

## Supplementary data

### Supplementary table 1

Spearman correlation coefficient of cathepsin D, L or cystatin B to baseline clinical and laboratory measurements in the MDC-CC cohort

	Cathepsin D	Cathepsin L	Cystatin B
	r	r	r
Age	0.10**	0.08*	0.17***
Smoking <sup>†</sup>	0.19**	0.12	0.10
BMI	0.24***	0.12***	0.22***
White blood cells	0.17***	0.08*	0.23***
Cholesterol	0.03	-0.01	0.08*
LDL	0.02	-0.01	0.09*
HDL	-0.26***	-0.12**	-0.16***
Triglycerides	0.30***	0.1**	0.18***
Fasting blood glucose	0.23***	0.09*	0.08*
HbA1c	0.22***	0.08*	0.17***
Plasma insulin	0.31***	0.14***	0.23***
Systolic BP	0.19***	0.08*	0.12**
Diastolic BP	0.20***	0.13***	0.11**

r indicates Spearman correlation coefficient; n.s. indicates not significant; BP, blood pressure;

<sup>†</sup>number of cigarette packages per year

\* $P < 0.05$

\*\*  $P < 0.01$

\*\*\*  $P < 0.001$

Supplementary table 2

Hazard ratios (95% confidence intervals, CI) for incident coronary events by tertiles of cathepsin D in non-diabetics in the MDC-CC cohort

	1 <sup>st</sup> tertile	2 <sup>nd</sup> tertile	3 <sup>rd</sup> tertile	<i>P</i> for trend
Cathepsin D (AU)	<144	144-193	>194	
Coronary events (no)	86	96	123	
Coronary events HR (95% CI) <sup>1</sup>	1	1.12 (0.83-1.50)	1.63 (1.23-2.14)	0.0005
Coronary events HR (95% CI) <sup>2</sup>	1	1.00 (0.74-1.35)	1.47 (1.10-1.97)	0.008

<sup>1</sup> Non-adjusted.

<sup>2</sup> Adjusted for age, sex, LDL/HDL ratio, triglycerides, BMI and hypertension.

Supplementary table 3

Hazard ratios (95% confidence intervals, CI) for incident coronary events by tertiles of cystatin B in non-diabetics in the MDC-CC cohort

	1 <sup>st</sup> and 2 <sup>nd</sup> tertile	3 <sup>rd</sup> tertile	<i>P</i>
Cystatin B (AU)	<49.8	>49.8	
Coronary events (no)	183	122	
Coronary events HR (95% CI) <sup>1</sup>	1	1.46 (1.16-1.84)	0.001
Coronary events HR (95% CI) <sup>2</sup>	1	1.39 (1.09-1.77)	0.008

<sup>1</sup> Non-adjusted.

<sup>2</sup> Adjusted for age, sex, LDL/HDL ratio, triglycerides, BMI and hypertension.

Supplementary table 4

Odds ratios (95% confidence intervals, CI) for incident coronary events per standard deviation of cathepsin D, L and cystatin B in non-smokers in the MDC-CC cohort

	Cathepsin D		Cathepsin L		Cystatin B	
	OR	<i>P</i>	OR	<i>P</i>	OR	<i>P</i>
OR (95% CI) <sup>1</sup>	1.25(1.05-1.49)	0.012	1.12(0.95-1.34)	0.18	1.19(1.00-1.41)	0.053
OR (95% CI) <sup>2</sup>	1.09(0.89-1.33)	0.39	1.06(0.88-1.27)	0.56	1.12(0.93-1.36)	0.24

<sup>1</sup> Non-adjusted.

<sup>2</sup> Adjusted for age, sex, LDL/HDL ratio, triglycerides, BMI, hypertension and glucose.

Supplementary table 5

Odds ratios (95% confidence intervals, CI) for incident coronary events per standard deviation of cystatin B in smokers in the MDC-CC cohort

	Cathepsin D		Cathepsin L		Cystatin B	
	OR	<i>P</i>	OR	<i>P</i>	OR	<i>P</i>
OR (95% CI) <sup>1</sup>	1.54(1.17-2.02)	0.002	1.29(0.99-1.68)	0.063	1.43(1.08-1.88)	0.012
OR (95% CI) <sup>2</sup>	1.42(1.02-1.97)	0.038	1.19(0.89-1.61)	0.24	1.27(0.94-1.73)	0.13

<sup>1</sup> Non-adjusted.

<sup>2</sup> Adjusted for age, sex, LDL/HDL ratio, triglycerides, BMI, hypertension and glucose.