Section/topic	Item No	Checklist item	Reported on Page Number/Line Number	Reported on Section/Paragraph
TITLE	•			•
Title	1	Identify the report as a systematic review, meta-analysis, or both.	1/1	Title/NA
ABSTRACT	•			•
Structured summary	2	Provide a structured summary including, as applicable: background; objectives; data sources; study eligibility criteria, participants, and interventions; study appraisal and synthesis methods; results; limitations; conclusions and implications of key findings; systematic review registration number.	2/34-57	Abstract/NA
INTRODUCTION	'		•	•
Rationale	3	Describe the rationale for the review in the context of what is already known.	4/61-79	Introduction/1-2
Objectives	4	Provide an explicit statement of questions being addressed with reference to participants, interventions, comparisons, outcomes, and study design (PICOS).	4/80-81	Introduction/3
METHODS	•		•	•
Protocol and registration	5	Indicate if a review protocol exists, if and where it can be accessed (e.g., Web address), and, if available, provide registration information including registration number.	5/85-6/114	Methods/1-5
Eligibility criteria	6	Specify study characteristics (e.g., PICOS, length of follow-up) and report characteristics (e.g., years considered, language, publication status) used as criteria for eligibility, giving rationale.	5/91-100	Methods/2
Information sources	7	Describe all information sources (e.g., databases with dates of coverage, contact with study authors to identify additional studies) in the search and date last searched.	5/86-87	Methods/1
Search	8	Present full electronic search strategy for at least one database, including any limits used, such that it could be repeated.	5/86-90	Methods/1
Study selection	9	State the process for selecting studies (i.e., screening, eligibility, included in systematic review, and, if applicable, included in the meta-analysis).	5/85-100	Methods/1-2
Data collection process	10	Describe method of data extraction from reports (e.g., piloted forms, independently, in duplicate) and any processes for obtaining and confirming data from investigators.	5/101-6/110	Methods/3-4
Data items	11	List and define all variables for which data were sought (e.g., PICOS, funding sources) and any assumptions and simplifications made.	5/101-6/110	Methods/3-4

Risk of bias in individual studies	12	Describe methods used for assessing risk of bias of individual studies (including specification of whether this was done at the study or outcome level), and how this information is to be used in any data synthesis.	6/111-114	Methods/5
Summary measures	13	State the principal summary measures (e.g., risk ratio, difference in means).	6/115-125	Methods/6
Synthesis of results	14	Describe the methods of handling data and combining results of studies, if done, including measures of consistency (e.g., l²) for each meta-analysis.	6/115-125	Methods/6
Risk of bias across studies	15	Specify any assessment of risk of bias that may affect the cumulative evidence (e.g., publication bias, selective reporting within studies).	6/115-125	Methods/6
Additional analyses	16	Describe methods of additional analyses (e.g., sensitivity or subgroup analyses, meta-regression), if done, indicating which were pre-specified.	NA	N/A
RESULTS				
Study selection	17	Give numbers of studies screened, assessed for eligibility, and included in the review, with reasons for exclusions at each stage, ideally with a flow diagram.	7/128-132	Results/1 (Figure 1)
Study characteristics	18	For each study, present characteristics for which data were extracted (e.g., study size, PICOS, follow-up period) and provide the citations.	7/133-143	Results/2 (Table 1,2)
Risk of bias within studies	19	Present data on risk of bias of each study and, if available, any outcome level assessment (see item 12).	7/143-145	Results/2 (S1 Table)
Results of individual studies	20	For all outcomes considered (benefits or harms), present, for each study: (a) simple summary data for each intervention group (b) effect estimates and confidence intervals, ideally with a forest plot.	7/146-9/188	Results/3-8 (Figure 2,3)
Synthesis of results	21	Present results of each meta-analysis done, including confidence intervals and measures of consistency.	7/146-9/188	Results/3-8 (Figure 2,3)
Risk of bias across studies	22	Present results of any assessment of risk of bias across studies (see Item 15).	9/189-191	Results/9 (Figure 4)
Additional analysis	23	Give results of additional analyses, if done (e.g., sensitivity or subgroup analyses, meta-regression [see Item 16]).	NA	NA
DISCUSSION				
Summary of evidence	24	Summarize the main findings including the strength of evidence for each main outcome; consider their relevance to key groups (e.g., healthcare providers, users, and policy makers).	9/193-200	Discussion/1
Limitations	25	Discuss limitations at study and outcome level (e.g., risk of bias), and at review-level (e.g., incomplete retrieval of identified research, reporting bias).	11/226-235	Discussion/6
Conclusions	26	Provide a general interpretation of the results in the context of other evidence, and implications for future research.	11/236-240	Discussion/7
			1	<u> </u>

FUNDING							
Funding	27		NA	NA			
		systematic review.					

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<sup>\*</sup>As the checklist was provided upon initial submission, the page number/line number reported may be changed due to copyediting and may not be referable in the published version. In this case, the section/paragraph may be used as an alternative reference.