

Hypertension series: Isolated diastolic hypertension

Jasmine Tan

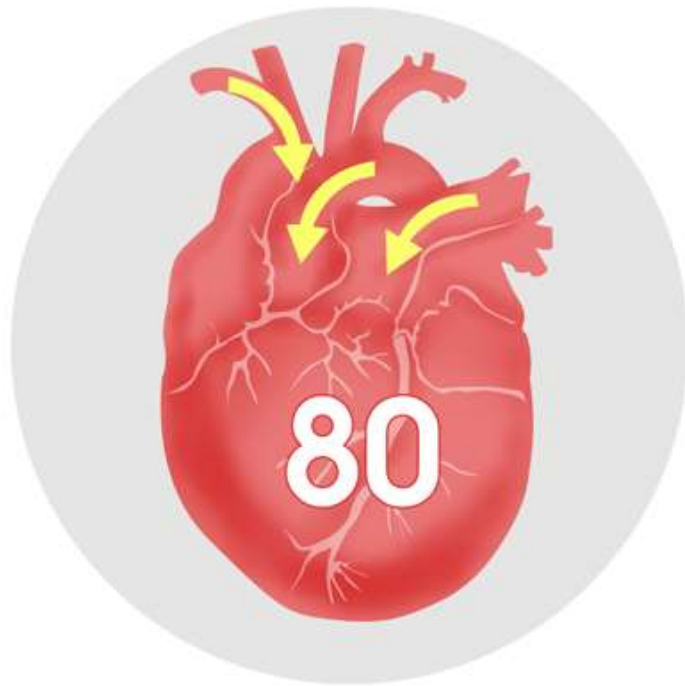
17th February 2024

GLMS symposium

Who would you consider treating?

Mr A	Mr B	Mr A
<p>35 M</p> <ul style="list-style-type: none">• HbA1 49 mmol/mol• CKD G2 A1• Dyslipidemia• Strong family history of hypertension, no CV events• Increased BMI• Mean BP 128/94	<p>62 M</p> <ul style="list-style-type: none">• Normal lipids• No diabetes, no CVD• Increased BMI• Normal kidney function• Non smoker• Hyperuricemia• Mean BP 130/94	<p>40 M</p> <ul style="list-style-type: none">• No diabetes or• Normal kidney function• No secondary causes • Mean BP 150/90 mmHg

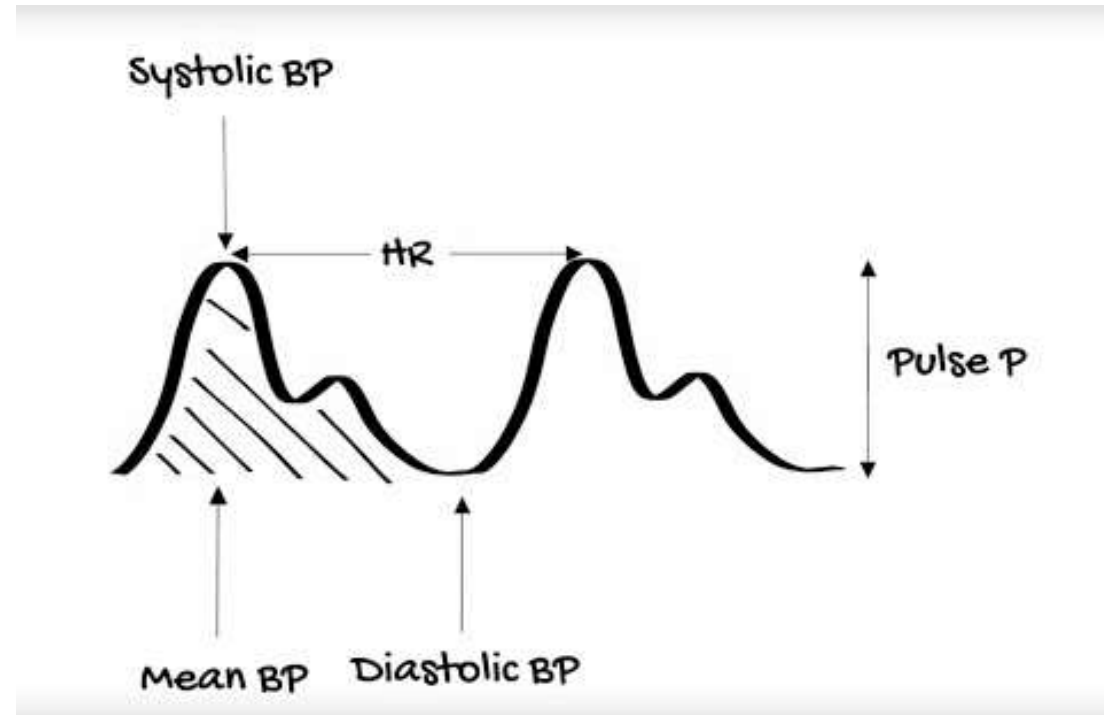
Blood pressure: force against blood vessel wall

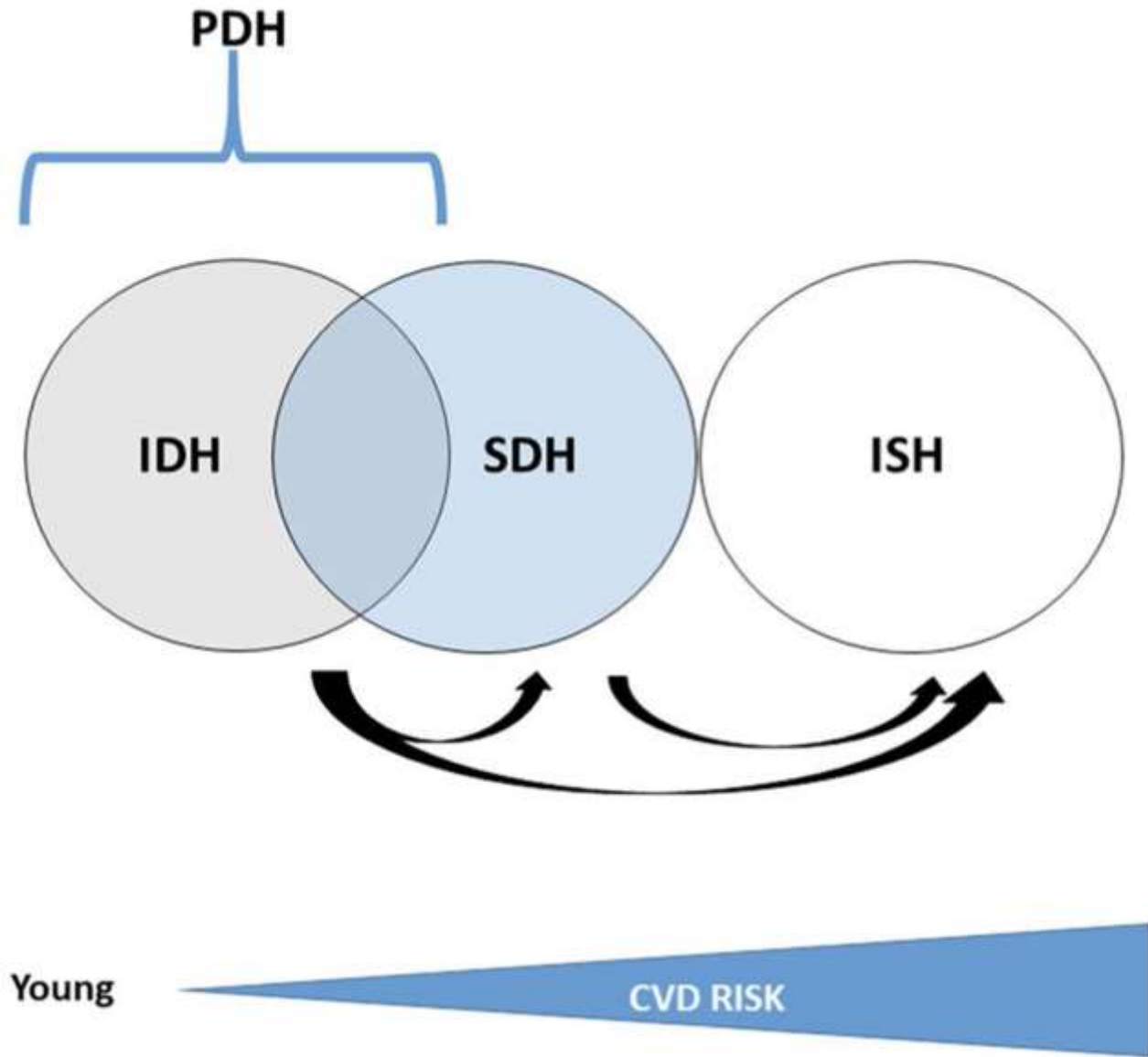


Systolic | Diastolic

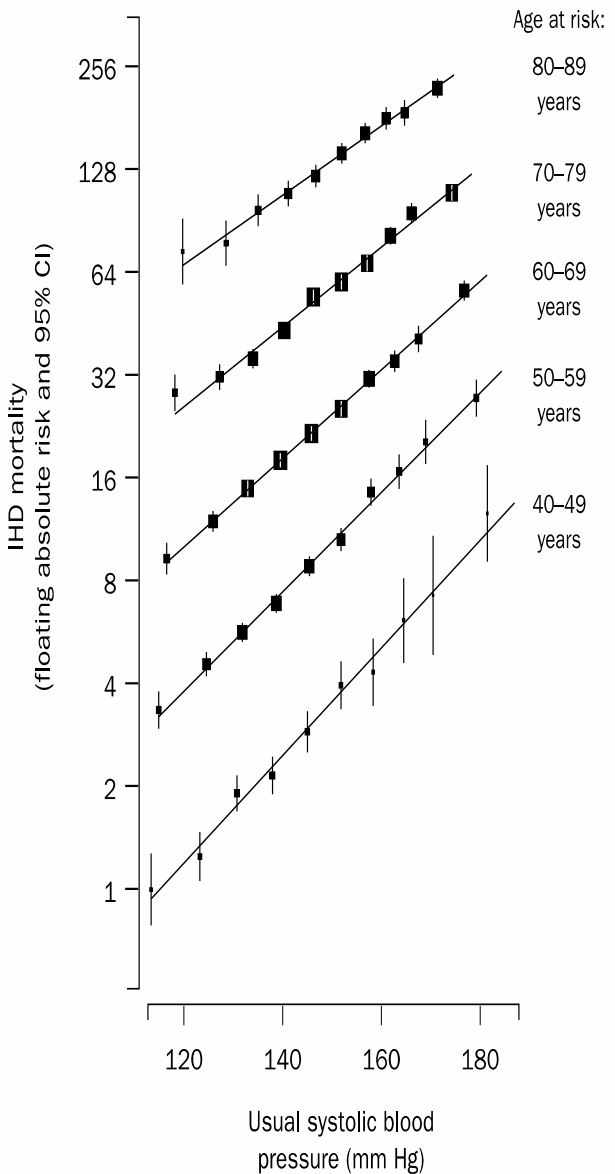
The highest, normal blood pressure reading: 120/80

- Systolic BP – maximal BP peak (cardiac ejection phase)
- Diastolic BP – minimal pressure (relaxed cardiac phase)

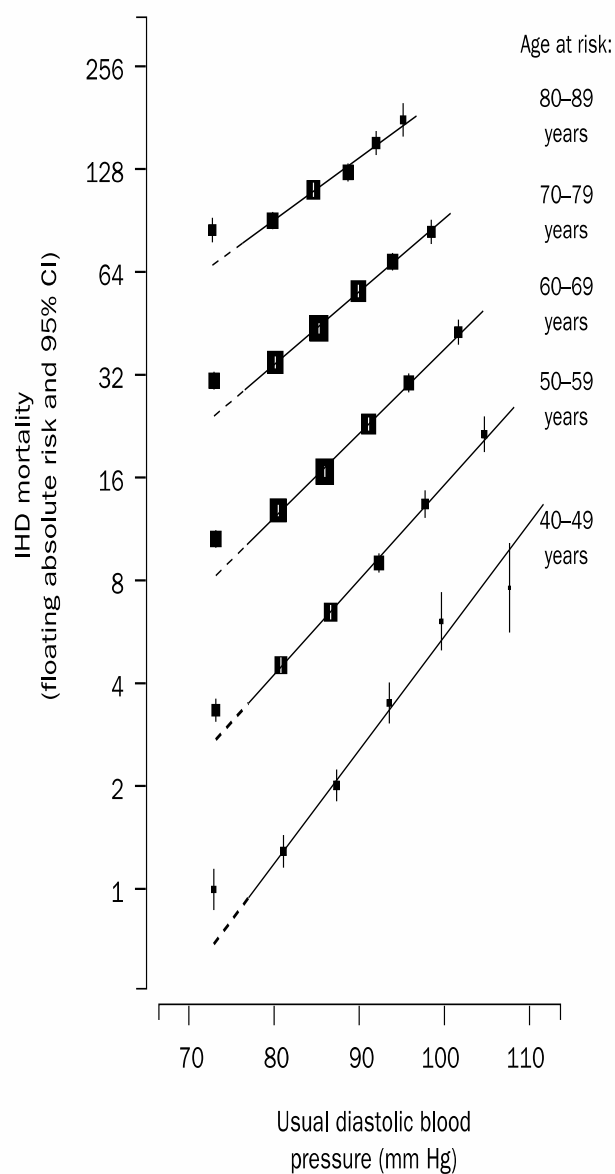




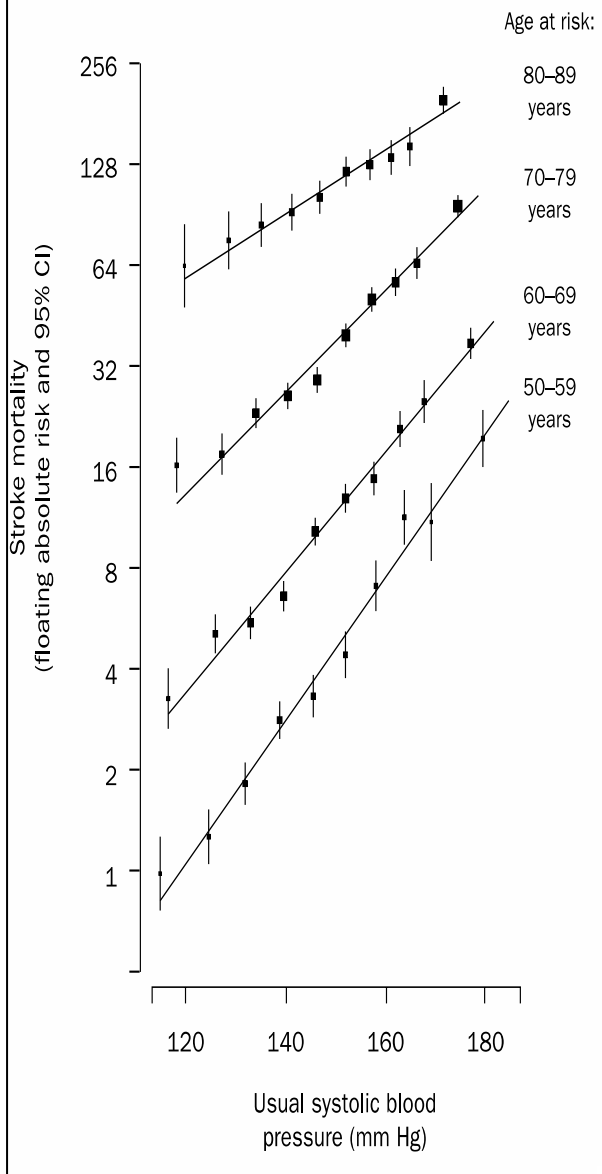
A: Systolic blood pressure



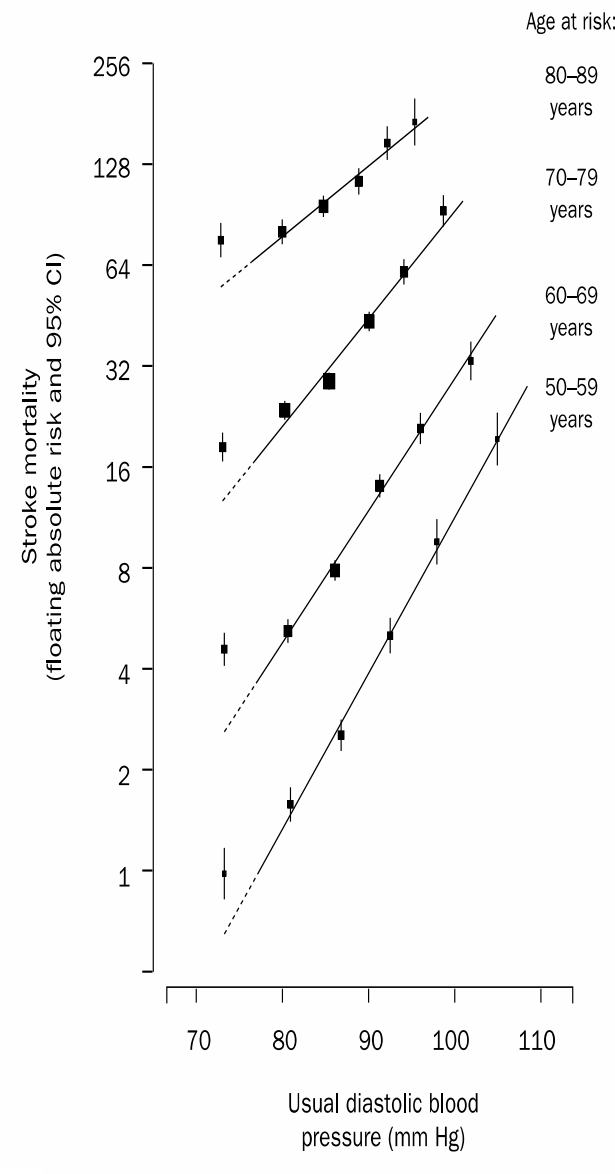
B: Diastolic blood pressure



A: Systolic blood pressure



B: Diastolic blood pressure



- Hypertension contribute to CV disease burden
- Systolic BP most important association to CV events
 - Minimum participant age in major hypertension trials is 45.
 - Elevated DBP 80 – 90 mmHg, in the context of well controlled SBP, is not associated with excess risk of CVD risk.
 - IDH in elderly (>90 mmHg) demonstrated weak influence on CV risk.
- Subgroups of people with IDH and ISH were unknown

Does Isolated diastolic hypertension (IDH) matter ?

	ISH	IDH	SDH
Overall prevalence	1.57	2.66	0.93
Male	2.23	4.09	1.42
Female	0.92	1.25	0.46
ACC/AHA 2017 increment (%)	-10	4.3	30.5

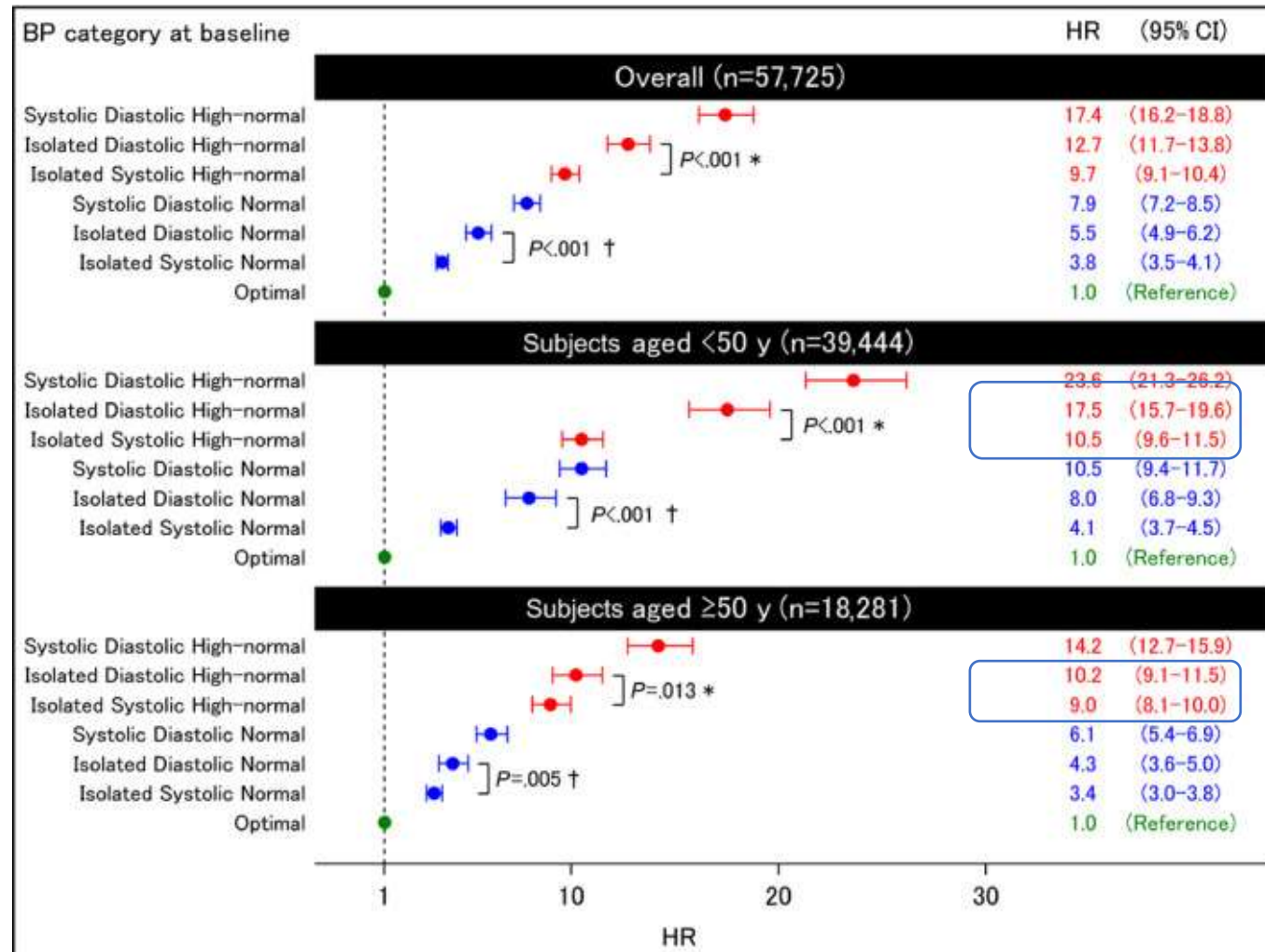
Isolated diastolic hypertension (IDH)

- Prevalence is 1.3 – 6.5% (DBP >90 mmHg and >80 mmHg respectively) in the US population; 7.8 – 24.7% in the Chinese population.
- More frequent hypertensive phenotype in people <50 years
 - Peak prevalence in 30 – 49 years age group
 - Prevalence in individuals <40 years 2.6%
 - Prevalence in individuals < 18 years 1.9%
- IDH increases the risk for the development of incident systolic hypertension
 - HR 23 over 10 years compared to normotensive patients

IDH – what is the risk?

IDH – risk of developing new hypertension

Ref <120/80 mmHg



Mean age 41 years
4.9 year follow up

ISN: SBP 120–129 & DBP <80 mmHg
 IDN: SBP <120 & DBP 80–84 mmHg
 SDN: SBP 120–129 & DBP 80–84 mmHg
 ISHN: SBP 130–139 & DBP <85 mmHg
 IDHN: SBP <130 & DBP 85–89 mmHg
 SDHN: SBP 130–139 & DBP 85–89 mmHg

Cardiovascular Risk of Isolated Diastolic Hypertension Defined by the 2017 American College of Cardiology/American Heart Association Blood Pressure Guideline

A Nationwide Age-Stratified Cohort Study

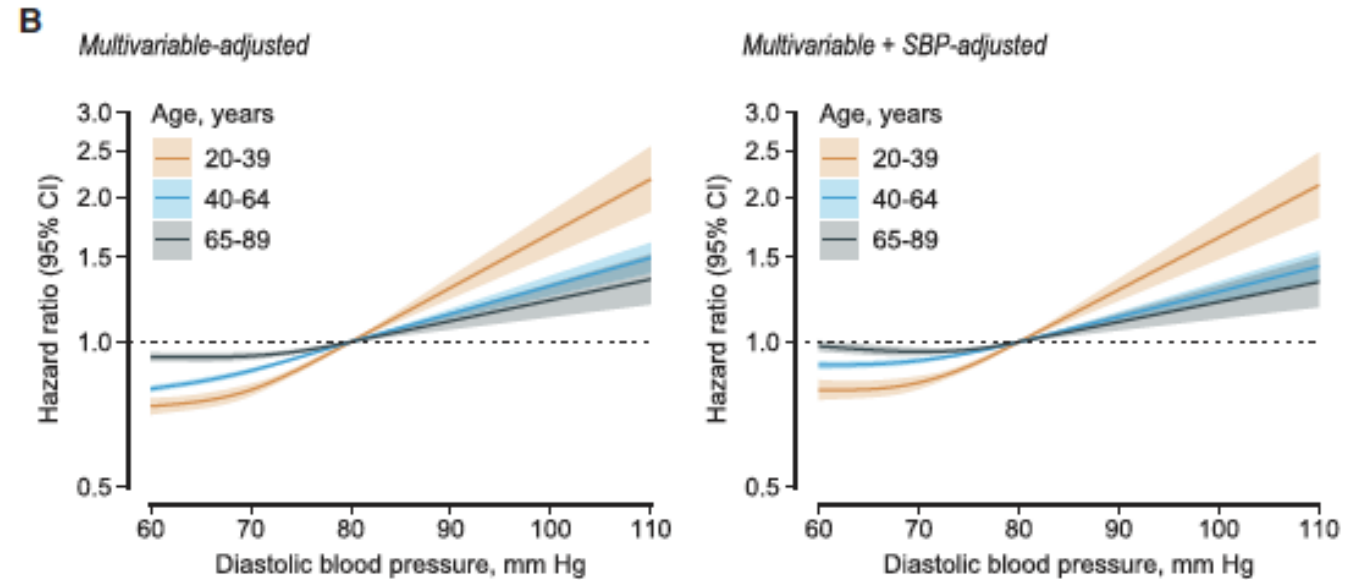
Hokyoo Lee, Yuichiro Yano, So Mi Jemma Cho, Sungha Park, Donald M. Lloyd-Jones, Hyeon Chang Kim

8, 109, 484 participants
 Median age 40 years; 49.5% men
 Follow up 15.3 years
 Composite CV events and mortality

A

Age, y	<130/<80 mm Hg		<130/≥80 mm Hg		<130/≥80 mm Hg vs. <130/<80 mm Hg			
	Events	Person-ys	Events	Person-ys	Low risk	High risk	HR (95% CI)	P interaction
<i>Multivariable-adjusted</i>								
20-39	13,417	39,497,472	9,855	16,777,233			1.30 (1.26-1.33)	<0.001
40-64	78,458	40,460,839	47,698	17,886,178			1.19 (1.17-1.20)	<0.001
65-89	39,626	3,120,911	18,223	1,319,282			1.09 (1.07-1.11)	<0.001
<i>Multivariable + SBP-adjusted</i>								
20-39	13,417	39,497,472	9,855	16,777,233			1.23 (1.19-1.26)	<0.001
40-64	78,458	40,460,839	47,698	17,886,178			1.11 (1.10-1.13)	<0.001
65-89	39,626	3,120,911	18,223	1,319,282			1.07 (1.05-1.09)	<0.001

Note: The forest plot shows HR values for each age group. The x-axis ranges from 1.0 to 1.4. A vertical line is at 1.0. Arrows indicate 'Low risk' to the left and 'High risk' to the right.



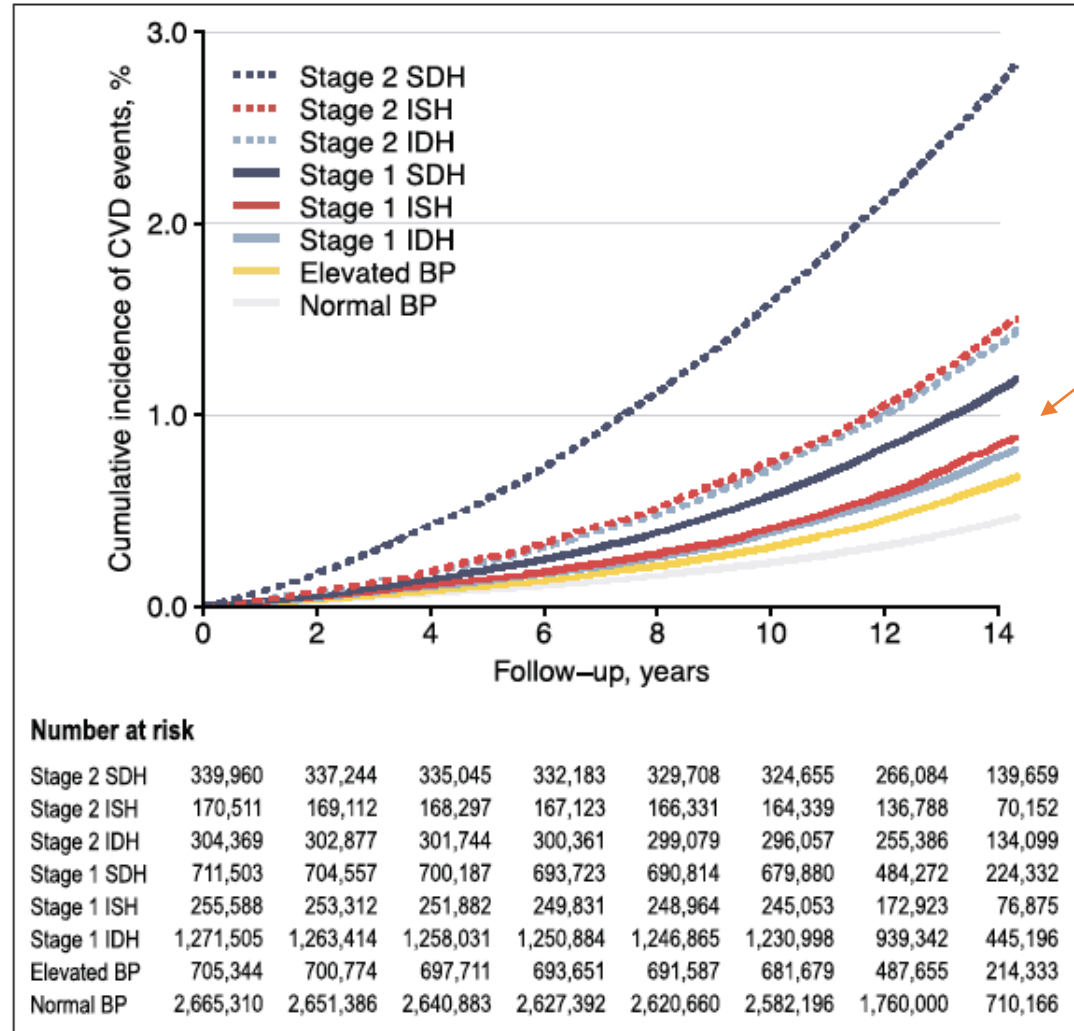
ORIGINAL RESEARCH ARTICLE

Cardiovascular Risk of Isolated Systolic or Diastolic Hypertension in Young Adults

Lee et al.

6, 424, 090 participants
 Median age 30 years; 61% men
 Follow up 13.2 years
 composite CV events and mortality

Normal BP	<120/80 mmHg	REF
Stage 1 ISH	130–139/ <80 mmHg	HR 1.32 (1.28–1.36)
stage 1 IDH	<130/ 80–89 mmHg	HR 1.36 (1.61–1.72)
Stage 1 SDH	130-139/ 80-89 mmHg	HR 1.67 (1.61–1.72)



Isolated Diastolic Hypertension in the IDACO Study: An Age-Stratified Analysis Using 24-Hour Ambulatory Blood Pressure Measurements

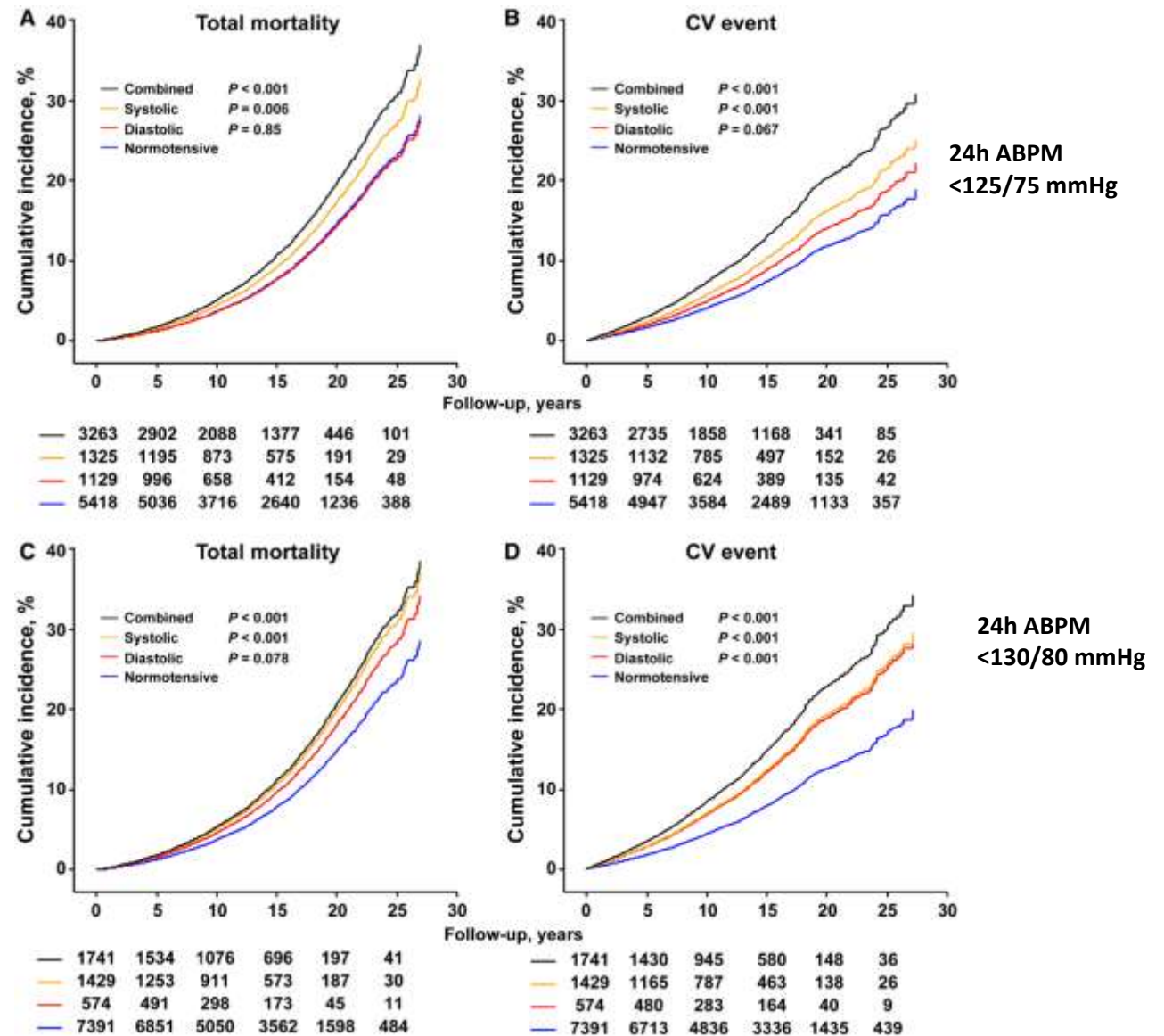
John W. McEvoy¹, Wen-Yi Yang², Lutgarde Thijs³, Zhen-Yu Zhang⁴, Jesus D. Melgarejo⁵, José Boggia⁶, Tine W. Hansen⁷, Kei Asayama⁸, Takayoshi Ohkubo⁹, Eamon Dolan, Katarzyna Stolarz-Skrzypek, Sofia Malyutina, Edoardo Casiglia¹⁰, Lars Lind¹¹, Jan Filipovský¹², Gladys E. Maestre¹³, Yan Li¹⁴, Ji-Guang Wang¹⁵, Yutaka Imai, Kalina Kawecka-Jaszcz, Edgardo Sandoya¹⁶, Krzysztof Narkiewicz¹⁷, Eoin O'Brien, Thomas Vanassche, Jan A. Staessen¹⁸; on behalf of the International Database on Ambulatory Blood Pressure in Relation to Cardiovascular Outcomes (IDACO) Investigators

11,135 participants

<50 vs >50 years old (median age 54.7, 51% men)

FU 13.8 years

2017 ACC/AHA and 2018 ESC guidelines



Isolated Diastolic Hypertension in the IDACO Study: An Age-Stratified Analysis Using 24-Hour Ambulatory Blood Pressure Measurements

John W. McEvoy¹, Wen-Yi Yang², Lutgarde Thijs³, Zhen-Yu Zhang⁴, Jesus D. Melgarejo⁵, José Boggia⁶, Tine W. Hansen⁷, Kei Asayama⁸, Takayoshi Ohkubo⁹, Eamon Dolan, Katarzyna Stolarz-Skrzypek, Sofia Malyutina, Edoardo Casiglia¹⁰, Lars Lind¹¹, Jan Filipovský¹², Gladys E. Maestre¹³, Yan Li¹⁴, Ji-Guang Wang¹⁵, Yutaka Imai, Kalina Kawecka-Jaszcz, Edgardo Sandoya¹⁶, Krzysztof Narkiewicz¹⁷, Eoin O'Brien, Thomas Vanassche, Jan A. Staessen¹⁸; on behalf of the International Database on Ambulatory Blood Pressure in Relation to Cardiovascular Outcomes (IDACO) Investigators

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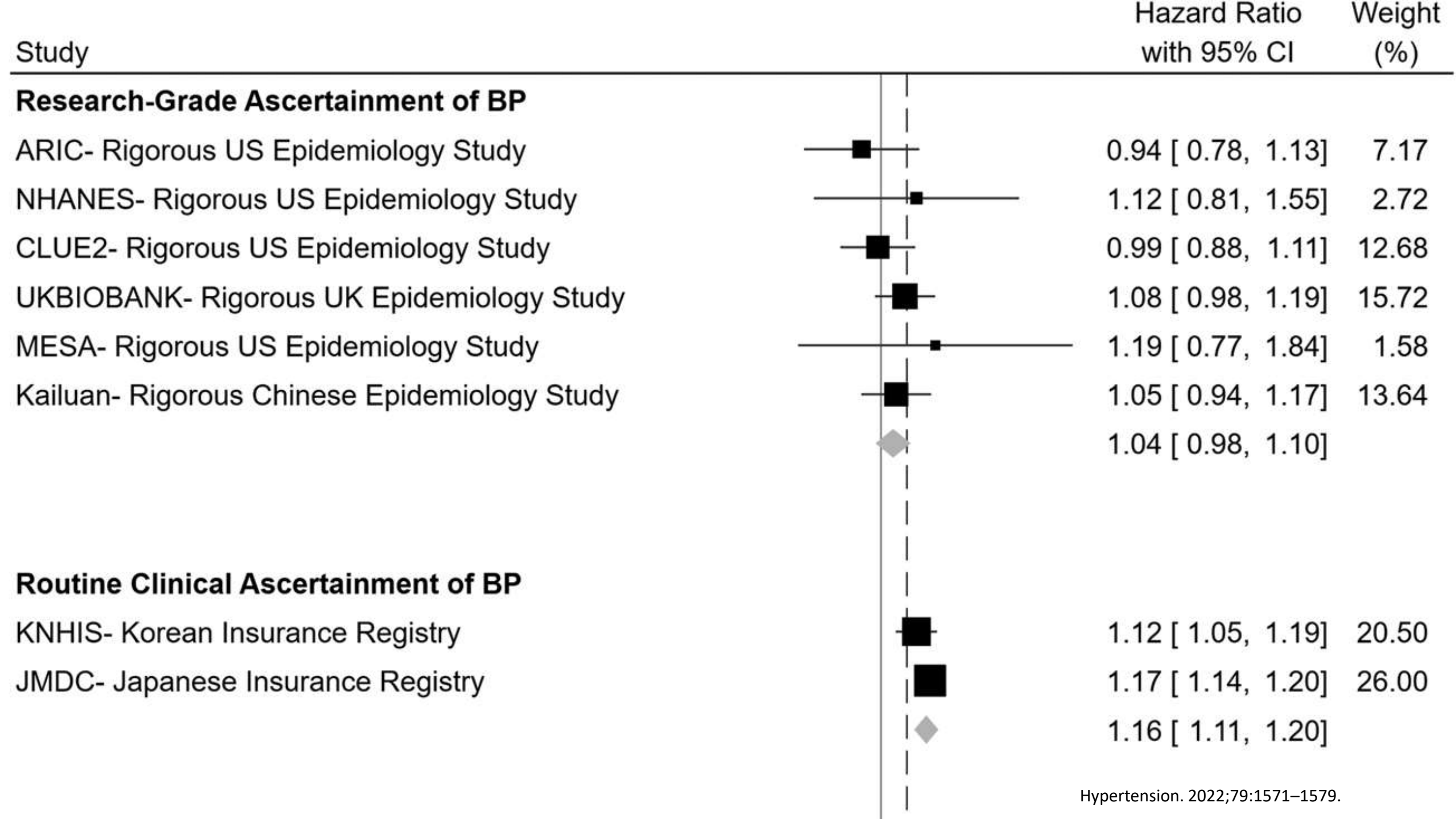
<50 vs >50 years old (median age 54.7, 49% F)

FU 13.8 years

2017 ACC/AHA and 2018 ESC guidelines

Table 4. Multivariable-Adjusted HR in Relation to Hypertension Categories After Stratification by Age

Characteristic	Total mortality				<i>P</i> _{int}	CV events				<i>P</i> _{int}
	Age<50		Age≥50			Age<50		Age≥50		
	HR (95% CI)	<i>P</i> value	HR (95% CI)	<i>P</i> value	HR (95% CI)	<i>P</i> value	HR (95% CI)	<i>P</i> value		
2017 AHA/ACC criteria										
Normotensive	Reference		Reference			Reference		Reference		
Isolated diastolic	1.66 (0.96–2.86)	0.068	0.91 (0.75–1.10)	0.31	0.076	2.87 (1.72–4.80)	<0.001	0.98 (0.78–1.23)	0.87	<0.001
Isolated systolic	0.68 (0.21–2.20)	0.52	1.14 (1.03–1.28)	0.015	0.28	0.74 (0.22–2.43)	0.62	1.33 (1.17–1.52)	<0.001	0.45
Combined	2.08 (1.33–3.25)	0.0013	1.33 (1.21–1.46)	<0.001	0.15	2.39 (1.47–3.89)	<0.001	1.68 (1.50–1.88)	<0.001	0.092



Cross-classification by systolic and diastolic blood pressure levels and chronic kidney disease, proteinuria, or kidney function decline

Tsukasa Suenaga¹ · Michihiro Satoh^{1,2} · Takahisa Murakami^{1,2,3} · Takuo Hirose^{4,5,6} · Taku Obara^{2,7} · Shingo Nakayama^{1,5} · Hideaki Hashimoto^{1,2,5} · Maya Toyama^{1,2,8} · Tomoko Muroya^{1,2,9} · Atsuhiro Kanno¹⁰ · Takefumi Mori⁵ · Takayoshi Ohkubo^{11,12} · Yutaka Imai¹² · Hirohito Metoki^{1,2,12}

1,492,291 participants

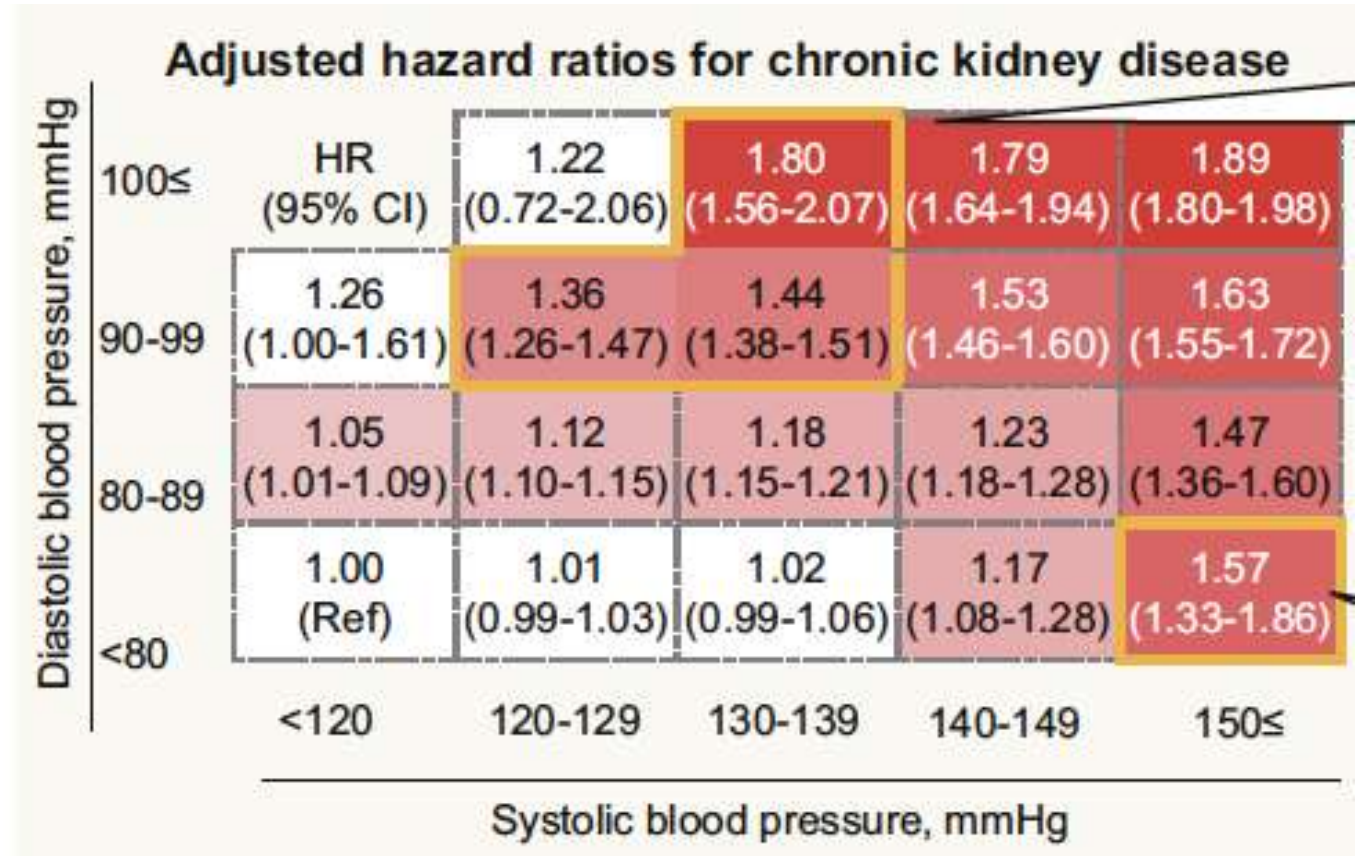
No CKD and no antihypertensive agents at baseline

Median age 41 years; 61% men

FU 3.2 years

2017 ACC/AHA ref (<120/<80 mmHg)

Incident CKD (eGFR <60 ml/min +/- proteinuria)



IDH – what is the risk?

- Age-specific risk with IDH
- IDH and ISH associated with 13 and 10 times risk of hypertension, respectively
- DBP threshold of <75 mmHg appear to be an adverse prognostic factor in people <50 years for CV events
- IDH (and ISH) associated with CKD risk
- Threshold for DBP target <75 – 80 mmHg

Who would you consider treating?

Mr A	Mr B	Mr C
<p>35 M</p> <ul style="list-style-type: none">• CKD G2 A2• HbA1 54 mmol/mol• Dyslipidemia• Strong family history of hypertension and CV events• Increased BMI• Mean BP 130/94	<p>62 M</p> <ul style="list-style-type: none">• Normal kidney function• No diabetes, no CVD• Normal lipids• Increased BMI• Mean BP 130/94	<p>40 M</p> <ul style="list-style-type: none">• No diabetes• Normal kidney function• Normal lipids• No secondary causes• Mean BP 150/90 mmHg

Therapeutic considerations

- Observational studies
- Guidelines do not address management of IDH as a distinct phenotype
- Approach patient
 - Age-specific (Low absolute risk vs cumulative risk in <50 yo)
 - Co-existing metabolic risks, CV risk scores
- What is too low?
 - No specific study addressing lower level of DBP alone on increased CV risk
 - HOT trial showed no J-shaped associated with CV events at DBP 70 mmHg
- Low risk: periodic testing, aggressive lifestyle intervention
- High risk: Pharmacotherapy may be considered. Theoretical benefit of the use vasodilators

Questions?

- Thanks for your attention