Association of dietary sodium and potassium intakes with albuminuria in normal-weight, overweight, and obese participants in the Reasons for Geographic and Racial Differences in Stroke (REGARDS) Study

Kristal J Aaron, Ruth C Campbell, Suzanne E Judd, Paul W Sanders, and Paul Muntner

INTRODUCTION

Albuminuria, a marker of kidney damage, is common among US adults (1). Studies have reported an overall albuminuria prevalence of 6% to 10% among US adults, 13% to 16% in individuals with hypertension, and 28% to 38% in individuals with diabetes mellitus (2–4). Several recent studies have documented a continuous positive relation between albuminuria and cardiovascular events and end-stage renal disease (5).

Dietary sodium and potassium intakes are also associated with cardiovascular disease (6). In addition, a higher sodium intake is associated with hypertension, and this relation is stronger in overweight and obese populations (7, 8). A higher potassium intake has been associated with blood pressure reduction and improved hypertension control and may be particularly beneficial in the setting of a high-sodium diet (9–12). Although the association between sodium and potassium and cardiovascular disease and hypertension is well established, the relation with albuminuria is not as well documented, especially for obese individuals.

Because hypertension is also a known risk factor for albuminuria (13), we hypothesized that high dietary sodium and low potassium intakes would be associated with a higher prevalence of albuminuria. To test this hypothesis, we used data from the baseline visit of the REGARDS Study—a large population-based cohort of US adults aged ≥45 y (14). Because other researchers have reported that sodium intake has a more adverse effect on cardiovascular disease among obese individuals, we examined the relation between dietary sodium and potassium intakes and albuminuria by BMI in the Reasons for Geographic and Racial Differences in Stroke (REGARDS) Study (n = 30,239 adults aged ≥45 y).

RESULTS: The prevalences of albuminuria were 11.5%, 11.6%, and 16.0% in normal-weight, overweight, and obese participants, respectively. The multivariable-adjusted ORs for albuminuria in a comparison of the highest with the lowest quintile of Na/K intake (≥1.12 to <0.70 for men and ≥1.07 to <0.62 for women) were 1.09 (95% CI: 0.65, 1.22), 1.08 (95% CI: 0.85, 1.36), and 1.06 (95% CI: 1.02, 1.06) in normal-weight, overweight, and obese participants, respectively. The highest quintile of dietary sodium was associated with an increased OR for albuminuria in obese participants (OR: 1.44; 95% CI: 1.00, 2.07) but not in normal-weight or overweight participants. Dietary potassium was not associated with albuminuria.

Conclusion: In obese adults, higher dietary Na/K and sodium intakes were associated with albuminuria. Am J Clin Nutr 2011;94:1071–8.

ABSTRACT

Background: Among obese adults, sodium intake has been associated with cardiovascular disease. Few data are available on sodium intake and albuminuria, a marker of kidney damage and risk factor for cardiovascular disease.

Objective: We examined the relation between dietary sodium and potassium intakes and the ratio of sodium to potassium (Na/K) with albuminuria by BMI in the Reasons for Geographic and Racial Differences in Stroke (REGARDS) Study (n = 30,239 adults aged ≥45 y).

Design: A modified Block 98 food-frequency questionnaire was used for dietary assessment in 21,636 participants, and nutritional variables were categorized by sex-specific quintiles. Normal weight, overweight, and obese were defined as BMI (in kg/m2) categories of 18.5–24.9, 25–29.9, and ≥30, respectively. Albuminuria was defined as a ratio (mg/g) of urinary albumin to creatinine of ≥30.

Results: The prevalences of albuminuria were 11.5%, 11.6%, and 16.0% in normal-weight, overweight, and obese participants, respectively. The multivariable-adjusted ORs for albuminuria in a comparison of the highest with the lowest quintile of Na/K intake (≥1.12 to <0.70 for men and ≥1.07 to <0.62 for women) were 1.09 (95% CI: 0.65, 1.22), 1.08 (95% CI: 0.85, 1.36), and 1.06 (95% CI: 1.02, 1.06) in normal-weight, overweight, and obese participants, respectively. The highest quintile of dietary sodium was associated with an increased OR for albuminuria in obese participants (OR: 1.44; 95% CI: 1.00, 2.07) but not in normal-weight or overweight participants. Dietary potassium was not associated with albuminuria.

Conclusion: In obese adults, higher dietary Na/K and sodium intakes were associated with albuminuria. Am J Clin Nutr 2011;94:1071–8.

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a priori decided to examine this association for normal-weight, overweight, and obese individuals separately (15–17).

SUBJECTS AND METHODS

Participants

The design and objectives of the REGARDS Study were described previously (14). Briefly, by design, the study oversampled blacks and included an approximately equal representation of men and women. Enrollment began in January of 2003 and concluded in October of 2007. In addition, ~50% of the participants were recruited from the “stroke belt” (North Carolina, South Carolina, Georgia, Tennessee, Alabama, Mississippi, Arkansas, and Louisiana). The REGARDS study participants were identified by using a commercial, nationwide list of >250 million individuals in the United States (Genesys Incorporated). A trained interviewer contacted households by telephone, and one resident per household aged ≥45 y was randomly screened for eligibility. The response rate (the percentage agreeing to be interviewed among known eligible candidates contacted after adjustment for those of unknown eligibility) was 41%. Exclusion criteria included race other than black or white, active treatment of cancer, medical conditions preventing long-term participation, cognitive impairment as judged by the telephone interviewer, residence in or on a waiting list for a nursing home, and inability to communicate in English.

Of the 30,239 participants enrolled in the REGARDS Study, 8603 participants who did not complete the modified Block 98 FFQ were excluded from the current analysis (Figure 1). In addition, 1430 participants with missing serum creatinine or urinary albumin or creatinine values, 115 participants with missing BMI values, and 207 underweight (BMI [in kg/m²] <18.5) participants were excluded. Excluded participants were younger, more likely to be women, had a higher mean BMI, and were less likely to be smokers or have diabetes than those of unknown eligibility). Of the 30,239 participants enrolled in the REGARDS Study, 8603 participants who did not complete the modified Block 98 FFQ were excluded from the current analysis (Figure 1). In addition, 1430 participants with missing serum creatinine or urinary albumin or creatinine values, 115 participants with missing BMI values, and 207 underweight (BMI [in kg/m²] <18.5) participants were excluded. Excluded participants were younger, more likely to be women, had a higher mean BMI, and were less likely to be smokers or have diabetes or hypertension than were the participants included in this analysis. After these exclusions, data for 19,884 participants were available for the current analyses. The REGARDS Study was approved by the institutional review boards of the participating institutions, and all participants provided verbal consent before the telephone interview was conducted and written informed consent before completion of the in-home study visit.

Data collection

Data were collected during a telephone interview, during an in-home visit, and via self-administered questionnaires that were left for participants to complete after the in-home visit. Of relevance to the current analysis, the telephone interview included information on age, race, sex, education, household income, physical activity, current smoking status, alcohol consumption, self-rated health, a prior diagnosis of hypertension and/or diabetes mellitus, self-report of a previous diagnosis of myocardial infarction, and use of antihypertensive, lipid-lowering, and/or antidiabetes medications.

During the in-home study visit, weight, height, and waist circumference were measured following a standardized protocol. BMI was calculated as weight (in kg) divided by height squared (in m). Participants were categorized as being normal weight (BMI: 18.5–24.9), overweight (BMI: 25.0–29.9), or obese (BMI ≥30.0). With a tape measure, waist circumference was measured midway between the lowest rib and the iliac crest while the participants were standing. Abdominal obesity was defined as a waist circumference ≥88 cm for women and ≥102 cm for men. On the basis of the mean of 2 seated blood pressure measurements, hypertension was defined as a systolic blood pressure ≥140 mm Hg, a diastolic blood pressure ≥90 mm Hg, or the use of antihypertensive medication.

Participants were asked to fast for 10 to 12 h before the in-home study visit. A blood sample was collected, centrifuged, refrigerated, and shipped overnight to the REGARDS central laboratory at the University of Vermont. Total cholesterol, HDL cholesterol, triglycerides, serum glucose, and serum creatinine were measured by colorimetric reflectance spectrophotometry by using the Ortho Vitros Clinical Chemistry System 950IRC instrument (Johnson & Johnson Clinical Diagnostics). LDL cholesterol was calculated by using the Friedewald equation for individuals with triglycerides <400 mg/dL. Dyslipidemia was defined as total cholesterol ≥240 mg/dL, LDL cholesterol ≥160 mg/dL, HDL cholesterol ≤40 mg/dL, or the use of lipid-lowering medication.

As described previously (18), an isotope dilution mass spectrometry–traceable equation was used in the calibration of REGARDS serum creatinine values for calculating the eGFR by

using the Modification of Diet in Renal Disease (MDRD) formula (19):

\[
\text{eGFR} = 175 \times \text{standardized creatinine}^{-1.154} \times \text{age}^{-0.203} \times 1.212 \times (\text{if black}) \times 0.742 \times (\text{if female}) \\
(1)
\]

Diabetes mellitus was defined as a fasting glucose concentration \(\geq 126\) mg/dL, \(\geq 200\) mg/dL for participants who did not fast) or self-reported current use of antidiabetes medication.

**Measurement of urinary albumin and creatinine**

Albumin from spot urine samples was measured in batches during enrollment by using the BN ProSpec Nephelometer (Dade Behring). The assay range is 2.4–76.9 mg/L on initial sampling. Automatic dilutions are performed on specimens with higher concentrations. The interassay CVs were 2.2% at 109.9 mg/L and 4.3% at 127.9 mg/L. Urinary creatinine was measured in batches during enrollment by using the Modular-P chemistry analyzer from Roche/Hitachi. The assay range was 1–650 mg/dL on initial sampling. Automatic dilutions were performed on specimens with higher concentrations. The interassay CVs were 2.6% at 66.6 mg/dL and 8.6% at 15.6 mg/dL. Albuminuria was defined as a urinary albumin–to–creatinine ratio \(\geq 30\) mg/g.

**Statistical analysis**

Because the association of sodium with cardiovascular disease has been reported to differ by BMI, we a priori decided to perform all analyses for normal-weight, overweight, and obese participants separately. Characteristics of the participant were calculated for each BMI category separately. Next, the prevalence and multivariable-adjusted ORs for albuminuria associated with sex-specific quintiles of Na/K were calculated. Multivariable adjusted ORs were calculated by using logistic regression with 4 levels of adjustment. The first models included adjustment for age and total energy intake. The second models included additional adjustment for sex and race. The third models included additional adjustment for education, income, smoking status, alcohol consumption, self-rated health, and physical activity. The fourth models were further adjusted for hypertension, dyslipidemia, and diabetes mellitus. As a secondary analysis, the ORs for albuminuria associated with Na/K intake were calculated for normal-weight, overweight, and obese participants stratified by race. The prevalence and ORs for albuminuria associated with sex-specific quintiles of sodium intake, and separately for potassium intake, were determined in secondary analyses. In sensitivity analyses, the ORs for albuminuria associated with sex-specific quintile of Na/K were calculated separately for participants with and without abdominal obesity. The analyses were performed with SAS version 9.2 software (SAS Institute Inc).

**RESULTS**

The mean ages for normal-weight, overweight, and obese participants were 66.1, 65.6, and 63.1 y, respectively. Individuals in the higher BMI categories were more likely to be black, to have less than a high school education, to have an annual income < $20,000, to have fair or poor self-rated health, to have a sedentary lifestyle, and to have dyslipidemia, hypertension, and diabetes mellitus. In addition, systolic and diastolic blood pressure levels were higher at higher BMI categories. In contrast, current smoking and current alcohol consumption were less common at higher BMI categories. The median sodium and Na/K intake was higher at higher BMI categories, whereas potassium intake and total energy intake were not substantially different by BMI (Table 1).

Overall, the prevalences of albuminuria were 11.5%, 11.6%, and 16.0% in normal-weight, overweight, and obese participants, respectively. Among participants in each BMI category, albuminuria was most common in the highest sex-specific quintile of Na/K (Figure 2). After adjustment for age and total energy intake, in each BMI category, individuals in the highest sex-specific quintile of Na/K had higher ORs for albuminuria (Table 2). After further adjustment for sex and race, among normal-weight individuals, the association between the highest sex-specific quintile of Na/K and albuminuria was attenuated. After adjustment for education, income, smoking status, alcohol consumption, self-reported health status, and physical activity category, no association was present between sex-specific quintiles of Na/K and albuminuria in normal-weight or overweight individuals. Among obese participants, the OR for albuminuria in those in the highest compared with the lowest sex-specific quintile of Na/K was 1.30 (95% CI: 1.04, 1.62). This association remained present after further adjustment for dyslipidemia, hypertension, and diabetes mellitus. Consistent with the main findings, after multivariable adjustment, no association was present between higher sex-specific quintiles of Na/K and albuminuria for normal-weight or overweight whites (see Supplementary Table 1 under “Supplemental data” in the online issue) or blacks...
Supplementary Table 2 under “Supplemental data” in the online issue). Among obese individuals, the OR for albuminuria associated with the highest compared with the lowest sex-specific quintile of Na/K was 1.48 (95% CI: 1.06, 2.06) for whites and 1.17 (95% CI: 0.86, 1.60) for blacks (P-interaction = 0.68).

### TABLE 1

Characteristics of participants in the REGARDS Study by BMI

<table>
<thead>
<tr>
<th>Characteristic</th>
<th>Normal weight (18.5–24.9)</th>
<th>Overweight (25–29.9)</th>
<th>Obese (≥30)</th>
</tr>
</thead>
<tbody>
<tr>
<td>No. of subjects</td>
<td>4979</td>
<td>7637</td>
<td>7268</td>
</tr>
<tr>
<td>Age (y)</td>
<td>66.1 ± 10.0^2</td>
<td>65.6 ± 9.2</td>
<td>63.1 ± 8.4</td>
</tr>
<tr>
<td>Black (%)</td>
<td>21.7</td>
<td>29.6</td>
<td>44.4</td>
</tr>
<tr>
<td>Women (%)</td>
<td>57.3</td>
<td>47.1</td>
<td>61.8</td>
</tr>
<tr>
<td>Less than high school education (%)</td>
<td>7.6</td>
<td>8.7</td>
<td>11.5</td>
</tr>
<tr>
<td>Annual income &lt;$20,000 (%)</td>
<td>13.4</td>
<td>13.0</td>
<td>18.8</td>
</tr>
<tr>
<td>Currently smoking (%)</td>
<td>17.4</td>
<td>12.2</td>
<td>11.5</td>
</tr>
<tr>
<td>Currently consumes alcohol (%)</td>
<td>45.3</td>
<td>42.9</td>
<td>32.7</td>
</tr>
<tr>
<td>Fair/poor self-reported health status (%)</td>
<td>11.2</td>
<td>11.7</td>
<td>22.3</td>
</tr>
<tr>
<td>Sedentary (%)</td>
<td>28.0</td>
<td>28.0</td>
<td>38.0</td>
</tr>
<tr>
<td>Dyslipidemia (%)</td>
<td>46.3</td>
<td>61.4</td>
<td>64.1</td>
</tr>
<tr>
<td>Hypertension (%)</td>
<td>42.9</td>
<td>53.8</td>
<td>69.2</td>
</tr>
<tr>
<td>Diabetes mellitus (%)</td>
<td>7.4</td>
<td>14.7</td>
<td>29.1</td>
</tr>
<tr>
<td>DBP (mm Hg)</td>
<td>123 ± 17.3</td>
<td>127 ± 15.7</td>
<td>130 ± 15.8</td>
</tr>
<tr>
<td>eGFR (mL · min^{-1} · 1.73 m^{-2})</td>
<td>84.3 ± 17.9</td>
<td>84.1 ± 18.5</td>
<td>86.5 ± 20.3</td>
</tr>
<tr>
<td>Ratio of urinary albumin to creatinine (mg/g)^4</td>
<td>7.1 (4.7, 13.5)^5</td>
<td>6.8 (4.4, 13.1)</td>
<td>7.5 (4.7, 17.0)</td>
</tr>
<tr>
<td>Dietary intake (mg/d)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Sodium</td>
<td>1996 (1492, 2687)</td>
<td>2103 (1532, 2787)</td>
<td>2123 (1536, 2911)</td>
</tr>
<tr>
<td>Potassium</td>
<td>2519 (1889, 3275)</td>
<td>2516 (1894, 3264)</td>
<td>2444 (1806, 3215)</td>
</tr>
<tr>
<td>Ratio of sodium to potassium</td>
<td>0.82 (0.66, 0.99)</td>
<td>0.85 (0.69, 1.03)</td>
<td>0.90 (0.72, 1.09)</td>
</tr>
<tr>
<td>Total energy (kcal)</td>
<td>1559 (1193, 2012)</td>
<td>1592 (1206, 2081)</td>
<td>1594 (1186, 2136)</td>
</tr>
</tbody>
</table>

1 DBP, diastolic blood pressure; eGFR, estimated glomerular filtration rate; REGARDS, Reasons for Geographic and Racial Differences in Stroke; SBP, systolic blood pressure.

2 Mean ± SD (all such values).

3 Defined as no self-reported weekly physical activity.

4 Albuminuria is defined as a urinary albumin to creatinine ratio ≥30 mg/g.

5 Median; 25th–75th percentile in parentheses (all such values).

(see Supplementary Table 2 under “Supplemental data” in the online issue). Among obese individuals, the OR for albuminuria associated with the highest compared with the lowest sex-specific quintile of Na/K was 1.48 (95% CI: 1.06, 2.06) for whites and 1.17 (95% CI: 0.86, 1.60) for blacks (P-interaction = 0.68).

![FIGURE 2](https://academic.oup.com/ajcn/article-abstract/94/4/1071/4598103)

FIGURE 2. Prevalence of albuminuria by sex-specific quintile of the ratio of dietary sodium to potassium (Na/K) for normal-weight, overweight, and obese participants in the REGARDS Study. Albuminuria is defined as a urinary albumin to creatinine ratio ≥30 mg/g. ^1^Normal weight is defined as a BMI (in kg/m^2^) of 18.5–24.9, overweight as a BMI of 25–29.9, and obese as a BMI of ≥30. Q, quintile; REGARDS, Reasons for Geographic and Racial Differences in Stroke.
Among participants without abdominal obesity, the age- and total energy intake–adjusted OR for albuminuria in a comparison of the highest with the lowest sex-specific quintile of Na/K was 1.37 (95% CI: 1.13, 1.66; Table 3). This association was attenuated after adjustment for sex and race (OR: 1.21; 95% CI: 0.99, 1.48) and was no longer present after further adjustment for education, income, smoking status, alcohol consumption, self-reported health status, and physical activity category.

After adjustment for age and total energy intake, dietary sodium intake alone was not associated with albuminuria (see Supplementary Table 3 under “Supplemental data” in the online issue). After further adjustment, an increased OR for albuminuria was present for obese individuals in the highest compared with the lowest sex-specific quintile of sodium intake. After adjustment for age and total energy intake, higher dietary potassium intake was associated with a lower OR for albuminuria among normal-weight, overweight, and obese individuals (see Supplementary Table 4 under “Supplemental data” in the online issue). These associations were no longer present after multivariable adjustment.

DISSCUSSION

In the current study, obese participants with diets both high in sodium and low in potassium had a higher prevalence of albuminuria. This finding was consistent after multivariable adjustment for potential confounders and when defining obesity on the basis of BMI or waist circumference. In contrast, no association was present between dietary sodium or potassium intake and albuminuria among normal-weight or overweight individuals.

Prior studies showed an association between sodium intake and urinary albumin (15, 26). In a group of 839 individuals (471 men and 368 women) aged 15–70 y, du Cailar et al (15) examined the relation of sodium intake with microalbuminuria using 2 consecutive measurements of 24-h urinary sodium excretion. In that study, an association was found between the relation of higher concentrations of urinary sodium excretion with microalbuminuria after adjustment for sex, age, BMI, and systolic blood pressure.

Fox et al (26) evaluated the relation between urinary sodium, based on a spot measure and albuminuria, among 2700 participants in the Framingham Offspring Study (53% women; mean age: 58 y) who underwent routine examination between 1995 and 1998. Using multivariate regression techniques, they found a robust correlation between log urinary sodium and the log albumin-to-creatinine ratio.

The relation between sodium intake and albuminuria by level of BMI was reported in at least one prior study (17). The PREVEND Study used cross-sectional data from 7850 adults aged 28–75 y from the Netherlands to study the relation between sodium intake (as determined by the mean sodium value from two 24-h urine collections) and albuminuria in normal-weight, overweight, and obese participants separately. This cohort was enriched with participants with elevated urinary albumin concentrations. The PREVEND investigators showed a positive relation between

<table>
<thead>
<tr>
<th>Quintile of dietary sodium to potassium ratio</th>
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<tbody>
<tr>
<td>1</td>
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<tr>
<td>---</td>
</tr>
<tr>
<td>Men (^2)</td>
</tr>
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<td>Women (^2)</td>
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<tr>
<td>Model 1 (^4)</td>
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<td>Model 2 (^5)</td>
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<td>Model 3 (^6)</td>
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<tr>
<td>Model 4 (^3)</td>
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</table>

\(^1\) Normal weight is defined as a BMI (in kg/m\(^2\)) of 18.5–24.9, overweight as 25–29.9, and obese as ≥30. Albuminuria is defined as a urinary albumin to creatinine ratio ≥30 mg/g. REGARDS, Reasons for Geographic and Racial Differences in Stroke.

\(^2\) Values are ranges.

\(^3\) Adjusted for age and total energy intake.

\(^4\) Adjusted for strategies in model 1 plus sex and race.

\(^5\) Adjusted for variables in model 2 plus education category, income category, smoking status, alcohol consumption, self-reported health status, and physical activity category.

\(^6\) Adjusted for variables in model 3 plus dyslipidemia, hypertension, and diabetes mellitus.

## Table 3

| Obese Reference | Overweight Reference | Normal weight Reference | Adjusted for variables in model 2 plus education category, income category, smoking status, alcohol consumption, self-reported health status, and physical activity category.
<table>
<thead>
<tr>
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</thead>
<tbody>
<tr>
<td>1.13 (0.91, 1.45)</td>
<td>0.88 (0.66, 1.17)</td>
<td>0.92 (0.69, 1.23)</td>
<td>0.85 (0.63, 1.16)</td>
</tr>
<tr>
<td>1.15 (0.91, 1.45)</td>
<td>0.83 (0.66, 1.05)</td>
<td>0.87 (0.68, 1.09)</td>
<td>1.05 (0.84, 1.32)</td>
</tr>
<tr>
<td>1.15 (0.91, 1.45)</td>
<td>0.79 (0.62, 1.00)</td>
<td>0.82 (0.65, 1.05)</td>
<td>0.98 (0.78, 1.24)</td>
</tr>
<tr>
<td>1.13 (0.89, 1.44)</td>
<td>0.88 (0.66, 1.17)</td>
<td>0.92 (0.69, 1.23)</td>
<td>0.85 (0.63, 1.16)</td>
</tr>
<tr>
<td>1.06 (0.83, 1.35)</td>
<td>1.08 (0.85, 1.36)</td>
<td>1.11 (0.88, 1.40)</td>
<td>1.28 (1.02, 1.61)</td>
</tr>
</tbody>
</table>

**DISCUSSION**

In the current study, obese participants with diets both high in sodium and low in potassium had a higher prevalence of albuminuria. This finding was consistent after multivariable adjustment for potential confounders and when defining obesity on the basis of BMI or waist circumference. In contrast, no association was present between dietary sodium or potassium intake and albuminuria among normal-weight or overweight individuals.

Prior studies showed an association between sodium intake and urinary albumin (15, 26). In a group of 839 individuals (471 men and 368 women) aged 15–70 y, du Cailar et al (15) examined the relation of sodium intake with microalbuminuria using 2 consecutive measurements of 24-h urinary sodium excretion. In that study, an association was found between the relation of higher concentrations of urinary sodium excretion with microalbuminuria after adjustment for sex, age, BMI, and systolic blood pressure.

Fox et al (26) evaluated the relation between urinary sodium, based on a spot measure and albuminuria, among 2700 participants in the Framingham Offspring Study (53% women; mean age: 58 y) who underwent routine examination between 1995 and 1998. Using multivariate regression techniques, they found a robust correlation between log urinary sodium and the log albumin-to-creatinine ratio.

The relation between sodium intake and albuminuria by level of BMI was reported in at least one prior study (17). The PREVEND Study used cross-sectional data from 7850 adults aged 28–75 y from the Netherlands to study the relation between sodium intake (as determined by the mean sodium value from two 24-h urine collections) and albuminuria in normal-weight, overweight, and obese participants separately. This cohort was enriched with participants with elevated urinary albumin concentrations. The PREVEND investigators showed a positive relation between...
...endothelial dysfunction, reduces vascular compliance and amplifies peripheral oxidant production, which results in unopposed excess vascular stiffening and arterial constriction (33–35). Thus, whereas a reduction of sodium in the diet results in lower blood pressure independent of weight loss (36), it also results in further reductions in blood pressure in the setting of antihypertensive therapy (37, 38) and improves hypertension control (39, 40).

The distinction of the Na/K ratio may be important because dietary potassium intake may play a protective role in the incidence of albuminuria. This theory was detailed by Meneely and Ball (41) >50 y ago and was based on their observations that rats fed toxic amounts of sodium chloride survived longer if they also received higher amounts of potassium. Dietary potassium intake and supplementation have also been found to counteract the harmful effects of dietary salt in humans (12, 36).

Although it appears contradictory, the protective effect of potassium is more intense in the presence of excess dietary sodium as compared with salt depletion (9). The mechanistic action by which dietary potassium attenuates the effect of dietary sodium excess is not fully understood; however, animal and clinical studies have shown that the effect may be associated with sympathetic nerve inhibition in salt-sensitive hypertension (42). A high-potassium diet improves vascular function with respect to generation of vasodilators—nitric oxide and other endothelium-derived factors. Experimental studies suggest that a potassium-rich diet may decrease cardiovascular disease risk through one or more of the following mechanisms (38, 43): inhibition of free radical formation from vascular endothelial cells and macrophages, reduced proliferation of vascular smooth muscle cells, decreases in platelet aggregation and arterial thrombosis, and reduced vascular stiffness and arterial transport of potassium to generation of vasodilators—nitric oxide and other endothelium-derived factors. Experimental studies suggest that a potassium-rich diet may decrease cardiovascular disease risk through one or more of the following mechanisms (38, 43): inhibition of free radical formation from vascular endothelial cells and macrophages, reduced proliferation of vascular smooth muscle cells, decreases in platelet aggregation and arterial thrombosis.
and a reduction of renal vascular resistance and a subsequent increase in GFR.

The current study had potential limitations. We were unable to assess causality because of the study’s observational cross-sectional design. In addition, the assessment of albuminuria was determined from spot urine samples as opposed to a timed or 24-h urine collection; however, spot urine measurement has been shown to perform well at detecting abnormal urinary albumin excretion, and the one time collection avoids error introduced by inadequate collections over time (44, 45). Another limitation of the current study was that sodium and potassium intakes were estimated by dietary recall rather than by 24-h urine collections—the gold standard for urinary electrolyte measurement. A well-known issue with dietary assessment instruments is the underreporting of nutritional intakes. Several factors (eg, age, sex, race, socioeconomic status, and other lifestyle factors) influence eating habits and the reporting of nutritional intake (37, 40). To address this limitation, dietary sodium, potassium, and Na/K were analyzed by using quintiles rather than by estimating daily intakes. In addition, dietary questionnaires provide acceptable classification of nutrient intakes on an individual level and are widely accepted in cohort studies with large sample sizes (39, 46). Some bias may have potentially been introduced because about one-third of the cohort was excluded, mainly because of missing FFQs. Despite these limitations, the REGARDS Study had several notable strengths, including the enrollment of a large sample of whites and blacks from the continental United States and the collection of dietary data with the use of a standardized and validated FFQ. In addition, urinary albumin and creatinine were measured at a central laboratory following standardized procedures.

In conclusion, in the current study obese individuals with a diet high in sodium and low in potassium were more likely to have albuminuria. Although the physiologic mechanisms are not fully understood, diets high in potassium may mediate the adverse effects of dietary sodium and serve to protect the vascular endothelium from sodium-induced injury. If the results from the current study are confirmed in the future, testing the effect of sodium reduction on albuminuria in randomized trials may be warranted.

The authors’ responsibilities were as follows—KJA and RCC: conceived of and designed the study, interpreted the data, and wrote and critically reviewed the manuscript; SEJ: analyzed and interpreted the data, critically reviewed the manuscript, and assisted with the writing of the manuscript; PM: critically reviewed and assisted with the writing of the manuscript; and PWS: assisted with the conception of the study and with the interpretation of the data and critically reviewed and edited the manuscript. None of the authors had a conflict of interest or any financial disclosures.

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