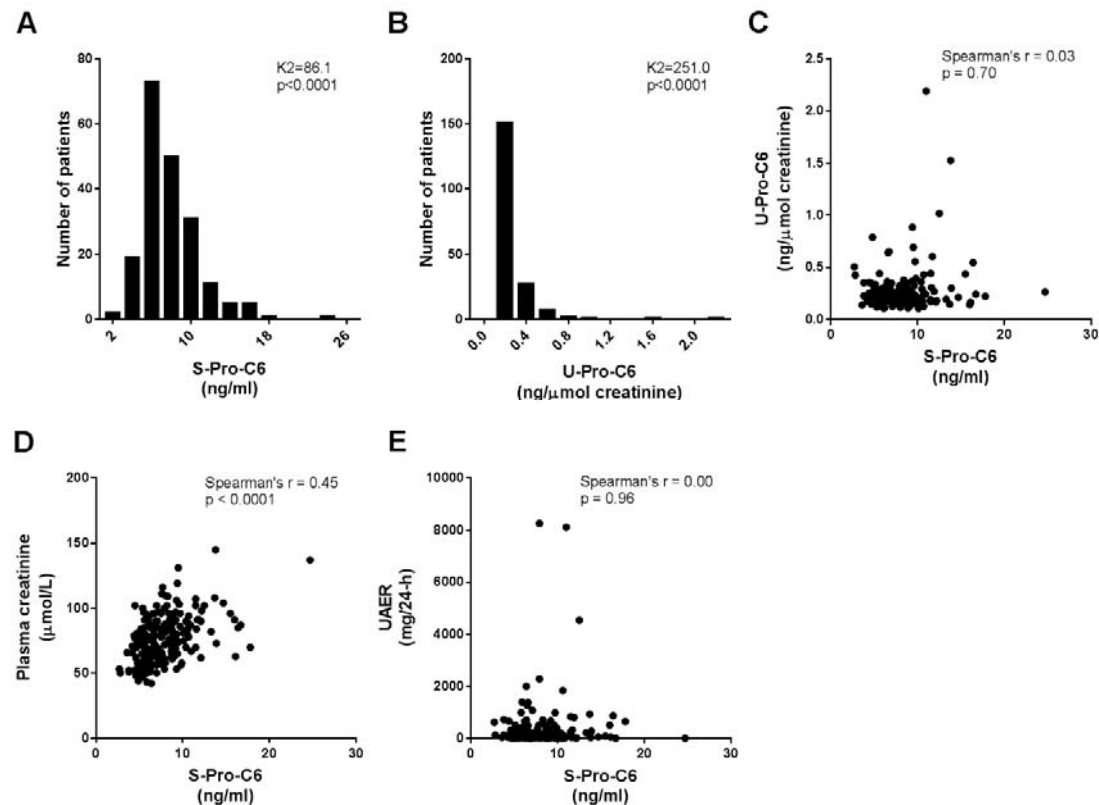


## SUPPLEMENTARY DATA

**Supplementary Figure S1. Distribution and correlations.** The distribution of (A) S-Pro-C6 (n=198) and (B) U-Pro-C6 (n=190) of the patients were plotted. Both distributions were positively skewed and were tested for Gaussian distribution by a D'Agostino & Pearson normality test. (A) S-Pro-C6 had a non-Gaussian distribution ( $K2=86.1$ ,  $p<0.0001$ ), and log-transformed data showed borderline Gaussian distribution ( $K2=6.0$ ,  $p=0.05$ ). (B) U-Pro-C6 had a non-Gaussian distribution for both non-transformed ( $K2=251.0$ ,  $p<0.0001$ ) and log-transformed data ( $K2=70.9$ ,  $p<0.0001$ ). (C) Due to the non-Gaussian distribution, a nonparametric Spearman's rank correlation analysis between S-Pro-C6 and U-Pro-C6 was performed. There was no correlation between S-Pro-C6 and U-Pro-C6 (Spearman's  $r=0.03$ ,  $p=0.70$ ). Nonparametric Spearman's rank correlation analysis between S-Pro-C6 and (D) plasma creatinine (Spearman's  $r=0.45$ ,  $p<0.0001$ ) and (E) UAER (Spearman's  $r=0.00$ ,  $p=0.96$ ) at baseline.



SUPPLEMENTARY DATA

**Supplementary Table S1.** Hazard ratios of S-Pro-C6 and conventional risk factors in the fully adjusted models

Variables	Cardiovascular events (n=38)		Mortality (n=26)		Disease progression <sup>‡</sup> (n=42)	
	HR(95% CI)	p-value	HR(95% CI)	p-value	HR (95% CI)	p-value
S-Pro-C6*	3.06 (1.31-7.14)	<b>0.01</b>	6.91 (2.96-16.11)	<b>&lt;0.0001</b>	4.81 (1.92-12.01)	<b>0.0008</b>
Age (years)	1.08 (1.01-1.15)	<b>0.02</b>	1.04 (0.98-1.11)	0.21	1.01 (0.96-1.05)	0.81
Sex=male	3.09 (0.93-10.26)	0.06	2.90 (0.83-10.17)	0.10	0.87 (0.36-2.12)	0.76
BMI (kg/m <sup>2</sup> )	0.95 (0.88-1.02)	0.13	1.00 (0.93-1.08)	0.92	0.91 (0.85-0.98)	<b>0.01</b>
Systolic BP (mmHg)	1.01 (0.99-1.03)	0.21	0.99 (0.97-1.02)	0.56	1.01 (0.99-1.03)	0.35
Smoking=yes	1.56 (0.77-3.14)	0.22	2.89 (1.25-6.66)	<b>0.01</b>	0.67 (0.30-1.50)	0.33
HbA <sub>1C</sub> (%)	1.15 (0.86-1.56)	0.35	0.82 (0.58-1.16)	0.26	1.12 (0.87-1.44)	0.38
LDL-cholesterol (mmol/L)	1.44 (0.96-2.16)	0.08	1.28 (0.77-2.11)	0.34	0.76 (0.48-1.22)	0.25
Plasma creatinine (μmol/L)	1.0 (0.97-1.02)	0.82	0.97 (0.94-1.00)	<b>0.04</b>	1.00 (0.98-1.03)	0.87
UAER (mg/24-h) <sup>†</sup>	1.30 (0.74-2.28)	0.36	1.17 (0.59-2.33)	0.65	2.11 (1.23-3.62)	<b>0.007</b>

Hazard ratios (HR) with 95% CI are listed. \*S-Pro-C6 hazard ratios are listed as per doubling. <sup>†</sup>UAER was log<sub>10</sub>-transformed prior to analysis. Cardiovascular events was defined as a composite of cardiovascular mortality, stroke, ischaemic cardiovascular disease and heart failure. <sup>‡</sup>Data on disease progression was only available for 175 out of 198 patients (88.4%). Disease progression was defined as a decline of eGFR of more than 30%. BMI, body mass index; UAER, urinary albumin excretion ratio.