.44 (1.23 to 1.68, p < 0.0001) Peripheral artery disease HR 1.15 (1.00 to 1.33, p = 0.05) Myocardial infarction HR 0.72 (0.57 to 0.92, p = 0.01) Hypercoagulable state HR	Interpretation and observations hyperlipidaemia, or both), previous cerebrovascular accident, having peripheral arterial disease, hypercoagulable state, OAG, AMD, and cataract are all associated with higher risk for incident CRVO in adults aged 55 years and over. Previous myocardial infarction is associated with lower risk of incident CRVO in adults aged 55 years and over.
	sociodemographic information. Individuals were included in the analysis if they met the following criteria:				Myocardial infarction HR 0.72 (0.57 to 0.92, $p = 0.01$ ) Hypercoagulable state HR 2.45 (1.40 to 4.28, $p = 0.002$ )	
	continuous enrolment in the medical plan for at least 2 years, ≥ 2 visits to an eye care provider				Ophthalmic disease: OAG HR 1.50 (1.30 to 1.72, p < 0.0001)	

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
	<ul> <li>(ophthalmologist or optometrist), and age ≥ 55 years. Individuals were excluded if they received a diagnosis of CRVO during the first 2 years they were enrolled in the plan to exclude non-incident cases.</li> <li>Quality rating: QUIPS RoB assessment 1. Study participation: Low risk of bias.</li> <li>2. Study attrition: Low risk of bias.</li> <li>3. Prognostic factor measurement: Low risk of bias.</li> <li>4. Outcome measurement: Low risk of bias.</li> <li>5. Adjustment for other prognostic factors: Low risk of bias.</li> <li>6. Statistical analysis and reporting: Low risk of bias.</li> </ul>				AMD HR 1.50 (1.31 to 1.72, p < 0.0001) Cataract HR 1.24 (1.08 to 1.42, p = 0.003) <b>Modelling method of</b> <b>analysis:</b> Multivariate Cox regression analysis <b>Adjustment factors used:</b> age, sex, ethnicity, education level, household net worth, region of residence, ocular co- morbidities, systemic co- morbidities, Charlson co- morbidity index	
Stingl et al. (2023) (Germany)	Study Design: Prospective, population- based cohort study Eligibility criteria/recruitment methods: Random sampling of residents of the State of Rhine- Palatine by the regional registration office.	Sample size: 10,175 participants (9978 right eyes, 9952 left eyes) Participants: Mean (± SD) age 53.5 ± 10.5 years, range 35-74 years 48.5% female Dates of data collection: 2007 to 2017	Outcome: 5-year change in refractive error Method of measurement: Refractive error measurement was conducted without cycloplegia. Refractive values were measured in spherical and cylindrical dioptres (D), cylindrical power was indicated in negative sign	Prognostic factor: Sex, age, smoker status, education, occupation Method of measurement: Method of measurement/ data collection not explicitly stated.	Adjusted prognostic effect (95% confidence interval): Myopic change at 5 years: Sex (female) OR 1.49 (1.28 to 1.73, p < 0.001) Age (per year) OR 0.52 (0.49 to 0.55, p <0.001) Baseline SER (per dioptre) OR 0.89 (0.87 to 0.91, p < 0.001) Myopic change no significant association with cardiovascular parameters,	Female sex, younger age and baseline myopic SER are associated with increased risk of having a myopic shift in refractive error at 5 years in adults aged 35 to 74 years. Increasing age and being a smoker are associated with an increased risk of a hyperopic shift in refractive error at 5 years

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
	Inclusion criteria: mental	5	convention. SER was	Prognostic	physical activity, smoking	in adults aged 35 to 74
	and physical ability to		computed as SER =	factor: Baseline	history, IOP, lens opacity,	years.
	visit the study centre and		sphere + 0.5 × cylinder.	SER	education, or occupation.	
	to pass through the		No refractive change was	Method of		Cardiovascular
	examinations, sufficient		defined as -0.50 to +0.50	measurement:	Hyperopic change at 5 years:	parameters, physical
	knowledge of the German		D change in SER, myopic	As for outcome.	Age (per year) OR 1.62 (1.52	activity, IOP, lens opacity,
	language.		shift as < −0.50 D and		to 1.72, p <0.001)	education, and
	All study participants with		hyperopic shift as $> +0.50$	Prognostic	Smoker OR 1.31 (1.14 to	occupation were not
	objective refraction		D.	factor: IOP	1.50, p < 0.001)	associated with increased
	measurement at both			Method of	Hyperopic change no	risk of either myopic or
	baseline and 5-year			measurement:	significant association with	hyperopic shift at 5 years
	follow-up examinations			Non-contact	sex, cardiovascular	in adults aged 35 to 74
	were included.			tonometry (Nidek	parameters, physical activity,	years.
	Exclusion criteria:			NT-2000)	baseline SER, IOP, lens	
	previous corneal or			Dramastia	opacity, education, or	
	cataract surgery,			Prognostic factor: Lens	occupation.	
	aphakia.				Modelling method of	
	Quality rating: QUIPS			opacity Method of	analysis: Multivariable logistic	
	RoB assessment			measurement:	regression analyses	
	1. Study participation:			Slit lamp	regression analyses	
	Low risk of bias.			examination	Adjustment factors used:	
	Prospective cohort			examination	Sex, age, SER, IOP, presence	
	2. Study attrition:			Prognostic	of cataract, cardiovascular	
	Moderate risk of bias.			factor: Physical	parameters, BMI, physical	
	12423/15010 (83%) of			activity	activity, smoking, education,	
	eligible participants had			Method of	occupation	
	follow-up data			measurement:		
	3. Prognostic factor			Completion of the		
	measurement: Low risk of			Short		
	bias.			Questionnaire to		
	4. Outcome			Assess Health-		
	measurement: Low risk of			enhancing physical		
	bias.			activity		
	5. Adjustment for other					
	prognostic factors:			Prognostic		
	Moderate risk of bias.			factor:		
	Family history of			Cardiovascular		
	refractive error and			parameters		
	ethnicity not included			Method of		
	6. Statistical analysis and			measurement:		

Citation	Study Details	Participants &	Outcomes	Prognostic	Key findings	Interpretation and
(Country)	-	setting		factors		observations
	reporting: Moderate risk of bias. Unclear whether strategy for model			Laboratory measurements		
	building is appropriate			Prognostic		
	and is based on a			factor: BMI		
	conceptual framework or			Method of		
	model			measurement:		
				Height and weight measured and BMI		
				calculated as		
				weight/height <sup>2</sup>		
		Sample size: 132,046	Outcome: Referral to a	Prognostic	Adjusted prognostic effect	Delayed attendance for
Wright et al.	Study Design:	participants, 444,045	GP	factor: Delayed	(95% confidence interval):	eye examinations is
(2020)	Retrospective cohort analysis	eye examinations,	Method of	attendance at eye	Aged 60-69:	associated with increased
(UK)	analysis	311,999 examination	measurement:	examination	Delayed eye exam attendance	risk of requiring a GP
(OII)	Eligibility	intervals	Data were extracted from	Method of	OR 1.30 (1.04 to 1.61)	referral for adults aged 60
	criteria/recruitment	Participants: Aged 60-	the Family Practitioner Services Ophthalmic	measurement: Data were	Early eye exam attendance OR 2.86 (2.36 to 3.46)	years and older. Early attendance is also
	methods: Information on	69 43.2% male	Database.	extracted from the	On-time eye exam attendance	associated with increased
	attendance at routine eye	Aged $\geq$ 70 40.7% male.	Dutababbi	Family Practitioner	(ref)	risk of referral for adults
	examinations was drawn	No other details given.		Services		aged 60 years or older,
	from the Family Practitioner Services			Ophthalmic	Aged ≥ 70:	though this is driven by
	Ophthalmic Database	Dates of data		Database.	Delayed eye exam attendance	early attendance usually
	(managed by the	collection: October		Examination	OR 1.07 (1.01 to 1.13)	being due to symptomatic
	Business Services	2009 to September 2014		intervals were split into three	Early eye exam attendance OR 2.72 (2.58 to 2.87)	problems or early recall suggested by the
	Organisation, NI			categories based	On-time eye exam attendance	optometrist.
	Department of Health),			on recommended	(ref)	optomotion
	an administrative			intervals for each		
	database used to manage payment to			age group. The	Modelling method of	
	service providers.			first category 'on-	analysis: Multivariable logistic	
	Records of eye			time' consisted of	regression	
	examinations of those			intervals	Adjustment factors used:	
	aged ≥ 60 years			conforming to recommendations	sex, religion, eligibility for NHS	
	conducted during a 5-			(24 months for	sight test on health grounds,	
1	year period (October			those aged 60–69,	eligibility for NHS sight test on	
	2009 to September 2014			12 months for	income grounds, general	
	inclusive) were extracted. The cohort consisted of			those aged $\geq$ 70).	health, household structure,	
	all community-dwelling			Longer intervals (>	tenure, household cars,	
				24 and > 12	practice density, drive time,	

Citation (Country)	Study Details	Participants & setting	Outcomes	Prognostic factors	Key findings	Interpretation and observations
	respondents to the 2011			months for the	highest qualification, carer	
	Census aged ≥ 60 years			younger and older	status, household adaptations	
	at the beginning of the			groups	for visual difficulties, area	
	study period that had			respectively) were	income deprivation	
	attended at least two free			classified as		
	eye examinations during			'delayed		
	the period. A longitudinal			attendance'. The		
	sequence of eye			third category		
	examinations was			'early recall'		
	constructed for each			consisted of		
	individual and from these,			intervals shorter		
	the analysis dataset of			than		
	311,999 examination			recommended.		
	intervals was calculated.			Early recall may		
				occur when an		
	Quality rating: QUIPS			individual returns		
	RoB assessment			to the optometrist		
	<ol> <li>Study participation:</li> </ol>			with visual		
	Low risk of bias. Cohort			symptoms or at the		
	2. Study attrition: Low risk			request of the		
	of bias.			optometrist to		
	<ol><li>Prognostic factor</li></ol>			monitor an ocular		
	measurement: Low risk of			condition that does		
	bias.			not warrant		
	4. Outcome			immediate GP		
	measurement: Low risk of			referral.		
	bias.			Classifications		
	5. Adjustment for other			were based on		
	prognostic factors: Low			calendar months.		
	risk of bias.					
	6. Statistical analysis and					
	reporting: Low risk of bias					

Abbreviations: AL: axial length; AMD: age-related macular degeneration; BMI: body mass index; CRVO: central retinal vein occlusion; D: dioptre; dB: decibel; EG: exfoliation glaucoma; EGS: exfoliation glaucoma suspect; HL: hearing level; HR: hazard ratio; IQR: interquartile range; logMAR: logarithm of the minimum angle of resolution; OAG: open-angle glaucoma; OHT: ocular hypertension; OR: odds ratio; QUIPS: Quality In Prognosis Studies tool; RBC: red blood cell; RoB: risk of bias; SD: standard deviation; SER: spherical equivalent refraction; VA: visual acuity; VI: visual impairment

### 6.3 Quality appraisal

#### Table 10: Quality in Prognostic factor Studies (QUIPS) tool for included primary studies

Citation	1. Study participation	2. Study attrition	3. Prognostic factor measurement	4. Outcome measurement	5. Adjustment for other prognostic factors	6. Statistical analysis and reporting
Barsam et al. (2017)	High	High	Low	Low	Low	Moderate
Ekström (2012)	Low	Low	Moderate	Low	Moderate	Low
Ekström & Hårleman (2023)	Low	Low	Moderate	Low	Low	Moderate
Elmore et al. (2022)	Low	Moderate	Moderate	Moderate	Low	Low
Fernandez-Montero et al. (2017)	Low	Moderate	Moderate	Moderate	Low	Low
Gopinath et al. (2014)	Moderate	Moderate	Low	Low	Moderate	Low
Guggenheim et al. (2012)	Low	Moderate	Moderate	Low	Low	Low
Hopf et al. (2022)	Low	Low	Low	Low	Low	Moderate
Irving et al. (2016)	Low	Low	Low	Moderate	Low	Low
Kang et al. (2012)	Moderate	Low	Moderate	Moderate	Low	Low
Keel et al. (2017)	Low	Low	Moderate	Low	Low	Moderate
Khachatryan et al. (2015)	Low	Low	Low	Low	Low	Low
Marcus et al. (2012)	Low	Moderate	Moderate	Low	Low	Low
Pasquale & Kang (2011)	Moderate	Low	Moderate	Moderate	Low	Low
Stem et al. (2013)	Low	Low	Low	Low	Low	Low
Stingl et al. (2023)	Low	Moderate	Low	Low	Moderate	Moderate
Wright et al. (2020)	Low	Low	Low	Low	Low	Low

**Signalling items: 1. Study participation** (a) Adequate participation in the study by eligible persons, (b) Description of the target population or population of interest, (c) Description of the baseline study sample, (d) Adequate description of the sampling frame and recruitment, (e) Adequate description of the period and place of recruitment, (f) Adequate description of inclusion and exclusion criteria; 2. Study attrition (a) Adequate response rate for study participants, (b) Description of attempts to collect information on participants who dropped out, (c) Reasons for loss to follow-up are provided, (d) Adequate description of participants lost to follow-up, (e) There are no important differences between participants who completed the study and those who did not; **3. Prognostic factor measurement** (a) A clear definition or description of the prognostic factor is provided, (b) Method of prognostic factor measurement is adequately valid and reliable, (c) Continuous variables are reported or appropriate cutpoints are used, (d) The method and setting of measurement of prognostic factor is the same for all study participants, (e) Adequate proportion of the study sample has complete data for the prognostic factor, (f) Appropriate methods of imputation are used for missing prognostic factor data; **4. Outcome measurement** (a) A clear definition of the outcome is provided, (b)

Method of outcome measurement used is adequately valid and reliable, (c) The method and setting of outcome measurement is the same for all; **5. Adjustment for other prognostic factors** (a) All other important prognostic factors are measured, (b) Clear definitions of the important prognostic factors measured are provided, (c) Measurement of all important prognostic factors is adequately valid and reliable, (d) The method and setting of prognostic factors measurement are the same for all study participants, (e) Appropriate methods are used to deal with missing values of prognostic factors, such as multiple imputation, (f) Important prognostic factors are accounted for in the study design, (g) Important prognostic factors are accounted for in the analysis; **6. Statistical analysis and reporting** (a) Sufficient presentation of data to assess the adequacy of the analytic strategy, (b) Strategy for model building is appropriate and is based on a conceptual framework or model, (c) The selected statistical model is adequate for the design of the study, (d) There is no selective reporting of results.

Study	1. Study eligibility criteria	2. Identification and selection of studies	3. Data collection and study appraisal	4. Synthesis and findings	5. Risk of bias in the review
Dinu et al. (2019)	Low	Low	Low	Unclear	Unclear
Kessel et al. (2015)	Low	Unclear	Low	Low	Low

#### Table 11: Risk of Bias in Systematic reviews (ROBIS) tool for included secondary studies

**Signalling items: Domain 1 Study eligibility criteria** 1.1 Did the review adhere to pre-defined objectives and eligibility criteria? 1.2 Were the eligibility criteria appropriate for the review question? 1.3 Were eligibility criteria unambiguous? 1.4 Were all restrictions in eligibility criteria based on study characteristics appropriate (e.g. date, sample size, study quality, outcomes measured)? 1.5 Were any restrictions in eligibility criteria based on sources of information appropriate (e.g. publication status or format, language, availability of data)? Domain 2 Identification and selection of studies 2.1 Did the search include an appropriate range of databases/electronic sources for published and unpublished reports? 2.2 Were methods additional to database searching used to identify relevant reports? 2.3 Were the terms and structure of the search strategy likely to retrieve as many eligible studies as possible? 2.4 Were restrictions based on date, publication format, or language appropriate? 2.5 Were efforts made to minimise error in selection of studies? Domain 3 Data collection and study appraisal 3.1 Were efforts made to minimise error in data collection? 3.2 Were sufficient study characteristics available for both review authors and readers to be able to interpret the results? 3.3 Were all relevant study results collected for use in the synthesis? 3.4 Was risk of bias (or methodological quality) formally assessed using appropriate criteria? 3.5 Were efforts made to minimise error in risk of bias assessment? Domain 4 Synthesis and findings 4.1 Did the synthesis include all studies that it should? 4.2 Were all pre-defined analyses reported or departures explained? 4.3 Was the synthesis appropriate given the nature and similarity in the research questions, study designs and outcomes across included studies? 4.4 Was between-study variation (heterogeneity) minimal or addressed in the synthesis? 4.5 Were the findings robust, e.g. as demonstrated through funnel plot or sensitivity analyses? 4.6 Were biases in

#### 6.4 Information available on request

The protocol, search strategies, and excluded studies for this rapid review are available on request.

### 7. ADDITIONAL INFORMATION

#### 7.1 **Conflicts of interest**

The authors declare they have no conflicts of interest to report.

#### 7.2 Acknowledgements

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## 8. APPENDIX

### **APPENDIX 1: Search Strategies**

#### **Searches for Secondary Research**

#### **Medline Search Strategy**

Ovid MEDLINE(R) ALL <1946 to August 03, 2023> Conducted 03.08.2023

#	Search Query	Results
1	exp Prognosis/	1922657
2	exp Incidence/	301423
3	exp Risk Assessment/	310857
4	exp Decision Support Techniques/	82250
5	(prognos* or prevention or progress* or diagnos* or detect* or prevalence or incidence or rate*).tw.	10451982
6	(risk* adj2 (assess* or factor*)).tw.	878756
7	((first or initial) adj (episode* or detection)).tw.	21793
8	1 or 2 or 3 or 4 or 5 or 6 or 7	11718920
9	exp Vision Screening/	2467
10	exp Vision Tests/	117699
11	exp Mass Screening/	144061
12	((eye* or sight or vision or visual) adj1 (test* or exam* or screen* or follow- up)).tw	17962
13	(asymptomatic adj2 (test* or exam* or screen*)).tw.	4556
14	(routine adj2 (test* or exam* or screen*)).tw.	43911
15	((eye* or sight or vision or visual or screening) adj3 (frequenc* or interval* or recall*)).tw.	8138
16	9 or 10 or 11 or 12 or 13 or 14 or 15	318610
17	exp Eye Diseases/	637918
18	exp Vision, Ocular/	29938
19	exp Visual Acuity/	92659
20	(ametrop* or emmetrop* or glaucoma or diabetic retinopathy or refractive error or macular degeneration or cateract* or presbyop* or amblyop* or myop* or hyperop* or hypermetrop* or astigmat* or anisometrop* or vision impairment* or vision loss* or vis* acuity).tw.	255933
21	exp Cataract/di, dg, pc [Diagnosis, Diagnostic Imaging, Prevention & Control]	4774
22	exp Glaucoma/di, dg, pc [Diagnosis, Diagnostic Imaging, Prevention & Control]	15607
23	exp Macular Degeneration/di, dg, pc [Diagnosis, Diagnostic Imaging, Prevention & Control]	9999
24	exp Diabetic Retinopathy/di, dg, pc [Diagnosis, Diagnostic Imaging, Prevention & Control]	9211
25	exp Refractive Errors/di, dg, pc [Diagnosis, Diagnostic Imaging, Prevention & Control]	7806

26	17 or 18 or 19 or 20 or 21 or 22 or 23 or 24 or 25	757960
27	(UK or United kingdom or England or Wales or Scotland or Northern Ireland or Ireland or Australia or Canada or New Zealand or USA or United States or Austria or Finland or Germany or Malta or Netherlands or Norway or Spain or Sweden or Switzerland).tw,cp.	29980939
28	exp "Systematic Review"/	234551
29	exp Meta-Analysis/	184949
30	exp Systematic Reviews as Topic/	10951
31	exp Meta-Analysis as Topic/	27324
32	(systematic review* or meta analys#s or review*).pt.	3356402
33	(systematic adj2 (review* or overview*)).ti,ab,kf.	310829
34	(quantitative adj2 (review* or overview* or synthes*)).ti,ab,kf.	6829
35	(meta-analy* or metaanaly* or meta- synthes#s or metasynthes#s).tw.	277814
36	rapid review*.tw.	1967
37	28 or 29 or 30 or 31 or 32 or 33 or 34 or 35 or 36	3455183
38	8 and 16 and 26 and 27 and 37	3178
39	limit 38 to (english language and yr="2009 -Current")	1967

#### **EMBASE Search Strategy**

Conducted 03.08.2023

#	Search Query	Results
1	exp Prognosis/	912238
2	exp Incidence/	670315
3	exp Risk Assessment/	737723
4	exp Decision Support Techniques/	34339
5	(prognos* or prevention or progress* or diagnos* or detect* or prevalence or incidence or rate*).tw.	13884935
6	(risk* adj2 (assess* or factor*)).tw.	1278642
7	((first or initial) adj (episode* or detection)).tw.	33689
8	1 or 2 or 3 or 4 or 5 or 6 or 7	14827598
9	exp Vision Screening/	43654
10	exp Vision Tests/	43654
11	exp Mass Screening/	307655
12	((eye* or sight or vision or visual) adj1 (test* or exam* or screen* or follow- up)).tw	24336
13	(asymptomatic adj2 (test* or exam* or screen*)).tw.	6538
14	(routine adj2 (test* or exam* or screen*)).tw.	66065
15	((eye* or sight or vision or visual or screening) adj3 (frequenc* or interval* or recall*)).tw.	11434
16	9 or 10 or 11 or 12 or 13 or 14 or 15	442099
17	exp Eye Diseases/	1095511

18	exp Vision, Ocular/	316640
19	exp Vision, Ocular/ exp Visual Acuity/	154663
20		
20	(ametrop* or emmetrop* or glaucoma or	327995
	diabetic retinopathy or refractive error or macular degeneration or cateract* or	
	presbyop* or amblyop* or myop* or	
	hyperop* or hypermetrop* or astigmat* or	
	anisometrop* or vision impairment* or	
	vision loss* or vis* acuity).tw.	
21	exp Cataract/di, dg, pc [Diagnosis,	5974
21	Diagnostic Imaging, Prevention & Control	5974
22	exp Glaucoma/di, dg, pc [Diagnosis,	14182
22	Diagnostic Imaging, Prevention & Control	14102
23	exp Macular Degeneration/di, dg, pc	2396
23	[Diagnosis, Diagnostic Imaging,	2390
	Prevention & Control]	
24	exp Diabetic Retinopathy/di, dg, pc	8275
24	[Diagnosis, Diagnostic Imaging,	0275
	Prevention & Control]	
25	exp Refractive Errors/di, dg, pc	6537
20	[Diagnosis, Diagnostic Imaging,	0007
	Prevention & Control]	
26	17 or 18 or 19 or 20 or 21 or 22 or 23 or	1297500
20	24 or 25	1257500
27	(UK or United kingdom or England or	30453772
21	Wales or Scotland or Northern Ireland or	00400772
	Ireland or Australia or Canada or New	
	Zealand or USA or United States or	
	Austria or Finland or Germany or Malta or	
	Netherlands or Norway or Spain or	
	Sweden or Switzerland).tw,cp.	
28	exp "Systematic Review"/	424400
29	exp Meta-Analysis/	288688
30	exp Systematic Reviews as Topic/	32284
31	exp Meta-Analysis as Topic/	53131
32	(systematic review* or meta analys#s or	3104591
-	review*).pt.	
33	(systematic adj2 (review* or	379724
	overview*)).ti,ab,kf.	-
34	(quantitative adj2 (review* or overview* or	7763
	synthes*)).ti,ab,kf.	
35	(meta-analy* or metaanaly* or meta-	353655
	synthes#s or metasynthes#s).tw.	
36	rapid review*.tw.	2326
37	28 or 29 or 30 or 31 or 32 or 33 or 34 or	3551601
-	35 or 36	
38	8 and 16 and 26 and 27 and 37	3030
39	limit 38 to (english language and yr="2009	2113
	-Current")	

#### **CINAHL Search Strategy**

Conducted 03.08.2023

#	Search Query	Results
1	(MH "Prognosis+")	550866
2	(MM "Incidence")	1707
3	(MM "Risk Assessment")	62988

	r	1
4	TX (prognos* or prevention or progress*	3479872
	or diagnos* or detect* or prevalence or	
	incidence or rate*)	
5	AB (risk* N2 (assess* or factor*))	239041
6	TX ((first or initial) N1 (episode* or	12,663
	detection))	
7	S1 OR S2 OR S3 OR S4 OR S5 OR S6	3727296
8	(MM "Vision Screening")	822
9	(MH "Vision Tests+")	7549
10	TX ((eye* or sight or vision or visual) N1	19149
	(test* or exam* or screen* or follow-up))	
11	TX (asymptomatic N2 (test* or exam* or	3751
10	screen*))	00540
12	TX (routine N2 (test* or exam* or	33516
40	screen*))	00.47
13	AB ((eye* or sight or vision or visual or	2647
	screening) N3 (frequenc* or interval* or	
4.4		C1110
14	S8 OR S9 OR S10 OR S11 OR S12 OR	61140
45		102000
15	(MH "Eye Diseases+")	103998
16	(MH "Diagnosis, Eye+")	23008
17	(MM "Vision, Subnormal")	1133
18	(MM "Visual Acuity")	3891
19	AB (ametrop* or emmetrop* or glaucoma	33038
	or diabetic retinopathy or refractive error	
	or macular degeneration or cateract* or	
	presbyop* or amblyop* or myop* or	
	hyperop* or hypermetrop* or astigmat* or	
	anisometrop* or vision impairment* or	
20	vision loss* or vis* acuity)	459
20	(MM "Cataract/DI/PC/PR") (MH "Glaucoma+/DI/PC/PR")	2292
21	· · · · · · · · · · · · · · · · · · ·	2292
22	(MH "Macular Degeneration+/DI/PC/PR")	
	(MM "Diabetic Retinopathy/DI/PC/PR")	1427
24 25	(MH "Refractive Errors+/DI/PC/PR") S15 OR S16 OR S17 OR S18 OR S19	1472 124965
25		124900
	OR S20 OR S21 OR S22 OR S23 OR S24	
26	TX (UK or United kingdom or England or	7581371
20	Wales or Scotland or Northern Ireland or	7001371
	Ireland or Australia or Canada or New	
	Zealand or USA or United States or	
	Austria or Finland or Germany or Malta or	
	Netherlands or Norway or Spain or	
	Sweden or Switzerland)	
27	(MM "Systematic Review")	1514
28	(MM "Meta Analysis")	1779
20	PT (systematic review* or meta analys#s	479918
23	or review*)	010017
30	AB (systematic N2 (review* or overview*))	84071
50		
	A R (auantitative NC) (reviews or evenuews	
31	AB (quantitative N2 (review* or overview*	2434
31	or synthes*))	
	or synthes*)) AB (meta-analy* or metaanaly* or meta-	72993
31 32	or synthes*)) AB (meta-analy* or metaanaly* or meta- synthes#s or metasynthes#s)	72993
31 32 33	or synthes*)) AB (meta-analy* or metaanaly* or meta- synthes#s or metasynthes#s) AB rapid review*	72993 1295
31 32	or synthes*)) AB (meta-analy* or metaanaly* or meta- synthes#s or metasynthes#s)	72993

35	S7 AND S14 AND S25 AND S26 AND S34	408
	Limiters - Published Date: 20090101- 20230831; Language: English	

#### **Epistemonikos**

Search Query	Results
(advanced_title_en:(advanced_title_en:((advanced_title_en:(vision screening) OR advanced_abstract_en:(vision screening))) OR advanced_title_en:(vision test) OR advanced_title_en:(eye test) OR advanced_title_en:(eye sight) OR advanced_title_en:(visual acuity)) OR	148
advanced_abstract_en:(advanced_title_en:((advanced_title_en:(vision screening)) OR advanced_abstract_en:(vision screening))) OR advanced_title_en:(vision test) OR advanced_title_en:(eye test) OR advanced_title_en:(eye test) OR advanced_title_en:(visual acuity))) AND (advanced_title_en:((prognos* OR prevention OR progress* OR diagnos* OR detect* OR prevalence OR incidence OR rate*)) OR advanced_abstract_en:((prognos* OR prevention OR progress* OR diagnos* OR detect* OR prevalence OR incidence OR rate*))) [Filters: classification=systematic-review, protocol=no, min_year=2009, max_year=2023]	

Database	Results
Medline (Ovid)	1967
EMBASE (Ovid)	2113
CINAHL (EBSCO)	408
Epistemonikos	148
Cochrane	5
TOTAL	4641

#### **Searches for Primary Research Medline Search Strategy**

Ovid MEDLINE(R) ALL <1946 to August 21, 2023> Conducted 21.08.2023

#	Search Query	Results
1	exp Prognosis/	1925629
2	Incidence/	301723
3	Risk Assessment/	306886
4	Decision Support Techniques/	22494
5	(prognos* or prevention or progress* or diagnos* or detect* or prevalence or incidence or rate*).tw.	10479336
6	(risk* adj2 (assess* or factor*)).tw.	881915
7	((first or initial) adj (episode* or detection)).tw.	21854
8	1 or 2 or 3 or 4 or 5 or 6 or 7	11702337
9	exp Vision Screening/	2468
10	exp Vision Tests/	117787
11	exp Mass Screening/	144174
12	((eye* or sight or vision or visual) adj1 (test* or exam* or screen* or follow- up)).ti,ab.	18010
13	(asymptomatic adj2 (test* or exam* or screen*)).tw.	4567

4.4	(reutine edi) (teets or everys er	44000
14	(routine adj2 (test* or exam* or screen*)).tw.	44026
15	((eye* or sight or vision or visual or screening) adj3 (frequenc* or interval* or	8155
16	recall*)).ti,ab. 9 or 10 or 11 or 12 or 13 or 14 or 15	210000
16 17		318989
18	Eye Diseases/	38697
	Vision, Ocular/	27018
19	Visual Acuity/	85126
20	(ametrop* or emmetrop* or glaucoma or diabetic retinopathy or refractive error or	256482
	macular degeneration or cateract* or	
	presbyop* or amblyop* or myop* or	
	hyperop* or hypermetrop* or astigmat* or	
	anisometrop* or vision impairment* or	
04	vision loss* or vis* acuity).tw.	4007
21	Cataract/di, dg, pc [Diagnosis, Diagnostic Imaging, Prevention & Control]	4607
22	Glaucoma/di, dg, pc [Diagnosis,	9563
	Diagnostic Imaging, Prevention & Control]	
23	Macular Degeneration/di, dg, pc	4672
	[Diagnosis, Diagnostic Imaging,	
	Prevention & Control]	
24	Diabetic Retinopathy/di, dg, pc	9241
	[Diagnosis, Diagnostic Imaging,	
	Prevention & Control]	
25	Refractive Errors/di, dg, pc [Diagnosis,	2309
	Diagnostic Imaging, Prevention & Control]	0.40070
26	17 or 18 or 19 or 20 or 21 or 22 or 23 or 24 or 25	348878
27	(UK or United kingdom or England or	1234304
	Wales or Scotland or Northern Ireland or	
	Ireland or Australia or Canada or New	
	Zealand or USA or United States or	
	Austria or Finland or Germany or Malta or	
	Netherlands or Norway or Spain or	
	Sweden or Switzerland).tw	2005
28	8 and 16 and 26 and 27	3985
29	(comment or editorial or letter).pt. or	2297837
20	(comment or editorial or letter).ti,ab.	2012
30	28 NOT 29	3912
31	(meta analysis or "review" or "systematic	4074667
	review").pt. or (meta analysis or "review"	
22	or "systematic review").ti,ab.	2252
32	30 NOT 31	3353
33	Limit 32 to (English Language and humans and yr= 22009-Current" and	2059
	English	

#### **EMBASE Search Strategy**

Conducted 21.08.2023

#	Search Query	Results
1	exp Prognosis/	914519
2	Incidence/	567003
3	Risk Assessment/	736472
4	Decision Support Techniques/	23770

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(prognos\* or prevention or progress\* or diagnos\* or detect\* or prevalence or

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m

	diagnos* or detect* or prevalence or	
	incidence or rate*).tw.	4000000
6	(risk* adj2 (assess* or factor*)).tw.	1282303
7	((first or initial) adj (episode* or detection)).tw.	33734
8	1 or 2 or 3 or 4 or 5 or 6 or 7	14825285
9	exp Vision Screening/	43893
10	exp Vision Tests/	43893
11	exp Mass Screening/	308708
12	((eye* or sight or vision or visual) adj1 (test* or exam* or screen* or follow- up)).ti,ab.	24543
13	(asymptomatic adj2 (test* or exam* or screen*)).tw.	6547
14	(routine adj2 (test* or exam* or screen*)).tw.	66221
15	((eye* or sight or vision or visual or screening) adj3 (frequenc* or interval* or recall*)).ti,ab.	11477
16	9 or 10 or 11 or 12 or 13 or 14 or 15	443705
17	Eye Diseases/	28376
18	Vision, Ocular/	97000
19	Visual Acuity/	135130
20	(ametrop* or emmetrop* or glaucoma or diabetic retinopathy or refractive error or macular degeneration or cateract* or presbyop* or amblyop* or myop* or hyperop* or hypermetrop* or astigmat* or anisometrop* or vision impairment* or vision loss* or vis* acuity).tw.	331096
21	Cataract/di, dg, pc [Diagnosis, Diagnostic Imaging, Prevention & Control]	4870
22	Glaucoma/di, dg, pc [Diagnosis, Diagnostic Imaging, Prevention & Control]	8600
23	Macular Degeneration/di, dg, pc [Diagnosis, Diagnostic Imaging, Prevention & Control]	619
24	Diabetic Retinopathy/di, dg, pc [Diagnosis, Diagnostic Imaging, Prevention & Control]	8050
25	Refractive Errors/di, dg, pc [Diagnosis, Diagnostic Imaging, Prevention & Control]	1373
26	17 or 18 or 19 or 20 or 21 or 22 or 23 or 24 or 25	486510
27	(UK or United kingdom or England or Wales or Scotland or Northern Ireland or Ireland or Australia or Canada or New Zealand or USA or United States or Austria or Finland or Germany or Malta or Netherlands or Norway or Spain or Sweden or Switzerland).tw	2164194
28	8 and 16 and 26 and 27	2477
29	(comment or editorial or letter).pt. or (comment or editorial or letter).ti,ab.	2215128
30	28 NOT 29	2433
31	(meta analysis or "review" or "systematic review").pt. or (meta analysis or "review"	4598180

32	30 NOT 31	2086
33	Limit 32 to (English Language and humans and yr= 22009-Current" and English	1571

#### **CINAHL Search Strategy**

Conducted 21.08.2023

#	Search Query	Results
1	(MM "Prognosis")	1921
2	(MM "Incidence")	1712
3	(MM "Risk Assessment")	63998
4	TX (prognos* or prevention or progress*	3490646
	or diagnos* or detect* or prevalence or	
	incidence or rate*)	
5	AB (risk* N2 (assess* or factor*))	239810
6	TX ((first or initial) N1 (episode* or	12704
	detection))	
7	S1 OR S2 OR S3 OR S4 OR S5 OR S6	3564033
8	(MM "Vision Screening")	825
9	(MH "Vision Tests+")	7553
10	TX ((eye* or sight or vision or visual) N1	19207
	(test* or exam* or screen* or follow-up))	
11	TX (asymptomatic N2 (test* or exam* or	3757
	screen*))	
12	TX (routine N2 (test* or exam* or	33621
	screen*))	
13	AB ((eye* or sight or vision or visual or	2663
	screening) N3 (frequenc* or interval* or	
	recall*))	
14	S8 OR S9 OR S10 OR S11 OR S12 OR	61314
	S13	
15	(MM "Eye Diseases")	6614
16	(MM "Vision, Subnormal")	1133
17	(MM "Visual Acuity")	3893
18	AB (ametrop* or emmetrop* or glaucoma	33125
	or diabetic retinopathy or refractive error	
	or macular degeneration or cateract* or	
	presbyop* or amblyop* or myop* or	
	hyperop* or hypermetrop* or astigmat* or	
	anisometrop* or vision impairment* or	
10	vision loss* or vis* acuity)	461
19 20	(MM "Cataract/DI/PC/PR") (MM "Glaucoma/DI/PC/PR")	461 1458
20	(MM "Macular Degeneration/DI/PC/PR")	1458
21	(MM "Diabetic Retinopathy/DI/PC/PR")	1433
22	(MM "Refractive Errors/DI/PC/PR")	290
23	S15 OR S16 OR S17 OR S18 OR S19	43169
27	OR S20 OR S21 OR S22 OR S23	50103
25	TX (UK or United kingdom or England or	7754118
20	Wales or Scotland or Northern Ireland or	110
	Ireland or Australia or Canada or New	
	Zealand or USA or United States or	
	Austria or Finland or Germany or Malta or	
	Netherlands or Norway or Spain or	
	Sweden or Switzerland)	
26	S7 AND S14 AND S24 AND S25	4259
27	PT ((comment or editorial or letter)) OR	759333
	AB (comment or editorial or letter))	
		1

28	26 NOT 27	4083
29	PT ((meta analysis or "review" or "systematic review")) OR TI ((meta analysis or "review" or "systematic review")) OR AB ((meta analysis or "review" or "systematic review"))	853902
30	28 NOT 29	3531
31	30 Limiters - Published Date: 20090101- 20231231	2926
32	31 Limited to English Language	2583

Database	Results
Medline (Ovid)	2059
EMBASE (Ovid)	1571
CINAHL (EBSCO)	2583
TOTAL	6213

### **APPENDIX 2: Grey Literature resources**

Websites	
National Eye Institute   National Eye Institute (nih.gov)	
American Academy of Ophthalmology: Protecting Sight. Empowering Lives - American Academy of	
Ophthalmology (aao.org)	
Help & Support For People Living With Glaucoma   Glaucoma UK	
RNIB   Home	
NICE   The National Institute for Health and Care Excellence	
Age-related macular degeneration - Macular Society	
AMDF - Saving Sight Through Research and Education (macular.org)	
Fight for Sight - Stopping sight loss through pioneering research	
Cataracts & Other Eye Conditions: What You Should Know (beyondcataracts.uk)	
Sightsavers   Protecting sight and fighting for disability rights	
Best Eye Charity in UK, Sight Loss Charity London   Mission 4 Vision NGO in UK	
General Optical Council (GOC)	
British and Irish Orthoptic Society	
Royal College of Opthamologists	
College of Optometrists	
European Council of Optometry and Optics (ECOO)	
Association of Optometrists	
International Agency for the Prevention of Blindness (IAPB)	
Association of British Dispensing Opticians (ABDO)	
Search terms	
Vision test, visual test, test frequency, test interval, prognosis, incidence, progression, vision screening	
Review, rapid review, systematic review, meta-analysis (for identification of secondary research only)	

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