APPENDIX A: SEARCH STRATEGY

SYSTEMATIC REVIEWS

1. Search for systematic reviews					
A. Bibliographic Databases:	-20-22; #	# Search Statement			
MEDLINE: Systematic Reviews Ovid MEDLINE(R) ALL 1946 to October 19, 2022	1	exp Orthopedic Procedures/ OR (orthop?edic procedure\$ OR orthop?edic surger\$ OR orthop?edic operation\$ OR arthroplast\$ OR hip replacement\$ OR knee replacement\$ OR spine fusion\$ OR carpal tunnel\$ OR spine decompression\$ OR rotator cuff OR meniscectom\$ OR meniscal repair OR anterior cruciate ligament OR ACL OR finger replacement\$ OR joint replacement\$ OR joint implantation\$ OR ankle replacement\$ OR elbow replacement\$ OR shoulder replacement\$ OR spine surgery OR lumbar surgery OR spondylodes\$).ti,ab,kw.	452810		
	2	Smoking Cessation/ OR exp "Tobacco Use Disorder"/ OR Smoking/ OR (smoking cessation OR nicotine patch OR nicotine polacrilex OR nicotine inhalant\$ OR nicotine device\$ OR nicotine nasal spray OR nicotine replacement\$ OR nicotine lozenge\$ OR smoking habit\$ OR smoking behavio?r\$ OR nicotine addiction\$ OR nicotine dependence\$ OR (nicotine disorder\$) OR (tobacco disorder\$) OR (smoking adj1 (quit OR quitting OR stop OR stopping or giving up))).ti,ab,kw.	190965		
	3	1 and 2	1106		
	4	(systematic review.ti. or meta-analysis.pt. or meta-analysis.ti. or systematic literature review.ti. or this systematic review.tw. or pooling project.tw. or (systematic review.ti,ab. and review.pt.) or meta synthesis.ti. or meta-analy*.ti. or integrative review.tw. or integrative research review.tw. or rapid review.tw. or umbrella review.tw. or consensus development conference.pt. or practice guideline.pt. or drug class reviews.ti. or cochrane database syst rev.jn. or acp journal club.jn. or health technol assess.jn. or evid rep technol assess summ.jn. or jbi database system rev implement rep.jn. or (clinical guideline and management).tw. or ((evidence based.ti. or evidence-based medicine/ or best practice*.ti. or evidence synthesis.ti,ab.) and (((review.pt. or diseases category/ or behavior.mp.) and behavior mechanisms/) or therapeutics/ or evaluation studies.pt. or validation studies.pt. or guideline.pt. or pmcbook.mp.)) or (((systematic or systematically).tw. or critical.ti,ab. or study selection.tw. or ((predetermined or inclusion) and criteri*).tw. or exclusion criteri*.tw. or main outcome measures.tw. or standard of care.tw. or standards of care.tw.) and ((survey or surveys).ti,ab. or overview*.tw. or review.ti,ab. or reviews.ti,ab. or appraisal.tw. or (reduction.tw. and (risk/ or risk.tw.) and (death or recurrence).mp.)) and ((literature or articles or publications or publication or bibliography or bibliographies or published).ti,ab. or references.tw. or scales.tw. or nates.tw. or tatabase.tw. or trials.ti,ab. or meta-analy*.tw. or (clinical and studies).ti,ab. or	584041		



		treatment outcome/ or treatment outcome.tw. or pmcbook.mp.))) not (letter or newspaper article).pt.	
	5 3 and 4		59
	6	limit 5 to english language	59
CDSR: Protocols	1	(Orthopedic Procedures).kw	14
and Reviews EBM Reviews - Cochrane Database of Systematic Reviews 2005 to October 19, 2022	2	(orthopedic procedure\$ OR orthopedic surger\$ OR orthopedic operation\$ OR arthroplast\$ OR hip replacement\$ OR knee replacement\$ OR spine fusion\$ OR carpal tunnel\$ OR spine decompression\$ OR rotator cuff OR meniscectom\$ OR meniscal repair OR anterior cruciate ligament OR ACL OR finger replacement\$ OR joint replacement\$ OR joint implantation\$ OR ankle replacement\$ OR elbow replacement\$ OR shoulder replacement\$ OR spine surgery OR lumbar surgery OR spondylodes\$).ti,ab,kw.	
	3	1 OR 2	149
	4	(Smoking Cessation OR "Tobacco Use Disorder" OR Smoking).kw.	122
	5	(smoking cessation OR nicotine patch OR nicotine polacrilex OR nicotine inhalant\$ OR nicotine device\$ OR nicotine nasal spray OR nicotine replacement\$ OR nicotine lozenge\$ OR smoking habit\$ OR smoking behavio?r\$ OR nicotine addiction\$ OR nicotine dependence\$ OR (nicotine disorder\$) OR (tobacco disorder\$) OR (smoking adj1 (quit OR quitting OR stop OR stopping or giving up))).ti,ab,kw.	124
	6	4 OR 5	140
	7	3 AND 6	1

2. Search for current systematic reviews							
Date Searched: 1	Date Searched: 10-20-22						
B. Non- bibliographic databases	Evidence	<u>Results</u>					
AHRQ: evidence reports, technology assessments, U.S Preventative Services Task Force Evidence Synthesis	http://www.ahrq.gov/research/findings/evidence-based- reports/search.html Search: (orthopedic surgery OR arthroplasty OR replacement) AND smoking	0					
CADTH	https://www.cadth.ca Search: (orthopedic surgery OR arthroplasty OR replacement) AND smoking	0					
HTA: Health Technology Assessments (UP TO 2016)	http://www.ohsu.edu/xd/education/library/ See CDSR search	0					
EPPI-Centre	http://eppi.ioe.ac.uk/cms/Default.aspx?tabid=62	0					



	Use browser search function [CNTL + F] for keyword search		
	Search: smoking		
NLM	http://www.ncbi.nlm.nih.gov/books	0	
	Search: (orthopedic surgery OR arthroplasty OR replacement) AND smoking		
VA Products -	A. http://www.hsrd.research.va.gov/research/default.cfm	0	
and HSR&D	B. <u>http://www.research.va.gov/research_topics/</u>		
	Search: (orthopedic surgery OR arthroplasty OR replacement) AND smoking		
ClinicalTrials.gov	https://www.clinicaltrials.gov/	5	
23)	Search: (orthopedic surgery OR arthroplasty OR joint replacement OR finger replacement OR shoulder replacement OR ankle replacement OR elbow replacement OR knee replacement OR ACL) AND smoking		
	Smoking Cessation Study		
	Surgeon-Patient Communication: A Randomized Controlled Trial Evaluating the Impact of an Orthopaedic Related Smoking Cessation Discussion After Fracture Surgery		
	Let's STOP Now Trial: Smoking in Trauma Orthopaedic Patients		
	The Impact of Orthopaedic Smoking Cessation Education on Cigarette Abstinence Self-Efficacy: A Randomized Controlled Trial		
	Smoking Cessation Intervention in Acute Orthopedic Surgery		
3. Search for sys protocols)	tematic reviews currently under development (includes forthcoming re	eviews &	
D. Under	Evidence:	Results:	
development:			
AHRQ topics in	Email Charli Armstrong Charlotte.Armstrong1@va.gov	0	
development (EPC Status Report)	Emailed on 10-20-22		
PROSPERO	http://www.crd.york.ac.uk/PROSPERO/	0	
(SR registry)	(orthopedic surgery OR arthroplasty OR replacement) AND smoking cessation		

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PRIMARY STUDIES

4. Se	earch for primary literature	
Date	searched: 11-03-22; updated 05-19-2023	
MED	LINE [OVID MEDLINE(R) ALL 1946 to November 02, 2022]	D K
#	Search Statement	Results
1	exp Orthopedic Procedures/ OR (orthop?edic procedure\$ OR orthop?edic surger\$ OR orthop?edic operation\$ OR arthroplast\$ OR hip replacement\$ OR knee replacement\$ OR spine fusion\$ OR carpal tunnel\$ OR spine decompression\$ OR rotator cuff OR meniscectom\$ OR meniscal repair OR anterior cruciate ligament OR ACL OR finger replacement\$ OR joint replacement\$ OR joint implantation\$ OR ankle replacement\$ OR elbow replacement\$ OR shoulder replacement\$ OR spine surgery OR lumbar surgery OR spondylodes\$).ti,ab,kw.	452810
2	Smoking Cessation/ OR (smoking cessation OR nicotine patch OR nicotine polacrilex OR nicotine inhalant\$ OR nicotine device\$ OR nicotine nasal spray OR nicotine replacement\$ OR nicotine lozenge\$ OR smoking behavio?r\$ OR (smoking adj1 (quit OR quitting OR stop OR stopping or giving up))).ti,ab,kw.	54598
3	1 and 2	236
4	limit 3 to English language	215
CINA	AL [EBSCO CINAHL Plus with Full Text]	
#	Search Statement	Results
1	(MH "Orthopedic Surgery+") OR TI ((orthop?edic procedure\$ OR orthop?edic surger\$ OR orthop?edic operation\$ OR arthroplast\$ OR hip replacement\$ OR knee replacement\$ OR spine fusion\$ OR carpal tunnel\$ OR spine decompression\$ OR rotator cuff OR meniscectom\$ OR meniscal repair OR anterior cruciate ligament OR ACL OR finger replacement\$ OR joint replacement\$ OR joint implantation\$ OR ankle replacement\$ OR elbow replacement\$ OR shoulder replacement\$ OR spine surgery OR lumbar surgery OR spondylodes\$)) OR AB ((orthop?edic procedure\$ OR orthop?edic surger\$ OR orthop?edic operation\$ OR arthroplast\$ OR hip replacement\$ OR knee replacement\$ OR spine fusion\$ OR carpal tunnel\$ OR spine decompression\$ OR rotator cuff OR meniscectom\$ OR meniscal repair OR anterior cruciate ligament OR ACL OR finger replacement\$ OR joint replacement\$ OR spine decompression\$ OR rotator cuff OR meniscectom\$ OR meniscal repair OR anterior cruciate ligament OR ACL OR finger replacement\$ OR joint replacement\$ OR spine decompression\$ OR rotator cuff OR meniscectom\$ OR meniscal repair OR anterior cruciate ligament OR ACL OR finger replacement\$ OR joint replacement\$ OR joint implantation\$ OR ankle replacement\$ OR elbow replacement\$ OR shoulder replacement\$ OR spine surgery OR lumbar surgery OR spondylodes\$))	157792
2	(MH "Smoking Cessation") OR TI ((smoking cessation OR nicotine patch OR nicotine polacrilex OR nicotine inhalant\$ OR nicotine device\$ OR nicotine nasal spray OR nicotine replacement\$ OR nicotine lozenge\$ OR smoking behavio?r\$ OR (smoking N1 (quit OR quitting OR stop OR stopping or giving up)))) OR AB ((smoking cessation OR nicotine patch OR nicotine polacrilex OR nicotine inhalant\$ OR nicotine device\$ OR nicotine device\$ OR nicotine device\$ OR nicotine device\$ OR nicotine nasal spray OR nicotine replacement\$ OR nicotine inhalant\$ OR nicotine device\$ OR nicotine nasal spray OR nicotine replacement\$ OR nicotine lozenge\$ OR smoking behavio?r\$ OR (smoking N1 (quit OR quitting OR stop OR stopping or giving up))))	30384
3	1 AND 2	138
4	limit 3 to English language	137

APPENDIX B: EXCLUDED STUDIES

Exclude reasons: 1=Ineligible population, 2=Ineligible intervention, 3=Ineligible comparator, 4=Ineligible outcome, 5=Ineligible timing, 6=Ineligible study design, 7=Ineligible publication type, 8=Outdated or ineligible systematic review.

Citation	Exclude Reason
Cut it out sooner than later. Nursing Review (1173-8014). 2014;14(3):16-16.	E7
Akhavan S, Nguyen LC, Chan V, Saleh J, Bozic KJ. Impact of smoking cessation counseling prior to total joint arthroplasty. Orthopedics. 2017;40(2):e323-e328.	E4
Alamanda VK, Springer BD. The prevention of infection: 12 modifiable risk factors. Bone & Joint Journal. 2019;101-B(1_Supple_A):3-9.	E8
Beaupre LA, Hammal F, DeSutter C, Stiegelmar RE, Masson E, Finegan BA. Impact of a standardized referral to a community pharmacist-led smoking cessation program before elective joint replacement surgery. Tobacco Induced Diseases. 2019;17:14.	E4
Bishop JY, Santiago-Torres JE, Rimmke N, Flanigan DC. Smoking predisposes to rotator cuff pathology and shoulder dysfunction: A systematic review. Arthroscopy. 2015;31(8):1598-1605.	E8
Boylan MR, Bosco JA, 3rd, Slover JD. Cost-effectiveness of preoperative smoking cessation interventions in total joint arthroplasty. Journal of Arthroplasty. 2019;34(2):215-220.	E4
Chen J, Cui Y, Li X, et al. Risk factors for deep infection after total knee arthroplasty: A meta-analysis. Arch Orthop Trauma Surg. 2013;133(5):675-687.	E8
Davis EM, Notier AE, Douaihy A, et al. A telemedicine pathway to increase tobacco cessation in patients undergoing total joint replacement surgery. NEJM Catalyst Innovations in Care Delivery. 2021;2(3):1-1.	E4
Deguchi M, Rapoff AJ, Zdeblick TA. Posterolateral fusion for isthmic spondylolisthesis in adults: Analysis of fusion rate and clinical results. Journal of Spinal Disorders. 1998;11(6):459-464.	E2
Glassman SD, Anagnost SC, Parker A, Burke D, Johnson JR, Dimar JR. The effect of cigarette smoking and smoking cessation on spinal fusion. Spine. 2000;25(20):2608-2615.	E3
Grau L, Orozco FR, Duque AF, Post ZD, Ponzio DY, Ong AC. A simple protocol to stratify pulmonary risk reduces complications after total joint arthroplasty. Journal of Arthroplasty. 2019;34(6):1233-1239.	E2
Grønkjær M, Eliasen M, Skov-Ettrup LS, et al. Preoperative smoking status and postoperative complications: A systematic review and meta-analysis. Ann Surg. 2014;259(1):52-71.	E8
Iannotti F, Prati P, Fidanza A, et al. Prevention of periprosthetic joint infection (pji): A clinical practice protocol in high-risk patients. Tropical Medicine & Infectious Disease. 2020;5(4):11.	E7
Jackson KL, 2nd, Devine JG. The effects of smoking and smoking cessation on spine surgery: A systematic review of the literature. Global Spine Journal. 2016;6(7):695-701.	E8

Citation	Exclude Reason
Kainiemi K, Malmivaara A, Sillman-Tetri S, et al. Smoking cessation intervention prior to orthopedic surgery: A study protocol to determine patient outcomes and feasibility. Tobacco Prevention & Cessation. 2022;8:33.	E3
Kang J, Shi C, Gu Y, Yang C, Gao R. Factors that may affect outcome in cervical artificial disc replacement: A systematic review. European Spine Journal. 2015;24(9):2023-2032.	E8
Kanneganti P, Harris JD, Brophy RH, Carey JL, Lattermann C, Flanigan DC. The effect of smoking on ligament and cartilage surgery in the knee: A systematic review. Am J Sports Med. 2012;40(12):2872-2878.	E8
Kim JH, Patel S. Is it worth discriminating against patients who smoke? A systematic literature review on the effects of tobacco use in foot and ankle surgery. Journal of Foot & Ankle Surgery. 2017;56(3):594-599.	E8
Kozak E. A preoperative smoking intervention decreased postoperative complications in elective knee or hip replacement. ACP Journal Club. 2002;137(1):7-7.	E7
Kunutsor SK, Whitehouse MR, Blom AW, Beswick AD. Re-infection outcomes following one- and two-stage surgical revision of infected hip prosthesis: A systematic review and meta-analysis. PLoS One. 2015;10(9):e0139166.	E8
Kwon BK, Hilibrand AS, Malloy K, et al. A critical analysis of the literature regarding surgical approach and outcome for adult low-grade isthmic spondylolisthesis. Journal of Spinal Disorders & Techniques. 2005;18 Suppl:S30-40.	E8
Lambers Heerspink FO, Dorrestijn O, van Raay JJ, Diercks RL. Specific patient- related prognostic factors for rotator cuff repair: A systematic review. J Shoulder Elbow Surg. 2014;23(7):1073-1080.	E8
Lee MS, Jimenez AE, Owens JS, et al. Comparison of outcomes between nonsmokers and patients who discontinued smoking 1 month before primary hip arthroscopy: A propensity-matched study with minimum 2-year follow-up. Orthopaedic Journal of Sports Medicine. 2022;10(6):1-8.	E2
Lindstrom D, Sadr Azodi O, Wladis A, et al. Effects of a perioperative smoking cessation intervention on postoperative complications: A randomized trial. Annals of Surgery. 2008;248(5):739-745.	E1
Liu J, Hai Y, Kang N, Chen X, Zhang Y. Risk factors and preventative measures of early and persistent dysphagia after anterior cervical spine surgery: A systematic review. European Spine Journal. 2018;27(6):1209-1218.	E8
Mak JC, Fransen M, Jennings M, et al. Evidence-based review for patients undergoing elective hip and knee replacement. ANZ Journal of Surgery. 2014;84(1-2):17-24.	E8
Malay DS. Cigarette smoking and foot and ankle surgery. Journal of Foot & Ankle Surgery. 2011;50(5):515-516.	E7
Meng J, Sun T, Zhang F, Qin S, Li Y, Zhao H. Deep surgical site infection after ankle fractures treated by open reduction and internal fixation in adults: A retrospective case-control study. International Wound Journal. 2018;15(6):971- 977.	E1
Murray EW. A preoperative smoking intervention decreased postoperative complications in elective knee or hip replacement. Evidence Based Nursing. 2002;5(3):84-84.	E7

₩ 4

Citation	Exclude Reason
Papas PV, Gold CA, Scuderi GR. The effects of smoking on postoperative complications after total joint arthroplasty. Techniques in Orthopaedics. 2019;34(3):183-186.	E7
Paulsen K, Carender CN, Noiseux NO, Elkins JM, Brown TS, Bedard NA. What is the fate of total joint arthroplasty patients who are asked to quit smoking prior to surgery? Arthroplasty Today. 2023;19:101087.	E3
Pearson RG, Clement RG, Edwards KL, Scammell BE. Do smokers have greater risk of delayed and non-union after fracture, osteotomy and arthrodesis? A systematic review with meta-analysis. BMJ Open. 2016;6(11):e010303.	E8
Ring J, Shoaib A, Shariff R. Smoking cessation advice in limb reconstruction: An opportunity not to be missed. Injury. 2017;48(2):345-348.	E1
Santiago-Torres J, Flanigan DC, Butler RB, Bishop JY. The effect of smoking on rotator cuff and glenoid labrum surgery: A systematic review. American Journal of Sports Medicine. 2015;43(3):745-751.	E8
Sullivan JP, Huston LJ, Zajichek A, et al. Incidence and predictors of subsequent surgery after anterior cruciate ligament reconstruction: A 6-year follow-up study. American Journal of Sports Medicine. 2020;48(10):2418-2428.	E2
Truntzer J, Comer G, Kendra M, Johnson J, Behal R, Kamal RN. Perioperative smoking cessation and clinical care pathway for orthopaedic surgery. JBJS Reviews. 2017;5(8):e11.	E7
Verweij JP, Houppermans PN, Gooris P, Mensink G, van Merkesteyn JP. Risk factors for common complications associated with bilateral sagittal split osteotomy: A literature review and meta-analysis. Journal of Cranio-Maxillo-Facial Surgery. 2016;44(9):1170-1180.	E8
Zhu Y, Zhang F, Chen W, Liu S, Zhang Q, Zhang Y. Risk factors for periprosthetic joint infection after total joint arthroplasty: A systematic review and meta-analysis. J Hosp Infect. 2015;89(2):82-89.	E8

APPENDIX C: EVIDENCE TABLES

CHARACTERISTICS OF INCLUDED PRIMARY STUDIES

Experimental Studies

Author Year	Study Design	Participant Characteristics	Surgery Type	Intervention	Comparator
	Follow-up				
N		Duration and Frequency of Smoking			
Moller 2002 ¹⁶	RCT	Mean age 65Primary elective hip orCour42.59% maleknee alloplasty	Counseling and NRT	Standard care (little or no information about the	
Moller 4 v 2003 ¹⁷	4 weeks	Race/ethnicity NR			risk of tobacco smoking or smoking cessation
N=120		Pack years (smoking years X daily consumption): 35 years for intervention group, 37 years for control group. 15 cigarettes per day for both groups			

Abbreviations. NR=not reported; NRT=nicotine replacement therapy; RCT=randomized controlled trial.

Observational Studies

Author Year	Study Design	Participant Characteristics	Surgery Type	Intervention	Comparator
	Follow-up				
Ν	·	Duration and Frequency of Smoking			
Herrero 2020 ¹⁸	Retrospective cohort	Mean age 57.6 51.74% male	Total joint arthroplasty	Smoking cessation program (assessment,	Patients who did not enroll in the voluntary
	90 days	34.83% Black		education, counseling,	smoking cessation
N=201		45.77% White		NRT)	program
		NR			

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Author Year	Study Design	Participant Characteristics	Surgery Type	Intervention	Comparator
N	Follow-up	Duration and			
		Frequency of Smoking			
Khalid 2022 ¹⁹	Retrospective cohort	Median age between 50- 54 years	Single-level anterior cervical discectomy and	Nicotine replacement (varenicline and/or	Patients with active smoking history
	90 days	39.26% male	fusion	nicotine gum, patches,	
N=5,769		Race/ethnicity NR		and lozenges)	
		NR			
Khalid 2022 ²⁰	Retrospective cohort	Median age between 45 and 64 years	Single-level lumbar fusion	Nicotine replacement (nicotine products or	Active smokers
	30 days, 90 days, or 2.5	39.58% male		varenicline along with	
N=31,935	years	Race/ethnicity NR		smoking cessation counseling)	
		NR			

Abbreviations. NR=not reported; NRT=nicotine replacement therapy.

OUTCOME DATA OF INCLUDED PRIMARY STUDIES

Experimental Studies

Author Year	Follow-up	Intervention	Complication Outcomes	Length of Stay Outcomes	Readmission/Revision Surgery Outcomes	
Moller 2002 ¹⁶ Moller 2003 ¹⁷	4 weeks	4 weeks	NRT and counseling	Relative risk (95% CI) (below 1 favors intervention): Any complications 0.34 (0.17- 0.58)	Median (range) length of stay 11 days (7-55) for intervention group and 13 days (8-65) for control group ($p = 0.41$)	Relative risk (95% CI): Secondary surgery 0.23 (0.05- 1.02)
			Wound-related complications 0.16 (0.05-0.52)	Rate of secondary surgery 4% for intervention group and 15% for control group ($p = 0.07$)		
			Overall complication rate 18% for intervention group and 52% for control group ($p = 0.0003$). Wound-related complication rate 5% for intervention group and 31% for control group ($p = 0.001$). Rates for other complication types were not significantly different between groups.			

Abbreviations. CI=confidence interval; NRT=nicotine replacement therapy.

Observational Studies

Author Year	Follow- up	Intervention	Complication Outcomes	Length of Stay Outcomes	Readmission/Revision Surgery Outcomes
Herrero 90 2020 ¹⁸	90 days	Smoking cessation program	Enrolled in SCP vs not enrolled: Infection rates 7.3% for SCP and 12.5% for non-SCP, $p = 0.27$	Enrolled in SCP vs not enrolled: Length of stay 2.47 days for SCP and 2.62 days for non- SCP, <i>p</i> = 0.52	Enrolled in SCP vs. not enrolled: Readmission rate 5.8% for SCP and 4.7% for non-SCP, $p = 0.73$
			Smoking vs quit (regardless of SCP enrollment):		Smoking vs. quit (regardless of SCP enrollment):
			Infection rates 3.7% for quitters and 12.5% for smokers, $p = 0.03$	Smoking vs quit (regardless of SCP enrollment): Length of stay 2.5 days for	Readmissions 4.9% for quitters and 9.2% for smokers, <i>p</i> = 0.26
				quitters and 2.6 days for smokers, <i>p</i> = 0.65	
Khalid 2022 ¹⁹	90 days	Nicotine replacement	Odds ratios (95% CI) for matched smoking <i>vs</i> smoking	NR	Readmission within 30 days 1.000 (0.691–1.447)
			cessation cohort: Composite surgical		Readmission within 90 days 0.932 (0.705–1.231)
			1.014)		Revision surgery 0.846 (0.726–
			Dysphagia 0.559 (0.330–0.947) Hematoma 1.589 (0.769–3.282)		0.988)
			Pseudarthrosis 1.034 (0.724– 1.479)		
			Instrumentation removal 0.750 (0.495–1.136)		

Author Year	Follow- up	Intervention	Complication Outcomes	Length of Stay Outcomes	Readmission/Revision Surgery Outcomes
Khalid 2022 ²⁰	halid 30 days, Nicotine Odds ratios (95% CI) for D22 ²⁰ 90 days, replacement matched smoking <i>vs</i> smoking		NR	Readmission within 30 days 1.026 (0.924-1.140)	
or 2.5		cessation cohort:		Readmission within 90 days	
	years Any complication 0.862 (0.804- 0.925) Pseudoarthritis 1.041 (0.929- 1.166)		1.102 (1.005-1.207)		
			Revision surgery 0.835 (0.694- 1.001)		
			1,811 complications ('any complication') in smoking cessation group compared to 2,045 in smoking group ($p < 0.05$). Group differences nonsignificant for pseudoarthritis.		Group differences non- significant for readmissions and revision surgery

Abbreviations. CI=confidence interval; NR=not reported; SCP=smoking cessation program.

QUALITY ASSESSMENT OF INCLUDED PRIMARY STUDIES

RCTs – RoB 2 Tool

Author Year	Risk of Bias from Randomization Process	Risk of Bias from Deviation from Intended Interventions (Assignment)	Risk of Bias from Deviation from Intended Interventions (Adherence)	Risk of Bias from Missing Outcome Data	Risk of Bias in Measurement of Outcome	Risk of Bias in Selection of Reported Result	Overall Bias (High, Low, Some Concerns)
Moller 2002 ¹⁶ Moller 2003 ¹⁷	Low Participants randomly assigned using stratified block randomization and assignments were concealed using opaque sealed envelopes. Baseline differences between groups do not suggest a problem with the randomization process.	Low Cannot blind participants or providers. Conducted ITT analysis that included all participants randomized except for 12 whose operations were postponed or cancelled.	Unclear Information about adherence to the intervention is not reported.	Unclear ITT analysis was conducted, but extent and handling of missing data is unclear.	Low An assessor masked to the intervention recorded all complications.	Unclear Protocol not identified.	Some concerns

Abbreviations. ITT=intent to treat.

Observational Studies – ROBINS-I Tool for Cohort Studies

Author Year	Selection Bias	Bias in Classification of Interventions	Bias Due to Departures from Intended Intervention	Bias Due to Measurement of Outcomes	Bias Due to Confounding	Bias Due to Missing Data	Bias in the Selection of Reported Results	Overall Bias (Low, Moderate, Serious, Critical, No Information)
Herrero 2020 ¹⁸	Serious Included consecutive patients meeting criteria identified from a database during a specified period. However, referral to the SCP was recommended but not required of attending surgeons and not all patients were offered the intervention.	Low Groups clearly defined and status obtained from medical records.	Moderate All patients who enrolled in SCP were included in SCP group, and a substantial number enrolled did not complete the program.	Low Complication outcomes were assessed and recorded as part of routine care.	Moderate Information was collected on smoking characteristics of sample but was not accounted for in analyses. No significant differences between groups were found for demographic variables collected.	No Information Unclear level and handling of missing data.	Low No pre-study protocol but appears to report outcomes and analyses mentioned in methods section.	Serious

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Auth Year	or Selection Bias	Bias in Classification of Interventions	Bias Due to Departures from Intended Intervention	Bias Due to Measurement of Outcomes	Bias Due to Confounding	Bias Due to Missing Data	Bias in the Selection of Reported Results	Overall Bias (Low, Moderate, Serious, Critical, No Information)
Khali 2022	 Low ⁹ Unclear if some characteristics used for matching/final selection were observed before or after start of NRT, but patients were matched exactly on these characteristics, but if bias exists, it seems likely to be non- differential. 	Low Information on NRT prescription provided in medical record and documented at start of intervention.	No information Adherence not reported; no information on how frequently patients skipped prescribed medication, were prescribed NRT and then stopped taking it prior to 90 days, or were prescribed NRT within 90 days of surgery.	Low Outcomes reported as ICD codes; unlikely to have been influenced by knowledge of outcomes.	Moderate Exact matching likely reduces bias due to confounding but could be residual confounding for socioeconomic factors.	No Information Unclear level and handling of missing data.	Low No pre-study protocol but appears to report outcomes and analyses mentioned in methods section.	Moderate

Author Year	Selection Bias	Bias in Classification of Interventions	Bias Due to Departures from Intended Intervention	Bias Due to Measurement of Outcomes	Bias Due to Confounding	Bias Due to Missing Data	Bias in the Selection of Reported Results	Overall Bias (Low, Moderate, Serious, Critical, No Information)
Khalid 2022 ²⁰	Low Appear to include all patients meeting criteria identified from health records during a specified period in their initial cohort. Unclear if some characteristics used for matching/final selection were observed before or after start of NRT, but no indication that bias would be differential between groups.	Low Information on NRT use and smoking cessation counseling provided in medical record and documented at start of intervention.	No information No information provided on adherence to NRT or smoking cessation counseling.	Low Outcomes reported as ICD codes; unlikely to have been influenced by knowledge of outcomes.	Moderate Exact matching likely reduces bias due to confounding but could be residual confounding for socioeconomic factors.	No Information Unclear level and handling of missing data.	Low No pre-study protocol but appears to report outcomes and analyses mentioned in methods section.	Moderate

Abbreviations. ICD=International Classification of Diseases; NRT=nicotine replacement therapy; SCP=smoking cessation program.

APPENDIX D: SYSTEMATIC REVIEWS ON ORTHOPEDIC SURGERY COMPLICATION RISKS ASSOCIATED WITH TOBACCO USE

Author, Year Search Timeframe	Surgery Type	Exposure(s) and Comparator(s)	Included Studies and Total Number of Participants (N)	Relevant Outcomes	Main Findings
Hip					
Emara 2021 ³⁵ Jan 1985-Jan 2020	Hip arthroscopy	Smoking vs not smoking	4 prospective and 3 retrospective cohorts; N = 18,585	Complication rates and rates of revision arthroscopy and/or conversion to THA	Increased risk of VTE and infection with smoking. No difference in need for revision arthroscopy and/or conversion to THA.
Teng 2015 ²⁴ Inception through Aug 2014	THA	Smoking vs not smoking	4 retrospective and 2 prospective cohorts; N = 8,181	LOS after THA and prothesis-related complications	Increased risk of aseptic loosening, deep infection, and all-cause revisions with smoking. No difference LOS after THA or the risk of implant dislocation.
Knee					
He 2022 ²⁵ Inception through Feb 2021	ТКА	Smoking (current or former) vs never smoked	9 cohorts; N = 2,109,482	Surgical complications, pneumonia, need for blood transfusion, VTE, UTI, need for revision, total complications, and mortality	Increased risk of surgical complications (superficial and deep wound infections), pneumonia, need for revision, and total complications with smoking. No difference in VTE, UTI, or mortality.
Kong 2016 ²⁶ Inception through Feb 2016	ТКА	Risk factors for infection including smoking vs absence of risk factors	4 observational studies (type NR); NR	SSI	Increased risk of infection with smoking.
Novikov 2016 ³⁶ Jan 1975-Feb 2016	ACL reconstruction	Smoking vs not smoking	5 observational studies (type NR); NR	Complications (infection, VTE), ipsilateral graft tear and contralateral ACL tear, need for early revision	Increased risk for infection, VTE, and ipsilateral graft tear with smoking.
Shoulder					
Fan 2022 ³⁷ Inception through Jan 2021	Patients who underwent arthroscopic or open rotator cuff repair	Smoking vs not smoking	6 cohort and 4 case series; N = 505 (retear risk) and N = 72,284 reoperation risk)	Retears and reoperations	Increased risk for retear and reoperation with smoking.

Author, Year Search Timeframe	Surgery Type	Exposure(s) and Comparator(s)	Included Studies and Total Number of Participants (N)	Relevant Outcomes	Main Findings
Spine					
Kong 2017 ²⁷ Inception through June 2016	Spinal surgery including of the cervical, thoracic, lumbar, and sacral vertebrae	Smoking (current and former) vs never smoking	15 cohort and 11 case-control studies; N = 67,405	SSI	Increased risk of infection with smoking.
Li 2021 ³⁰ Inception through Mar 2021	Spinal fusion surgery including of the cervical, thoracic, lumbar, and sacral vertebrae	Smoking (variable definitions) vs not smoking	22 cohort and 4 case- control studies; N = 4,409	Fusion rate	Lower rates of spinal fusion with smoking.
Xing 2013 ²⁸ Inception through Jun 2012	Spinal surgery	Smoking vs not smoking	2 cohorts, 2 case- control studies; N = 858	SSI	Increased risk of infection with smoking.
Zheng 2022 ²⁹ Inception through Jul 2021	Cervical spine surgery	Smoking (variable definitions) vs not smoking	16 cohorts and 27 case-control studies; N = 10,020	Operating time, LOS, estimated blood loss, complications, fusion, and reoperation	Increased risk of respiratory complications, dysphagia, wound infection, and reoperation with smoking. No difference in operating time, LOS, estimated blood loss, or fusion.
Multiple					
Bedard 2019 ²¹ Inception through Jan 2018	THA or TKA	Current tobacco use (within 30 days before surgery), non-current tobacco use (past use but not in the 30 days before surgery), any history of tobacco use, and no history of tobacco use	14 observational studies (type NR); N = 227,289	Any wound complication and PJI	Increased risk of wound complications and PJI with any tobacco use compared to no use. Increased risk of PJI among current tobacco users compared to former tobacco users.
Kunutsor 2016 ²² Inception through Sep 2015	Primary or revision total joint arthroplasty (THA, TKA, total shoulder arthroplasty, total elbow arthroplasty, total ankle arthroplasty)	Smoking vs not smoking	8 observational studies (type NR); N = 20,689	PJI	Increased risk of PJI with smoking.

Author, Year Search Timeframe	Surgery Type	Exposure(s) and Comparator(s)	Included Studies and Total Number of Participants (N)	Relevant Outcomes	Main Findings
Singh 2011 ²³ Inception through Mar 2010	THA or TKA	Smoking (current or former) vs not smoking	15 cohort, 6 case- control studies; NR	Any postoperative complication, reoperation or revision, implant loosening, deep infections, mortality	Increased risk of complications and death among current and former smokers compared to non-smokers.

Abbreviations. ACL=anterior cruciate ligament; BMI=body mass index; NR=not reported; LOS=length of stay; OA=osteoarthritis; PJI=periprosthetic joint infection; RA=rheumatoid arthritis; SSI=surgical site infection; THA=total hip arthroplasty; TJA=total joint arthroplasty; TKA=total knee arthroplasty; UTI=urinary tract infection; VTE=venous thromboembolism.

APPENDIX E: PEER REVIEW DISPOSITION

Comment #	Reviewer #	Comment	Author Response
Are the object	ives, scope, an	d methods for this review clearly described?	
1	3	Yes	None
2	4	Yes	None
3	5	Yes	None
4	7	Yes	None
5	8	Yes	None
Is there any in	dication of bias	in our synthesis of the evidence?	
6	3	No	None
7	4	Yes - see comments	
8	5	Yes - See comment below. There was one sentence that sounded a little biased.	
9	7	No	None
10	8	No	None
Are there any	published or un	published studies that we may have overlooked?	
11	3	No	None
12	4	No	None
13	5	No	None
14	7	No	None
15	8	Yes - It could be worth searching clinicaltrials.gov to comment on the presence or absence of ongoing clinical trials that could contribute to this literature in the next few years. If no relevant trials are currently being conducted, the absence of current trials would be worth including the report. This protocol paper describing a current trial might be worth briefly describing in the report: Kainiemi K, Malmivaara A, Sillman-Tetri S, Lasander M, Heinonen M, Korhonen T, Sand J, Laatikainen T, Kyrö A. Smoking cessation intervention prior to orthopedic surgery: A study protocol to determine patient outcomes and	Thank you for this suggestion. In response, we conducted a search of clinicaltrials.gov for ongoing trials and identified no ongoing trials that meet our criteria. We identified the Kainiemi 2022 protocol during our initial literature search. This study planned to include adults undergoing orthopedic surgery, regardless of smoking status, and all the tobacco users received the smoking cessation intervention. We excluded this study because the comparison group was not tobacco users who did not receive a smoking cessation intervention or alternate type of smoking cessation intervention.

Comment #	Reviewer #	Comment	Author Response
		feasibility. Tob Prev Cessat. 2022 Aug 29;8:33. doi: 10.18332/tpc/152608. PMID: 36118614; PMCID: PMC9422999.	
Additional sug	gestions or con	nments can be provided below. If applicable, please indicate th	e page and line numbers from the draft report.
16	3	This systematic review involves three important research questions: 1) What is the effect of smoking on orthopedic surgical complications? 2) What is the effect of smoking cessation on orthopedic surgical complications? and 3) What is the effect of FDA-approved NRT on orthopedic surgical complications? These questions are highly relevant for the field. The main focus of this ESP review is on the 2nd question and is the focus of Key Question 1. The first question is addressed but not until the end of the review; waiting until the end of the review is puzzling. The third question is implied or embedded in Key Question 1. I think it would be helpful if in the introduction there was an overview these three questions are being addressed in the review and how.	Thank you for this comment. The Key Question of this systematic review was the effect of smoking cessation interventions on orthopedic surgical complications, including NRT. We did not synthesize evidence on the effect of smoking on orthopedic surgical complications in general, but thought that this background provided important context, which is why we included this information in the discussion. We did not discuss this background in depth in the introduction because we did not want to create confusion about the Key Question of the systematic review, which was specifically on the effect of smoking cessation interventions.
17	3	The justification for how and why the vast majority of articles were excluded needs more explanation. When looking at the titles of the articles of those excluded, it seems like some would be appropriate for inclusion in this review. More detailed explanation is needed.	Thank you for this comment regarding the exclusion of studies. Reasons for exclusion are provided in the literature flow diagram of the report (Figure 1) and in the list of excluded studies in the Supplementary Materials (Appendix B). Some studies were excluded because they looked at the effect of smoking on orthopedic surgery complications rather than the effect of smoking cessation interventions. Other studies did examine smoking cessation interventions but did not include a comparison group of tobacco users who received no intervention or an alternate intervention, or only examined outcomes related to smoking behaviors and not complications. We added this detail regarding excluded studies to the Methods section.
18	3	With respect to smoking cessation and orthopedic surgical complications, the bottom line of this review concludes that the evidence is quite limited and more research is needed. On page 14, this ESP review seems to question the benefit of smoking cessation and whether or not smoking cessation	Thank you for this suggestion. We agree that the overall benefits of tobacco cessation should be highlighted and not confused with the questions raised by this report. We have revised the

Comment #	Reviewer #	Comment	Author Response
		should be recommended prior to orthopedic surgery. However, this message is inconsistent with US national clinical practice guidelines and current evidence and I suspect not what the authors intend. A suggestion to address this is to consider this review in the context of the extensive existing literature and evidence (four clinical practice guidelines, multiple Surgeon General Reports) regarding the harmful effects of smoking on health and the beneficial effects of smoking cessation on health. Orthopedic surgical complications are only one set of outcomes and are short-term outcomes. Yes, the evidence regarding orthopedic surgical complications is limited. However, evidence of the beneficial effects of smoking cessation on health outcomes such as cardiovascular outcomes (e.g., reduced heart attack risk at one year), cancer risk, COPD, etc are well-established. A surgical procedure represents an opportunity to advise and encourage a patient to quit smoking. US Public Health Service Tobacco Use guidelines recommend, at every clinical encounter, patients who smoke should be advised to quit and be offered smoking cessation assistance.	Discussion text accordingly and cite the 2020 US Surgeon General report.
19	3	For future research: consider recommending a comparative effectiveness trial of NRT versus non-nicotine treatment (e.g., varenicline) in order to assess whether nicotine in NRT is an independent risk factor for worse orthopedic surgical outcomes.	Thank you for this suggestion. We have added this suggestion to the future research section of the Discussion.
20	4	This is a well done and written manuscript. I agree with the concepts and conclusions.	Thank you for this comment.
21	4	The author's methodology is somewhat limited. Systematic reviews over broad policy tend to be more adaptable with numerous stakeholders input. For example, patients, endocrinology, pharmacists.	Thank you for this comment. We aimed to synthesize the available evidence related to this review's key questions. The process of creating clinical guidelines (which is not this review's purpose) often incorporates input from a variety of stakeholders. While we make a suggestion regarding how to apply the report's findings to clinical practice, we defer to VHA leadership to use this report to formally inform clinical guidance as appropriate.

Comment #	Reviewer #	Comment	Author Response
22	4	There is orthopedic surgeon in the author listing -this presents inherent bias in that the review of articles with limited by no clinical expertise on the subject matter Limiting to English language should be considered a limitation in the study design Why not contrast the positive effects of smoking cessation compared to complications for cessation?	Thank you for your comments on limitations of the review. While none of the report authors have a background in orthopedic surgery, the scope of this report was developed in consultation with representatives of the VHA National Surgery Office and peer reviewers include those practicing orthopedic surgery. We do not view the authors' backgrounds as a source of bias.
			Regarding limiting the search to English-language studies, while we agree that this limitation can be a concern, we think it is unlikely that we missed a study with high relevance to the review question and VHA population. Of note, we searched multiple databases and did not exclude any studies for non- English language.
			Regarding your last comment, we focused on surgical complications because whether tobacco cessation mitigates surgical complication risks was the key question of this review. We agree that tobacco cessation has many positive effects and have added text highlighting that point in the Discussion section.
23	4	I believe the terms used to define search are restricting the number of papers. Most orthopedic surgery is elective and often in the research papers/abstracts this word is not used (elective). In addition spine surgery is also done through neurologic surgery. Hand surgery with plastic surgeons SO restricting to word orthopedics limits the articles for orthopedic procedures	Thank you for these comments on the search strategy. We did not require the term 'elective' to be present in our literature searches. In addition to search terms for 'orthopedic surgery,' we included search terms for specific procedures (see Appendix A of the Supplemental Materials).
24	4	In methodology why did you combine ortho procedures/smoking cessation with 4(systematic reviews) does this not limit your set to 1 and 2 that are included in a review? (I maybe misunderstanding this)	Thank you for this question on the search strategy. Our search procedure involves first searching for systematic reviews ('part 1' of the search strategy in Appendix A) and then conducting a search for all study types. This search is described in 'part 4' of the search strategy.

Comment #	Reviewer #	Comment	Author Response
25	5	This is a very well-written and conducted review. Thank you for inviting me to comment.	Thank you for this comment.
26	5	The background section is very well-written and provides excellent information on current recommendations and practices and their rationale. These sentences, in particular, are very helpful: "The evidence most often cited to recommend tobacco cessation before surgery compares outcomes between those who never smoked and current or former tobacco users. Less is known about the difference in risk among those who smoked but quit or reduced their use before surgery and those who continued to smoke their usual amount before surgery."	Thank you for this comment.
27	5	The additional context on the association between tobacco use and surgical complications is helpful as are the findings from the VA's EXTEND QUERI project. On page 15 line 30, should that be "compared with patients not currently smoking or having an unknown smoking status?"	Thank you for this comment. We have corrected the text to indicate that the VA EXTEND QUERI comparison was between current tobacco users and those not currently using tobacco or with an unknown smoking status.
28	5	Page 15, line 37. I'm not following how the first 2 sentences lead to the conclusion "These observations suggest that restrictive policies". I can see how the data in the review "suggest" this, but not necessarily the observations made between tobacco use (former and current) and complications.	Thank you for this comment. We agree that the text starting "These observations suggest" was not adequately supported by our summary of the evidence. We have revised the text in this section to be more explicit about what can and cannot be concluded from the available evidence and what questions remain.
29	3	Page 15, lines 40-41: "Instead," Do we know this? Maybe tone down a bit to say: "It is unclear to what extent cessation requests or requirements may unintentionally" This paragraph sounds a bit biased. The following sentence is great.	Thank you for this comment. We agree that the text you refer to was not adequately supported by our summary of the evidence. We have revised the text in this section to be more explicit about what can and cannot be concluded from the available evidence and what questions remain.
30	7	It appears that the literature search was limited to terms including orthopedic. While it is not known if a search for "orthopaedic" vs. "orthopedic" would result in additional articles for inclusion, this should be assessed. These are often used interchangeably and many societies/academies/associations use orthopaedic (e.g., The American Academy of Orthopaedic Surgeons (AAOS),	Thank you for this comment regarding the search strategy. We agree that the alternate spelling should be included and updated our searches with inclusion of this term. We did not identify additional relevant studies.

Comment #	Reviewer #	Comment	Author Response
		American Orthopaedic Association. If there is a reason it was limited to 'orthopedic', this should be mentioned.	
31	7	This study "Effects of a perioperative smoking cessation intervention on postoperative complications: a randomized trial" (David Lindström, Omid Sadr Azodi, Andreas Wladis, Hanne Tønnesen, Stefan Linder, Hans Nåsell, Sari Ponzer, Johanna Adami) was excluded due to Ineligible Population (p 25); however, it does examine whether an intervention with smoking cessation starting 4 weeks before general and orthopedic surgery (hip or knee prosthesis) would reduce the frequency of postoperative complications. Was it not included because it was not solely focused on orthopedic surgery? If so, this should be noted.	Thank you for this comment. You are correct that this study was excluded because it included patients undergoing surgery types other than orthopedic surgery and outcomes were not reported separately for patients who had orthopedic surgery who did and did not receive the smoking cessation intervention. We added further explanation of why we excluded studies to the Methods section (please also see the response to comment #17).
32	7	There are times in the review that statements seem to be generalizing to all surgeries (for example, p 13, lines 46-54 when discussing NRT and surgical complications), not just elective orthopedic surgeries.	Thank you for this comment. We edited text throughout the report to clarify that we are referring to elective orthopedic surgery rather than all surgeries.
33	8	The eligibility table is generally clear, though the inclusion criteria could benefit from some additional context. The table lists a comparator as an eligibility criteria, but that could mean either 1) the study must have a comparator to be included in the review, or 2) the comparator must meet the eligibility criteria, but only if there is a comparator.	Thank you for this comment regarding eligible comparators. We required that studies have a comparison condition, which could have been either a no-treatment comparison or an alternate intervention. We have added text to the criteria specifying that studies without a comparison condition were excluded.
34	8	On page 15, lines 35-37, the statement about people who stopped smoking shortly before surgery could share a similar level of complication risk as people who continue smoking could be balanced by a statement that the lone trial testing this question found that those who stopped smoking had	Although the text of this comment is incomplete, we assume that the comment suggested referring to the trial in Denmark finding a lower risk of wound- related complications among patients who quit smoking 6-8 weeks before elective knee and hip replacement. We agree with this point and have revised this section of the Discussion to highlight the findings of this trial.
35	8	Though the report appropriately has a narrow focus on post-surgical complications, I suggest include some comment on the overall impact of smoking cessation on a Veteran's health. Since tobacco use is the leading preventable cause of mortality, there is substantial value to the Veterans and to VHA of providing and encouraging	Thank you for this comment. We agree and revised the Discussion text accordingly (please also see the response to comment #18).

Comment #	Reviewer #	Comment	Author Response
		tobacco cessation treatment. In addition, it is noteworthy that major medical events like surgery have the potential to motivate patients to make a smoking cessation attempt that they had been considering.	
36	8	This report does a good job of noting the limitations of the small literature on pre-surgical smoking cessation interventions specifically. It also makes good recommendations about the type of trials that could most effectively inform the types of interventions that work best, and the duration of smoking abstinence typically required to confer benefit for reduced post-surgical complications.	Thank you for this comment.
37	8	Considering the evidence presented, the skepticism about the benefits vs. risks of requiring smoking cessation before surgery seems warranted. The risks of restricting access to surgery have to be weighed against the risks of smoking, especially when people often take many attempts over a long period of time before they are ultimately able to stop smoking. Alternatively, recommending smoking cessation and providing access to evidence-based smoking cessation aids still seems worthwhile from an overall risk vs. benefit perspective. I suggest adding a brief comment to the conclusions stating something about how smoking cessation provides substantial health benefits generally, though the current evidence is insufficient to conclude that smoking cessation shortly before orthopedic surgery results in reduced post-surgical complications. t's possible that clinical decision-makers could read the current conclusion as a recommendation to discou	Thank you for this comment. We agree and have revised the text in this section to be more explicit about what can and cannot be concluded from the available evidence and what questions remain. We also highlight the benefits of tobacco cessation in general and importance of promoting tobacco cessation.