

SUPPLEMENTARY DATA

**Supplementary Table 1.** Baseline characteristics across categories of CRP according to diabetes status

Categories for CRP	Overall				No diabetes				Diabetes			
	<1mg/l	1-3 mg/l	>3mg/l	p-value	<1mg/l	1-3 mg/l	>3mg/l	p-value	<1mg/l	1-3 mg/l	>3mg/l	p-value
N	7890	9421	8668		7708	8975	8013		182	446	655	
Median CRP ( mg/L)	0.5	1.7	5.7	N/A	0.5	1.7	5.7	N/A	0.6	1.8	6.0	N/A
Age (years)	51.4 (12.5)	56.1 (13)	59 (13.1)	<0.0001	51.2 (12.4)	55.7 (12.9)	58.7 (13.1)	<0.0001	62.4 (12.8)	63.5 (11.9)	63.3 (12)	0.54
Women (%)	54.1	49.9	58.6	<0.0001	54.5	50.4	59.0	<0.0001	34.1	40.1	54.0	<0.0001
Current smoking (%)	19.4	24.2	29.2	<0.0001	19.6	24.4	29.7	<0.0001	13.8	19.1	22.7	0.02
Systolic blood pressure (mm Hg)	129 (18)	136 (19)	139 (21)	<0.0001	129 (18)	135 (19)	139 (21)	<0.0001	141 (21)	143 (20)	142 (23)	0.70
Resting heart rate (bpm)	69 (11)	70 (11)	73 (12)	<0.0001	69 (10)	70 (11)	72 (11)	<0.0001	71 (12)	71 (12)	75 (13)	<0.0001
Body mass index (kg/m <sup>2</sup> )	25.2 (3.4)	27.5 (4.2)	29.4 (5.4)	<0.0001	25.2 (3.4)	27.4 (4.2)	29.2 (5.3)	<0.0001	26.7 (3.6)	29.1 (4.4)	32 (6.1)	<0.0001
Waist circumference (cm)	85.5 (11.5)	92.3 (12.5)	96.3 (13.6)	<0.0001	85.3 (11.4)	92 (12.4)	95.5 (13.2)	<0.0001	92.7 (11.9)	99.3 (13)	105.6 (14)	<0.0001
Waist/hip ratio	0.85 (0.08)	0.88 (0.09)	0.89 (0.08)	<0.0001	0.85 (0.08)	0.88 (0.09)	0.89 (0.08)	<0.0001	0.90 (0.08)	0.93 (0.08)	0.94 (0.08)	<0.0001
Total cholesterol (mmol/l)	5.6 (1.0)	5.9 (1.2)	5.9 (1.1)	<0.0001	5.6 (1.0)	5.9 (1.2)	6.0 (1.1)	<0.0001	5.2 (1.0)	5.6 (1.0)	5.7 (1.1)	<0.0001
HbA1c (%)	5.2 (0.5)	5.4 (0.7)	5.6 (0.9)	<0.0001	5.2 (0.3)	5.3 (0.4)	5.4 (0.38)	<0.0001	6.9 (1.1)	7.5 (1.5)	7.7 (1.6)	<0.0001
Ethnicity, Caucasians (%)	97.4	97.8	97.9	0.06	97.5	98.0	98.1	0.04	92.9	93.9	96.5	0.05

p-values are from the analysis of variance (ANOVA)

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**Supplementary Table 2.** Hazard ratio (and 95% confidence interval) for all-cause (left panel) and cardiovascular disease (right panel) mortality across levels of C-reactive protein overall and by gender and diabetes status

Subgroup	CRP levels	All-cause mortality			CVD mortality		
		Overall	No diabetes	Diabetes	Overall	No diabetes	Diabetes
Overall							
	<1 mg/l	1.00 (0.91-1.10)	1.00 (0.91-1.10)	1.00 (0.73-1.38)	1.00 (0.84-1.20)	1.00 (0.83-1.21)	1.00 (0.57-1.76)
	1-3 mg/l	1.05 (0.98-1.12)	1.05 (0.98-1.13)	0.90 (0.73-1.12)	1.31 (1.17-1.47)	1.27 (1.12-1.44)	1.38 (1.01-1.89)
	>3 mg/l	1.86 (1.76-1.96)	1.83 (1.73-1.93)	1.65 (1.42-1.91)	2.49 (2.28-2.70)	2.40 (2.19-2.63)	2.44 (1.96-3.04)
Men							
	<1 mg/l	1.00 (0.88-1.14)	1.00 (0.87-1.15)	1.00 (0.67-1.49)	1.00 (0.79-1.26)	1.00 (0.78-1.28)	1.00 (0.45-2.00)
	1-3 mg/l	1.14 (1.04-1.24)	1.15 (1.04-1.26)	0.91 (0.69-1.21)	1.26 (1.04-1.46)	1.23 (1.04-1.46)	1.20 (0.79-1.83)
	>3 mg/l	2.11 (1.97-2.27)	2.10 (1.95-2.27)	1.75 (1.43-2.13)	2.69 (2.40-3.01)	2.55 (2.25-2.89)	2.82 (2.15-3.69)
Women							
	<1 mg/l	1.00 (0.87-1.14)	1.00 (0.87-1.15)	1.00 (0.59-1.69)	1.00 (0.75-1.32)	1.00 (0.75-1.34)	1.00 (0.37-2.68)
	1-3 mg/l	0.96 (0.86-1.06)	0.95 (0.85-1.06)	0.88 (0.62-1.25)	1.40 (1.17-1.66)	1.34 (1.11-1.61)	1.67 (1.04-2.69)
	>3 mg/l	1.60 (1.49-1.72)	1.56 (1.44-1.69)	1.48 (1.19-1.84)	2.25 (1.98-2.55)	2.21 (1.94-2.53)	1.95 (1.36-2.81)

Cox models are stratified by cohort and adjusted for sex and age. The 95% confidence intervals are from floating absolute risk methods.

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**Supplementary Table 3.** Hazard ratios (and 95% confidence intervals) for 1 Standard deviation log<sub>e</sub> change in CRP levels and all-cause (left panel) and cardiovascular disease (right panel) mortality by diabetes status within subgroups of participants

Variables	Subgroups	All-cause mortality				Cardiovascular disease mortality			
		No diabetes	Diabetes	p-int	p 3-way	No diabetes	Diabetes	p-int	p 3-way
Smoking	Non smoker	1.32 (1.26-1.39)	1.43 (1.25-1.64)	0.23	0.62	1.36 (1.25-1.47)	1.34 (1.10-1.63)	0.90	0.68
	Smokers	1.33 (1.24-1.44)	1.52 (1.18-1.95)	0.18		1.67 (1.45-1.91)	2.08 (1.32-3.30)	0.30	
Age (years)	≤55	1.46 (1.30-1.64)	0.92 (0.58-1.45)	0.07	0.12	1.63 (1.29-2.07)	1.59 (0.77-3.29)	0.92	0.61
	>55	1.36 (1.31-1.42)	1.49 (1.33-1.68)	0.09		1.46 (1.36-1.56)	1.42 (1.18-1.70)	0.84	
BMI (kg/m <sup>2</sup> )	≤28.6	1.44 (1.36-1.52)	1.85 (1.46-2.35)	0.01	0.20	1.51 (1.36-1.67)	1.50 (0.97-2.32)	0.76	0.38
	>28.6	1.38 (1.29-1.48)	1.38 (1.18-1.61)	0.84		1.47 (1.31-1.65)	1.29 (1.03-1.62)	0.37	
WC (cm)	≤91.1	1.36 (1.28-1.45)	1.30 (1.03-1.64)	0.82	0.02	1.44 (1.29-1.61)	1.00 (0.61-1.62)	0.29	0.09
	>91.1	1.37 (1.29-1.45)	1.46 (1.27-1.67)	0.24		1.47 (1.34-1.62)	1.39 (1.15-1.68)	0.65	
TC (mmol/l)	≤5.8	1.39 (1.31-1.47)	1.60 (1.32-1.95)	0.12	0.07	1.46 (1.31-1.63)	1.05 (0.76-1.46)	0.44	0.43
	>5.8	1.34 (1.26-1.42)	1.38 (1.12-1.69)	0.57		1.44 (1.30-1.60)	1.26 (0.93-1.71)	0.53	
SBP (mmHg)	≤132	1.39 (1.29-1.49)	1.58 (1.28-1.95)	0.16	0.29	1.38 (1.22-1.56)	1.60 (1.17-2.19)	0.45	0.69

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	>132	1.37 (1.30-1.44)	1.39 (1.20-1.61)	0.76			1.50 (1.38-1.64)	1.39 (1.10-1.75)	0.44	
Physical activity (sessions)	≤1/week	1.36 (1.29-1.42)	1.58 (1.38-1.81)	0.04	0.31		1.43 (1.32-1.55)	1.36 (1.11-1.68)	0.64	0.12
	>1/week	1.34 (1.23-1.45)	0.93 (0.66-1.30)	0.05			1.54 (1.32-1.80)	2.31 (1.27-4.19)	0.17	
Ethnicity	Caucasians	1.41 (1.35-1.47)	1.55 (1.38-1.75)	0.06	0.06		1.52 (1.42-1.64)	1.55 (1.29-1.86)	0.78	0.75
	Non Caucasians	1.06 (0.60-1.88)	0.30 (0.08-1.15)	0.12			0.36 (0.10-1.34)	0.40 (0.06-2.68)	0.98	
Overall		1.41 (1.35-1.47)	1.53 (1.35-1.72)	0.08			1.51 (1.41-1.63)	1.54 (1.28-1.85)	0.80	

p-int, p-value for interaction for the association by diabetes status within subgroups;p 3-way, 3-ways interaction p-values for the associations across complementary subgroups. Cox models are stratified by cohort and adjusted for sex and age.