

Outpatient Hypertension Management

UW Medicine | Valley Medical Center Grand Rounds

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Disclosures

- None

Objectives

- Epidemiology of hypertension
- Timeline of hypertension management guidelines
- Measuring blood pressure
- Current guidelines for hypertension management
 - 2017 ACC/AHA
 - 2017 ACP/AAFP in older adults
 - 2021 KDIGO guidelines for blood pressure management in CKD
 - 2022 AAFP
- Young adults?

Epidemiology

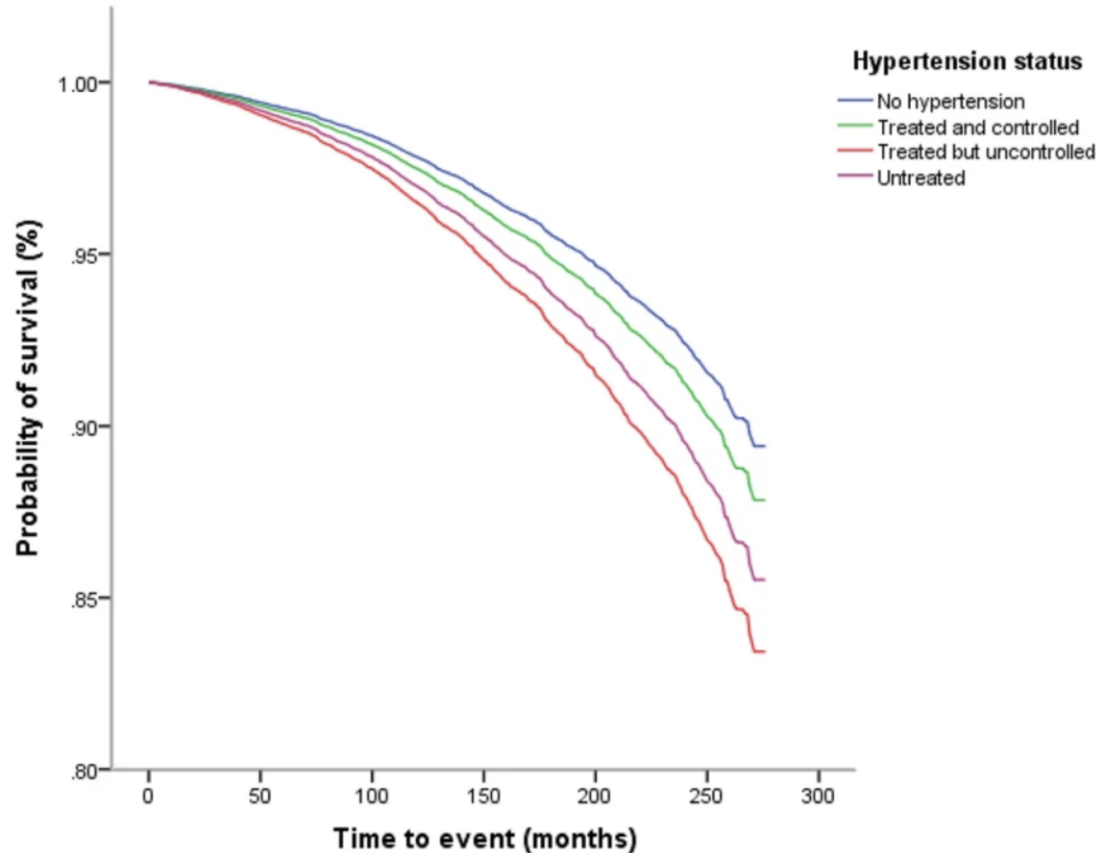
- 1.28 billion adults world-wide aged 30-79 yrs have HTN (>140/90)
- ~ 46% of adults with HTN are unaware
- <42% of adults with HTN are diagnosed and treated
- 1/5 adults with HTN have it under control
- HTN is the major cause of premature death world-wide

Uncontrolled HTN*: All-cause & CVD mortality

*JNC 8: <60 yrs BP <140/90, ≥ 60 yrs BP <150/90

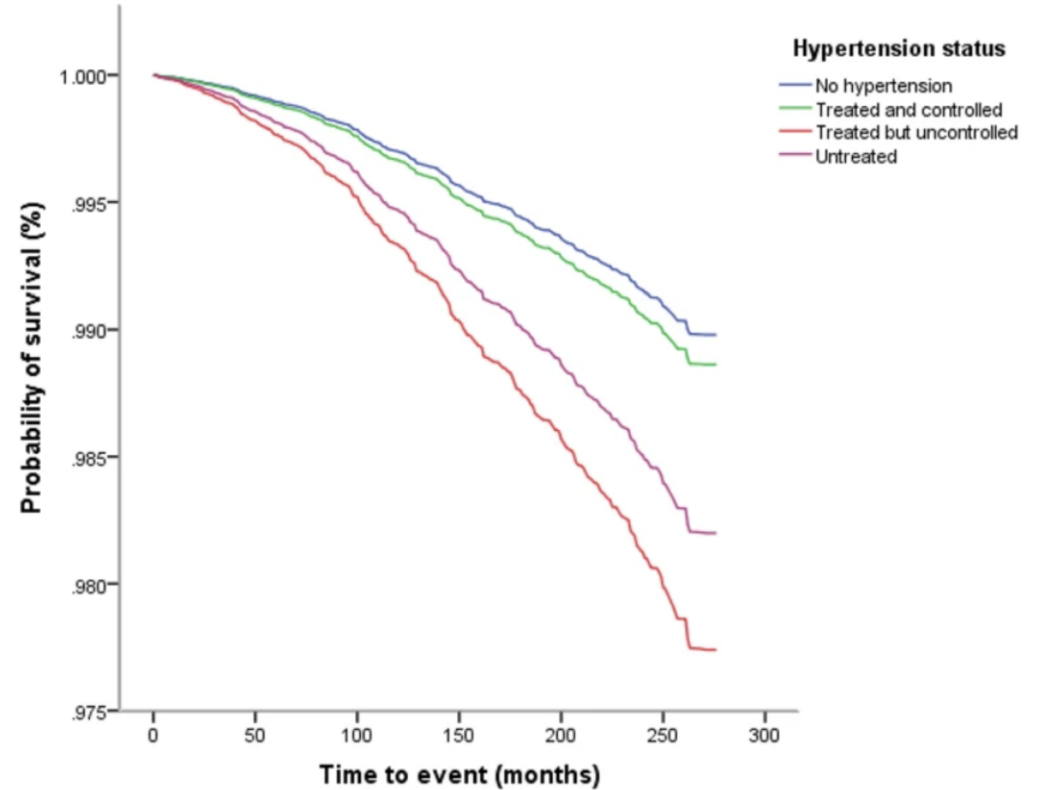
A.

All-cause

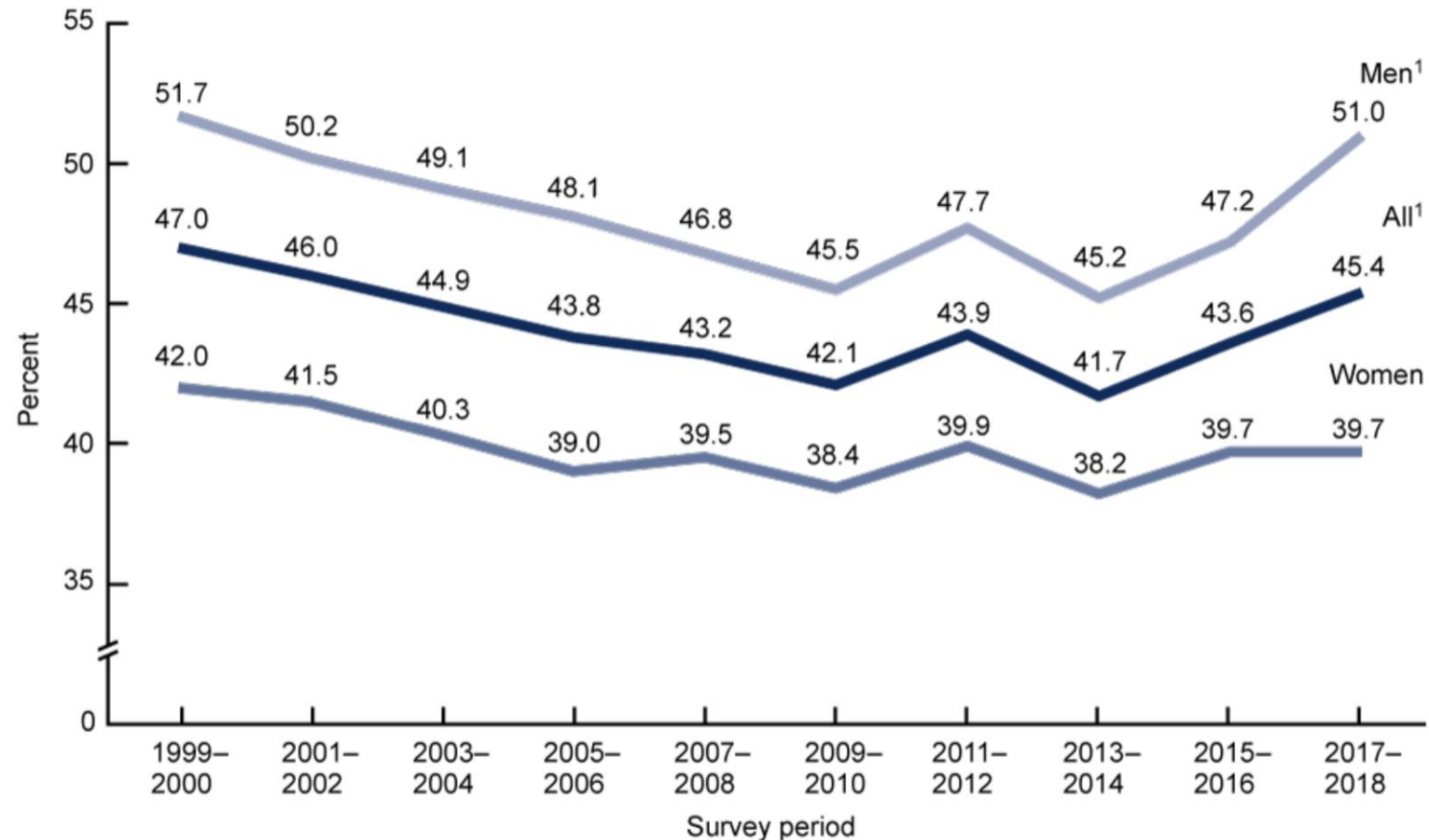


B.

CVD

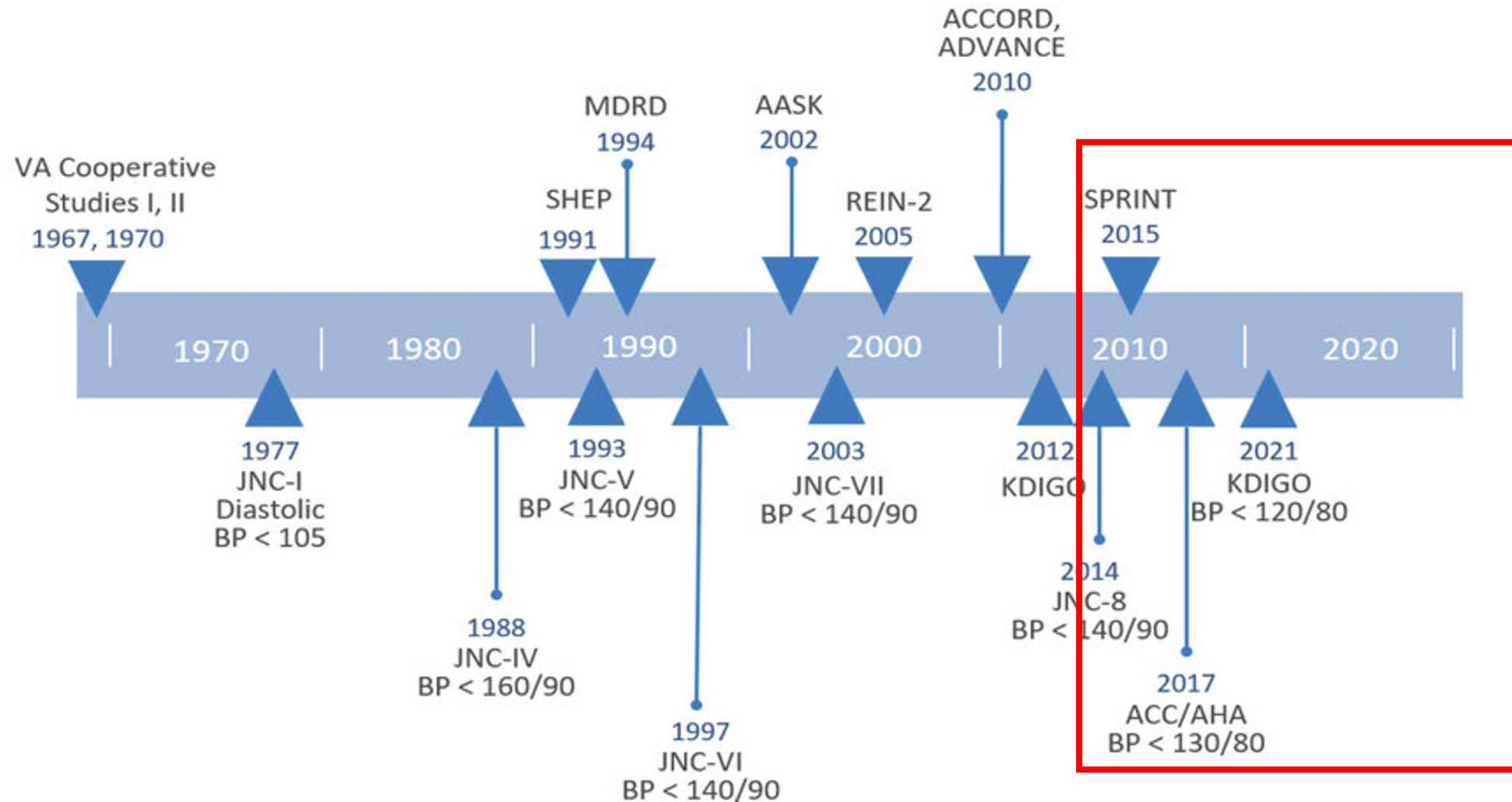


Age-adjusted trend in HTN prevalence among US adults; 1999-2018 (NHANES)

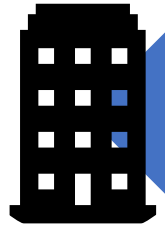


HTN defined as SBP >130, DBP >80 or being on anti-hypertensive medications

HTN management trials/guidelines



BP measurement



Office (OBPM)
(1) Standardized (2) Routine

≠ standardized BP protocols in RCT



Home (HBPM)

The USPSTF (2021), Level A



24-h ambulatory (ABPM)

Labile, nocturnal, white-coat, masked hypertension

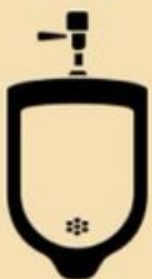
STANDARDISED OFFICE BLOOD PRESSURE MEASUREMENT



PREPARATION



30 min prior



Empty bladder



>5 minutes

TECHNIQUE



- ✓ Validate
- ✓ Calibrate
- ✓ Manual or Auto



- ✓ Arm supported
- ✓ Cuff at RA level
- ✓ Correct size
- ✓ No clothing under cuff



- ✓ Palpate to estimate SBP
- ✓ Inflate 20-30mmHg higher
- ✓ Deflate 2mmHg/sec
- ✓ Korotkoff sounds

MEASUREMENT



- ✓ Both arms
- ✓ Higher arm noted for future use
- ✓ Repeat 1-2 minutes later



- ✓ Record SBP/DBP
- ✓ Average
 - ≥2 readings
 - ≥2 occasions
- ✓ Note time of BP medication



Tell patient the BP verbally & in writing

Out of Office BPM can complement Standardised BPM



Home-based BPM

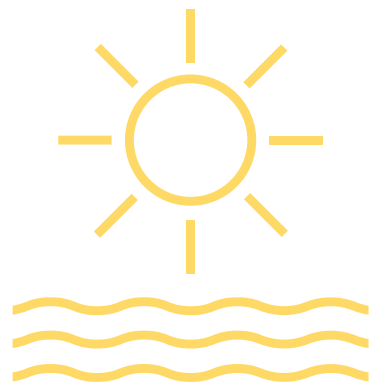


Ambulatory BPM

Can be used to detect white coat HTN / effect and masked HTN

HBPM

2



2



3 days



<https://www.validatebp.org/>

How to measure your blood pressure at home

TARGET:BP™



Follow these steps for an accurate blood pressure measurement

1. PREPARE

Avoid caffeine, smoking and exercise for 30 minutes before measuring your blood pressure.

Wait at least 30 minutes after a meal.

If you're on blood pressure medication, measure your BP *before* you take your medication.

Empty your bladder beforehand.

Find a quiet space where you can sit comfortably without distraction.

2. POSITION



3. MEASURE

Rest for five minutes while in position before starting.

Take two or three measurements, one minute apart, twice daily for seven days.

Keep your body relaxed and in position during measurements.

Sit quietly with no distractions during measurements—avoid conversations, TV, phones and other devices.

Record your measurements when finished.

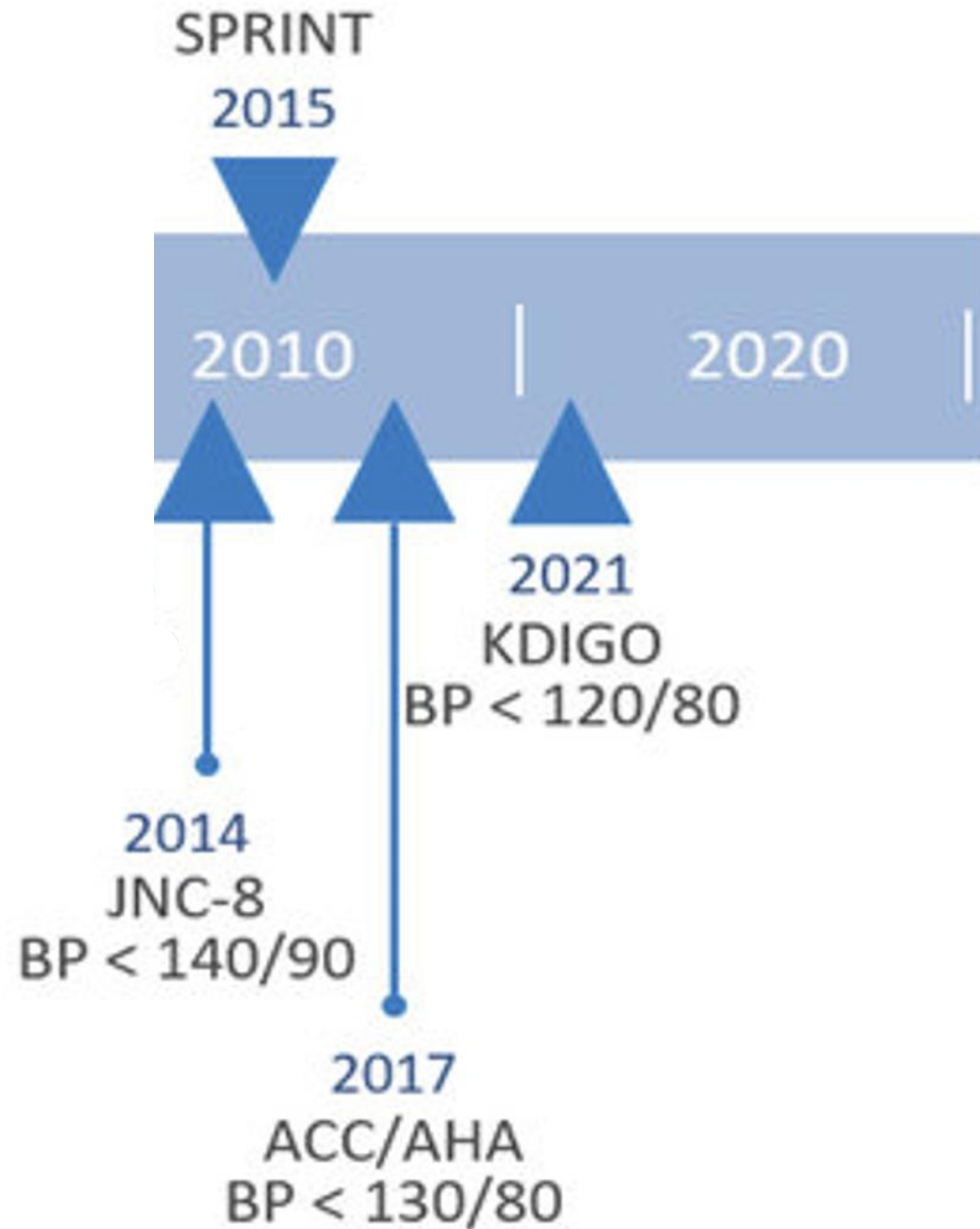
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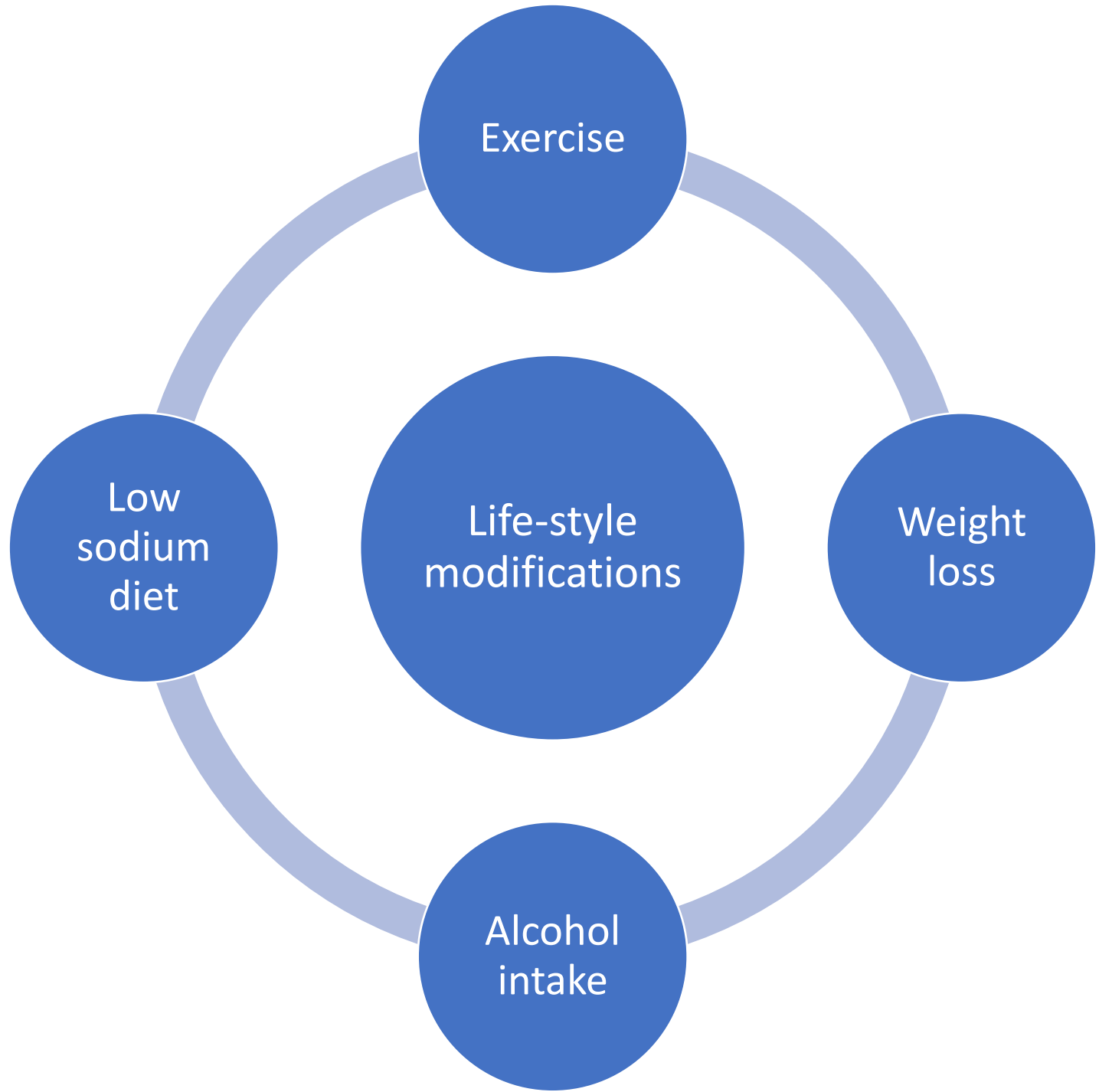


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10/20 MRG15940-6B

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HTN management trials/guidelines





Exercise

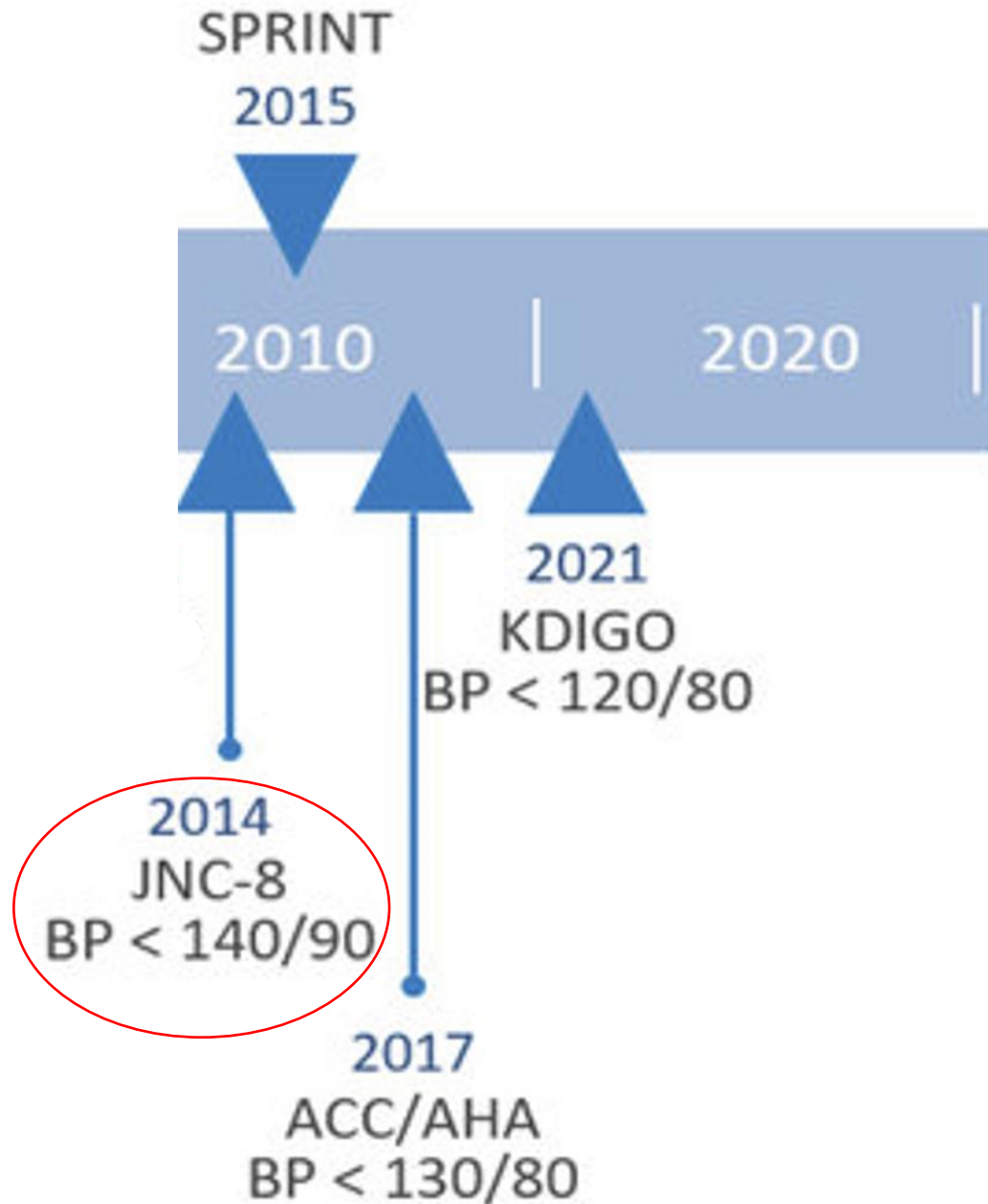
Weight
loss

Life-style
modifications

Alcohol
intake

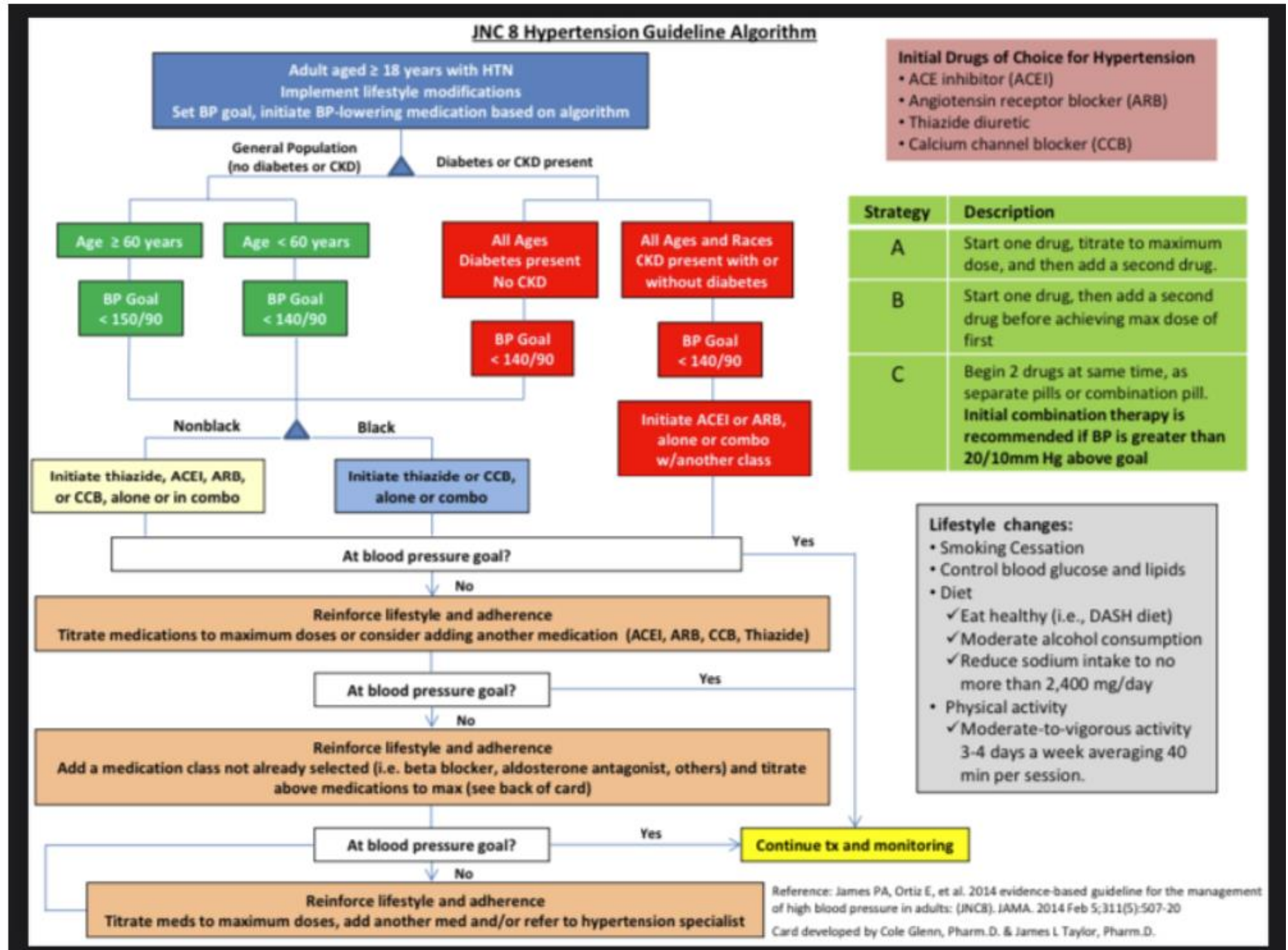
Low
sodium
diet

HTN management trials/guidelines



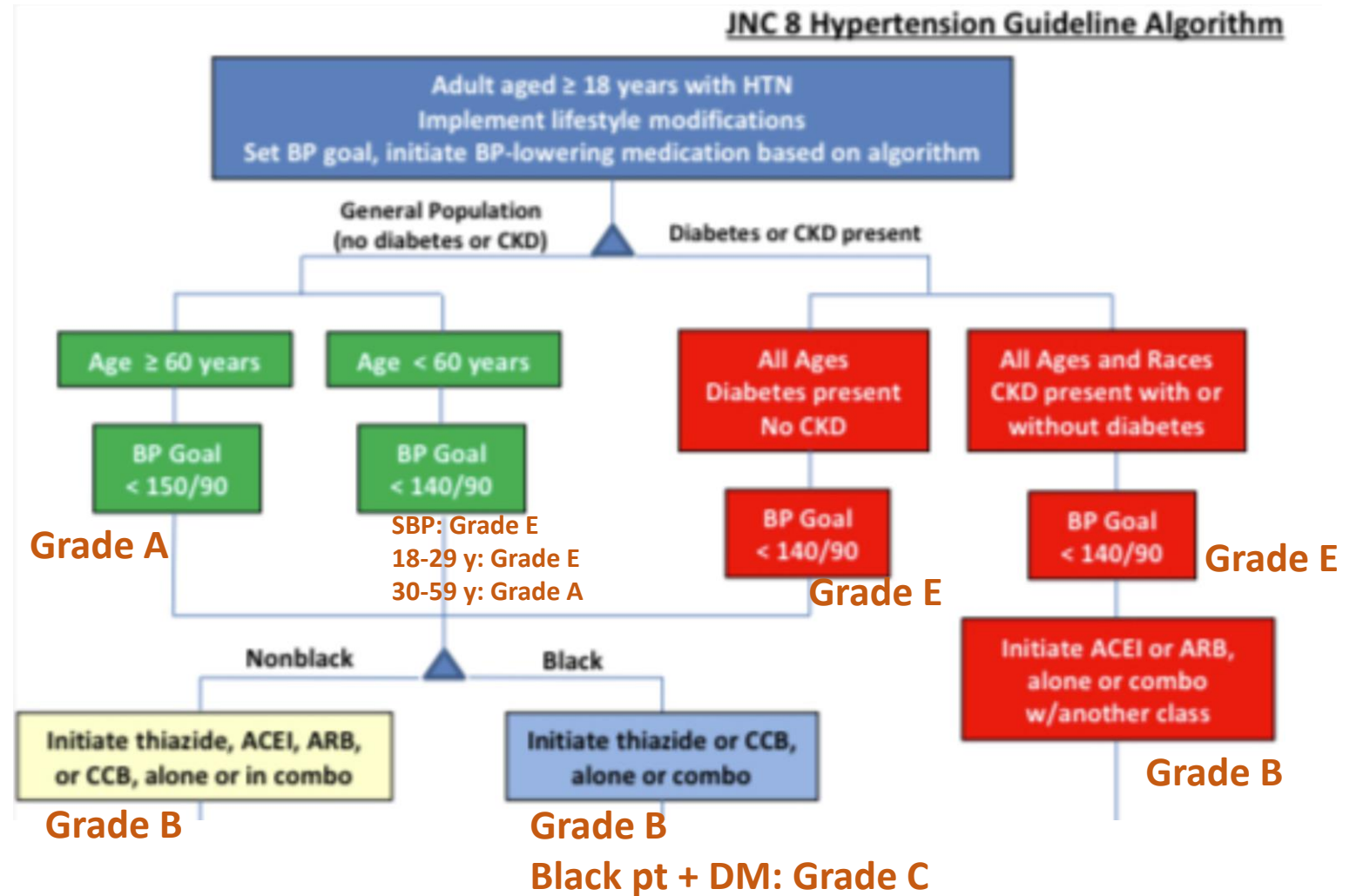
JNC-8 (2014)

Evidence based recommendations for management of high blood pressure in adults

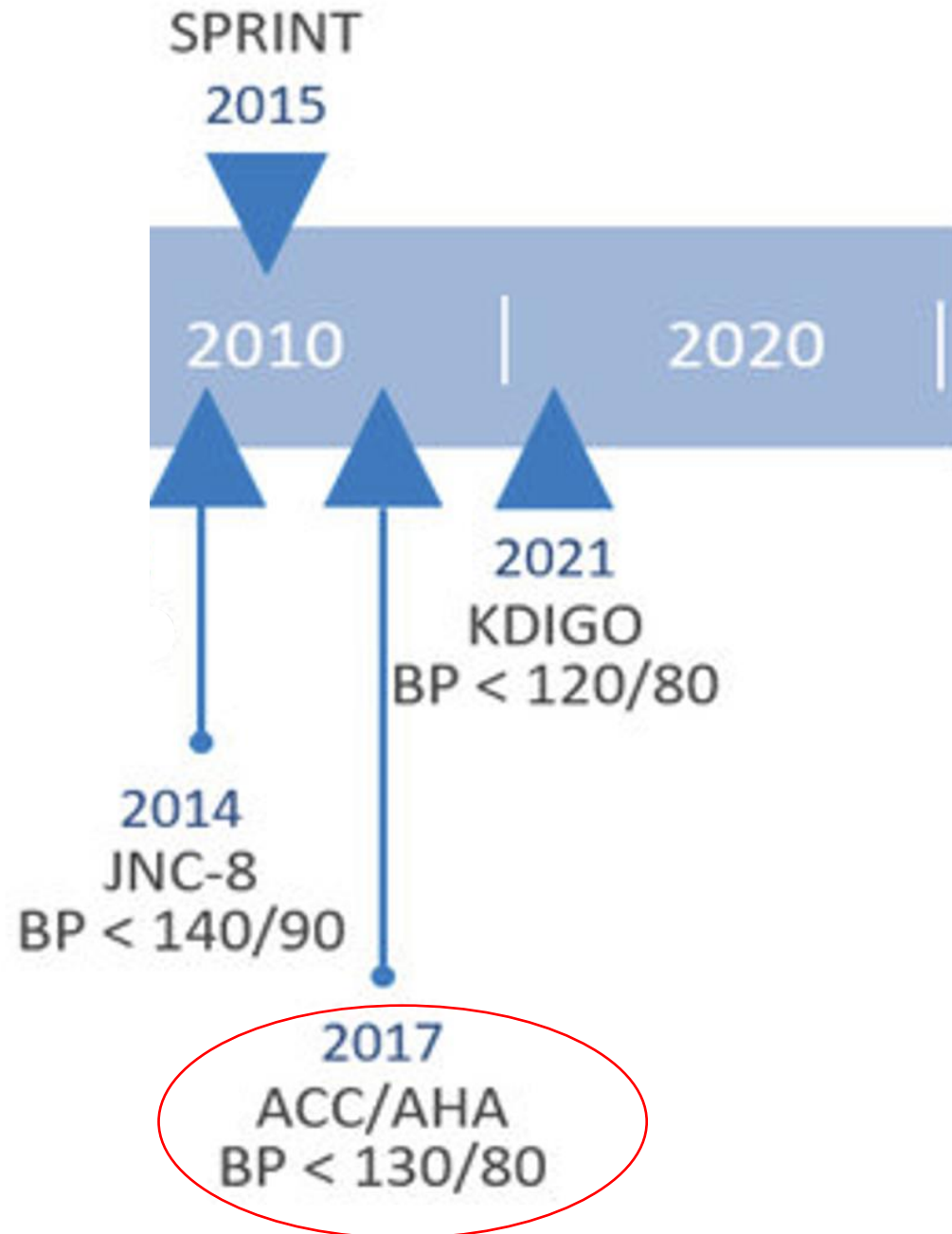


JNC-8 controversies

- Panel members withdrew
- Not accepted by ACC/AHA, NHLBI, ASH



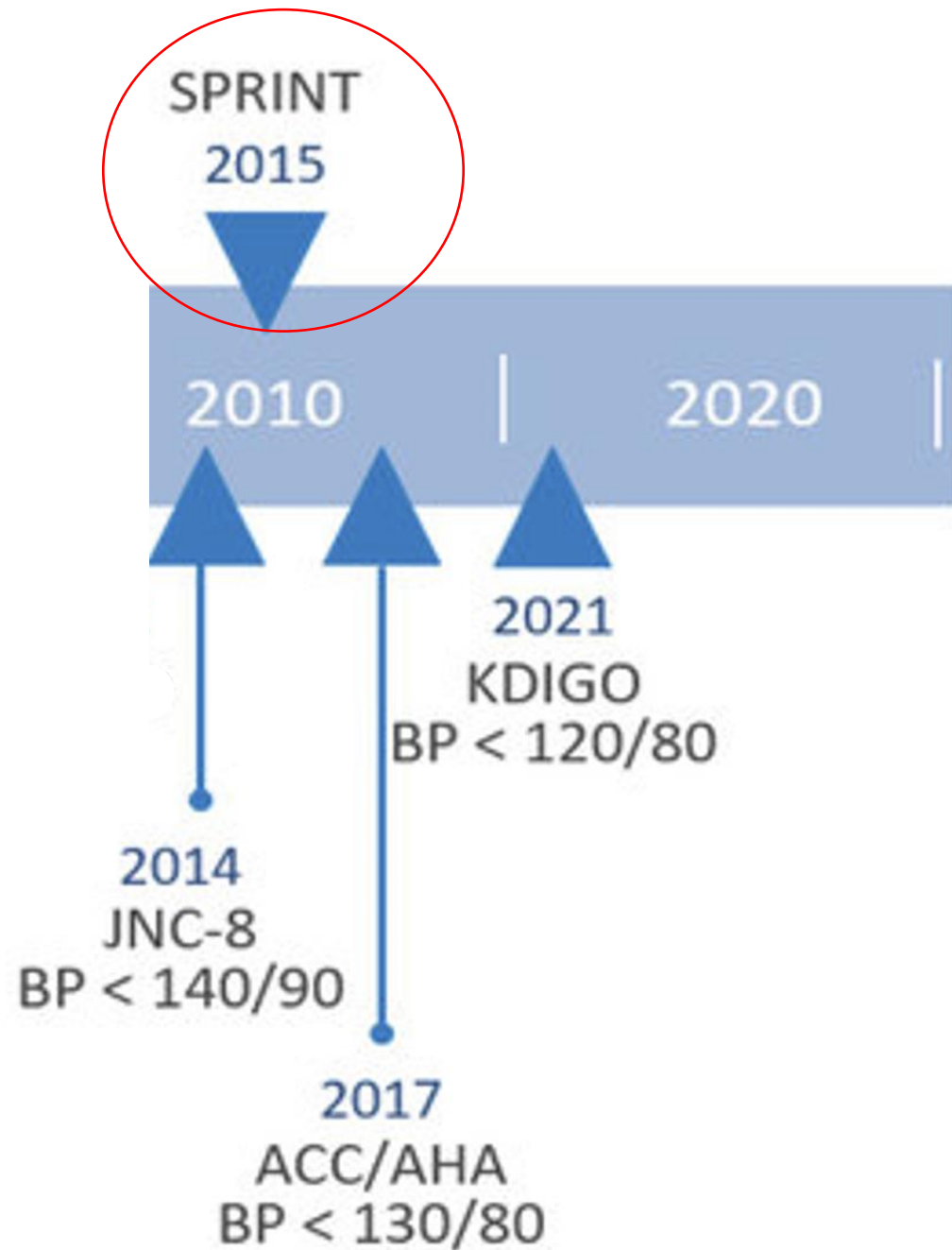
HTN management trials/guidelines



2017 ACC/AHA: <130/80

- SHEP: Systolic hypertension in the elderly program
- HYVET: Hypertension in the very elderly trial
- FEVER: Felodipine event reduction study
- JATOSH: Japanese trial to assess optimum systolic blood pressure in the elderly hypertensive patients
- VALISH: Valsartan in elderly isolated systolic hypertension study
- ACCORD: Action to control cardiovascular risk in diabetes
- SPS-3: Secondary prevention of subcortical strokes study
- **SPRINT: Systolic blood pressure intervention trial**

HTN management trials/guidelines



Systolic Blood Pressure Intervention Trial (SPRINT) - Randomized, open-label, multi-center trial



Age \geq 50 yr, SBP \geq 130mm Hg + one risk factor of heart disease
(CV dz, CKD, 10-yr Framingham CVD risk score \geq 15% or age \geq 75 years)
Notable exclusions: eGFR $<$ 20 mL/min/1.72m², h/o stroke, DM, LVEF $<$ 35%,



N = 4678

Intensive treatment SBP $<$ 120 mmHg

N = 4683

Standard treatment SBP $<$ 140 mmHg



Median follow-up 3.26 years

1.65% per yr **Primary composite outcome of MI, ACS, CVA, HF or death from CV cause** 2.19% per yr

HR = 0.75, 95% CI [0.64-0.89], p $<$ 0.001

1.03% per yr

Death from any cause

1.4% per yr

HR = 0.73, 95% CI [0.6-0.9], p=0.003

0.25% per yr

Death from CV cause

0.43% per yr

HR = 0.57, 95% CI [0.38-0.85], p=0.005

38.3% of patients

Serious adverse events

37.1 % of patients

HR = 1.04, p=0.25

SPRINT - Conclusion

In high-risk, non-diabetic individuals with HTN, including elderly, intensive BP control of SBP <120 vs. <140, reduced CV and all-cause mortality

2017 ACC/AHA guidelines

Office BP vs. out-of-office BP

Optimal target BP

Choice of initial medication

Monotherapy vs. combination therapy

2017 ACC/AHA guidelines

Office BP vs. out-of-office BP

- Out-of-office BP measurements for diagnosis and titration of medications (COR/LOE:1/A)

Optimal target BP

Choice of initial medication

Monotherapy vs. combination therapy

2017 ACC/AHA guidelines

Office BP vs. out-of-office BP

- Out-of-office BP measurements for diagnosis and titration of medications (COR/LOE: Grade 1/A)

Optimal target BP

- HTN + known CVD or 10-yr ASCVD risk $\geq 10\%$: SBP <130 (COR/LOE: 1/B-R), DBP <80 (COR/LOE:1/C-EO)
- HTN & no additional risk factors: SBP <130 (COR/LOE: 2b/B-NR), DBP <80 (COR/LOE: 2b/C-EO)

Choice of initial medication

Monotherapy vs. combination therapy



2017 ACC/AHA guidelines

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- HTN & no additional risk factors: SBP <130 (COR/LOE: 2b/B-NR), DBP <80 (COR/LOE: 2b/C-EO)

Choice of initial medication

- Initiation first line agents: Thiazide diuretics, ACEi/ARB, CCB (COR/LOE: 1/A)

Monotherapy vs. combination therapy

2017 ACC/AHA guidelines

Office BP vs. out-of-office BP

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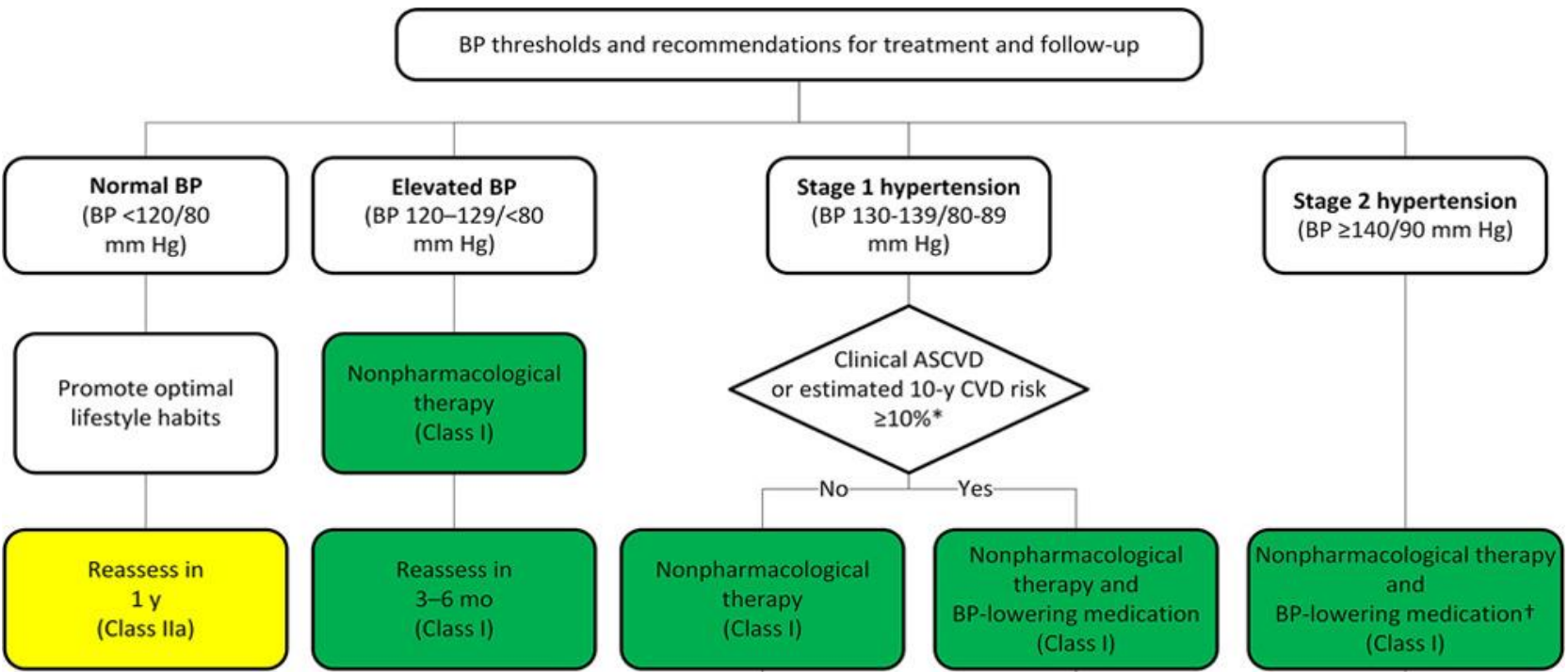
Choice of initial medication

- Initiation first line agents: Thiazide diuretics, ACEi/ARB, CCB (COR/LOE: 1/A)

Monotherapy vs. combination therapy

- Co-morbid conditions
- 2 drugs separate or combination for stage 2 HTN & avg BP >20/10mmHg above target (COR/LOE: 1/C-EO)
- Stage I HTN, monotherapy acceptable to keep <130/80(COR/LOE:2a/C-EO)

2017 ACC/AHA guidelines



Blood Pressure Categories



BLOOD PRESSURE CATEGORY	SYSTOLIC mm Hg (upper number)		DIASTOLIC mm Hg (lower number)
NORMAL	LESS THAN 120	and	LESS THAN 80
ELEVATED	120 – 129	and	LESS THAN 80
HIGH BLOOD PRESSURE (HYPERTENSION) STAGE 1	130 – 139	or	80 – 89
HIGH BLOOD PRESSURE (HYPERTENSION) STAGE 2	140 OR HIGHER	or	90 OR HIGHER
HYPERTENSIVE CRISIS (consult your doctor immediately)	HIGHER THAN 180	and/or	HIGHER THAN 120



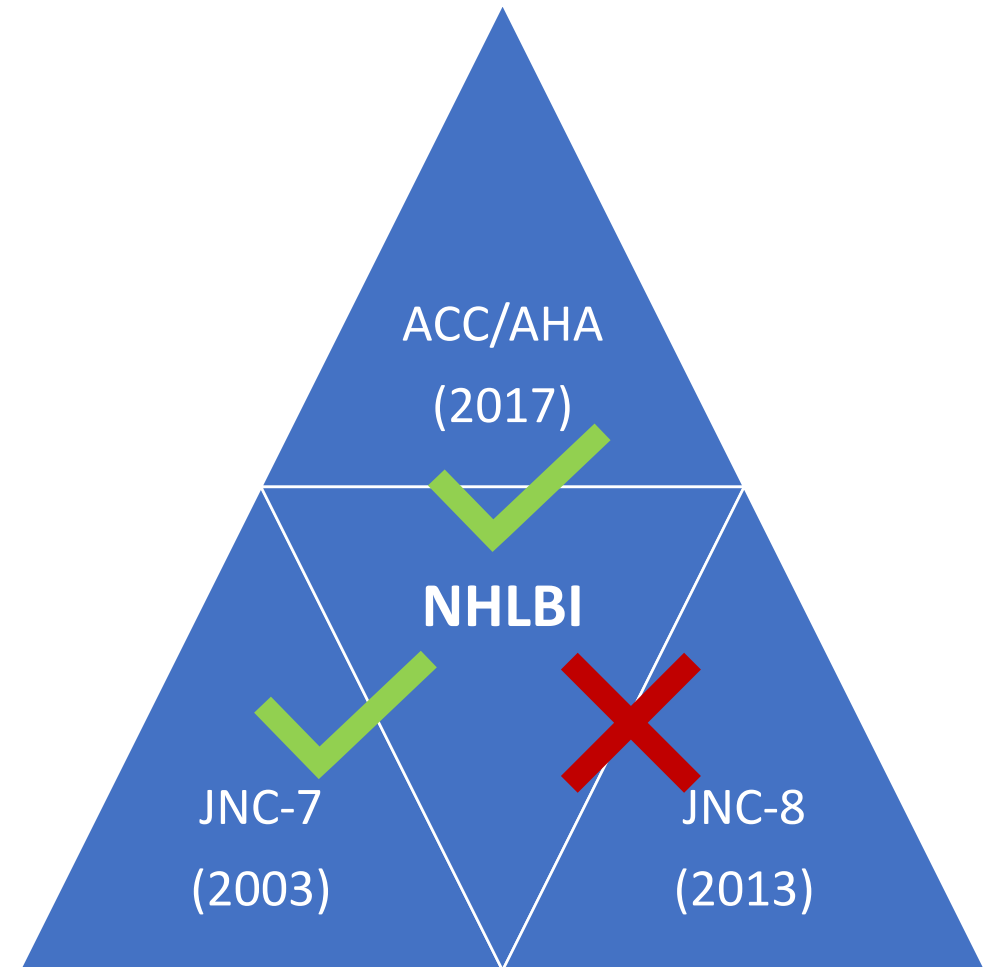
2017 ACP/AAFP

Concern for:

- Conflicts of interest
- Deviation from IOM recommended practices for developing trustworthy guidelines

ACP/AAFP

- Primary care groups declined to participate in ACC/AHA guideline panel
- Did not endorse 2017 ACC/AHA HTN guidelines
- 2017 ACP/AAFP HTN guidelines for older adults with hypertension



2017 ACC/ACP older adults (≥ 60 yrs)

Systematic review of RCT for primary outcome: all-cause mortality, morbidity and mortality related to stroke, major cardiac events (fatal & non-fatal MI, SCD)

Observational data for harms

Treatment of higher (<150 mmHg) vs. lower (≤ 140 mmHg) SBP targets for adults aged ≥ 60 yrs with HTN

2017 ACC/ACP older adults (≥ 60 yrs)

1

Initiate treatment to target SBP <150 , to reduce risk of stroke, cardiac events and possibly death (strong recommendation)

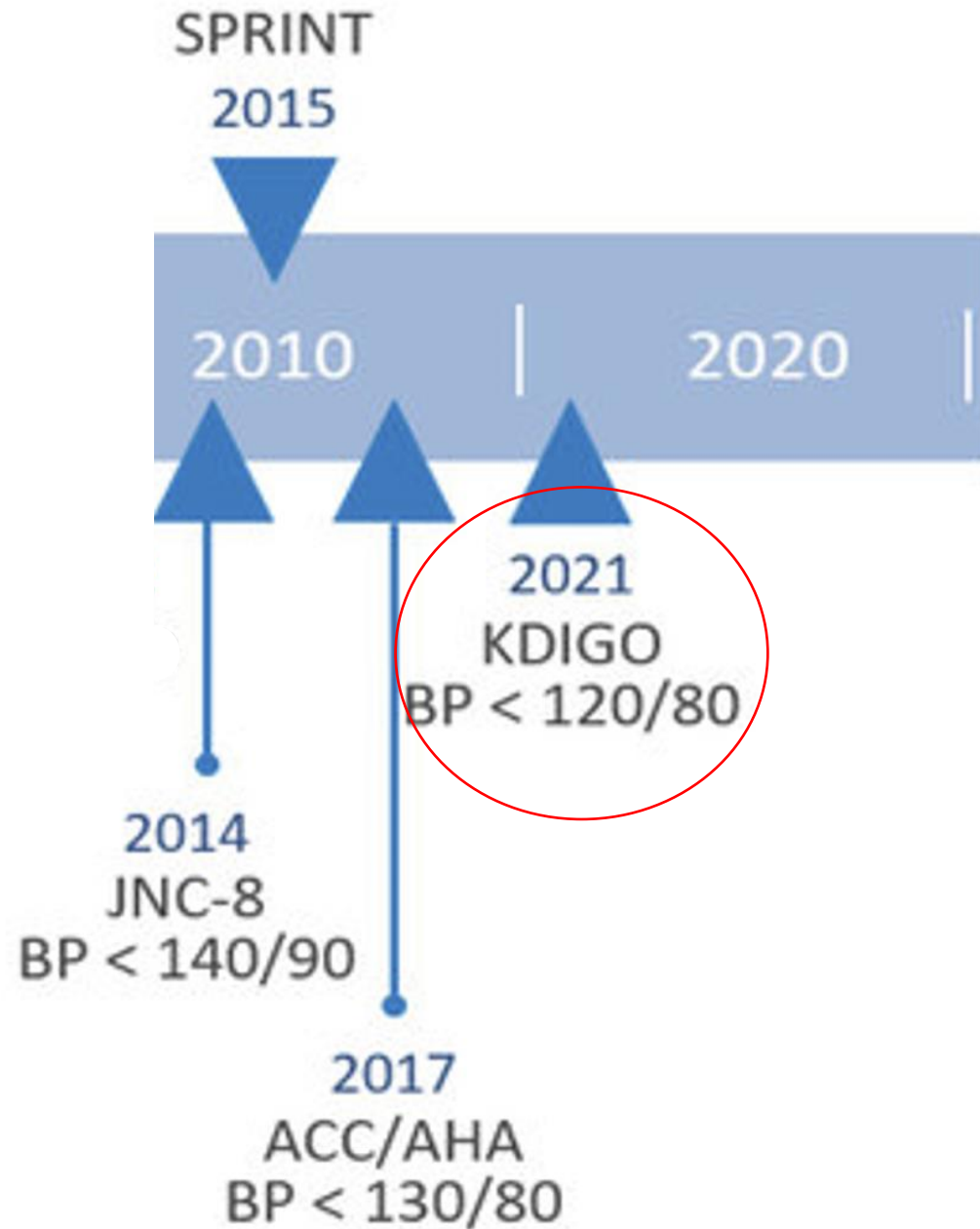
2

Initiate or intensify treatment to target SBP <140 in those with h/o stroke or TIA to reduce recurrence risk (weak recommendation)

3

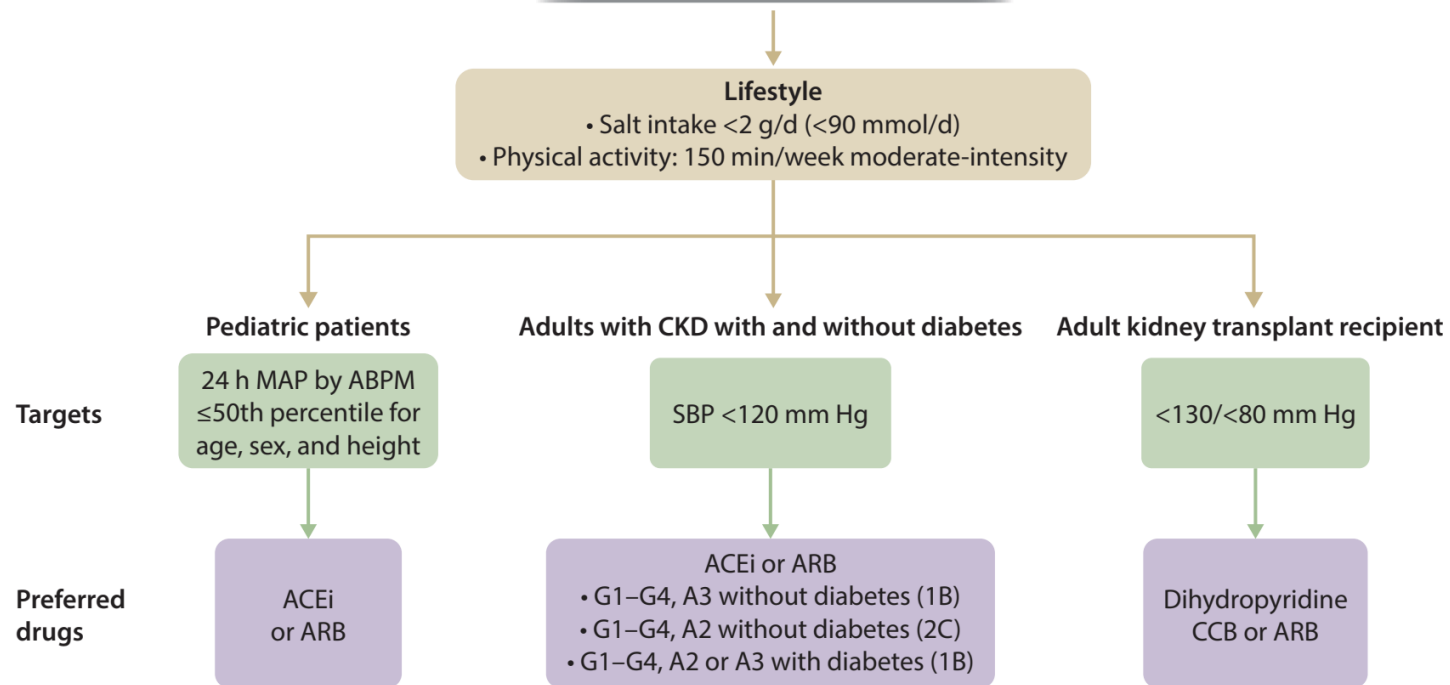
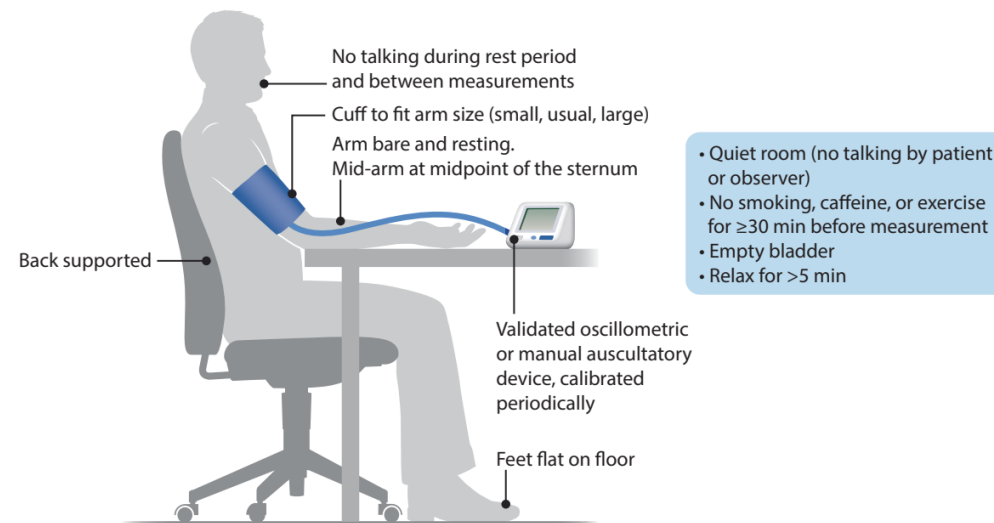
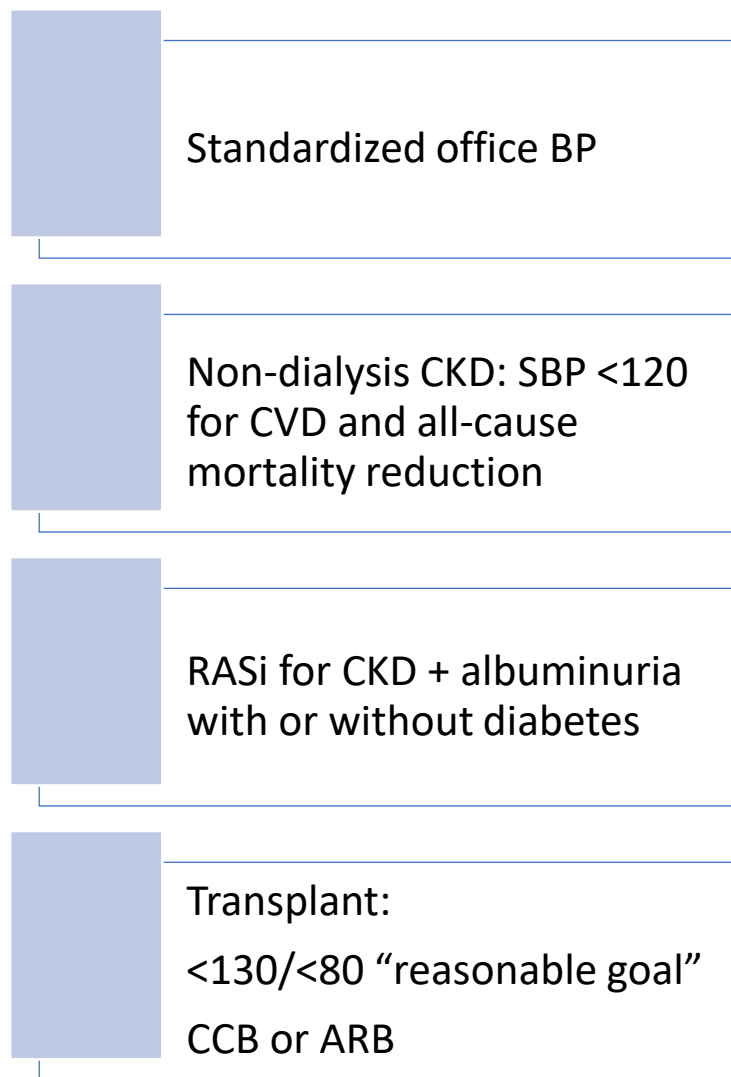
Initiate or intensify treatment to target SBP <140 those with high cardiovascular risk based on individual assessment (weak recommendation)

HTN management trials/guidelines

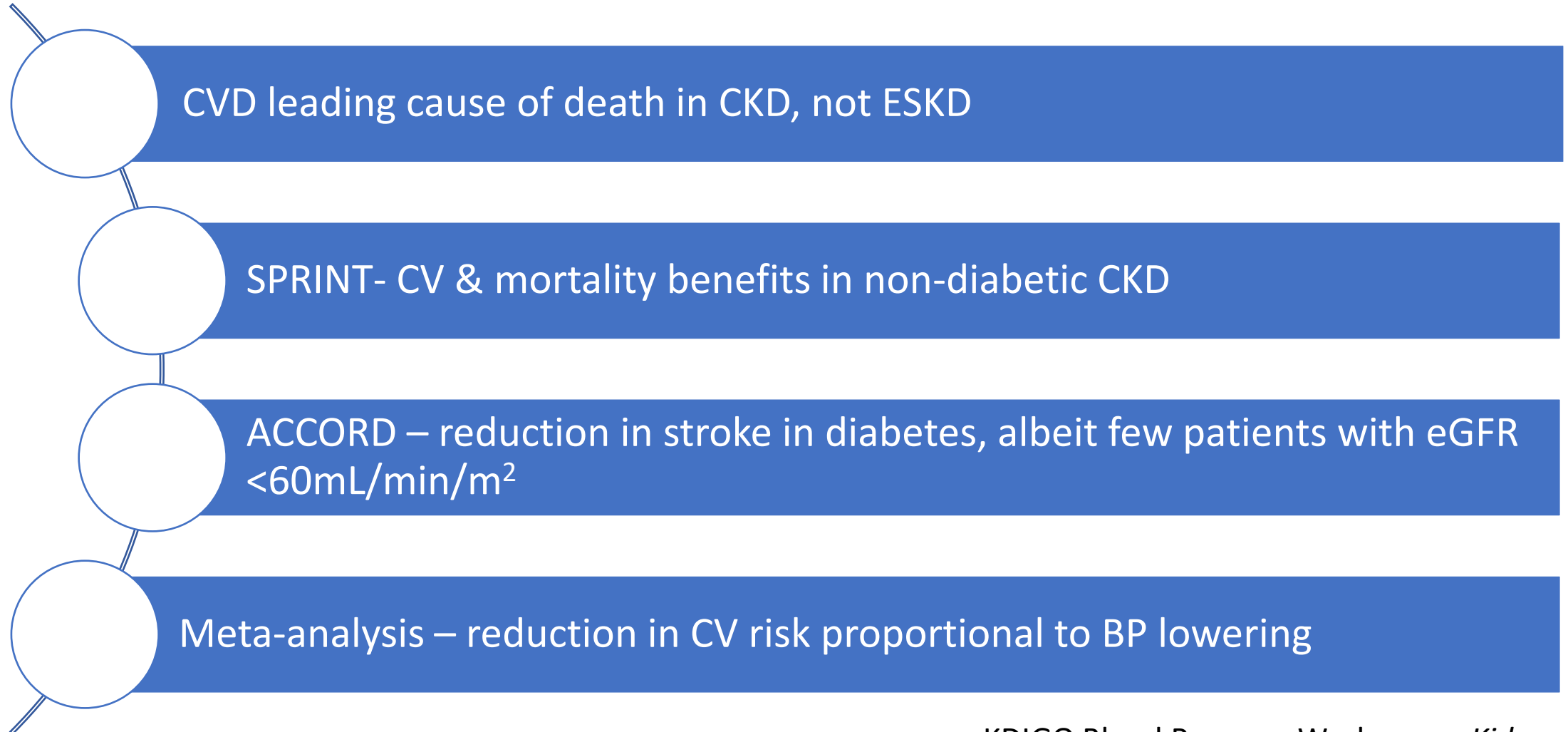


2021 KDIGO

Central Illustration for KDIGO 2021 Guideline for the Management of Blood Pressure in Patients with CKD, not receiving dialysis



2021 KDIGO – rationale for SBP <120mmHg



Intensive BP Lowering Reduces GFR But Does Not Increase Injury Biomarkers

978 of 2,646
Participants
With CKD



Mean age
72 years






INTERVENTION



Intensive BP Lowering
(SBP < 120 mm Hg)
Baseline mean eGFR 44.8
N = 519

Standard BP Lowering
(SBP < 140 mm Hg)
Baseline mean eGFR 45.2
N = 459

1-YEAR OUTCOMES

Marker	Ratio of Intensive to Standard BP Arm	P
 eGFR	↓ 7%	< 0.001
 ACR	↓ 32%	< 0.001
 B2M	↓ 29%	0.005
 A1M	↓ 24%	0.002
 YKL-40, IL-18, UMOD, MCP-1, KIM-1, NGAL	<i>No Difference Between Groups</i>	

ARTICLE: Effects of Intensive BP Lowering on Kidney Tubule Injury in CKD: A Longitudinal Subgroup Analysis in SPRINT

Rakesh Malhotra, Timothy Craven, Walter T. Ambrosius, et al

Am J Kidney Dis (ePub Oct 2, 2018) | DOI: 10.1053/j.ajkd.2018.07.015 | © National Kidney Foundation

Intensive BP Lowering Reduces GFR But Does Not Increase Injury Biomarkers

529 of 4,733
 Participants



- Age 62 ± 6.5 years
- Type 2 Diabetes

INTERVENTION



Intensive BP Lowering
 (SBP < 120 mm Hg)
 Baseline mean eGFR 85.9
 N = 260

Standard BP Lowering
 (SBP < 140 mm Hg)
 Baseline mean eGFR 85.4
 N = 269

2-YEAR OUTCOMES

Marker	Ratio of Intensive to Standard BP Arm	P
eGFR	↓ 11%	< 0.001
ACR	↓ 30%	0.004
IL-18	↓ 14%	0.04
KIM-1	↓ 12%	NS
MCP-1	↓ 7%	NS
YKL-40	↓ 15%	NS

Effect of Intensive Blood Pressure Lowering on Kidney Tubule Injury: Findings From the ACCORD Trial Study Participants

Girish N. Nadkarni, Kinsuk Chauhan, Veena Rao, et al

Am J Kidney Dis (ePub Oct 2, 2018) | DOI: 10.1053/j.ajkd.2018.07.016 | © National Kidney Foundation

2022 AAFP

1

All adults, BP target <140/90mmHg to reduce the risks of all-cause and cardiovascular **mortality** (strong recommendation)

2

Consider a BP target <135/85 mmHg to reduce the risk of myocardial infarction (weak recommendation)



Potential harms: cost of medications, drug-drug interactions, adverse events (falls, syncope, “AKI”)

Guidelines	BP target
2014 JNC-8	<150/90, <140/90
2017 ACC/AHA	<130/80
2017 ACP/AAFP (≥ 60 years)	<150, ≤ 140
2021 KDIGO for CKD	<120/80, <130/80 (transplant)
2022 AAFP (all adults)	<140/90, <135/85

Study objectives

The STEP study was conducted to provide new evidence on the benefits of BP lowering in older patients with hypertension. Specifically, it examined whether intensive treatment targeting a SBP below 130 mmHg could reduce the risk of cardiovascular disease compared with a SBP target below 150 mmHg.

Where?

China



42 clinical sites

Follow-up

BP measuring (months)



Home BP

Data centre



Primary endpoint

Composite of stroke, acute coronary syndrome, acute decompensated heart failure, coronary revascularisation, atrial fibrillation, or death from CV causes



Relative risk reduction: 26%

HR: 0.74 (95% CI: 0.60-0.92)

Who and what?

Patients (60-80 years old)

SBP 140-190 mmHg during 3 screening visits or taking antihypertensive medication. No prior stroke



Median follow-up 3.34 years

Results

Average decrease in SBP from baseline (mmHg)

19.4 10.1

Average SBP reached (mmHg)

126.7 135.9

Secondary endpoints

33% ↓ relative risk of stroke
95% CI: 0.47-0.97

33% ↓ relative risk of acute coronary syndrome
95% CI: 0.47-0.94

Safety outcomes and renal outcomes

No difference between

Except hypotension Rate (%)



STEP Study - 2021

STEP study

#ESCCongress

Intensive vs. standard blood pressure control among older hypertensive patients

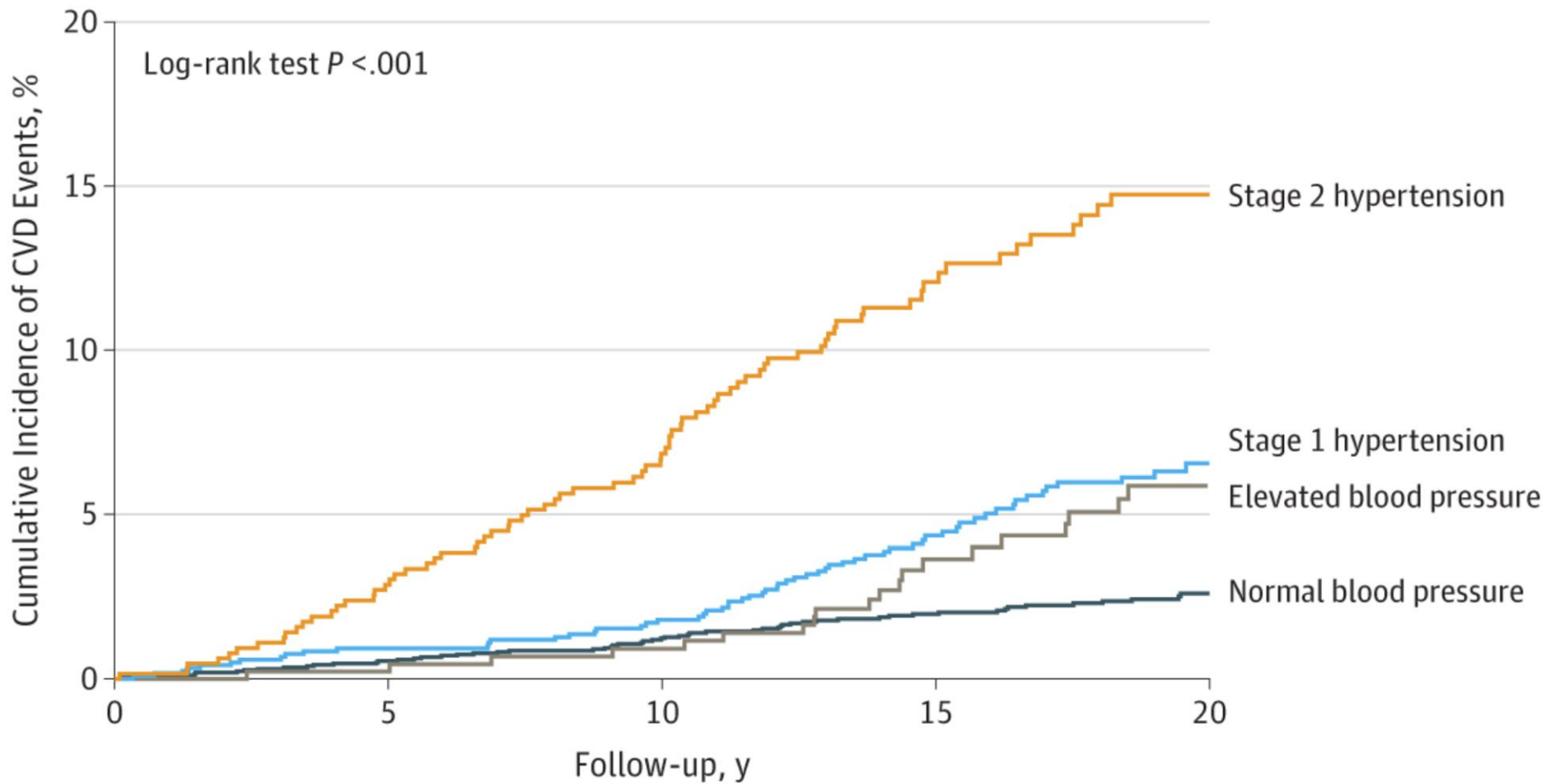
Conclusion



Active control of systolic blood pressure (SBP) to below 130 mmHg in older hypertensive patients, as compared with below 150 mmHg, resulted in a lower incidence of major CV events, with no increase in renal injuries.

Young adults

- Hypertension without elevated 10- yr ASCVD risk
 - Understudied, lower-risk group will require very long follow-up to see clinically significant results
 - No trial evidence supporting decrease in CVD morbidity & mortality
- Observational data: high life-time risk, benefit of BP control early in life
- In the absence of clear guidelines for younger patients with low 10-yr risk
 - Focus on life-time risk
 - Prevention of progression of HTN



No. at risk

Blood pressure

Normal	2574	2522	2351	1824	813
Elevated	445	437	409	286	99
Hypertension					
Stage 1	1194	1158	1078	732	284
Stage 2	638	603	519	314	105

Young adults

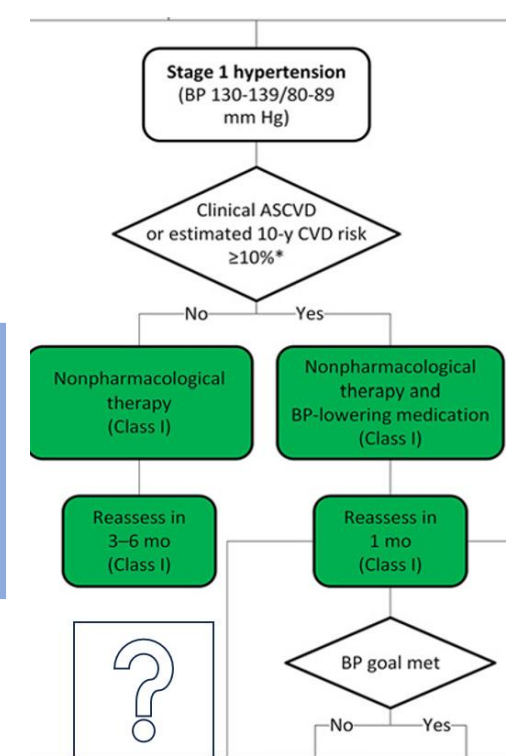
2017 ACC/AHA

Adults with an **elevated BP or stage 1 hypertension** who have an **estimated 10-year ASCVD risk less than 10%** should be managed with nonpharmacological therapy and have a repeat BP evaluation within 3 to 6 months

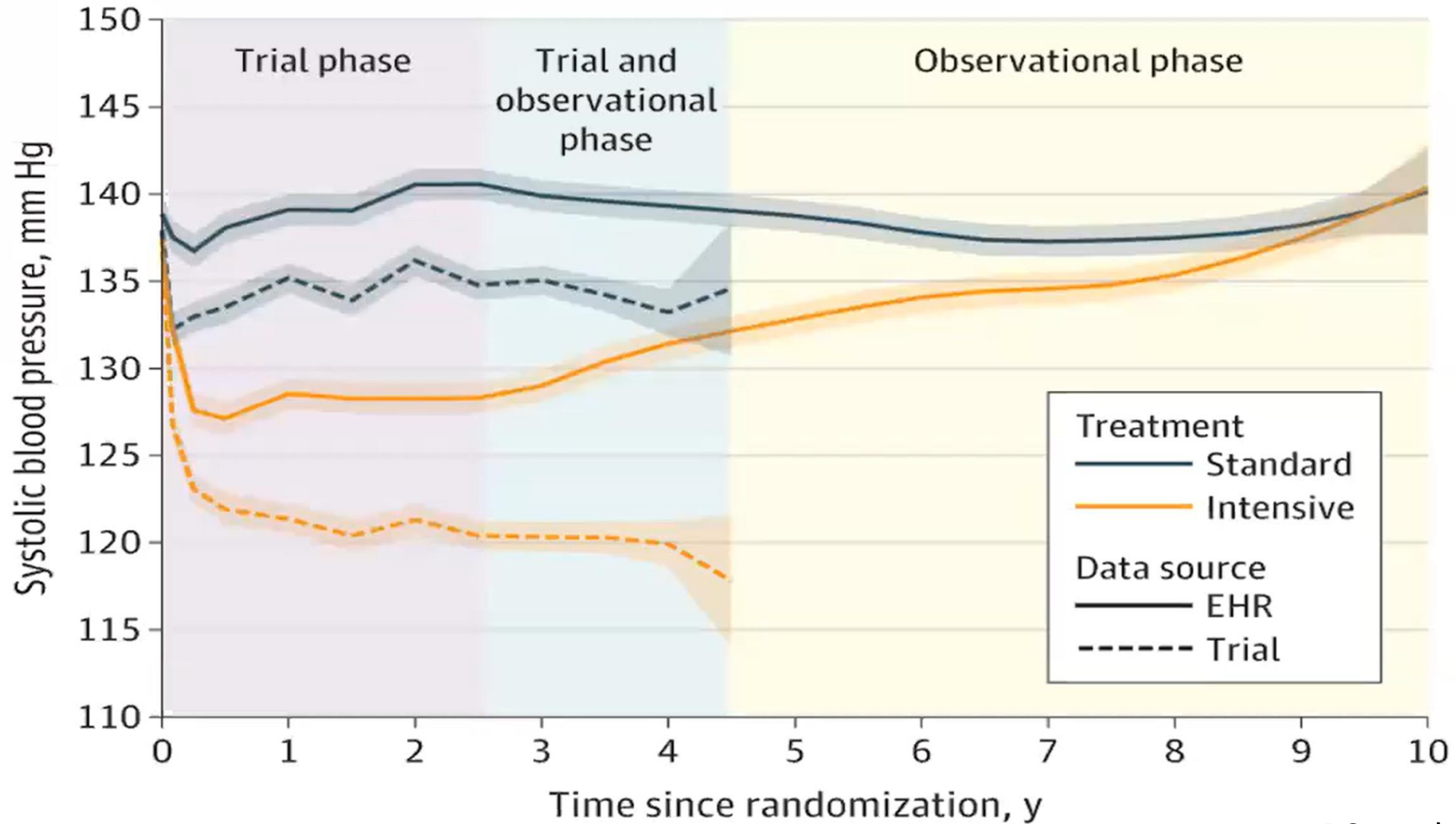


2021 AHA Scientific statement

Medications if BP >130/80 mmHg after 6 months of lifestyle therapy
System changes are necessary



5 years post-SPRINT



Bottom-line

Most adults <130/80
CKD <120/80
(standardized office BP,
out-of-office BP)

Lifestyle modifications
+/- pharmacological
therapy

Elevated BP or stage I:
Single agent (ARB, CCB,
thiazide)
Low dose combination

Stage 2 and/or
escalation: Combination
(ARB/CCB, CCB/thiazide,
ARB/thiazide)

*co-morbid conditions, drug allergies, resistant hypertension, individualized risk vs. benefit if AE, race – social construct

Take home points



Shared decision making

Effective guidelines are followed by patients and practitioners



Guidelines should not replace clinical judgement



Standardized office blood pressure > routine office blood pressure



Combination therapy of anti-HTN to achieve target BP in an individual patient

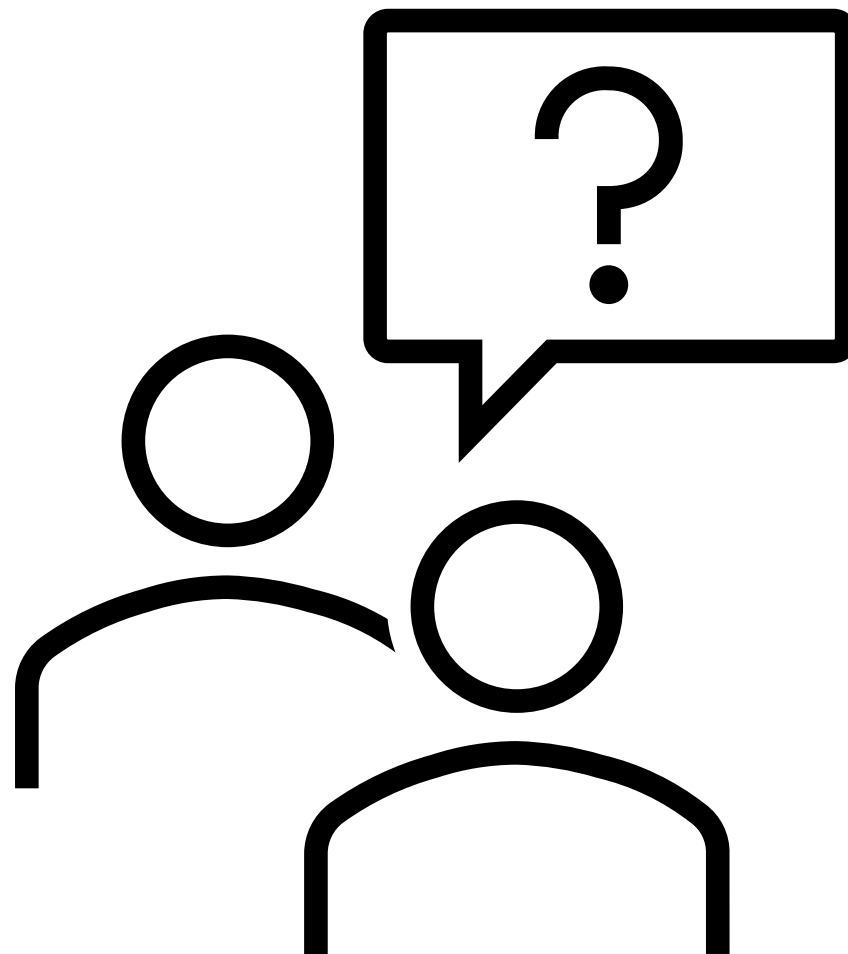


Lifestyle modifications



Overcoming clinical inertia for improved patient outcomes

Questions



Supplementary slides

SPRINT MIND

QUESTION Does intensive blood pressure control compared with standard control reduce the occurrence of dementia?

CONCLUSION This randomized clinical trial of adults with hypertension found that intensive systolic blood pressure (SBP) control (target <120 mm Hg) did not significantly reduce the risk of probable dementia.

POPULATION

6029 Men
3332 Women



Adults aged ≥50 years with hypertension and without diabetes or stroke

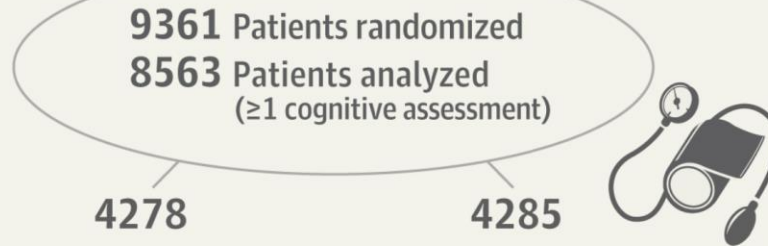
Mean age: 68 years

LOCATIONS

102 US sites (including Puerto Rico)



INTERVENTIONS



Intensive control (Target SBP <120 mm Hg) **Standard control** (Target SBP <140 mm Hg)

Median treatment period, 3.3 years

PRIMARY OUTCOME

Occurrence of adjudicated probable dementia

SECONDARY OUTCOMES

Adjudicated mild cognitive impairment (MCI)
Composite outcome of MCI or probable dementia

FINDINGS

PRIMARY OUTCOME: Adjudicated probable dementia

Intensive control

149 patients
(7.2 cases/1000 person-years)

Standard control

176 patients
(8.6 cases/1000 person-years)

Hazard ratio: **0.83** (95% CI, 0.67-1.04)

SECONDARY OUTCOME: Adjudicated MCI

Intensive control

287 patients
(14.6 cases/1000 person-years)

Standard control

353 patients
(18.3 cases/1000 person-years)

Hazard ratio: **0.81** (95% CI, 0.69-0.95)

SECONDARY OUTCOME: Composite outcome

Intensive control

402 patients
(20.2 cases/1000 person-years)

Standard control

469 patients
(24.1 cases/1000 person-years)

Hazard ratio: **0.85** (95% CI, 0.74-0.97)

Heart failure events in a clinical trial on arterial hypertension: new insights into the SPRINT trial

SPRINT trial population



post-hoc analysis

9361 participants

Intervention



intensive
systolic blood pressure
lowering < 120 mmHg

randomization 1:1



standard
systolic blood pressure
lowering < 140 mmHg

Outcome

1) composite clinical endpoint: myocardial infarction, stroke, acute coronary syndrome, cardiovascular death = primary SPRINT outcome excluding heart failure events

2) cardiovascular deaths (excluding heart failure)

3) all-cause deaths (excluding heart failure)

Results

Composite clinical endpoint

4.4%

5.5%

Cardiovascular death (excluding heart failure)

0.4%

1.2%

All-cause death (excluding heart failure)

3.1%

4.3%

Hazard ratio:

0.79, 95% CI 0.66–0.95

0.52, 95% CI 0.33–0.81

0.73, 95% CI 0.59–0.90

■ intensive SBP lowering <120 mm Hg ■ standard SBP lowering <140 mm Hg

Conclusions

Intensive lowering of systolic blood pressure (to <120 mm Hg) using the SPRINT methodology is associated with a 21% reduced risk ($p = 0.012$) of a composite endpoint (myocardial infarction, acute coronary syndrome other than myocardial infarction, stroke, and cardiovascular death) compared with the standard treatment (lowering the target SBP to <140 mm Hg)

2010

ACCORD BP TRIAL



Effects of intensive blood-pressure control in type 2 diabetes mellitus

Multicenter, randomized, controlled, open-label trial



Objective: In patients with T2DM at high risk for CV events, does intensive BP control (SBP <120 mmHg) reduce rates of nonfatal MI, nonfatal stroke, or CV mortality when compared to standard BP control (SBP <140 mmHg)?

4,733
patients

Inclusion criteria: Patients with type 2 diabetes mellitus, hemoglobin A1C $\geq 7.5\%$ and either active CVD or at risk for CVD.



Intensive BP control
(n=2,362)

VS

Conventional BP control
(n=2,371)



PRIMARY OUTCOME

1.87

Nonfatal MI, nonfatal stroke, or CV mortality %/yr
HR 0.88; 95% CI 0.73-1.06; P=0.20

2.09

SECONDARY OUTCOME

1.13

Nonfatal MI %/yr
HR 0.87; 95% CI 0.68-1.10; P=0.25

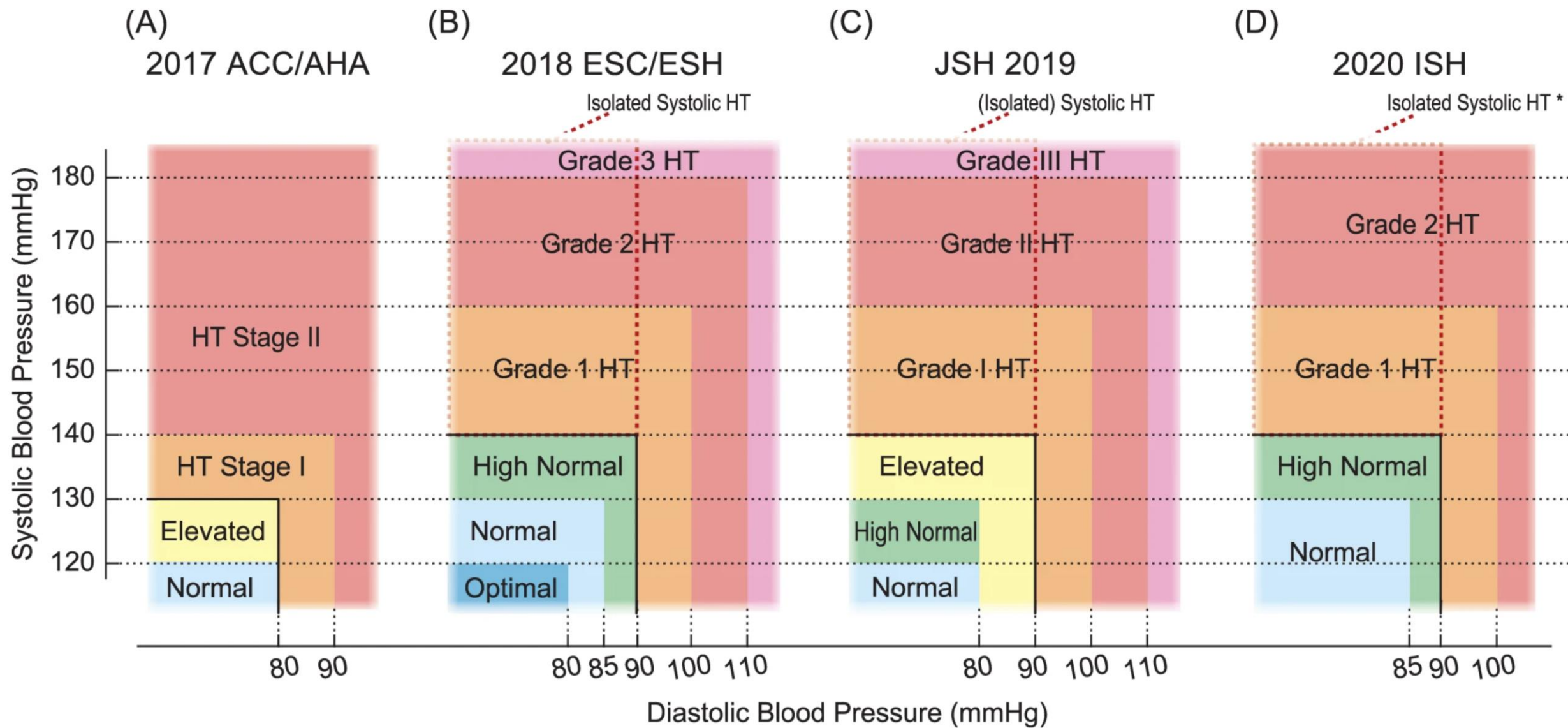
1.28

0.32

Any stroke %/yr
HR 0.59; 95% CI 0.39-0.89; P=0.01; NNT 476/yr

0.53

Conclusion: In patients with type 2 diabetes at high risk for CV events, targeting a systolic blood pressure of less than 120 mm Hg, as compared with less than 140 mm Hg, did not reduce the rate of a composite outcome of fatal and nonfatal major cardiovascular events.



STOP-ACEi

Does the discontinuation of RAS inhibitors improve eGFR in patients with advanced CKD?



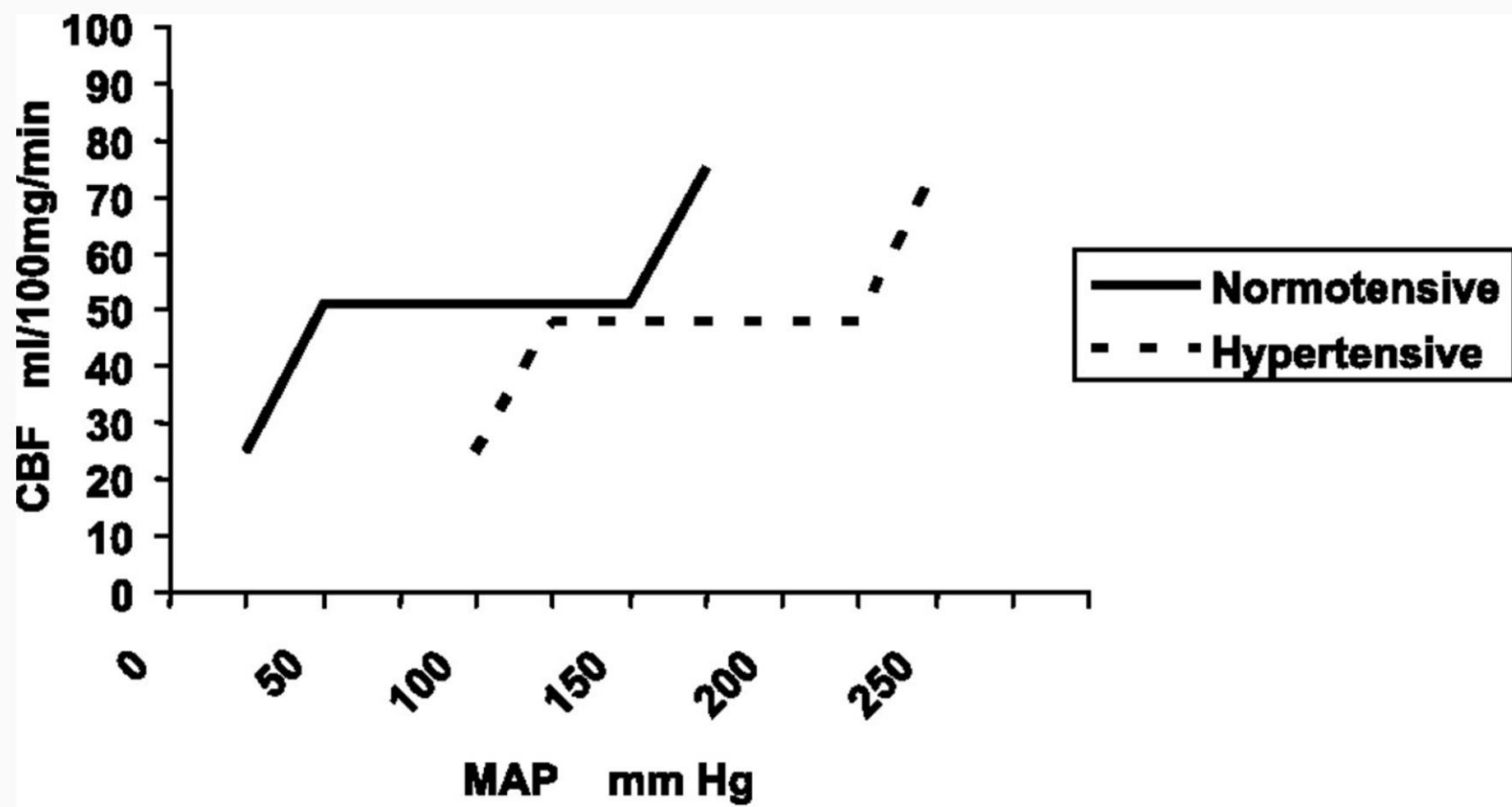
		1° Outcome eGFR (by MDRD*)	2° Outcome (ESKD or RRT)	MACE
Open-Label Randomized Control Trial 39 Centers United Kingdom 411 Adults Stage 4 or 5 CKD (eGFR < 30 mL/min/1.73m ²) > 2 mL/min/1.73m ² per year eGFR decline over 2 year RAS inhibitor > 6 months (ACEi or ARB)	Continue RAS inhibitor n=205	 13.3±0.6 mL/min/1.73m ²	 56% (115/205)	 43% (88/205)
	3 years	P = 0.42 (-2.5 to 1.0)	HR = 1.28 (0.99 to 1.65)	Similar
	Discontinue RAS inhibitor n=206	12.6±0.7 mL/min/1.73m ²	62% (128/206)	52% (108/206)

Conclusion: Among patients with advanced and progressive chronic kidney disease, the discontinuation of RAS inhibitors was not associated with a significant between-group difference in the long-term rate of eGFR decline.

*Modification of Diet in Renal Disease

Reference: STOP ACEi trial investigators, Renin-Angiotensin System Inhibition in Advanced Chronic Kidney Disease.

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CBF=cerebral blood flow; MAP=mean arterial pressure