#### **On-Line Data Supplement**

# Longitudinal patterns of blood pressure, incident cardiovascular events and allcause mortality in normotensive diabetic persons

Authors: Zhijun Wu<sup>1</sup>, Cheng Jin<sup>2</sup>, Anand Vaidya<sup>3</sup>, Wei Jin<sup>1</sup>, Zhe Huang<sup>2</sup>, Shouling Wu<sup>2</sup>, Xiang Gao<sup>4</sup>

### Affiliations:

<sup>1</sup>Department of Cardiology, Ruijin Hospital, Shanghai Jiaotong University School of Medicine, Shanghai, People's Republic of China

<sup>2</sup> Department of Cardiology, Kailuan Hospital, Tangshan, People's Republic of China

<sup>3</sup> Division of Endocrinology, Diabetes, and Hypertension, Brigham and Women's

Hospital and Harvard Medical School, Boston, MA

<sup>4</sup> Department of Nutritional Sciences, Pennsylvania State University, State college, PA

Short title: blood pressure and diabetes

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#### Table S1 Adjusted hazard ratios (HRs) and 95% confidence intervals (95% CIs)

### for all-cause mortality and primary cardiovascular outcomes based on blood

	BP<120/80 versus BP of 120-139/80-89mmHg , HR (95%CI)						
Outcomes	All-cause	All-cause Cardiovascular		Stroke			
	mortality	events	infarction				
Case number <sup>*</sup> , n	81/166	40/137	12/36	28/106			
Model $1^{\dagger}$	1.46 (1.12-1.91)	0.85 (0.60-1.22)	1.02 (0.53-1.96)	0.76 (0.50-1.16)			
Model 2 <sup>‡</sup>	1.42 (1.08-1.88)	0.84 (0.58-1.20)	0.99 (0.50-1.93)	0.75 (0.49-1.15)			
Model 3 <sup>§</sup>	1.46 (1.10-1.93)	0.83 (0.58-1.19)	0.96 (0.49-1.88)	0.75 (0.49-1.14)			
Sensitivity analysis							
Model $4^{\parallel}$	1.39 (0.99-1.95)	1.07 (0.71-1.62)	1.16 (0.56-2.40)	0.94 (0.57-1.55)			
Model 5 <sup>¶</sup>	1.50 (1.12-2.01)	0.79 (0.54-1.16)	0.80 (0.39-1.66)	0.74 (0.47-1.17)			
Model 6 <sup>#</sup>	1.49 (1.12-1.98)	0.81 (0.56-1.17)	0.96 (0.49-1.89)	0.72 (0.47-1.11)			

pressure statu	s at baseline	(2006)	) during	2006-2014
		·	,	

Abbreviations: BP, blood pressure; HR, hazard ratios; CI, confidence interval.

\*The case number in BP<120/80 group versus that in BP of 120-139/80-89mmHg group.

<sup>†</sup>Model1: adjusted for age, sex and glucose-lowering therapy.

<sup>‡</sup>Model2: adjustments as in model1 plus smoking status (never, former, occasional or daily smoker), body mass index, physical activity (inactive, moderately active or vigorously active), low density lipoprotein cholesterol (mg/dL), high density lipoprotein cholesterol (mg/dL), triglyceride (mg/dL) and high sensitivity C-reactive protein (mg/L). <sup>§</sup>Model3: adjustments as in model2 plus alcohol consumption (never, former, occasional or daily), educational level (primary, middle/high school or college/university), occupation (white collar, coalminer or blue collar), fasting blood glucose and estimated Glomerular Filtration Rate (eGFR).

<sup>I</sup>Model4: adjusted for model3 after excluding individuals receiving glucose-lowering therapy.

<sup>#</sup>Model5: adjusted for model3 after excluding individuals using lipid-lowering drugs. <sup>#</sup>Model6: adjusted for model3 after excluding individuals with eGFR<30 ml/minute/1.73 m<sup>2</sup>.

	Change in BP status, mmHg (from 2006 to 2008), HR (95%CI)						
	Stable	<120/80→	<120/80→	120-139/80-89	Stable 120-	120-139/80-89	
	<120/80	120-139/80-89	≥140/90	→<120/80	139/80-89	→≥140/90	
Case	12/200	11/260	8/148	17/236	23/725	40/742	
number/total,							
n							
Model 1 <sup>*</sup>	2.05 (1.01-	1.39 (0.68-	1.71 (0.76-	2.70 (1.44-	1	1.46 (0.88-	
	4.15)	2.85)	3.83)	5.07)	(reference)	2.45)	
Model 2 <sup>†</sup>	2.44 (1.16-	1.23 (0.58-	1.79 (0.79-	2.81 (1.47-	1	1.55	
	5.12)	2.61)	4.06)	5.37)	(reference)	(0.92-2.62)	
Model 3 <sup>‡</sup>	2.35 (1.10-	1.29 (0.60-	2.05 (0.89-	3.04 (1.56-	1	1.63	
	5.01)	2.77)	4.70)	5.92)	(reference)	(0.96-2.77)	
Sensitivity anal	ysis						
Model 4 <sup>§</sup>	0.92 (0.20-	1.24 (0.44-	3.05 (1.14-	2.75 (1.21-	1	1.65	
	4.25)	3.51)	8.18)	6.24)	(reference)	(0.87-3.14)	
Model 5 <sup>∥</sup>	2.44 (1.08-	1.21 (0.54-	2.44 (1.04-	2.81 (1.38-	1	1.64 (0.95-	
	5.51)	2.70)	5.68)	5.75)	(reference)	2.83)	
Model 6 <sup>¶</sup>	2.39 (1.09-	1.36 (0.63-	2.20 (0.95-	3.20 (1.63-	1	1.69 (0.98-	
	5.21)	2.93)	5.08)	6.28)	(reference)	2.89)	
Model 7 <sup>#</sup>	1.95 (0.81-	1.43 (0.63-	2.14 (0.87-	3.89 (1.93-	1 (reference)	1.50 (0.84-	

## Table S2 Adjusted hazard ratios (HRs) and 95% confidence intervals (95% CIs)

of change in blood pressure with risk of all-cause mortality during 2008-2014

	4.69)	3.24)	5.23)	7.85)			2.70)
Model 8**	1.61 (0.69-	0.91 (0.38-	2.02 (0.82-	2.26 (1.09-	1	(reference)	1.23 (0.64-
	3.75)	2.17)	4.97)	4.71)			2.33)
Model 9***	2.25 (1.05-	1.36 (0.63-	2.17 (0.95-	2.89 (1.48-	1	(reference)	1.63 (0.96-
	4.86)	2.93)	4.98)	5.64)			2.77)

Abbreviations: BP, blood pressure; HR, hazard ratios; CI, confidence interval.

\*Model1: adjusted for age, sex and glucose-lowering therapy.

<sup>†</sup>Model2: adjustments as in model1 plus smoking status (never, former, occasional or daily smoker), body mass index, physical activity (inactive, moderately active or vigorously active), low density lipoprotein cholesterol (mg/dL), high density lipoprotein cholesterol (mg/dL), triglyceride (mg/dL) and high sensitivity C-reactive protein (mg/L).

<sup>‡</sup>Model3: adjustments as in model2 plus alcohol consumption (never, former, occasional or daily), educational level (primary, middle/high school or college/university), occupation (white collar, coalminer or blue collar), fasting blood glucose and estimated Glomerular Filtration Rate (eGFR).

<sup>§</sup>Model4: adjusted for model3 after excluding individuals receiving glucose-lowering therapy.

<sup>I</sup>Model5: adjusted for model3 after excluding individuals using lipid-lowering drugs.

Model6: adjusted for model3 after excluding individuals with eGFR<30 ml/minute/1.73 m<sup>2</sup>.

<sup>#</sup>Model7: adjusted for model3 after excluding individuals who suffered cardiovascular

diseases during 2008-2012.

\*\*Model8: adjusted for model3 after excluding individuals who developed hypertension during 2008-2012.

\*\*\*Model9: adjusted for model3 after excluding individuals who with hypotension (SBP<90mmHg or DBP<60mmHg).

## Table S3 Adjusted hazard ratios (HRs) and 95% confidence intervals (95% CIs)

## of change in blood pressure with risk of primary cardiovascular outcomes

during 2008-2014								
Change in BP status, mmHg (from 2006 to 2008), HR (95%CI)								
	Stable	<120/80→	<120/80→	120-139/80-89	Stable 120-	120-139/80-89		
	<120/80	120-139/80-89	≥140/90	→<120/80	139/80-89	→≥140/90		
Case	7/200	8/260	8/148	9/236	28/725	55/742		
number/total,								
n								
Model 1*	0.97 (0.42-	0.82 (0.37-	1.38 (0.63-	1.03 (0.49-	1	1.81 (1.15-		
	2.23)	1.81)	3.04)	2.18)	(reference)	2.86)		
Model 2 <sup>†</sup>	1.04 (0.44-	0.81 (0.37-	1.38 (0.62-	0.97 (0.45-	1	1.89		
	2.45)	1.79)	3.05)	2.07)	(reference)	(1.19-3.02)		
Model 3 <sup>‡</sup>	1.06 (0.45-	0.84 (0.38-	1.46 (0.65-	0.99 (0.46-	1	1.98		
	2.50)	1.87)	3.29)	2.12)	(reference)	(1.24-3.17)		
Sensitivity anal	ysis							
Model 4 <sup>§</sup>	1.11 (0.31-	1.54 (0.65-	3.09 (1.29-	1.00 (0.36-	1	2.28		
	3.97)	3.65)	7.40)	2.77)	(reference)	(1.27-4.09)		
Model 5 <sup>∥</sup>	0.62 (0.21-	0.86 (0.38-	1.56 (0.69-	1.03 (0.48-	1	1.85 (1.14-		
	1.82)	1.92)	3.53)	2.23)	(reference)	3.00)		
Model 6 <sup>¶</sup>	0.91 (0.37-	0.84 (0.38-	1.47 (0.65-	0.98 (0.46-	1	1.99 (1.24-		
	2.27)	1.86)	3.31)	2.11)	(reference)	3.18)		

Abbreviations: BP, blood pressure; HR, hazard ratios; CI, confidence interval.

\*Model1: adjusted for age, sex and glucose-lowering therapy.

<sup>†</sup>Model2: adjustments as in model1 plus smoking status (never, former, occasional or daily smoker), body mass index, physical activity (inactive, moderately active or vigorously active), low density lipoprotein cholesterol (mg/dL), high density lipoprotein cholesterol (mg/dL), triglyceride (mg/dL) and high sensitivity C-reactive protein (mg/L).

<sup>‡</sup>Model3: adjustments as in model2 plus alcohol consumption (never, former, occasional or daily), educational level (primary, middle/high school or college/university), occupation (white collar, coalminer or blue collar), fasting blood glucose and estimated Glomerular Filtration Rate (eGFR).

<sup>§</sup>Model4: adjusted for model3 after excluding individuals receiving glucose-lowering therapy.

<sup>II</sup>Model5: adjusted for model3 after excluding individuals using lipid-lowering drugs. <sup>II</sup>Model6: adjusted for model3 after excluding individuals with eGFR<30 ml/minute/1.73 m<sup>2</sup>.

	Change in BP status, mmHg (from 2006 to 2008), HR (95%CI)						
	Stable	<120/80→	<120/80→	120-139/80-89	Stable 120-	120-139/80-89	
	<120/80	120-139/80-89	≥140/90	→<120/80	139/80-89	→≥140/90	
Case	2/200	2/260	1/148	0/236	6/725	17/742	
number/total,							
n							
Model 1 <sup>*</sup>	1.42 (0.28-	0.99 (0.20-	0.85 (0.10-	—	1	2.67 (1.05-	
	7.12)	4.93)	7.05)		(reference)	6.78)	
Model 2 <sup>†</sup>	1.87 (0.36-	0.94 (0.19-	0.82 (0.10-	—	1	2.84	
	9.82)	4.73)	6.91)		(reference)	(1.10-7.37)	
Model 3 <sup>‡</sup>	1.69 (0.31-	0.76 (0.14-	0.68 (0.07-	_	1	3.34	
	9.11)	4.24)	6.59)		(reference)	(1.27-8.83)	
Sensitivity anal	ysis						
Model 4 <sup>§</sup>	4.69 (0.68-	2.17 (0.33-	2.14 (0.21-	—	1	5.15	
	32.5)	14.1)	22.4)		(reference)	(1.40-19.0)	
Model $5^{\parallel}$	0.65 (0.07-	0.78 (0.14-	0.69 (0.07-	—	1	3.16 (1.18-	
	5.95)	4.39)	6.58)		(reference)	8.42)	
Model 6 <sup>¶</sup>	1.69 (0.31-	0.76 (0.14-	0.67 (0.07-	—	1	3.33 (1.26-	
	9.10)	4.23)	6.54)		(reference)	8.80)	

## Table S4 Adjusted hazard ratios (HRs) and 95% confidence intervals (95% CIs)

of change in blood pressure with risk of myocardial infarction during 2008-2014

Abbreviations: BP, blood pressure; HR, hazard ratios; CI, confidence interval.

\*Model1: adjusted for age, sex and glucose-lowering therapy.

<sup>†</sup>Model2: adjustments as in model1 plus smoking status (never, former, occasional or daily smoker), body mass index, physical activity (inactive, moderately active or vigorously active), low density lipoprotein cholesterol (mg/dL), high density lipoprotein cholesterol (mg/dL), triglyceride (mg/dL) and high sensitivity C-reactive protein (mg/L).

<sup>‡</sup>Model3: adjustments as in model2 plus alcohol consumption (never, former, occasional or daily), educational level (primary, middle/high school or college/university), occupation (white collar, coalminer or blue collar), fasting blood glucose and estimated Glomerular Filtration Rate (eGFR).

<sup>§</sup>Model4: adjusted for model3 after excluding individuals receiving glucose-lowering therapy.

<sup>II</sup>Model5: adjusted for model3 after excluding individuals using lipid-lowering drugs. <sup>II</sup>Model6: adjusted for model3 after excluding individuals with eGFR<30 ml/minute/1.73 m<sup>2</sup>.

	Change in BP status, mmHg (from 2006 to 2008), HR (95%CI)					
	Stable	<120/80→	<120/80→	120-139/80-89	Stable 120-	120-139/80-89
	<120/80	120-139/80-89	≥140/90	→<120/80	139/80-89	→≥140/90
Case	5/200	6/260	7/148	9/236	24/725	40/742
number/total,						
n						
Model 1 <sup>*</sup>	0.78 (0.30-	0.72 (0.29-	1.39 (0.60-	1.20	1	1.51 (0.91-
	2.06)	1.75)	3.23)	(0.56-2.59)	(reference)	2.50)
Model 2 <sup>†</sup>	0.82 (0.30-	0.70 (0.28-	1.40 (0.60-	1.13	1	1.57
	2.21)	1.73)	3.29)	(0.52-2.47)	(reference)	(0.94-2.65)
Model 3 <sup>‡</sup>	0.86 (0.32-	0.75 (0.30-	1.53 (0.65-	1.17 (0.53-	1	1.65
	2.33)	1.87)	3.64)	2.58)	(reference)	(0.97-2.78)
Sensitivity anal	ysis					
Model 4 <sup>§</sup>	0.45 (0.06-	1.44 (0.54-	3.17 (1.23-	1.16 (0.41-	1	1.74
	3.51)	3.81)	8.18)	3.30)	(reference)	(0.90-3.37)
Model 5 <sup>∥</sup>	0.54 (0.16-	0.77 (0.31-	1.66 (0.69-	1.25 (0.56-	1	1.50 (0.87-
	1.87)	1.91)	3.99)	2.75)	(reference)	2.58)
Model 6 <sup>¶</sup>	0.69 (0.23-	0.75 (0.30-	1.55 (0.65-	1.16	1	1.65 (0.97-
	2.05)	1.86)	3.68)	(0.53-2.55)	(reference)	2.78)

of change in blood pressure with risk of stroke during 2008-2014

Table S5 Adjusted hazard ratios (HRs) and 95% confidence intervals (95% CIs)

Abbreviations: BP, blood pressure; HR, hazard ratios; CI, confidence interval.

\*Model1: adjusted for age, sex and glucose-lowering therapy.

<sup>†</sup>Model2: adjustments as in model1 plus smoking status (never, former, occasional or daily smoker), body mass index, physical activity (inactive, moderately active or vigorously active), low density lipoprotein cholesterol (mg/dL), high density lipoprotein cholesterol (mg/dL), triglyceride (mg/dL) and high sensitivity C-reactive protein (mg/L).

<sup>‡</sup>Model3: adjustments as in model2 plus alcohol consumption (never, former, occasional or daily), educational level (primary, middle/high school or college/university), occupation (white collar, coalminer or blue collar), fasting blood glucose and estimated Glomerular Filtration Rate (eGFR).

<sup>§</sup>Model4: adjusted for model3 after excluding individuals receiving glucose-lowering therapy.

<sup>II</sup>Model5: adjusted for model3 after excluding individuals using lipid-lowering drugs. <sup>II</sup>Model6: adjusted for model3 after excluding individuals with eGFR<30 ml/minute/1.73 m<sup>2</sup>.



Figure S1 Flow chart of the participants with diabetes utilized in this study.



Figure S2 Adjusted hazard ratios (HRs) and 95% confidence intervals (95% CIs) of all-cause mortality during 2008-2014 according to blood pressure trajectories from 2006 to 2008 after exclusion of new onset cancers (Figure S2A), annual weight loss >5% from 2006 to 2008 (Figure S2B) and body mass index <18.5 kg/m<sup>2</sup> at baseline (Figure S2C). All models were adjusted for age (years), sex, smoking status (never, former, occasional or daily), glucose-lowering therapy, body mass index (kg/m<sup>2</sup>), high density lipoprotein cholesterol (mmol/L), low density lipoprotein cholesterol (mmol/L), triglyceride (mmol/L), alcohol consumption (never, former, occasional or daily), educational level (primary, middle/high school or college/university), occupation (white collar, coalminer or blue collar), physical activity (inactive, moderately

active or vigorously active), high sensitivity C-reactive protein (mg/L), fasting blood glucose (mmol/L) and estimated Glomerular Filtration Rate

(eGFR)( mL/min/1.73m<sup>2</sup>).



Figure S3 Adjusted hazard ratios (HRs) and 95% confidence intervals (95% CIs) of cardiovascular events during 2008-2014 according to blood pressure trajectories from 2006 to 2008 in individuals diagnosed prior to recruitment (Figure S3A) and in undiagnosed diabetic persons (Figure S3B). Primary cardiovascular outcomes included fatal or non-fatal myocardial infarction events and fatal or non-fatal stroke events. All models were adjusted for age (years), sex, smoking status (never, former, occasional or daily), glucose-lowering therapy, body mass index (kg/m<sup>2</sup>), high density lipoprotein cholesterol (mmol/L), low density lipoprotein cholesterol (mmol/L), triglyceride (mmol/L), alcohol consumption (never, former, occasional or daily), educational level (primary, middle/high school or college/university), occupation (white collar, coalminer or blue collar), physical activity (inactive, moderately active or vigorously active), high sensitivity C-reactive protein (mg/L), fasting blood glucose (mmol/L) and estimated Glomerular Filtration Rate  $(eGFR)(mL/min/1.73m^2).$