



2016

Hypertension Canada CHEP Guidelines for the Management of Hypertension

What's new in the treatment of hypertension?
What's still really important?



Presenter Disclosure

- Relationships with commercial interests:
 - Grants/Research Support:
 - Speakers Bureau/Honoraria:
 - Consulting Fees:
 - Data Safety and Monitoring:



Mitigating Potential Bias

- The information presented is based on recent information that is explicitly “evidence-based”.
- This presentation and all the recommendations involving clinical medicine are based on evidence that was vetted by the Canadian Hypertension Education Program. The presentation has been developed for dissemination by Hypertension Canada.



Hypertension Canada

Mission:

- Advancing health through the prevention and control of high blood pressure and its complications.

Vision:

- Canadians will have the healthiest blood pressure in the world.



Evidence-based Annual Guidelines

- Canada has the world's highest reported national blood pressure control rates
- CHEP is known as the most credible source for evidence-based hypertension guidelines, with annual updates, a well-validated review process and effective dissemination techniques across Canada



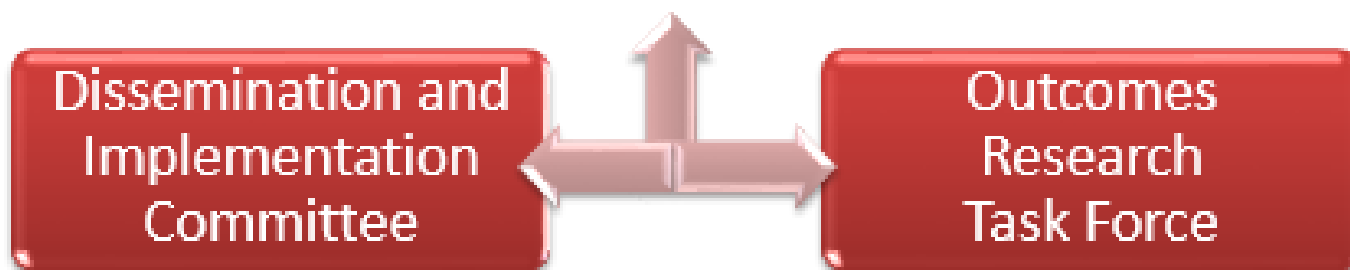
2016 CHEP Guidelines Task Force





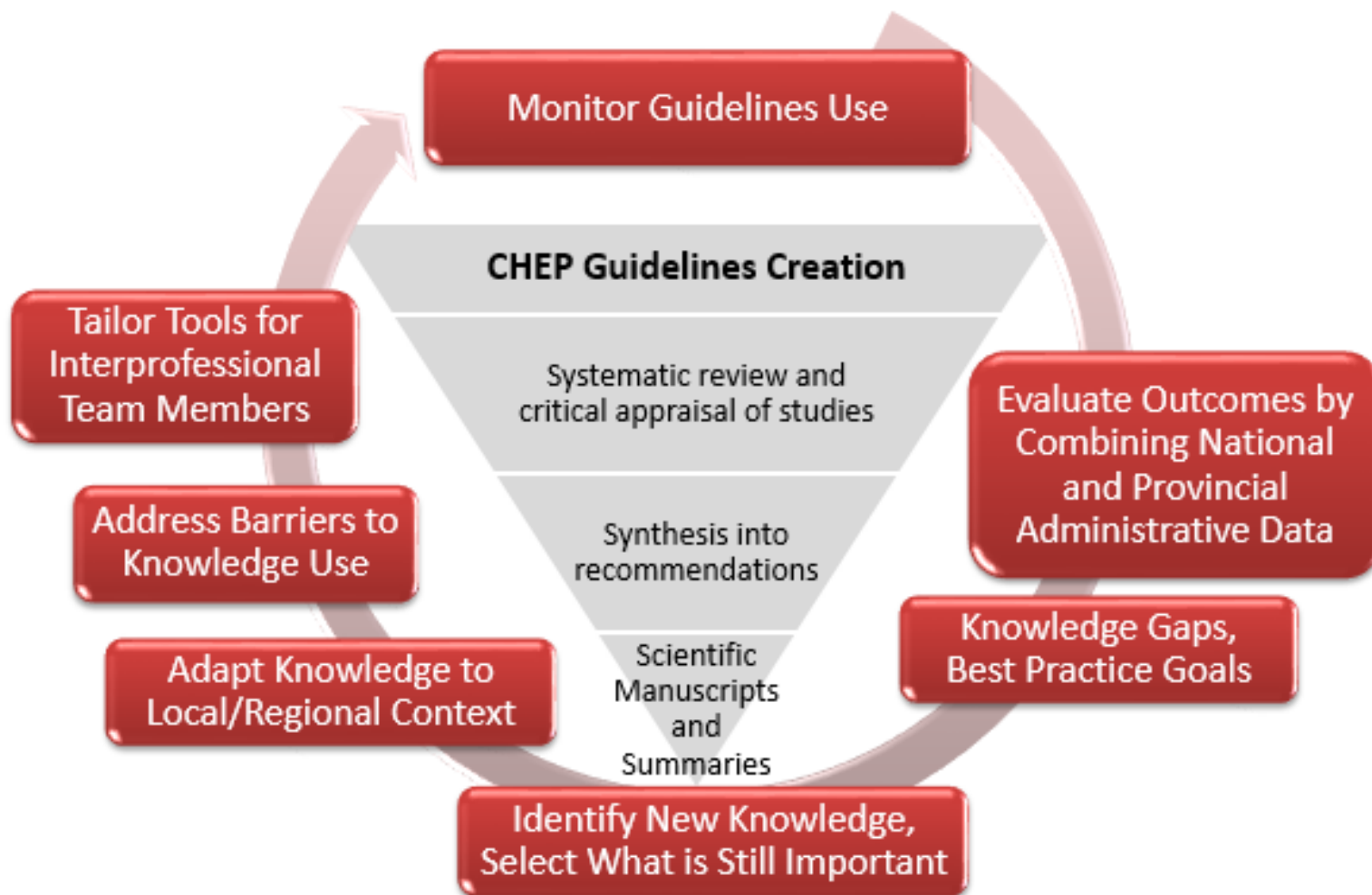
CHEP Guidelines Organizational Chart

Guidelines Task Force





Hypertension Canada's Annual *KT* Cycle for Developing Management Guidelines





CHEP 2016 Guidelines

What's new?

- **New thresholds and targets for high risk patients (SPRINT)**
- ***Assessing*** clinic blood pressures using **automatic electronic** (oscillometric) monitors
- ***Adopting*** healthy behaviours is integral to the management of hypertension (focus on potassium supplementation)
- ***Updating*** the evaluation of patients with suspected secondary forms of hypertension (focus on primary hyperaldosteronism)
- ***Updating*** the treatment of patients with hypertension with concurrent coronary artery disease
- ***New*** recommendations on the diagnosis and management of hypertension in pediatric patients (*NOT the focus of this presentation*)



CHEP 2016 Guidelines

What's still important?

- The diagnosis of hypertension should be based on **out-of-office** measurements
- The management of hypertension is all about global cardiovascular risk management and vascular protection
- The most important step in prescription of antihypertensive therapy is achieving patient “buy-in” and adherence



Usual Office BP Threshold Values for Initiation of Pharmacological Treatment

Population	SBP	DBP
High Risk (SPRINT population)	≥ 130	<u>NA</u>
Diabetes	≥ 130	≥ 80
Moderate-to-high risk (TOD or CV risk factors)*	≥ 140	≥ 90
Low risk (no TOD or CV risk factors)	≥ 160	≥ 100

TOD = target organ damage

*AOBP threshold $\geq 135/85$



Recommended Office BP Treatment Targets

Treatment consists of health behaviour \pm pharmacological management

Population	SBP	DBP
High Risk	≤ 120	NA
Diabetes	< 130	< 80
All others*	< 140	< 90

* Target BP with AOBP $< 135/85$



New thresholds/targets for the high risk patient post-SPRINT: *who does this apply to??*

- Clinical or sub-clinical cardiovascular disease
OR
- Chronic kidney disease (non-diabetic nephropathy, proteinuria <1 g/d, *estimated glomerular filtration rate 20-59 mL/min/1.73m²)
OR
- †Estimated 10-year global cardiovascular risk ≥15%
OR
- Age ≥ 75 years

Patients with one or more clinical indications should consent to intensive management.

* Four variable MDRD equation

† Framingham Risk Score, D'Agastino, Circulation 2008



New thresholds/targets for the high risk patient post-SPRINT: *who does this NOT apply to??*

Limited or No Evidence:

- Heart failure (EF <35%) or recent MI (within last 3 months)
- Indication for, but not currently receiving a beta-blocker
- Frail or institutionalized elderly

Inconclusive Evidence:

- Diabetes mellitus
- Prior stroke
- eGFR < 20 ml/min/1.73m²

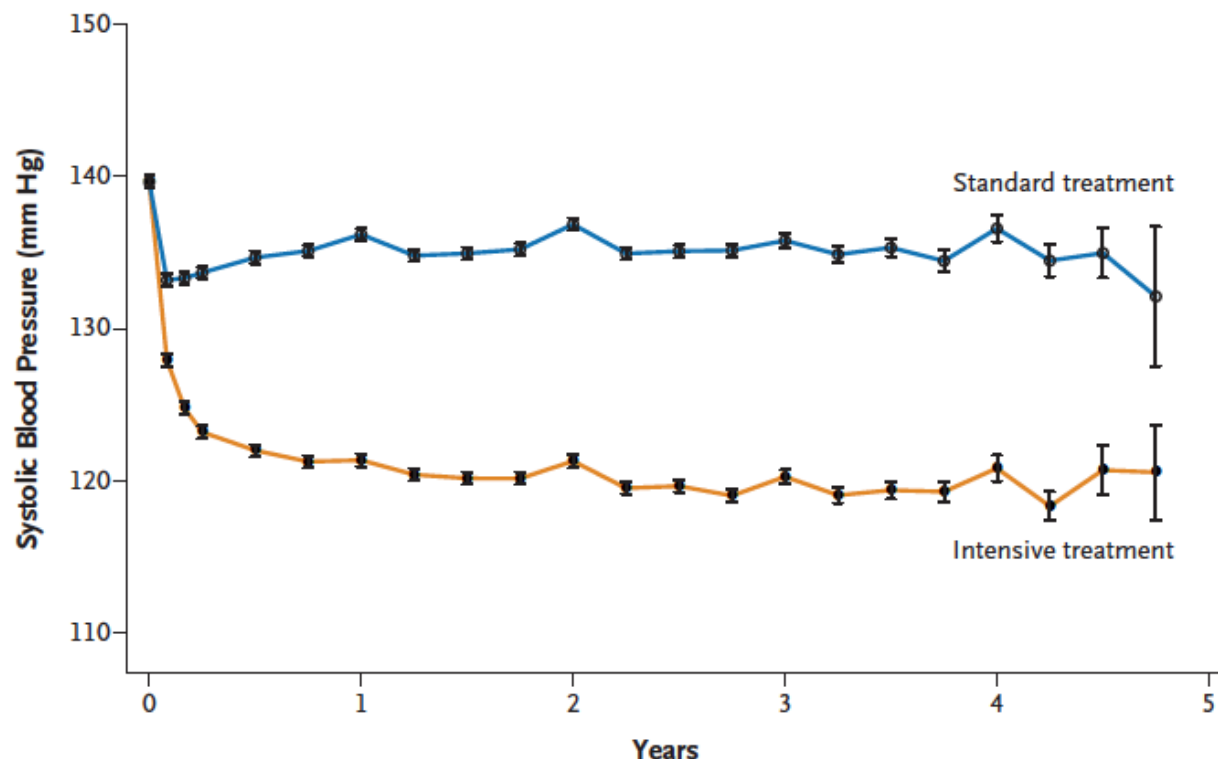
Contraindications:

- Patient unwilling or unable to adhere to multiple medications
- Standing SBP <110 mmHg
- Inability to measure SBP accurately
- Known secondary cause(s) of hypertension



SPRINT: SBPs Achieved

Average no. of medications
Intensive care: 2.8
Standard care: 1.8



No. with Data

Standard treatment	4683	4345	4222	4092	3997	3904	3115	1974	1000	274
Intensive treatment	4678	4375	4231	4091	4029	3920	3204	2035	1048	286

Mean No. of Medications

Standard treatment	1.9	1.8	1.8	1.8	1.8	1.8	1.8	1.8	1.8	1.9
Intensive treatment	2.3	2.7	2.8	2.8	2.8	2.8	2.8	2.8	2.8	3.0

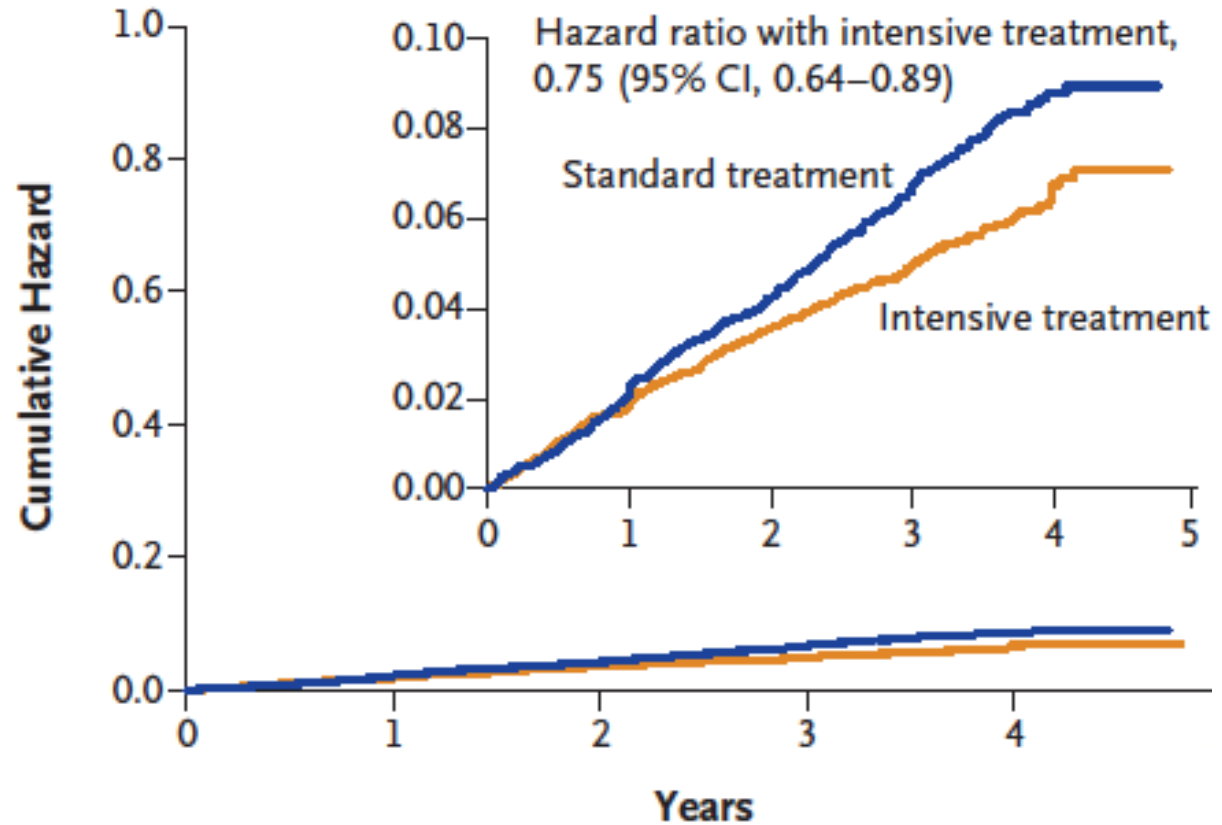
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The SPRINT Research Group, NEJM, Nov 9th, 2015



NNT=61

Primary Outcome



No. at Risk

Standard treatment	4683	4437	4228	2829	721
Intensive treatment	4678	4436	4256	2900	779



New Guideline post-SPRINT

New 2016

For high-risk patients, aged ≥ 50 years, with systolic BP levels ≥ 130 mm Hg, intensive management to target a systolic BP ≤ 120 mm Hg should be considered.

Intensive management should be guided by automated office BP measurements.

Patient selection for intensive management is recommended and caution should be taken in certain high-risk groups.



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Office BP Measurement Methods

Office (attended):

- Auscultatory (mercury, aneroid) – not recommended
- Non-automated oscillometric (electronic)

Automated office (unattended): AOBP

- Oscillometric (electronic)



2015 Recommendation on BP Measurement

New 2015

AOBP:

- Measurement using electronic (oscillometric) devices in the upper arm
- Provider outside the room/area (mitigates white coat effect)
- Multiple readings
- Mean automatically calculated



New 2016 Recommendation BP Measurement

- Automated office blood pressure (AOBP) is the preferred method of performing in-office BP measurement.

Automated Office (unattended, AOBP)
Oscillometric (electronic)





Comparison of Automated Office, Ambulatory and Pharmacy BP measurements

AOBP is Not Affected by the Setting in Which BP is Recorded

- Readings recorded in an ABPM unit or in an office waiting room are similar to AOBP recorded in a physician's examination room

Myers MG, et al. Blood Press Monit 2009;14:108-11

Greiver M, et al. Blood Press Monit 2012;17:137-8

Armstrong D, et al. Blood Press Monit 2015;20:204-8

-
- AOBP results obtained in the pharmacy were comparable with AOBP results from the physician's office

Chambers LW, et al. CMAJ Open 2013;1:E37-42



Comparisons of blood pressure readings obtained in clinical settings using different methods of blood pressure measurement

	Mean blood pressure* (mmHg)		
	Centre for Studies in Primary Care ₁	ABPM referral unit ₂	CAMBO trial ₃
Routine manual office BP	151/83	152/87	150/81
Automated office BP	140/80	132/75	135/77
Awake ambulatory BP	142/80	134/77	133/74

*The automated office blood pressure (BP) and awake ambulatory BP were similar, and both were lower than the routine manual BP obtained in community practice.

1. Beckett L et al, BMC Cardiovasc. Disord. 2005; 5: 18. 2. Myers MG et al, J. Hypertens. 2009; 27: 280. 3. Myers MG, et al. BMJ 2011; 342: d286.

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Predictive value of AOBP

AOBP predicts end-organ damage

- Systolic AOBP correlates with **LVMI** similarly to awake ABPM
- AOBP and 24-h ABPM have similar predictive ability for **microalbuminuria**
- AOBP is more strongly associated with **cIMT** (compared to OBPM)

cIMT: Carotid Intima Media Thickness

LVMI: Left Ventricular Mass Index

Campbell NRC, et al. J Hum Hypertens 2007;21:588-90; Andreadis EA, et al. Am J Hypertens 2011;24:661-6; Andreadis EA, et al. Am J Hypertens 2012;25:969-73.



Predictive Value of AOBP

The CHAP Study

AOBP Predicts Cardiovascular Events

- 3627 community-dwelling residents, aged >65 yrs, untreated for hypertension – part of the CHAP trial
- BpTRU® device in community pharmacies
- f/u 4.9 ± 1.0 yrs for **fatal and non-fatal CV events**

Myers MG, et al. Hypertension 2015;66:489-95. Kaczorowski J, et al. 2008 Preventive Medicine 46: 537–544



Predictive Value of AOBP

Cardiovascular Events

Systolic Blood Pressure

Systolic BP	Hazard Ratio
< 110	1.35 (0.8-2.3)
110-119 (referent)	1.00
120-129	1.08 (0.7-1.7)
130-139	1.30 (0.9-2.0)
135-144	1.66 (1.1-2.5)
140-149	1.79 (1.2-2.8)
150-159	1.96 (1.2-3.2)
160+	2.06 (1.3-3.4)

Diastolic Blood Pressure

Diastolic BP	Hazard Ratio
< 60	1.06 (0.6-1.9)
60-69 (referent)	1.00
70-79	1.15 (0.8-1.6)
80-89	1.72 (1.2-2.5)
90 +	2.07(1.3-3.2)

Myers MG, et al. *Hypertension* 2015;66:489-95.



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Health Behaviour Management

Intervention	Target
Reduce foods with added sodium	→ 2000 mg /day
Weight loss	BMI <25 kg/m ²
Alcohol restriction	≤ 2 drinks/day
Physical activity	30-60 minutes 4-7 days/week
Dietary patterns	DASH diet
Smoking cessation	Smoke-free environment
Waist circumference	Men < 102 cm Women < 88 cm
Potassium supplementation	NEW RECOMMENDATION



New 2016 Recommendation: Health Behaviours

Potassium intake:

- In patients *not* at risk of hyperkalemia, increase dietary potassium intake to reduce blood pressure.

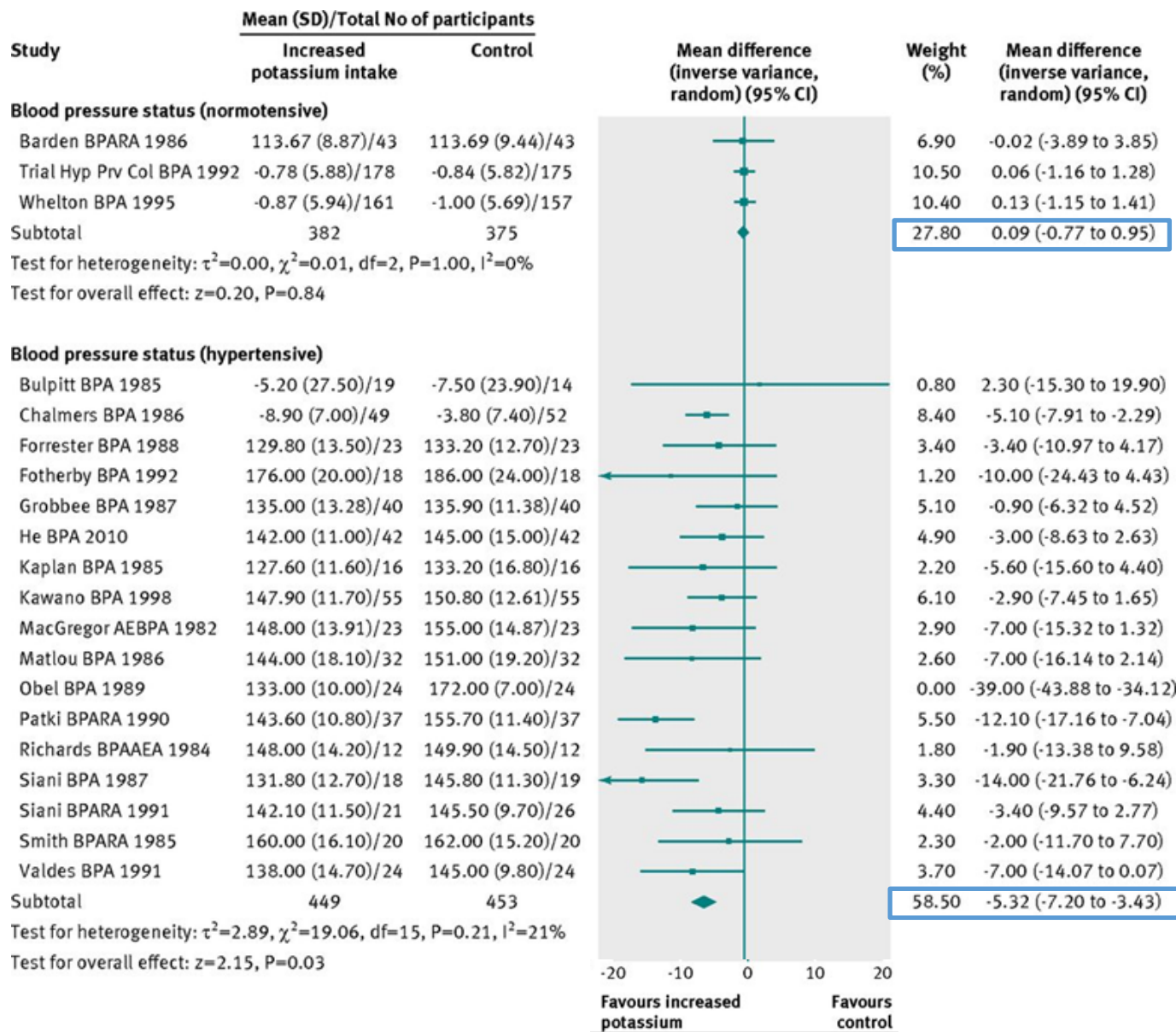


Systematic Reviews Showing Significant Effect of Potassium on BP

Author	Year	RCTs	Total N	Pooled effect SBP	Pooled effect DBP	Notes
Cappuccio	1991	19	586	-5.9 (-6.6 to -5.2)	-3.4 (-4.0 to -2.8)	Mixed status, 5-112 days, 10-150 participants; ?all RCTs
Whelton	1997	33	2609	-3.11 (-4.3 to -1.9)	-1.97 (-3.4 to -0.5)	Mixed status; 4d-3yrs; 10-484 N
Geleijnse	2003	27	NR	-2.4 (-3.8 to -1.1)	-1.57 (-2.6 to -0.5)	Mixed status; >2 wks duration
Dickinson	2006	5	425	-3.9 (-8.6 to 0.8) -11.2 (-25.2 to 2.7)	-1.5 (-6.2 to 3.1) -5.0 (-12.5 to 2.4)	Cochrane; hypertensive only; >8wks; 12-212 N; still significant heterogeneity; one trial not pooled – no ss dec in BP
van Bommel	2012	10	563	-7.12 (-8.5 to -5.7) -9.5 (-10.8 to -8.1)	-4.9 (-5.8 to -4.0) -6.4 (-7.3 to -5.6)	Hypertensive pts with high Na diet; heterogeneity dec. after exc. of outlier
Aburto	2013	22	1606	-3.49 (-5.2 to -1.8)	-1.96 (-3.1 to -0.9)	Mixed status; >4 wks; measured urinary K
Binia	2015	15	917	-4.7 (2.4 to -7)	- 3.5 (1.3 to 5.7)	Pts not on anti-htn Rx; mixed status; >=4wks;



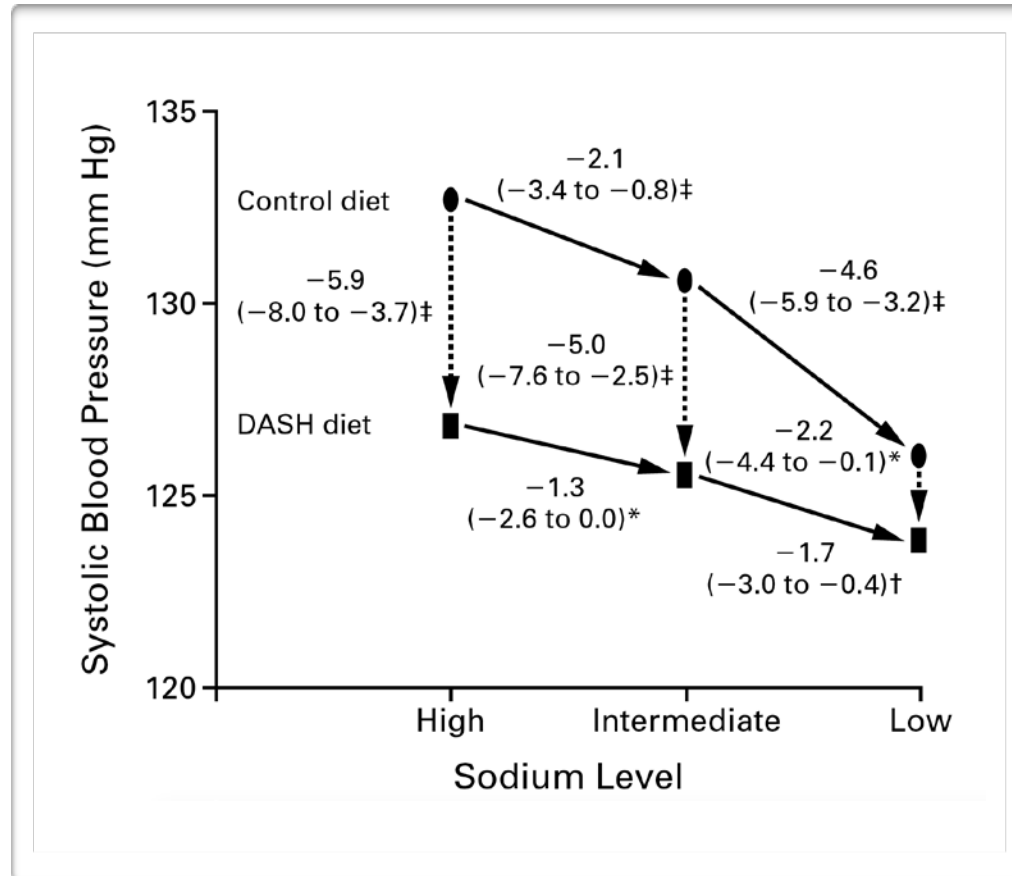
Increased Potassium intake decreases BP:



Effect of increased potassium intake on cardiovascular risk factors and disease: systematic review and meta-analyses. Aburto et al, BMJ 2013.



A K rich diet has additive effects to Na restriction



Sacks et al. *N Engl J Med*, Vol. 344, No. 1 · January 4, 2001



Enriching dietary potassium lowers BP: Summary

- Potassium supplementation leads to a decrease in BP
- Effect most consistently seen in patients with hypertension
- Effect of K is modified by Na intake, with greater effect at higher baseline Na



Risk of Hyperkalemia with K Supplementation

**Identify those at Risk of Hyperkalemia
with Potassium supplementation**

Prior to advising increase in potassium intake, the following kinds of patients – who are at high risk of hyperkalemia, should be assessed for suitability, and monitored closely:

- Patients taking renin-angiotensin-aldosterone inhibitors
- Patients on other drugs that can cause hyperkalemia (trimethoprim and sulfamethoxazole, amiloride, triamterene)
- Patients with CKD (GFR < 45mL/min)
- Patients with baseline serum potassium > 4.5 mmol/L



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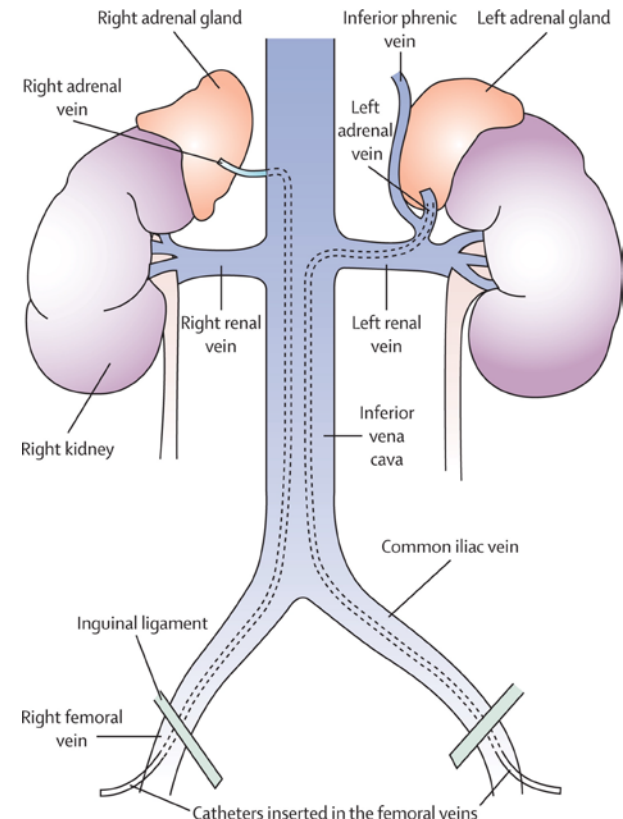
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New 2016 Recommendation: Endocrine Hypertension

- In patients with primary hyperaldosteronism and a definite adrenal mass who are eligible for surgery, adrenal venous sampling (AVS) is recommended to assess for lateralization of aldosterone secretion. AVS should be performed exclusively by experienced teams working in specialized centres.



adapted from Monticone S et al. *The Lancet*
Volume 3, No. 4, p296–303, April 2015

DOI: [http://dx.doi.org/10.1016/S2213-8587\(14\)70069-5](http://dx.doi.org/10.1016/S2213-8587(14)70069-5)



Role for adrenal venous sampling in primary aldosteronism

- Cohort of 203 patients from Mayo Clinic with PA, receiving CT and AVS
- 41% of those with normal CT had unilateral hypersecretion
- Only 51% of those with unilateral micronodules (<1 cm) had ipsilateral hypersecretion; 66% of those with unilateral macronodules (>1 cm) had ipsilateral hypersecretion
- 15% with micronodules and 3% with macronodules had contralateral hypersecretion
- 49% with bilateral micronodules and 33% with bilateral macronodules had unilateral hypersecretion

CT alone would incorrectly exclude 21% of patients for surgery
Based on CT alone, 25% would have received unnecessary surgery



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Change in 2016 Recommendation: Hypertension in CAD

Prior recommendation:

For patients with stable angina, β -blockers are preferred as initial therapy. *CCBs may also be used.*

New recommendation:

For patients with hypertension and stable angina pectoris but without prior HF, MI or coronary artery bypass surgery, *either a beta blocker or a calcium channel blocker can be used as initial therapy.*

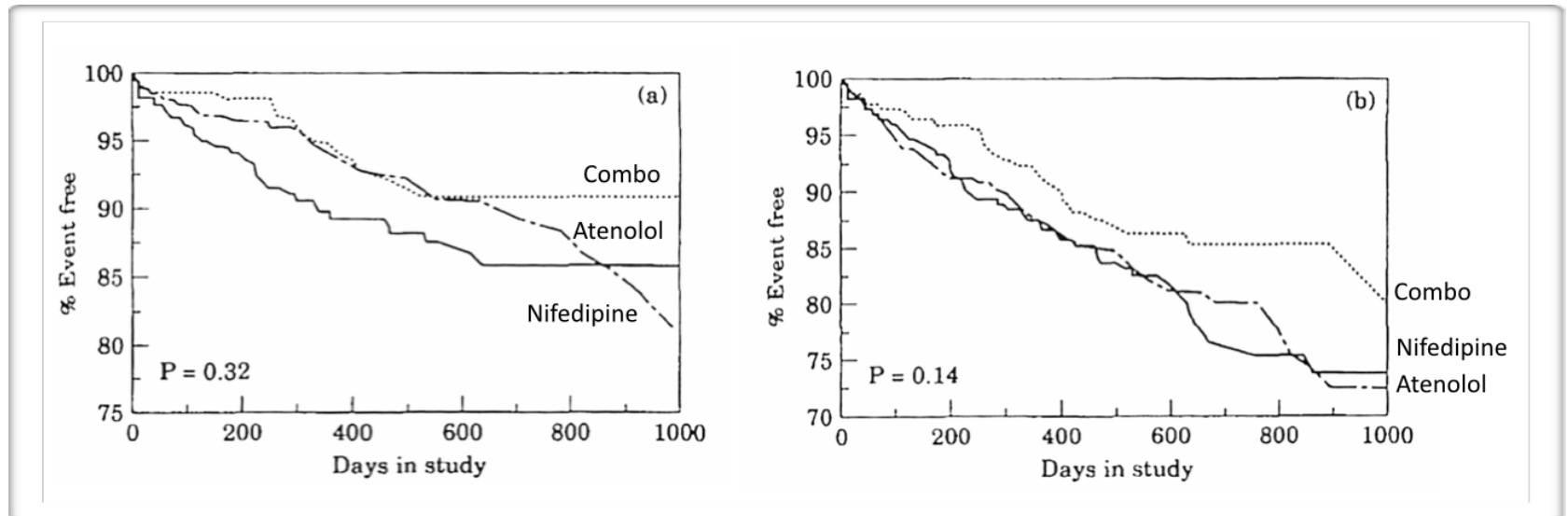


Beta blocker vs CCB in Treatment of CAD

The TIBET Trial

Events = MI, CV death, HF, ACS

Hard events + ischemic ST changes on 24 h ECG

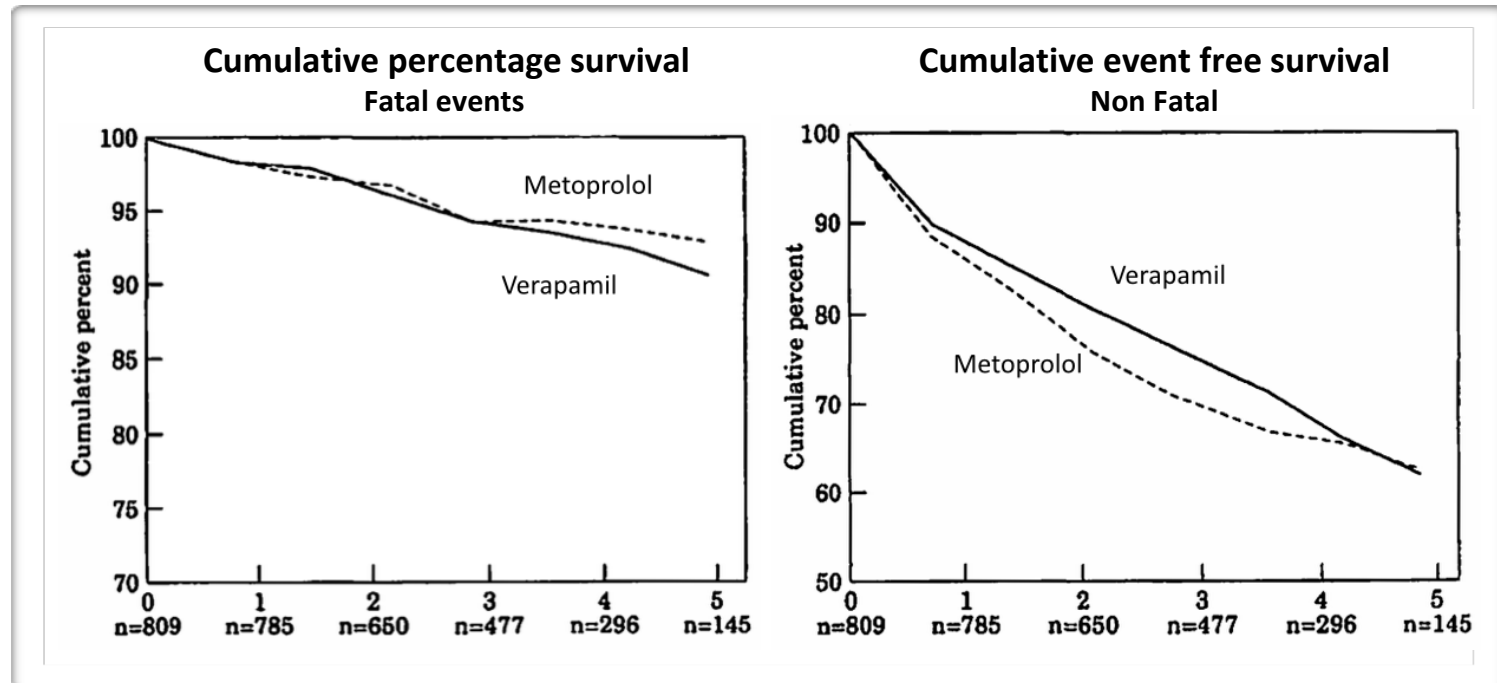


Total Ischaemic Burden European Trial (TIBET): Effects of atenolol (N=226), nifedipine SR (N=232) or combination (N=224) on outcome in chronic stable angina. Dargie et al. EHJ 1996;17:104-112



Beta blocker vs. CCB in treatment of CAD

The APSIS Trial



The curves were not extended beyond 5 years as few patients were followed thereafter

APSIS: metoprolol vs verapamil in stable angina pectoris

No difference: CV events (30.8% v 29.3%) CV mortality 4.7% vs 4.7%), Non-fatal CV events 26.1 v 24.3%

Hjemdahl P. et al. Favourable long term prognosis in stable angina pectoris: an extended follow up of the angina prognosis study in Stockholm (APSIS); Heart 2006;92:177-182



CCB vs. Non-CCB in treatment of CAD

The INVEST trial

As required to achieve
blood pressure control:

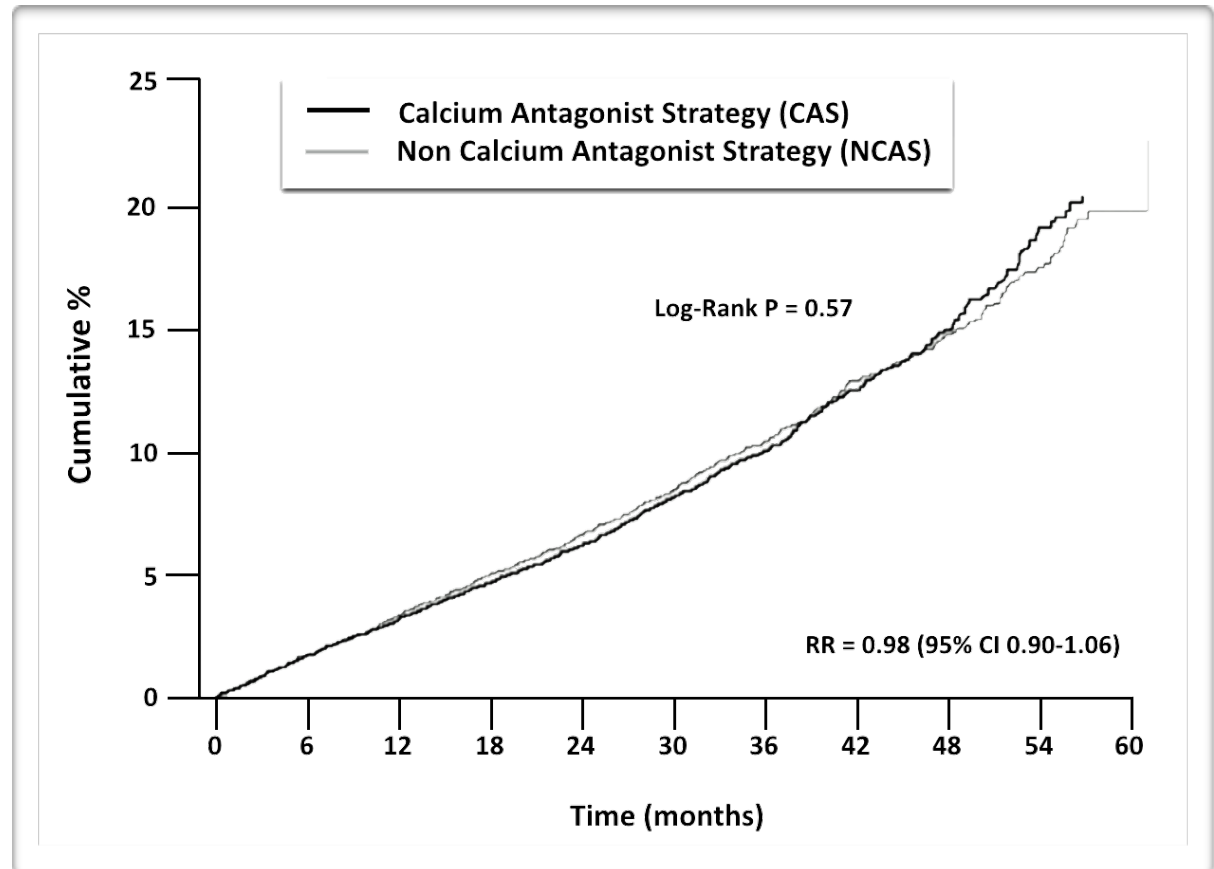
CCB strategy:

Verapamil sustained release +
Trandolapril + HCTZ

Non-CCB strategy:

Atenolol + HCTZ, + Trandolapril

-
- 22,000 HT patients with CAD
 - Primary Outcome:
Alive, Free of MI or Stroke
 - Total FU: 61,807 pt-y, mean FU 2.7y,
 - Annual event rate = 3.6%





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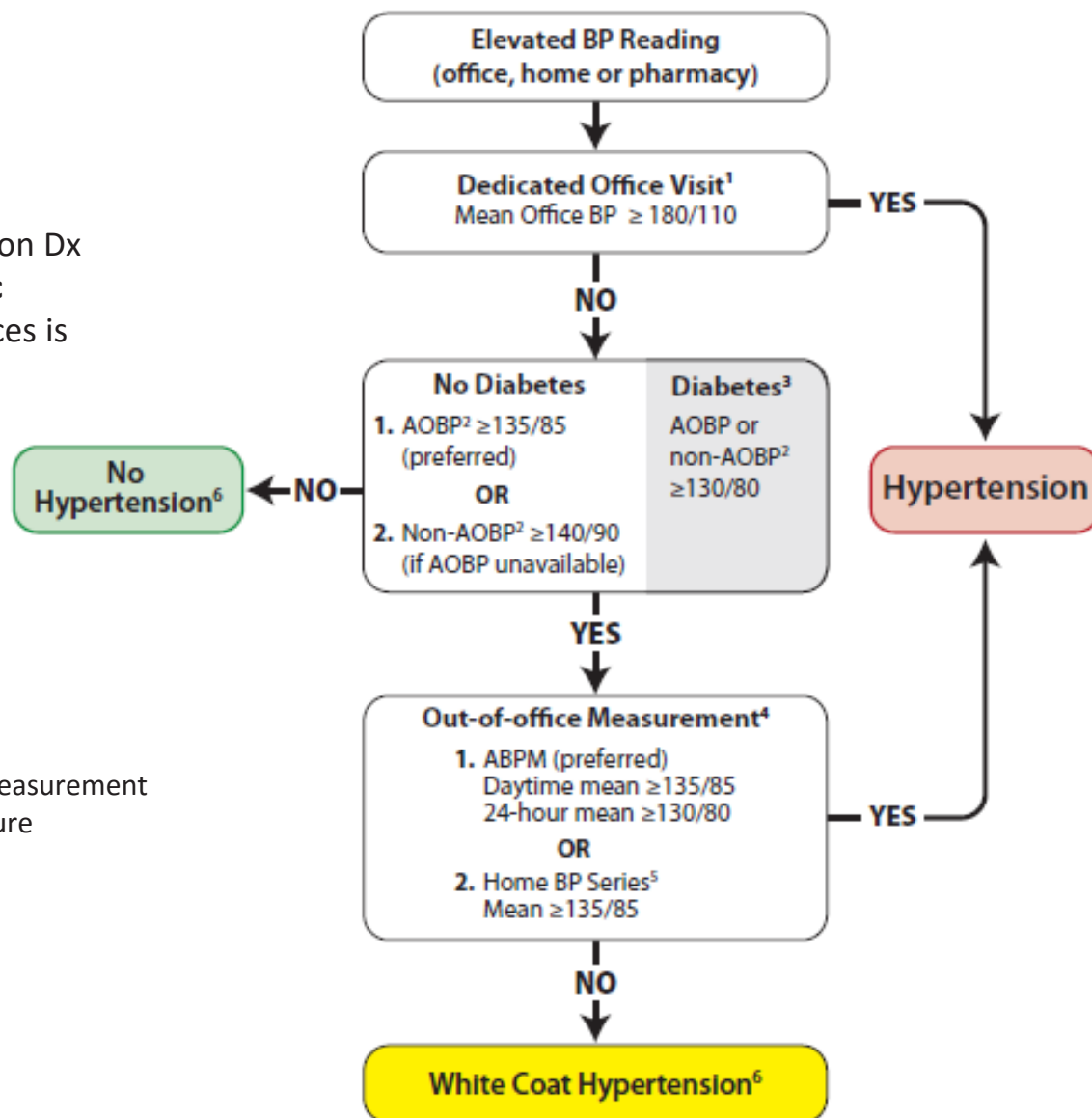
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Hypertension Diagnostic Algorithm



1. **Out of office** assessment is the preferred means of hypertension Dx
2. **Measurement using electronic** (oscillometric) upper arm devices is preferred over auscultation



ABPM: Ambulatory Blood Pressure Measurement

AOBP: Automated Office Blood Pressure

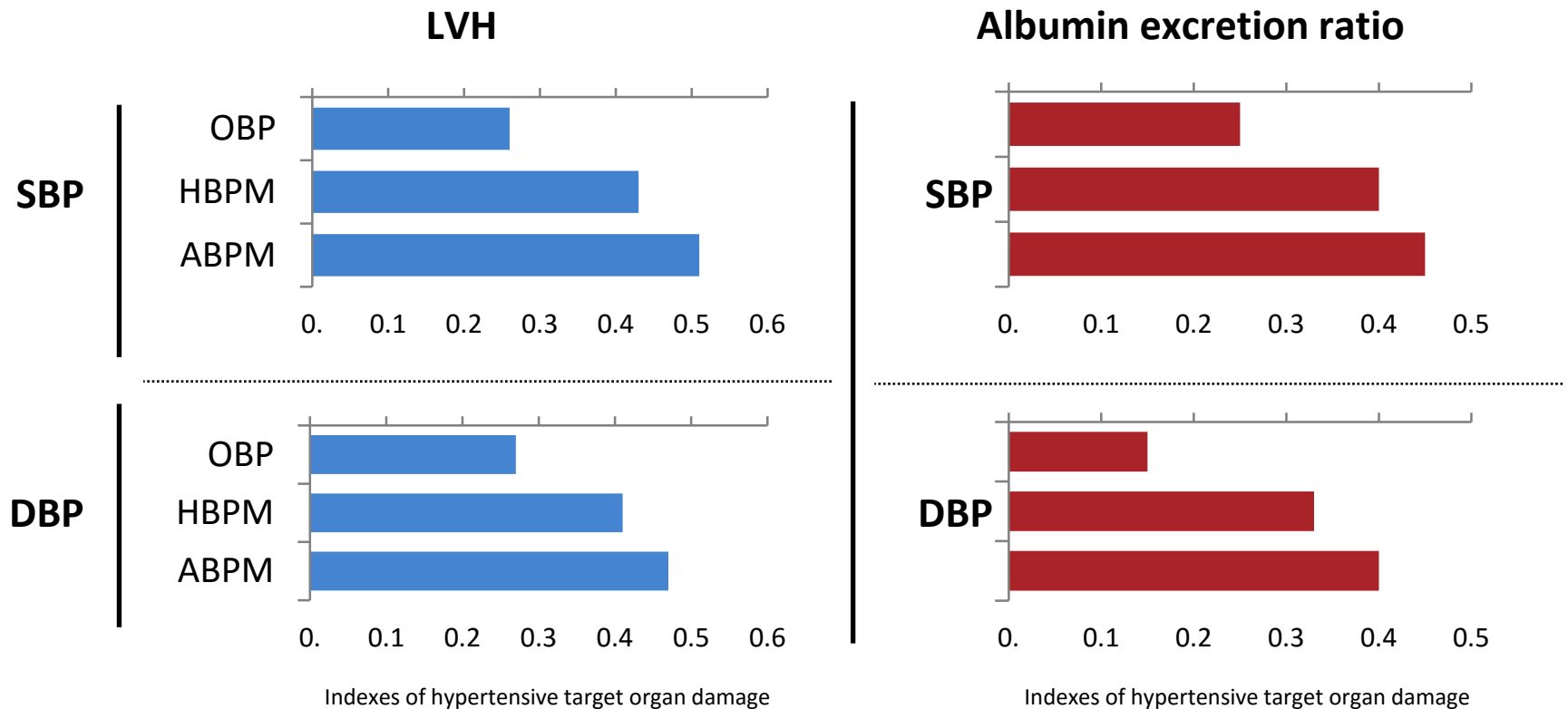


Out-of-Office BP Measurements

- ABPM has better predictive ability than OBPM and is the recommended out-of-office measurement method.
- HBPM has better predictive ability than OBPM and is recommended if ABPM is not tolerated, not readily available or due to patient preference.
- Identifies white coat hypertension and masked hypertension.



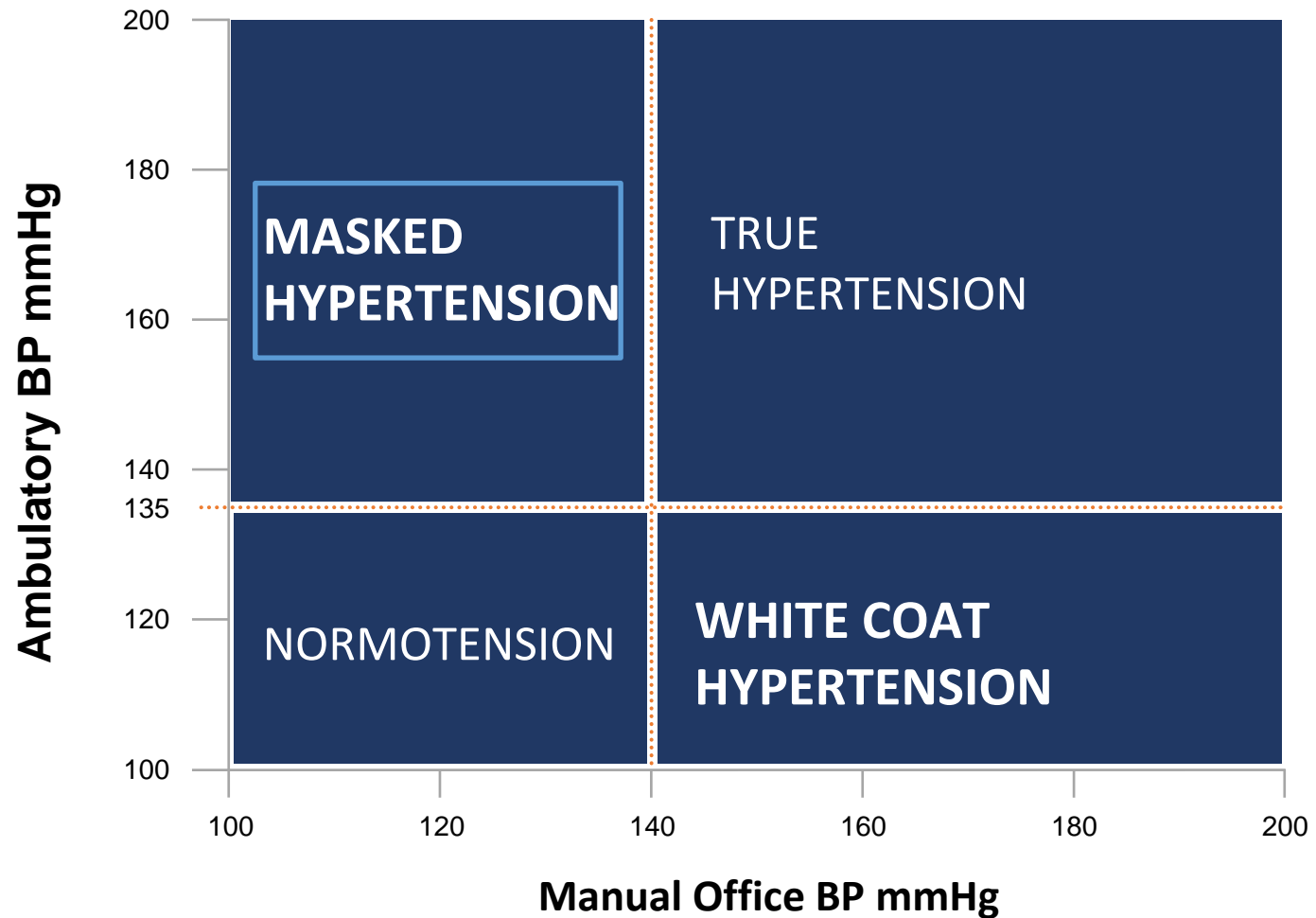
Out-of-office BP measurements are more highly correlated with BP-related risk



Value of Home Blood Pressures as Predictor of Target Organ Damage in Mild Arterial Hypertension
Mulè G. et al. *J Cardiovasc Risk* 2002;9:123-9.



White Coat and Masked Hypertension



Adapted from CHEP. www.hypertension.ca. Derived from Pickering *et al. Hypertension* 2002; 40: 795-796.



Criteria for the Diagnosis of Masked Hypertension

	BP (mm Hg)
Office BP	< 140/90
Awake Ambulatory	$\geq 135/85$
24-hour Ambulatory BP	$\geq 130/80$



Prevalence of Masked Hypertension

about

10%

in the general
population

about

30%

in treated
hypertensive
patients*

higher

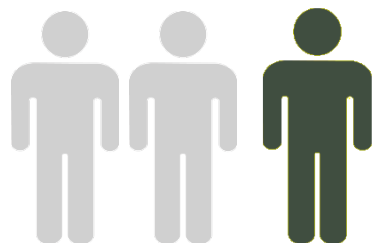
in patients with

diabetes

and

chronic kidney

disease patients

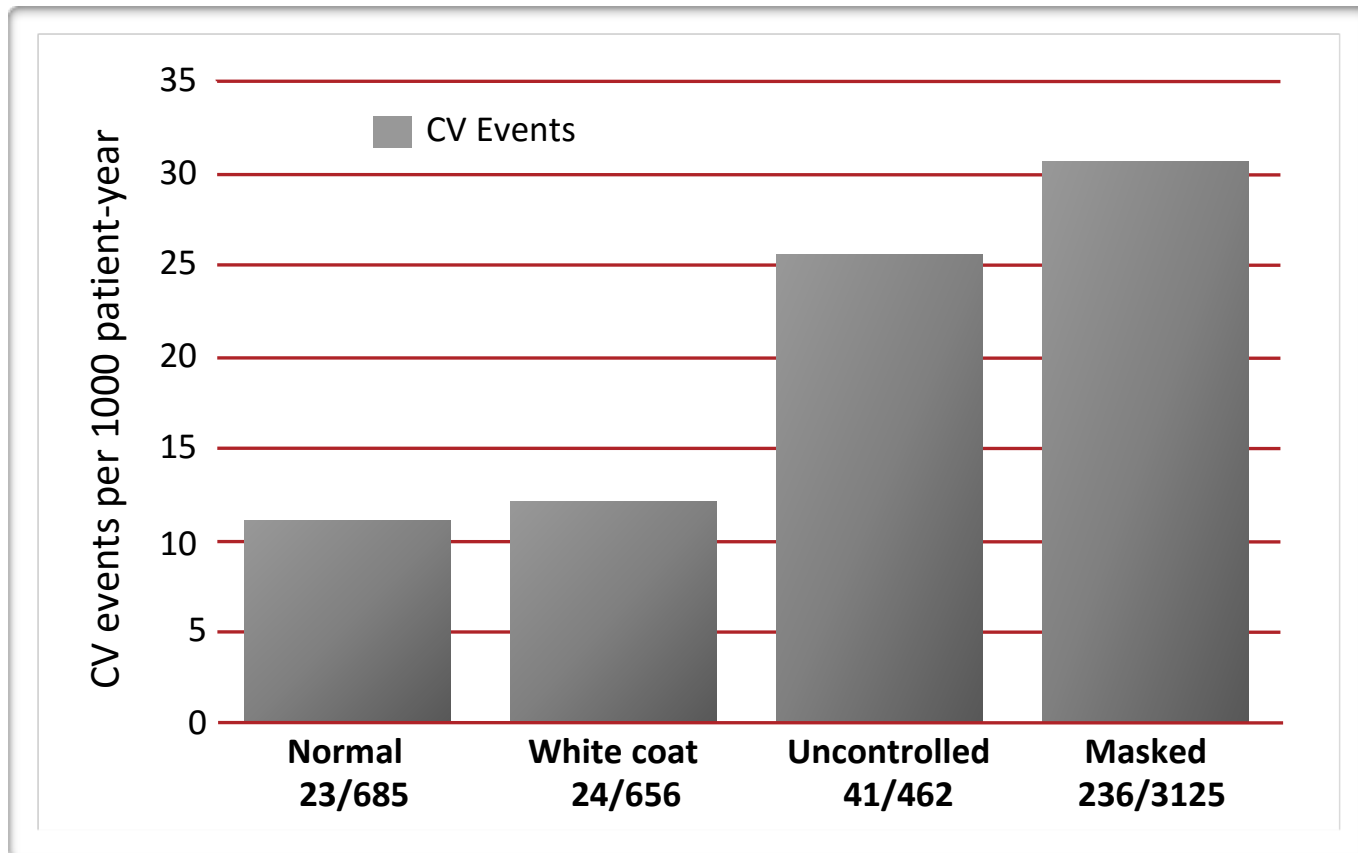


*One out of three treated hypertensive patients
has masked hypertension*

*Andalib A et al. Int Med J 2012; 42:260-6



The prognosis of *white coat* and *masked* hypertension



Okhubo et al. *J. Am. Coll. Cardiol.* 2005;46;508-515



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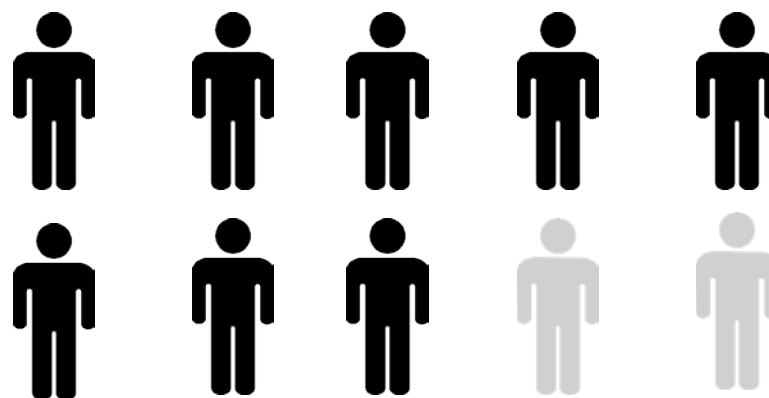
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Cardiovascular Risk Factors in Hypertensive Patients

**8 out of 10 hypertensive patients
have at least 1 additional risk factor**

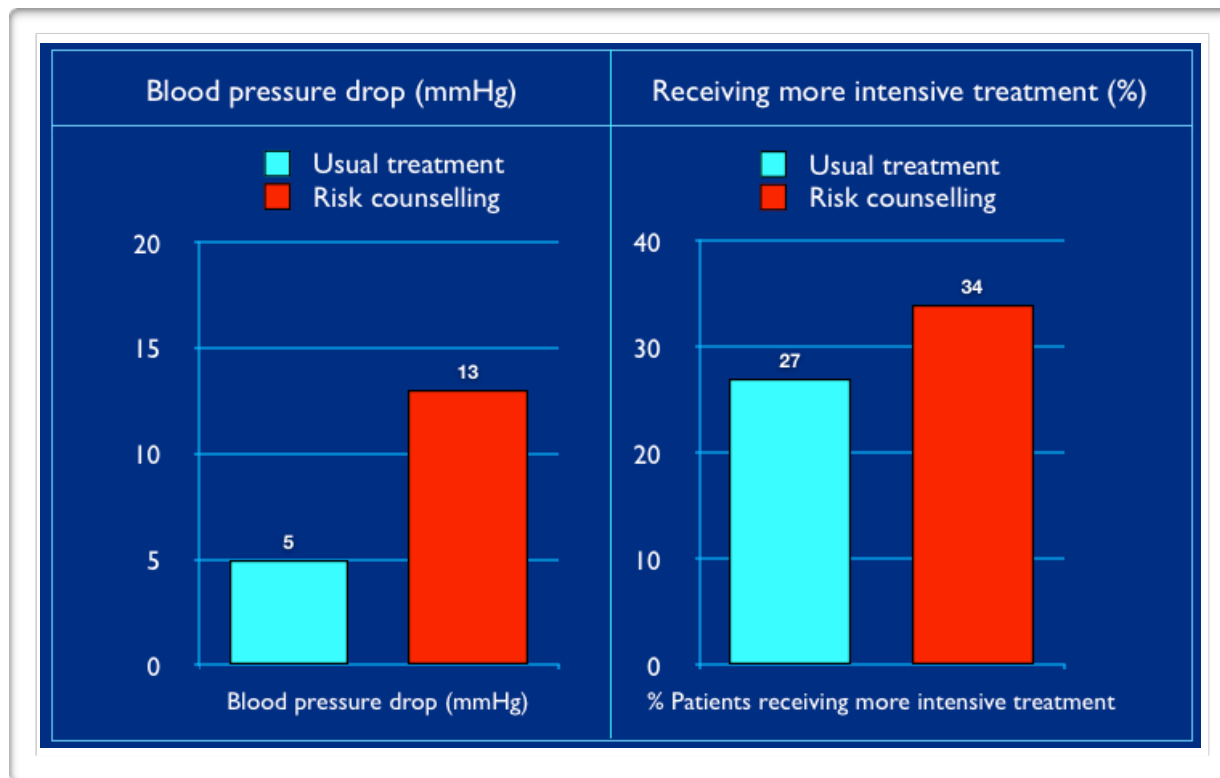


Gee ME, Bienek A, McAlister FA, et al. Factors Associated With Lack of Awareness and Uncontrolled High Blood Pressure Among Canadian Adults With Hypertension. Can J Cardiol. 2012;28:375-382.



Impact of discussing coronary risk with patients on blood pressure treatment

Informing Patients of Their Global Risk improves BP Control
Cardiovascular Age™ www.myhealthcheckup.com



Grover SA, et al. *J Gen Intern Med* 2009;24(1):33-9.



Vascular protection: **Statins for high risk hypertensive patients**

Statins are recommended in high risk hypertensive patients based on having established atherosclerotic disease or at least 3 of the following:

- Male
- 55 y or older
- Smoking
- Type 2 Diabetes
- Total-C/HDL-C ratio of 6 or higher
- Premature Family History of CV disease
- Previous Stroke or TIA
- LVH
- ECG abnormalities
- Albuminuria or CKD
- Peripheral Vascular Disease

The Treatment of Hypertension is All About Vascular Protection



Vascular protection: ASA for hypertensive patients

**Low dose ASA in hypertensive patients
is recommended for patients ≥ 50 years**

Caution should be exercised if BP is not controlled.

Hansson L, Zanchetti A, Carruthers SG, et al. Effects of intensive blood-pressure lowering and low-dose aspirin in patients with hypertension: principal results of the Hypertension Optimal Treatment (HOT) randomised trial. Lancet 1998;351:1755-1762.



Strong Evidence for Vascular Protection: Smoking Cessation

New 2015

- **Tobacco use status** of all patients should be updated on a regular basis and health care providers should clearly advise patients to quit smoking.
- **Advice** in combination with pharmacotherapy (e.g., varenicline, bupropion, nicotine replacement therapy) should be offered to all smokers with a goal of smoking cessation.



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Adherence in hypertensive patients

Adherence Can Be Improved by a Multi-Pronged Approach

- Educate patients and patients' families about their disease/treatment regimens verbally and in writing
- Use an interdisciplinary care approach coordinating with work-site health care givers and pharmacists if available
- Healthcare practitioner-based telephone contact, particularly, over the first three months of therapy
- Encourage greater patient responsibility/autonomy in regular monitoring of their blood pressure



Adherence in hypertensive patients

Adherence Can Be Improved by a Multi-Pronged Approach

- Assess adherence to pharmacological and health behaviour therapies at every visit
- Teach patients to take their pills on a regular schedule associated with a routine daily activity e.g. brushing teeth.
- Simplify medication regimens using long-acting once-daily dosing
- Utilize single pill combinations
- Utilize unit-of-use packaging e.g. blister packaging



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CHEP 2016 Recommendations

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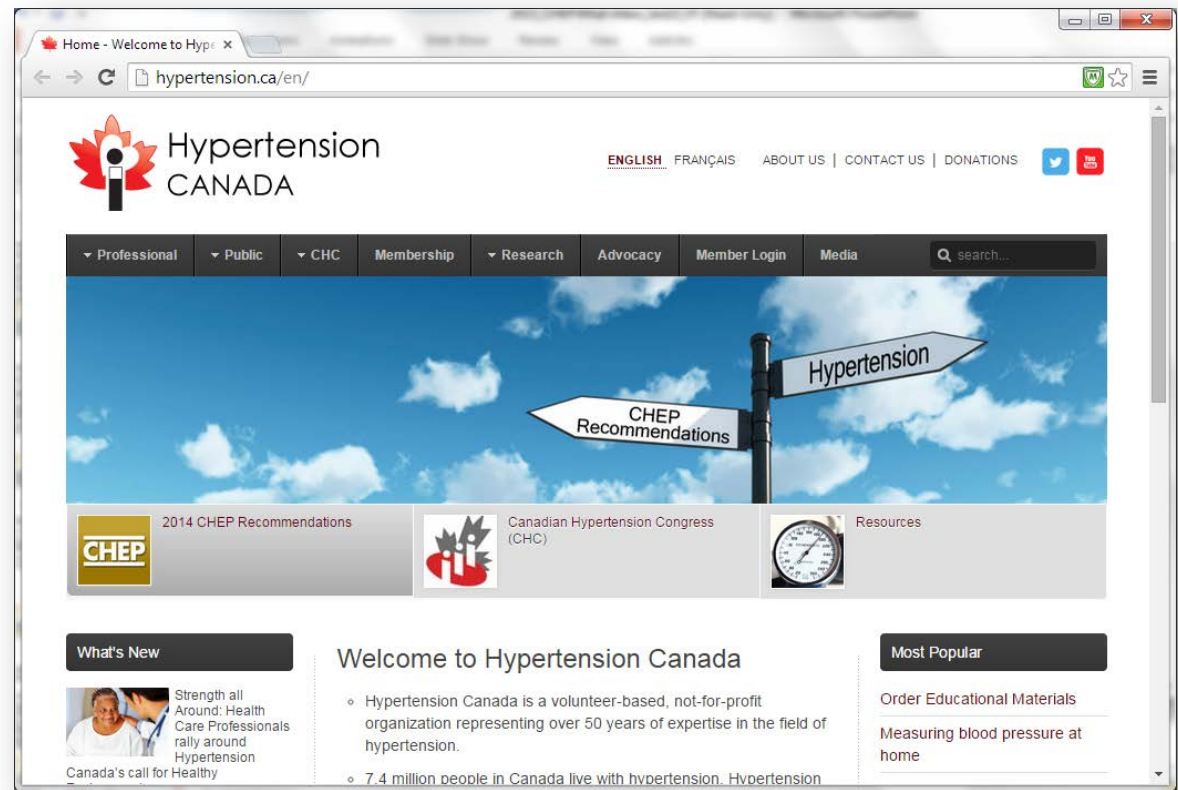
hypertension.ca

For patients:

- Free access to the latest information and resources

For professionals:

- Accredited 15.5 hour interdisciplinary training program
- Free monthly news updates, featured research and educational resources
- Become a member for special privileges and savings





2016

Hypertension Canada CHEP Guidelines for the Management of Hypertension

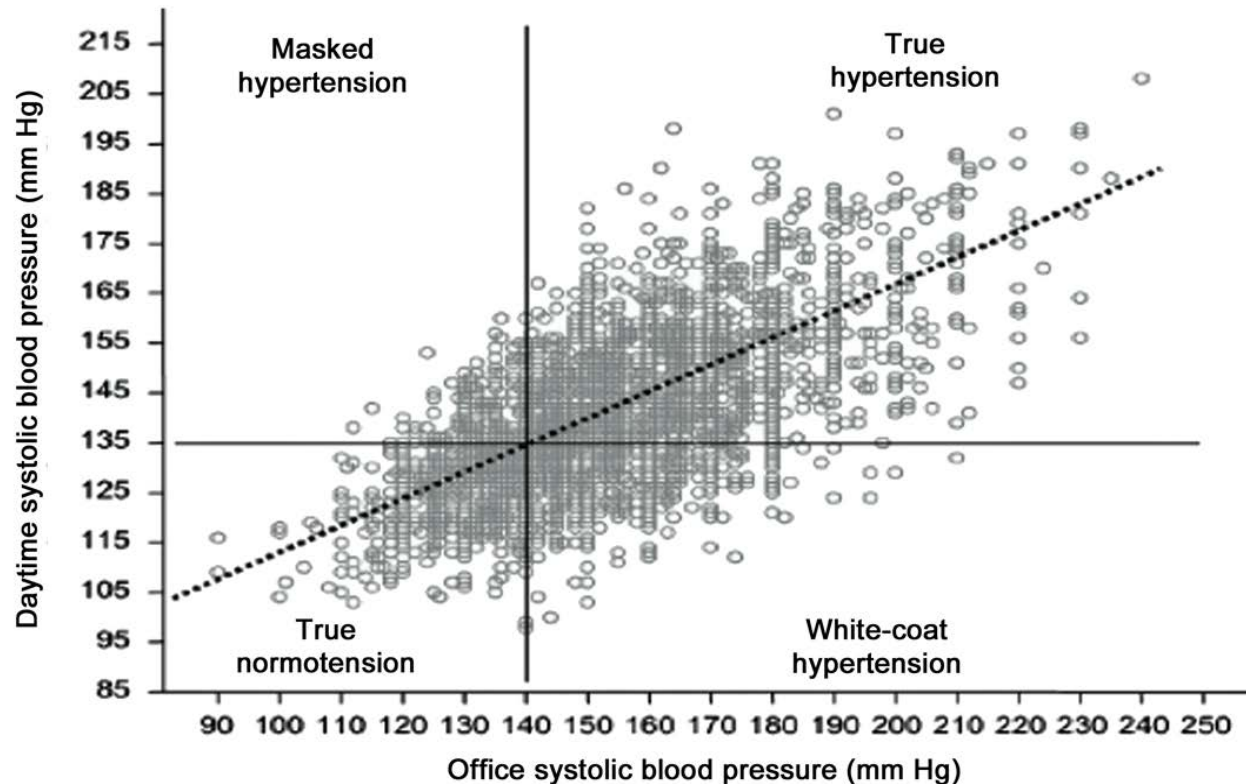
What's new in the treatment of hypertension?
What's still really important?



Additional Data Slides



Office BP and ABPM: different evaluation of Blood Pressure



Verdecchia P et al, The clinical significance of white-coat and masked hypertension.
Blood Press Monit 2007; 12: 387–389



Correlation of *AOBP* and *OBPM* with *ABPM*

Stronger Correlation with *ABPM* for *AOBP* than for *OBPM*

Study	<i>n</i>	Patient population	Coefficient of correlation (<i>r</i>) office BP vs awake ambulatory BP	
			Automated office BP	Manual office BP
Beckett and Godwin ²⁶	481	Community family practice	0.57/0.61	0.15/0.32
Myers <i>et al.</i> ²⁷	309	Ambulatory BP unit	0.62/0.72	0.32/0.48
Myers <i>et al.</i> ²⁹	303	Community family practice (research study)	0.34/0.56	0.10/0.40

Beckett L, Godwin M. *BMC Cardiovasc Disord* 2005;5:18;Myers MG, *et al.* *J Hypertens* 2009; 27: 280-6;
Myers MG, *et al.* *BMJ* 2011;342:d286



Studies comparing AOBP measurement with Awake ABPM

AOBPs Equivalent to Awake ABPs

Table 2. Studies comparing AOBP measurement with AABP measurement: Mean overall AOBP was 137/79 mm Hg, and mean overall AABP was 137/79 mm Hg.

STUDY	NO. OF PATIENTS	POPULATION	AOBP, mm Hg	AABP, mm Hg
Myers et al, ⁵ 2009	309	ABPM unit	132/75	134/77
Myers et al, ²¹ 2008	200	ABPM unit	133/72	135/76
	200	ABPM unit	132/76	134/77
Myers et al, ²² 2010	139	ABPM unit	141/82	142/81
Beckett and Godwin, ²⁵ 2005	481	Family practice	140/80	142/80
Myers et al, ²⁶ 2009	62	Hypertension clinic	140/77	141/77
Myers, ²⁷ 2010	254	ABPM unit	133/80	135/81
Godwin et al, ²⁸ 2011	654	Family practice	139/80	141/80
Myers et al, ²⁹ 2011	303	Family practice	135/77	133/74
Andreadis et al, ³⁰ 2011	90	Research unit	140/88	136/87

AABP—awake ambulatory BP, ABPM—24-hour ambulatory BP monitoring, AOBP—automated office BP, BP—blood pressure.

Myers MG, et al. *Can Fam Physician* 2014;60:127-32; Myers MG. *J Clin Hypertens* 2014;16:83-6; Myers MG. *J Hypertens* 2012;30:1894-8; Myers MG, et al. *Family Practice* 2012;29:376-82; Myers MG, et al. *BMJ* 2011;342:d286; Myers MG. *J Hypertens* 2010;28:703-8; Myers MG, et al. *J Hypertens* 2009;27:280-6; Myers MG, et al. *Blood Press Monit* 2009;14:108-11; Myers MG, Godwin M. *J Clin Hypertens* 2007;9:267-70