

## Supplementary Materials: Associations of serum betaine with blood pressure and hypertension incidence in middle-aged and older adults: a prospective cohort study

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**Supplemental Table 1. Baseline characteristics of hypertensive and nonhypertensive participants**

□ Variables	Hypertensive (n=657)	Nonhypertensive (n=1339)	P-value
Serum betaine, μmol/L	49.9 (39.8-60.8)	51.9 (42.5-61.9)	0.013
Age, years	58.4 (5.0)	56.4 (4.6)	<0.001
Women, n (%)	440 (67.0)	987 (73.7)	0.002
Education attainment, n (%)			<0.001
≤9 years	221 (33.6)	346 (25.8)	
9-12 years	273 (41.6)	687 (51.3)	
≥12 years	163 (24.8)	306 (22.9)	
Household income, n (%)			0.237
≤1500 Yuan/Month/Person	211 (32.1)	475 (35.5)	
1501-3000 Yuan/Month/Person	312 (47.5)	623 (46.5)	
>3000 Yuan/Month/Person	134 (20.4)	241 (18.0)	
Smoker, n (%)	98 (14.9)	179 (13.4)	0.384
Alcohol drinker, n (%)	40 (6.1)	66 (4.9)	0.328
Total energy intake, kcal/d	1813 (512)	1810 (492)	0.905
Chinese Healthy Eating Index	71.1 (7.6)	71.4 (8.1)	0.323
Physical activity, MET-h/d	42.8 (16.1)	43.0 (15.8)	0.836
Waist circumference, cm	85.1 (8.5)	81.0 (8.6)	<0.001
Serum metabolites			
TG, mmol/L	1.45 (1.04-2.07)	1.20 (0.85-1.71)	<0.001
TC, mmol/L	5.48 (1.01)	5.42 (1.05)	0.302
HDL-C, mmol/L	1.31 (0.31)	1.41 (0.33)	<0.001
LDL-C, mmol/L	3.69 (0.89)	3.60 (0.89)	0.025
Choline, μmol/L	20.1 (14.8-26.2)	18.9 (13.8-25.3)	0.023
Homocysteine, μmol/L	13.5 (11.6-17.3)	12.8 (11.0-16.0)	<0.001
Abdominal obesity, n (%)	282 (42.9)	372 (27.8)	<0.001
Prediabetes or diabetes, n (%)	90 (13.7)	73 (5.5)	<0.001
Antihypertension medication use, n (%)	311 (47.3)	0 (0.0)	<0.001
SBP, mmHg	139.5 (15.8)	115.4 (11.3)	<0.001
DBP, mmHg	85.9 (10.0)	73.4 (7.5)	<0.001
Pulse pressure, mmHg	53.6 (13.2)	42.0 (8.4)	<0.001

Data were presented as mean (standard deviation) or median (interquartile range) for continuous variables; and frequency (percentage) for categorical variables.

Abbreviations: MET, metabolic equivalent of task; TG, triglycerides; TC, total cholesterol; HDL-C, high-density lipoprotein cholesterol; LDL-C, low-density lipoprotein cholesterol; SBP, systolic blood pressure; DBP, diastolic blood pressure.

**Supplemental Table 2. Stratified HRs and 95% CIs for incident hypertension according to baseline serum betaine levels (quartiles)**

□	N	Sex-Specific Quartiles of Serum Betaine Levels				P-interaction <sup>a</sup>
		Q1	Q2	Q3	Q4	
<b>Age, years</b>						0.999
≤55.5	669	Ref.	0.84 (0.53, 1.33)	0.78 (0.49, 1.25)	1.01 (0.65, 1.56)	
>55.5	670	Ref.	0.83 (0.58, 1.20)	0.75 (0.51, 1.08)	0.93 (0.64, 1.36)	
<b>Sex</b>						0.873
men	352	Ref.	0.79 (0.48, 1.30)	0.84 (0.51, 1.38)	1.05 (0.63, 1.75)	
women	987	Ref.	0.78 (0.55, 1.11)	0.72 (0.50, 1.03)	0.86 (0.61, 1.22)	
<b>Smoker</b>						0.359
no	1160	Ref.	0.80 (0.59, 1.09)	0.74 (0.54, 1.02)	0.84 (0.62, 1.15)	
yes	179	Ref.	0.74 (0.34, 1.61)	0.79 (0.36, 1.74)	1.51 (0.70, 3.26)	
<b>Alcohol drinker</b>						0.791
no	1273	Ref.	0.82 (0.61, 1.10)	0.75 (0.56, 1.01)	0.94 (0.70, 1.26)	
yes	66	Ref.	0.30 (0.05, 1.76)	0.65 (0.15, 2.84)	0.40 (0.08, 1.98)	
<b>CHEI</b>						0.772
≤72.0	670	Ref.	0.74 (0.48, 1.12)	0.82 (0.55, 1.23)	0.98 (0.66, 1.48)	
>72.0	669	Ref.	0.83 (0.56, 1.22)	0.68 (0.45, 1.03)	0.90 (0.60, 1.35)	
<b>Abdominal obesity</b>						0.217
no	967	Ref.	0.76 (0.53, 1.09)	0.61 (0.42, 0.90)	0.78 (0.55, 1.12)	
yes	372	Ref.	0.85 (0.53, 1.37)	1.01 (0.64, 1.59)	1.22 (0.76, 1.96)	
<b>TG, mmol/L</b>						0.244
<1.7	996	Ref.	0.66 (0.46, 0.95)	0.75 (0.53, 1.07)	0.91 (0.65, 1.30)	
≥1.7	343	Ref.	1.12 (0.70, 1.78)	0.76 (0.45, 1.31)	0.94 (0.56, 1.58)	
<b>HDL-C, mmol/L</b>						0.557
<1.0	115	Ref.	0.65 (0.23, 1.87)	0.24 (0.09, 0.65)	2.07 (0.73, 5.82)	
≥1.0	1224	Ref.	0.82 (0.61, 1.11)	0.78 (0.57, 1.06)	0.92 (0.68, 1.25)	
<b>Prediabetes or diabetes</b>						0.306
no	1266	Ref.	0.84 (0.63, 1.13)	0.80 (0.59, 1.08)	0.95 (0.70, 1.28)	
yes	73	Ref.	0.19 (0.04, 0.81)	0.20 (0.05, 0.86)	0.56 (0.13, 2.35)	
<b>Serum choline, μmol/L</b>						0.173
≤18.9	670	Ref.	0.95 (0.64, 1.42)	0.93 (0.62, 1.37)	0.83 (0.54, 1.27)	
>18.9	669	Ref.	0.67 (0.45, 1.01)	0.59 (0.38, 0.92)	0.99 (0.67, 1.46)	

<sup>a</sup> Interactions between serum betaine and covariates were tested by the likelihood ratio test using the multiplicative interaction term. In the multivariable-adjusted model, confounding factors including age, education attainment, household income, smoker, alcohol drinker, energy intake, CHEI, physical activity, WC, TG, HDL-C, prediabetes or diabetes (yes or no), and serum choline. The medians of age, CHEI, and serum choline were cut-off points.