			Certainty a	ssessment			№ of patients		Effect			
№ of studies	Study design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Risk factors		Relative (95% CI)	Absolute (95% CI)	Certainty	Importance
ender - co	hort study											
9	observational studies	serious ^a	not serious	not serious	not serious	publication bias strongly suspected ^b	6654/30087 (22.1%)	0/0	OR 1.10 (0.98 to 1.24)	1 fewer per 1,000 (from 1 fewer to 1 fewer)	OVERY IOW	CRITICAL
gender - ca	se-control study											
8	observational	serious ^a	serious ^c	not serious	not serious	publication bias strongly	2674 cases 2829 controls		OR 1.05	-	⊕OOO	CRITICAL
	studies					suspected ^b	-	0.0%	(0.99 to 1.12)	0 fewer per 1,000 (from 0 fewer to 0 fewer)	Very low	
ge - cohor	t study											
4	observational studies	serious ^a	not serious	not serious	not serious	none	4014/6413 (62.6%)	-/0	OR 1.04 (1.02 to 1.06)	1 fewer per (from 1 fewer to 1 fewer)	OCO Very low	CRITICAL
age - case-	control study								•			
7	observational	serious ^a	not serious	not serious	not serious	none	3098 cases	3243 controls	OR 1.01	-	⊕000	CRITICAL
	studies						-	0.0%	(1.00 to 1.01)	O fewer per (from 0 fewer to 0 fewer)	Very low	
BMI - cohori	study		<u> </u>			l l		I.				<u> </u>
5	observational studies	serious ^a	not serious	not serious	not serious	publication bias strongly suspected ^b	4085/7797 (52.4%)	-/0	OR 1.06 (1.04 to 1.09)	1 fewer per 1,000 (from 1 fewer to 1 fewer)	⊕OOO Very low	CRITICAL
BMI - case-o	ontrol study								I	I I		1
6	observational	serious ^a	serious ^c	not serious	not serious	publication bias strongly	2765 cases 3183 controls		OR 1.04	_	⊕OOO	CRITICAL
	studies	serious	Senous			suspected ^b	-	0.0%	(1.00 to 1.08)	O fewer per (from 0 fewer to 0 fewer)	Very low	
RMI>24ka/r	n2 - cohort study									1 22 2 22 2		
6	observational studies	serious ^a	not serious	not serious	not serious	publication bias strongly suspected ^b	3106/22915 (13.6%)	-/0	OR 1.29 (1.02 to 1.63)	1 fewer per (from 2 fewer to 1 fewer)	OCO Very low	CRITICAL
BMI≥24kg/n	n2 - case-control	study	<u>l</u>			!		<u>I</u>	<u> </u>	-		!
5	observational	serious ^a	not serious	not serious	not serious	publication bias strongly	1974 cases	1734 controls	OR 1.32	_	⊕OOO	CRITICAL
	studies	serious				suspected ^b	-	0.0%	(1.00 to 1.74)	0 fewer per	Very low	
								0.070		(from 0 fewer to 0 fewer)		
Trauma hist	tory - case-contro	ol study										
2	observational	serious ^a r	not serious not serious	not serious	not serious	publication bias strongly	1517 cases 1648 controls		OR 1.53	-	⊕OOO	IMPORTAN
	studies					suspected ^b	-	0.0%	(0.71 to 3.33)	O fewer per (from 0 fewer to 0 fewer)	Very low	
Trauma hist	tory - cohort stud	ly	•					•	-	-		•
4	observational studies	serious ^a	not serious	not serious	not serious	publication bias strongly suspected ^b	1825/15481 (11.8%)	-/0	OR 1.34 (0.99 to 1.82)	1 fewer per (from 2 fewer to 1 fewer)	⊕OOO Very low	IMPORTAN

3	observational studies	serious ^a	not serious	not serious	not serious	publication bias strongly suspected ^b	1622 cases :	1768 controls	OR 1.94 (0.38 to 9.96)	-	Ocery low	NOT IMPORTANT
							-	0.0%	(0.38 to 9.96)	O fewer per (from 0 fewer to 0 fewer)		
ase-contr	ol study - sunless					•						
2	observational	serious ^a	not serious	not serious	not serious	publication bias strongly	1506 cases :	1621 controls	OR 2.03	-	⊕OOO	NOT IMPORTANT
	studies					suspected ^b	-	0.0%	(0.33 to 12.47)	O fewer per (from 0 fewer to 0 fewer)	Very low	
ase-contr	ol study - moist+s	unless										
2	observational studies	serious ^a	not serious	not serious	not serious	publication bias strongly	1506 cases 1621 controls		OR 2.60 (0.12 to 56.64)	-	\oplus OOO	NOT IMPORTANT
	studies					suspected ^b	-	0.0%	(6.12 to 36.6 t)	O fewer per (from 0 fewer to 0 fewer)	Very low	
ase-contr	ol study - cold											
1	observational studies	serious ^a	not serious	not serious	not serious	publication bias strongly	116 cases 147 controls		OR 1.95	-	⊕OOO	NOT IMPORTANT
	studies					suspected ^b	-	0.0%	(0.00 to 4285259.42)	O fewer per (from 0 fewer to)	Very low	
	ol study - Junior hi	gh school		T		<u></u>				 		
2	observational studies	serious ^a	not serious	not serious	not serious	publication bias strongly suspected ^b	1496 cases :	1647 controls	OR 1.21 (0.62 to 2.36)	-	October 10 November 10 Novembe	NOT IMPORTANT
						Suspected	-	0.0%		O fewer per (from 0 fewer to 0 fewer)		
case-contr	ol study - High sch	nool						•	•	•		
2	observational studies	serious ^a	not serious	serious not serious	not serious	publication bias strongly suspected ^b	1496 cases :	1647 controls	OR 0.49 (0.30 to 0.79)	-	⊕OOO Very low	NOT IMPORTANT
							-	0.0%		O fewer per (from 0 fewer to 0 fewer)		
case-contr	ol study - Universi	ty	1	1	1	I L		1		1		_1
2	observational	serious ^a	serious ^d	not serious	not serious	publication bias strongly	1496 cases 1647 controls		OR 0.22	-	⊕OOO	NOT IMPORTANT
	studies					suspected ^b	-	0.0%	(0.06 to 0.86)	O fewer per (from 0 fewer to 0 fewer)	Very low	
case-contr	ol study - OA fami	ly history		•				•				•
4	observational		not serious	not serious	not serious	publication bias strongly	1800 cases 2223 controls		OR 1.04	-	⊕OOO	IMPORTANT
	studies					suspected ^b	-	0.0%	(0.98 to 1.10)	0 fewer per 1,000 (from 0 fewer to 0 fewer)	Very low	
case-contr	ol study - manual	labour	<u>l</u>	1	1			1		ı l		1
	observational studies	nal _{serious} a not se	not serious	ot serious not serious	not serious	publication bias strongly suspected ^b	720 cases 783 controls		OR 1.11	-	⊕OOO	IMPORTANT
4							-	0.0%	(0.93 to 1.31)	0 fewer per	Very low	
4	1									1,000 (from 0 fewer to 0 fewer)		
4												
	ol study - agriculto	ural labor										
	ol study - agriculti observational studies	u ral labor serious ^a	not serious	not serious	not serious	publication bias strongly suspected ^b	269 cases :	323 controls	OR 1.17 (0.54 to 2.56)	-	⊕OOO Very low	IMPORTANT

cohort stud	y - manual labour	r										
3	observational studies	serious ^a	not serious	not serious	not serious	none	1272/18346 (6.9%)	-/0	OR 1.09 (0.88 to 1.35)	1 fewer per 1,000 (from 1 fewer to 1 fewer)	⊕OO Very low	IMPORTANT
ohort stud	y - drinking											
3	observational studies	serious ^a	not serious	not serious	not serious	publication bias strongly suspected ^b	3563/10710 (33.3%)	-/0	OR 0.98 (0.79 to 1.21)	1 fewer per 1,000 (from 1 fewer to 1 fewer)	⊕OOO Very low	NOT IMPORTANT
case-contro	l study - drinking	l							•			•
2	observational studies	serious ^a	not serious	not serious	not serious	none	287 cases 313 controls		OR 1.60 (0.40 to 6.45)	-	\oplus OOO	NOT IMPORTANT
							-	0.0%	(0.40 to 6.45)	0 fewer per 1,000 (from 0 fewer to 0 fewer)	Very low	
ohort stud	y - smoking								-			
4	observational studies	serious ^a	not serious	not serious	not serious	none	3170/20608 (15.4%)	-/0	OR 0.88 (0.74 to 1.04)	1 fewer per 1,000 (from 1 fewer to 1 fewer)	⊕OOO Very low	NOT IMPORTANT
case-contro	l study - smoking	j				•						
3	observational studies	l serious ^a n	serious ^a not serious not serious	us not serious	not serious	publication bias strongly	987 cases 1013 controls		OR 1.09 (0.84 to 1.41)	-	⊕OOO Very low	NOT IMPORTAN
					suspected ^b	-	0.0%	0 fewer per 1,000 (from 0 fewer to 0 fewer)				
case-contro	l study - exercise)	•			'			*	-1		1
4	observational	serious ^a	not serious	s not serious	not serious	publication bias strongly	3713 cases 3995 controls		OR 0.75	-	⊕OOO	IMPORTANT
	studies					suspected ^b	-	0.0%	(0.62 to 0.91)	0.91) 0 fewer per 1,000 (from 0 fewer to 0 fewer)	Very low	

CI: confidence interval; OR: odds ratio

Explanations

a. Most information is from studies at low or unclear risk of bias. Potential limitations are likely to lower confidence in the estimate of effect. b. Funnel plot asymmetry c. 1²=55% d. 1²=65%