# ORIGINAL ARTICLE

# Projected Effect of Dietary Salt Reductions on Future Cardiovascular Disease

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# ABSTRACT

## BACKGROUND

The U.S. diet is high in salt, with the majority coming from processed foods. Reducing dietary salt is a potentially important target for the improvement of public health.

# METHODS

We used the Coronary Heart Disease (CHD) Policy Model to quantify the benefits of potentially achievable, population-wide reductions in dietary salt of up to 3 g per day (1200 mg of sodium per day). We estimated the rates and costs of cardiovascular disease in subgroups defined by age, sex, and race; compared the effects of salt reduction with those of other interventions intended to reduce the risk of cardiovascular disease; and determined the cost-effectiveness of salt reduction as compared with the treatment of hypertension with medications.

#### RESULTS

Reducing dietary salt by 3 g per day is projected to reduce the annual number of new cases of CHD by 60,000 to 120,000, stroke by 32,000 to 66,000, and myocardial infarction by 54,000 to 99,000 and to reduce the annual number of deaths from any cause by 44,000 to 92,000. All segments of the population would benefit, with blacks benefiting proportionately more, women benefiting particularly from stroke reduction, older adults from reductions in CHD events, and younger adults from lower mortality rates. The cardiovascular benefits of reduced salt intake are on par with the benefits of population-wide reductions in tobacco use, obesity, and cholesterol levels. A regulatory intervention designed to achieve a reduction in salt intake of 3 g per day would save 194,000 to 392,000 quality-adjusted life-years and \$10 billion to \$24 billion in health care costs annually. Such an intervention would be cost-saving even if only a modest reduction of 1 g per day were achieved gradually between 2010 and 2019 and would be more cost-effective than using medications to lower blood pressure in all persons with hypertension.

#### CONCLUSIONS

Modest reductions in dietary salt could substantially reduce cardiovascular events and medical costs and should be a public health target.

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This article (10.1056/NEJMoa0907355) was published on January 20, 2010, at NEJM.org.

N Engl J Med 2010. Copyright © 2010 Massachusetts Medical Society. THE U.S. DIET IS HIGH IN SALT. THE DEpartments of Agriculture and Health and Human Services recommend daily intake of less than 5.8 g of salt (2300 mg of sodium), with a lower target of 3.7 g of salt per day for most adults (persons over 40 years of age, blacks, and persons with hypertension).<sup>1</sup> Despite these guidelines, during the period from 2005 through 2006, the average man in the United States is estimated to have consumed 10.4 g of salt per day and the average woman 7.3 g per day amounts that exceed those in preceding years.<sup>2</sup>

Reducing dietary salt lowers blood pressure and the risk of cardiovascular disease.3,4 Accomplishing this reduction is challenging, however, in part because 75 to 80% of the salt in the U.S. diet comes from processed foods, not from salt added during food preparation or consumption.5,6 Many countries, including Japan, the United Kingdom, Finland, and Portugal, have reduced population-wide salt intake through a combination of regulations on the salt content in processed foods, labeling of processed and prepared foods, public education, and collaboration with the food industry.7 To explore the potential impact of a modest reduction in dietary salt on population health, we used the Coronary Heart Disease (CHD) Policy Model, a computer simulation of heart disease in U.S. adults 35 to 84 years old, and an extension of the model that is used to assess stroke. We estimated the effects in different segments of the U.S. population, compared these projections with the health benefits expected from a range of other public health and clinical interventions aimed at reducing cardiovascular disease, and analyzed the relative cost-effectiveness of salt reduction as compared with treatment of hypertension with medication.

#### METHODS

## STRUCTURE OF THE MODEL

The CHD Policy Model is a computer-simulation, state-transition (Markov cohort) model of the incidence and prevalence of CHD and of the mortality and costs associated with CHD in U.S. residents 35 years of age or older. The model has been used to describe trends in CHD and the effects of interventions intended to reduce the risk of CHD.<sup>8,9</sup> (For an overview of the model, see the Supplementary Appendix, available with the full text of this article at NEJM.org.) The model has three submodels: demographic-epidemiologic, bridge, and disease-history. The demographicepidemiologic submodel predicts the incidence of CHD and the rates of death due to causes other than CHD among persons without a history of CHD. The risk of CHD is categorized according to age, sex, and the following six factors: systolic blood pressure, use or nonuse of antihypertensive medications, smoking status, level of highdensity lipoprotein (HDL) cholesterol, level of low-density lipoprotein (LDL) cholesterol, and presence or absence of diabetes mellitus. For persons in whom CHD develops, the bridge submodel characterizes the initial CHD event and its sequelae for 30 days. The disease-history submodel then predicts the rate of subsequent CHD events and rates of death from CHD and of deaths not related to CHD among simulated subjects with CHD, with each category stratified according to age, sex, and history of events. The data in the CHD Policy Model are derived from national data sets and calibrated to national event-rate estimates.

In addition to using the standard CHD Policy Model of the entire U.S. population, we created race-specific versions of the model for the black and nonblack populations. We derived the racespecific distribution of risk factors for CHD from the National Health and Nutrition Examination Survey. Beta coefficients derived from the Framingham Heart Study and the Framingham Offspring Study were applied to all three population groups, but the overall age-specific incidence rates of CHD were specific to each population.<sup>10</sup> The average incidence rates for the black and nonblack subpopulations were validated with national data.11 In sensitivity analyses we also examined beta coefficients specific to the black population.<sup>12</sup> We did not assign a coefficient to use of antihypertensive medications; rather, we used systolic blood pressure or treatment with antihypertensive medications to define the population of persons with hypertension, who might have a greater response to a reduction in dietary salt. Finally, we extended the model to estimate the incidence of stroke using beta coefficients derived from the Framingham Heart Study and the Framingham Offspring Study and published rates on the incidence of stroke.13,14

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#### MODELING APPROACH AND UNDERLYING ASSUMPTIONS

We assumed that the effect of salt reduction on blood pressure reduction was linear over the range of 0 to 3 g per day.<sup>15</sup> We used a lower estimate for the effect of salt reduction on systolic blood pressure on the basis of the findings of a large metaanalysis<sup>3,16</sup> and used a higher estimate for this effect on the basis of data from clinical trials.<sup>17,18</sup> We modeled an accentuated response to salt reduction among blacks, persons with hypertension, and persons 65 years of age or older (Table 1).<sup>17,19-22</sup> We compared the effects of a salt-restricted diet on numbers of CHD events with the effects of other interventions aimed at reducing cardiovascular risk by modeling a 50% reduction in smoking and environmental tobacco exposure,23 a 5% reduction in body-mass index among obese adults,8 treatment of persons at low or intermediate risk with statins in accordance with the guidelines in the Adult Treatment Panel III of the National Cholesterol Education Program,9 and treatment of hypertension as described in the Antihypertensive and Lipid-Lowering Treatment to Prevent Heart Attack Trial (ALLHAT; ClinicalTrials .gov number, NCT00000542).24,25

We conducted simulations for the entire U.S. population and for black and nonblack subgroups and estimated annual reductions in the incidence of CHD, new and recurrent myocardial infarctions, the incidence of stroke, and death from any cause as a result of reductions in dietary salt for the entire population and for subgroups defined according to age, sex, and race. We projected annual cost savings in health care and annual gains in quality-adjusted life years (QALYs) that would result from an intervention to reduce salt for the entire U.S. population and for Medicare enrollees, on the basis of the World Health Organization estimate that the cost of such a national effort would be \$1 per person annually.<sup>26</sup> We made the same projections for population-wide treatment of hypertension with antihypertensive medications.<sup>25</sup> We also report the cumulative costs and effectiveness for a gradual introduction of the intervention over the decade from 2010 through 2019.

# SENSITIVITY ANALYSES

We used Monte Carlo simulations to estimate the uncertainty of our projections for both the high and low estimates of the effects of salt reduction on systolic blood pressure. Beta coefficients for the association of systolic blood pressure, LDL and HDL cholesterol, and diabetes with both CHD events and deaths not associated with CHD were assumed to have a normal probability distribution, with standard errors derived from the fitted regression. We generated covariance matrixes for each of these beta coefficients. On the basis of the evidence of minimal correlation between factors, we assumed that the effects were independent. For each simulation, we report the mean

Group	Salt Reduct	ion, 1 g/day	Salt Reduct	ion, 3 g/day	Reference No.
	Low Estimate of SBP Decrease	High Estimate of SBP Decrease	Low Estimate of SBP Decrease	High Estimate of SBP Decrease	
		mm	n Hg		
Entire U.S. population					
Persons with hypertension†	1.20	1.87	3.60	5.61	3, 15
Persons ≥65 yr old	1.20	1.87	3.60	5.61	17, 19–22
All others	0.60	1.17	1.80	3.51	3, 15
Black subpopulation					
Persons with hypertension†	1.80	3.03	5.40	9.10	3, 17, 19–22
Persons ≥65 yr old	1.20	1.87	3.60	5.61	17, 19–22
All others	1.20	1.87	3.60	5.61	17, 19–22

\* SBP denotes systolic blood pressure.

† Hypertension was defined as a systolic blood pressure of 140 mm Hg or higher, a diastolic blood pressure of 90 mm Hg or higher, or use of an antihypertensive medication.

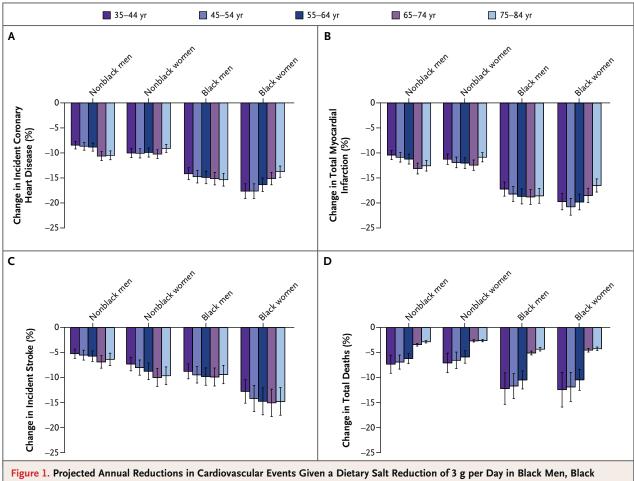
(±SE) for 1000 simulations. We conducted sensitivity analyses in which we varied the effect of salt reduction on changes in cardiovascular risk on the basis of estimates suggesting that the risk of cardiovascular disease associated with blood pressures lowered by means of salt reduction or medication is not as low as the risk associated with similar, naturally occurring blood pressures.<sup>27,28</sup>

# RESULTS

A population-wide reduction in dietary salt of 3 g per day (1200 mg of sodium per day) is projected to reduce the annual number of new cases of CHD by 60,000 to 120,000, stroke by 32,000 to 66,000, and myocardial infarction by 54,000 to 99,000

and to reduce the annual number of deaths from any cause by 44,000 to 92,000. Since the relationship between reductions in salt and the projected declines in event rates is linear over the range examined, even a more modest reduction of 1 g of salt per day is projected to result in large declines in annual rates of cardiovascular events and deaths (with new cases of CHD declining by 20,000 to 40,000, new and recurrent cases of myocardial infarction by 18,000 to 35,000, new cases of stroke by 11,000 to 23,000, and deaths from any cause by 15,000 to 32,000).

All adult age groups, both sexes, and blacks and nonblacks would be expected to benefit from reductions in salt intake (Fig. 1). The anticipated relative benefits among blacks would be greater



Women, Nonblack Men, and Nonblack Women, According to Age Group. Panel A shows changes in the incidence of coronary heart disease, Panel B changes in the rates of new and recurrent myocardial infarc-

tions, Panel C changes in the incidence of stroke, and Panel D changes in the rate of death from any cause. The projections are based on a reduction in dietary salt of 3 g per day and on the high estimate for the effect of salt reduction on systolic blood pressure. Total myocardial infarction includes new and recurrent myocardial infarctions. I bars indicate standard errors of the Monte Carlo simulation.

than those among nonblacks across all age groups and both sexes. The projected reductions in stroke would be greater among women than among men, with rates decreasing by 9 to 15% among black women and by 5 to 9% among nonblack women. All age groups would be expected to benefit, with middle-aged and older populations projected to have larger relative reductions in the incidence of CHD and in rates of new and recurrent myocardial infarction and stroke. A large relative reduction in mortality is projected for young and middle-aged adults, with mortality rates among blacks between 35 and 64 years of age reduced by 7 to 11% and those among nonblacks in this age range reduced by 3 to 6%. blood pressure without such intervention, the expected health benefit of salt reduction will be decreased (Table 2). If persons 65 years of age or older have the same degree of salt sensitivity as those younger than 65 years of age, the total estimated benefits of salt reduction would also be somewhat decreased. If blacks have no greater salt sensitivity than nonblacks, the magnitude of the anticipated effects on blacks will be reduced, but there will still be greater reductions in cardiovascular events and deaths among blacks because they have a higher prevalence of hypertension.

## COMPARISON WITH OTHER INTERVENTIONS

#### SENSITIVITY ANALYSES

If a lower blood pressure achieved through reduced salt intake is not as advantageous as the same

In a projection assuming a reduction in dietary salt of 3 g per day calculated using the high estimate for the effects of salt reduction on systolic blood pressure, the expected reductions in cardiovascular events would be similar in magnitude to

 Table 2. Projected Estimates of Annual Reductions in Rates of Cardiovascular Disease with a Dietary Salt Reduction of 3 g per Day,

 in the Main Simulation and According to Various Assumptions about Differential Salt Sensitivity in the Sensitivity Analyses.\*

Group	Incidence	e of CHD	Total	ΜI†	Incidence	of Stroke	Death from	Any Cause
	Low	High	Low	High	Low	High	Low	High
	Estimate	Estimate	Estimate	Estimate	Estimate	Estimate	Estimate	Estimate
		reduction	in rate per 10,	000 persons (%	6)‡			
Main simulation								
U.S. population	4.7±0.4	8.3±0.8)	3.7±0.3	6.2±0.6	2.4±0.3	3.9±0.5	3.3±0.5	5.4±0.8
	(6.1)	(10.7)	(7.7)	(12.8)	(5.2)	(8.2)	(2.7)	(4.4)
Nonblack population subgroup	4.3±0.4	7.0±0.6	3.4±0.3	5.3±0.4	2.2±0.3	3.4±0.5	3.1±0.4	4.9±0.7
	(6.1)	(9.8)	(7.7)	(12.0)	(5.1)	(8.0)	(2.7)	(4.3)
Black population subgroup	7.9±0.7	12.6±1.0	5.8±0.5	9.3±0.7	4.2±0.5	6.7±0.9	5.3±0.7	8.5±1.2
	(9.8)	(15.8)	(11.8)	(18.7)	(7.7)	(12.4)	(4.4)	(7.0)
Sensitivity analyses								
U.S. population								
Diminished risk reduction with	2.8±0.3	4.6±0.4	2.5±0.2	4.0±0.3	1.6±0.2	2.6±0.3	2.3±0.3	3.6±0.5
blood-pressure lowering∬	(4.1)	(6.6)	(5.2)	(8.3)	(3.5)	(5.5)	(1.8)	(2.9)
No increased salt sensitivity	4.5±0.4	7.3±0.6	3.5±0.3	5.6±0.5	2.2±0.3	3.6±0.5	3.2±0.4	5.2±0.7
with advanced age	(5.9)	(9.6)	(7.2)	(11.5)	(4.7)	(7.6)	(2.5)	(4.1)
Black subpopulation								
No increased salt sensitivity	5.2±0.5	8.2±0.7	4.0±0.3	6.1±0.5	2.8±0.4	4.4±0.6	3.5±0.5	5.5±0.8
	(6.5)	(10.3)	(8.0)	(12.4)	(5.2)	(8.1)	(2.9)	(4.6)
Black race–specific beta co-	6.4±2.8	10.0±4.2	4.6±2.0	7.0±2.9	4.2±0.6	6.8±0.9	4.7±1.2	7.4±1.8
efficients¶	(8.0)	(12.5)	(9.0)	(13.9)	(7.7)	(12.5)	(3.8)	(6.1)

\* Plus-minus values are means ±SE from the Monte Carlo simulations. CHD denotes coronary heart disease, and MI myocardial infarction. Results based on a dietary salt reduction of 1 g per day are available with the full text of this article at NEJM.org.

† Total myocardial infarction includes new and recurrent myocardial infarctions.

‡ Rates for the nonblack and black populations were adjusted in accordance with the age of the U.S. population.

If The cardiovascular benefit for a person with blood pressure that was lowered by reducing dietary salt was assumed to be equivalent to two thirds of the benefit for a person whose natural blood pressure was at that level.<sup>27,28</sup>

¶ The black race-specific beta coefficients used for all CHD risk factors are based on a published analysis from the Atherosclerosis Risk in Communities study.<sup>12</sup>

or greater than those projected for interventions targeting tobacco, obesity, primary prevention with statins, and pharmacologic treatment of hypertension, based on simulations for the same time frame and overall population (Table 3). For example, a reduction in dietary salt of 3 g per day would have approximately the same effect on rates of CHD events as a 50% reduction in tobacco use, a 5% reduction in body-mass index among obese adults, or the use of statins to treat persons at low or intermediate risk for CHD events. Salt reduction would have a much greater benefit with respect to stroke prevention than these other interventions. A population-wide reduction of salt intake of 3 g per day is expected to result in the same reduction in death rates as the use of medical treatment to control hypertension in all persons with the condition.

3 g per day would result in an estimated annual gain of 194,000 to 392,000 QALYs and estimated savings of \$10 billion to \$24 billion in health care costs. Even if salt targets were achieved gradually over the years 2010 through 2019 (Table 4) and a reduction in dietary salt of only 1 g were achieved only by the end of the decade, such an intervention is expected to result in cost savings. A saltreduction strategy is projected to compare favorably with the provision of antihypertensive therapy for all persons with hypertension — a strategy that would result in more QALYs gained but at a cost of \$6,000 to \$26,000 for each additional QALY. Even if the federal government were to bear the entire cost of a regulatory program designed to reduce salt consumption, the government would still be expected to realize cost savings for Medicare, saving \$6 to \$12 in health care expenditures for each dollar spent on the regulatory program. The strategy of providing antihypertensive medications for all persons with hy-

## COST-EFFECTIVENESS

A national effort to decrease salt consumption by

Table 3. Projected Estimates of C	Comparative Effect of Vario	us Population Interventions	on Annual Reductions in O	Cardiovascular Events.*
Intervention	Incidence of CHD	Total MI†	Incidence of Stroke	Death from Any Cause
	redi	uction in absolute number of e	vents (% change from expe	cted)
Salt reduction				
l g/day				
Low estimate	22,000±2000 (2.0)	20,000±1800 (2.6)	13,000±1800 (1.7)	17,000±2400 (0.9)
High estimate	37,000±3300 (3.3)	32,000±2900 (4.2)	20,000±2900 (2.7)	28,000±3800 (1.4)
2 g/day				
Low estimate	44,000±4000 (4.0)	39,000±3500 (5.1)	25,000±3500 (3.4)	34,000±4600 (1.7)
High estimate	71,500±6300 (6.4)	62,500±5400 (8.1)	40,000±5400 (5.3)	55,000±7500 (2.8)
3 g/day				
Low estimate	66,000±5800 (5.9)	58,000±5100 (7.6)	37,000±5100 (5.0)	51,000±7100 (2.6)
High estimate	110,000±9200 (9.6)	92,000±7800 (12.0)	59,000±8100 (7.8)	81,000±11,000 (4.1)
Smoking cessation;	41,000±10,000 (3.7)	92,000±14,000 (11.9)	32,000±13,000 (4.4)	84,000±9300 (4.3)
Weight loss§	59,000±3500 (5.3)	61,000±3200 (8.0)	5600±600 (0.7)	36,000±2000 (2.0)
Statin therapy for primary pre- vention¶	52,000±5600 (5.3)	17,000±1800 (2.9)	6600±200 (0.9)	5400±540 (0.3)
Pharmacologic treatment of hypertension	100,000±11,000 (9.3)	100,000±9700 (13.1)	69,000±11,000 (9.3)	80,000±10,000 (4.1)

\* Plus-minus values are means ±SE from the Monte Carlo simulations. CHD denotes coronary heart disease, and MI myocardial infarction. † Total myocardial infarction includes new and recurrent myocardial infarctions.

 $m \pm$  Smoking cessation was defined as elimination of 50% of use of or exposure to tobacco.

Weight loss was defined as a 5% reduction in body-mass index in an obese adult.

Pcholesterol treatment for primary prevention was defined according to the National Cholesterol Education Program Adult Treatment Panel III guidelines as treatment with statins in persons with a 10-year risk of CHD of less than 20%.

Pharmacologic treatment of hypertension was defined on the basis of treatment of all persons with hypertension to the degree described in the Antihypertensive and Lipid-Lowering Treatment to Prevent Heart Attack Trial.<sup>24</sup>

Table 4. Projected Estimates of the Cost and Effectiveness of Salt Reduction and Hypertension Treatment Annually and Cumulatively, 2010–2019.*	of the Cost an	d Effectiveness of	f Salt Reduction	ו and Hypertensi	on Treatment A	nnually and Cur	nulatively, 2010–2	2019.*		
Intervention			U.S. Population				Med	Medicare Population	5	
	Cost of Intervention (billions of dollars)	Reduction in Health Care Costs (billions of dollars)†	Gain in QALYs (thousands)	Cost per QALY Gained (dollars)‡	Cost Saved per Dollar Spent on Intervention (dollars)‡	Cost of Intervention (billions of dollars)	Reduction in Health Care Costs (billions of dollars)†	Gain in QALYs (thousands)	Cost per QALY Gained (dollars)‡	Cost Saved per Dollar Spent on Intervention (dollars)‡
Reduction in dietary salt										
1 g/day										
Low estimate	0.3§	4.1±0.8	75±9	Cost savings	15.4±3.0	0.3	0.7±0.1	46±4	Cost savings	2.5±0.2
High estimate	0.3§	7.0±1.4	120±15	Cost savings	26.1±5.2	0.3	1.0±0.1	72±6	Cost savings	3.8±0.4
3 g/day										
Low estimate	0.3§	12.1±2.4	220±26	Cost savings	45.2±9.1	0.3	2.0±0.2	135.0±12	Cost savings	7.3±0.7
High estimate	0.3§	20.4±4.1	350±42	Cost savings	76.0±15.4	0.3	3.0±0.3	208.3±19	Cost savings	11.1±1.1
Hypertension treatment	19.5±0.1	$14.2\pm 2.7$	360±42	15,800±9,900	0.7±0.1	9.3±0.03	3.4±0.3	260±24	23,300±3,600	0.4±0.04
Gradual reduction in dietary salt, 2010–2019										
1 g/day										
Low estimate	2.7§	18.9±3.8	220±27	Cost savings	7.0±1.4	2.7	4.3±0.4	220±20	Cost savings	1.6±0.2
High estimate	2.7§	31.6±6.5	350±43	Cost savings	11.8±2.4	2.7	6.1±0.6	240±21	Cost savings	2.3±0.2
3 g/day										
Low estimate	2.7§	56.9±11.5	650±78	Cost savings	21.2±4.3	2.7	$12.1\pm 1.2$	420±37	Cost savings	4.5±0.5
High estimate	2.7§	95.6±19.6	1000±127	Cost savings	35.6±7.3	2.7	18.5±1.9	665±58	Cost savings	6.9±0.7
<ul> <li>Plus-minus values are means ±SE from the Monte Carlo simulations. QALY denotes quality-adjusted life years.</li> <li>The reduction in health care costs is for the U.S. population of persons 35 years of age or older. Costs were discounted at 3% over the course of the decade.</li> <li>These values represent the ratio of the cost of the intervention in dollars to the number of QALYs gained as a result of the intervention. In some cases this calculation results in a negative number because the savings in health care costs as a result of the intervention are greater than the total cost of the intervention is described as cost savings. The column "Cost saved per dollar spent on the intervention" provides an estimate of the magnitude of these savings.</li> <li>The cost of a population-wide regulatory approach to salt reduction is estimated at \$1 per person per year, discounted at 3% over the course of the decade, according to the World Health Organization, and the total U.S. population was 306,913,687 persons as of July 2009, according to the U.S. Census.</li> <li>Hypertension treatment was defined on the basis of treatment of all persons with hypertension to the degree described in the Antihypertensive and Lipid-Lowering Treatment to Prevent Heart Attack Tria<sup>124</sup>; the cost-effectiveness analysis was also based on the results of the total reduction active of the total reduction accurs a cumulative effect, with one third of the total reduction accurse and the intervention to the foral reduction accurse of the final reatment to Prevent Heart Attack Tria<sup>124</sup>; the cost-effectiveness analysis was also based on the results of the total reduction accurse of the final third in 2012, another third in 2019.</li> </ul>	$s \pm SE$ from th costs is for th costs is for th stito of the cost ings in health Cost saved pe e regulatory al e total U.S. po defined on th the cost-effec 2010 to 2019 r	e Monte Carlo si e U.S. populatior t of the intervent care costs as a ru care costs as a ru er dollar spent on pproach to salt re pulation was 306 e basis of treatm tiveness analysis epresents a cum	mulations. QA n of persons 35 ion in dollars t ssult of the intreventi cduction is esti ,913,687 perso ent of all perso was also baset ulative effect, v	Carlo simulations. QALY denotes quality-adjusted life years. pulation of persons 35 years of age or older. Costs were discounted at netevention in dollars to the number of QALYs gained as a result of the its as a result of the intervention are greater than the total cost of the in spent on the intervention" provides an estimate of the magnitude of the to salt reduction is estimated at \$1 per person per year, discounted at 3 was 306,913,687 persons as of July 2009, according to the U.S. Census. of treatment of all persons with hypertension to the degree described in analysis was also based on the results of this trial. <sup>25</sup> is a cumulative effect, with one third of the total reduction achieved in 2.	ty-adjusted life older. Costs we QALYs gained atter than the tre atter than the tre estimate of the person per yeau 9, according to 19, according to 19, according to 19, according to 19, according to 10, according to 10, according to 11, according to 11, according to 11, according to 12, according to 12, according to 13, according to 14, accord	years. re discounted a: re discounted a: as a result of the tral cost of the ii magnitude of th magnitude of th discounted at the U.S. Censu the U.S. Censu sree described in	Carlo simulations. QALY denotes quality-adjusted life years. opulation of persons 35 years of age or older. Costs were discounted at 3% over the course of the decade. intervention in dollars to the number of QALYs gained as a result of the intervention. In some cases this calculation results in a nega intervention in dollars to the number of QALYs gained as a result of the intervention itself. In these cases, the intervention is described spent on the intervention are greater than the total cost of the intervention itself. In these cases, the intervention is described spent on the intervention" provides an estimate of the magnitude of these savings. to salt reduction is estimated at \$1 per person per year, discounted at 3% over the course of the decade, according to the World was 306,913,687 persons as of July 2009, according to the U.S. Census. of treatment of all persons with hypertension to the degree described in the Antihypertensive and Lipid-Lowering Treatment to analysis was also based on the results of this trial. <sup>25</sup> ts a cumulative effect, with one third of the total reduction achieved in 2012, another third in 2015, and the final third in 2019.	rse of the deca some cases thi In these cases, se of the decad isive and Lipid- rd in 2015, and	de. s calculation res the intervention e, according to t Lowering Treatrr the final third in	ults in a nega- i is described :he World nent to 1 2019.

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pertension would still be cost-effective if it were added to a successful program of population-wide dietary salt reduction, but the number of persons requiring treatment with such medications would be markedly reduced; the number of women with hypertension would be reduced by 16 to 24%, and the number of men by 22 to 34%, with a savings of \$3 billion to \$6 billion annually in the cost of treating hypertension.

#### DISCUSSION

Despite evidence linking salt intake to hypertension and cardiovascular disease, dietary salt intake in the U.S. is on the rise. This worsening trend has led to calls for population-wide interventions to reduce salt in the U.S. diet<sup>29</sup> — interventions already adopted in other countries.7 Our findings provide evidence to support these calls for action. Our postulated dietary reduction of 3 g of salt per day, which is within the range targeted by other developed countries, is projected to benefit the entire U.S. population and yield substantial reductions in morbidity, mortality, and health care costs. The population-wide benefits of salt reduction are similar in magnitude to the benefits that would accrue from other public health and clinical interventions and would reduce costs even if only a more modest reduction of 1 g of salt per day were achieved gradually over time.

Changes in behavior are notoriously difficult to achieve, and attempts to lower dietary salt intake on an individual basis have largely proved to be ineffective.<sup>5</sup> Nevertheless, cholesterol levels in the U.S. population fell before the widespread use of cholesterol-lowering medications, and smoking rates have fallen substantially as the result of a combination of regulatory, public health, and individual approaches to smoking cessation. The large and growing burden of hypertension, despite improved medical therapies,<sup>30</sup> and increased awareness that dietary salt reduction can help prevent and treat hypertension reinforce the urgent need for dietary change.

There is a considerable body of literature linking higher salt intake with higher blood pressure and increased cardiovascular risk,<sup>16,31</sup> and randomized trials have shown that a diet that is lower in salt reduces both blood pressure<sup>17,32</sup> and cardiovascular risk.<sup>31</sup> Despite concerns about the accurate assessment of salt intake, the difficulty of adhering to a low-salt diet, and the theoretical

increase in the risks associated with diets that are very low in salt, several large meta-analyses and reports from the Institute of Medicine<sup>3,5,16,27,34</sup> have concluded that reducing dietary salt would lower blood pressure and reduce cardiovascular risk. Professional societies, including the American Medical Association, the American Heart Association, the American Society of Hypertension, and the World Health Organization, have endorsed population-wide efforts to reduce salt intake.

The results of our study are similar to those of other analyses<sup>35,36</sup> and extend the findings of these analyses in important ways. We incorporated updated information on distributions of the prevalence of cardiovascular risk factors, particularly hypertension, in the entire U.S. population and in the black and nonblack subpopulations. We considered current levels of hypertension treatment, treatment and control of other cardiovascular risk factors, and competing and ongoing risks among persons in whom death was averted. Our expectations concerning the cardiovascular benefits of salt reduction are similar to those anticipated for established public health targets such as reducing tobacco use, obesity, and levels of LDL cholesterol. Targeted interventions have very large per-person effects, but their benefits are restricted to the smaller numbers of affected persons who are at increased risk. Lowering salt in the U.S. diet would result in small but measurable reductions in blood pressure across the entire U.S. population, thereby reducing rates of cardiovascular disease among all adults at risk.

A national regulatory effort to lower dietary salt intake would be cost-saving even if only a modest salt reduction were achieved after a decade. If a population-wide approach to lowering salt intake were sponsored by the federal government, the savings in expenditures for Medicare, the major federally sponsored health care program, would be greater than the cost of the regulatory intervention itself, even without the incremental benefits for younger persons not covered by Medicare. Some costs, such as those borne by the food industry in reformulating processed foods, are not considered in these analyses. However, as salt intake is reduced, people appear to prefer food with less salt,<sup>15</sup> a phenomenon that is probably related to the accommodation of taste receptors over the course of weeks to months.37 In the United Kingdom, a population-wide reduction in dietary salt of 10% was achieved in 4 years<sup>38</sup> without a reduction in sales of the food products included in the initial effort and without consumer complaints about taste. The magnitude of the health benefit suggests that salt should be a regulatory target of the Food and Drug Administration, which currently designates salt as a food additive that is "generally regarded as safe."<sup>29</sup>

We projected that similar levels of salt reduction may be of proportionately greater benefit to certain subpopulations. Blacks have high rates of hypertension and cardiovascular diseases that contribute to racial disparities in mortality<sup>39</sup>; the benefits of a diet lower in salt could narrow these disparities. Similarly, women would have a proportionately greater benefit than men because women have a higher lifetime risk of stroke.<sup>11</sup> Young and middle-aged adults could benefit because of the relative importance of blood-pressure elevations in younger adults without other major risk factors for cardiovascular disease. Blood-pressure elevations in young adulthood increase the likelihood that atherosclerosis9 and other illnesses will develop by middle age,<sup>40</sup> yet younger adults with hypertension are less likely to receive treatment for it.<sup>41</sup> The benefits of salt reduction may be even greater than we have projected if hypertension is completely prevented or its onset delayed by lowering salt intake even earlier, during childhood and adolescence.42

Projections such as ours are limited by any uncertainty concerning the data entered into the model. We modeled the effects of salt reduction on blood pressure in accordance with published data and assumed that the health benefits of salt reduction were mediated through these bloodpressure reductions. We did not account fully for possible effects of salt reduction that are unrelated to control of blood pressure — for exam-

ple, potential improvements in outcomes for the increasing numbers of patients with heart failure or prevention of other serious conditions, such as end-stage renal disease. Our estimates of the differential effects of salt reduction according to age and race were extrapolated from clinical-trial data, and there is uncertainty about these effects on the total population; however, sensitivity analyses suggest that our primary findings are not dependent on variations in these assumptions. We modeled only the linear effects of reduced salt intake on reductions in blood pressure. It has been suggested that these effects may be nonlinear,17 with greater reductions in blood pressure at lower levels of salt intake; such an assumption would result in larger reductions in cardiovascular disease than we present here.

Even with these limitations, our simulations suggest that modest reductions in dietary salt would yield substantial health benefits across the U.S. population of adults by lowering rates of cardiovascular events and death and reducing medical costs. Our findings underscore the need for an urgent call to action that will make it possible to achieve these readily attainable cardiovascular benefits.

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#### REFERENCES

2. U.S. Department of Agriculture, Agricultural Research Service. 2008. Nutrient intakes from food: mean amounts consumed per individual, one day, 2005– 2006. (Accessed January 15, 2010, at http://www.ars.usda.gov/SP2UserFiles/ Place/12355000/pdf/0506/usual\_nutrient\_

intake\_vitD\_ca\_phos\_mg\_2005-06.pdf.) 3. He FJ, MacGregor GA. Effect of longer-term modest salt reduction on blood pressure. Cochrane Database Syst Rev 2004;3:CD004937.

4. Panel on Dietary Reference Intakes for Electrolytes and Water, Standing Committee on the Scientific Evaluation of Dietary Reference Intakes, Food and Nutrition Board, Institute of Medicine. Dietary reference intakes for water, potassium, sodium, chloride, and sulfate. Washington, DC: National Academies Press, 2005.

5. Hooper L, Bartlett C, Davey SG, Ebrahim S. Advice to reduce dietary salt for prevention of cardiovascular disease. Cochrane Database Syst Rev 2004;1:CD003656. **6.** Mattes RD, Donnelly D. Relative contributions of dietary sodium sources. J Am Coll Nutr 1991;10:383-93.

**7.** He FJ, Macgregor GA. A comprehensive review on salt and health and current experience of worldwide salt reduction programmes. J Hum Hypertens 2008;23: 363-84.

**8.** Bibbins-Domingo K, Coxson P, Pletcher MJ, Lightwood J, Goldman L. Adolescent overweight and future adult coronary heart disease. N Engl J Med 2007;357: 2371-9.

9. Pletcher MJ, Lazar L, Bibbins-Domin-

<sup>1.</sup> Application of lower sodium intake recommendations to adults — United States, 1999–2006. MMWR Morb Mortal Wkly Rep 2009;58:281-3.

go K, et al. Comparing impact and costeffectiveness of primary prevention strategies for lipid-lowering. Ann Intern Med 2009;150:243-54.

**10.** D'Agostino RB, Grundy S, Sullivan LM, Wilson P. Validation of the Framingham coronary heart disease prediction scores: results of a multiple ethnic groups investigation. JAMA 2001;286:180-7.

Lloyd-Jones D, Adams R, Carnethon M, et al. Heart disease and stroke statistics — 2009 update: a report from the American Heart Association Statistics Committee and Stroke Statistics Subcommittee. Circulation 2008;119(3):e21-e181.
 Chambless LE, Heiss G, Shahar E, Earp MJ, Toole J. Prediction of Ischemic Stroke Risk in the Atherosclerosis Risk in Communities Study. Am J Epidemiol 2004;160:259-69.

**13.** Williams GR, Jiang JG, Matchar DB, Samsa GP. Incidence and occurrence of total (first-ever and recurrent) stroke. Stroke 1999;30:2523-8.

**14.** Kissela B, Schneider A, Kleindorfer D, et al. Stroke in a biracial population: the excess burden of stroke among blacks. Stroke 2004;35:426-31.

**15.** He FJ, MacGregor GA. How far should salt intake be reduced? Hypertension 2003;42:1093-9.

16. Idem. Effect of modest salt reduction on blood pressure: a meta-analysis of randomized trials: implications for public health. J Hum Hypertens 2002;16:761-70.
17. Sacks FM, Svetkey LP, Vollmer WM, et al. Effects on blood pressure of reduced dietary sodium and the Dietary Approaches to Stop Hypertension (DASH) diet. N Engl J Med 2001;344:3-10.

**18.** MacGregor GA, Markandu ND, Sagnella GA, Singer DR, Cappuccio FP. Double-blind study of three sodium intakes and long-term effects of sodium restriction in essential hypertension. Lancet 1989;2:1244-7.

**19.** Bray GA, Vollmer WM, Sacks FM, Obarzanek E, Svetkey LP, Appel LJ. A further subgroup analysis of the effects of the DASH diet and three dietary sodium levels on blood pressure: results of the DASH-Sodium Trial. Am J Cardiol 2004; 94:222-7.

**20.** Vollmer WM, Sacks FM, Ard J, et al. Effects of diet and sodium intake on blood pressure: subgroup analysis of the DASH-sodium trial. Ann Intern Med 2001; 135:1019-28.

**21.** Cappuccio FP, Markandu ND, Carney C, Sagnella GA, MacGregor GA. Doubleblind randomised trial of modest salt restriction in older people. Lancet 1997; 350:850-4.

**22.** Swift PA, Markandu ND, Sagnella GA, He FJ, MacGregor GA. Modest salt reduction reduces blood pressure and urine protein excretion in black hypertensives: a randomized control trial. Hypertension 2005;46:308-12.

**23.** Lightwood JM, Coxson PG, Bibbins-Domingo K, Williams LW, Goldman L. Coronary heart disease attributable to passive smoking: CHD Policy Model. Am J Prev Med 2009;36:13-20.

24. Major outcomes in high-risk hypertensive patients randomized to angiotensin-converting enzyme inhibitor or calcium channel blocker vs diuretic: the Antihypertensive and Lipid-Lowering Treatment to Prevent Heart Attack Trial (ALLHAT). JAMA 2002;288:2981-97. [Errata, JAMA 2003;289:178, 2004;291:2196.] 25. Heidenreich PA, Davis BR, Cutler JA, et al. Cost-effectiveness of chlorthalidone, amlodipine, and lisinopril as first-step treatment for patients with hypertension: an analysis of the Antihypertensive and Lipid-Lowering Treatment to Prevent Heart Attack Trial (ALLHAT). J Gen Intern Med 2008:23:509-16.

26. Reducing salt intake in populations: report of a WHO forum and technical meeting, 5–7 October 2006 Paris, France. Geneva: World Health Organization; 2007.
27. Law MR, Frost CD, Wald NJ. By how much does dietary salt reduction lower blood pressure? III. Analysis of data from trials of salt reduction. BMJ 1991;302:819-24. [Erratum, BMJ 1991;302:939.]

**28.** Asaria P, Chisholm D, Mathers C, Ezzati M, Beaglehole R. Chronic disease prevention: health effects and financial costs of strategies to reduce salt intake and control tobacco use. Lancet 2007; 370:2044-53.

**29.** Havas S, Dickinson BD, Wilson M. The urgent need to reduce sodium consumption. JAMA 2007;298:1439-41.

**30.** Chobanian AV. Shattuck Lecture: the hypertension paradox — more uncontrolled disease despite improved therapy. N Engl J Med 2009;361:878-87. [Erratum, N Engl J Med 2009;361:1516.]

**31.** Stamler J, Rose G, Stamler R, Elliott P, Dyer A, Marmot M. INTERSALT study findings: public health and medical care

implications. Hypertension 1989;14:570-7.

**32.** Maruthur NM, Wang NY, Appel LJ. Lifestyle interventions reduce coronary heart disease risk: results from the PREMIER Trial. Circulation 2009;119: 2026-31.

**33.** Cook NR, Cutler JA, Obarzanek E, et al. Long term effects of dietary sodium reduction on cardiovascular disease outcomes: observational follow-up of the trials of hypertension prevention (TOHP). BMJ 2007;334:885-8.

**34.** Otten JJ, Hellwig JP, Meyers LD, eds. Dietary reference intakes: the essential guide to nutrient requirements. Washington, DC: Institute of Medicine, 2006.

**35.** Danaei G, Ding EL, Mozaffarian D, et al. The preventable causes of death in the United States: comparative risk assessment of dietary, lifestyle, and metabolic risk factors. PLoS Med 2009;6(4):e1000058.

**36.** Palar K, Sturm R. Potential societal savings from reduced sodium consumption in the U.S. adult population. Am J Health Promot 2009;24:49-57.

**37.** Blais CA, Pangborn RM, Borhani NO, Ferrell MF, Prineas RJ, Laing B. Effect of dietary sodium restriction on taste responses to sodium chloride: a longitudinal study. Am J Clin Nutr 1986;44:232-43.

**38.** Dietary sodium levels surveys. Aberdeen, United Kingdom: Food Standards Agency, 2008. (Accessed January 14, 2010, at http://www.food.gov.uk/science/dietarvsurveys/urinary.)

**39.** Wong MD, Shapiro MF, Boscardin WJ, Ettner SL. Contribution of major diseases to disparities in mortality. N Engl J Med 2002;347:1585-92.

**40.** Bibbins-Domingo K, Pletcher MJ, Lin F, et al. Racial differences in incident heart failure among young adults. N Engl J Med 2009;360:1179-90.

**41.** Hajjar I, Kotchen TA. Trends in prevalence, awareness, treatment, and control of hypertension in the United States, 1988-2000. JAMA 2003;290:199-206.

**42.** He FJ, MacGregor GA. Importance of salt in determining blood pressure in children: meta-analysis of controlled trials. Hypertension 2006;48:861-9.

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