



# Cochrane and systematic reviews



#### **Jenny McCleery**

Joint Co-ordinating editor Cochrane dementia and cognitive improvement group





# What is a systematic review?

## Archie Cochrane (1909-1988)

In medicine, we don't have evidence for most of what we do

Some things we do are probably harmful







# What is a systematic review?

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In medicine, we don't have evidence for most of what we do

Some things we do are probably harmful

We need "a critical summary, adapted periodically, of all ... relevant, randomised controlled trials"









Started in perinatal medicine in 1980s

- 1. Computerised register of RCTs
- 2. Methods to combine data from different trials to create overall estimates of effects
- 3. An international collaboration to prepare and maintain the "critical summaries" (systematic reviews) of the RCTs in the register



















# Cochrane Methods









## Writing (or reading) a systematic review – start with a question

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## What do I want to know?

Should I be offering aducanumab to my patients with dementia?





# **Structure the question: PICO for interventions**

**P** – Participants (Patients with dementia? With dementia due to AD? With mild dementia due to AD? With mild dementia due to AD and positive amyloid markers?)

I – Intervention (Low or high dose aducanumab?)

**C** – Comparison (Placebo? Placebo and a cholinesterase inhibitor?)

**O** – Outcomes (Cognition? Function? Cognition and function combined? Which scales? Which harms?)



# **Process of a systematic review**

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# Literature search

On 27 Jan 2020, PubMed included >30 million citations & abstracts

In the 10 years to 31 Dec 2019, an average of nearly 1 million new records were added to PubMed each year

Cochrane's Central Register of Controlled Trials is a highly concentrated source of reports of RCTs

Cochrane is pioneering the use of 'crowd' methods and machinelearning to identify RCTs





# **Quality assessment**

An essential part of a good systematic review

Cochrane Risk of Bias tool – risk of bias in individual RCTs

- Selection bias (random sequence generation, allocation concealment)
- Performance bias (blinding of participants and study personnel)
- Detection bias (blinding of outcome assessors)
- Attrition bias (incomplete outcome data)
- Reporting bias (selective outcome reporting)
- Other biases

Different quality assessment tools for other study types (e.g. QUADAS-2 for diagnostic test accuracy studies to assess risk of bias and external validity)





# Risk of bias assessment: Pharmacotherapies for sleep disturbances in dementia



Camargos 2014 Dowling 2008 Herring 2020 Morales-Delgado 2018 NCT00325728 NCT03001557 Serfaty 2002 Singer 2003 Wade 2014





# **Data synthesis**

Meta-analysis is a common but not essential part of a systematic review

## Systematic review

Rigorous, scientific approach to identifying, appraising, synthesising and interpreting information Meta-analysis Statistical approach to synthesising information to obtain a summary estimate of effect



# **Interpreting results** *How confident can I be that this review gives me the right answer to my question?*

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GRADE – overall certainty of the evidence related to each outcome

- Risk of bias in included studies
- Imprecision of results
- Inconsistency between studies
- Indirectness in relation to question
- Publication bias

Critical for interpretation of results





# **Interpreting results**

• Endovascular thrombectomy for acute ischaemic stroke:

"Treatment increased the chance of achieving a good functional outcome, defined as a modified Rankin Scale score of 0 to 2: risk ratio (RR) 1.50 (95% confidence interval (CI) 1.37 to 1.63; 3715 participants, 18 RCTs; *high-certainty evidence*)."

[Roaldsen et al. Endovascular thrombectomy and intra-arterial interventions for acute ischaemic stroke. Cochrane Database of Systematic Reviews 2021, Issue 6. Art. No.: CD007574. DOI: 10.1002/14651858.CD007574.pub3. Accessed 27 June 2021]

- Discontinuing cholinesterase inhibitors:
- "Compared to continuing cholinesterase inhibitors, discontinuing treatment may be associated with worse cognitive function in the short term (standardised mean difference (SMD) -0.42, 95% confidence interval (CI) -0.64 to -0.21; 4 studies; *low certainty*), but the effect in the medium term is very uncertain (SMD -0.40, 95% CI -0.87 to 0.07; 3 studies; *very low certainty*).

• [Parsons et al. Withdrawal or continuation of cholinesterase inhibitors or memantine or both, in people with dementia. CochraneDatabase of Systematic Reviews 2021, Issue 2. Art. No.: CD009081. DOI: 10.1002/14651858.CD009081.pub2. Accessed 27 June 2021]





# A warning .....





### John Ionnadis (2016) – "The Mass Production of Redundant, Misleading, and Conflicted Systematic Reviews and Meta-analyses"

The Millbank Quarterly, 94 (3), 485-514

Annual publications between 1991 and 2014 increased 2,728% for systematic reviews and 2,635% for meta-analyses versus only 153% for all PubMed-indexed items. Currently, probably more systematic reviews of trials than new randomized trials are published annually.

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Most topics addressed by meta-analyses of randomized trials have overlapping, redundant metaanalyses ..... Some fields produce massive numbers of meta-analyses; for example, 185 meta-analyses of antidepressants for depression were published between 2007 and 2014. These meta-analyses are often produced either by industry employees or by authors with industry ties and results are aligned with sponsor interests.

Many ... meta-analyses have serious flaws. Of the remaining, most have weak or insufficient evidence to inform decision making. Few systematic reviews and meta-analyses are both non-misleading and useful.

<u>Conclusions</u>: The production of systematic reviews and meta-analyses has reached epidemic proportions. Possibly, the large majority of produced systematic reviews and meta-analyses are unnecessary, misleading, and/or conflicted.





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➤Conflicted?

No, robust conflict of interest policy.

➤ Misleading?

No, we hope not, rigorous methods and quality control.

➤Unnecessary?

Maybe some. Increasing emphasis on prioritisation.





# **Cochrane Dementia Greatest Hits** (some of them)

# Terry Quinn,

Co-ordinating Editor Cochrane Dementia @CochraneDCIG

@DrTerryQuinn





# **COCHRANE** 7,500 reviews

- 14 languages
- 53 review groups
- 30,000 volunteers
- 7.89 CDSR Impact factor 2019



#### Figure 3: Average number of Full Text Accesses received by Cochrane Review Groups in 2019

#### Figure 5: Average number of guideline cites to reviews (published anytime) for each Cochrane Review Group







#### Table 9: Top 10 Altmetric scores for reviews published in 2019

Score	Review title	CD Number	Publication date	CRG	CCA number	B	T	N	F	W	М
774	Exercise for preventing falls in older people living in the community	CD012424.pub2	Jan-2019	Bone, Joint and Muscle Trauma Group	2469	6	949	29	17	0	355
641	General health checks in adults for reducing morbidity and mortality from disease	CD009009.pub3	Jan-2019	Effective Practice and Organisation of Care Group	1598	4	1058	3	11	0	105
420	Constraint-induced movement therapy in children with unilateral cerebral palsy	CD004149.pub3	Apr-2019	Developmental, Psychosocial and Learning Problems Group	-	0	99	44	1	1	141
355	Environmental interventions to reduce the consumption of sugar-sweetened beverages and their effects on health	CD012292.pub2	Jun-2019	Public Health Group	-	4	246	26	10	1	269
307	Different doses, durations and modes of delivery of nicotine replacement therapy for smoking cessation	CD013308	Apr-2019	Tobacco Addiction Group	2626	8	226	28	8	2	91
304	Incentives for smoking cessation	CD004307.pub6	Jul-2019	Tobacco Addiction Group	1533	3	165	30	2	1	194
290	Paracetamol versus placebo for knee and hip osteoarthritis	CD013273	Feb-2019	Musculoskeletal Group	2520	2	467	2	7	1	110
224	Mediterranean-style diet for the primary and secondary prevention of cardiovascular disease	CD009825.pub3	Mar-2019	Heart Group	2536	4	347	2	5	2	224
211	Memantine for dementia	CD003154.pub6	Mar-2019	Dementia and Cognitive Improvement Group	2645	0	367	1	2	2	403
147	C-reactive protein for diagnosing late-onset infection in newborn infants	CD012126.pub2	Jan-2019	Neonatal Group	-	1	291	0	6	0	64

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B=Bloggers T=Tweeters N=News outlets F=Facebook mentions W=Wikipedia pages M=Mendeley readers





# **Reviews of drugs**







Cochrane Database of Systematic Reviews

#### Tacrine for Alzheimer's disease (Review)

Qizilbash N, Birks J, López Arrieta J, Lewington S, Szeto S

Qizilbash N, Birks J, López Arrieta J, Lewington S, Szeto S. Tacrine for Alzheimer's disease. *Cochrane Database of Systematic Reviews* 1999, Issue 1. Art. No.: CD000202. DOI: 10.1002/14651858.CD000202.





Objectives: To determine the clinical efficacy of tacrine for the symptoms of Alzheimer's disease.

**Search strategy:** The Cochrane Dementia Group Register of Clinical Trials was searched using the terms 'tacrine', 'tetrahydroaminoacridine' and 'THA' (see the Group's search strategy for full details).

**Selection criteria:** All unconfounded, double-blind, randomized trials in which treatment with tacrine was administered for more than a day and compared to placebo in patients with dementia of the Alzheimer's type.

**Data collection and analysis:** Data were extracted independently by two reviewers, pooled if appropriate and possible, and the pooled odds ratios (95%CI) or the average differences (95%CI) were estimated. Where possible, intention-to-treat data were used.

Main results: This review produced no clear results. The results were compatible with tacrine producing improvement, no change or even harm for those with Alzheimer's disease. It was not possible to use many of the published results in a combined analysis. For measures of overall clinical improvement, the intention-to-treat analyses failed to detect any difference between tacrine and placebo (OR 0.87; 95%Cl 0.61 - 1.23). Behavioural disturbance, as measured by the Alzheimer's Disease Assessment Scale-noncognitive, failed to detect any difference between tacrine and placebo (SMD -0.04; 95%CI -0.52 - 0.43). For cognition function, the effect of tacrine was not statistically significantly different from placebo for the MiniMental State Examination score (0-30; high = good) (SMD 0.14; 95%CI -0.02 - 0.30) and was barely statistically significantly in favour of treatment for the Alzheimer's Disease Assessment Scale-cognitive scale (SMD -0.22; 95%CI -0.32 - -0.13). Adverse events were not reported in a systematic way in the different trials, making formal comparison difficult. Raised serum liver enzymes was the major reason for withdrawal. The odds ratio for withdrawal due to an adverse event was significantly different from one, the control group experienced fewer events (OR 5.7; 95%CI 4.1-7.9). Gastrointestinal side effects (diarrhoea, anorexia, dyspepsia and abdominal pain) were the other major cause of adverse events and for withdrawal, and the odds ratio for withdrawal was also significantly different from one in favour of the control group (OR 3.8; 95%Cl 2.8-5.1). No deaths were reported in any of the studies during the trial period, up to six months.

#### Clear objective(s) - protocol

Search strategy

#### **Paired reviewers**

Evidence synthesis Pre-specified outcomes Meta-analysis, benefits/harms

> No assessment of bias No GRADE









Cochrane Database of Systematic Reviews

### Memantine for dementia (Review)

McShane R, Westby MJ, Roberts E, Minakaran N, Schneider L, Farrimond LE, Maayan N, Ware J, Debarros J



#### Figure 1. Study flow diagram of studies identified



	Random sequence generation (selection bias)	Allocation concealment (selection bias)	Blinding (performance bias and detection bias)	Incomplete outcome data (attrition bias): Primary Outcomes	Incomplete outcome data (attrition bias): Safety Data	Selective reporting (reporting bias)	Other bias			
Aarsland 2009	•	•	•	?	•	?	•			
Asada 2011 (MA3301)	?	?	•	•	•	?	?			
Asada 2011a (IE3501)	?	?	•	•	•	?	?			
Ashford 2011 (95722)	•	•	•	•		?	?			
Std. Mean Difference					w	elght	s	td. Mean Diff	erence	

Study or subgroup	Memantine		P	lacebo	Std. Mean Difference	Weight	Std. Mean Difference
	N	Mean(SD)	N	Mean(SD)	Fixed, 95% CI		Fixed, 95% CI
Asada 2011a (IE3501)	190	4.5 (1.1)	177	4.6 (1)	-+	13.33%	-0.11[-0.31,0.1]
Bakchine 2008 (99679) SG	145	4.2 (1.1)	65	4.4 (1.2)		6.52%	-0.14[-0.43,0.15]
Dysken 2014 SG	50	0 (0)	63	0 (0)			Not estimable
Forest 2006 (MD-22)	131	3.6 (1.4)	129	3.8 (1.3)		9.44%	-0.15[-0.4,0.09]
Grossberg 2008 (MD-50)	269	3.8 (1.1)	272	4.1 (1.2)	_ <b></b>	19.53%	-0.26[-0.43,-0.09]
Homma 2007 (IE2101)	84	4.4 (1.6)	87	4.7 (1.3)		6.19%	-0.22[-0.52,0.08]
Howard 2012 (DOMINO-AD)	77	0 (0)	66	0 (0)			Not estimable
Peskind 2004 (MD-10) SG	106	4.4 (1)	116	4.7 (1.1)	<b>-</b>	7.97%	-0.31[-0.57,-0.04]
Porsteinsson 2008(MD-12)S	135	4.5 (1)	125	4.5 (1)		9.45%	-0.04[-0.28,0.2]
Relsberg 2003 (9605)	97	4.4 (1.1)	84	4.7 (1.1)		6.5%	-0.27[-0.56,0.03]
Tarlot 2004 (MD-02)	172	4.4 (1.1)	152	4.6 (1.1)		11.67%	-0.24[-0.46,-0.02]
van Dyck 2007 (MD-01)	134	4.3 (1.1)	127	4.6 (1)	<b>-</b>	9.4%	-0.28[-0.53,-0.04]
Wang 2013	13	0 (0)	13	0 (0)			Not estimable
Total ***	1603		1476		•	100%	-0.2[-0.28,-0.13]
Heterogeneity: Tau <sup>2</sup> =0; Chl <sup>2</sup> =4.7,	df=9(P=0.86	); I <sup>2</sup> =0%					
Test for overall effect: Z=5.32(P<	0.0001)						
			Favou	rs memantine -1	-0.5 0 0.5	1 Favours pl	acebo





#### SUMMARY OF FINDINGS

#### Summary of findings for the main comparison. Moderate-to-severe AD, six to seven months

Memantine 20 mg or equivalent compared to placebo for moderate-to-severe Alzheimer's disease (AD) 24- to 30-week data. OC

Population: Alzheimer's disease (AD), moderate-to-severe Intervention: memantine 20 mg or equivalent

Comparison: placebo

Continuous out- comes	Score with placebo (median)	Mean improvement in change score be- tween memantine and placebo	SMD (95% CI) meta- analysis findings	№ of partici- pants (stud- ies)	Certain- ty of the evi- dence (GRADE)	Comments
Clinical Global (CIBIC+)	Median CIBIC+ score was 4.60 <sup>3</sup>	MD: 0.21 (0.14 to 0.30)	-0.20 (-0.28 to	2797 (10	⊕⊕⊕⊕ HIGH	SMD as a negative outcome
7-point Likert scale	(i.e. deterioration with time)		-0.13)	RCTs)		(Analysis 1.1)
						Converted to CIBIC+ scale; median SD(pooled) = 1.06.
Cognitive Function	Median SIB score at baseline: 75.2.	MD: 3.11 (2.42 to 3.92)	-0.27	3337	0000	SMD as a negative outcome (Analysis 1.2).
(SIB) 100-point scale	Median change from baseline (posi- tive scale): -2.4 <sup>4</sup>		(-0.34 to -0.21)	(13 RCTs)	HIGH	Converted to SIB scale (and scale direction inverted); median SD (pooled) = 11.53.
	(i.e. deterioration with time)					
Functional perfor- mance on activities	Median ADCS-ADL19 score at base- line: 33.2	MD: 1.09 (0.62 to 1.64)	-0.16 (-0.24 to	2687 (11 PCTs)	0000 HIGH <sup>1</sup>	SMD for decline in ADL (a negative out- come)
CS-ADL19	Median change from baseline (pos-		-0.03)	KCT3)		(Analysis 1.3).
54-point scale	itive scale): -2.8 > (i.e. deterioration with time)					Converted to ADCS-ADL19 scale (and scale direction inverted); median SD(pooled) = 6.84.

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### Coronavirus (COVID-19) resources

### Memantine as a treatment for dementia

Published:

20 March 2019

Our evidence

#### Authors:

McShane R, Westby MJ, Roberts E, Minakaran N, Schneider L, Farrimond LE, Maayan N, Ware J, Debarros J

Primary Review Group: Dementia and Cognitive Improvement Group

#### **Review question**

We reviewed the evidence on memantine, which is one of the main drugs for treating people with dementia. We wanted to find out if memantine can slow down the course of dementia and if it is harmful in any way. We also wanted to know if adding memantine to other dementia drugs gives an extra effect.

#### Background

The commonest type of dementia is Alzheimer's disease (AD), followed by vascular dementia. About one or two people in 100 have AD at age 65, and this rate doubles every five years. Dementia involves loss of memory, difficulty thinking and often changes in mood and behaviour.



Who is talking about this article?

Video: Systematic reviews explained

How our health evidence can help you

Read in different languages







Cochrane Database of Systematic Reviews

Cholinesterase inhibitors for vascular dementia and other vascular cognitive impairments: a network meta-analysis (Review)

Battle CE, Abdul-Rahim AH, Shenkin SD, Hewitt J, Quinn TJ



### **Network plot: Cognition**



Figure 5. Forest plot (Bayesian model) network meta-analysis results: Cognition.





Cochrane Database of Systematic Reviews

Withdrawal versus continuation of long-term antipsychotic drug use for behavioural and psychological symptoms in older people with dementia (Review)

Van Leeuwen E, Petrovic M, van Driel ML, De Sutter AIM, Vander Stichele R, Declercq T, Christiaens T



Cochrane Database of Systematic Reviews

Antihypertensive withdrawal for the prevention of cognitive decline (Review)

Jongstra S, Harrison JK, Quinn TJ, Richard E





# **Non-drug Reviews**







Cochrane Database of Systematic Reviews

### Aromatherapy for dementia (Review)

Ball EL, Owen-Booth B, Gray A, Shenkin SD, Hewitt J, McCleery J



**UK Cochr**a

#### Aromatherapy versus control (placebo aromatherapy / no intervention) for dementia

#### Patient or population: Dementia

Setting: Care facilities or hospital wards Intervention: Aromatherapy Comparison: Control (placebo aromatherapy / no intervention)

Outcomes	Impact	№ of participants (studies)	Certainty of the evidence (GRADE)
Agitation assessed with: CMAI, PAS, in- dividual study assessment tools follow up: range 1 to 12 weeks	5 trials provided either no usable data or data in which our confidence was very low. Of the remaining 5 trials, 4 reported no statistically significant effect on agitation and 1 reported a significant benefit.	593 (10 RCTs)	⊕ooo VERY LOW <sup>1234</sup>
Overall behavioural and psy- chological symptoms assessed with: NPI follow up: range 2 to 12 weeks	3 trials provided either no useable data or data in which our confidence was very low. Of the remain- ing 5 trials, 4 trials reported a significant reduction in overall behavioural and psychological symptoms and 1 trial did not find a significant effect of aromathera- py.	346 (8 RCTs)	⊕ooo VERY LOW <sup>1345</sup>
Adverse effects follow up: range 1 to 12 weeks	Adverse effects were reported in only 4 of 12 trials. None reported any adverse effects.	206 (4 RCTs)	⊕©©© VERY LOW <sup>34</sup>
Quality of life assessed with: Blau Quality of Life, Dementia Care Map- ping follow up: range 4 to 12 weeks	1 trial reported a significant beneficial effect of aro- matherapy on quality of life. The other trial did not find any significant effect of aromatherapy on quality of life.	134 (2 RCTs)	⊕ooo VERY LOW 13467
Mood assessed with: CSDD-C, PG- CARS follow up: range 1 to 9 weeks	1 trial reported no significant effect of aromatherapy on mood. The other trial reported a statistically signif- icant beneficial effect of aromatherapy on depressive symptoms.	120 (2 RCTs)	⊕©©© VERY LOW <sup>1348</sup>
Sleep	1 trial provided no useable data.	21 (1 RCT)	-
Activities of daily living assessed with: Barthel Index for Activities of Daily Living, follow up: 12 weeks	1 trial provided no useable data. 1 trial found no sig- nificant effect of aromatherapy on activities of daily living.	91 (2 RCTs)	⊕⊙⊝© VERY LOW <sup>3 4 10</sup>







Cochrane Database of Systematic Reviews

# Interventions for preventing delirium in hospitalised non-ICU patients (Review)

Siddiqi N, Harrison JK, Clegg A, Teale EA, Young J, Taylor J, Simpkins SA

Summary of findings for the main comparison. A multi-component delirium prevention intervention compared to usual care for hospitalised non-ICU patients

Multi-component delirium prevention intervention compared to usual care for hospitalised non-ICU patients

Outcomes	Illustrative comparative	risks* (95% CI)	Relative	No of Par-	Quality of the	Com-
	Assumed risk	Corresponding risk	(95% CI)	(studies)	(GRADE)	ments
		A multi-component delirium prevention intervention	,			
Incidence of deliri- um validated instru- ments <sup>1</sup>	<b>209 per 1000</b> <sup>2</sup>	<b>144 per 1000</b> (123 to 172)	<b>RR 0.69</b> (0.59 to 0.81)	1950 (7 studies <sup>3</sup> )	⊕⊕⊕© moderate 4,5,6	
Duration of deliri- um (days)	The mean duration of delirium in the control groups ranged from <b>2.1 to 10.2 days</b>	The mean duration of delirium in the intervention groups was <b>1.16 days shorter</b> (2.96 shorter to 0.64 longer)		244 (4 studies)	⊕©©© <b>very low</b> 4,6,7,8,9	
Severity of deliri- um DRS-R-98 and CAM- S <sup>10</sup>		The standardised mean severity of delirium in the interven- tion groups was <b>1.04 standard deviations lower</b> (1.65 to 0.43 lower) <sup>11</sup>		67 (2 studies)	⊕⊕©© low <sup>4,12</sup>	
<b>Length of admis- sion</b> Days	The mean length of ad- mission in the control groups ranged from <b>5 to 38 days</b>	The mean length of admission in the intervention groups was <b>0.01 days longer</b> (0.48 days shorter to 0.51 days longer)		1920 (6 studies)	⊕⊕⊕⊙ moderate 4,6,7	

Intervention: A multi-component delirium prevention intervention versus usual care





# **Diagnosis Reviews**







Cochrane Database of Systematic Reviews

# AD-8 for detection of dementia across a variety of healthcare settings (Review)

Hendry K, Green C, McShane R, Noel-Storr AH, Stott DJ, Anwer S, Sutton AJ, Burton JK, Quinn TJ



Figure 3. Summary ROC plot of AD-8 informant cut-off score 2. The dark point is a summary point, the other points individual studies; the broken line represents 95% confidence region.







# META-DTA v2.0 Crsu.shinyapps.io/dta\_ma/

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Suzanne Freeman, Clareece Nevill, Amit Patel, Nicola Cooper, Terry Quinn, Alex Sutton For feedback/questions about this app please contact Alex Sutton at ajs22@leicester.ac.uk App powered by Rshiny with statistical analyses performed using the package Ime4:

https://CRAN.R-project.org/package=Ime4

#### **UK Cochrane Centre**





ELSEVIER



Journal of Clinical Epidemiology 99 (2018) 64-74

#### Journal of Clinical Epidemiology

#### ORIGINAL ARTICLE

### Network meta-analysis of diagnostic test accuracy studies identifies and ranks the optimal diagnostic tests and thresholds for health care policy and decision-making

Rhiannon K. Owen<sup>a,\*</sup>, Nicola J. Cooper<sup>a</sup>, Terence J. Quinn<sup>b</sup>, Rosalind Lees<sup>b</sup>, Alex J. Sutton<sup>a</sup>

<sup>a</sup>Department of Health Sciences, University of Leicester, Leicester, UK <sup>b</sup>Institute of Cardiovascular and Medical Sciences, University of Glasgow, Glasgow, UK Accepted 7 March 2018; Published online 13 March 2018

MMSE <27/30	· · · · ·						
MoCA <26/30	MMSE <25/30	Sensitivity (95% Crl)	Specificity (95% Crl)	Rank best sensitivity (95% Crl)	<i>P</i> (Best) sensitivity	Rank best specificity (95% Crl)	P (Best) specificity
MoCA <22/30	Without threshold	constraints					
	MMSE <25	0.72 (0.61, 0.82)	0.84 (0.79, 0.89)	4 (3,4)	0	1 (1, 2)	0.97
	MMSE <27	0.89 (0.81, 0.95)	0.58 (0.45, 0.70)	2 (2,3)	0.01	3 (3, 3)	0
	MoCA <22	0.82 (0.70, 0.91)	0.77 (0.67, 0.85)	3 (2,4)	0	2 (1, 2)	0.03
	MoCA <26	0.97 (0.94, 0.99)	0.35 (0.23, 0.48)	1 (1,1)	0.99	4 (4, 4)	0





# **Prognosis Reviews**









**Prognosis research in Cochrane** 

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Fundamental prognosis

What is the natural history of X

Prognostic factor

Is exposure to X associated with development of Y

**Prediction models** 

Can a model that includes a,b,c predict development of X







Cochrane Database of Systematic Reviews

Anticholinergic burden (prognostic factor) for prediction of dementia or cognitive decline in older adults with no known cognitive syndrome (Review)

Taylor-Rowan M, Edwards S, Noel-Storr AH, McCleery J, Myint PK, Soiza R, Stewart C, Loke YK, Quinn TJ





# **Promoting EBM & Raising Standards**

#### Reporting standards for studies of diagnostic test accuracy in dementia The STARDdem Initiative

Anna H. Noel-Storr, MSc ABSTRACT Jenny M. McCleery, MB, BS Edo Richard, MD Craig W. Ritchie, MD Leon Flicker, MD Sarah J. Cullum, MBChB, MRCPsych, PhD Daniel Davis, MD Terence J. Quinn, MD Chris Hyde, MBBS Anne W.S. Rutjes, PhD Nadja Smailagic, MD Sue Marcus, MSc Sandra Black, MD Kaj Blennow, MD Carol Brayne, MD Mario Fiorivanti, MD Julene K. Johnson, PhD standards, and ad Sascha Köpke, PhD

Objective: To prov dementia disorder Methods: An inter impairment (STAR tivity and specifici 4 rounds of conse consensus meetin plement the gene their use in demer Results: More than most risk of inaded supplement the ST reference standar Conclusion: STAF checklist is elabor ders. Its adoption nostic tests in Alz



Cochrane Database of Systematic Reviews

Neuropsychological tests for the diagnosis of Alzheimer's disease dementia and other dementias: a generic protocol for crosssectional and delayed-verification studies (Protocol)

Davis DHJ, Creavin ST, Noel-Storr A, Quinn TJ, Smailagic N, Hyde C, Brayne C, McShane R, Cullum

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Age and Ageing 2018; 47: 349-355 doi: 10.1093/ageing/afy023 Published electronically 8 March 2018 © The Author(s) 2018. Published by Oxford University Press on behalf of the British Geriatrics Society. All rights reserved. For permissions, please email: journals.permissions@oup.com

#### REVIEW OF RESEARCH METHODS

## **Review of Diagnostic Test Accuracy (DTA)** studies in older people

Yemisi Takwoingi<sup>1</sup>, Terence | Quinn<sup>2</sup>

<sup>1</sup>Institute of Applied Health Research, University of Birmingham, UK <sup>2</sup>Institute of Cardiovascular and Medical Sciences, University of Glasgow, UK

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Alzheimer's & Dementia Volume 9, Issue 3, May 2013, Pages e96-e105



Online Exclusive

# Systematic review of the body of evidence for the use of biomarkers in the diagnosis of dementia

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## When is Alzheimer's not dementia—Cochrane commentary on The National Institute on Ageing and Alzheimer's Association Research Framework for Alzheimer's Disease @

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# Our next review..... Prioritisation











Categories	Definition
Prevention	Prevention of dementia and understanding relevant risk
	factors
Pathology	Understanding disease mechanisms, causes or stages of
	disease
Diagnosis	Role of identification of the disease and diagnostic tools
Drugs and	Using drugs and other interventions to manage disease
Interventions	
Support	Supporting people with dementia in daily life and disease
	management
Caregivers	Addressing the needs of caregivers, and how to support
	them
Awareness &	Educating and raising awareness of dementia and
Education	dementia-related issues for people living with dementia,
	care-givers, lay public and professionals
Research Methods	To improve the design, conduct, reporting and
	implementation of primary dementia research