Original Report: Cardiovascular Disease and Risk Factors VALIDATION OF AN ALBUMINURIA SELF-ASSESSMENT TOOL IN THE MULTI-ETHNIC STUDY OF ATHEROSCLEROSIS (MESA)

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Objective: We previously developed an 8-item self-assessment tool to identify individuals with a high probability of having albuminuria. This tool was developed and externally validated among non-Hispanic Whites and non-Hispanic Blacks. We sought to validate it in a multi-ethnic cohort that also included Hispanics and Chinese Americans.

Design: This is a cross-sectional study.

Setting: Data were collected using standardized questionnaires and spot urine samples at a baseline examination in 2000-2002. The 8 items in the self-assessment tool include age, race, gender, current cigarette smoking, history of diabetes, hypertension, or stroke, and self-rated health.

Participants: Of 6,814 community-dwelling adults aged 45-84 years participating in the Multi-Ethnic Study of Atherosclerosis (MESA), 6,542 were included in the primary analysis.

Main Outcome Measures: Albuminuria was defined as urine albumin-to-creatinine ratio \geq 30 mg/g at baseline.

Results: Among non-Hispanic Whites, non-Hispanic Blacks, Hispanics, and Chinese Americans, the prevalence of albuminuria was 6.0%, 11.3%, 11.6%, and 10.8%, respectively. The c-statistic for discriminating participants with and without albuminuria was .731 (95% CI: .692, .771), .728 (95% CI: .687, .761), .747 (95% CI: .709, .784), and .761 (95% CI: .699, .814) for non-Hispanic Whites, non-Hispanic Blacks, Hispanics, and Chinese Americans, respectively. The self-assessment tool over-estimated the probability of albuminuria for non-Hispanic Whites and Blacks, but was well-calibrated for Hispanics and Chinese Americans.

INTRODUCTION

established Albuminuria, an marker of kidney disease, affects more than 10 million US adults.¹ It is a strong independent risk factor for cardiovascular disease (CVD), end-stage renal disease (ESRD), and all-cause mortality.²⁻⁴ Despite its high prevalence and prognostic importance, most individuals with albuminuria are unaware of having this condition.⁵ This low awareness is present in the general population and in high-risk groups, including individuals with diabetes.6-8 Research has shown that populationwide screening for albuminuria is

Conclusions: The albuminuria self-assessment tool maintained good test characteristics in this large multi-ethnic cohort, suggesting it may be helpful for increasing awareness of albuminuria in an ethnically diverse population. *Ethn Dis*.2015;25(4):427-434; doi:10.18865/ed.25.4.427

Keywords: Albuminuria, Screening, Kidney Disease, Self-assessment, Race, Ethnicity, Racial Disparities

¹Department of Epidemiology, University of Alabama at Birmingham, Birmingham, AL ²The George Institute for Global Health, Nuffield Department of Population Health, University of Oxford, Oxford, UK ³The George Institute for Global Health, University of Sydney, Sydney, Australia not cost-effective; however, screening for albuminuria may be beneficial among high-risk individuals.⁷⁻⁹

Using data from two large population-based studies, we previously developed and externally validated an 8-item self-assessment tool designed to identify individuals with a high probability of having albuminuria.¹⁰ This tool is available for use on the National Kidney Foundation website.¹¹ While this tool has demonstrated strong test characteristics and has implications for increasing the awareness, detection, and treatment of albuminuria, only non-Hispanic White and non-Hispanic Black adults were included in the

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Address correspondence to Paul Muntner, PhD; Department of Epidemiology; University of Alabama at Birmingham; 1665 University Boulevard, Suite 230J; Birmingham, AL 35294; 205. 975.8077; pmuntner@uab.edu tool's development and validation. Validation of the albuminuria selfassessment tool among Hispanic and Asian adults is necessary before it can be recommended for use in these populations. Therefore, we sought to assess the test characteristics (sensitivity, specificity, and positive and negative predictive values), discrimination, and calibration of the albuminuria self-assessment tool in a multi-ethnic cohort that included Hispanics and Chinese Americans. We also assessed the characteristics of the self-assessment tool for men and women, separately. To do so, we analyzed data from the Multi-Ethnic Study of Atherosclerosis (MESA).

METHODS

Study Population

Details of the MESA study design and recruitment have been published previously.¹² In brief, between July 2000 and August 2002, 6,814 community-dwelling adults aged 45-84 years who were free of clinically apparent cardiovascular disease were enrolled in MESA. Participants from four race/ethnicity groups (non-Hispanic Whites, non-Hispanic Blacks, Hispanics, and Chinese Americans) were recruited from six US communities: Baltimore City and Baltimore County, Maryland; Chicago, Illinois; Forsyth County, North Carolina; Los Angeles County, California; Northern Manhattan and the Bronx, New York; and St. Paul, Minnesota. The MESA study was approved by the institutional review boards of all participating centers, and written in...we sought to assess the test characteristics (sensitivity, specificity, and positive and negative predictive values), discrimination, and calibration of the albuminuria selfassessment tool in a multiethnic cohort that included Hispanics and Chinese Americans.

formed consent was obtained from all participants, including consent to use the data for scientific purposes. Approval to conduct the current analysis was granted by the institutional review board at the University of Alabama at Birmingham.

Data Collection

MESA data for the primary analysis were collected through a screening/recruitment questionnaire and a baseline examination. Additional data for the secondary analysis were collected during a follow-up examination occurring approximately 18 months after baseline. The eight items in the self-assessment tool collected at baseline included age, sex (male or female), race-ethnicity (non-Hispanic Black, non-Hispanic White,

Hispanic, or Chinese American), cigarette smoking (current vs former or never), self-reported history of diabetes, hypertension, or stroke, and self-rated health (excellent, very good, good, fair, or poor). Demographic information and data on tobacco use, hypertension, diabetes, and self-rated health were obtained using standardized questionnaires. Current cigarette smoking was defined as having smoked cigarettes in the last 30 days. Hypertension and diabetes were defined by self-report of a prior diagnosis by a doctor or health professional. Individuals with a history of cardiovascular disease at baseline were not enrolled in MESA.

Definition of Albuminuria and Persistent Albuminuria

Using spot urine samples collected at baseline and exam 2, urinary albumin concentration was determined by nephelometry using the Array 360 CE Protein Analyzer (Beckman Instruments, Inc). Urinary creatinine was measured by the Jaffé rate method using the Vitros 950IRC instrument (Johnson & Johnson Clinical Diagnostics Inc). For our primary analysis, albuminuria was defined as an albumin-tocreatinine ratio $\geq 30 \text{ mg/g}$ at the baseline exam. In a secondary analysis, the outcome was persistent albuminuria, defined as having an albumin-to-creatinine ratio $\geq 30 \text{ mg/g}$ at both the baseline exam and exam 2.

Statistical Analysis

Of the 6,814 MESA participants who attended the baseline exam, we excluded those without information on components of the selfassessment tool (n=94) or urinary albumin or creatinine (n=38), or who self-reported a prior diagnosis of kidney disease (n=140), resulting in the inclusion of 6,542 participants for the primary analysis. Using the albuminuria self-assessment tool developed in the REasons for Geographic And Racial Differences in Stroke (REGARDS) study,10 each MESA participant was assigned a predicted probability of having albuminuria at baseline using the following equation: exp(X)/(1+exp[X])where X = -5.1062 + .0318 * (age in years) + .000994 * (age in years - $(65)^2 + .3476 * (1 \text{ for non-Hispanic})^2$ Blacks, 0 for other race-ethnicities) + .2515 * (1 for men, 0 for women) + .4383 * (1 for current smokers, 0 for nonsmokers) + .2615 * (1 if selfrated health is good) + .5043 * (1 if self-rated health is fair) + .4545 * (1 if self-rated health is poor) + .8515 * (1 for self-report of a diagnosis of diabetes, 0 for those not reporting diabetes) + .4755 * (1 for self-report of a diagnosis of hypertension, 0 for those not reporting hypertension)

+ .3536 * (1 for prior stroke, 0 for those without a prior stroke). As the development of the self-assessment tool did not include Hispanics and Chinese Americans, we included them in the "other race-ethnicities" group. All analyses were conducted overall and separately for each race-ethnicity group (non-Hispanic Whites, non-Hispanic Blacks, Hispanics, and Chinese Americans).

Participant characteristics, including the eight components of the self-assessment tool, were calculated for individuals with and without albuminuria, separately. We calculated sensitivity, specificity, and positive and negative predictive values for albuminuria associated with a probability $\geq 10\%$, the previously published cut-point for identifying a high probability of albuminuria on the self-assessment tool.¹⁰ We then calculated the c-statistic for discriminating MESA participants with and without albuminuria associated with the probability of having albuminuria on the self-assessment tool. The c-statistic is a measure of how well a model discriminates between individuals with and without a condition (eg, albuminuria). For the current analysis, the c-statistic is the proportion of all pairwise comparisons of participants, one with albuminuria and one without albuminuria, for which the participant with albuminuria had a higher predicted probability on the self-assessment tool. We used a 1,000 iteration bias-corrected bootstrap to create 95% confidence intervals (CI) around the c-statistics. This approach involves resampling (with replacement) from the available data to create a large number of samples to approximate the distribution of the c-statistic. The cstatistics in this sample are ordered from lowest to highest and the 25th and 975th values are considered the boundary for the 95% CI around the c-statistic. To assess calibration of the self-assessment tool, the observed and predicted prevalence of albuminuria was calculated by quintile of predicted probability. Since some people with albuminuria do not have it on re-retesting, we con-

	Non-Hispanic White (n=2,538)		Non-Hispanic Black (n=1,819)		Hispanic (n=1,417)		Chinese-American (n=768)	
	No albuminuria (n=2,387)	Albuminuria (n=151)	No albuminuria (n=1,613)	Albuminuria (n=206)	No albuminuria (n=1,253)	Albuminuria (n=164)	No albuminuria (n=685)	Albuminuria (n=83)
Age in years, mean (SD)	62.1 (10.2)	68.8 (9.6)	61.7 (10.0)	65.9 (9.5)	60.7 (10.1)	65.9 (10.9)	61.5 (10.2)	68.3 (9.7)
Male, %	47.8	51.7	43.9	50.5	48.7	47.0	48.3	42.2
Current smoker, %	11.3	13.3	17.9	16.5	13.7	12.8	5.6	4.8
Hypertension, %	30.4	56.3	52.3	74.8	31.0	64.6	29.1	60.2
Diabetes, %	4.7	13.9	12.7	41.3	12.8	42.7	7.9	34.9
History of stroke, %	0	0	0	0	0	0	0	0
Self-rated health, %								
Excellent	28.1	12.6	11.8	6.3	9.2	6.7	4.2	2.4
Very good	44.3	46.4	31.9	21.4	23.5	16.5	29.4	14.5
Good	24.4	32.4	44.2	46.6	50.8	58.5	64.2	79.5
Fair	3.1	8.6	11.4	23.3	15.3	14.6	2.2	3.6
Poor	.1	0	.7	2.4	1.2	3.7	0	0

Table 2. Test characteristics for albuminuria associated with a predicted probability of albuminuria ≥10% on the self-assess-
ment tool and c-statistics for discriminating MESA participants with albuminuria

	Sensitivity (95% CI)	Specificity (95% CI)	PPV (95% CI)	NPV (95% CI)	c-statistic (95% CI)
Overall	.72 (.68, .76)	.59 (.58, .61)	.17(.15, .18)	.96 (.95, .96)	.743 (.722, .763)
Race-ethnicity					
Non-Hispanic White	.55 (.47, .63)	.79 (.77, .81)	.14 (.11, .17)	.97 (.96, .97)	.731 (.692, .771)
Non-Hispanic Black	.87 (.82, .91)	.41 (.38, .43)	.16 (.14, .18)	.96 (.94, .97)	.728 (.687, .761)
Hispanic	.71 (.63, .78)	.68 (.65, .70)	.22 (.19, .26)	.95 (.93, .96)	.747 (.709, .784)
Chinese-American	.69 (.58, .78)	.74 (.71, .78)	.25 (.19, .31)	.95 (.93, .97)	.761 (.699, .814)

CI, confidence interval; PPV, positive predictive value; NPV, negative predictive value.

ducted a secondary analysis wherein we examined the discrimination and test characteristics of the screening tool for identifying persistent albuminuria. These analyses were limited to the 5,911 MESA participants who attended, and had valid albuminuria measures, at both baseline and exam 2. We also assessed the test characteristics (sensitivity, specificity, and positive and negative predictive values), discrimination and calibration of the self-assessment tool for men and women, separately. All analyses were conducted using SAS version 9.2 (SAS Institute, Cary, NC).

RESULTS

Participant Characteristics

Among non-Hispanic Whites, non-Hispanic Blacks, Hispanics, and Chinese Americans, the prevalence of albuminuria was 6.0%, 11.3%, 11.6%, and 10.8%, respectively. On average, participants with albuminuria were older than those without albuminuria (Table 1). Participants with albuminuria were also more likely to self-report a history of hypertension or diabetes. Individuals with vs without albuminuria were more likely to report fair or poor health.

Validation of the Self-assessment Tool by Race/ethnicity

Test characteristics for the selfassessment tool identifying MESA participants with albuminuria are shown by race/ethnicity in Table 2. In the overall population, the sensitivity, specificity, positive predictive value, and negative predictive value were .72 (95% CI: .68, .76), .59 (95% CI: .58, .61), .17 (95% CI: .15, .18), and .96 (95% CI: .95, .96), respectively. The c-statistic for discriminating participants with and without albuminuria was .743 (95% CI: .722, .763). The discrimination was higher for Chinese Americans (c-statistic .761; 95% CI: .699, .814) and Hispanics (c-statistic .747; 95% CI: .709, .784), compared with non-Hispanic Whites (cstatistic .731; 95% CI: .692, .771) and non-Hispanic Blacks (.728; 95% CI: .687, .761). The self-assessment tool over-estimated the probability of albuminuria for non-Hispanic Whites and Blacks, but was well-calibrated for Hispanics and Chinese Americans (Figure 1).

Validation of the Selfassessment Tool for Persistent Albuminuria

The prevalence of persistent albuminuria was 3.5%, 6.8%, 7.3%, and 6.4% among non-Hispanic Whites, non-Hispanic Blacks, Hispanics, and Chinese Americans, re-

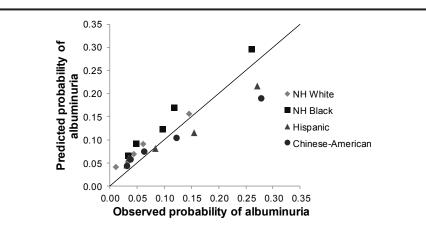


Figure 1. Observed vs predicted probability of albuminuria by race-ethnicity NH,non-Hispanic

spectively. The sensitivity, specificity, positive predictive value, and negative predictive value for persistent albuminuria associated with a predicted probability ≥10% were .72 (95% CI: .67, .77), .66 (95% CI: .65, .67), .11 (95% CI: .10, .12), and .98 (95% CI: .97, .98), respectively (supplemental data available from lead author). The c-statistic for discriminating participants with and without persistent albuminuria was .756 (95% CI: .727, .780). The discrimination was higher for Hispanics (c-statistic .774; 95%CI: .728, .816) and Chinese Americans (c-statistic .758; 95% CI: .675, .827), compared with non-Hispanic Whites (c-statistic .740; 95% CI: .682, .789) and Blacks (c-statistic .743; 95% CI: .687, .790). The self-assessment tool over-predicted persistent albuminuria for each race-ethnicity group (supplemental data available from lead author).

Validation of the Selfassessment Tool by Sex

The prevalence of albuminuria was 9.5% and 9.0% among men and women, respectively, while the prevalence of persistent albuminuria was 6.6% and 4.6% among men and women, respectively. The c-statistics for albuminuria were .763 (95% CI: .733, .787) and .730 (95% CI: .701, .759) among men and women, respectively (data not shown). The c-statistic for persistent albuminuria was .759 (95% CI: .717, .793) for men and .723 (95% CI: .683, .768) for women. For both men and women, the self-assessment tool was well-calibrated for albuminuria but over-estimated the probability of persistent albuminuria (supplemental data available from lead author).

DISCUSSION

Using data from a large multiethnic cohort, we validated a selfassessment tool designed to identify individuals with a high probability of albuminuria in non-Hispanic Whites, non-Hispanic Blacks, Hispanics and Chinese Americans. Although the tool demonstrated only moderate sensitivity and specificity, it maintained good discrimination in each race-ethnicity. The self-assessment tool was well-calibrated in Hispanics and Chinese Americans. Although the tool over-predicted albuminuria for non-Hispanic Whites and non-Hispanic Blacks in our study, it was well-calibrated among Whites and Blacks in our prior validation study using National Health and Nutrition Examination Survey (NHANES) data.¹⁰ When evaluated in men and women, the selfassessment tool performed well in each group with good discrimination and calibration statistics. Therefore, this tool could be used to increase albuminuria awareness among individuals with a high risk.

The early stages of CKD are usually asymptomatic and thus, awareness is low both in the general population and in high-risk populations.^{5,13-17} In a prior analysis of MESA data, albuminuria was associated with higher mean end-diastolic left ventricular mass and coronary artery calcification scores.¹⁸ In other studies, albuminuria has been associated with an increased risk for CVD

events, ESRD, and death.^{3,4} These data suggest that early albuminuria detection could provide an opportunity for risk reduction. In 2000, the National Kidney Foundation developed a free voluntary community-based kidney health screening program (Kidney Early Evaluation Program [KEEP]) to identify individuals at greatest risk of CKD, particularly those with diabetes or hypertension.^{6,19} Since its inception, KEEP has screened approximately 185,000 high-risk individuals.²⁰ Although awareness of CKD is increasing, overall rates remain low.5

While screening for albuminuria is feasible in a primary care setting, population-wide albuminuria screening is not recommended and studies show it is likely not cost-effective.²¹⁻²³ However, data have suggested it is cost-effective to screen for albuminuria in high-risk populations.^{9,24} The American Diabetes Association recommends screening for CKD in all patients with diabetes and several organizations, including the Joint National Committee on Prevention. Detection, Evaluation, and Treatment of High Blood Pressure (JNC), recommend screening patients being treated for hypertension.²⁵ The self-assessment tool provides an opportunity to identify high-risk individuals for albuminuria screening.

The prevalence of albuminuria was higher among Hispanics and Chinese Americans compared with non-Hispanic Whites in our study, highlighting the importance of increasing awareness among these groups. Among Hispanics, a recent study of NHANES participants demonstrated a higher odds of albuminuria compared to Whites (multivariable adjusted odds ratio: 1.65; 95% CI: 1.07, 2.54).²⁶ Furthermore, the incidence of ESRD among Hispanics in the United States is 1.8 times higher than that of Whites.²⁷ Among Asians, data suggest that the risk of ESRD is similar to that of Whites.²⁸ Hispanics, but not Asians, are also more likely to forego medical visits than Whites, particularly in the wake of the "Great Recession" of 2007-2009.29 As such, a self-assessment tool may be especially important for increasing awareness of CKD in Hispanics.

Framingham study data show that men have an increased risk for incident albuminuria compared to women (OR: 1.89; 95% CI: 1.37, 2.56).³⁰ However, men are also less likely than women to utilize many health care services.^{31,32} In a cohort of 1,742 US adults participating in targeted, free, community-based CKD screenings, 70% were women.33 This suggests a potential unmet need for screening strategies for men at high risk for CKD. The self-assessment tool validated in our study demonstrated good discrimination and calibration for both men and women, suggesting it may be useful for increasing albuminuria awareness, particularly among men who otherwise would not utilize screening services.

The albuminuria self-assessment tool validated in our study includes eight items that are individually associated with albuminuria.³⁴⁻³⁷ In development of the self-assessment tool, the items in aggregate provided significantly better discrimination than any of the items individually.¹⁰ Although self-assessment tools such as the one validated in our study allow for widespread use, they do not provide clinical diagnoses. Individuals with a high probability for albuminuria based on the self-assessment tool should be recommended for further screening using spot urine testing for albuminuria. The best approach for effectively communicating the need for screening for individuals who test positive on the albuminuria self-assessment tool warrants further study. Also, the overall

The prevalence of albuminuria was higher among Hispanics and Chinese Americans compared with non-Hispanic Whites in our study...

sensitivity and specificity were modest (72% and 59%, respectively) and future studies may identify screening tools with improved test characteristics.

Our study has a number of strengths. These include the use of a large multiethnic study population that provided an adequate sample size to validate the albuminuria self-assessment tool in four race-ethnicity groups. Also, MESA included extensive data collection following rigorous standardized protocols. The availability of albuminuria at a follow-up examination allowed us to validate the self-assessment tool for persistent albuminuria. Despite these strengths, the findings of our study should be interpreted in the context of certain limitations. The self-assessment tool was developed using intervieweradministered questionnaires and thus, are subject to the inherent limitations of self-reported data. Evaluating the test characteristics of the tool using other modes of administration may be useful. For the current validation study, the eight items of the tool were part of several different questionnaires collected as part of the baseline MESA study visit, and the eight items were not administered in an uninterrupted succession. Also, the MESA study was limited to adults aged ≥45 years. Although albuminuria is not common in younger adults, future efforts may be warranted to validate the self-assessment tool for adults ≥45 years of age. The exclusion of individuals with a history of cardiovascular disease from MESA and the fact that the Asian participants were primarily of Chinese descent may limit the generalizability of our findings. Additionally, individuals with weight >300 lbs were excluded from MESA and thus, our results may not be generalizable to the severely obese.

CONCLUSION

In conclusion, albuminuria is an important risk factor for ESRD, CVD, and mortality. Early detection and treatment may reduce the risk of these events. However, despite the prognostic importance of albuminuria and the availability of effective treatments for it, awareness of albuminuria in the general population is low. In our study, we have externally validated an albuminuria self-assessment tool among non-Hispanic Whites, non-Hispanic Blacks, Hispanics and Chinese Americans. The tool maintained good discrimination and calibration in this large multi-ethnic cohort, suggesting it may be helpful for increasing awareness of albuminuria in an ethnically diverse population.

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AUTHOR CONTRIBUTIONS

Research concept and design: Tanner, Woodward, Muntner. Acquisition of data: Katz, Muntner. Data analysis and interpretation: Warnock, Tanner, Kramer, Katz, Shimbo, Gutiérrez, Woodward, Muntner, Peralta. Manuscript draft: Warnock, Tanner, Kramer, Katz, Shimbo, Gutiérrez, Woodward, Muntner, Peralta. Statistical expertise: Tanner, Kramer, Katz, Woodward, Muntner. Acquisition of funding: Warnock. Administrative: Shimbo, Gutiérrez, Muntner, Peralta. Supervision: Shimbo, Muntner

References

- Coresh J, Selvin E, Stevens LA, et al. Prevalence of chronic kidney disease in the United States. *JAMA*. 2007;298(17):2038-2047. http://dx.doi.org/10.1001/jama.298.17.2038. PMID:17986697.
- Warnock DG, Muntner P, McCullough PA, et al; REGARDS Investigators. Kidney function, albuminuria, and all-cause mortality in the REGARDS (Reasons for Geographic and Racial Differences in Stroke) study. *Am J Kidney Dis.* 2010;56(5):861-871. http:// dx.doi.org/10.1053/j.ajkd.2010.05.017. PMID:20692752.
- Sarnak MJ, Katz R, Newman A, et al; Health ABC Study. Association of urinary injury biomarkers with mortality and cardiovascular events. J Am Soc Nephrol. 2014;25(7):1545-1553. http://dx.doi.org/10.1681/

ASN.2013070713. PMID:24511130.

- Bentata Y, Abouqal R. Does albuminuria predict renal risk and/or cardiovascular risk in obese type 2 diabetic patients? *Am J Cardiovasc Dis.* 2014;4(1):26-30. PMID:24551483.
- Plantinga LC, Boulware LE, Coresh J, et al. Patient awareness of chronic kidney disease: trends and predictors. *Arch Intern Med.* 2008;168(20):2268-2275. http:// dx.doi.org/10.1001/archinte.168.20.2268. PMID:19001205.
- Whaley-Connell A, Sowers JR, McCullough PA, et al; KEEP Investigators. Diabetes mellitus and CKD awareness: the Kidney Early Evaluation Program (KEEP) and National Health and Nutrition Examination Survey (NHANES). *Am J Kidney Dis.* 2009;53(4)(suppl 4):S11-S21. http://dx.doi.org/10.1053/j.ajkd.2009.01.004. PMID:19285607.
- Vijan S, Hofer TP, Hayward RA. Cost-utility analysis of screening intervals for diabetic retinopathy in patients with type 2 diabetes mellitus. *JAMA*. 2000;283(7):889-896. http://dx.doi. org/10.1001/jama.283.7.889. PMID:10685713.
- Kiberd BA, Jindal KK. Screening to prevent renal failure in insulin dependent diabetic patients: an economic evaluation. *BMJ*. 1995;311(7020):1595-1599. http:// dx.doi.org/10.1136/bmj.311.7020.1595. PMID:8555801.
- Boulware LE, Jaar BG, Tarver-Carr ME, Brancati FL, Powe NR. Screening for proteinuria in US adults: a cost-effectiveness analysis. *JAMA*. 2003;290(23):3101-3114. http://dx.doi.org/10.1001/jama.290.23.3101. PMID:14679273.
- Muntner P, Woodward M, Carson AP, et al. Development and validation of a selfassessment tool for albuminuria: results from the Reasons for Geographic and Racial Differences in Stroke (REGARDS) Study. *Am J Kidney Dis.* 2011;58(2):196-205. http:// dx.doi.org/10.1053/j.ajkd.2011.01.027. PMID:21620547.
- National Kidney Foundation. Quick Kidney Risk Check. 2013; http://www.kidney.org/ kidneydisease/selfAssessmentQuiz.cfm. Last accessed July 31, 2014.
- Bild DE, Bluemke DA, Burke GL, et al. Multi-ethnic study of atherosclerosis: objectives and design. *Am J Epidemiol.* 2002;156(9):871-881. http://dx.doi.org/10.1093/aje/kwf113. PMID:12397006.
- Coresh J, Byrd-Holt D, Astor BC, et al. Chronic kidney disease awareness, prevalence, and trends among U.S. adults, 1999 to 2000. *J Am Soc Nephrol.* 2005;16(1):180-188. http:// dx.doi.org/10.1681/ASN.2004070539. PMID:15563563.
- Li C, Wen XJ, Pavkov ME, et al. Awareness of kidney disease among US adults: findings from the 2011 behavioral risk factor surveillance system. *Am J Nephrol.* 2014;39(4):306-

313. http://dx.doi.org/10.1159/000360184. PMID:24732234.

- Tuot DS, Plantinga LC, Hsu CY, et al; Centers for Disease Control Chronic Kidney Disease Surveillance Team. Chronic kidney disease awareness among individuals with clinical markers of kidney dysfunction. *Clin J Am Soc Nephrol.* 2011;6(8):1838-1844. http://dx.doi. org/10.2215/CJN.00730111. PMID:21784832.
- Tuot DS, Plantinga LC, Judd SE, et al; REGARDS Investigators. Healthy behaviors, risk factor control and awareness of chronic kidney disease. *Am J Nephrol.* 2013;37(2):135-143. http://dx.doi.org/10.1159/000346712. PMID:23392070.
- Garcia-Garcia G, Gutiérrez-Padilla AJ, Renoirte-Lopez K, et al. Chronic kidney disease in homeless persons in Mexico. *Kidney Int Suppl (2011)*. 2013;3(2):250-253. http://dx.doi.org/10.1038/ kisup.2013.25. PMID:25018992.
- Kramer H, Jacobs DR Jr, Bild D, et al; The Multi-Ethnic Study of Atherosclerosis. Urine albumin excretion and subclinical cardiovascular disease. *Hypertension*. 2005;46(1):38-43. http://dx.doi.org/10.1161/01. HYP.0000171189.48911.18. PMID:15956113.
- Brown WW, Peters RM, Ohmit SE, et al. Early detection of kidney disease in community settings: the Kidney Early Evaluation Program (KEEP). *Am J Kidney Dis.* 2003;42(1):22-35. PMID: 12830453.
- National Kidney Foundation. *Kidney Early Evaluation Program Publications*. 2013; http://www. kidney.org/news/keep/. Last accessed October 9, 2014.
- Wu HY, Huang JW, Peng YS, et al. Microalbuminuria screening for detecting chronic kidney disease in the general population: a systematic review. *Ren Fail*. 2013;35(5):607-614. http:// dx.doi.org/10.3109/0886022X.2013.779907. PMID:23534678.
- 22. Atthobari J, Asselbergs FW, Boersma C, et al; PREVEND IT Study Group. Costeffectiveness of screening for albuminuria with subsequent fosinopril treatment to prevent cardiovascular events: A pharmacoeconomic analysis linked to the prevention of renal and vascular endstage disease (PREVEND) study and the prevention of renal and vascular endstage disease intervention trial (PREVEND IT). *Clin Ther.* 2006;28(3):432-444. http:// dx.doi.org/10.1016/j.clinthera.2006.03.012. PMID:16750458.
- Hoerger TJ, Wittenborn JS, Segel JE, et al; Centers for Disease Control and Prevention CKD Initiative. A health policy model of CKD: 2. The cost-effectiveness of microalbuminuria screening. *Am J Kidney Dis.* 2010;55(3):463-473. http://dx.doi.org/10.1053/j.ajkd.2009.11.017. PMID:20116910.
- Ewald B, Attia J. Which test to detect microalbuminuria in diabetic patients? A systematic review. Aust Fam Physician. 2004;33(7):565-

Albuminuria screening in MESA - Tanner et al

567, 571. PMID:15301181.

- Moyer VA. Screening for chronic kidney disease: US Preventive Services Task Force recommendation statement. *Ann Intern Med.* 2012;157(8):567-570. http://dx.doi. org/10.7326/0003-4819-157-8-201210160-00533. PMID:22928170.
- Sinha SK, Shaheen M, Rajavashisth TB, Pan D, Norris KC, Nicholas SB. Association of race/ethnicity, inflammation, and albuminuria in patients with diabetes and early chronic kidney disease. *Diabetes Care*. 2014;37(4):1060-1068. http://dx.doi.org/10.2337/dc13-0013. PMID:24550221.
- US Renal Data System. 2012 Annual Data Report: Atlas of Chronic Kidney Disease and End-stage Renal Disease in the United States. 2012. http://www.usrds.org/atlas12.aspx. Last accessed: September 14, 2015.
- Bellary S, O'Hare JP, Raymond NT, et al. Premature cardiovascular events and mortality in south Asians with type 2 diabetes in the United Kingdom Asian Diabetes Study - effect of ethnicity on risk. *Curr Med Res Opin*. 2010;26(8):1873-1879. http://dx.doi.org/10.11 85/03007995.2010.490468. PMID:20528107.
- Burgard SA, Hawkins JM. Race/Ethnicity, educational attainment, and foregone health care in the United States in the 2007-2009 recession. *Am J Public Health*. 2014;104(2):e134-e140. http://dx.doi.org/10.2105/AJPH.2013.301512. PMID:24328647.
- O'Seaghdha CM, Hwang SJ, Upadhyay A, Meigs JB, Fox CS. Predictors of incident albuminuria in the Framingham Offspring cohort. *Am J Kidney Dis*. 2010;56(5):852-860. http://dx.doi.org/10.1053/j.ajkd.2010.04.013. PMID:20599306.
- Vaidya V, Partha G, Howe J. Utilization of preventive care services and their effect on cardiovascular outcomes in the United States. *Risk Manag Healthc Policy*. 2011;4:1-7. http://dx.doi.org/10.2147/RMHP.S15777. PMID:22312222.
- Gerritsen AA, Devillé WL. Gender differences in health and health care utilisation in various ethnic groups in the Netherlands: a cross-sectional study. *BMC Public Health.* 2009;9(1):109. http://dx.doi.org/10.1186/1471-2458-9-109. PMID:19379499.
- 33. Harward DH, Bomback AS, Jennette CE, Amamoo MA, Falk R. The Kidney Education Outreach Program's community-based screenings: participants' demographics and screening results. N C Med J. 2009;70(6):507-512. PMID:20198833.
- 34. Islam TM, Fox CS, Mann D, Muntner P. Age-related associations of hypertension and diabetes mellitus with chronic kidney disease. *BMC Nephrol.* 2009;10(1):17. http:// dx.doi.org/10.1186/1471-2369-10-17. PMID:19563681.
- 35. Chen J, Muntner P, Hamm LL, et al. The metabolic syndrome and chronic kidney disease in

U.S. adults. Ann Intern Med. 2004;140(3):167-174. http://dx.doi.org/10.7326/0003-4819-140-3-200402030-00007. PMID:14757614.

- 36. Hogan SL, Vupputuri S, Guo X, et al. Association of cigarette smoking with albuminuria in the United States: the third National Health and Nutrition Examination Survey. *Ren Fail.* 2007;29(2):133-142. http:// dx.doi.org/10.1080/08860220601098888. PMID:17365926.
- Muntner P, He J, Hamm L, Loria C, Whelton PK. Renal insufficiency and subsequent death resulting from cardiovascular disease in the United States. *J Am Soc Nephrol.* 2002;13(3):745-753. PMID:11856780.