Supplemental Online Content

Hussain SM, Ebeling PR, Barker AL, Beilin LJ, Tonkin AM, McNeil JJ. Association of plasma high-density lipoprotein cholesterol level with risk of fractures in healthy older adults. *JAMA Cardiol*. Published online January 18, 2023. doi:10.1001/jamacardio.2022.5124

- **eTable 1.** General characteristics of participants stratified by sex: overall, and by quintiles of HDL-C
- **eTable 2.** Association between baseline non-HDL-C (mg/dL) and at least one fracture occurring after recruitment (HR, 95% CI)
- eTable 3. Association between baseline HDL-C (mg/dL) and at least one fracture occurring after recruitment (HR, 95% CI)
- **eTable 4**. Association between baseline HDL-C (mg/dL) and at least one fracture occurring after recruitment based on statin use status (HR, 95% CI)

This supplemental material has been provided by the authors to give readers additional information about their work.

eTable 1: General characteristics of participants stratified by sex: Overall, and by quintiles of HDL-C

	Overall	Q1	Q2-4	Q5	P Value
Males					
N	7319	1976	4188	1155	
Participants with at least 1	450 (6)	107 (5)	247 (6)	96 (8)	<.001
fracture, n (%)	. ,				
Types of fracture, n (%)					.012
Minimal trauma fracture, n	192 (3)	43 (2)	111 (3)	38 (3)	
(%)	. ,				
Other trauma fracture, n	258 (4)	64 (3)	136 (3)	58 (5)	
(%)	` ´		. ,		
Age in years	75 (4)	75 (4)	75 (4)	76 (5)	<.001
Low activity, n (%)	389 (5)	141 (7)	207 (5)	41 (3)	<.001
BMI (kg/m2), mean (SD)	28 (4)	29 (4)	28 (4)	26 (34)	<.001
Waist Circumference	102 (11)	105 (10)	102 (10)	98 (10)	<.001
(cm), mean (SD)					
Current/former smoking, n	4164 (57)	1081 (55)	2371 (57)	712 (62)	<.001
(%)	,				
Current alcohol use, n (%)	6254 (85)	1542 (78)	3643 (87)	1060 (92)	<.001
Education, n (%)	` /				<.001
<12 years of schooling	4236 (58)	1239 (63)	2364 (57)	633 (55)	1
≥12 years of schooling	3083 (42)	737 (37)	1824 (44)	522 (45)	1
Hypertension, n (%)	5557 (76)	1560 (79)	3118 (75)	879 (76)	.001
Chronic kidney disease	1196 (17)	452 (24)	614 (15)	130 (12)	<.001
(eGFR <60ml/min), n (%)					
Diabetes, n (%)	869 (12)	389 (20)	402 (10)	78 (7)	<.001
Prefrail/Frail, n (%)	2708 (37)	783 (40)	1526 (36)	399 (35)	.009
On trial medication, n (%)	3656 (50)	982 (50)	2066 (49)	608 (53)	.13
Osteoporotic medication,	143 (2)	35 (2)	72 (2)	36 (3)	.008
n (%)					
Total Cholesterol (mg/dL),	195 (36)	182 (36)	197 (34)	208 (34)	<.001
mean (SD)					
HDL-C (mg/dL), mean	54 (15)	38 (5)	55 (6)	80 (10)	
(SD)	, ,				
Non-HDL-C (mg/dL),	141 (35)	144 (36)	142 (34)	128 (34)	<.001
mean (SD)	, ,	, ,	, ,	l ` ´	
Lipid lowering	2124 (31)	642 (34)	1147 (29)	335 (31)	.002
medications, n (%)					
Females					
N	8945	2548	4892	1505	
Participants with at least 1	1209 (14)	305 (12)	660 (14)	244 (16)	.001
fracture, n (%)	<u> </u>				
Types of fracture, n (%)					.004
Minimal trauma fracture, n	519 (6)	124 (5)	290 (6)	105 (7)	
(%)					
Other trauma fracture, n	690 (8)	181 (7)	370 (8)	139 (9)	
(%)					

Overall	Q1	Q2-4	Q5	P Value
75 (4)	75 (4)	75 (4)	76 (5)	.10
655 (7)	241 (10)	337 (7)	77 (5)	<.001
28 (5)	30 (5)	28 (5)	26 (5)	<.001
93 (13)	98 (13)	92 (12)	87 (12)	<.001
3044 (34)	843 (33)	1612 (33)	589 (39)	<.001
6590 (74)	1668 (66)	3674 (75)	1248 (83)	<.001
				<.001
5719 (64)	1721 (68)	3115 (64)	883 (59)	
3225 (36)	827 (33)	1777 (36)	621 (41)	
6639 (74)	2042 (80)	3556 (73)	1041 (69)	<.001
1677 (19)	577 (23)	873 (18)	227 (15)	<.001
719 (8)	373 (15)	288 (6)	58 (4)	<.001
3521 (39)	1053 (41)	1873 (38)	595 (40)	.04
4453 (50)	1266 (50)	2428 (50)	759 (50)	0.86
953 (11)	210 (8)	534 (11)	209 (14)	<.001
211 (38)	199 (40)	213 (36)	227 (34)	<.001
67 (18)	48 (6)	68 (7)	96 (13)	
144 (37.5)	151 (40)	145 (37)	131 (34)	<.001
3244 (37)	1087 (44)	1697 (36)	460 (32)	<.001
	75 (4) 655 (7) 28 (5) 93 (13) 3044 (34) 6590 (74) 5719 (64) 3225 (36) 6639 (74) 1677 (19) 719 (8) 3521 (39) 4453 (50) 953 (11) 211 (38) 67 (18)	75 (4) 75 (4) 655 (7) 241 (10) 28 (5) 30 (5) 93 (13) 98 (13) 3044 (34) 843 (33) 6590 (74) 1668 (66) 5719 (64) 1721 (68) 3225 (36) 827 (33) 6639 (74) 2042 (80) 1677 (19) 577 (23) 719 (8) 373 (15) 3521 (39) 1053 (41) 4453 (50) 1266 (50) 953 (11) 210 (8) 211 (38) 199 (40) 67 (18) 48 (6)	75 (4) 75 (4) 75 (4) 655 (7) 241 (10) 337 (7) 28 (5) 30 (5) 28 (5) 93 (13) 98 (13) 92 (12) 3044 (34) 843 (33) 1612 (33) 6590 (74) 1668 (66) 3674 (75) 5719 (64) 1721 (68) 3115 (64) 3225 (36) 827 (33) 1777 (36) 6639 (74) 2042 (80) 3556 (73) 1677 (19) 577 (23) 873 (18) 719 (8) 373 (15) 288 (6) 3521 (39) 1053 (41) 1873 (38) 4453 (50) 1266 (50) 2428 (50) 953 (11) 210 (8) 534 (11) 211 (38) 199 (40) 213 (36) 67 (18) 48 (6) 68 (7) 144 (37.5) 151 (40) 145 (37)	75 (4) 75 (4) 75 (4) 76 (5) 655 (7) 241 (10) 337 (7) 77 (5) 28 (5) 30 (5) 28 (5) 26 (5) 93 (13) 98 (13) 92 (12) 87 (12) 3044 (34) 843 (33) 1612 (33) 589 (39) 6590 (74) 1668 (66) 3674 (75) 1248 (83) 5719 (64) 1721 (68) 3115 (64) 883 (59) 3225 (36) 827 (33) 1777 (36) 621 (41) 6639 (74) 2042 (80) 3556 (73) 1041 (69) 1677 (19) 577 (23) 873 (18) 227 (15) 719 (8) 373 (15) 288 (6) 58 (4) 3521 (39) 1053 (41) 1873 (38) 595 (40) 4453 (50) 1266 (50) 2428 (50) 759 (50) 953 (11) 210 (8) 534 (11) 209 (14) 211 (38) 199 (40) 213 (36) 227 (34) 67 (18) 48 (6) 68 (7) 96 (13) 144 (37.5) 151 (40) 145 (37)

Abbreviation: HDL-C, high density lipoprotein cholesterol; BMI, body mass index; GFR, glomerular filtration rate; SD, standard deviation

To convert cholesterol to mmol/L, multiply values by 0.0259.

eTable 2: Association between baseline non-HDL-C (mg/dL) and at least one fracture occurring after recruitment (HR, 95% CI)

	Non-HDL	Non-HDL-C quantiles			
	(mg/dL)	Q1	Q2-Q4	Q5	
	continuous	(reference)			
All participants					
IR per 1000 PY	26 (24, 27)	28 (25, 31)	26 (24, 28)	23 (20, 26)	
(95% CI)					
Model 1 ^a	0.92 (0.88, 0.97)	1	0.96, 0.85, 1.08)	0.85 (0.73, 0.99)	
Model 2 ^b	0.92 (0.88, 0.97)	1	0.95 (0.85, 1.06)	0.85 (0.72, 0.99)	
Model 3 ^c	0.92 (0.88, 0.97)	1	0.96 (0.85, 1.08)	0.85 (0.73, 0.99)	
Model 4 ^d	0.92 (0.87, 0.97)	1	0.95 (0.83, 1.08)	0.85 (0.71, 1.00)	
Analyses stratifie	ed by sex ^e				
Males					
IR per 1000 PY	15 (14, 17)	17 (14, 21)	16 (14, 18)	13 (10, 16)	
(95% CI)					
Model 1	0.92 (0.83, 1.02)	1	0.89 (0.71, 1.12)	0.72 (0.53, 0.97)	
Model 2	0.92 (0.83, 1.01)	1	0.89 (0.71, 1.12)	0.72 (0.53, 0.97)	
Model 3	0.92 (0.84, 1.02)	1	0.89 (0.71, 1.12)	0.72 (0.53, 0.97)	
Model 4	0.91 (0.82, 1.02)	1	0.87 (0.67, 1.11)	0.71 (0.51, 0.98)	
Females					
IR per 1000 PY	34 (32, 36)	35 (31, 40)	35 (32, 37)	32 (28, 36)	
(95% CI)					
Model 1	0.93 (0.88, 0.98)	1	0.98 (0.86, 1.13)	0.90 (0.75, 1.08)	
Model 2	0.92 (0.87, 0.98)	1	0.98 (0.85, 1.12)	0.90 (0.75, 1.07)	
Model 3	0.93 (0.87, 0.98)	1	0.98 (0.85, 1.13)	0.90 (0.75, 1.08)	
Model 4	0.92 (0.86, 0.98)	1	0.98 (0.84, 1.15)	0.91 (0.74, 1.11)	

Abbreviation: IR, Incidence Rate; PY, Person Year; BMI, body mass index;

^a Model 1: adjusted for age, sex;

^b Model 2: adjusted for age, sex, physical activity, alcohol use;

^c Model 3: adjusted for age, sex, physical activity, alcohol use, prefrailty/frailty status; Model

^d 4: adjusted for age, sex, physical activity, alcohol use, prefrailty, frailty status, education, BMI, smoking status, Aspirin, diabetes, chronic kidney disease, lipid lowering medication, OP medications;

^e not adjusted for sex

eTable 3: Association between baseline HDL-C (mg/dL) and at least one fracture occurring after recruitment (HR, 95% CI)

	HDL (mg/dL)	HDL quantiles			
	continuous	Q1	Q2-Q4	Q5	
		(reference)			
Minimal traumatic	Minimal traumatic fractures (N=14764)				
All	1.10 (1.02, 1.20)	1	1.22 (1.02, 1.48)	1.42 (1.11, 1.81)	
Males ^a	1.11 (0.94, 1.33)	1	1.15 (0.80, 1.66)	1.31 (0.82, 2.09)	
Females ^a	1.10 (1.00, 1.20)	1	1.24 (1.01, 1.55)	1.44 (1.09, 1.91)	
Participants not on	Participants not on osteoporotic medications ^b (N=15168)				
All	1.15 (1.09, 1.22)	1	1.06 (0.93, 1.21)	1.32 (1.12, 1.58)	
Males ^a	1.22 (1.08, 1.38)	1	1.04 (0.82, 1.32)	1.41 (1.04, 1.91)	
Females ^a	1.13 (1.06, 1.21)	1	1.07 (0.92, 1.25)	1.30 (1.07, 1.58)	
Who never-smoked	Who never-smoked and reported that they did not drink alcohol ^c (N=2445)				
All	1.22 (1.07, 1.40)	1	1.10 (0.83, 1.45)	1.59 (1.07, 2.35)	
Males ^a	1.15 (0.75, 1.74)	1	0.60 (0.29, 1.21)	1.76 (0.72, 4.35)	
Females ^a	1.24 (1.08, 1.43)	1	1.26 (0.92, 1.72)	1.60 (1.03, 2.49)	
Who walked outside less than 30 minutes and reported no participation in					
moderate/vigorous physical activity ^d (N=5931)					
All	1.18 (1.09, 1.28)	1	1.09 (0.90, 1.31)	1.42 (1.11, 1.80)	
Males ^a	1.13 (0.93, 1.39)	1	0.96 (0.65, 1.42)	1.31 (0.79, 2.18)	
Females ^a	1.18 (1.08, 1.29)	1	1.12 (0.90, 1.34)	1.45 (1.10, 1.90)	

Adjusted for age, sex, physical activity, alcohol use, prefrailty, frailty status, education, body mass index, smoking status, Aspirin, diabetes, chronic kidney disease, lipid lowering medication, osteoporosis medications;

^a not adjusted for sex;

^b not adjusted for osteoporosis medications;

^c not adjusted for smoking and alcohol use status;

^d not adjusted for physical activity

eTable 4: Association between baseline HDL-C (mg/dL) and at least one fracture occurring after recruitment based on statin use status (HR, 95% CI)

	HDL (mg/dL)	HDL quantiles				
	continuous	Q1	Q2-Q4	Q5		
		(reference)				
Participants not on	Participants not on statins (n=10,170)					
All	1.13 (1.06, 1.21)	1	1.05 (0.90, 1.22)	1.36 (1.11, 1.61)		
Males ^a	1.22 (1.07, 1.39)	1	0.99 (0.74, 1.30)	1.49 (1.05, 2.10)		
Females ^a	0.97 (0.81, 1.61)	1	1.07 (0.99, 1.29)	1.27 (1.02, 1.60)		
Participants on statins (n=5368)						
All	1.16 (1.06, 1.27)	1	1.17 (0.95, 1.43)	1.27 (1.00, 1.67)		
Males ^a	1.19 (0.98, 1.47)	1	1.27 (0.89, 1.95)	1.21 (0.70, 2.23)		
Females ^a	1.15 (1.04, 1.27)	1	1.14 (0.90, 1.43)	1.29 (0.98, 1.77)		

Adjusted for age, sex, physical activity, alcohol use, prefrailty, frailty status, education, body mass index, smoking status, Aspirin, diabetes, chronic kidney disease, osteoporosis medications;

^a not adjusted for sex;