## **Supplementary Material**

**Table S1** Associations between change in self-reported protein intake and change in kidney function from baseline to 1 year.

Δ Protein Intake (g/kg/day)						
Variable	n	Unadjusted	<i>p</i> -Value	n	Adjusted	<i>p</i> -Value
		(ß ± SE)			(ß ± SE)	
<b>Δ Creatinine clearance</b> (mL/min)	233	$-6.70 \pm 9.85$	0.497	177	$-1.70 \pm 10.91$	0.876
$\Delta \text{ eGFR} (\text{mL/min}/1.73\text{m}^2)$	242	$0.43 \pm 2.14$	0.842	183	$-0.24 \pm 2.55$	0.925
$\Delta$ Albumin/Creatinine ratio (ACR)	242	$-2.62 \pm 7.13$	0.714	183	$-1.09 \pm 9.52$	0.908
$\Delta$ Urea/Creatinine Ratio (UCR)	242	$7.77 \pm 3.76$	0.04	183	$9.93 \pm 4.45$	0.03
<b>Δ S-Creatinine</b> (μmol/L)	242	$-1.72 \pm 2.13$	0.420	183	$-1.31 \pm 2.51$	0.602
$\Delta$ S-Urea (mmol/L)	242	$0.51 \pm 0.26$	0.05	183	$0.62 \pm 0.31$	0.05

All values are beta ± SE. Statistical differences between changes are based on analysis of covariance.

Beta is the slope coefficient of outcome measures per 1 unit change in protein intake (g/kg/day)

Model Adjusted for age, gender, physical activity (moderate and vigorous), and study site (UCPH, HEL, UOA, UNAV, and UNOTT). Abbreviations: eGFR: estimated glomerular filtration rate; S-Creatinine: serum creatinine; S-Urea: serum urea.

Table S2. Associations between change in urea excretion and change in kidney function from baseline to 1 year

Δ Urea Excretion (mmol/day)						
Variable	n	Unadjusted	<i>p</i> -Value	n	Adjusted	<i>p</i> -Value
		(ß ± SE)	-		$(\mathbf{\hat{B}} \pm \mathbf{SE})$	-
<b>Δ Creatinine clearance</b> (mL/min)	294	$0.22 \pm 0.01$	<.0001	177	$0.21\pm0.01$	<.0001
$\Delta$ eGFR (mL/min/1.73m <sup>2</sup> )	309	$0.003 \pm 0.004$	0.366	183	$0.008 \pm 0.04$	0.07
$\Delta$ Albumin/Creatinine ratio (ACR)	309	$-0.0004 \pm 0.02$	0.978	183	$0.0005 \pm 0.02$	0.975
$\Delta$ Urea/Creatinine Ratio (UCR)	309	$0.03 \pm 0.007$	<.0001	183	$0.04\pm0.008$	<.0001
$\Delta$ S-Creatinine ( $\mu$ mol/L)	309	$-0.003 \pm 0.004$	0.408	183	$-0.008 \pm 0.004$	0.08
$\Delta$ S-Urea (mmol/L)	309	$0.002 \pm 0.0004$	<.0001	183	$0.002 \pm 0.0005$	<.0001

All values are beta ± SE. Statistical differences between changes are based on analysis of covariance.

Beta is the slope coefficient of outcome measures per 1 unit change in urea excretion (mmol/day)

Model Adjusted for age, gender, physical activity (moderate and vigorous), and study site (UCPH, HEL, UOA, UNAV, and UNOTT). Abbreviations: eGFR: estimated glomerular filtration rate; S-Creatinine: serum creatinine; S-Urea: serum urea.

**Table S3** Associations between change in self-reported protein intake and change in kidney function from baseline to 1 year. Additionally adjusted for change in body weight.

Δ Protein Intake (g/kg/day)						
Variable	n	Unadjusted (ß ± SE)	<i>p</i> -Value	n	Adjusted (ß ± SE)	<i>p</i> -Value
<b>Δ Creatinine clearance</b> (mL/min)	233	$-6.70 \pm 9.85$	0.497	193	$-2.87 \pm 10.12$	0.777
$\Delta$ eGFR (mL/min/1.73m <sup>2</sup> )	242	$0.43 \pm 2.14$	0.842	202	$-1.13 \pm 2.36$	0.633
$\Delta$ Albumin/Creatinine ratio (ACR)	242	$-2.62 \pm 7.13$	0.714	202	$-021 \pm 8.59$	0.981
$\Delta$ Urea/Creatinine Ratio (UCR)	242	$7.77 \pm 3.76$	0.04	202	$6.91 \pm 4.29$	0.108
<b>Δ S-Creatinine</b> (μmol/L)	242	$-1.72 \pm 2.13$	0.420	202	$-0.43 \pm 2.32$	0.852
$\Delta$ S-Urea (mmol/L)	242	$0.51 \pm 0.26$	0.05	202	$0.45\pm0.29$	0.125
All values are beta ± SE. Statistical differences between changes are based on analysis of covariance.						

Beta is the slope coefficient of outcome measures per 1 unit change in protein intake (g/kg/day)

Model Adjusted for age, change in body weight from baseline to 1 year follow-up, gender, physical activity (moderate

and vigorous), and study site (UCPH, HEL, UOA, UNAV, and UNOTT)

Abbreviations: eGFR: estimated glomerular filtration rate; S-Creatinine: serum creatinine; S-Urea: serum urea.

$\Delta$ Urea Excretion (mmol/day)						
Variable	n	Unadjusted (ß ± SE)	<i>p</i> -Value	n	Adjusted (ß ± SE)	<i>p</i> -Value
<b>Δ Creatinine clearance</b> (mL/min)	294	$0.22 \pm 0.01$	<.0001	219	$0.20 \pm 0.01$	<.0001
$\Delta$ eGFR (mL/min/1.73m <sup>2</sup> )	309	$0.003 \pm 0.004$	0.366	230	$0.007\pm0.004$	0.06
$\Delta$ Albumin/Creatinine ratio (ACR)	309	$-0.0004 \pm 0.02$	0.978	230	$0.004\pm0.014$	0.786
$\Delta$ Urea/Creatinine Ratio (UCR)	309	$0.03 \pm 0.007$	<.0001	230	$0.03 \pm 0.007$	<.0001
<b>Δ S-Creatinine</b> (µmol/L)	309	$-0.003 \pm 0.004$	0.408	230	$-0.007 \pm 0.004$	0.04*
Δ S-Urea (mmol/L)	309	$0.002 \pm 0.0004$	<.0001	230	$0.002 \pm 0.0004$	<.0001

**Table S4.** Associations between change in urea excretion and change in kidney function from baseline to 1 year. Additionally adjusted for change in body weight.

All values are beta ± SE. Statistical differences between changes are based on analysis of covariance.

Beta is the slope coefficient of outcome measures per 1 unit change in urea excretion (mmol/day)

Model Adjusted for age, gender, change in body weight from baseline to 1 year follow-up, physical activity (moderate and vigorous), and study site (UCPH, HEL, UOA, UNAV, and UNOTT). Abbreviations: eGFR: estimated glomerular filtration rate; S-Creatinine: serum creatinine; S-Urea: serum urea.

**Table S5** Associations between change in self-reported protein intake and change in kidney function from baseline to 1 year. Individuals who improved their glycemic control (n = 191) are excluded from the statistical analysis.

$\Delta$ Protein Intake (g/kg/day)						
Variable	n	Unadjusted (ß ± SE)	<i>p</i> -Value			
Δ Creatinine clearance (mL/min)	49	$6.27 \pm 4.97$	0.213			
$\Delta$ eGFR (mL/min/1.73m <sup>2</sup> )	47	$13.24 \pm 17.2$	0.446			
$\Delta$ Albumin/Creatinine ratio (ACR)	49	$-0.85 \pm 2.20$	0.701			
$\Delta$ Urea/Creatinine Ratio (UCR)	49	$14.19\pm8.65$	0.108			
$\Delta$ S-Creatinine ( $\mu$ mol/L)	49	$-8.44 \pm 4.75$	0.08			
Δ S-Urea (mmol/L)	49	$0.62 \pm 0.55$	0.263			

All values are beta ± SE. Statistical differences between changes are based on analysis of covariance.

Beta is the slope coefficient of outcome measures per 1 unit change in protein intake (g/kg/day)

Abbreviations: eGFR: estimated glomerular filtration rate; S-Creatinine: serum creatinine; S-Urea: serum urea.

**Table S6.** Associations between change in urea excretion and change in kidney function from baseline to 1 year. Individuals who improved their glycemic control (n = 191) are excluded from the statistical analysis.

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$\Delta$ Urea Excretion (mmol/day)								
Variable	n Unadjusted		<i>p</i> -Value					
		$(\mathbf{B} \pm \mathbf{SE})$						
<b>Δ Creatinine clearance</b> (mL/min)	59	$0.17 \pm 0.02$	<.0001					
$\Delta \text{ eGFR} (\text{mL/min}/1.73\text{m}^2)$	61	$0.01 \pm 0.009$	0.137					
$\Delta$ Albumin/Creatinine ratio (ACR)	61	$0.007\pm0.004$	0.076					
$\Delta$ Urea/Creatinine Ratio (UCR)	61	$0.03 \pm 0.01$	0.06					
<b>Δ S-Creatinine</b> (μmol/L)	61	$-0.01 \pm 0.008$	0.146					
$\Delta$ S-Urea (mmol/L)	61	$0.001 \pm 0.001$	0.147					

All values are beta ± SE. Statistical differences between changes are based on analysis of covariance.

Beta is the slope coefficient of outcome measures per 1 unit change in urea excretion (mmol/day)

Abbreviations: eGFR: estimated glomerular filtration rate; S-Creatinine: serum creatinine; S-Urea: serum urea.

Protein Intake (g/kg/day)						
Variable	n	Unadjusted (ß ± SE)	<i>p</i> -Value			
Creatinine clearance (mL/min)	262	$-6.71 \pm 9.02$	0.458			
<b>eGFR</b> (mL/min/1.73m <sup>2</sup> )	266	$-0.94 \pm 3.11$	0.764			
U-Albumin/U-Creatinine ratio (ACR)	266	$-26.51 \pm 15.88$	0.09			
Urea/Creatinine Ratio (UCR)	266	$17.34 \pm 4.03$	<.0001			
S-Creatinine (µmol/L)	266	$-1.75 \pm 3.70$	0.636			
S-Urea (mmol/L)	266	$1.32 \pm 0.32$	<.0001			

All values are beta ± SE. Statistical analysis is based on analysis of covariance

Beta is the slope coefficient of outcome measures per 1 unit change in protein intake (g/kg/day)

Model Adjusted for age, gender, physical activity (moderate and vigorous), and study site (UCPH, HEL, UOA, UNAV, and UNOTT). Abbreviations: eGFR: estimated glomerular filtration rate; S-Creatinine: serum creatinine; S-Urea: serum urea.

Table S8 Associations between self-reported protein intake and markers of the kidney function after 1 year (CID4).

Variable	n	Unadjusted (ß ± SE)	<i>p</i> -Value	
Creatinine clearance (mL/min)	258	$5.19 \pm 7.69$	0.50	
eGFR (mL/min/1.73m <sup>2</sup> )	266	$-0.73 \pm 2.26$	0.75	
U-Albumin/U-Creatinine ratio (ACR)	266	$-31.78 \pm 17.91$	0.07	
Urea/Creatinine Ratio (UCR)	266	$7.55 \pm 3.54$	0.03	
S-Creatinine (µmol/L)	266	$-1.70 \pm 2.61$	0.52	
S-Urea (mmol/L)	266	$0.46 \pm 0.25$	0.06	

All values are beta ± SE. Statistical analysis is based on analysis of covariance.

Beta is the slope coefficient of outcome measures per 1 unit change in protein intake (g/kg/day)

Model Adjusted for age, gender, physical activity (moderate and vigorous), and study site (UCPH, HEL, UOA, UNAV, and UNOTT). Abbreviations: eGFR: estimated glomerular filtration rate; S-Creatinine: serum creatinine; S-Urea: serum urea.

Table S9. Associations between baseline urea excretion and baseline markers of the kidney function (CID1).

Urea Excretion (mmol/day)					
Variable	n	Unadjusted (ß ± SE)	<i>p</i> -Value		
Creatinine clearance (mL/min)	303	$0.18 \pm 0.01$	<.0001		
eGFR (mL/min/1.73m <sup>2</sup> )	309	$0.003 \pm 0.005$	0.559		
U-Albumin/U-Creatinine ratio (ACR)	309	$-0.03 \pm 0.02$	0.216		
Urea/Creatinine Ratio (UCR)	309	$0.01 \pm 0.006$	0.04*		
S-Creatinine (µmol/L)	309	$0.02 \pm 0.006$	< 0.01		
S-Urea (mmol/L)	309	$0.002 \pm 0.005$	<.0001		

All values are beta ± SE. Statistical analysis is based on analysis of covariance.

Beta is the slope coefficient of outcome measures per 1 unit change in urea excretion (mmol/day)

Model Adjusted for age, gender, physical activity (moderate and vigorous), and study site (UCPH, HEL, UOA, UNAV, and UNOTT). Abbreviations: eGFR: estimated glomerular filtration rate; S-Creatinine: serum creatinine; S-Urea: serum urea.

Table S10. Associations between urea excretion and markers of the kidney function after 1 year (CID4).

Urea Excretion (mmol/day)						
Variable	n	Unadjusted (ß ± SE)	<i>p</i> -Value			
Creatinine clearance (mL/min)	303	$0.19\pm0.009$	<.0001			
eGFR (mL/min/1.73m <sup>2</sup> )	309	$0.005\pm0.004$	0.207			
U-Albumin/U-Creatinine ratio (ACR)	309	$-0.03 \pm 0.03$	0.267			
Urea/Creatinine Ratio (UCR)	309	$0.02 \pm 0.006$	<.0001			
S-Creatinine (µmol/L)	309	$0.02 \pm 0.006$	0.052			
S-Urea (mmol/L)	309	$0.002 \pm 0.005$	<.0001			

All values are beta ± SE. Statistical analysis is based on analysis of covariance.

Beta is the slope coefficient of outcome measures per 1 unit change in urea excretion (mmol/day)

Model Adjusted for age, gender, physical activity (moderate and vigorous), and study site (UCPH, HEL, UOA, UNAV, and UNOTT). Abbreviations: eGFR: estimated glomerular filtration rate; S-Creatinine: serum creatinine; S-Urea: serum urea.