SUPPLEMENTARY APPENDIX

Proportions and incidence of locally advanced cervical cancer: a global systematic literature review

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Supplementary Methods

The search strategies shown below were created to support this locally advanced cervical cancer epidemiology systematic literature review as well as one focused on the natural history of locally advanced cervical cancer. Only the epidemiology publications are reported in this article. EMBASE, MEDLINE (PubMed), and Cochrane databases were searched using the search strategies below. Because some studies are not appropriately indexed in electronic databases, bibliographic searching and pearl growing techniques were used to identify any potentially relevant studies that were not captured by database searches.

Embase search strategy run on June 10, 2020

Search	Query	Results		
number				
#1	'uterine cervix cancer'/syn	114 434		
#2	'cervical tumor' OR 'cervical neoplasm' OR 'cervical tumour' OR 'cervical cancer'	67 038		
#3	cervi* NEAR/5 (cancer* OR oncolog* OR neoplas* OR carcinom* OR malignan*	159 269		
	OR tumor* OR tumour* OR mass* OR growth* OR cyst* OR adenocarcinom* OR			
	squamous)			
#4	#1 OR #2 OR #3	159 271		
#5	'natural history'/exp OR 'natural history':ab,ti,kw OR 'natural course'	423 926		
#6	'natural history study'			
#7	('observational' OR 'prospective' OR 'retrospective' OR 'cross-sectional' OR 'cross	2 513 311		
	sectional' OR 'longitudinal') NEAR/3 ('study' OR 'studies' OR analys*)			
#8	#5 AND #7	18 090		
#9	'disease course':ab,ti,kw OR 'clinical course' OR ('natural history' NEAR/2	113 467		
	prognos*)			
#10	'inception cohort' OR 'disease exacerbation'/syn OR 'disease progression' OR	246 377		
	'outcome assessment':ab,ti,kw			
#11	#5 OR #6 OR #8 OR #9 OR #10	770 081		
#12	#4 AND #11	3494		
#13	'locally advanced' OR 'local advanced' OR (local* NEAR/2 'advanced') OR 'stage	103 225		
	one' OR 'stage two' OR 'stage three' OR 'stage four' OR 'stage ib2' OR 'stage iib'			

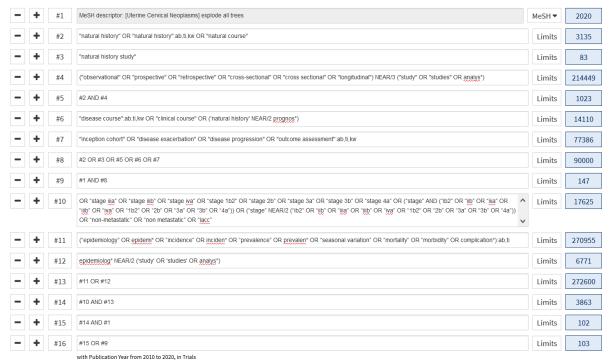
	OR 'stage iiia' OR 'stage iiib' OR 'stage iva' OR 'stage 1b2' OR 'stage 2b' OR 'stage 3a' OR 'stage 3b' OR 'stage 4a' OR ('stage' AND ('ib2' OR 'iib' OR 'iiia' OR 'iiib' OR 'iva' OR '1b2' OR '3a' OR '3b' OR '4a')) OR ('stage' NEAR/2 ('ib2' OR 'iib' OR 'iiia' OR 'iiib' OR 'iva' OR '1b2' OR '2b' OR '3a' OR '3b' OR '4a')) OR 'non-metastatic' OR 'non metastatic' OR 'lacc'	
#14	'epidemiology':ab,ti OR epidemi*:ab,ti OR 'incidence':ab,ti OR inciden*:ab,ti OR 'prevalence':ab,ti OR prevalen*:ab,ti OR 'seasonal variation':ab,ti OR 'mortality':ab,ti OR 'morbidity':ab,ti OR complication*:ab,ti	4 471 505
#15	epidemiolog* NEAR/2 ('study' OR 'studies' OR analys*)	136 119
#16	#14 OR #15	4 476 280
#17	#13 AND #16	26 494
#18	#4 AND #17	2549
#19	#12 OR #18	5995
#20	#19 AND ([conference abstract]/lim OR [conference paper]/lim OR [conference review]/lim OR [letter]/lim OR [note]/lim OR [review]/lim)	2540
#21	#19 AND [animals]/lim	126
#22	#20 OR #21	2598
#23	#19 NOT #22	3397
#24	#23 AND [2010-2020]/py	1671
#25	#24 AND [english]/lim	1551

PubMED search strategy run on June 10, 2020

Search number	Query	Results
#1	Search: 'uterine cervix cancer'[MeSH Terms]	74 490
#2	Search: "cervical tumor" OR "cervical neoplasm" OR "cervical tumour" OR "cervical cancer"	47 147
#3	Search: cervi* AND (cancer* OR oncolog* OR neoplas* OR carcinom* OR malignan* OR tumor* OR tumour* OR mass* OR growth* OR cyst* OR adenocarcinom* OR squamous)	161 507
#4	Search: #1 OR #2 OR #3	161 507
#5	Search: ('natural history'[MeSH Terms]) OR ('natural history'[Title/Abstract]) OR 'natural course'	84 236
#6	Search: "natural history study"	776
#7	Search: ("observational" OR "prospective" OR "retrospective" OR "cross-sectional" OR "cross sectional" OR "longitudinal") AND ("study" OR "studies" OR analys*)	2 332 275
#8	Search: #5 AND #7	17 334
#9	Search: "disease course":[Title/Abstract] OR "clinical course" OR ("natural history" AND prognos*)	81 583
#10	Search: (("inception cohort") OR ("disease exacerbation"[MeSH Terms]) OR "disease progression") OR ("outcome assessment"[MeSH Terms])	218 726
#11	Search: #5 OR #6 OR #8 OR #9 OR #10	360 274
#12	Search: #4 AND #11	4419
#13	Search: "locally advanced" OR "local advanced" OR (local* AND "advanced") OR "stage one" OR "stage two" OR "stage three" OR "stage four" OR "stage ib2" OR "stage iib" OR "stage iiia" OR "stage iiib" OR "stage iva" OR "stage 1b2" OR "stage 2b" OR "stage 3a" OR "stage 3b" OR "stage 4a" OR ("stage" AND ("ib2" OR "iib" OR "iiia" OR "iiib" OR "iva" OR "1b2" OR "2b" OR "3a" OR	100 166

	"3b" OR "4a")) OR ("stage" AND ("ib2" OR "iib" OR "iiia" OR "iiib" OR "iva" OR "1b2" OR "2b" OR "3a" OR "3b" OR "4a")) OR "non-metastatic" OR "non-metastatic" OR "lacc"	
#14	Search: "epidemiology"[Title/Abstract]OR epidemi*[Title/Abstract]OR "incidence"[Title/Abstract]OR inciden*[Title/Abstract]OR "prevalence"[Title/Abstract]OR prevalen*[Title/Abstract]OR "seasonal variation"[Title/Abstract]OR "mortality"[Title/Abstract]OR "morbidity"[Title/Abstract]OR complication*[Title/Abstract]	3 291 414
#15	Search: epidemiolog* AND (study OR studies OR analys*)	1 520 332
#16	Search: #14 OR #15	3 926 004
#17	Search: #13 AND #16	25 044
#18	Search: #4 AND #17	2117
#19	Search: #12 OR #18	6471
#20	Search: (#19 AND (inprocess[sb] OR pubstatusaheadofprint))	131

Cochrane search strategy run on June 10, 2020



The following conferences were also searched for relevant abstracts from meetings held between January 2017 and June 2020: American Society of Clinical Oncology, European Society for Medical Oncology, European Society of Gynaecological Oncology, Society of Gynaecologic Oncology, American Association for Cancer Research, International Society for Pharmacoeconomics and Outcomes Research, International Gynecologic Cancer Society. The publication timeframe for conference searching was limited to the previous 3 years

based on the assumption that research presented at conferences is usually published within 3–4 years as a full-text article or indexed in different biomedical literature databases as a conference paper, conference review, etc.

Inclusion and exclusion criteria used to identify relevant studies are shown in the table below.

Parameter	Inclusion/exclusion criteria
Study design	Retrospective observational study
	Prospective observational study
	Case-control studies
	Surveys and cross-sectional studies
	Registry/database studies
	Excluded: controlled trials (randomized controlled trial, non-
	randomized controlled study, or single-arm study)
Population	Adult population (aged ≥18 years)
	Any race
	Locally advanced cervical cancer: stages IB2-IVA per any
	version of the FIGO staging criteria
	Excluded: studies that only include patients with early-stage or
	metastatic cervical cancer
Line of therapy	Not restricted
	Studies of patients with locally advanced cervical cancer (both
	untreated and treated)
Countries	Not restricted
Language	• English ^a
Time-frame	• 2010–2020
Data reported	Proportion of patients with cervical cancer by disease stage

Incidence of cervical cancer by disease stage

^aEnglish language was a criterion from the beginning of the systematic literature review process and was used as an exclusion criterion in database search queries.

FIGO, International Federation of Gynecology and Obstetrics.

Data extraction

The following information was extracted from the final set of published reports, where available: study details (sample size, inclusion/exclusion criteria, disease stage, stage classification criteria, treatment details, study limitations, time-frame of data collection, data source, location), patient demographics (age, race/ethnicity), clinical characteristics (histology, prior therapy), the proportion of patients with locally advanced stages of cervical cancer, prevalence (rate, odds ratio, risk ratio), and incidence (rate, risk ratio).

Calculation of the Proportion of Locally Advanced Cervical Cancer
The Surveillance, Epidemiology, and End Results summary stage categorizes the extent of
cancer spread in a basic set of criteria. In the past, this classification system has also been
referred to as General Stage, California Stage, historic stage, and Surveillance,
Epidemiology, and End Results Stage. Summary stage uses all information available via
medical records (ie, both clinical and pathologic documentation). Below are the criteria as per
the most recent version (v2.0) published in 2020; however, studies included in the systematic
literature review may have used older versions of the criteria. A summary of changes
between the last available version (v1.7) and version 2.0 is available at
https://seer.cancer.gov/tools/ssm/change-log.pdf. The 2020 criteria were used to determine
which SEER Summary stages were equivalent to FIGO stage IB2-IVA.

2018 Surveillance, Epidemiology, and End Results summary staging criteria for cervical cancer[1]

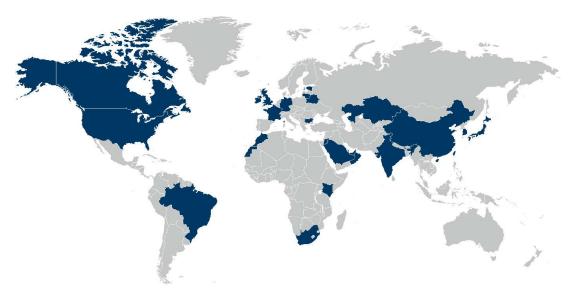
Code	Stage	Definition
0	In situ	Noninvasive, intraepithelial lesions. Includes cancer in situ with endocervical gland involvement, cervical intraepithelial neoplasia Grade III, preinvasive.
1	Localized	Clinically visible lesion (macroscopic), including superficial invasion.

		Confined to cervix uteri or uterus NOS, except corpus uteri NOS, including if not clinically visible or unknown if clinically visible.
		Measured stromal invasion less than 5 mm from the base of the epithelium AND horizontal spread of 7.0 mm or less.
		Includes FIGO stage IA1, IA2, IA NOS, IB1, IB2, IB NOS, I NOS.
2	Regional (direct extension)	Extension to the bladder wall; bladder NOS excluding mucosa; bullous edema of bladder mucosa; confined to corpus uteri, size, depth and horizontal spread unknown; corpus uteri NOS; Cul de sac (rectouterine pouch); fallopian tube(s); "frozen pelvis" (clinically described); hydronephrosis or nonfunctioning kidney; invasion beyond uterus NOS; ligament(s) (broad, cardinal, uterosacral); ovary/ovaries; parametrial (paracervical soft tissue) invasion; pelvic wall(s); rectal wall; rectum NOS excluding mucosa; upper two-thirds of vagina including fornices; ureter (intra- and extramural); urethra; vagina (lower third [not extending into pelvic wall], NOS); vaginal wall NOS; vulva. Includes FIGO stage IIA, IIB, II NOS, IIIA, IIIB, III NOS.
3	Regional (lymph node involvement only)	Localized tumor WITH regional lymph node involvement. Involvement of the following types of lymph nodes: para-aortic, iliac NOS, paracervical, parametrial, sacral NOS, regional NOS. Includes FIGO stages IIIC1, IIIC2, IIIC NOS.
4	Regional (both direct extension and regional lymph nodes involved)	Any combination of codes 2 and 3 above.
7	Distant (sites or lymph nodes)	Cervical cancer that has metastasized. Includes bladder mucosa, rectal mucosa, sigmoid colon, small intestine, inguinal (femoral) lymph node, mediastinal lymph node, scalene lymph node, supraclavicular lymph node; or cancers labeled as carcinomatosis or distant metastasis with or without distant lymph nodes. Includes FIGO stage IVA, IVB, IV NOS.
9	Unknown	Unknown if extension or metastasis.

FIGO, International Federation of Gynecology and Obstetrics; NOS, not otherwise specified.

Supplementary Fig 1. Countries represented by the studies included in the systematic

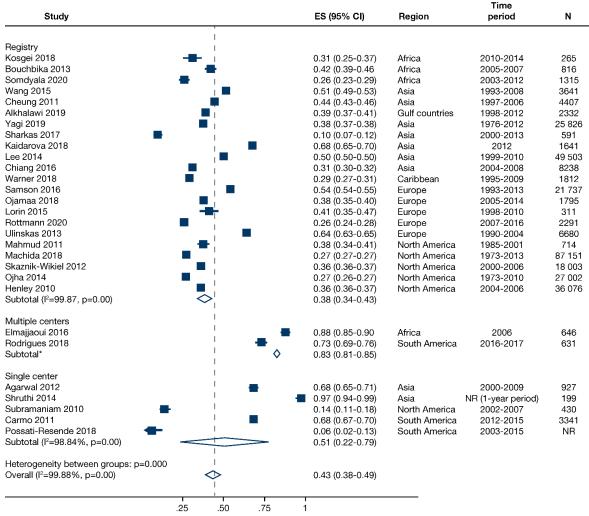
literature review.



Included countries are the United States, Canada, China, India, Japan, Jordan, collective Gulf countries [Saudi Arabia, United Arab Emirates, Qatar, Oman, Kuwait, Bahrain], Kazakhstan, Korea, Taiwan, Belarus, Bulgaria, Estonia, France, Germany, Ireland, Lithuania, the United Kingdom, Morocco, Kenya, South Africa, Brazil, and Trinidad & Tobago.

Supplementary Fig 2. Estimated proportion of locally advanced cervical cancer by type of

data source



Proportion of cervical cancer patients with locally advanced disease

Estimated proportion for each study (ES) and the 95% confidence intervals are plotted according to data source (registry, multicenter institution, or single institution). Overlapping timeframes and duplicate data from the same study have been removed. Red triangles represent the range of the subtotal estimated proportion, and the red dashed line represented the overall estimated proportion of locally advanced cervical cancer from this dataset. Heterogeneity of studies is reflected in the I^2 value; a score of >60% = high heterogeneity. Single center studies provided the most unreliable data with the largest variance (estimated range, 6–97%). N indicates the total number of women with cervical cancer. NR, not reported.

Supplementary Table 1. Strengthening the Reporting of Observational studies in Epidemiology checklist items

Section	Item No	Recommendation
Title and abstract	1	(a) Indicate the study's design with a commonly used term in the title or the abstract
		(b) Provide in the abstract an informative and balanced summary of what was done and what
		was found
Introduction		
Background/rationale	2	Explain the scientific background and rationale for the investigation being reported
Objectives	3	State specific objectives, including any prespecified hypotheses
Methods		
Study design	4	Present key elements of study design early in the paper
Setting	5	Describe the setting, locations, and relevant dates, including periods of recruitment, exposure,
		follow-up, and data collection
Participants	6	(a) Cohort study—Give the eligibility criteria, and the sources and methods of selection of
		participants. Describe methods of follow-up
		Case-control study—Give the eligibility criteria, and the sources and methods of case
		ascertainment and control selection. Give the rationale for the choice of cases and controls

		Cross-sectional study—Give the eligibility criteria, and the sources and methods of selection of
		participants
		(b) Cohort study—For matched studies, give matching criteria and number of exposed and
		unexposed
		Case-control study—For matched studies, give matching criteria and the number of controls per
		case
Variables	7	Clearly define all outcomes, exposures, predictors, potential confounders, and effect modifiers.
		Give diagnostic criteria, if applicable
Data sources/ measurement	8*	For each variable of interest, give sources of data and details of methods of assessment
		(measurement). Describe comparability of assessment methods if there is more than one group
Bias	9	Describe any efforts to address potential sources of bias
Study size	10	Explain how the study size was arrived at
Quantitative variables	11	Explain how quantitative variables were handled in the analyses. If applicable, describe which
		groupings were chosen and why
Statistical methods	12	(a) Describe all statistical methods, including those used to control for confounding
		(b) Describe any methods used to examine subgroups and interactions
		(c) Explain how missing data were addressed

	Case-control study—If applicable, explain how matching of cases and controls was addressed
	Cross-sectional study—If applicable, describe analytical methods taking account of sampling
	strategy
	(<u>e</u>) Describe any sensitivity analyses
13*	(a) Report numbers of individuals at each stage of study—eg numbers potentially eligible,
	examined for eligibility, confirmed eligible, included in the study, completing follow-up, and
	analysed
	(b) Give reasons for non-participation at each stage
	(c) Consider use of a flow diagram
14*	(a) Give characteristics of study participants (eg demographic, clinical, social) and information on
	exposures and potential confounders
	(b) Indicate number of participants with missing data for each variable of interest
	(c) Cohort study—Summarise follow-up time (eg, average and total amount)
15*	Cohort study—Report numbers of outcome events or summary measures over time
	Case-control study—Report numbers in each exposure category, or summary measures of
_	14*

		exposure
		Cross-sectional study—Report numbers of outcome events or summary measures
Main results	16	(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and their
		precision (eg, 95% confidence interval). Make clear which confounders were adjusted for and
		why they were included
		(b) Report category boundaries when continuous variables were categorized
		(c) If relevant, consider translating estimates of relative risk into absolute risk for a meaningful
		time period
Other analyses	17	Report other analyses done—eg, analyses of subgroups and interactions, and sensitivity
		analyses
Discussion		
Key results	18	Summarise key results with reference to study objectives
Limitations	19	Discuss limitations of the study, taking into account sources of potential bias or imprecision.
		Discuss both direction and magnitude of any potential bias
Interpretation	20	Give a cautious overall interpretation of results considering objectives, limitations, multiplicity of
		analyses, results from similar studies, and other relevant evidence
Generalisability	21	Discuss the generalisability (external validity) of the study results

Other information		
Funding	22	Give the source of funding and the role of the funders for the present study and, if applicable, for
		the original study on which the present article is based

^{*}Information should be given separately for cases and controls in case-control studies and, if applicable, for exposed and unexposed groups in cohort and cross-sectional studies.

Items as reported on the Strengthening the Reporting of Observational studies in Epidemiology website (https://www.equator-network.org/reporting-quidelines/strobe/).

Supplementary Table 2. Strengthening the Reporting of Observational studies in Epidemiology – Abstract version – checklist items

Item	Recommendation
Title	Indicate the study's design with a commonly used term in the title (e.g cohort, case-control, cross sectional)
Authors	Contact details for the corresponding author
Study design	Description of the study design (e.g cohort, case-control, cross sectional)
Objective	Specific objectives or hypothesis
Methods	
Setting	Description of setting, follow-up dates or dates at which the outcome events occurred or at
	which the outcomes were present, as well as any points or ranges on other time scales for
	the outcomes (e.g., prevalence at age 18, 1998-2007).
Participants	Cohort study—Give the most important eligibility criteria, and the most important sources
	and methods of selection of participants. Describe briefly the methods of follow-up
	Case-control study—Give the major eligibility criteria, and the major sources and
	methods of case ascertainment and control selection
	Cross-sectional study—Give the eligibility criteria, and the major sources and methods of
	selection of participants
	Cohort study—For matched studies, give matching and number of exposed and

	unexposed
	Case-control study—For matched studies, give matching criteria and the number of
	controls per case
Variables	Clearly define primary outcome for this report.
Statistical methods	Describe statistical methods, including those used to control for confounding
Results	
Participants	Report Number of participants at the beginning and end of the study
Main results	Report estimates of associations. If relevant, consider translating estimates of relative risk into absolute risk for a
	meaningful time period
	Report appropriate measures of variability and uncertainty (e.g., odds ratios with confidence intervals)
Conclusions	General interpretation of study results

Checklist items were obtained from the EQUATOR network website: https://www.equator-network.org/reporting-guidelines/strobe-abstracts/.

Supplementary Table 3. Strengthening the Reporting of Observational studies in Epidemiology checklist for included studies[2-30]

Study name		e & tract		ro- tion							Meth	nods							Results								Discu	ssio	n	Other infor-mation				
	1a	1b	2	3	4	5	6a	6b	7	8	9	10	11	12a	12b	12c	12d	12e	13a	13b	13c	14a	14b	14c	15	16a	16b	16c	17	18	19	20	21	22
Henley 2010	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ	Υ	Υ	Υ	Υ	N	N	Υ	Υ	N	Υ	Υ	N	Υ	Υ	Υ	N	N	Υ	Υ	Υ	N	N
Garg 2011	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ	Υ	Υ	Υ	N	Ν	N	Υ	Ν	Ν	Υ	N	N	Υ	Υ	Υ	Υ	N	Υ	Υ	Υ	N	N
Skaznik- Wikiel 2012	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ	Υ	Υ	Υ	N	N	N	Υ	N	N	Υ	Υ	N	Υ	Υ	Υ	N	N	Υ	Υ	Υ	N	N
Ojha 2014	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	N	N	Ν	Ν	Υ	Ν	N	Υ	Υ	Υ	Υ	Υ	Υ	N	N	Υ	Υ	Υ	Ζ	N
Machida 2018	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ	Υ	Υ	Υ	Υ	N	Υ	Υ	Υ	N	Υ	Υ	N	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ
Zahnd 2018	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Ν	Υ	Υ	Υ	Υ	N	N	N	Υ	N	N	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ	Υ	Ν	Ζ	N
Hou 2019	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Ν	Υ	Υ	Υ	Υ	Υ	Ν	Ν	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ	Υ	Υ	Ν	Ν
Bruegl 2020	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ	Υ	Υ	Υ	Υ	N	N	Υ	Υ	N	Υ	Υ	N	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ
Tian 2020	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Ν	Υ	Υ	Υ	Υ	N	Ν	Ν	Υ	N	N	Υ	N	Ν	Υ	Υ	Υ	N	Υ	Υ	Υ	Υ	Ν	Υ
Mahmud 2011	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ	Υ	Υ	Υ	N	N	N	Υ	N	N	Υ	N	N	Υ	Υ	N	N	N	Υ	Υ	Υ	N	Υ
Carmo 2011	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	N	Υ	N	N	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ	Υ	Υ	Υ	N	N
Possati- Resende 2018	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Ν	Υ	Υ	N	N	N	N	N	N	N	Ν	N	N	Υ	N	N	Υ	Υ	N	N	N	N	Υ	Υ	Υ	Z	N
Warner 2018	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Ν	Υ	Υ	Υ	N	N	N	N	Υ	N	Υ	Υ	Υ	Ν	Υ	N	N	Ν	N	Υ	Υ	Υ	Υ	Υ
Lorin 2015	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Ν	Ν	Ν	N	Ν	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ	N	N	Υ	N
Samson 2016	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ	Υ	Υ	Υ	N	Ν	N	Υ	N	N	Υ	N	N	Υ	N	N	N	N	Υ	Υ	Υ	N	Υ
Ojamaa 2018	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Ν	Υ	Υ	Υ	Υ	N	Ν	N	Υ	N	N	Υ	Ν	Ν	Υ	Υ	Υ	Ν	N	Υ	Υ	Υ	Ν	Υ
Ulinskas 2013	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ	Υ	Υ	Υ	N	Ν	N	N	N	N	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ	N	N	N	Υ
Bouchbika 2013	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	N	Υ	Υ	Υ	N	N	N	Υ	N	N	Υ	Υ	N	Υ	Υ	Υ	N	Υ	Υ	Υ	Υ	Υ	N

Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ	Υ	Υ	Υ	N	Ν	N	Υ	N	Υ	Υ	N	Υ	Υ	N	N	N	Ν	Υ	N	N	Υ	N
Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ	Υ	Υ	Υ	Υ	Ν	N	Υ	N	N	Υ	Υ	N	Υ	Υ	Υ	N	Υ	Υ	Υ	Υ	N	Υ
Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ	Υ	N	Υ	Υ	Υ	N	N	Ν	N	Υ	N	N	Υ	Υ	N	Υ	N	N	N	N	Υ	N	N	N	N
Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ	Υ	Υ	Υ	N	Ν	N	Υ	N	N	Υ	N	N	Υ	N	N	N	N	Υ	N	N	N	N
Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ	Υ	Υ	Υ	N	Ν	N	Υ	N	N	Υ	N	N	Υ	N	N	N	N	Υ	N	N	N	N
Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ	Υ	N	Υ	N	Ν	N	Υ	N	N	Υ	N	N	Υ	N	N	N	N	Υ	N	N	N	N
Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Ν	Υ	Υ	Υ	Υ	Ν	Z	Ν	Υ	Ν	Υ	Υ	Ν	Ν	Υ	Υ	Υ	Ν	Ν	Υ	Ν	Υ	Ν	Υ
Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ	Υ	Υ	Υ	N	Ν	N	Υ	N	N	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ	Υ	N	N	N	N
Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	Υ	Υ	Υ	Υ	Υ	Ν	N	Υ	Ν	N	Υ	N	N	Υ	Υ	Υ	Ν	N	Υ	Υ	Υ	N	N
Υ	N	Υ	Υ	Υ	Υ	Υ	N	Υ	Υ	N	Υ	Υ	Υ	Υ	N	N	N	Υ	N	N	N	Υ	Υ	N	Υ	N	N	Υ	Υ	Υ	Υ	Υ	Υ
Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	N	N	Υ	Υ	Υ	N	Ν	N	N	N	N	N	N	Υ	Υ	Υ	N	N	Υ	Υ	Υ	N	N	Υ
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Checklist items are explained in Online Supplementary Table 1.

Study Name	Title	Authors	Study	Objective		Me	thods		Results -	Main results	Conclusions
			Design		Setting	Participants	Variables	Statistics	Participants		
Subramaniam 2010	N	Υ	Υ	Υ	Y	Y	Y	Y	Y	N	Y
McLean 2012 (Int J Gyn Can)	N	Υ	N	Υ	Y	Y	Y	Y	Y	N	Y
McLean 2012 (Gyn Oncol)	N	Υ	Υ	Υ	Y	Y	Y	Y	Y	Υ	Y
Popadiuk 2010	N	N	N	Υ	Y	Y	N	N	Y	N	Y
Rodrigues 2018	N	Y	Υ	Υ	Υ	Y	N	Y	Y	Y	Y
Nathani 2012	N	Υ	N	Υ	Y	Y	Y	N	Y	N	Y
Garry 2018	N	Y	Υ	Υ	Y	Y	N	N	Y	N	Y
Rottmann 2020	Υ	Y	Υ	Υ	Y	Y	Y	Y	Y	Y	Υ
Litvinova 2017	N	Y	N	Υ	Y	Y	Y	N	Y	N	N
Kosgei 2018	N	Y	N	Υ	Y	Y	Y	N	Y	N	Y
Kaidarova 2018	N	Υ	N	Υ	Υ	Y	Y	Y	Y	N	Υ

Checklist items are explained in Online Supplementary Table 2.

Supplemental material

Supplementary Table 5. Study characteristics

Study / publication type	Location	Study design & data source type	Time-frame	Data source	Classification criteriaª	Stages of cervical cancer included	Population	Total patients with cervical cancer
Henley 2010[11] Manuscript	USA	Retrospective cohort, National registry	2004–2006	National Program of Cancer Registries and SEER database	ICD-O-3: C53 Collaborative Stage classification	Localized, regional, distant	Invasive cervical, breast, or colon/rectum cancers ≥20 years of age for cervical cancer	36 076
Subramaniam 2010[41] Congress abstract	Birmingham, Alabama, USA	Retrospective cohort, Single center institution	2002–2007	University-based gynecologic oncology program	NR	I, II, III, IV	Invasive cervical cancer	430
Garg 2011[10] Manuscript	USA	Retrospective cohort, National registry	1988–2005	SEER database 17 registries used	NR FIGO staging	IIA (IIA1, IIA2)	Stage IIA cervical cancer Primary treatment with RH or RT	560
McLean 2012[36] Congress abstract	USA	Retrospective cohort, Healthcare database	1992–2007	SEER-Medicare database	NR	I, II, III, IV	Any stage cervical cancer Aged 65–100 years	6718
McLean 2012[35] Congress abstract	USA	Case-control, Healthcare database	1992–2007	SEER-Medicare database	NR	I, II, III, IV	Diagnosed with cervical cancer after age 70 (n=734) Matched non-cancer controls (n=2936)	734
Skaznik-Wikiel 2012[22] Manuscript	USA	Retrospective cohort, National registry	2000–2006	SEER database 17 registries used	NR FIGO staging	I, II, III, IV	Cervical cancer diagnosis	18 003

Study / publication type	Location	Study design & data source type	Time-frame	Data source	Classification criteria ^a	Stages of cervical cancer included	Population	Total patients with cervical cancer
Ojha 2014[17] Manuscript	USA	Longitudinal/cohor t, National registry	1973–2010	SEER database Only 9 registries used: Atlanta, Connecticut, Detroit, Hawaii, Iowa, New Mexico, San Francisco- Oakland, Seattle- Puget Sound, Utah SEER-PAYA cancer survivors' cohort	NR SEER summary staging	1 – localized 2/3 – locally advanced 4 – metastatic	PAYA: females diagnosed with any cancer before age 30 years, had survived ≥5 years post-diagnosis, and were later diagnosed with invasive cervical cancer (n=46) Females in the general population aged ≤56 years at primary cervical cancer diagnosis (n=26,956)	27 002
Machida 2018[14] Manuscript	USA	Retrospective cohort, National registry	1973–2013	SEER database	ICD-O-3 and WHO classifications (histology) TNM: AJCC 7 th ed. staging	I, II, III, IV	Cervical cancer diagnosis	87 151
Zahnd 2018[29] Manuscript	USA	Retrospective cohort, National registry	2009–2013	North American Association of Central Cancer Registries	NR SEER summary stage	Localized and distant	All stageable cancer types combined HPV-associated cancers Tobacco-associated cancers	NR

Study / publication type	Location	Study design & data source type	Time-frame	Data source	Classification criteriaª	Stages of cervical cancer included	Population	Total patients with cervical cancer
							Individual cancers with screening recommendation from the United States Preventive Services Task Force and has current recommendations (colorectal, female breast, cervical, and lung) Cancers for which screening was recommended for most of the study period (prostate) Cancers with insufficient evidence for recommended screening but for which screening may be performed regularly in clinical practice (skin and oral)	
Hou 2019[12] Manuscript	USA	Retrospective cohort, National registry	1988–2011	SEER database	ICD-O-3: C53.0- 53.9 FIGO staging	I, II, III, IV	White and Asian-American patients with cervical cancer	58 780

Study / publication type	Location	Study design & data source type	Time-frame	Data source	Classification criteria ^a	Stages of cervical cancer included	Population	Total patients with cervical cancer
Bruegl 2020[5] Manuscript	Idaho, Oregon, Washington, USA	Retrospective cohort, Regional registry	1996–2016	Cancer Data Registry of Idaho Oregon State Cancer Registry Washington State Cancer Registry	ICD-O-3: C53.0- 53.9 NR	Localized, regional, distant	Non-Hispanic White and American Indian/Alaskan Native women diagnosed with a gynecological cancer	7222
Tian 2020[24] Manuscript	USA	Retrospective cohort, National registry	2010–2015	SEER database 18 registries used	NR FIGO staging	IB2-IVA	Cervical cancer stages IB2 to IVA Pathological biopsy confirmed SCC and AC No distant metastases Aged 20–69 years	4131
Mahmud 2011[15] Manuscript	Saskatchewan, Canada	Retrospective cohort, Regional registry	1987–2001	Provincial cancer registry- Saskatchewan	NR FIGO staging	I, II, III, IV	Cervical cancer diagnosis	714
Popadiuk 2010[38] Congress abstract	Newfoundland, Canada	Retrospective cohort, Regional registry	1992–2008	Newfoundland Cancer Registry	NR	IA, IB, IIB, IIIB, IVA	Invasive cervical cancer Aged 19–29 years	37
Carmo 2011[6] Manuscript	Rio de Janeiro, Brazil	Retrospective cohort, Single center institution	1999–2004	Brazilian National Cancer Institute	NR FIGO staging	I, II, III, IV	Cervical cancer diagnosis	3341
Rodrigues 2018[39] Congress abstract	Brazil	Prospective, Multiple institutions	2016–2017	16 sites, representing 5 Brazilian regions	NR	I, II, III, IV	 Invasive cervical cancer Aged ≥18 years 	631
Possati-Resende 2018[18] Manuscript	Barretos, Brazil	Retrospective cohort, Single center institution	2003–2015	Prevention Institute at Barretos Cancer Hospital	NR	I, II, III, IV	Cervical cancer diagnosis	NR

Study / publication type	Location	Study design & data source type	Time-frame	Data source	Classification criteria ^a	Stages of cervical cancer included	Population	Total patients with cervical cancer
Warner 2018[27] Manuscript	Trinidad and Tobago	Retrospective cohort, National registry	1995–2009	Dr. Elizabeth Quamina Cancer Registry (aka National Cancer Registry of Trinidad and Tobago)	ICD-10: C53 NR	Localized, regional, distant	Any cancer diagnosis	1812
Nathani 2012[37] Congress abstract	Bradford, UK	Retrospective cohort, Single center institution	2007–2011	Bradford Royal Infirmary database	NR	IA1, IA2, IB1, III	Cervical cancer diagnosis Aged 19–30 years	19
Garry 2018[31] Congress abstract	Dublin, Ireland	Retrospective cohort, Single center institution	2006–2015	Electronic case report forms from a tertiary oncology center	NR FIGO staging	IA, IB, II, III, IV	Cervical cancer diagnosis Aged ≥60 years	119
Lorin 2015[13] Manuscript	Côte-d'Or, France	Retrospective cohort, Regional registry	1998–2010	Côte d'Or gynecological registry	NR FIGO staging	I, II, III, IV	Invasive cervical cancer	311
Rottmann 2020[40] Congress abstract + poster	Upper Bavaria, Germany	Retrospective cohort, Regional registry	2007–2016	Munich Cancer Registry	NR	IA1–IV, M1	Cervical cancer diagnosis	2291
Litvinova 2017[34] Congress abstract	Minsk City, Belarus	Retrospective cohort, National registry	2012–2016	National Cancer Registry	NR	IIB, III, IVA	Unresectable cervical cancer diagnosis Only young women discussed for proportions of disease by stage	324
Samson 2016[19] Manuscript	Bulgaria	Retrospective cohort, National registry	1993–2013	Bulgarian National Cancer Registry	ICD-O: C53.0, C53.1, C53.8, and C53.9	I, II, III, IV	Cervical cancer diagnosis	21 737
Ojamaa 2018[16] Manuscript	Estonia	Retrospective cohort, National registry	1968–2014	Estonian Cancer Registry	ICD-O-3: C53.0; C53.1, C53.8, and C53.9	I, II, III, IV	Invasive cervical cancer	3403

Study / publication type	Location	Study design & data source type	Time-frame	Data source	Classification criteria ^a	Stages of cervical cancer included	Population	Total patients with cervical cancer
					TNM (AJCC 7 th ed) for staging			
Ulinskas 2013[25] Manuscript	Lithuania	Retrospective cohort, National registry	1990–2004	Lithuanian cancer registry	ICD-10: C53.0, C53.1, C53.8, and C53.9	I, II, III, IV	Cervical cancer diagnosis	6680
Kosgei 2018[33] Congress abstract	Uasin Gishu, Kenya	Retrospective cohort, Regional registry	2010–2014	Eldoret Cancer Registry	NR	I, II, III, IV	Cervical cancer diagnosis	265
Bouchbika 2013[4] Manuscript	Casablanca, Morocco	Retrospective cohort, Regional registry	2005–2007	Greater Casablanca Registry	ICD-O-3, converted to ICD- 10: C53	Localized, regional, distant	Any cancer diagnosis	816
Elmajjaoui 2016[9] Manuscript	Morocco	Retrospective cohort, Multiple institutions	2006	National Institute of Oncology, Mohammed V Hospital, Rabat Cheikh Khalifa Ibn Zaid Hospital, Université Mohammed VI des Sciences de la Santé, Casablanca	NR FIGO staging	I, II, III, IV	Invasive cervical cancer	646
Somdyala 2020[23] Manuscript	Eastern Cape Province, South Africa	Retrospective cohort, Regional registry	1998–2012	Eastern Cape Cancer Registry	ICD-O: C53.0– C53.9	I, II, III, IV	Cervical cancer diagnosis	1315
Sharkas 2017[20] Manuscript	Jordan	Retrospective cohort, National registry	2000–2013	Jordan Cancer Registry	ICD-10: C53 TNM staging	Localized, regional, distant	Cervical cancer diagnosis Only women who were "Jordanian"	591
Kaidarova 2018[32] Congress abstract	Kazakhstan	Retrospective cohort, National registry	2012	Kazakhstan Cancer Registry	NR	IA, IB, IIA, IIB, III	Cervical cancer diagnosis	1641

Study / publication type	Location	Study design & data source type	Time-frame	Data source	Classification criteria ^a	Stages of cervical cancer included	Population	Total patients with cervical cancer
Alkhalawi 2019[3] Manuscript	Gulf countries	Retrospective cohort, Multinational registry	1998–2012	Gulf Centre for Cancer Control and Prevention Database	ICD-O-3: C53.0, C53.2, C53.8, C53.9	Localized, regional, distant	Invasive cervical cancer	2332
					SEER summary staging			
Agarwal 2012[2] Manuscript	Delhi, India	Retrospective cohort, Single center institution	2000–2009	Guru Teg Bahadur Hospital	NR FIGO staging	I, II, III, IV	Any primary gynecologic cancer diagnosis	927
Shruthi 2014[21] Manuscript	Kolar, India	Retrospective cohort, Single center institution	NR 1-year period	Sri Devaraj Urs Medical College, Sri Devaraj Urs Academy of Higher Education and Research	NR TNM staging	I, II, III, IV	Cervical cancer diagnosis	199
Wang 2015[26] Manuscript	Beijing, China	Retrospective cohort, Regional registry	1993–2008	Statistics Database of Beijing Cancer Registry	ICD-O FIGO staging	I, II, III, IV	Cervical cancer diagnosis Beijing residents only	3641
Cheung 2011[7] Manuscript	Hong Kong, China	Retrospective cohort, Regional registry	1997–2006	Hong Kong Cancer Registry	NR FIGO and TNM staging	I, II, III, IV	Cervical cancer diagnosis	4407
Yagi 2019[28] Manuscript	Osaka Prefecture, Japan	Retrospective cohort, Regional registry	1976–2012	Osaka Cancer Registry	C53, C54, C55 (C55 later sorted to C53 or C54 using a multiple imputation estimation)	Localized (T1N0M0), regional lymph nodes (N1), adjacent organs (T2, 3, 4), distant (M1)	Cervical cancer diagnosis	25 826
Seol 2014[30] Congress abstract	Korea	Retrospective cohort, National registry	1999–2010 (total population) 1999-2004 (with stage information)	Korea Central Cancer Registry	NR FIGO staging	IA1-IVB	Cervical cancer diagnosis	49 503 (total population) 19 282 (with stage information)

Study / publication type	Location	Study design & data source type	Time-frame	Data source	Classification criteria ^a	Stages of cervical cancer included	Population	Total patients with cervical cancer
				Gynecologic Oncology Committee of Korean Society of Obstetrics and Gynecology				
Chiang 2016[8] Manuscript	Taiwan	Retrospective cohort, National registry	2002–2012	Taiwan Cancer Registry	ICD-O-3: C53 TNM staging	I, II, III, IV	Any invasive cancerAge ≥15 years	8238

^aTwo types of classifications were found in the included studies. Disease coding classification criteria was used to identify patients with cervical cancer in large registries and databases and included different versions of the ICD or ICD-O criteria. The specific codes used to identify cervical cancer patients are also summarized where available. The second classification types found in the included studies were used to determine the stage of disease, and included FIGO, TNM, SEER summary, and Collaborative Stage criteria.

AC, adenocarcinoma; AJCC, American Joint Committee on Cancer; FIGO, International Federation of Gynecology and Obstetrics; HPV, human papillomavirus; *ICD-10, International Classification of Diseases, 10th edition; ICD-O, International Classification of Diseases, Oncology*; NR, not reported; PAYA, pediatric and young adult cancers; RH, radical hysterectomy; RT, radiotherapy; SCC, squamous cell carcinoma; SEER, Surveillance, Epidemiology, and End Results; TNM, tumor, node, metastasis; USA, United States of America; WHO, World Health Organization.

Supplementary Table 6. Studies reporting incidence by stage of cervical cancer.

Reference	Region	Location / data collection period	N	Incidence of cervical cancer by stage	Incidence of locally advanced cervical cancer
Bruegl 2020[5] ^a	North America	USA Idaho, Oregon, Washington 1996–2016	7222	Age- standardized rate per 100 000 population	Age- standardized rate per 100 000 population
		1000 2010		American Indian/Alaskan Natives Localized, 4.3 Regional, 3.6 Distant, 1.8 Unknown, 0.9	American Indian/Alaskan Natives Regional, 3.6
				Non-Hispanic White Localized, 3.7 Regional, 2.0 Distant, 0.7 Unknown, 0.5	<i>Non-Hispanic</i> <i>White</i> Regional, 2.0
Henley 2010[11]	North America	USA 2004–2006	36,076	Age- standardized rate per 100 000 population	Age- standardized rate per 100 000 population
				Localized, 5.3 Regional, 4.0 Distant, 1.2 Unknown, 0.9	Regional, 4.0
Zahnd 2018[29] ^b	North America	USA 2009–2013	Not reported	Age- standardized rate per 100 000 population	Not calculable
				Rural Localized, 3.7 Distant, 1.1	
				<i>Urban</i> Localized, 3.4 Distant, 1.0	
McClean 2012[36]	North America	USA 1992–2007	6718	Women aged 65–100 years, Age-adjusted incidence rate	Women aged 65–100 years, Age-adjusted incidence rate
				Stage I, decreased by 2.4% per year	Stage III, increased by 2.0% per year

				Stage III, increased by 2.0% per year	
Litvinova 2017[34]	Europe	Belarus Minsk City 2012–2016	324	Incidence per 100 000 female population	Incidence per 100 000 female population
				IIB, decreased from 3.8 to 1.9 III, decreased from 3.2 to 2.3 IVA, increased from 0.4 to 0.7	IIB, decreased from 3.8 to 1.9 III, decreased from 3.2 to 2.3 IVA, increased from 0.4 to 0.7

^aThe Bruegl 2020 study only included patients who were *American Indian/Alaskan Natives* or non-Hispanic White.

USA, United States of America.

^bIn the Zahnd 2018 study, only the incidence of localized and distant cervical cancer was compared in urban and rural areas; neither of these stages was considered locally advanced disease according to our method of estimation (ie, only "regional" disease is considered).

Section and Topic	Item #	PRISMA Checklist item	Location where item is reported
TITLE			
Title	1	Identify the report as a systematic review.	Page 1
ABSTRACT			
Abstract	2	See the PRISMA 2020 for Abstracts checklist.	Page 3
INTRODUCTION	1		
Rationale	3	Describe the rationale for the review in the context of existing knowledge.	Page 5
Objectives	4	Provide an explicit statement of the objective(s) or question(s) the review addresses.	Page 5
METHODS			
Eligibility criteria	5	Specify the inclusion and exclusion criteria for the review and how studies were grouped for the syntheses.	Pages 5-6, Supplementary Appendix pages 4-5
Information sources	6	Specify all databases, registers, websites, organisations, reference lists and other sources searched or consulted to identify studies. Specify the date when each source was last searched or consulted.	Page 5, Supplementary Appendix pages 1-3
Search strategy	7	Present the full search strategies for all databases, registers and websites, including any filters and limits used.	Supplementary Appendix pages 1-3
Selection process	8	Specify the methods used to decide whether a study met the inclusion criteria of the review, including how many reviewers screened each record and each report retrieved, whether they worked independently, and if applicable, details of automation tools used in the process.	Page 6
Data collection process	9	Specify the methods used to collect data from reports, including how many reviewers collected data from each report, whether they worked independently, any processes for obtaining or confirming data from study investigators, and if applicable, details of automation tools used in the process.	Page 6
Data items	10a	List and define all outcomes for which data were sought. Specify whether all results that were compatible with each outcome domain in each study were sought (e.g. for all measures, time points, analyses), and if not, the methods used to decide which results to collect.	Supplementary Appendix page 5
	10b	List and define all other variables for which data were sought (e.g. participant and intervention characteristics, funding sources). Describe any assumptions made about any missing or unclear information.	Supplementary Appendix pages 5
Study risk of bias assessment	11	Specify the methods used to assess risk of bias in the included studies, including details of the tool(s) used, how many reviewers assessed each study and whether they worked independently, and if applicable, details of automation tools used in the process.	N/A
Effect measures	12	Specify for each outcome the effect measure(s) (e.g. risk ratio, mean difference) used in the synthesis or presentation of results.	N/A
Synthesis methods	13a	Describe the processes used to decide which studies were eligible for each synthesis (e.g. tabulating the study intervention characteristics and comparing against the planned groups for each synthesis (item #5)).	Supplementary Appendix page 5

Section and Topic	Item #	PRISMA Checklist item	Location where item is reported
	13b	Describe any methods required to prepare the data for presentation or synthesis, such as handling of missing summary statistics, or data conversions.	Page 6-7, Supplementary Appendix page 5-6
	13c	Describe any methods used to tabulate or visually display results of individual studies and syntheses.	Page 6-7, Supplementary Appendix page 5-6
	13d	Describe any methods used to synthesize results and provide a rationale for the choice(s). If meta-analysis was performed, describe the model(s), method(s) to identify the presence and extent of statistical heterogeneity, and software package(s) used.	Page 6-7, Supplementary Appendix page 5-6
	13e	Describe any methods used to explore possible causes of heterogeneity among study results (e.g. subgroup analysis, meta-regression).	Supplementary Appendix page 8
	13f	Describe any sensitivity analyses conducted to assess robustness of the synthesized results.	N/A
Reporting bias assessment	14	Describe any methods used to assess risk of bias due to missing results in a synthesis (arising from reporting biases).	N/A
Certainty assessment	15	Describe any methods used to assess certainty (or confidence) in the body of evidence for an outcome.	N/A
RESULTS			
Study selection	16a	Describe the results of the search and selection process, from the number of records identified in the search to the number of studies included in the review, ideally using a flow diagram.	Page 7 and Figure 1
	16b	Cite studies that might appear to meet the inclusion criteria, but which were excluded, and explain why they were excluded.	N/A
Study characteristics	17	Cite each included study and present its characteristics.	Pages 7-8, Supplementary figure 1, Supplementary Tables 3, 4, 5
Risk of bias in studies	18	Present assessments of risk of bias for each included study.	N/A
Results of individual studies	19	For all outcomes, present, for each study: (a) summary statistics for each group (where appropriate) and (b) an effect estimate and its precision (e.g. confidence/credible interval), ideally using structured tables or plots.	Pages 8-11, Figure 2, Tables 1-3, Supplementary Figure 2

Section and Topic	Item #	PRISMA Checklist item	Location where item is reported
Results of syntheses	20a	For each synthesis, briefly summarise the characteristics and risk of bias among contributing studies.	Pages 8-11, Figure 2, Tables 1-3
	20b	Present results of all statistical syntheses conducted. If meta-analysis was done, present for each the summary estimate and its precision (e.g. confidence/credible interval) and measures of statistical heterogeneity. If comparing groups, describe the direction of the effect.	Pages 8-11, Figure 2, Tables 1-3, Supplementary Figure 2
	20c	Present results of all investigations of possible causes of heterogeneity among study results.	Page 9, Supplementary Figure 2
	20d	Present results of all sensitivity analyses conducted to assess the robustness of the synthesized results.	N/A
Reporting biases	21	Present assessments of risk of bias due to missing results (arising from reporting biases) for each synthesis assessed.	N/A
Certainty of evidence	22	Present assessments of certainty (or confidence) in the body of evidence for each outcome assessed.	N/A
DISCUSSION			
Discussion	23a	Provide a general interpretation of the results in the context of other evidence.	Pages 11-12
	23b	Discuss any limitations of the evidence included in the review.	Page 12-13
	23c	Discuss any limitations of the review processes used.	Page 12-13
	23d	Discuss implications of the results for practice, policy, and future research.	Page 13
OTHER INFORMA	TION		
Registration and	24a	Provide registration information for the review, including register name and registration number, or state that the review was not registered.	Page 6
protocol	24b	Indicate where the review protocol can be accessed, or state that a protocol was not prepared.	Page 6, Supplementary Appendix page 1
	24c	Describe and explain any amendments to information provided at registration or in the protocol.	N/A
Support	25	Describe sources of financial or non-financial support for the review, and the role of the funders or sponsors in the review.	Page 15
Competing interests	26	Declare any competing interests of review authors.	Page 15
Availability of data, code and other materials	27	Report which of the following are publicly available and where they can be found: template data collection forms; data extracted from included studies; data used for all analyses; analytic code; any other materials used in the review.	N/A

From: Page MJ, McKenzie JE, Bossuyt PM, Boutron I, Hoffmann TC, Mulrow CD, et al. The PRISMA 2020 statement: an updated guideline for reporting systematic reviews. BMJ 2021;372:n71. doi: 10.1136/bmj.n71

For more information, visit: http://www.prisma-statement.org/

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