AMSTAR 2: a critical appraisal tool for systematic reviews that include randomised or non-randomised studies of healthcare interventions, or both

For Yes	:	Optional (recommended)		
	<u>Population</u> Intervention <u>C</u> omparator group <u>O</u> utcome	□ Timeframe for follow-up		Yes No
2.		ntain an explicit statement that the review t of the review and did the report justify an		
The aut	ial Yes: hors state that they had a written l or guide that included ALL the	For Yes: As for partial yes, plus the protocol should be registered and should also have specified:		
	review question(s) a search strategy inclusion/exclusion criteria a risk of bias assessment	 a meta-analysis/synthesis plan, if appropriate, and a plan for investigating causes of heterogeneity justification for any deviations from the protocol 		Yes Partial Yes No
3.	Did the review authors explain	their selection of the study designs for incl	usion i	n the review?
	, the review should satisfy ONE of Explanation for including only R OR Explanation for including on OR Explanation for including bo	CTs ly NRSI th RCTs and NRSI	X	Yes No
4. For Part	Did the review authors use a co ial Yes (all the following):	mprehensive literature search strategy? For Yes, should also have (all the		
	searched at least 2 databases (relevant to research question) provided key word and/or search strategy justified publication restrictions (e.g. language)	following): searched the reference lists / bibliographies of included studies searched trial/study registries included/consulted content experts in the field where relevant, searched for grey literature conducted search within 24 months of completion of the review 	X	Yes Partial Yes No
5. For Yes	and achieved consensus on which OR two reviewers selected a same	ntly agreed on selection of eligible studies	X	Yes No

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01 103	, either ONE of the following:			
		onsensus on which data to extract from	×	Yes No
	OR two reviewers extracted data	from a sample of eligible studies <u>and</u> st 80 percent), with the remainder		
7.	Did the review authors provide	a list of excluded studies and justify the ex	clusion	ns?
For Par	tial Yes:	For Yes, must also have:		
	provided a list of all potentially relevant studies that were read in full-text form but excluded from the review	☐ Justified the exclusion from the review of each potentially relevant study		Yes Partial Yes No
8.	Did the review authors describe	e the included studies in adequate detail?		
For Par	tial Yes (ALL the following):	For Yes, should also have ALL the following:		
	described populations	□ described population in detail	X	Yes
	described interventions	□ described intervention in		Partial Yes
	described comparators	detail (including doses where		No
	described outcomes	relevant)		
	described research designs	 described comparator in detail (including doses where relevant) 		
		□ described study's setting		
		\Box timeframe for follow-up		
9.	Did the review authors use a sa individual studies that were inc	tisfactory technique for assessing the risk of	of bias	(RoB) in
RCTs	individual studies that were inc	tisfactory technique for assessing the risk of	of bias	(RoB) in
RCTs For Par	individual studies that were inc	tisfactory technique for assessing the risk of cluded in the review? For Yes, must also have assessed RoB from:		
RCTs For Part from	individual studies that were inc tial Yes, must have assessed RoB unconcealed allocation, <i>and</i>	tisfactory technique for assessing the risk of cluded in the review? For Yes, must also have assessed RoB from:	×	Yes
RCTs For Part from	individual studies that were inc tial Yes, must have assessed RoB unconcealed allocation, <i>and</i> lack of blinding of patients and	 tisfactory technique for assessing the risk of cluded in the review? For Yes, must also have assessed RoB from: allocation sequence that was not truly random, and 	×	Yes Partial Yes
RCTs For Part from	individual studies that were inc tial Yes, must have assessed RoB unconcealed allocation, <i>and</i> lack of blinding of patients and assessors when assessing	tisfactory technique for assessing the risk of cluded in the review? For Yes, must also have assessed RoB from: allocation sequence that was not truly random, and selection of the reported result	X	Yes Partial Yes No
RCTs For Part from	individual studies that were inc tial Yes, must have assessed RoB unconcealed allocation, <i>and</i> lack of blinding of patients and	 tisfactory technique for assessing the risk of cluded in the review? For Yes, must also have assessed RoB from: allocation sequence that was not truly random, and 	×	Yes Partial Yes
RCTs For Part from	individual studies that were inc tial Yes, must have assessed RoB unconcealed allocation, <i>and</i> lack of blinding of patients and assessors when assessing outcomes (unnecessary for objective outcomes such as all- cause mortality)	 tisfactory technique for assessing the risk of cluded in the review? For Yes, must also have assessed RoB from: allocation sequence that was not truly random, and selection of the reported result from among multiple measurements or analyses of a specified outcome 	X	Yes Partial Yes No Includes only
RCTs For Part from	individual studies that were inc tial Yes, must have assessed RoB unconcealed allocation, <i>and</i> lack of blinding of patients and assessors when assessing outcomes (unnecessary for objective outcomes such as all-	tisfactory technique for assessing the risk of cluded in the review? For Yes, must also have assessed RoB from: allocation sequence that was not truly random, and selection of the reported result from among multiple measurements or analyses of a specified outcome For Yes, must also have assessed RoB:	X	Yes Partial Yes No Includes only NRSI
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RCTs For Part from	individual studies that were inc tial Yes, must have assessed RoB unconcealed allocation, <i>and</i> lack of blinding of patients and assessors when assessing outcomes (unnecessary for objective outcomes such as all- cause mortality) tial Yes, must have assessed from confounding, <i>and</i>	 tisfactory technique for assessing the risk of cluded in the review? For Yes, must also have assessed RoB from: allocation sequence that was not truly random, and selection of the reported result from among multiple measurements or analyses of a specified outcome For Yes, must also have assessed RoB: methods used to ascertain exposures and outcomes, and 	X	Yes Partial Yes No Includes only NRSI Yes Partial Yes
RCTs For Part from	individual studies that were inc tial Yes, must have assessed RoB unconcealed allocation, <i>and</i> lack of blinding of patients and assessors when assessing outcomes (unnecessary for objective outcomes such as all- cause mortality) tial Yes, must have assessed	tisfactory technique for assessing the risk of cluded in the review? For Yes, must also have assessed RoB from: allocation sequence that was not truly random, and selection of the reported result from among multiple measurements or analyses of a specified outcome For Yes, must also have assessed RoB: methods used to ascertain exposures and outcomes, and selection of the reported result		Yes Partial Yes No Includes only NRSI Yes Partial Yes No
RCTs For Part from	individual studies that were inc tial Yes, must have assessed RoB unconcealed allocation, <i>and</i> lack of blinding of patients and assessors when assessing outcomes (unnecessary for objective outcomes such as all- cause mortality) tial Yes, must have assessed from confounding, <i>and</i>	 tisfactory technique for assessing the risk of cluded in the review? For Yes, must also have assessed RoB from: allocation sequence that was not truly random, and selection of the reported result from among multiple measurements or analyses of a specified outcome For Yes, must also have assessed RoB: methods used to ascertain exposures and outcomes, and 	X	Yes Partial Yes No Includes only NRSI Yes Partial Yes
RCTs For Part from	individual studies that were inc tial Yes, must have assessed RoB unconcealed allocation, <i>and</i> lack of blinding of patients and assessors when assessing outcomes (unnecessary for objective outcomes such as all- cause mortality) tial Yes, must have assessed from confounding, <i>and</i> from selection bias	 For Yes, must also have assessed RoB from: allocation sequence that was not truly random, <i>and</i> selection of the reported result from among multiple measurements or analyses of a specified outcome For Yes, must also have assessed RoB: methods used to ascertain exposures and outcomes, <i>and</i> selection of the reported result from among multiple methods used to ascertain exposures and outcomes, <i>and</i> selection of the reported result from among multiple methods used to ascertain exposures and outcomes, <i>and</i> 		Yes Partial Yes No Includes only NRSI Yes Partial Yes No Includes only RCTs
RCTs For Part from	individual studies that were inc tial Yes, must have assessed RoB unconcealed allocation, <i>and</i> lack of blinding of patients and assessors when assessing outcomes (unnecessary for objective outcomes such as all- cause mortality) tial Yes, must have assessed from confounding, <i>and</i> from selection bias	 tisfactory technique for assessing the risk of cluded in the review? For Yes, must also have assessed RoB from: allocation sequence that was not truly random, and selection of the reported result from among multiple measurements or analyses of a specified outcome For Yes, must also have assessed RoB: methods used to ascertain exposures and outcomes, and selection of the reported result from among multiple measurements or analyses of a specified outcome 		Yes Partial Yes No Includes only NRSI Yes Partial Yes No Includes only RCTs

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11. If meta-analysis was performed did the review authors use appropriate combination of results?	metho	ods for statistical
RCTs For Yes:		
		Yes
,		No
AND they used an appropriate weighted technique to combine study results and adjusted for heterogeneity if present.	×	No meta-analysis
AND investigated the causes of any heterogeneity		conducted
For NRSI		
For Yes:		
The authors justified combining the data in a meta-analysis		Yes
AND they used an appropriate weighted technique to combine		No
study results, adjusting for heterogeneity if present	X	No meta-analysis
AND they statistically combined effect estimates from NRSI that		conducted
were adjusted for confounding, rather than combining raw data,		
or justified combining raw data when adjusted effect estimates		
were not available		
AND they reported separate summary estimates for RCTs and		
NRSI separately when both were included in the review		
12. If meta-analysis was performed, did the review authors assess the poter individual studies on the results of the meta-analysis or other evidence s		
For Yes:		
□ included only low risk of bias RCTs		
□ OR, if the pooled estimate was based on RCTs and/or NRSI at variable		
RoB, the authors performed analyses to investigate possible impact of		
RoB on summary estimates of effect.		conducted
13. Did the review authors account for RoB in individual studies when into results of the review?	erpreti	ng/ discussing the
For Yes:		
□ included only low risk of bias RCTs		
OR, if RCTs with moderate or high RoB, or NRSI were included the	Σ	No No
review provided a discussion of the likely impact of RoB on the results		
14. Did the review authors provide a satisfactory explanation for, and disc heterogeneity observed in the results of the review?	ussion	of, any
For Yes:		
□ There was no significant heterogeneity in the results		
OR if heterogeneity was present the authors performed an investigation of	_	Yes
sources of any heterogeneity in the results and discussed the impact of this on the results of the review		No
15. If they performed quantitative synthesis did the review authors carry o investigation of publication bias (small study bias) and discuss its likely the review?		
For Yes:		
□ performed graphical or statistical tests for publication bias and discussed		Yes
the likelihood and magnitude of impact of publication bias		No
		No meta-analysis
		conducted

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16.	16. Did the review authors report any potential sources of conflict of interest, including any funding they received for conducting the review?				
For Yes:					
	The authors reported no competing interests OR	X	Yes		
	The authors described their funding sources and how they managed		No		
	potential conflicts of interest				

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