

# BMJ Open Vision impairment and cognitive decline among older adults: a systematic review

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## ABSTRACT

**Objectives** There has been increasing epidemiological research examining the association between vision impairment (VI) and cognitive impairment and how poor vision may be a modifiable risk factor for cognitive decline. The objective of this systematic review is to synthesise the published literature on the association of VI with cognitive decline, cognitive impairment or dementia, to aid the development of interventions and guide public policies pertaining to the relationship between vision and cognition. **Methods** A literature search was performed with Embase, Medline and Cochrane library databases from inception to March 2020, and included abstracts and articles published in peer-reviewed journals in English. Our inclusion criteria included publications that contained subjective/objective measures of vision and cognition, or a diagnosis of VI, cognitive impairment or dementia. Longitudinal or cross-sectional studies with ≥100 participants aged >50 years were included. The search identified 11 805 articles whose abstracts underwent screening by three teams of study authors. Data abstraction and quality assessment using the Effective Public Health Practice Project Quality Assessment Tool were performed by one author (NN). 10% of the articles underwent abstraction and appraisal by a second author (LA/VV), results were compared between both and were in agreement. **Results** 110 full-text articles were selected for data extraction, of which 53 were cross-sectional, 43 longitudinal and 14 were case–control studies. The mean age of participants was 73.0 years (range 50–93.1). Ninety-one (83%) of these studies reported that VI was associated with cognitive impairment. **Conclusion** Our systematic review indicates that a majority of studies examining the vision–cognition relationship report that VI is associated with more cognitive decline, cognitive impairment or dementia among older adults. This synthesis supports the need for additional research to understand the mechanisms underlying the association between VI and cognitive impairment and to test interventions that mitigate the cognitive consequences of VI.

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## INTRODUCTION

Dementia is among the most pressing public health challenges of the 21st century.<sup>1</sup> In 2015, 46.8 million people were living with dementia, and the number is expected to

## Strengths and limitations of this study

- There was heterogeneity in the measurement of cognitive and visual function among all included studies.
- The quality assessment tool used for assessing quality of included studies penalised longitudinal studies that lost over 40% of participants due to drop-outs/withdrawals, which is common in studies that span over many years.
- Majority of the included studies were cross-sectional, and these are prone to selection bias.

double every 20 years.<sup>2</sup> Vision impairment (VI), another major global health problem, affects at least 2.2 billion people worldwide,<sup>3</sup> most of whom are aged 50 years and older.<sup>4</sup> Both cognitive and VI are projected to affect an increasing number of people over time, primarily due to population ageing.<sup>4,5</sup>

Prior work has suggested that cognition and vision are associated,<sup>6,7</sup> and while there are shared risk factors (neuropathological/vascular),<sup>8</sup> there is also longitudinal evidence that VI is associated with cognitive changes.<sup>9</sup> The mechanisms underlying the vision–cognition relationship have not yet been fully characterised, but it is hypothesised that sensory loss, such as hearing impairment and VI, may lead to increased cognitive load, structural and functional changes in the brain, and decreased emotional and social well-being, all of which could potentially increase the risk of cognitive impairment.<sup>9,10</sup> While the role of treating hearing loss in preventing cognitive impairment has been acknowledged, VI has not yet been recognised as a potentially modifiable risk factor for cognitive impairment.<sup>1,11</sup>

Since the majority of VI is due to correctable conditions, namely refractive error and cataract,<sup>12</sup> establishing the existence of an association between vision and cognitive impairment could present an additional opportunity to prevent cognitive impairment



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and dementia through interventions that optimise vision. In this systematic review of the literature, we examined the association between cognitive and vision impairment among older adults in existing observational studies. This qualitative review summarises the existing research examining the vision–cognition relationship, providing insight on data gaps and areas for continued investigation, as well as highlighting differences in methodological approaches that may impact the interpretation of results across studies.

## METHODS/LITERATURE SEARCH

We followed the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines (online esupplemental file 1).

Cross-sectional and longitudinal studies reporting a measure of association between visual function and cognitive impairment were included if they had  $\geq 100$  participants aged of  $\geq 50$  years (mean) at baseline. Reasons for exclusion of studies were: (1) Outcome measure was not vision or cognition, (2) Association between vision and cognition was not explored, (3) Sample size  $< 100$ , (4) Publication not in English, (5) Mean age  $< 50$  years, (6) No cognitive measure, (7) No vision measure and (8) Outcome was not part of inclusion criteria.

An academic librarian searched: Ovid Medline, Embase, Cochrane and PubMed from their inception to March 2020, and developed a search strategy that combined controlled vocabulary and keywords related to geriatrics, cognition and vision (online esupplemental file 2-complete Ovid Medline search strategy). Searches were limited to human studies published in English. Conference and poster abstracts, and short oral presentations were also included.

Search results were exported to Covidence (Veritas Health Innovation, Melbourne, Australia). Three teams of two reviewers each worked independently and in duplicate to screen titles, abstracts and full-text articles to determine inclusion (NN and MM; EC and YS; VV and LA). Disagreements were adjudicated by a member of the other study team.

Data were extracted from the included publications by one author (NN), and another (LA) extracted data from a random sample of 10% to compare results. Any discrepancies were adjudicated by a third author (VV). Data collected for each publication included: study design, participant characteristics, vision and cognition assessment methods and the summary measure that described the vision–cognition association.

The methodological quality of included studies was assessed by one author (NN) using the Effective Public Health Practice Project Quality Assessment Tool (EPHPP)<sup>13</sup> and the global quality ratings and findings were summarised qualitatively. A random 10% sample was reviewed by another author (VV) to ascertain consistency in quality assessment (QA).

**Table 1** Patient demographics and study characteristics

Demographics and study characteristics	No
Total no of studies	110
Total no of participants	9 799 329
Mean age, year, IQR	73.0 (50–93.06)
Study design	
Cross-sectional	53 (48%)
Longitudinal	43 (39%)
Case-control	14 (13%)
Country	
USA	30 (27%)
Japan	6 (5%)
UK	6 (5%)
China	6 (5%)
Australia	5 (4.5%)
France	5 (4.5%)
Germany	5 (4.5%)
Singapore	5 (4.5%)
Ireland	3 (3%)
Canada	3 (3%)
Spain	2 (2%)
Taiwan	2 (2%)
Sweden	1 (<1%)
Nordic Countries	1 (<1%)
New Zealand	1 (<1%)
Switzerland	1 (<1%)
Netherland	1 (<1%)
Others	27 (25%)

## RESULTS

### Study Selection

Online esupplemental file 3 is a PRISMA flow chart that describes the results of the search strategy of articles that examined the association between VI and cognitive impairment or decline. Of the 11 805 studies that were imported for screening, 110 articles were included in our final systematic review.<sup>7 14–122</sup>

### Description of included studies

Table 1 describes the characteristics of the studies included. The total number of participants in this review was 9 799 329 (range: 112–7 210 535 per study), with a mean age of 73.0 years, (range: 50.0–93.1). Of the total 110 studies included, 53 were cross-sectional, 43 were longitudinal and 14 had a case–control study design. The range of follow-up time for the longitudinal studies was 2 months to 10 years.

Of the 110 studies included, 51 reported findings from participants enrolled in population-based studies. There were five studies each from the following large population

**Table 2** Measures of vision and cognition assessed in studies

Outcome measures	No of studies
<b>Cognition (n, %)</b>	
Objective assessment n=89 (81%)	
MMSE	42 (47%)
MoCA	5 (6%)
Global cognition scale	6 (7%)
Others	36 (40%)
Other assessment methods n=13 (12%)	
Self-report	4 (31%)
ICD diagnosis/from records	9 (69%)
Combination of Objective+Subjective (5%)	7
No info (<1%)	1
<b>Vision (n, %)</b>	
Objective n=66 (70%)	
Acuity	42 (65%)
Acuity+others	16 (23%)
Others	8 (12%)
Other assessment methods n=34 (22%)	
Self-report	24 (71%)
ICD diagnosis/from records	10 (29%)
Combination of Objective+Subjective (7.5%)	8
No info (<1%)	2

MMSE, Mini-Mental State Examination; MoCA, Montreal-Cognitive Assessment.

based longitudinal studies: English Longitudinal Study of Aging (ELSA) and The Three-City Study. Three studies each from Fujiwara-Kyo Study, Salisbury Eye Evaluation Study, Irish Longitudinal Study on Aging and Singapore Epidemiology of Eye Diseases. Two studies each came from The Newcastle 85+ study, Study of Osteoporotic Fractures, Blue Mountain Eye Study, Australian Longitudinal Study on Aging, Health and Retirement Study, National Health and Aging Trends Study (NHATS), Leiden 85+ study, Health ABC study and the Singapore Malay Eye Study. Additionally, 10 studies used insurance claims data from different countries.

The studies in this review included participants from over 17 different countries (table 1), 30 studies (27%) from the USA, followed by 25 studies from Europe (23%) including the UK, Germany, Ireland, Finland, Switzerland, France and Netherlands. Ninety of the studies were published between 2009 and 2020. All papers provided a description of sampling methods. 16 studies were included which were either conference abstracts or short oral presentations.<sup>24 32 45 49–51 53 56 92 93 96 112–116</sup>

### Assessment of cognitive function

To assess cognitive function, 89 studies used objective assessments, 13 used other assessment methods such

as self-report and diagnosis codes, and 7 studies used a combination of both (table 2); one study did not provide information about cognitive function assessment.<sup>92</sup> Mini-Mental State Examination (MMSE) was the most commonly used objective method to assess cognitive function (42 studies). Other methods used to objectively measure cognition included: Montreal-Cognitive Assessment test,<sup>28 32 34 53 93</sup> Addenbrooke's Cognitive Examination-Revised,<sup>45 61 79</sup> Cognitive Performance Scale,<sup>42 49 65</sup> Blessed-Orientation-Memory-Concentration test,<sup>25 62</sup> Abbreviated Mental Test,<sup>72 74 89 100 110 119</sup> Blessed Dementia Scale,<sup>86</sup> Digit Symbol Substitution Test<sup>22 122</sup> and Cambridge Cognitive Examination test.<sup>88</sup> For the studies that used other assessment methods, four used self-reported cognitive measures,<sup>18 22 39 105</sup> and nine used diagnostic codes to define cognitive decline and/or dementia.<sup>19–21 51 69 99 103 115 117</sup> Among the studies that used a combination of objective and subjective methods, four used self-reported cognitive function along with an objective measure.<sup>22 30 48 55</sup>

### Assessment of visual function

In order to assess visual function, 66 studies used objective assessments, 34 used other assessment methods such as self-report and diagnosis codes, and 8 studies used a combination of both (table 2); no information was available from two studies.<sup>24 116</sup> Visual acuity (VA) was the most commonly measured visual function (42 studies), of which the Snellen acuity chart was the most commonly used method (18 studies). VA was also measured in combination with other visual functions, including: visual fields (VF) (six studies),<sup>20 51 70 90 102 118</sup> contrast sensitivity (CS) (eight studies),<sup>29 52 86 93 94 102 118 122</sup> macular pigment optical density (two studies)<sup>14 34</sup> and fundus photography (two studies).<sup>63 118</sup> Other methods used to objectively measure visual functions included: colour vision,<sup>17</sup> VF only,<sup>28</sup> CS only,<sup>35 76</sup> fundus photo with grading<sup>38 100</sup> and autorefraction.<sup>74</sup> Other assessment methods included: self-reported vision (24 studies) and diagnostic codes or patient records to define VI (10 studies).<sup>19 21 46 69 83 99 104 115 117 121</sup> The studies that used a combination of methods, eight studies used self-report along with an objective measure of visual function.<sup>22 40 53 59 68 71 107 113</sup>

### Quality of studies

The methodological quality of included studies was assessed using the EPHPP.<sup>13</sup> The tool assessed each study on five domains: (1) Selection bias, (2) Study design, (3) Confounders, (4) Data collection methods and (5) Analysis. For each included study the five relevant domains were ranked on a three-point Likert scale with three representing a low risk of bias ('strong'), two a possible risk of bias ('moderate') and one a high risk of bias ('weak'). An overall rating was derived following the EPHPP methodology. A study consisting of at least one 'weak' rating in a domain received an overall rating of 'moderate,' while those with two or more domains with 'weak' ratings were automatically classified as 'weak' overall. We present

Table 3 Studies with a 'strong' rating

Author, title and year	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	CI: type of measurement/ evaluation	VI: Type of measurement/ evaluation	Point estimates and analysis performed	Summary of association	Quality of study
Dearborn <i>et al</i> <sup>27</sup>	US (Maine-Syracuse Longitudinal Study (MSLS))	Longitudinal cohort, wave 6&7	655	Normal VA 60.34	40%/60%	Cognition	Visual-Spatial Organisation and Memory, Scanning and Tracking, Verbal Episodic and Working memory	Acuity: Snellen eye test (log transformed)	Cross-sectional (CSA) and prospective analysis (PA) using multiple linear and multiple logistic regression	Poorer VA associated with lower cognitive function and 5 year decline	Strong
2018	compared sixth (2001–2006) wave and			VI 70.6	40%/60%		Global composite score-loaded equally across domains, z-transformed		Global OR: CSA 1.512 (0.71 to 3.23), PA 1.539 (0.74 to 3.21), VSMO OR: CSA 1.28 (0.63 to 2.59), PA 2.26 (1.16 to 4.4)	in cognitive function over a range of domains incl the global measurements	
	seventh (2006–2010) wave								Working OR: CSA 1.73 (0.85 to 3.51), PA 1.55 (0.78 to 3.08), VEM OR: CSA 0.55 (0.25 to 1.24), PA 1.43 (0.73 to 2.81)	Visual-spatial organisation, verbal episodic memory	
Dimiz-Filho <i>et al</i> <sup>28</sup>	USA	Longitudinal cohort	115	67.4		Cognitive Decline	Montreal-Cognitive Assessment test (MoCA); assesses atten and concentration, executive functions	Standard Automated Perimetry (SAP) using 24–2 SITA	Univariate: 5-point decline in MoCA=0.18 dB increase of residuals of SAP MD, R <sup>2</sup> =4.3% (0.06 to 0.30) p<0.003	Statistically significant association between change in	Strong
2017							memory, language, visuoconstructional skills, conceptual thinking, calculations and orientation	<33% fixation loss and <15% FP only included	Multivariate: 5-point decline in MoCA=0.23 dB inc of residuals of SAP MD, (0.11 to 0.35) p<0.001	MoCA scores and VF variability over time	
Fischer <i>et al</i> <sup>35</sup>	USA (The Epidemiology of Hearing Loss Study)	Longitudinal cohort. Baseline info was from EHLS-2 (1998–2000), CI data from EHLS-3 (2003–05) and EHLS-4 (2009–10)	1884	66.7	40.9/49.1	Cognition	MMSE	Contrast sensitivity using Pelli-Robson letter charts. VI was defined as	Cox discrete time proportional hazard analyses performed to model relationship btwn CI and Sensory Imp.	Hearing, Visual and Olfactory impairment were independently	Strong
2016							score<1.55 log units in the better eye		Vision: HR=2.05 (1.24, 3.38); Olfaction: HR=3.92 (2.45, 6.26), Hearing: HR=1.90 (1.11, 3.26)	associated with cognitive impairment risk	

Continued

**Table 3** Continued

Author, title and year	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	CI: type of measurement/ evaluation	VI: Type of measurement/ evaluation	Point estimates and analysis performed	Summary of association	Quality of study
Hall <sup>43</sup>	USA (Impact of Cataract on Mobility (ICOM) study)	Longitudinal cohort study. 3 groups identified: No cataract, with cataract and undergoing surgery, with cataract and declined surgery	No Cat: 92	66.8	48.9/51.1	Cognition	Mattis Organic Mental Syndrome Screening Examination (MOM SSE)-evaluates 14 domains (abstraction, orientation, memory, speech, comprehension). Scored 0 to 28 (lower score higher functioning)	Acuity: Dist VA ETD RS chart, Log Contrast sensitivity: Pelli-Robson chart	Cat+no Sx & cat+surgery grps significantly less CI (p<0.001 and p=0.009, respectively) than at baseline.	Cataract surgery does not affect cognitive function	Strong
2005			W Cat+Sx: 122	70.9	41.8/58.2			Ophthalmologist/ optometrist graded cataract at clinic visit	No cataract group=no change in Cog status, association between change in VA in the better and worse eye and		
			W cat+no Sx: 87	71.1	59.8/40.2				change in MOM SSE observed (p=0.003 & p=0.03, respectively)		
Lin <i>et al</i> <sup>47</sup>	USA (Study of Osteoporotic Fractures)	Longitudinal cohort	Vision testing sample:1668	75.9	0/100	Cognition	3MS-Modified version of MMSE	Acuity: Bailey Lovie Target	VI (worse than 20/40): Cog decline - 1.78 (1.21 to 2.61), greater odds of Functional decline- 1.79 (1.15 to 2.79)	Vision impairment is associated with greater odds of cognitive and functional decline	Strong
2004			Overall 6112	76.1					DSI: Cog decline - 2.19 (1.26 to 3.81), Functional decline- 1.87 (1.01 to 3.47)	functional decline over time in older women	
Jefferis <i>et al</i> <sup>46</sup>	UK	Longitudinal cohort.	112	Normal cog: 80 (3.8)	20/80	Vision	Addenbrooke's Cognitive Examination (which includes the MMSE)	Acuity: LogMAR, Other: VFQ-25	Normal cog group: Mean at baseline 0.13 (0.09), 1 year FU 0 (0.09), 95% CI for difference: 0.09 to 0.16<0.001	Patients with impaired cognition benefit from cataract surgery, but not to the same extent as patients with normal cognition. Cognitive impairment may, however, limit visual improvements	Strong
2015				Impaired cog: 81.2 (3.9)	30/70					Impaired cog group: following cataract surgery	
										Mean at baseline 0.18 (0.14), 1 year FU 0.06 (0.11), 95% CI for difference: 0.08 to 0.16	

Continued



**Table 3** Continued

Author, title and year	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ci: type of measurement/ evaluation	Vi: Type of measurement/ evaluation	Point estimates and analysis performed	Summary of association	Quality of study
Reyes-Ortiz <i>et al</i> <sup>67</sup>	USA (Hispanic Established Populations for Epidemiologic Studies of the Elderly)	Longitudinal cohort.	2140	Near Vi: 73.5 (M/F)	39.3/60.7 (M/F)	Cognition	MMSE-blind	Acuity: subjects hold cards at least seven inches from their eyes and asking them to read the numbers	Near vision impairment; -0.62±0.29, .03, Interaction btwn near Vi & time=-0.13 +/- 0.07, .045	Results showed that near vision impairment was predictive of	Strong
2005				Distance Vi: 75.3	35/65			Each card had 7-digit "telephone numbers" of three different type sizes: 7, 10, and 23 points	Distance vision impairment=-0.06 +/- 0.36, .87, Interaction btwn distance Vi & time, -0.12±0.08, .14	cog decline in older Mexican Americans independent of other health factors	
Haan <i>et al</i> <sup>67</sup>	USA (Women's Health Initiative MS-MRI)	Longitudinal cohort	511	69	0/100	Cognition	Modified Mini Mental State (3 MS)	Acuity: Snellen with pinhole	Acuity slope in model of 3MSE score (β slope // 0.33, p=0.06). Covariate-adjusted mean 3MSE scores for women	Retinopathy as a marker of small vessel disease is a risk factor for cerebrovascular disease that may influence cognitive performance and related brain changes	Strong
2012									with retinopathy compared with others throughout follow-up: mean (SE) difference 1.01 (0.43), p=0.019	Very mild association between acuity and 3MSE scores	
Trick <i>et al</i> <sup>70</sup>	USA	Case-control. Visually and cognitively normal controls (n=61), and senile dementia of Alzheimer's type (SDAT) n=61	122	SDAT: 73.4	NA	Cognition	each patient was assigned a clinical dementia rating (CDR) based on a scale developed and tested at Washington Uni	Acuity: ETDRS. VF analysis using 30-2 HFA	Patient groups, Foveal threshold, Mean deviation, Pattern SD, Corrected pattern SD.	We found that the frequency of potentially unreliable VF was significantly higher (chisquare,	Strong
1995				Control: 72.8			i.e CDR scale ranges from 0.0 (individual without dementia) to 3.0 (severely demented).		Control (n=44): 33.2±3.05, -2.81+/-3.45, 3.48±2.05, 2.58±2.30 SDAT (n=34): 30.79±4.00 (p=0.0027), -5.11+/-3.17(p=0.0057), 4.461+/-2.49(NS), 3.91+/-2.73(p=0.0408) CDR=0.5(n=25): 31.12+/-3.54, -4.25+/-2.23, 4.47+/-2.60, 3.83+/-2.89	p=0.0232) in SDAT patients (44.3%) than in controls (27.9%)	

Continued

**Table 3** Continued

Author, title and year	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	CI: type of measurement/ evaluation	VI: Type of measurement/ evaluation	Point estimates and analysis performed	Summary of association	Quality of study
Helmer <i>et al</i> <sup>60</sup>	France (The Three-City–Alienor Cohort)	Longitudinal, cohort	812	79.7	35/65	Cognition	MMSE; Free and Cued Selective Reminding Test Isaacs Set Test the Trail Making Test parts A and B Then suspected cases were assessed by a neurologist	Acuity: No chart name provided, IOP: pneumotonometer	Association of OAG and incident dementia: Model 1: OR 4.3 95% CI 1.7 to 10.8 p 0.0020, Model 2: OR 4.2 95% CI 1.6 to 10.9 p 0.0030	Yes. Results show that OAG participants were at an increased risk for developing dementia during the 3 year follow-up time period.	Strong
2013									Model 3: OR 3.9 95% CI 1.5 to 10.4 p 0.0054, Association of Vertical cup:disk ratio and incident dementia: $\geq 0.65$ : OR 3.7 95% CI 1.4 to 9.7 p 0.0083 $\geq 0.70$ : OR 4.4 95% CI 1.5 to 12.5 p 0.0055 Association of Minimal rim:disk ratio = < 0.1 and incident dementia: OR 2.7 95% CI 1.005–7.1 p 0.0489		
de la Fuente <i>et al</i> <sup>67</sup>	UK (English Longitudinal Study of Ageing)	Longitudinal, cohort	3508	69	43.2/56.8	Cognition	Four measured tests of verbal fluency, processing speed, and short-term and long-term memory	Three self-reported items covering eyesight in far, near, and general vision	Visual ( $\beta=0.140$ , $p<0.001$ ) difficulties predicted cognitive difficulties 8 years later	Yes. Visual difficulties were identified as predictors of later	Strong
2018								Dichotomized, collapsing "Excellent," "Very good," and "Good" as "Absence of difficulties".	The latent increase in cognitive difficulties was steeper in people with visual impairment ( $d=0.52$ , $p<0.001$ )	subsequent cognitive decline in the old age	
Hamalainen <i>et al</i> <sup>68</sup>	Canada (Canadian Longitudinal Study on Ageing)	Cross-sectional cohort	30 029	No info	No info	Cognition	five cognitive tests: Mental Alternation Test, Animal Fluency test, Controlled Oral Word Association Test, Stroop test and Rey Auditory Verbal Learning Test with immediate and 5 min recall	Vision was measured as the better-seeing eye pinhole-corrected VA (reported in logMAR)	VA was a predictor of only executive function ( $b=-0.785$ , $p<0.001$ ) but not memory	Yes, greater executive function scores were associated with better vision	Strong
2019											
Chuanying Huang	UK (English Longitudinal Study of Ageing)	Cross-sectional cohort	4197	Optimal:70.21	58.6/41.4	Cognition	Memory (immediate and delayed), verbal fluency, and time orientation	Self-report: 'Is your eyesight (using glasses or corrective lens as usual) excellent, very good, good, fair, poor, or registered blind?'	poor vision (= $-1.309$ , $p=0.004$ ), and poor dual sensory function (= $-2.442$ , $p=0.013$ ) was associated with worse cognition	Yes.	Strong
2019				Good: 71.84	51.2/48.8		Global cognition was the sum of scores for the three domains, and higher score indicates better cognitive performance	Answer of 'excellent' or 'very good' was defined as optimal vision. Answer of 'fair' or 'poor' was			

Continued

**Table 3** Continued

Author, title and year	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	CI: type of measurement/ evaluation	Vi: Type of measurement/ evaluation	Point estimates and analysis performed	Summary of association	Quality of study
Virginie Naël	France (The Three-City-Alienor Cohort)	Longitudinal, cohort	7736	Poor: 74.75 Incident dem: 76.9	47.2/52.8 34.8/65.2	Cognition	3-step procedure: 1. MMSE, the Isaacs set test and the Benton Visual Retention Test	Near VA assessed using the Parinaud scale. Mild near VI was classified by Parinaud three or 4 (Snellen equivalent 20/30–20/60)	defined as poor vision. Thus vision was classified as optimal, good, and poor vision Moderate to severe near VI was associated with an increased risk of dementia in the first 2 years (HR2.0, 95% CI 1.2 to 3.3)	Yes. Near VI may represent an indicator of dementia risk at	Strong
2019				No inci dem: 73.6	39.2/60.8		2. senior neurologist to establish a clinical diagnosis, 3. an independent committee of	and moderate to severe near VI by Parinaud>4 (Snellen equivalent<20/60)	and from 2 to 4 years (HR 1.8, 95% CI 1.1 to 3.1). distance VF loss was associated with an increased risk beyond 4 years (HR 1.5, 95% CI 1.1 to 2.0)	short and middle-term, mostly in depressed elderly people	
Zhi Wei Lim	Singapore (Singapore Epidemiology of Eye Diseases)	Longitudinal cohort	2478	67.6	50.7/49.3	Cognition	neurologists and geriatricians reviewed all potential cases of dementia obtain a consensus on the diagnosis and aetiology, according to the DSM-IV and the NINCDS-ADRDA criteria	Dist VA was selfreported, defined as an inability or difficulty in recognising a familiar face at 4 m	but the association was no longer significant after taking into account baseline cognitive performances	Yes, poor vision was independently associated with a decline in cognitive function	Strong
2020							A locally validated Abbreviated Mental Test (AMT)	VA was measured at 4 m using the logMAR number chart	Baseline VI was associated with a decrease in AMT score over 6 years ( $\beta = -0.27$ ; 95% CI, $-0.37$ to $-0.17$ ; $p < 0.001$ )		
Bernard Michalowsky	Germany (NA)	Case-control study	122 708	81	39/61	Cognition	ICD-10 diagnosis code for dementia	(Lighthouse International).VI was defined as presenting VA worse than 20/40.	When change in vision over 6 years was evaluated, unchanged or deteriorated VI was associated with a decrease in AMT score over 6 years ( $\beta = -0.29$ ; 95% CI, $-0.40$ to $-0.18$ ; $P < .001$ )	No, but DSI was not significantly associated with dementia (OR=0.97, risk of dementia CI=95% 0.97 to 1.02, p=0.219)	Strong

Continued



Table 3 Continued

Author, title and year	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ci: type of measurement/ evaluation	Vi: Type of measurement/ evaluation	Point estimates and analysis performed	Summary of association	Quality of study
2019									Combination of both visual and hearing impairments and the risk of dementia (OR=1.14, CI=95% 1.04 to 1.24, p=0.005)		
Jung-Yu Liao <sup>121</sup>	Taiwan (National Health Insurance Research database)	Case-control study	9200	NA	LOAD 36.7/63.3	Cognition	LOAD cases diagnosed as dementia and prescribed any acetylcholinesterase inhibitors (ACHEIs)	ICD-9 codes	Disorders of refraction and accommodation with LOAD OR for Year 1,2,3,4: 1.2, Year 6:1.3	Yes, in the path analysis model, disorders of ref and acc	Strong
2020	1997–2013					Without LOAD 36.7/63.3			Total effect reflects an association between prior diseases and LOAD incidence via all paths in the model: 0.042	had a significant positive effect on LOAD incidence	

DSI, dual sensory impairment; MMSE, Mini-Mental State Examination; VA, visual acuity; VF, visual field.

our studies in three different tables which is categorised based on the overall ratings, with ‘strong’, ‘moderate’ and ‘weak’ studies in [tables 3–5](#), respectively. In our sample, 17 studies received a rating of strong, 70 moderate and 23 weak.

### Study findings

Of the 110 studies included ([tables 3–5](#)), 91 found a significant positive association between VI and cognitive decline, cognitive impairment or dementia, and 13 studies found no significant association.<sup>26 30 43 44 60 61 68 81 82 90 94 115 117</sup> There were six studies that were inconclusive.<sup>23 37 67 70 79 83</sup> Of the 91 studies that found a significant association, 77 used objective methods to assess their vision or cognitive outcome. Of the 43 longitudinal studies, 35 found a significant association between VI and cognitive decline, cognitive impairment or dementia. The most commonly presented statistical measures were ORs and HRs. The random 10% of the study sample that was separately extracted by an independent author (LA) was found to be similar to elements from the primary extraction.

### DISCUSSION

In this systematic review, we evaluated and synthesised the literature examining the association between VI and cognitive function among older adults, and found strong agreement that VI is associated with cognitive impairment, cognitive decline or dementia. Results from the longitudinal studies that found a positive association between vision and cognition supports our hypothesis that VI may be a risk factor for cognitive impairment, cognitive decline or dementia.

Ninety-one studies reported associations between decline in visual and cognitive functions. Garin *et al*,<sup>40</sup> who received a ‘moderate’ rating in the QA, performed a cross-sectional analysis in a representative sample of Spanish population and measured cognition objectively. They also measured distance and near vision and found that objective and subjectively measured poor distance and near VA were associated with worse cognitive functioning. Lin *et al*<sup>47</sup> used data from a large longitudinal cohort study of older women and found that VI was associated with greater odds of cognitive and functional decline over 2 years. This study used objective measures of assessment for both vision and cognition and received a ‘strong’ rating in the bias assessment. Luo *et al*,<sup>48</sup> who received a ‘moderate’ rating in QA, performed a cross-sectional analysis on a large population sample from China. They reported that those with VI and Dual Sensory Impairment (DSI) were more likely to have severe to extremely severe dementia compared with those without any sensory impairment. Another longitudinal study that received a ‘moderate’ rating in QA from Germany by Hajek *et al*<sup>58</sup> with a large sample size (n=2394) showed that the onset of severe VI was associated with a decline in cognitive function scores. Uhlmann *et al*<sup>64</sup> in their paired case–control study between VI and dementia patients

**Table 4** Studies with a 'moderate' rating

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ci: type of measurement/evaluation	Vi: Type of measurement/evaluation	Point estimates and analysis performed	Summary of association	Quality of study
Ajana <i>et al</i> <sup>14</sup>	France (ALIENOR study, 3 City-Bordeaux cohort) France	Cross-sectional study	184	82.3	31.5%/68.5%	Cognition	MMSE, Isaacs Set test (IST15), Benton Visual Retention test, Free and Cues Selective Reminding Test	Acuity: Measured but no mention of method	MPOD 0.5 degree, Z score (b)=-0.12 (0.01, 0.23), MPOD 1 degree, Z score (b)=-0.11 (0.01, 0.22)	Yes- Between MPOD and Global cognitive z score	Moderate
2018							Summarised by a composite global cognitive Z-score	MPOD: Measured using confocal scanning laser Ophthalmoscope	Regression models		
Anstey <i>et al</i> <sup>15</sup>	Australia (Australian Longitudinal Study of Ageing)	Longitudinal	1823	77.77	51.2%/49.8%	Cognition and vision	MMSE, Verbal ability, Processing speed and Memory	Acuity: Distance using Snellen chart at 3 m, Near at 20 cm	N/A	Yes- significant moderate-sized association between rates of	Moderate
2003						Analysis of change using growth curves		using chart containing short passages in font size 5 to 18. Left and Right eye were tested separately. Score was the smallest font size	Latent growth curve analysis	change in Memory and Vision	
Anstey <i>et al</i> <sup>16</sup>	Australia (Australian Longitudinal Study of Ageing)	Longitudinal	3766	76.34	Wave 1: 56%/44%	Cognition	Verbal ability, Processing speed and Memory	Acuity: Distance using Snellen chart at 3 m	Sensory status x group interaction for memory comparing participants who declined on Vision vs not declined	Yes- Decline in visual performance had a significant effect on decline in memory performance	Moderate
2001					Wave 3: 33%/67%			Wilks A=0.98, F=2.94, p<0.05, n2=0.054			
Amaoutoglou <i>et al</i> <sup>17</sup>	Greece	Longitudinal, Case-control	Total 103 AD group n=32	AD 73.3	38%/62%	Vision	MMSE, Dementia blood screening, Reisberg AD scale, MRI and NINDS-AIREN to diagnose VaD	Colour vision using Ishihara colour deficiency test	Difference in ischi-hara scores between two groups F(1, 161)=9.558, p=0.003, AUC 0.819 (0.70, 0.93)	Yes- Ischi-hara a promising test to differentiate AD and	Moderate
2017			VaD group n=36	VaD 75.6			Consists of 38 pseudoisochromatic plates		ANOVA to estimate any sig variations between three groups within their age, MMSE, Resiberg and total Ischi-hara score	VaD	
			Healthy controls n=35	Controls 74.2			six errors suggests definite colour blindness				

Continued

**Table 4** Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ct: type of measurement/ evaluation	Vi: Type of measurement/ evaluation	Point estimates and analysis performed	Summary of association	Quality of study
Arrighi <i>et al</i> <sup>18</sup>	US (National Health interview survey (NHIS)2001–05)	Cross-sectional study using nationally available survey data	23 474	With dementia 80.72	With dementia 36%/64%	Vision	Self reported surveys. Was categorised as 'With dementia if patient reported 'senile''	Self reported vision problems	Prevalence OR 1.46 vision problems with functional limitations among americans aged>60	Yes- Higher odds of vision problems being prevalent	Moderate
			Person with dementia 443	Without dementia 72.3	Without 39%/61%			with and without limitations related to dementia		among people with functional limitation	
Bauer <i>et al</i> <sup>19</sup>	German statutory health insurance fund data (2006)	Cross-sectional study using claims data from insurance company	37 753	80.1	28.4%/71.6%	Comorbidity complex	ICD codes with diagnoses of dementia	ICD codes with diagnoses of severe vision reduction	OR 0.59 persons with severe vision problems were less likely to have dementia	Yes- significantly lesser odds of vision problems among	Moderate
			Without dementia 23 031								
2014	Dementia group 9139	Patient having a diagnosis of vision prob	81.6	25.4%/74.6%						cases with dementia	
			Without dementia 28 614	79.6	29.4%/70.6%	Using ICD codes					
Chen <i>et al</i> <sup>21</sup>	Taiwan National Health Insurance	Case-control. Randomised sample data of one million patients who made claims from insurance	Eye disease 4097	50 y	AMD 60.9%/39.1%	Alzheimer's disease	ICD codes with diagnoses of Alzheimer's Disease	ICD codes with diagnoses of AMD, Diabetic retinopathy and Glaucoma	Cumulative IR of 1.22% ADE among eye disease group vs 0.04% among controls	AMD, DR and Glaucoma were asso with an increased	Moderate
			Controls 20 745		DR 52.9%/47.1%					HR for DR 39.31 (4.79, 332.67), HR for AMD 36.94 (4.62, 295.46), HR for Glau 34.08 (13.37, 86.84)	risk of Alzheimer's disease
2016	Research database (HIRD)		20 745		Glaucoma 45.9%/54.1%						
			NHAMES 2975	72	48%/52%	Cognition	NHAMES: Cog testing done using Digit symbol substitution test (DSST). </=28 indicated poor cognition	NHAMES: Dist-Snellen VA chart, Near- 5-linr near vision card	NHAMES:Dist VI b=-5.1 (-8.6--1.6), OR 2.8 (1.1, 6.7), Near VI b=-3.8*, OR 1.7*, Subj VI b=-5.3*, fn OR 2.7*	Yes, VI significantly associated with worse cognitive	Moderate
Chen <i>et al</i> <sup>22</sup>	US (NHAMES and NHATS)	Cross-sectional analysis of two datasets: 1999–2002 NHAMES data and 2011–2015 NHATS data									

Continued

Table 4 Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ct: type of measurement/evaluation	Vi: Type of measurement/evaluation	Point estimates and analysis performed	Summary of association	Quality of study
2017			NHATS 30202	NA	42%/58%		NHATS: Probable/possible dementia vs No dementia based on NHATS classification scheme (self-report)	NHATS: Vision tested via questionnaires	NHATS: Subj Near Vi: OR 2.6 (2.2–3.1), Subj Dist Vi: OR 3.9 (3.4–4.4)		
Chiqui <sup>23</sup>	Canada	Longitudinal cohort	150	86.7	27%/73%	Vision	MMSE	Dist vision- Shellen chart, 6 m. Vi was Vn<6/12	Presenting Vi: 37.3 (29.1–46.1)	37.3% of dementia participants had Vi.	Moderate
2017								Near vision- Topcon paragon/lighthouse	Vi after refraction: 20.1 (13.7–27.9)	Just prevalence study, no correlation tested	
De Celis <i>et al</i> <sup>25</sup>	US (Hurria <i>et al</i> JCO 2011 & 2016)	Cross-sectional analysis of 2 prospective studies	750	Median age 72 (65–94)	No info	Cognition	Blessed OMC test (>11 on this test)	self-reported visual impairment (fair, poor or blind)	OR for association between Vi and Cognition deficit.	Older pts with cancer and hearing/visual impairment	Moderate
2017									Univariate model: 2.0 (0.8–4.7), Multivariate model: 1.9 (0.75–5)	at higher risk of functional and cognitive deficits	
de Kok <i>et al</i> <sup>26</sup>	New Zealand (Life and Living in Advanced Age: A Cohort Study in New Zealand (LILACS NZ))	Cross-sectional baseline data	661	Maori: 82.3 Non-Maori: 84.6	40%/60%	Cognitive decline	MMSE-blind version, cognitive test for the visually impaired	Acuity: ETDRS	Generalised linear models and structural equation modelling	No direct association found in this population	Moderate
2017									Maori: Dis VA (logmar) b=0.103 (-0.21,0.42), Self reported Vi b=0.110 (-0.030, 0.251) Non-Maori: Dis VA (logmar) b=0.197 (-0.12, 0.51), Self-report Vi b=0.065 (-0.056, 0.187)		
Elliott <i>et al</i> <sup>29</sup>	US	Cross-sectional cohort	238	No info	26%/74%	Vision	MMSE	Dist VA: ETDRS. Near VA: Lighthouse near VA card. Vi<20/40 one eye	β MMSE in Multiple linear Regression for Predicting Vn: Dist VA: -0.015, Near VA: -0.013 both (p 0.00)	Cognitive status contributed to the prediction of Vi	Moderate
2015								Mars CS to assess CS. Score worse than 1.50 in one eye	Contrast sen: 0.17 (p0.00)		
Elliott <i>et al</i> <sup>30</sup>	US	Longitudinal case-control	78	REC: 79.2	23.1%/76.9%	Cognition	Self-report and MMSE	Distance: ETDRS	Intervention group: MMSE baseline 20.2 (4.4), follow-up 19.4 (4.8), p=0.05	Vision-enhancing interventions did not lead to short	Moderate

Continued

**Table 4** Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ct: type of measurement/evaluation	Vi: Type of measurement/evaluation	Point estimates and analysis performed	Summary of association	Quality of study
2009			64	Delayed corrc: 78	25%/75%			Near: Lighthouse Near VA Test	Control group: MMSE baseline 21.7 (4.5), follow-up 20.5 (5.5), p=0.015	term improvements in physical function/cognitive status in sample.	
			30	Cataract Sx: 81	26.7%/73.3%				Pre&post-intervention measures of Phy in &cognitive status compared within treatmnt groups paired t-tests		
			15	No cat sx: 87	13.3%/86.7%						
Elyashiv <i>et al</i> <sup>31</sup>	US (The Religious Orders Study and The Memory snd Ageing Project)	Longitudinal cohort study of ageing and AD	2716	78.13	28.6%/71.4%	Cognition	Global cognitive scale (GCS), summary of raw scored of 19 tests incl episodic, semantic and working mem	Tested with both eyes open with a card held at 14 inches and measured	Mixed model for assoc with global cog score: VA b=-0.03312 (p<0.0001)	Significant positive correlation between GCS and VA	Moderate
2014							Scores converted, avergd and normalised- categorised into no,mild,mild Cl+otherdiag, AD and AD+diag	at 7 increments from 20/40 or better to 20/400 or worse	Pearson and spearman correlation coeff to assess relationship btwn GCS and VA		
Elliott <i>et al</i> <sup>30</sup>	US	Cross-sectional	382	Range: 60 to<=100	No Impairmnt 26.8/73.2	Vision	MMSE	Acuity- Lighthouse near VA test	Multiple linear regression used to estimate assoc btwn vision and cognition	Vi affected HFQoL both with and without CI	Moderate
2009									NHVoQL, mean: No Vi+CI=72.8, Vi only=75.2, CI only=76.5, Vi+CI=65.2, p=0.02		
Feeney <i>et al</i> <sup>32</sup>	Ireland (The Irish Longitudinal Study on Ageing)	Cross-sectional cohort	4453	fourth quartile 62.8	42.5/57.5	Cognition	MMSE, MoCA, Neuropsych battery, CAPI for word-recall, CAMDEX-R for picture memory	Acuity:ETDRS, MPOD measured using Macular metrics densitometer	Multivariate regression to analyse assoc btwn Cog test performance and MPOD(continuous variable)	Lower MPOD associated with poorer performance in MMSE & MoCA	Moderate
2013				third quartile 62.4	46.3/53.7			Pt was asked if diagnosed with ARMD and Diabetes	MMSE: b=0.48 (0.06, 0.90) p=0.026, MoCA: b=0.83 (0.15, 1.50) p=0.016, Memory: OR=2.24 (1.2, 4.15)p0.01		
				second quartile 62.2	53.9/46.1						

Continued



**Table 4** Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Cl: type of measurement/ evaluation	Vi: Type of measurement/ evaluation	Point estimates and analysis performed	Summary of association	Quality of study
				first quartile 62.3	53.8/46.2						
Formiga et al <sup>36</sup>	Spain (NonaSantfeliu population-based study)	Cross-sectional design	186	93.06	23.5/76.5	Cognition	MEC (Spanish version of the MMSE), Functional status: Lawton-Brody Index (LI) and Barthel Index (BI)	Acuity: Snellen Eye test	T-test and chi-square test. MEC ( $\pm$ SD): group with visual deficit: $16.5 \pm 11$ , w/o visual deficit: $23.6 \pm 11$ , $p < 0.001$	Functional status and cognition significantly asso with sensory	Moderate
2006									Lower LI was asso with: VI OR=1.86 (1.44, 2.39) and combined impairment OR=1.995 (1.32, 3.01)	Impairment	
Friedman et al <sup>37</sup>	US (The Salisbury Eye Evaluation in Nursing Home	Cross-sectional	656	83.4	25/75	Vision	MMSE	ETDRS charts or Lea symbols and grating acuity tests using Teller cards	Bland-Altman plot to measure how strongly grating acuity and recognition acuity relate to one another.	Grating acuity testing appears to be useful in	Moderate
2002	Groups (SEEING)								MMSE (5-point decline)with Teller acuity cards: OR=2.67 (2.14, 3.31). Teller and recognition ICC=0.79	cognitively impaired individuals	
Frost et al <sup>38</sup>	Australia (Australian Imaging Biomarkers and Lifestyle study of ageing (AIBL))	Cross-sectional case control	123	AD 70.2	59.1/40.9	Vision (ARMD)	MMSE and PET scan	Digital, non-stereoscopic retinal colour photographs, both eyes with a	Logistic regression: Medium Drusen:(N (%)): CN: 13 (13), AD: 9 (41), OR: 4.69 (1.67–13.13)	A logistic model for early AMD found a	Moderate
2016				CN 71.3	39.6/60.4			Canon CR-1 Non-Mydiatic retinal camera with digital Canon camera at the back	Early AMD:(N (%)): CN: 3 (3.0), AD: 8 (36), OR: 18.67 (4.42–78.80)	significant association with AD diagnosis ( $p < 0.0001$ )	
Fuller et al <sup>39</sup>	US (The American Community Survey (ACS))	Cross-sectional survey data	7 210 535	61.31	47.1/52.9	Cognition	Self-report: Because of a physical, mental or emotional condition, does this person have difficulty	Grader (masked) evaluated images for signs of ARMD	Cog Imp: NSI reference group, SHI: 45–64y: 3.55–3.67, 65–79y: 2.86–2.95, 80y+: 1.81–1.86, SVI alone: 45–64y:	Cognitive difficulty greatly associated with sensory impairment when	Moderate

Continued

**Table 4** Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ct: type of measurement/ evaluation	Vi: Type of measurement/ evaluation	Point estimates and analysis performed	Summary of association	Quality of study
2018							concentrating, remembering or making decisions? Yes/No	he/she have serious difficulty seeing even when wearing glasses?yes/no	4.42–4.57, 65–79y: 4.71–4.91, 80y+: 2.39–2.44, DSI: 45–64y: 12.03–13.77, 65–79y: 11.19–12.84, 80y+:4.69–5.35	compared with NSI, esp greatest with SVI alone.	
Garin <i>et al</i> <sup>40</sup>	Spain (COURAGE)	Cross-sectional cohort	50–64: 1760	57	47.7/52.3	Vision	Global standardised score from five performance tests: learning and short-term mem, attention & working	Obj: Snellen, Subjective: two questions to the participant, 1. how much	Asso btwn poor VA and Cognition (OR): Dist VA: 1.47 (1.17–1.86), Near VA: 1.72 (1.46–2.02)	Objective and subjective poor distance and near VA	Moderate
2014							mem and language. Lower score indicates worse cognitive functioning	difficulty seeing object or recognising at 20 m 2. at arm's length	Subjective distance VA: 2.05 (1.44–2.90), Subjective near VA: 2.96 (1.92–4.55)	were associated with worse cognitive functioning.	
Gaynes <i>et al</i> <sup>41</sup>	US (Rush memory and ageing project (1997–2007))	Cross-sectional cohort	>65: 1865	74.9	44.95/55.05	Cognition	Global cog fn summary constructed from 19 neuropsychological tests that assessed episodic, semantic,	Near VA: Rosenbaum Pocket Vision Screener positioned 14 inches from eye. Vi was defined as BCVA<20/40	X2 test, pooled t-test for equal variance. Divided analytic cohort into Vi present and Vi not present. Linear reg	Each unit higher in neuroticism level worsened association	Moderate
2014							working memory, perceptual speed and visuospatial ability. Raw scores converted to Z-scores & averaged	the eye. Vi was defined as BCVA<20/40	Global cog fn: Vi b=-0.147, SE=0.086, p=0.09, Neuroticism b=-0.001, p=0.04, VixNeurot b=-0.017, SE=0.005, p0.001	between vision impairment and lower global cognitive func	
Guthrie <i>et al</i> <sup>42</sup>	Canada (Resident Assessment Instrument for Home Care)	Cross-sectional study	Home care: 291824	82.8	38.9/61.1	Physical, social and emotional	Cognitive Performance Scale (CPS) which has four terms pertaining to short-term memory, independence	Data from RAI-HC, participants interviewed on the use of glasses	HC: CI+VI highest mem problems (56.1%) and highest prevalence of depression (28.2%)	CI+DSI had highest rate of functional impairments, communication	Moderate
2018	(RAI-HC)		Longterm care: 110578	86.9	30.5/69.5	functioning	in eating, expressive communication and decision making. Scale of 0–6, intact to very severe impairment	0 adequate Vh and four severely impaired(can see only lights and colours)	LTC: All three impairments showed highest rates on indicators of delirium.	problems and difficulty understanding others	
Hofer <sup>44</sup>	Nordic Research on Ageing Study (NORA)	Cross-sectional study	1041	75 y	42.7/57.3	Cognition, Vision	WAIS-R: Digit Symbol test, Raven's Progressive Matrices- test of inductive reasoning,	Acuity: Computerised refractometer (Topcon RMA2300).	Of the correlations with the set of cognitive variables, the highest correlations were with corrected vision	no consistent associations were found across sensory, balance,	Moderate

Continued

**Table 4** Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ci: type of measurement/evaluation	Vi: Type of measurement/evaluation	Point estimates and analysis performed	Summary of association	Quality of study
2003							Word Fluency Task, The Digit Span Forward and Backward test, Reaction time to vis/audi stimuli	Composite scores derived using avg of right and left vision scores	between .20 and .40 (visual choice reaction time), with the highest asso on visuomotor performance tests.	strength, and cognitive domains.	
Jefferis <i>et al</i> <sup>45</sup>	UK (Newcastle 85+ Cohort study)	Cross-sectional study	839	Diag cataract: 85.5	30/70	Cognition	Standardised MMSE, Mmblind (blind version)	Review of family practice records, any pre-existing diagnosis of glauco,	Diagnosed Glau: 1.76 (1.05, 2.95), Diagnosed Glau or drops: 1.73 (1.05, 2.85)	Association between recorded glaucoma diagnosis and cognitive	Moderate
2013								cataract, cataract surgery (uni/ or bilateral) sight impaired registration	scores and record of previous cataract surgery & better cognition		
Luo <i>et al</i> <sup>48</sup>	China (Second National Sample Survey on Disability)	Cross-sectional study	250752	Total: 72.9 Diag Glaucoma: 85.4	47.5/52.5	Cognition	Dementia ascertained using combo of self-report/family member report and on-site psychiatrist eval	Acuity: Trained Ophthalmologist assessed VI according to WHO BCVA	Dementia: Only VI- 1.54 (1.27-1.86), p<0.001, Only HI- 1.04 (0.88-1.22), DSI- 1.63 (1.25-2.11), p<0.001	VI & DSI were more likely to have severe to extremely severe	Moderate
2018							Questionnaire tested: poor memory, diff concentrating, diff controlling emotions, strange language/beha	criteria (low vision: 0.05≤BCVA ≤ 0.29; blindness: no light perception		dementia than those without sensory impairment.	
Rovner <i>et al</i> <sup>62</sup>	US (AMD Trial)	Cross-sectional study	241	82.8	36.5/63.5	Vision function	Animal Fluency Test-brief assessment of cognitive function relevant to the completion of daily activities	Acuity: BCVA using ETDRS chart, NEI-VFQ near vision subscale	Predictors of Visual function: VA b=-1.93(-2.59 to -1.27), <0.001, Animal fluency b=-0.054 (0.019 to 0.090), .003	Visual ability was highly correlated with VA. Multivariate	Moderate
2011							Name as many animals as possible in 60 secs. Requires semantic knowledge, speed mental processing	CS using Pelli-Robson chart	CS b=-0.20 (-0.210 to 0.610), .33, PHQ-9 severity: -.0032 (-0.100 to 0.035), .35, Age b= -.0008 (-0.033 to 0.016)	model revealed coping strategies and cog fn contributed to Near Vn	

Continued



**Table 4** Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ct: type of measurement/ evaluation	Vi: Type of measurement/ evaluation	Point estimates and analysis performed	Summary of association	Quality of study
Spieler <i>et al</i> <sup>64</sup>	US	Cross-sectional study	190	81.6	30.5/69.5	Cognition	MMSE for the visually impaired (removed eight items from the original MMSE) max score of 22	Corrected near VA was tested using the Jaeger chart. Refraction using	Asso VA & High MMSE blind score: Bad near VA (>J3): 1.0, good near VA (<J3); OR=3.18 (1.57–6.43) p<0.001	Good VA and Wearing reading glasses were significantly associated	Moderate
2016				Range 75–101				Autorefractometer		with high MMSE-blind score	
Varin <i>et al</i> <sup>65</sup>	Canada	Cross-sectional case-cohort	303	AMD 83.6	30.2/69.8	Total cognitive activities	Victoria Activity Questionnaire, which is a 70-item survey assessing older adults' participation in various	Acuity: ETDRS chart, VF: Humphrey frequency doubling	Total cognitive activities: Normal vision b=0 (ref), AMD: -4.19 (-5.96, -2.43) *p<0.05.	Patients with AMD and glaucoma participated in fewer cognitive	Moderate
2017				Glaucoma 77.5	40.9/59.1		cognitive activities over the past 2 years. Ex. Gardening, exercise, prep meals, doing homework, etc	technology full-threshold N-30 testing each eye. Medical chart reviewed	Glaucoma: -1.80 (-3.34, -0.26) *p<0.05.	activities compared with older adults with normal vision potentially	
Gussekloo <i>et al</i> <sup>67</sup>	Netherlands (Leiden 85+study)	Cross-sectional cohort	459	85	34/66	Cognition	MMSE, The 12-Word Learning Test was used as a visual test of long-term memory. Cognitive speed	Acuity: ETDRS chart	MMSE and VI: beta=-1.2 (-1.6 to -0.84), Immediate word-learning test beta=-1.3 (-1.8 to -0.69)	VI was associated with lower MMSE scores. Increasing VI was asso with	Moderate
2005				Normal 72.8	48.3/51.7		Cognition was tested with the MMSE-blind version	for diagnosis of AMD and any co-existing eye disease		putting them at risk for cognitive impairment	
2005							was measured with the Letter Digit Coding Test (processing speed)		Delayed word-learning test beta=-0.62 (-0.9 to -0.35), Letter digit coding test beta=-2.1 (-2.8 to -1.4)	with poorer scored on memory and cog speed.	
Hajek <i>et al</i> <sup>68</sup>	Germany (German Study on Ageing, Cognition and Dementia in Primary Care Patients)	Longitudinal cohort	2394	82.5	34.2/65.8	Cognition	Cognitive activities included reading, writing, solving crossword puzzles, memory training, games	Patient graded their vision on three level Likert scale	Onset of mild visual impairment: beta (SE)=-0.065 (0.026)	Linear fixed-effects regression showed that the onset of severe	Moderate
2016							(eg, card games, board games), playing music, and social engagement (eg, in church, volunteering, in a club)	(with optical aid if necessary) as no impairment, mild, severe or profound VI	Onset of severe or profound visual impairment: beta (SE)=-0.376 (0.062)	VI was associated with a decline in physical & cognitive function score in the total sample	

Continued



**Table 4** Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ci: type of measurement/ evaluation	Vi: Type of measurement/ evaluation	Point estimates and analysis performed	Summary of association	Quality of study
Hidalgo <i>et al</i> <sup>69</sup>	Spain	Cross-sectional cohort	1160	73.3	44.1/55.9	Vision	Short portable Mental Status questionnaire	Acuity: Essilor visiotest instrument with snellen optotypes	VA <6/18 in the better eye -0.344 (-0.392 to -0.295), Self-reported vision: average/poor/very poor	Using multiple regression analysis, the variables associated with VF status	Moderate
2009								Self reported vision as very good, good, average, poor and very poor.	-0.077 (-0.101 to -0.054), Dependence in activities of daily living: -0.121 (-0.160 to -0.081), Cognitive impairment -0.073 (-0.117 to -0.029)	were: visual impairment, self-reported poor vision, dependence in daily activities, cognitive	
Pham TQ <i>et al</i> <sup>63</sup>	Australia (The Blue Mountains Eye Study)	Cross-sectional control	3509	Late AMD: 79	34/66	Cognition	Modified MMSE: omitted vision-dependent items, with a max score of 22.	Acuity: LogMAR chart, Macula photo using Zeiss FF3 fundus camera	OR for MMSE 24-27 and: NO AMD 1, Late AMD 2.3 (1.1-5.0), Early AMD 1.0 (0.7-1.4)	Significant association found between Late AMD and cognitive impairment	Moderate
2006				Early AMD: 75.1	38.1/61.9			Amd lesions were assessed from the photos based on Wisconsin AMD grading system	For MMSE 0-23 and: NO AMD 1, Late AMD3.7 (1.3-10.6), Early AMD 1.4 (0.8-2.6)		
				No AMD: 65.7	43.7/56.3						
Uhlmann <i>et al</i> <sup>64</sup>	US	Cross-sectional control	174	77	42/58	Cognition	MMSE	Acuity: was measured with the Snellen and Rosenbaum methods for far and near vision.	Relative Odds for Dementia of Far-vision Impairment: 1.9 (95% CI=0.8-4.6), RR of dementia and near Vn:	Visual impairment is associated with both an increased	Moderate
1991								20/100-20/200 1.5 (0.5-4.7), >=20/200 1.8 (0.5-6.1), RR of dementia and far Vn: 20/70-20/100 1.5 (0.3-6.1).	risk and an increased clinical severity of Alzheimer's disease		
								>=20/200 1.5 (0.3-6.0)		Consistent with the hypotheses that V1 exacerbates Cog dysfunction in dementia	

Continued

Table 4 Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ct. type of measurement/evaluation	Vt. Type of measurement/evaluation	Point estimates and analysis performed	Summary of association	Quality of study
Guthrie <i>et al</i> <sup>65</sup>	four countries	Cross-sectional	Canada: Comparison: 138 650, DSI 48 210	NA	F 67.2, 67.9	Cognition	CPS, captures issues with memory, independence in eating and decision-making	inter RAI assessment includes items measuring the presence of impairments	individuals with DSI were more likely than all others to have moderate to severe cognitive impairment (between 3.8% and 9.0% higher rate).	Yes. DSI significantly associated with CI	Moderate
2016			US: comparison 60 971, DSI 6577		F 66.8, 68.6		(Established validity against MMSE)	in vision. The vision item captures the person's ability to see close objects in adequate light while using their typical assistive device. Scored on a five-point scale where 0=adequate, 1=impaired, 2=moderately impaired, 3=highly impaired and 4=severely impaired	highly compromised independence with ADLs (5.2% to 11.4% higher), instrumental ADLs (6.8% to 21.4% higher). Those with DSI were not more likely to experience social isolation		
			Belgium: comparison 500, DSI 256		F 74.9, 78.9						
			Finland: comparison 4765 DSI 1358		F 70.9, 76.2						
Harrabi <i>et al</i> <sup>66</sup>	Canada	Cross-sectional cohort	420	AMD 82.5	25/75	Cognition	MMSE	Acuity: ETDRS chart with illuminated light box at 2 m, at 1 m if patient could	Model 1*: Binocular Logmar acuity, per 0.1 unit, -0.11 (-0.18, -0.04),	People with AMD, Fuch's corneal dystrophy, and glaucoma had lower cognitive	Moderate
2015			Fuch's 78.5	17/83				not see at 2m. Letter by letter scoring done, converted to logMAR	Model 2*: Log CS, per 0.1 unit, 1.40, (0.72, 2.13)	Scores, on average, than controls (p<0.05).	
			Glaucoma 76.6	42/58				CS measured using Pelli-Robinson chart 1m	Model 3*: VF in better eye, per 1 dB, 0.04, (-0.003, 0.08)	Scores were between 0.7 and 0.8 units lower than the control group	
			Control 74.0	41/59				VF: Humphrey frequency doubling technology (FDT)			

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Table 4 Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ci: type of measurement/evaluation	Vi: Type of measurement/evaluation	Point estimates and analysis performed	Summary of association	Quality of study
Holmen <i>et al</i> <sup>68</sup>	Sweden (Kungsholmen project)	Case-control	Cog impaired 82 55	82		Loneliness, but measured interaction between	MMSE	Acuity: Snellen, self-report questions: Do you have any problems with your sight? Yes or No" was used to assess the problems	There were no significant differences in the cognitive groups between subjects with slight to moderate	NO. In a multiple regression analysis, higher MMSE score and visual improvement were significantly related to lower levels of self-reported loneliness among the elderly with their cognition intact, but not	Moderate
1994			Cog intact 92	83		Cognition & vision		habitual VA (ie, those who could read newspapers with glasses and visually orient themselves without difficulty), and subjects with pronounced to severe visual impairment with residual vision ( $\chi^2=2.28$ , $p=0.13$ ). The same differences were found when corrected VA was compared analytically between the cognitive groups ( $\chi^2=1.15$ , $p=0.28$ )	among the subjects with impaired cognition.		
Su <i>et al</i> <sup>69</sup>	Taiwan (Taiwan National Health Insurance programme)	Cross-sectional cohort	Glaucoma 6509	59.4	45.5/56.5	Cognition	Dementia patients enrolled using ICD codes	Glaucoma cohort: formed enrolling patients who were newly diagnosed with it	HR=1.13 (1.01–1.27)	The patients with glaucoma exhibited a significantly higher risk of	Moderate
2016			Control 26036	59.2	42.4/54.6				dementia than the individuals without glaucoma dld (HR (HR)/4.13, 95% CI (CI)/4.01–1.27)		
Miyata <i>et al</i> <sup>71</sup>	Japan (Fujiwara-kyo Study)	Cross-sectional cohort	2764	76.3	52.6/47.4	Cognition	MMSE	Acuity, Landolt ring chart at 5 m, converted to logMAR, refractive errors were measured by an autorefr/keratometer	OR for MCI: no cataract surgery 1.00 (ref) / cataract surgery 0.79 (0.64–0.97) p 0.025	The subjects who had prior cataract surgery had significantly lower OR of having MCI	Moderate
2018								A prior cataract surgery was determined by the self-administered questionnaire	OR for dementia: no cataract surgery 1.00 (ref) / cataract surgery 1.20 (0.64–0.97) p 0.36	Cataract surgery may be important in reducing the risk of developing MCI but not for dementia independently	

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Table 4 Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Cl: type of measurement/ evaluation	Vi: Type of measurement/ evaluation	Point estimates and analysis performed	Summary of association	Quality of study
Ong <i>et al</i> <sup>72</sup>	Singapore (Singapore Malay Eye Study (SAMES))	Cross-sectional cohort	1032	Cognitive dysfunction 70.5	24/76	Cognition	Abbreviated Mental Test (AMT)	Acuity: number chart at a distance of 4 m converted to logMAR. VI logMAR>0.3 in the better eye	Cog Dysfunction OR: Model 2 :Myopia 1.78 (1.02–3.10) Hyperopia 1.11 (0.68–1.80) Emmetroopia 1.0	Yes. Compared with individuals with emmetroopia, persons with myopia were	Moderate
2013				No Cog dysfunction 67.7	66.5/33.5		[cognitive dysfunction was defined as a score less than or equal to 6 of 10 for those with 0 to 6 years of formal education		Model 3 : Myopia 1.86 (1.01–3.42) Hyperopia 0.92 (0.54–1.55) Emmetroopia 1.0	almost twice as likely to have cognitive dysfunction	
Soler <i>et al</i> <sup>73</sup>	France	Cross-sectional cohort	1648	82.6	35.6/64.4	Vision	MMSE	Acuity, Snellen decimal chart for distant vision, Parnaud chart for near vision	model one for distant vision: MMSE OR (CI) 0.94 (0.92–0.97) p<0.001	Yes. Results show that visual impairment is independently associated	Moderate
2016						and less than or equal to 8 of 10 for those with more than 6 years of formal education.]		Amsler grid testing	model two for near vision: MMSE OR (CI) 0.92 (0.89–0.94) p<0.001	with lower educational level, cognitive impairment, and lower ADL-assessed autonomy	
Sun <i>et al</i> <sup>74</sup>	China	Cross-sectional cohort	4123	67.5	48.8/51.2	Cognition	ADL, IADL, Fried score, SPPB class (Short Physical Performance Battery), AMT	Objective refraction was measured using an autorefractor (Canon RK-5)	OR of cog dysfunction for myopia: Before propensity score matching, OR 1.98 (1.61–2.44)	Results of this PSM analysis support a small, but positive association of	Moderate
2016							Spherical equivalent (SE) (Myopia: a SE of less than –0.50 dioptre (D) in the right eye. High myopia: a SE of less than –6.00 D.)	After matching: OR 1.31 (1.00–1.71). Multivariate logistic regression	myopia with cognitive dysfunction among elderly Chinese in China		
							model on the association between myopia and cognitive dysfunction, OR (95% CI):				
									Myopia (Yes vs No) 1.52 (1.23–2.06), Age-related cataract (Yes vs No) 2.06 (1.57–2.98)		

Continued



Table 4 Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ct: type of measurement/ evaluation	Vt: Type of measurement/ evaluation	Point estimates and analysis performed	Summary of association	Quality of study
Zheng <i>et al</i> <sup>75</sup>	US (The Salisbury Eye Evaluation Study)	Longitudinal cohort	2520	73.5	42/58	Cognition	MMSE	Acuity, ETDRS chart refraction was performed on participants with worse baseline MMSE score	Worse baseline VA was associated with worse baseline MMSE score	Worse baseline VA associated with worse baseline MMSE score	Moderate
2018								20/32 VA or worse using a forced-choice procedure, associated with worse VA was associated with worse VA (r = -0.226; 95% CI, -0.291 to -0.16; p<0.001). The rate of worsening VA was associated with the rate of declining MMSE score (r = -0.139; 95% CI, -0.261 to -0.017; p=0.03).	The rate of worsening VA was associated with a significant declining MMSE score	Moderate	
Ward <i>et al</i> <sup>76</sup>	US (Study of Osteoporotic Fractures)	Longitudinal cohort	1352	77.7	0/100	Vision	Global cognition (3MS), verbal memory (California Verbal Learning Test II, short form (CVLT)), executive function (Trails B, Digit Span Forward, Digit Span Backward), and semantic memory (category and verbal fluency)	CS, using a VISTECH (Hartford, CT) VCTS 6500 wall chart and light meter	OR of MCI/dementia for lowest quartile of baseline CS compared with women in the highest quartile: OR 2.16 (95% CI=1.58-2.96)	These data support an association between impaired CS and future MCI/dementia.	Moderate
2018											
Mitoku, K, Masaki N <i>et al</i> <sup>78</sup>	Japan (Gujo City Long-Term Care Insurance database)	Cross-sectional cohort	1754	80.89/82.42	34.5/65.5	Cognition	The assessment measures include communication, short-term memory,	Acuity: No chart name provided	Logistic regression: OR of CI was 1.389 (95% CI 1.04 to 1.85) in those with VI	Yes. Supports the relationship between DSI and CI. increased risk of mortality in those with DSI & CI.	Moderate
2016											
								Other: Vision at baseline five levels, "normal sight", "able to see chart at 1 m" "able to see chart in front, "very little sight", "indeterminable"	1.581 (95 % CI 1.24 to 2.01) in those with HI, and 2.349 (95 % CI 1.79 to 3.09) in those with DSI		
								season of year in addition to 18 problem behaviours			
								CI classified as none, mild, moderate, severe			

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Table 4 Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ct: type of measurement/evaluation	Vt: Type of measurement/evaluation	Point estimates and analysis performed	Summary of association	Quality of study
Jefferis JM <i>et al</i> <sup>79</sup>	UK	Longitudinal, cohort	112	80.7	45/55	Cognition	The ACE-R (which includes the MMSE), is scored from 0 (worst cognition) to 100 (best cognition);	logMAR VA in the better eye corrected with up-to-date refraction and/or pinhole	Group, Mean±SD: Baseline, Postop, 1 Year. P Value*: 3-Way†, Baseline vs Postop, Baseline vs 1 Year, Postop vs 1 Year	Significant improvements in ACE-R scores were seen between baseline and 1 year	Moderate
2015						38 of these points pertain to vision-dependent items and 62 to vision-independent items			Normal (n=46): 92.5±2.8, 92.6±3.8, 93.7±3.4, 0.018, 0.9, 0.009, 0.004:: Mildly impaired cognition (n=22): 84.5±1.4, 84.8±4.3, 88.0±3.5, <0.001, 0.75, <0.001 0.002	postoperatively (95% CI for improvement, 0.5–2.8; P Z .005). Improved	
									Moderately impaired cognition (n=23): 72.6±8.7, 73.6±10.4, 73.3±14.4, 0.75, --, --, --	cognition did not correlate with improved VA (r Z 0.13, P Z .22)	
Lindenberger <i>et al</i> <sup>81</sup>	Germany (BASE Berlin Ageing Study)	Longitudinal, cohort	516	85	NA	Cognition	MMSE	Acuity: Snellen decimal units. Close VA (close vision) was measured separately for the left and the right eye with a standard reading table presented at reading distance. Distance VA (distant vision) was assessed binocularly with a reading table presented to the participants at a minimum distance of 2.5 metres	Multivariate Results With Control for Age, Time to Death, and Risk of Dementia Construct, Mean (Level, Linear change, Quadratic change), Variance (Level, Linear change), Age, Dementia status, Age X Linear Change, Residual variance.	Contrary to expectations, the correlations between cognitive and sensory declines were only moderate in size, underscoring the need to delineate both domain-general and function-specific mechanisms of behavioral when controlling for age at first measurement, distance to death, and risk of dementia senescence	Moderate
									All participants (n=91): 85.6±9.5, 85.9±10.0, 87.2±11.4, 0.004, 0.45, 0.005, 0.006, *p<0.05 statistically significant		
2009									Close vision: 47.33 (0.30), -1.37 (0.11), ---, 34.17 (2.95), 0.83 (0.16), -0.44 (0.04), ---, -0.03 (0.01), 25.87 (1.42)		

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Table 4 Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ci: type of measurement/ evaluation	Vi: Type of measurement/ evaluation	Point estimates and analysis performed	Summary of association	Quality of study
Hong <i>et al</i> <sup>82</sup>	Australia (Blue Mountain Eye Study)	Longitudinal, cohort	2334	No Sensory impairment 66.9 (7.4),	No SI 58.4/41.6	Cognition	MMSE	Acuity: Distance VA at 24.4cm followed by pinhole acuity (VA was recorded as the number of letters read correctly) If no letters could be read count fingers, hand movements, light perception, or no light perception	Distant vision: -48.47 (0.32), -0.56 (0.11), ---, 28.14 (3.17), 1.20 (0.25), -0.59 (0.04), ---, ---, 44.79 (2.45) association between Decline>=3 MMSE blind score and visual impairment (adjusted for age and sex). OR after 5 years: Visual impairment 0.84 95% CI 0.40 to 1.79 DSI 1.41 95% CI 0.54 to 3.72 after 10 years: Visual impairment 1.09 95% CI 0.52 to 2.30 DSI 1.15 95% CI 0.28 to 4.73	No. The presence of VI, HL or DSI was not associated with possible cognitive decline over 5 years or 10 years.	Moderate
2016				Visual impairment 74.3 (8.4),	VI 65.8/34.2					There were no changes to these findings after adjustment for other potential confounder	
Mandas <i>et al</i> <sup>85</sup>	Italy	Case-Control	1168	Dual impairment 80.4 (7.0)	Dual impairment 29/71	Cognition	MMSE	Acuity: Snellen Chart	Total(n=1168), Control1 (n=436), All types of dementia (n=732), MCI (n=181), AD (n=230), MD (n=126), VD (n=195)		Moderate
2014				78.4	29/71			IOP, fundus examination under slit-lamp and fundu photograph	All vision disorders: No, n=297, n=433, n=117, n=135, n=68, n=113 yes, n=139, n=299, n=64, n=95, n=58, n=82, 2, 9.37, 0.70, 5.87, 8.60, 6.12 P-value, 0.002, 0.40, 0.015, 0.003, 0.013 OD (95% CI), 1.5 (1.1-1.9), 1.2 (0.8-1.7), 1.5 (1.1-2.1), 1.8 (1.2-2.7), 1.5 (1.1-2.2)	Subjects with any type of age-related vision disorders(n D438), cataract,	

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Table 4 Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ci: type of measurement/evaluation	Vi: Type of measurement/evaluation	Point estimates and analysis performed	Summary of association	Quality of study
Mendola <i>et al</i> <sup>86</sup>	US	Case-Control	188	NA	NA	Vision	Blessed Dementia Scale (BDS)	Acuity: Snellen, CS; two-alternative forced-choice format,	Test, Prevalence in AD, Correlation with BDS, % variance from BDS.	The results indicate that visual dysfunction, especially on Backward Masking, is a common sign of AD.	Moderate Macular degeneration, glaucoma, and diabetic retinopathy, were more likely to be depressed and had significantly lower MMSE scores than Control 2
1995								Stereopsis: binocular disparity stereopsis, as a clue to depth perception(Stereo Optical)	Backward Pattern Masking: 58% r=0.22; p=0.106; df=55, 13%; Gollin Incomplete Pictures: 30%, r=0.33; p=0.03; df=17, 6%		
								funduscopy, evaluation of fixation, pupillary function, extraocular movement,	Luria Mental Rotation: 39%, rb=0.74; p=0.004; df=36, 13%		
								Backward Masking, Gollin Incomplete Pictures, City University Colour Vision.	Log CS at one cpd: 39%, rb=0.69; p=0.002; df=17, 52%, Log CS at 2cpd: 6%, rb=0.55; p=0.022; df=17, 7%		
								Luria Mental Rotation. Moeny Road Map. Local speed discrimination and Global motion detection. Critical Flicker fusion.	Money Road Map: 29%, rb=0.37; p=0.028; df=35, 17%		
Magalhães <i>et al</i> <sup>88</sup>	Brazil	Cross-sectional cohort	466	71.4	44.4/55.6	Cognition	Cambridge Cognitive Examination (CAMCOG)	Self-report	logistic regression model for dementia:	Yes. Study showed strong association between dementia and sight-impairment	Moderate
2008							Dementia was diagnosed in individuals who fulfilled CAMDEX criteria for definitive, probable or possible dementia through a combination of clinical criteria, organicity indices $\geq 5$ and CAMCoG <80.		sight impairment OR (95% CI): 1.83 (1.15–2.91)		

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**Table 4** Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ct: type of measurement/ evaluation	Vt: Type of measurement/ evaluation	Point estimates and Summary of analysis performed association	Quality of study	
Ong <i>et al</i> <sup>89</sup> 2012	Singapore (SAMES)	Cross-sectional cohort	1179	with cognitive dysfunction: 71.2 Without cognitive dysfunction: 68.0	21/79 67.7/33.3	Cognition	The AMT is a 10-question test of general cognitive function  Items assess orientation (three points), semantic knowledge (1point), episodic memory (three points), delayed recall	Acuity: LogMAR number chart (lighthouse international)  Cataracts were assessed from lens photographs, Glaucoma diagnosed and classified using International Society Geographical and Epidemiological Ophthalmology scheme based on gonioscopy	Pvalue 0.011  Age-related eye disease, model 1, model 2, model 3.  Cataract: 1.52 (1.02–2.27), 1.42 (0.93–2.16), 1.44 (0.94–2.22), AMD: 1.71 (0.89–3.29), 1.44 (0.71–2.91), 1.36 (0.34–5.45)  Yes. VI due to cataract (OR=2.75; 95% CI, 1.35 to 5.63)	Moderate	
Mangione <i>et al</i> <sup>90</sup> 1993	US	Cross-sectional cohort	472	73	40/60	Cognition	(one point), picture naming (one point), and attention (one point)	Acuity: Snellen (results were transformed into percentages of functional central vision loss following the formula outlined in the Physician's Desk Reference for Ophthalmology)	Moderate or severe DR: 2.43 (1.13–5.20), 2.26 (1.02–5.00), 5.57 (1.56–19.91)  Glaucoma: 1.68 (0.81–3.46), 1.61 (0.75–3.45), 2.09 (0.52–8.31)  multivariate linear regression, percent increase in ODDS of abnormal TICS score (95% CI):  binocular vision loss(per 10% decline in vision) –9 (–23.9), percent field loss (per one category decline) 13 (–13,48) cataract (per eye) 18 (–12,58) ocular miotics 37 (–17,126)	were more likely to have cognitive dysfunction. Only moderate to severe diabetic retinopathy was independently associated with cognitive dysfunction  No. An association between low TICS score and visual disorders were not seen	Moderate
VF: Humphrey field analyser									Continued		

Table 4 Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Clt: type of measurement/ evaluation	Vi: Type of measurement/ evaluation	Point estimates and analysis performed	Summary of association	Quality of study	
Wittich <i>et al</i> <sup>83</sup>	Canada (COMPASS-ND study)	Longitudinal, cohort	109	72.94	No info	Cognition	MoCA	Reading acuity (MNRead) and CS (Mars Test)	Individuals with AD had significantly lower CS than those with MCI and SCI ( $p=0.04$ , $\omega^2=0.04$ ), after adjusting for age, sex and education. No differences in VA were observed $p=0.46$ , $\omega^2=0.004$ .	Yes. Declines seen in CS in individuals with AD relative to those with MCI and SCI.	Moderate	
2019									Study also observed higher rates of reduced reading acuity in the AD and MCI groups relative to those reported the general population.			
Brenowitz <i>et al</i> <sup>64</sup>	US (Health ABC Study)	Longitudinal, cohort	1810	76.7	39.4/60/6	Cognition	Dementia was defined as meeting one or more of the following criteria through study	Bailey-Lovie distance VA test, Pelli-Robson CS	Multivariable model for incident dementia with VA: HR=1.26 (0.90–1.77), CS: HR=1.11 (0.88–1.38)	No. Vision impairment and CS independently significantly associated with incident dementia	Moderate	
2019							Year 15 (2011–2012); (a) hospitalisation with dementia as a primary or secondary diagnosis, (b) documented use of dementia medication, or (c) clinically meaningful Modified MMSE (3MS) decline from Health ABC baseline (1.5 SD, race-stratified)					
Davies-Kershaw <i>et al</i> <sup>65</sup>	UK (English Longitudinal Study of Ageing)	Longitudinal cohort	7685	NA	44/56	Cognition	three methods to identify individuals with dementia: a physician diagnosis of dementia	Rate their eyesight (using glasses or corrective lenses as usual)	Cross-sectional analysis using Wave seven measures adjusted for covariates: compared with normal VA, moderate OR=2.04 (1.36–3.07)	Yes. Older adults with vision impairment have higher rates of dementia crosssectionally (all ages) and are at greater risk of incident dementia longitudinally (<70 only)	Moderate	
2918	Wave 7						that the participant or a caregiver reported between Wave 3 (2006–7) and 7 (2014–15);		poor OR=4.02 (2.64–6.13) $p<0.001$			

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Table 4 Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ci: type of measurement/ evaluation	Vi: Type of measurement/ evaluation	Point estimates and analysis performed	Summary of association	Quality of study	
							a score less than 3.5 on the adaptive Informant Questionnaire on Cognitive Decline in the Elderly (IQCODE)	Combined groups into three categories for analysis (excellent or very good=normal, moderate (HR=1.78, 95% CI=1.04-3.04) and poor (HR=3.60, 95% CI=1.10-11.78) self-rated vision were at greater risk of developing dementia than those with normal self-rated vision	Longitudinal analysis using Wave two measures individuals in the younger group (60-69) and with moderate (HR=1.78, 95% CI=1.04-3.04) and poor (HR=3.60, 95% CI=1.10-11.78) self-rated vision were at greater risk of developing dementia than those with normal self-rated vision			
							or prescriptions for anticholinesterase inhibitors, N-methyl-D-aspartic acid receptor antagonists, and other relevant medication (galantamine, rivastigmine, memantine, donepezil, tacrine) to indicate dementia	good or fair=moderate, poor or registered or legally blind=poor or blind				
Gui-Ying Cao	China (China Health and Retirement Longitudinal Study)	Longitudinal, cohort	7269	60.2	56.2/43.8	Cognition	Episodic memory (measured as the average of immediate and delayed recall scores of ten Chinese nouns), mental intactness (measured using some components of the Telephone Interview of Cognitive Status (TICS) battery), and global cognition (the sum of episodic memory and mental intactness scores)— was evaluated and followed up every 2 years	Self-rated vision measure from Wave 7 (2014-15) for cross-sectional analysis and from Wave 2 (2004-05) for the longitudinal analysis	Compared with older people with no VI, (1) those with DVI only were associated with poor episodic memory ( $\beta=-0.076$ , $p<0.0001$ )	Yes. VI is associated with an increased risk of poor cognitive function	Moderate	
2018							mental intactness (measured using some components of the Telephone Interview of Cognitive Status (TICS) battery), and about near VA for reading newspapers. Individuals were assigned to one of four categories: no VI, distance vision impairment (DVI), near vision impairment (NVI), or both distance and near vision impairment (DNVI)	VI was assessed by two self-reported questions, about distance VA for seeing faces on the other side of the street and about near VA for reading newspapers. Individuals were assigned to one of four categories: no VI, distance vision impairment (DVI), near vision impairment (NVI), or both distance and near vision impairment (DNVI)	Those with NVI only were associated with poor mental intactness ( $\beta=-0.031$ , $p=0.0001$ ) and global cognition ( $\beta=-0.032$ , $p=0.0224$ )			

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Table 4 Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	CI: type of measurement/ evaluation	VI: Type of measurement/ evaluation	Point estimates and Summary of analysis performed association	Quality of study
All <i>et al</i>	US (Medicare Claims data from 2014)	Cross-sectional cohort	472871	>65	34/66	Cognition	ICD-9 diagnosis codes for dementia or cognitive impairment	ICD-9 diagnosis codes for blindness/ low vision	Yes. There was a significant association between low vision/blindness and dementia or AD  Among the low vision/blindness group, the prevalence of dementia or CI was 50%.  Those with DNVI were associated with poor episodic memory ( $\beta=0.106$ , $p<0.0001$ ), mental intactness ( $\beta=-0.107$ , $p<0.0001$ ), global cognition ( $\beta=-0.105$ , $p<0.0001$ )	Moderate
2019									Blindness/low vision was also associated with a greater odds of Alzheimer's disease (AOR 1.44 95% CI: 1.415 to 1.464)	
Preeti Gupta <i>et al</i>	Singapore (Singapore Epidemiology of Eye Disease (SEED) Study)	Longitudinal cohort	682	67.3	55.6/44.4	Cognition	CI was assessed using the validated AMT defined as scores of	Fundus images using Canon DGI camera, graded by trained graders at University of Sydney	and all-cause dementia (AOR 1.239, 95% CI: 1.223 to 1.254)  Those with any DR had higher odds of incident CI (OR=2.32, 95% CI 1.07 to 5.03), and those with moderate or worse DR also had a higher likelihood of	Moderate
2019									increased risk of developing CI, independent of vision and other risk factors  using the modified Airie House classification system and categorised as none (Early Treatment of Diabetic Retinopathy Study (ETDRS) level 10), minimal/mild (level 20–35) and moderate or worse DR, (level 43–90) using data from the better eye	
Jost <i>et al</i>	China (The Beijing Eye Study)	Cross-sectional cohort	3127	64.2	43.4/56.6	Cognition	MMSE, assessed as cognitive function score (cfs). Categorized as mild (cfs 23–19), moderate (cfs 18–10)	Autorefractometry, Presenting, uncorrected and best corrected VA, associated with better best corrected VA ( $V2=0.38$ ),	Yes  Better cognition (ie, higher CFS) was significantly associated with better best corrected VA ( $V2=0.38$ ),	Moderate

Continued



Table 4 Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ct: type of measurement/evaluation	Vi: Type of measurement/evaluation	Point estimates and analysis performed	Summary of association	Quality of study
2018							severe (cfs<9)	fundus photo, SD-OCT	smaller amount of undercorrected VA, lower prevalence of primary angle-closure glaucoma, and thicker subfoveal choroidal thickness		
Allen <i>et al</i>	China	Longitudinal, cohort	15 576	74.5	36.2/63.8	Cognition	ICD-10 codes or Clinical dementia rating 1 to 3	Acuity: Snellen E chart, converted to LogMAR	Participants with incident dementia had poorer VA at baseline than those without (adjusted HR=5.88, 95% CI=4.04–8.57).	Yes, Moderate-to-severe visual impairment could be a	Moderate
2020									Incident Dementia at years 4–6 of Follow-up in Participants With Mild (HR=1.19, 95% CI 0.86 to 1.65, p=0.31)	potential predictor and possibly a risk factor for dementia	
									Moderate (HR=2.09, 95% CI=1.47–2.97) or Severe VI (HR=8.66, 95% CI=4.60–16.30) at baseline		
Cecilia <i>et al</i>	US (Adult Changes in Thought (ACT))	Longitudinal, cohort	3877	NA	42/58	Cognition	Cognitive Abilities Screening Instrument scores 85 underwent a standardised diagnostic evaluation, including physical and neurologic examinations and a neuropsychological test battery	ICD-9 codes for diagnosis of glaucoma, AMD, DR and cataract	The recent and established HR were 1.46 (P 5.01) and 0.87 (P 5.19) for glaucoma,	Yes, Increased AD risk was found for recent glaucoma diagnoses,	Moderate
2019							Dementia diagnoses using NINCDS-APDR criteria, Our primary outcome was either probable or possible late-onset clinical AD		1.20 (P 5.12) and 1.50 (P.001) for AMD, and 1.50 (P 5 .045) and 1.50 (P 5 .03) for DR	established AMD diagnoses, and both recent and established DR	
Moon Jeong Lee <i>et al</i>	US (National Health and Nutrition Examination Survey (1999–2006 cycles))	Cross-sectional cohort	5795	Non-VI 70.3	45/55	Cognition	Memory or confusion complaints were defined as present; if participants responded “yes” to the question —	VA was assessed using the built-in chart in an auto-refractor (NIDEK ARK-780; Nidek Co Ltd, Tokyo, Japan)	Individuals with VI were more likely to report cognitive complaints as compared with those without VI	Yes	Moderate

Continued

Table 4 Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ci: type of measurement/ evaluation	Vi: Type of measurement/ evaluation	Point estimates and analysis performed	Summary of association	Quality of study
2019				VI 77.8	39/61		“(Are you/’s survey participant) limited in any way because of difficulty remembering or because (you/’s/he) experience(s) periods of confusion?”	VI was defined as autorefractor corrected VA worse than 20/40 in the better-seeing eye			
Ruby Yu <i>et al</i>	China	Longitudinal, cohort	1949	76.1	23.1/76.9	Cognition	5-item Abbreviated Memory Inventory for Chinese (AMIC). Scored range from 0 to 5 (1 point for each item; 0 – best to 5 – worse). An AMIC score $\geq 3$ is predictive of MCI	Questionnaire: “Do you have any difficulty seeing things?” “very good” and “good” = “robust”; “fair”, “not too well”, “poor”, and “very poor” = “poor”	Poor vision (OR 2.2 95% CI 1.8 to 2.7) at baseline was significantly associated with an increased risk of at least 3 SMCs at follow-up	Yes	Moderate
2019											
Michio Maruta <i>et al</i>	Japan	Longitudinal cohort	2190	78.9	20.6/79.4	Cognition	Dementia Scale labelled Level 0 to Level IV and level M based on symptoms and the	VA assessed at five levels: normal vision (there is no hindrance in daily life); “able to see vision testing chart at a distance of about 1 m,”	DSI associated with a higher cumulative dementia incidence compared with no sensory impairment (log-rank $\chi^2=39.92$ ; $p<0.001$ )	Yes.	Moderate
2020							necessity for care. Level II and greater during the 8 year follow-up period was considered	“able to see vision testing chart at a distance of in front,” “very poor eyesight,”	DSI is the greatest risk factor for developing dementia among sensory impairments (HR, 1.45; 95% CI, 1.22 to 1.71; $p<0.001$ )	Older adults with sensory impairments have a high incidence of dementia, with DSI presenting the greatest risk	
							“incident dementia”	and “undecidable due to difficulty in communication.	Older adults with VI were found to be more likely to have day-night reversal symptoms when dementia occurs		
Ann <i>et al</i>	UK (English Longitudinal Study of Ageing)	Longitudinal cohort	4621	64.9	45/55	Cognition	Working memory, Executive function. Results from the three cognitive tests available were summed,	VA: Self-report asking participants whether their eyesight was excellent, very good, good, fair, or poor using glasses or corrective lens if they normally do so	Compared with people with good vision, poor vision was asso with worse cognitive function	Yes, ageing adults with individual and combined impairments in hearing and vision had greater risks of worse cognitive performance at 6 year follow-up compared with those with good sensory function	Moderate

Continued



**Table 4** Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ct: type of measurement/ evaluation	Vt: Type of measurement/ evaluation	Point estimates and Summary of analysis performed association	Quality of study
2020										
Stephanie Chen <i>et al</i>	US (NHATS)	Longitudinal cohort	7075	NA	40/60	Cognition	1. Doctor told the sample person that he/she had dementia or AD, 2. A score that indicates probable dementia on the AD8 Dementia Screening Interview	Self-reported distance and near VI	Participants with self-reported visual impairment were at significantly higher risk of developing	Moderate
2019										
All <i>et al</i>	US (Medicare beneficiaries)	Cross-sectional cohort	773905	NA	NA	Cognition	3. Cognitive tests that evaluate the sample person's memory and orientation	ICD-9 codes and procedure code-based algorithms	probable or possible dementia over subsequent follow-up (HR=4.4, CI: 3.9 to 5.0, p<0.001), compared with those without visual impairment  This association persisted after full adjustments for covariates (HR=2.1, CI: 1.8 to 2.5, p<0.001)	Moderate
2019										
Tien <i>et al</i>	Singapore (SEED)	Cross-sectional cohort	10 020	58.9	49.3/50.7	Vision	AMT, which consists of 10 questions of general cognitive function	ICD-9 codes and procedure code-based algorithms	Low vision was associated with greater odds of incident hip fracture (AOR 1.13, 95% CI: 1.04 to 1.22 and incident anxiety (AOR 1.11, 95% CI: 1.05 to 1.18 but not incident depression or dementia.	Moderate

Continued



Table 4 Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ci: type of measurement/evaluation	Vi: Type of measurement/evaluation	Point estimates and analysis performed	Summary of association	Quality of study
2019									was a significantly associated risk factor. CI was also significantly associated with higher risk of presenting bilateral VI or blindness (OR=2.15; 95% CI=1.75–2.63)		
Bonnielin Swenor <i>et al</i> <sup>12</sup>	US (Health ABC study)	Longitudinal cohort	2444	74	47.8/52.2	Cognition	DSST and 3MS. Incident cognitive impairment was defined as	VA: Bailey-love chart, CS: Pelli-Robson chart	VA impairment HR=1.55; 95% CI 1.12, 2.14 vs No VA impairment, CS impairment HR=1.33	Yes, VA, CS, and stereo acuity are	Moderate
2019							a 3MS score < 80 or a decline in 3MS > 5 points following Year 3		95% CI 1.13, 1.55 vs No CS impairment	risk factors for cognitive decline	

ACE-R, Addenbrooke's Cognitive Examination-Revised; AMD, age-related macular degeneration; CS, contrast sensitivity; DSI, dual sensory impairment; MMSE, Mini-Mental State Examination; MoCA, Montreal-Cognitive Assessment; MPOD, macular pigment optical density; NHATS, National Health and Aging Trends Study; VA, visual acuity; VF, visual field.

concluded that VI was associated with both an increased risk and an increased clinical severity of AD. Although Frost *et al*<sup>88</sup> found a strong association between early age-related macular degeneration and AD, their study was cross-sectional, and the sample size was too low to derive an inference. Both these studies received 'moderate' rating in QA. Davies-Kershaw *et al*<sup>95</sup> in their longitudinal analysis using the ELSA wave 2 and wave 7 data found that individuals in the younger group (50–69 years) and with moderate and poor self-rated vision were at greater risk of developing dementia than those with normal self-rated vision. Hamedani *et al*<sup>99</sup> used Medicare claims data from 2014 consisting of 472 871 participants and concluded that blindness/low vision was associated with a greater odd of Alzheimer's disease and all-cause dementia. Both these studies also received 'moderate' rating in QA. Soto-Perez-de-Celis *et al*<sup>62</sup> in their cross-sectional case-control study found that DSI was significantly associated with possible CI. However, the study received an overall 'weak' rating in QA.

Of the 91 studies that found an association between VI and cognitive function, 35 were longitudinal, 46 were cross-sectional and 10 were case-control studies. Of the 13 studies that found no association between VI and cognitive function, 6 were longitudinal, 5 were cross-sectional and 2 were case-control studies. Ihle *et al*<sup>60</sup> performed a cross-sectional analysis using a sample of 2812 participants from Switzerland. They objectively measured cognition and vision and concluded that their data did not support an increased relation of cognitive and sensory abilities in old age. This study received a 'weak' rating in QA. Hong *et al*<sup>82</sup> used data from Blue Mountain Eye Study, a longitudinal study from Australia that studied associations between VI and a decline in MMSE scores over a duration of 10 years. The study concluded that VI was not associated with cognitive decline over 5 years or 10 years. Although the study included a large number of participants overall (n=2334), only 152 individuals with VI were included in this analysis, suggesting that there may have been survival bias. Brenowitz *et al*<sup>94</sup> in their longitudinal study using the Health ABC data concluded that VA and CS independently were not significantly associated with incident dementia. However, Swenor *et al*<sup>122</sup> used data from the same study and found that impaired VA, CS and stereo acuity had a greater risk of incident cognitive impairment.<sup>122</sup> These three studies received a 'moderate' rating in QA. Michalowsky *et al*,<sup>117</sup> who received a 'strong' rating in their case-control study concluded that VI was not significantly associated with dementia, a combination of both visual and hearing impairments was significantly associated with the risk of dementia.

There was considerable heterogeneity in the measurement and reporting of cognitive function. Studies measured cognitive function using a variety of instruments with the most common being MMSE. The MMSE is a paper-based test with a maximum score of 30, with lower

Table 5 Studies with a 'weak' rating

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ci: type of measurement/evaluation	Vi: Type of measurement/evaluation	Point estimates and analysis performed	Summary of association	Quality of study
Bayer <sup>20</sup>	Germany	Case control	228	With AD+Glau 72.9	10%/19%	Glaucoma	Diagnosis of Alzheimer's Disease according to the NINADS-ADRDRA classification	Acuity - Snellen	25.9% occurrence rate of probable glaucoma among AD patients	Yes- Significantly higher occurrence rate of glaucoma	Weak
2002			AD 112	With AD+no Glau 71.4	28%/55%			VF - Humphrey Field Analyser	5.2% occurrence rate of probable glaucoma among healthy controls	among AD patients	
			Control 116	W/o AD+No glau 68.1	38%/72%			Optic disc evaluation			
			W/o AD+Glau 70.1	2%/4%							
Cigolle <sup>24</sup>	US (1995-2010 of HRS)	Longitudinal cohort, Adults->65y	8847	No info	No info	Cognition	Performance based measure (Telephone interview) to determine cognitive function (0-27)	No info	Vision and hearing predicted cognitive decline (p<0.001)	Cognitive function declines in an accelerating fashion, with older age	Weak
2013	Abstract Only									and visual impairment predicting decline over time	
Feeney <sup>34</sup>	Ireland (The Irish Longitudinal study on Ageing)	Cross-sectional study	4281	No info	No info	Cognition	MMSE, MoCA, CTT, CRT, SART, Prospective memory, word recall and visual reasoning	heterochromatic flicker photometry (HFP) - a non-invasive method of	Linear, Negative binomial and logistic regression. MPOD men=0.20, SD=0.15.	MPOD is significantly associated with cognitive function	Weak
2013	Abstract only							gauging the density of Macular Pigment	One SD inc in MPOD assoc with few errors on MoCA: b=-0.03 p<0.01 & MMSE: b=-0.05 p=<0.05	among older adults.	
									Faster time to complete CRT: b=-0.06 p<0.01, better word recall: b=0.07 p<0.05		
Jefferis <sup>45</sup>	UK	Cross-sectional study	112	80.7	NA	Cognition	Revised Addenbrook's Cognitive Examination (ACE-R)	Acuity: LogMar chart, Cataract density graded using LOC III system	Fewer SRT omission errors: b=-0.04 p<0.05, success on prospective memory task: b=0.14 p<0.01	Better general cognition (ACE-R total score) was associated with requiring vision, both subscores	Weak

Continued

**Table 5** Continued

Author and (Study name if applicable) title	Country	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	CI: type of measurement/ evaluation	VI: Type of measurement/ evaluation	Point estimates and Summary of analysis performed association	Quality of study	
2013	Abstract Only								were significantly and similarly associated with VA (p=0.008 and p=0.001 respectively).	better vision (p=0.001)	
Maharani <i>et al</i> <sup>49</sup>	Survey of Health, Ageing and Retirement in Europe 2002–2014	Cross-sectional study	24 116	NA	NA	Cognition	Using a summary cognitive score from the questions on episodic memory, verbal fluency, and numeracy.	Self-reported hearing and visual quality. Sensory function recoded into 2:	Older adults with single SI: b=-0.064 (-0.072 to -0.056), with DS: b=-0.229 (-0.247 to -0.211)	Older adults with single and dual SI showed significantly lower	Weak
2017	Conference abstract							Excellent, v good & good into fair & poor into Poor sens	Good sens fn and cognitive function compared with those without SI		
Miyata <sup>50</sup>	Japan (Fujiwara-Kyo study)	Cross-sectional study	668	76.3	NA	Cognition	MMSE	Cataract surgery history- obtained using a self-reported questionnaire	CI for Cataract surgery group OR: 0.82 (0.68, 0.99) p=0.042 when compared with no cataract sx group	Cat Sx group significantly lower odds for CI than the no Cat Sx group	Weak
2016	Conference abstract										
Nael <sup>51</sup>	France (Three-City Study)	Longitudinal cohort study	7722	Range: 65 & older	NA	Cognition	Incident dementia over the 12 year follow-up was actively screened for (diagnosis code)	Dist VF: loss self-reported, as inability/ difficulty recognising a face@ 4 m.	Cox regression models: Near VI: HR=1.30 (1.05, 1.61), Dist VF loss: HR=1.47 (1.02, 2.11) compared with those without visual loss	Both near VI and distant VF loss were associated with an increased risk of dementia	Weak
2017	Short oral presentation							Near VI: presenting binoc VA <20/30 @ reading distance of 33 cm.			
Settl <sup>53</sup>	Ireland (The Irish Longitudinal Study on Ageing (TILDA))	Cross-sectional study	5021	NA	NA	Cognition	MMSE and MoCA	Vision measured subjectively and objectively (no other info)	Poor hearing and immediate recall of orally presented words: b=-0.46, poor hearing and delayed recall b=-0.60	Vision was a significant predictor of visually presented tasks	Weak
2013	Abstract only							Poor hearing and category fluency b=-1.96, poor hearing & memory (OR 0.23)& absent-mindedness (OR2.18).		but also of category fluency.	

Continued

**Table 5** Continued

Author and (Study name if applicable) title	Country	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	CI: type of measurement/ evaluation	VI: Type of measurement/ evaluation	Point estimates and Summary of analysis performed association	Quality of study
Zheng <sup>56</sup>	US (The Salisbury Eye Evaluation Study)	Longitudinal cohort	2520	Range 65–84		Vision	MMSE	Acuity: ETDRS chart	VA and MMSE score worsened over time (VA (Log MAR) intercept=0.004, slope=0.022 for VA; MMSE intercept=27.3, slope=-.59; all p<0.001)	Weak
2017	Conference abstract								The intercept of VA trajectory is statistically significantly associated with the intercept of MMSE trajectory (f=-.267, SE=0.029, p<0.001) suggesting that worse baseline VA is associated with worse baseline MMSE score.	
Ihle et al <sup>60</sup>	Switzerland (LIVES study)	Cross-sectional cohort	2812	77.9	52.7/47.3	Cognition	Processing speed- Was assessed by the Trail Making Test part A, Cognitive flexibility- Was assessed by the Trail Making Test part B	Self-report on a 3-point Likert-type scale whether their cognitive and sensory abilities across the age tranches in old age were not moderated by general cognitive ability, educational level, nor general health status (all ps>0.225)	No. Present data do not support the view of a generally increased relation of cognitive and sensory abilities in old age.	Weak
2015				range 65–101			Verbal abilities- Was assessed by administering the Mill Hill vocabulary scale	current eyesight allowed them to read a newspaper by choosing one of the following 'yes, without difficulties,' 'yes, but with difficulties'; or 'no'		
Soto-Perez-de-Celis et al <sup>62</sup>	US	Cross-sectional case control	750	72	56/44	Cognition	Score of 11 in the Blessed Orientation-Memory-Concentration test.	Self-reported visual impairment based on their rating of their eyesight (with glasses) as "excellent," "good," "fair," "poor," "totally blind"	OR for VI and: IADL Dependence: 1.9 (1.2–3.2)<0.01, Poor Physical Function: 1.9 (1.1–3.3) .03, and Possible CI: 1.9 (0.7–4.8) 0.20, Depression: 2.5 (1.4–4.3)<0.01, Anxiety: 1.4 (0.8–2.4) .19. OR for DSI and: IADL dep 2.8 (1.5–5.3)<0.01	Weak
2018				Range 65–94						
									Poor Physical Function: 1.7 (0.9–3.4) .10, Possible CI: 3.2 (1.3 to 8.1) .01, Depression: 2.5 (1.3–4.8)<0.01, Anxiety: 2.3 (1.2–4.2) .01	

Continued

**Table 5** Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	CI: type of measurement/evaluation	VI: Type of measurement/evaluation	Point estimates and analysis performed	Summary of association	Quality of study	
Maharani <i>et al</i> <sup>77</sup>	HRS, ELSA, SHARE	Cross-sectional cohort	13 123 hours	67.8/32.2	41.8/58.2	Cognition	Episodic memory: In all surveys, the interviewer reads a list of 10 common nouns to the respondent	In HRS and ELSA, self-reported vision quality was collected in all seven waves using the question: 'Is your eyesight (using glasses or corrective lens as usual) excellent (1), very good (2), good (3), fair (4) or poor (5)?.'	HRS: Single impairment $\beta$ -0.15 (0.02)*	Yes. Those with sensory impairment are at a greater risk of developing cognitive impairment	Weak	
2018			11 417 ELSA	64.8/35.2	45.6/54.4		then asks the respondent to recall as many words as possible from the list in any order twice: immediately after the respondent heard the complete list (immediate recall) and at the end of the cognitive function module (delayed recall).	In SHARE, we used the two self-reported measures of visual function	Dual impairment $\beta$ -0.25 (0.04)*	and may show a faster trajectory of cognitive decline that those without sensory impairment		
			21 265 SHARE	64.8/35.2	45.5/54.5			that are present in all waves: distance eyesight and reading eyesight.	ELSA: Single impairment $\beta$ -0.14 (0.02)*  Dual impairment $\beta$ -0.35 (0.05)*			
									SHARE: Single impairment $\beta$ -0.26 (0.01)*  Dual impairment $\beta$ -0.68 (0.03)*			
Jefferis <i>et al</i> <sup>83</sup>	UK (Newcastle 85+study)	Cross-sectional cohort	839	no info	no info	Cognition	sMMSE	Data collection of general practice records on whether the participant was	Median (inter-quartile range) sMMSE scores were 25 (22-29) for SI and 28 (25-29) for non-SI participants (p=0.006).	It is important to consider the possibility of vision impairment in older	Weak	
2012							registered as blind (severely sight impaired) or partially sighted (sight impaired), or neither (by a consultant ophthalmologist)		SI participants had lower subscale scores on tasks requiring vision (p<0.001 for each) but also for some subscale	people when carrying out the MMSE and to consider using the MMblind.		

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Table 5 Continued

Author and (Study name if applicable)	Country	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	CI: type of measurement/evaluation	VI: Type of measurement/evaluation	Point estimates and analysis performed	Summary of association	Quality of study
MacDonald SWS <i>et al</i> <sup>64</sup>	Canada	Longitudinal, cohort	408	NIC 75.44	NIC 35.5/64.5	Cognition	Episodic memory- The word recall task (Hultsch, Hertzog, & Dixon, 1990) was used to assess episodic memory	Acuity: Vision-Binocular-corrected distance VA was measured at three metres using the Snellen chart.	Age-related change in sensory function.	Decline in VA was also a notable predictor	Weak
2018				SA-MCI :76.71	SA-MCI: 40.5/59.5		Inductive reasoning- The Letter Series test (Thurstone, 1962) was used to assess inductive reasoning.	Snellen acuity fractions ranged from 0.20 to 1.	Variables, Intercept $\gamma$ 00, Slope $\gamma$ 10, SE Slope, p	of being classified as SA-MCI or MA-MCI	
				MA-MCI: 75.68	MA-MCI: 48.2/51.8		Perceptual speed-The WAIS-R Digit Symbol Substitution task (Wechsler, 1958) was used to assess perceptual processing speed		Olfaction: 7.05-0.043, 0.008,<0.001		
							Verbal fluency-The Controlled Associations test from the ETS kit of factor-referenced cognitive tests (Ekstrom, French, Harman, & Dermen, 1976) was used to assess verbal fluency.		Distance Vision: 0.946,-0.004, 0.001,<0.001		
							Vocabulary-A recognition vocabulary test, combining three 18-item multiple-choice tests from the ETS kit of factor-referenced cognitive tests (Ekstrom <i>et al.</i> , 1976), was used to assess vocabulary		Audition: 32.84, 0.796, 0.059,<0.001, $\gamma$ 00=Average sensory function centred at the grand mean of age (74.17 years; SD=9.20); $\gamma$ 10=slope reflecting the average rate of linear change per additional year of age		
Miyata <i>et al</i> <sup>67</sup>	Japan (HEIJO-KYO Cohort)	Cross-sectional cohort	945	71.7	46.8/53.2	Cognition	MMSE	Acuity: using the Landolt ring chart. A better value of LogMAR was used for analysis.	Logistic Regression Analysis for the Association Between Ocular Status and Cognitive Impairment	Yes. ORs for cognitive impairment were significantly lower in the pseudophakic group than in the phakic group	Weak
2015							Questionnaire asking participants about pseudophakia/phakia		Age-adjusted, Multivariate OR for cognitive impairment: Model 1, Model2, Model 3		
							Confirmed by ophthalmologist using slit-lamp		Phakia (no previous cataract surgery): 1.00 (ref), 1.00 (ref), 1.00 (ref), 1.00 (ref), 1.00 (ref)		
									Pseudophakia (previous cataract surgery): 0.66 (0.45, 0.98), 0.66 (0.44, 0.98), 0.64 (0.43, 0.95), 0.64 (0.43, 0.96)		
									p: 0.038, 0.039, 0.026, 0.031		

Continued

**Table 5** Continued

Author and title	Country (Study name if applicable)	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ci: type of measurement/ evaluation	Vi: Type of measurement/ evaluation	Point estimates and analysis performed	Summary of association	Quality of study
Mine <i>et al</i> <sup>61</sup>	Japan (Fujiwara-kyo Study)	Cross-sectional cohort	2818	76.3	52.7/47.3	Cognition	MMSE; analysed the MMSE excluding the following vision-related five items: "naming two objects," "following a 3-step command," "reading and following instruction," "writing a sentence," and "visual reconstruction" and the maximum score for this was 22 points.	Acuity: Landolt ring chart at 5 m (converted to logMAR)(mild visual impairments >0.2 logMAR)	Associations between VA and Cognitive Impairment (multivariate regression model):	Yes. Subjects with mild visual impairments had 2.4 times higher odds of	Weak
2016									OR (CI) for different BCVA in the better eye (logMAR) groups: ≤0: 1.0 (reference), 0-0.1: 1.8 (1.2-2.7) p 0.007 0.1-0.2: 1.7 (0.8-3.5) p 0.187 >0.2: 3.3 (2.1-5.4) p 0.005	having cognitive impairment than those without visual impairment	
Steffi G. Riedel-Heller <sup>62</sup>	Germany (LEILA 75-and AgeCoDe)	Longitudinal, cohort	3199	79.3	34.7/65.3	Cognition	No data	Self-report at baseline	Vision impairment (HR 1.19, 95% CI 1.01 to 1.42, p=0.043) and the combination of both (HR 1.47, 95% CI 1.18 to 1.83, p=0.001)	Yes. Vision impairment is independently associated with incident dementia. Looking at the combination we could show that individuals suffering from both are at highest risk for developing dementia	Weak
2019									on dementia incidence adjusted for baseline gender, age, education, marital status, depression, diabetes and cardiovascular risks (smoking, hypertension)		
Asri Maharani <sup>109</sup>	US (HRS)	Longitudinal cohort	19 618	57.8	44.6/55.4	Cognition	Telephone interview for Cognitive Status (TICS): include episodic memory, serial of 7 subtraction, and counting backward tests	Self report: "Is your eyesight (using glasses or corrective lens as usual) excellent (1), very good (2), good (3), fair (4) or poor (5)?"	VI and risk of possible CIND HR 1.351 (1.267, 1.441) p<0.001 when compared with no VI. VI and risk of probable	Yes. Self-assessed sensory impairment is independently associated with cognitive decline and incident possible CIND and probable dementia	Weak
2019	Waves 3 (1996) to 12 (2014)					categorised those scoring 0 to 6 points on the 27-point TICS scale as having probable dementia	Further categorised sensory impairment in the simultaneous model into no impairment.		dementia HR 1.255 (1.074, 1.466) p=0.004 when compared with no VI		

Continued



**Table 5** Continued

Author and (Study name if applicable)	Country	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ci: type of measurement/ evaluation	Vi: Type of measurement/ evaluation	Point estimates and Summary of analysis performed association	Quality of study
Phillip Hwang <i>et al</i> <sup>112</sup>	US (GEM study)	Longitudinal cohort	2827	NA	NA	Cognition	Incident dementia over 7 years of follow-up was based on a clinical diagnosis of dementia using DSM-IV criteria	Self-report	For all-cause dementia, the adjusted HR was 1.27 (95% CI: 1.02, 1.59) for single sensory visual or hearing impairment, all-cause dementia and AD	Weak
2019							Alzheimer's disease was determined using NINCDS-ADRD criteria		and 1.70 (95% CI: 1.18, 2.45) for dual visual and hearing impairment, compared with no sensory impairment	
Virginie Nael <sup>113</sup>	France (The Three-City-Alienor Cohort)	Longitudinal cohort	7460	NA	NA	Cognition	four cognitive domains (global cognition (MMSE and MMSE-blind), verbal fluency (IST), executive function (TMT) and visuospatial abilities (BVRT)) were assessed up to 6 times over 12 years of follow-up	At baseline, near VI was measured using the Parinaud scale	Participants with near VI and distance VF loss had lower baseline performances in verbal fluency,	Weak
2019							at a standardised reading distance of 33 cm and distance VF loss was	global cognition, executive function and visuospatial abilities. Regarding changes over time,	visual loss had lower baseline performances in several cognitive tests	
M.Q.LJ <sup>116</sup>	China	Cross-sectional cohort	10 116	NA	NA	Vision	AD8 questionnaire	No info	Risk factor for cognitive impairment: VA loss 1.383 (1.188, 1.610)	Weak
2019							self-reported, defined as inability or difficulty in recognising a familiar face at 4 m		MMSE where participants with mild near VI exhibited a faster cognitive decline (b=-0.02, p=0.04)	
Melanie Varin <sup>118</sup>	US	Cross-sectional cohort	365	AMD:83.1	32.4/67.6	Cognition	six cognitive tests orally; the 1 min verbal fluency test (letter and category versions)	Binocular VA was measured using the ETDRS chart at 2 m	People with glaucoma showed lower scores on three cognitive tests than the group with normal vision:	Weak

Continued



Table 5 Continued

Author and (Study name if applicable)	Country	Study design	No of participants	Mean age (M/F)	Gender (M/F)	Outcome variable	Ci: type of measurement/evaluation	Vi: Type of measurement/evaluation	Point estimates and analysis performed	Summary of association	Quality of study
2019				Glaucoma:78.1	43.7/56.3		the digit span test (forward and backward versions), and the logical memory test (immediate and 30 min delayed recall)	VF measured using HFA, CS measured using the Pelli Robson chart at 1 m	the digit span forward and backward tests (b=-0.8, 95% CI -1.5 to -0.2 and b=-0.7, 95% CI -1.3 to -0.1, respectively)	People with glaucoma showed lower scores on cognitive tests	
Peiyuan Qiu <sup>123</sup>	China (Chinese Longitudinal Healthy Longevity Survey)	Longitudinal cohort	3859	74.5	48.7/51.3	Cognition	Chinese MMSE (C-MMSE)	Self-report	NA	Yes, visual impairment was a risk factor for cognitive decline	Weak
2019											

AMD, age-related macular degeneration; CS, contrast sensitivity; DSI, dual sensory impairment; ELSA, English Longitudinal Study of Aging; HRS, Health and Retirement Study; MMSE, Mini-Mental State Examination; MoCA, Montreal-Cognitive Assessment; MPOD, macular pigment optical density; sMMSE, standardised MMSE; VA, visual acuity; VF, visual field.

scores indicating more severe cognitive impairment. A score of 24 is often used as a threshold to define ‘normal’ cognitive function.<sup>123</sup> The MMSE has been found to be a valid and reliable tool as assessed by many studies.<sup>123 124</sup> Several studies used self-report, diagnosis codes and data from existing records to define cognitive status. Similarly, visual function was also assessed by various methods including self-report. While VA was assessed most commonly, there was significant variation in the charts and tools used to assess it. The parameters used to define cognitive decline and VI may have impacted results across and within studies.

Our systematic review has found that there is a strong consensus in the literature that VI is associated with cognitive decline, cognitive impairment or dementia. Two hypotheses may help explain this association. The first one is that a common pathological process (eg, vascular disease) might be responsible for both the sensory and cognitive impairment in older adults. The second one is that by increasing cognitive load, sensory impairments such as VI might cause cognitive impairment.<sup>125</sup> Literature also suggests that vision rehabilitation in the form of cataract surgery slows the rate of cognitive decline, and therefore, early vision interventions could potentially reduce risk of dementia.<sup>126</sup>

Our review evaluated bias for all of the 110 included studies. The majority of studies included in our review were cross-sectional, and according to EPHPP guidelines, cross-sectional studies can only receive a low or moderate rating in the bias assessment. Cross-sectional studies are also prone to selection bias, thus yielding estimates that may not reflect true associations in the target population. Studies receiving a strong rating were all longitudinal. However, the tool penalises longitudinal studies that lose >40% of participants due to dropouts/withdrawals. This may, perhaps unfairly, affect longer longitudinal studies to a greater extent since they collect data over many years and can have more drop-outs due to deaths since they are conducted among older adults.

This review has several important implications. First, it highlights the need for standardised methods to assess and define both visual and cognitive function that will aid future research on these emerging public health issues. Second, it brings into focus the consistent association of VI with cognitive impairment in older adults and the need to better understand the mechanisms underlying this relationship. Third, as the longitudinal results support the sensory consequence theory, and suggest that VI may be a risk factor for cognitive decline, this points to a need for formulating preventive measures and vision rehabilitation models, such as prescription glasses, cataract surgery, low vision rehabilitation, etc, that could have the potential to improve overall health and well-being of older adults.

### Limitations

Given the large number of studies included in this review and the heterogeneity of measures used to assess



the outcome, it was not possible to compare and meta-analyse results across studies. Although 35 longitudinal studies found a positive association between VI and cognitive decline, we cannot establish temporality between this relationship due to the heterogeneous nature of the studies. The studies included diverse populations, with different disease processes, and variation in definitions of both cognitive and VI. There is also potential bias associated with studies that used different protocols for cognitive and sensory measurements. The MMSE, which was the most commonly used assessment method for testing cognition is sensory dependent and therefore one can argue that the results may be confounded with VI.<sup>127</sup> Further studies should examine the impact of using vision independent cognitive tests on the vision–cognition relationship. Our review examined all cause VI and dementia, and further study is needed to examine the vision–cognition relationship by dementia subtype and by different vision pathology. However, despite the heterogeneity in studies and assessment methods, we synthesised the evidence qualitatively and by taking into account study quality assessed using a validated tool. While our search strategy was robust, it may have been limited by the exclusion of studies that were not published in English.

## CONCLUSION

The number of older adults with VI and dementia is increasing globally, and therefore, the elucidation of the relationship between vision and cognition is of particular public health importance. This systematic review found that the positive association of VI with cognitive decline, cognitive impairment or dementia is largely consistent across studies using different measures of vision and cognition, as well as between countries and cohorts. This overall agreement in the literature suggests that poor visual and cognitive function are associated, and that additional research is now needed to move beyond documenting these associations. The focus of this area of research should now turn to identifying the factors and strategies that mediate the vision–cognition relationship and identifying potential interventions, such as vision rehabilitation models and strategies tailored to people with VI, that may mitigate the cognitive implications of VI.

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