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## ORIGINAL ARTICLE

# Comparative Cost-Effectiveness of Hypertension Treatment in Non-Hispanic Blacks and Whites According to 2014 Guidelines: A Modeling Study 

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## BACKGROUND

We compared the cost-effectiveness of hypertension treatment in nonHispanic blacks and non-Hispanic whites according to 2014 US hypertension treatment guidelines.

## METHODS

The cardiovascular disease (CVD) policy model simulated CVD events, quality-adjusted life years (QALYs), and treatment costs in 35- to 74-yearold adults with untreated hypertension. CVD incidence, mortality, and risk factor levels were obtained from cohort studies, hospital registries, vital statistics, and national surveys. Stage 1 hypertension was defined as blood pressure $140-149 / 90-99 \mathrm{~mm} \mathrm{Hg}$; stage 2 hypertension as $\geq 150 / 100 \mathrm{~mm}$ Hg. Probabilistic input distribution sampling informed 95\% uncertainty intervals (Uls). Incremental cost-effectiveness ratios (ICERs) $<\$ 50,000 /$ QALY gained were considered cost-effective.

## RESULTS

Treating 0.7 million hypertensive non-Hispanic black adults would prevent about 8,000 CVD events annually; treating 3.4 million


#### Abstract

non-Hispanic whites would prevent about 35,000 events. Overall 2014 guideline implementation would be cost saving in both groups compared with no treatment. For stage 1 hypertension but without diabetes or chronic kidney disease, cost savings extended to non-Hispanic black males ages 35-44 but not same-aged non-Hispanic white males (ICER \$57,000/QALY; 95\% UI \$15,000$\$ 100,000$ ) and cost-effectiveness extended to non-Hispanic black females ages 35-44 (ICER \$46,000/QALY; \$17,000-\$76,000) but not same-aged non-Hispanic white females (ICER \$181,000/QALY; \$111,000-\$235,000).


## CONCLUSIONS

Compared with non-Hispanic whites, cost-effectiveness of implementing hypertension guidelines would extend to a larger proportion of non-Hispanic black hypertensive patients.

Keywords: blood pressure; cost-effectiveness; hypertension; race.
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Hypertension awareness and treatment improved in the United States over the last 2 decades, contributing to a nearly $50 \%$ improvement in the proportion of hypertensive patients with controlled blood pressure (BP). ${ }^{1}$ Hypertension treatment, control, and awareness improved in non-Hispanic blacks, but hypertension prevalence and the proportion of uncontrolled BP in non-Hispanic black hypertensive patients remain higher compared with non-Hispanic whites. Rates of deaths attributable to hypertension have increased over time in non-Hispanic black men, in whom mortality rates are double that of non-Hispanic white men. ${ }^{2}$ Hypertension awareness and treatment also continue to be lowest in young adults, ${ }^{1}$ and hypertension screening and control have been especially difficult to implement in young non-Hispanic black men. ${ }^{3,4}$

Our recent cost-effectiveness modeling study found that implementing 2014 US hypertension guidelines would be cost saving overall in untreated hypertensive adults aged 35-74 years old, but cost-effectiveness did not extend to primary of 35 - to 44 -year-old women with stage 1 hypertension for primary cardiovascular disease (CVD) prevention. ${ }^{5}$ Given their higher CVD risk, we hypothesized that hypertension treatment would be more cost-effective in young and middle-aged non-Hispanic blacks compared with sameaged non-Hispanic whites. We therefore compared the costeffectiveness of implementing 2014 hypertension guidelines in untreated non-Hispanic black and non-Hispanic white patients overall and by age, sex, hypertension stage, and CVD, diabetes, and chronic kidney disease (CKD) status.

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## METHODS

## CVD policy model

The CVD policy model is a computer-simulation, statetransition model of incidence, prevalence, mortality, and costs of CVD in the United States (Supplementary Figure). The primary study population included non-Hispanic black and white men and women aged 35-74 in 2014 who were not receiving pharmacotherapy for hypertension. We restricted our analysis to currently untreated hypertensive patients because we were not able to discern among different causes of uncontrolled BP in already treated patients (financial obstacles, nonadherence, side effects, or contraindications due to comorbidities). Adults aged 75 years and older were excluded from this analysis due to highly variable rates of medication-related adverse events. ${ }^{6}$ We were unable to analyze Hispanic Americans or Asian Americans at this time due to insufficient contemporary CVD incidence data for these heterogeneous groups.

Means or proportions and joint distributions of risk factors, including systolic and diastolic BP, cholesterols, and hypertension medication use, smoking, diabetes, and CKD status, were estimated from pooled National Health and Nutrition Examination Surveys (NHANES) 2007-2010 by 10 -year age categories, sex, and race/ethnic group. NHANES 2007-2010 systolic BP was categorized into 3 main categories: $<140,140-159$ (stage 1 hypertension), and $\geq 160 \mathrm{~mm} \mathrm{Hg}$ (stage 2 hypertension). To assess 2014 guideline treatment recommendations in older adults (ages 60-74 years), stage 1 systolic BP was further divided into the intervals 140-149 and $150-159 \mathrm{~mm}$ Hg. Diastolic BP categories were <90, $90-99$ (stage 1), and $\geq 100 \mathrm{~mm} \mathrm{Hg}$ (stage 2). Hypertensive patients were considered untreated if they answered no to both NHANES questions: "Because of your high blood pressure/hypertension, have you ever been told to take prescribed medicine?" and "Are you currently taking medication to lower your blood pressure?"

Default multivariate stroke and coronary heart disease incidence functions were estimated in an original Framingham Heart Study analysis. The CVD policy model simulated CVD events, non-CVD deaths, quality-adjusted life years (QALYs), and hypertension and CVD treatment costs of implementing 2014 US treatment guidelines in untreated hypertensive non-Hispanic black and white adults aged 35-74 years compared with no treatment.

## Race/ethnic-specific CVD policy models

While the structure of both the non-Hispanic black and non-Hispanic white CVD policy model versions is identical to that of the national model, several inputs were re-populated with data specific to non-Hispanic blacks and nonHispanic whites (for details, see Supplementary Text).

Population projections for the years 2014-2024 were obtained from 2010 US Census Bureau. National Health Interview Survey data were used to estimate the proportion of adults with known coronary heart disease, stroke, or both these conditions. All beta coefficients for the effect of non-BP risk factors on incident stroke (diabetes, CKD, and smoking)
and on incident coronary heart disease (cholesterols, diabetes, CKD, and smoking) were estimated from Framingham Heart Study data. Numbers of 2010 CVD deaths (stroke (ICD-10 codes I60-I69), coronary heart disease (I20-I25 and two-thirds of I49, I50, and I51), and hypertensive heart disease deaths (I11.0, I11.9)) and non-CVD deaths (remainder of ICD codes) reported by the Centers for Disease Control (CDC WONDER) for 2010 were used as model calibration targets.

Hospitalized stroke incidence in non-Hispanic whites was based on rates observed in the 2010 National Hospital Discharge Survey (NHDS). Since NHDS is missing over $15 \%$ of data on self-reported race/ethnicity, we were unable to similarly determine the number of total stroke events in non-Hispanic blacks. For non-Hispanic black stroke incidence rates, we first adjusted national NHDS-based stroke incidence rates for risk factor differences between the United States and racial subpopulations and the stronger association between BP and stroke in non-Hispanic blacks. ${ }^{7}$ Stroke incidence rate inputs for both groups were calibrated until simulated stroke deaths predicted 2010 national stroke mortality numbers within 1\% (Supplementary Table 1). The resulting incidence of hospitalized stroke approximated ageand sex-specific stroke incidence rates observed in stroke cohorts and surveillance studies of non-Hispanic blacks and whites. ${ }^{8-13}$

Non-Hispanic white age-range/gender-specific coronary heart disease incidence rates were estimated by calibrating incidence to match with cause-specific coronary heart disease mortality in 2010. Coronary heart disease incidence for nonHispanic blacks was estimated by multiplying white age and sex incidence rates by black/white incidence ratios observed during the 2005-2011 interval in the Atherosclerosis Risk in Communities (ARIC) surveillance study. Coronary heart disease incidence rate inputs for both groups were calibrated until simulated coronary heart disease deaths predicted 2010 national mortality numbers within 1\% (Supplementary Table 1).

Thirty-day acute myocardial infarction case fatality was assumed to be similar for non-Hispanic blacks and whites in 2010. ${ }^{14}$ Stroke 30-day case fatality rates for non-Hispanic blacks were assumed to be higher than national averages based on an observed difference between white and black case fatality rates in the ARIC cohort study. ${ }^{8}$ Annual probabilities of stroke after myocardial infarction ${ }^{15}$ and the probability of coronary heart disease in stroke patients were based on natural history studies and were assumed to be the same for both race/ethnic groups (Supplementary Text). Deaths, disability, and costs of heart failure due to coronary heart disease were tabulated by the CVD policy model. Deaths attributable to heart failure due to hypertensive heart disease (and not due to coronary heart disease) were tabulated, but chronic costs and nonfatal disability due to hypertensive heart disease were not.

## Hypertension treatment model inputs

The 2014 US hypertension guidelines ("JNC 8") recommended a goal of $<140 / 90 \mathrm{~mm} \mathrm{Hg}$ for diabetes and/or CKD, diastolic BP $<90$ if age under 60 years and BP $<150 / 90$ if
age $>60$ years without diabetes or CKD. ${ }^{16}$ The number of US non-Hispanic black and white adults eligible for treatment under JNC 8 was estimated using these categories and BP and treatment status information in NHANES 2007-2010.

BP change due to antihypertensive medications was determined by pretreatment BP and the number of standard doses of medications needed to reach the guideline BP goal, according to a trials-based formula. ${ }^{17} \mathrm{BP}$ changes were calculated based on pretreatment BP, age, and sex. We assumed the same BP reduction per standard dose of the main drug classes (Table 1; Supplementary Text). ${ }^{17,18}$ Quality of life penalties were applied for side effects. ${ }^{17}$ Treatment costs included hypertension monitoring costs, drug side effects, and the average wholesale cost of drugs. We assumed the $75 \%$ adherence observed in the same clinical trials metaanalysis that provided the association between BP reduction and CVD risk reduction. ${ }^{18}$ In our prior analysis, lower adherence reduced effectiveness and costs proportionally; thus, lower adherence reduced population impact but had little effect on cost-effectiveness. ${ }^{5}$

We assumed CVD risk reduction is due to BP reduction, ${ }^{18}$ and that BP is lowered to a similar extent across classes when comparing per-class standard doses. ${ }^{17,27}$ We started with observational Prospective Studies Collaboration age-specific relative risks and $95 \%$ confidence intervals for coronary heart disease and stroke per 10 mm Hg change in systolic BP or 5 mm Hg diastolic BP (Table 1). ${ }^{28}$ Age-specific relative risk inputs were calibrated to be within $\leq 0.02$ of these estimates and overall relative risks within $95 \%$ confidence interval bounds of the summary estimate from a large metaanalysis of randomized clinical trials of hypertension treatment (Supplementary Tables 2 and 3). ${ }^{18}$ The stroke relative risk estimate was found to be close to the pooled estimate from the East Asian trials included in that analysis (0.59 (0.49-0.71); Supplementary Text). The resulting relative risk assumptions were validated for treatment of systolic BP in ages $60-74$ years by simulating the treatment and placebo groups of the Systolic Hypertension in the Elderly Program (SHEP) trial and comparing simulated relative rate ratios with those observed in the trial (Supplementary Text; Supplementary Table 4).

## Main analysis

A status quo simulation projected CVD events, CVD deaths, and heart failure deaths, costs, and QALYs for nonHispanic black and white adults aged 35-74 years with untreated hypertension from 2014 to 2024. We then simulated CVD outcomes treating to JNC 8 BP targets in nonHispanic blacks and whites with subgroup analysis stratified by age groups (under and over 60 years) and status of CVD, CKD, or diabetes. Incremental cost-effectiveness ratios (ICERs) were calculated as change in costs divided by incremental change in QALYs. ICERs < \$50,000/QALY gained were considered cost-effective, $\geq \$ 50,000$ and $<\$ 150,000$ of intermediate value, and $\geq \$ 150,000$ of low value. ${ }^{29}$ All analyses were approached from a payer's perspective. Future costs and QALYs were discounted at 3\% per year. ICERs compared costs and costs savings associated with treating patients to

JNC 8 guidelines vs. no treatment within age groups and by CVD, diabetes, and CKD status.

In an exploratory analysis, we simulated alternative scenarios applying nonpharmacologic diet and exercise (lifestyle) interventions alone to lower BP in stage 1 hypertension without diabetes or CKD based on the upper and lower 95\% confidence interval bounds of a summary effectiveness estimate from lifestyle intervention trials. ${ }^{30}$ Reduction in total cardiovascular events due to lifestyle interventions was estimated, but cost analysis for lifestyle intervention was not done due to poor and inconsistent data available on the cost of these interventions.

## Multivariate, probabilistic sensitivity analysis

We did not perform 1-way sensitivity analyses since uncertainties surrounding each model input were explored extensively in our prior analysis. ${ }^{5}$ Uncertainty distributions of the main inputs, including clinical effectiveness of BP lowering, relative risk of CVD associated with BP change, quality of life decrements due to side effects, and costs related to side effects, medications, and monitoring, were randomly sampled 1,000 times in probabilistic simulations. From these results, we calculated $95 \%$ uncertainty intervals (UIs) for all model outputs.

## RESULTS

About 130,000 non-Hispanic blacks and 570,000 non-Hispanic whites with CVD and hypertension but not currently treated with medications would be eligible for secondary prevention every year during the 2014-2024. Treating these patients was estimated to prevent about 2,000 and 10,000 CVD events in black and white populations, respectively (Table 2). Additionally, 1.7 million non-Hispanic blacks and 5.4 million non-Hispanic whites aged $35-74$ years with hypertension were eligible for primary prevention treatment. Treating these patients to 2014 guideline targets was projected to prevent about 6,000 and 25,000 CVD events in non-Hispanic black and non-Hispanic white populations, respectively (Table 2 ).

Hypertension treatment was cost saving overall in both non-Hispanic blacks and non-Hispanic whites compared with no treatment (Table 2). Incremental addition of treatment of stage 1 hypertension for primary prevention to treatment of CVD and stage 2 patients was also cost-effective in both groups overall.

When treatment was assessed in progressively less costeffective strategies, cost savings were observed in a greater proportion of subgroups in the non-Hispanic black population compared with the non-Hispanic white adults. In non-Hispanic blacks, treating hypertension for primary prevention in males of all ages and BP stages was cost saving, regardless of diabetes or CKD status (Figure 1). Treatment was also cost saving in all non-Hispanic black women with stage 2 hypertension and in those aged $45-74$ years with stage 1 hypertension. Treating non-Hispanic black women aged 35-44 years with stage 1 hypertension was cost-effective both in those with diabetes or CKD ( $\$ 21,000 /$ QALY;

Table 1. Main assumptions and uncertainty ranges used for the comparative effectiveness analysis of US BP treatment guidelines

| Variable | Estimate (range in upper and lower bound estimates if a variation assumed according to age and/or sex) |  |  | Sources |
| :---: | :---: | :---: | :---: | :---: |
|  | Main | Lower | Upper |  |

Effectiveness
Average $R$ R per 5 mm Hg reduction in diastolic $B P$ or 10 mm Hg reduction in systolic $B P$, ages $35-59$ years $^{\text {a }}$

| CHD | $0.73(0.72-0.74)$ | $0.70(0.67-0.72)$ | $0.77(0.76-0.78)$ | Law, Morris, and Wald meta-analysis ${ }^{18}$ |
| :--- | :---: | :---: | :---: | :---: |
| Stroke | $0.64(0.61-0.66)$ | $0.59(0.54-0.63)$ | $0.69(0.68-0.69)$ |  |
| All-cause mortality | $0.86(0.83-0.89)$ | 0.76 | 0.95 |  |

Average $R$ R per 10 mm Hg reduction in systolic $B P$ or 5 mm Hg reduction in diastolic $B P$, ages $60-74$ years $^{\text {a }}$

| CHD | $0.77(0.74-0.78)$ | $0.74(0.72-0.76)$ | $0.79(0.76-0.81)$ |
| :--- | :---: | :---: | :---: | :---: |
|  |  |  |  |
| Stroke | $0.69(0.66-0.71)$ | $0.64(0.62-0.64)$ | $0.74(0.69-0.78)$ |
| All-cause mortality | $0.91(0.91-0.92)$ | 0.80 | 1.02 |

Average systolic BP-lowering effect ( mm Hg$)^{\text {b }}$

## Stage 2 hypertension

| Age $<60$ years, pretreatment $\geq$ | $31.0-34.7$ | $26.0-29.4$ | $36.0-39.9$ |
| :--- | :--- | :--- | :--- | 160 mm Hg target 140 mm Hg , 3-4 standard dose medications

Age $\geq 60$ years, pretreatment $\geq$ 160 mm Hg target 150 mm Hg , 2-3 standard dose medications
Stage 1 hypertension
Age <60 years, pretreatment $140-159 \mathrm{~mm} \mathrm{Hg}$ target 140 mm $\mathrm{Hg}, 0.5-2.0$ standard dose medications

Age $\geq 60$ years, pretreatment
7.1
3.2
11.0
$150-159 \mathrm{~mm}$ Hg target 150 mm $\mathrm{Hg}, 0.5$ standard dose medications

Diastolic BP-lowering effect $(\mathrm{mm} \mathrm{Hg})^{\text {b }}$

$$
\text { All ages, stage } 2
$$

17.1
12.0
22.2
aw, Morris, and Wald meta-analysis, ${ }^{18}$ Systolic Hypertension in the Elderly Program (SHEP) all-cause mortality upper bound $95 \%$ interval for upper estimate ${ }^{19}$

All ages (target $90 \mathrm{~mm} \mathrm{Hg}, 3$ standard dose medications

All ages, stage 1 hypertension 5.3 3.7
6.9

All ages (target 90 mm Hg ,
1 standard dose medication
Annual costs per person treated (2010 costs; inflated to 2014 costs in all results)
Physician office visit
Treatment monitoring visits (number)

Stage 2 hypertension

Stage 1 hypertension
Cost per routine monitoring visit Hospitalization

4

3
\$71

3

2
Not modeled

5 7 recommendation Outpatient visit, Medicare Physician Fee Schedule (code 99213, nonfacility limiting charge) ${ }^{20}$

Table 1. Continued

| Variable | Estimate (range in upper and lower bound estimates if a variation assumed according to age and/or sex) |  |  | Sources |
| :---: | :---: | :---: | :---: | :---: |
|  | Main | Lower | Upper |  |
| Average cost (used for infrequent hospitalized drug-related adverse events) | \$12,000 | Not modeled | Not modeled | National Inpatient Sample survey |
| High cost (used for rare hospitalized drug-related adverse events) | \$21,000 |  |  |  |
| Laboratory test (electrolytes monitoring on treatment) |  |  |  |  |
| Number of tests | 1 | 1 | 2 | U.S. Joint National Committee 7 recommendation Centers for Medicare and Medicaid laboratory fee schedule |
| Cost per test | \$15 | Not modeled | Not modeled |  |
| Antihypertensive drug costs (total daily doses) ${ }^{\text {c }}$ |  |  |  |  |
| 0.5 standard dose | \$124 | Not modeled | \$296 | Average wholesale prices reported by manufacturers ("Red Book"; 2010), ${ }^{21}$ see Methods text for estimation method |
| 1.0 standard dose | \$166 |  | \$363 |  |
| 1.5 standard doses | \$215 |  | \$409 |  |
| 2.0 standard doses | \$238 |  | \$567 |  |
| 3.0 standard doses | \$357 |  | \$850 |  |
| 3.5 standard doses | \$430 |  | \$1,311 |  |
| 4.0 standard doses | \$496 |  | \$1,374 |  |
| Pharmacy dispensing fees ${ }^{\text {d }}$ | \$27 |  | \$33 |  |
| Acute and chronic CVD treatment costs |  |  |  |  |
| Myocardial infarction hospitalization |  |  |  |  |
| Nonfatal | \$33,000 |  |  | California Office of Statewide Health Planning and Development (OSHPD) hospital data, $2008^{22}$ |
| Fatal | \$46,000 |  |  |  |
| Coronary revascularization procedures |  |  |  |  |
| Percutaneous coronary intervention | \$21,000-\$23,000 |  |  |  |
| Coronary artery bypass graft surgery | \$57,000-\$59,000 |  |  |  |
| Stroke |  |  |  |  |
| Fatal | \$21,000-\$26,000 |  |  |  |
| Nonfatal | \$15,000-\$21,000 |  |  |  |
| Chronic CHD costs |  |  |  |  |
| First year | \$11,000 |  |  | US Medical Expenditure Panel Surveys (MEPS), 1998-2008 |
| Subsequent years | \$2,000 |  |  |  |
| Chronic post-stroke costs |  |  |  |  |
| First year | \$16,000 |  |  |  |
| Subsequent years | \$5,000 |  |  |  |
| Inflation from 2010 to 2014 costs | 9\% |  | 11\% | Main = change in general US consumer price index; upper = change in medical component |

Serious adverse effects of medications (incidence per 100,000 person-years)
Common, outpatient management

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Table 1. Continued

| Variable | Estimate (range in upper and lower bound estimates if a variation assumed according to age and/or sex) |  |  | Sources |
| :---: | :---: | :---: | :---: | :---: |
|  | Main | Lower | Upper |  |
| 3 standard doses | 10,039.20 | 6,950.21 | 12,742.06 | Based on Law $2003{ }^{17}$ |
| 2 standard doses | 7,572.41 | 5,242.43 | 9,611.13 |  |
| 1 standard dose | 5,200.00 | 3,600.06 | 6,600.00 |  |
| 0.5 standard dose | 2,600.00 | 1,800.00 | 3,300.00 |  |
| Infrequent, hospitalized |  |  |  |  |
| 3 standard doses | 193.06 | 19.31 | 965.31 | Trials, medication labels, postmarketing reports |
| 2 standard doses | 145.62 | 14.56 | 728.12 |  |
| 1 standard dose | 100.00 | 10.00 | 500.00 |  |
| 0.5 standard dose | 50.00 | 5.00 | 250.00 |  |
| Rare, hospitalized/severe |  |  |  |  |
| 3 standard doses | 1.93 | 0.0193 | 19.31 |  |
| 2 standard doses | 1.46 | 0.0146 | 14.56 |  |
| 1 standard dose | 1.00 | 0.0100 | 10.00 |  |
| 0.5 standard dose | 0.50 | 0.0050 | 5.00 |  |
| Death |  |  |  |  |
| 3 standard doses | 0.0193 | 0.0002 | 0.1931 |  |
| 2 standard doses | 0.0146 | 0.0001 | 0.1456 |  |
| 1 standard dose | 0.0100 | 0.0001 | 0.1000 |  |
| 0.5 standard dose | 0.0050 | 0.0001 | 0.0500 |  |
| Utility (QALY weight penalty, duration) |  |  |  |  |
| Drug side effects managed as outpatient (1 day) | 0.23 |  |  | Montgomery ${ }^{23}$ |
| Drug side effect requiring hospitalization (1 day) | 0.50 |  |  | Clinical judgment |
| Acute stroke (1 month) | 0.86 |  |  | Global Burden of Disease 2010 Study ${ }^{24}$ |
| Chronic stroke survivors (12 months) | 0.85-0.88 |  |  |  |
| Acute myocardial infarction (1 month) | 0.91 |  |  |  |
| Acute unstable angina (1 month) | 0.95 |  |  |  |
| Chronic CHD (12 months) | 0.91-0.98 |  |  |  |
| Death | 1.00 |  |  |  |
| Disutility due to taking daily medications | 0.00 |  | 0.01-0.02 | Past cost-effectiveness analyses ${ }^{20,25,26}$ |
| Adherence to medications (percent of patients continuing prescribed treatment) | 75\% | $25 \%$ or $50 \%$ lower than observed in trials | Not modeled | Law, Morris, and Wald meta-analysis for main estimate ${ }^{18}$ |
| Annual discount rate | 3\% | Not modeled | Not modeled | Assumed |

Abbreviations: $\mathrm{BP}=$ blood pressure; $\mathrm{CHD}=$ coronary heart disease; $\mathrm{CVD}=$ cardiovascular disease; $\mathrm{QALY}=$ quality-adjusted life year.
${ }^{\text {a Relative risk reductions vary by age and sex category, see Methods and Supplementary Appendix for details. }}$
${ }^{\text {b }}$ Change in BP dependent on age- and sex-specific distribution of baseline BPs within stage 1 or stage 2 category and number of standard dose antihypertensive medications.
${ }^{\text {c Standard }}$ dose medications used to estimate costs: captopril 25 mg twice daily, nifedipine 30 mg daily, amlodipine 5 mg daily, hydrochlorothiazide 25 mg daily, and atenolol 50 mg daily. Combination medications equivalent to two-half standard doses were amlodipine/benazepril $2.5 / 10 \mathrm{mg}$ daily and captopril/hydrochlorothiazide $25 / 25 \mathrm{mg}$ daily. Combination medications equivalent to 1.5 standard doses were captopril/ hydrochlorothiazide $25 / 25 \mathrm{mg}$ once daily, benazepril/amlodipine $10 / 2.5 \mathrm{mg}$ daily, or hydrochlorothiazide/propranolol $25 / 80 \mathrm{mg}$ daily. Combination medications equivalent to 2 standard doses were atenolol/chlorthalidone $50 / 25 \mathrm{mg}$ daily and captopril/hydrochlorothiazide $50 / 25 \mathrm{mg}$ daily. Of the medications listed, only captopril (angiotensin converting enzyme inhibitor) taken divided doses ( 25 mg twice daily).

Table 2. Annual population treated and cost-effectiveness of implementing JNC 8 hypertension treatment guidelines in previously untreated non-Hispanic black (NHB) and nonHispanic white (NHW) adults with hypertension aged 35-74 years. Average annual results from a years 2014-2024 simulation, the CVD policy model.

| Strategy | Annual incremental results |  |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: |
|  | Number of people newly treated | CVD events averted compared with status quo | CVD costs compared with status quo (\$US, thousands) | ICER compared with status quo (\$US per QALY gained) | ICER compared with prior strategy (\$US per QALY gained) |
| NHB patients |  |  |  |  |  |
| NHB women ages 35-74 years |  |  |  |  |  |
| Treat CVD population | 35,000 | -1,000 (1,000; 1,000) | -48,000 (-70,000; -31,000) | Cost saving ${ }^{\text {a }}$ | Cost saving ${ }^{\text {a }}$ |
| Treat stage 2 along with CVD | 484,000 | -2,000 (2,000; 3,000) | -103,000 (-142,000; -73,000) | Cost saving ${ }^{\text {a }}$ | Cost saving ${ }^{\text {a }}$ |
| Treat stage 1 with diabetes or CKD, along with CVD and stage 2 | 703,000 | -3,000 (2,000; 3,000) | -112,000 (-157,000; -76,000) | Cost saving ${ }^{\text {a }}$ | Cost saving ${ }^{\text {a }}$ |
| Treat all hypertension | 855,000 | -3,000 (3,000; 4,000) | -115,000 (-164,000; -76,000) | Cost saving ${ }^{\text {a }}$ | Cost saving (cost saving; 4,000) |
| NHB men ages 35-74 years |  |  |  |  |  |
| Treat CVD population | 96,000 | -1,000 (1,000; 2,000) | -100,000 (-148,000; -67,000) | Cost saving ${ }^{\text {a }}$ | Cost saving ${ }^{\text {a }}$ |
| Treat stage 2 along with CVD | 592,000 | -4,000 (3,000; 5,000) | -258,000 (-336,000; -193,000) | Cost saving ${ }^{\text {a }}$ | Cost saving ${ }^{\text {a }}$ |
| Treat stage 1 with diabetes or CKD, along with CVD and stage 2 | 698,000 | -4,000 (4,000; 6,000) | -294,000 (-386,000; -219,000) | Cost saving ${ }^{\text {a }}$ | Cost saving ${ }^{\text {a }}$ |
| Treat all hypertension | 940,000 | -5,000 (4,000; 7,000) | -337,000 (-446,000; -244,000) | Cost saving ${ }^{\text {a }}$ | Cost saving ${ }^{\text {a }}$ |
| NHW patients |  |  |  |  |  |
| NHW women ages 35-74 years |  |  |  |  |  |
| Treat CVD population | 86,000 | -4,000 (3,000; 5,000) | -163,000 (-240,000; -105,000) | Cost saving ${ }^{\text {a }}$ | Cost saving ${ }^{\text {a }}$ |
| Treat stage 2 along with CVD | 785,000 | -10,000 (8,000; 12,000) | -286,000 (-431,000; -160,000) | Cost saving ${ }^{\text {a }}$ | Cost saving ${ }^{\text {a }}$ |
| Treat stage 1 with diabetes or CKD, along with CVD and stage 2 | 1,527,000 | -11,000 (9,000; 13,000) | -273,000 (-440,000; -130,000) | Cost saving ${ }^{\text {a }}$ | 6,000 (-4,000; 18,000) |
| Treat all hypertension | 2,559,000 | -13,000 (10,000; 15,000) | -203,000 (-399,000; -43,000) | Cost saving ${ }^{\text {a }}$ | 19,000 (9,000; 31,000) |
| NHW men ages 35-74 years |  |  |  |  |  |
| Treat CVD population | 482,000 | -6,000 (5,000; 8,000) | -455,000 (-702,000; -292,000) | Cost saving ${ }^{\text {a }}$ | Cost saving ${ }^{\text {a }}$ |
| Treat stage 2 along with CVD | 1,368,000 | -16,000 (12,000; 20,000) | -884,000 (-1,269,000; -597,000) | Cost saving ${ }^{\text {a }}$ | Cost saving ${ }^{\text {a }}$ |
| Treat stage 1 with diabetes or CKD, along with CVD and stage 2 | 1,961,000 | -19,000 (14,000; 24,000) | -964,000 (-1,399,000; -632,000) | Cost saving ${ }^{\text {a }}$ | Cost saving ${ }^{\text {a }}$ |
| Treat all hypertension | 3,436,000 | -22,000 (18,000; 28,000) | -971,000 (-1,473,000; -581,000) | Cost saving ${ }^{\text {a }}$ | Cost saving ${ }^{\text {a }}$ |

Abbreviations: CKD = chronic kidney disease; CVD = cardiovascular disease; ICER = incremental cost-effectiveness ratio; QALY = quality-adjusted life year.
a Upper and lower uncertainty interval bounds of ICERs were cost saving.
Non-Hispanic black

| Age group (years) | stage 2 | Males <br> stage 1 <br> diabetes <br> or CKD | stage 1 no diabetes or CKD |
| :---: | :---: | :---: | :---: |
| 35-44 | cost saving | cost saving | cost saving |
| 45-59 | cost saving | cost saving | cost saving |
| 60-74 | cost saving | cost saving | cost saving |


| Females <br> stage 1 <br> diabetes |  |  |
| :--- | ---: | ---: |
| stage 2 | stage 1 no <br> diabetes or |  |
| cost saving | $\$ 21,000$ | CKD |
| cost saving | cost saving | cost saving |
| cost saving | cost saving | cost saving |



Non-Hispanic white

| Age group (years) | stage 2 | Males <br> stage 1 <br> diabetes <br> or CKD | stage 1 no <br> diabetes <br> or CKD | stage 2 | Females <br> stage 1 <br> diabetes <br> or CKD | stage 1 no diabetes or CKD |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 35-44 | cost saving | \$5,000 | \$57,000 | \$38,000 | \$87,000 | \$181,000 |
| 45-59 | cost saving | cost saving | cost saving | cost saving | \$8,000 | \$21,000 |
| 60-74 | cost saving | cost saving | cost saving | cost saving | \$5,000 | \$7,000 |

cost saving<br>cost-effective (ICER < \$50,000)<br>intermediate value (ICER $\geq \$ 50,000$ and $<\$ 150,000$ )<br>low value (ICER $\geq \$ 150,000$ )



Figure 1. Incremental cost-effectiveness of implementing JNC 8 hypertension treatment guidelines in previously untreated non-Hispanic black (NHB) and non-Hispanic white (NHW) adults with hypertension aged 35-74 years, by age, sex, hypertension stage, and status of diabetes or chronic kidney disease (CKD). Within age and sex row, each group is compared with the higher risk group to its left; stage 2 hypertension is compared with patients living with cardiovascular disease (CVD) and hypertension. Average annual results from a years 2014-2024 simulation, the CVD policy model. Abbreviation: ICER = incremental cost-effectiveness ratio.

95\% UI \$3,000-\$43,000) and in those without diabetes or CKD (\$46,000/QALY; 95\% UI \$17,000-\$76,000).

Treating hypertension was cost saving in all non-Hispanic white men with stage 2 hypertension and those aged $45-74$ years with stage 1 hypertension. Treatment was costeffective in non-Hispanic white males aged 35-44 years with stage 1 hypertension and diabetes or CKD with an ICER of about $\$ 5,000 /$ QALY gained ( $95 \%$ UI from cost saving to $\$ 30,000$ ), but of only intermediate value in those with stage 1 hypertension but without diabetes or CKD (ICER of about \$57,000/QALY; 95\% UI \$15,000-\$100,000). In non-Hispanic white females, hypertension treatment was only cost saving in those 45-74 years old and with stage 2 hypertension. Treating non-Hispanic white women aged 35-44 years with stage 2 hypertension or those aged $45-74$ years with stage 1 hypertension was cost-effective (Figure 1). Treating hypertension in white women aged 35-44 years with stage 1 hypertension was of intermediate or low value (ICERS $\geq$ \$50,000/QALY; 95\% UI \$46,000-\$121,000 with diabetes or CKD, \$111,000-\$235,000 without diabetes or CKD).

For non-Hispanic whites with stage 1 hypertension but without diabetes or CKD, lifestyle interventions alone would result in about 1,000-3,000 CVD events avoided per year, less than the approximately 5,000 events prevented by pharmacologic treatment alone in the same patients according to JNC 8. For non-Hispanic blacks, lifestyle intervention
alone in stage 1 patients without diabetes or CKD would prevent $<1,000$ CVD events (about 300-700 fewer), less than the approximately 1,000 events that would be prevented by pharmacologic treatment alone.

## DISCUSSION

Implementation of 2014 hypertension treatment guideline goals would reduce CVD morbidity and mortality while saving costs overall in both Non-Hispanic whites and blacks with untreated hypertension. Compared with non-Hispanic whites, cost savings and cost-effectiveness extended to younger ages in non-Hispanic black patients with stage 1 hypertension. In the short term, implementing guideline goals would be cost-effective in 35 - to 44 -year-old non-Hispanic black females with stage 1 hypertension, but appeared to be of intermediate or low value in non-Hispanic white females of the same age and hypertension stage.

Over $41 \%$ of non-Hispanic blacks aged $\geq 18$ years have hypertension, more than 10 percentage points higher than other US race/ethnic groups. ${ }^{32}$ Awareness and treatment have improved in all US adults in the past 2 decades, ${ }^{1}$ but even now only about a half of non-Hispanic whites have controlled hypertension (50.3\%), and the proportion controlled is almost 9 percentage points lower in non-Hispanic blacks (41.4\%). ${ }^{33}$ Uncontrolled hypertension is a leading cause of death and
disability in the non-Hispanic black population, and nonHispanic black men have the highest rates of hypertensionrelated death (hypertensive heart disease, other heart diseases, and stroke) of any race/ethnic group. ${ }^{3}$ Hypertension treatment and control in non-Hispanic black adults is therefore an urgent national public health priority. Our results suggest that implementing hypertension treatment guidelines in younger non-Hispanic black adults is a high-value strategy for health system payers, yielding cost savings in older and higher risk patients, and cost-effectiveness in younger patients.

Clinical practice-based studies have demonstrated that intensive focus on hypertension treatment can lead to remarkable improvements in BP control. ${ }^{34}$ Investment in a teambased approach-supplementing usual care with intensive interactions between patient and a nonphysician health care provider-may achieve substantial gains. ${ }^{35}$ Non-Hispanic black men aged 21-54 years old in inner-city Baltimore who received educational-behavioral-pharmacologic intervention from a team including a nurse practitioner and community health worker had a $>10 \%$ higher proportion of controlled hypertension after a 36-month intervention, compared with those receiving usual care. ${ }^{36}$ A systematic clinic-based quality improvement intervention in the 15 Veteran's Administration medical centers achieved a $>30 \%$ point improvement in hypertension control over 10 years, with similar improvements in non-Hispanic black and white patients. ${ }^{37}$ A multicomponent intervention combining patient education, home BP monitoring, and lifestyle counseling in low-income non-Hispanic black patients ( $72 \%$ female) did not improve hypertension control compared with usual care-in part perhaps because the complexity of the intervention led to $<50 \%$ adherence to component interventions. ${ }^{38}$ The results of the SPRINT trial add to decades of evidence demonstrating that BP-lowering therapies save lives. ${ }^{18,39}$ Further research is needed on the most effective ways to deliver this highly effective and cost-effective therapy to all patients seen in clinical practice, including non-Hispanic black patients. Lifestyle interventions alone in stage 1 patients without diabetes or CKD would prevent a more modest number of CVD events compared with pharmacologic therapy alone; however, lifestyle measures are likely to augment prevention benefits when combined with pharmacologic treatment or serve as an alternative for patients who decline to take medications.

Due in part to documented barriers to accessing and accepting treatment, a significant proportion of non-Hispanic black patients do not regularly seek hypertension and other preventive care in medical clinics. ${ }^{3}$ Recent initiatives have tested the strategy of diagnosing and initiating treatment in non-Hispanic black adults outside of the clinic and in other community settings, including barbershops, community centers, and faith-based organizations. ${ }^{3}$ In the intervention arm of a trial in Dallas, barbers in 17 black-owned barbershops were trained to offer BP checks, deliver health education, and refer hypertensive patients to medical care. ${ }^{40}$ After 10 months, hypertension control was 8.8 percentage points higher in the intervention, compared with the usual care arm. The Georgia Stroke and Heart Attack Prevention Program (SHAPP) is a statewide initiative focused on hypertension screening and control in 16 predominantly rural public health districts that are about $56 \%$ non-Hispanic
black. Through provider training, affordable medications, adherence support, and community partnerships, ${ }^{41}$ SHAPP achieved approximately $60 \%$ hypertension control and was projected to save costs compared with no treatment. ${ }^{42}$ Our analysis provides further evidence supporting the value of investing in both clinic- and community-based interventions to improve hypertension control in non-Hispanic blacks and other high-risk groups with high hypertension prevalence.

The methods and reporting of this study conform to Consolidated Health Economic Evaluation Reporting Standards (CHEERS; see checklist in the Supplementary Text), ${ }^{43}$ and Quality of Health Economic Studies Instrument standards recommended for cost-effectiveness analyses of US CVD risk factor guidelines. ${ }^{29}$ Effectiveness assumptions were grounded in a large meta-analysis of randomized antihypertensive medication treatment trials and inputs for nonHispanic whites and blacks were estimated from national data. However, like all computer-simulation analyses, this analysis was limited by reliance on multiple assumptions and inputs derived from multiple data sources and study types. Because of our conservative approach, we did not fully capture disability and health care costs attributed to hypertensive heart disease or end-stage renal disease-both of which are more common in non-Hispanic blacks-so that we may have underestimated cost savings and QALY gains with hypertension treatment compared with no treatment and in between-group comparisons. Additionally, while the 2014 national guidelines recommended different first-line agents for hypertension control in non-Hispanic blacks (diuretics or calcium channel blockers) and whites (the same agents or renin-angiotensin system blockers), ${ }^{16}$ the BP response to different agents is similar in most patients, ${ }^{44}$ and we chose not to model differential use of antihypertensive medication classes in order not to bias cost-of-treatment inputs. Our projections of future CVD incorporated demographic trends but did not account for any secular or clinical practice changes in the next decade. Our analysis is further limited by its exclusion of other race/ethnic groups, including Hispanic Americans and Asian Americans due to the lack of highquality data on stroke and coronary heart disease incidence in these groups. Further research is needed to elucidate hypertension treatment cost-effectiveness in these groups as higher quality data becomes available.

Hypertension treatment cost savings and cost-effectiveness extend to younger ages among non-Hispanic black, compared with non-Hispanic white patients. Given their the higher hypertension prevalence and heavier burden of hypertension-related disease compared with any other US race/ethnic group, investment in effective clinic- and community-based hypertension control programs in nonHispanic black Americans is a particularly high-value investment in population health.

## SUPPLEMENTARY MATERIAL

Supplementary materials are available at American Journal of Hypertension (http://ajh.oxfordjournals.org).

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## DISCLOSURE

The authors declared no conflict of interest.

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