

# Health Behaviors in Younger and Older Adults With CKD: Results From the CRIC Study



Sarah J. Schrauben<sup>1,2</sup>, Jesse Y. Hsu<sup>2,3</sup>, Julie Wright Nunes<sup>4</sup>, Michael J. Fischer<sup>5</sup>, Anand Srivastava<sup>6</sup>, Jing Chen<sup>7</sup>, Jeanne Charleston<sup>8</sup>, Susan Steigerwalt<sup>4</sup>, Thida C. Tan<sup>9</sup>, Jeffrey C. Fink<sup>10</sup>, Ana C. Ricardo<sup>5</sup>, James P. Lash<sup>5</sup>, Myles Wolf<sup>11</sup>, Harold I. Feldman<sup>1,2,3</sup> and Amanda H. Anderson<sup>2,3</sup>; and the CRIC Study Investigators<sup>12</sup>

<sup>1</sup>Division of Renal, Electrolyte, and Hypertension, University of Pennsylvania, Philadelphia, Pennsylvania, USA; <sup>2</sup>Center for Clinical Epidemiology and Biostatistics, Perelman School of Medicine at the University of Pennsylvania, Philadelphia, Pennsylvania, USA; <sup>3</sup>Department of Biostatistics, Epidemiology, and Informatics, Perelman School of Medicine at the University of Pennsylvania, Philadelphia, Pennsylvania, USA; <sup>4</sup>Department of Medicine, University of Michigan, Ann Arbor, Michigan, USA; <sup>5</sup>Department of Medicine, University of Illinois at Chicago, and Center of Management for Complex Chronic Healthcare, Jesse Brown Veterans Affairs Medical Center, Chicago, Illinois, USA; <sup>6</sup>Department of Medicine, Northwestern University Feinberg School of Medicine, Chicago, Illinois; <sup>7</sup>Department of Medicine, Tulane University, New Orleans, Louisiana, USA; <sup>8</sup>Johns Hopkins University, Baltimore, Maryland, USA; <sup>9</sup>Kaiser Permanente, Oakland, California, USA; <sup>10</sup>Department of Medicine, University, Durham, North Carolina, USA;

**Introduction**: A cornerstone of kidney disease management is participation in guideline-recommended health behaviors. However, the relationship of these health behaviors with outcomes, and the identification of barriers to health behavior engagement, have not been described among younger and older adults with chronic kidney disease.

**Methods**: Data from a cohort study of 5499 individuals with chronic kidney disease was used to identify health behavior patterns with latent class analysis stratified by age <65 and  $\geq 65$  years. Cox models, stratified by diabetes, assessed the association of health behavior patterns with chronic kidney disease (CKD) progression, atherosclerotic events, and death. Logistic regression was used to assess for barriers to health behavior engagement.

**Results:** Three health behavior patterns were identified: 1 "healthy" pattern, and 2 "less healthy" patterns comprising 1 pattern with more obesity and sedentary activity and 1 with more smoking and less obesity. Less healthy patterns were associated with an increased hazard of poor outcomes. Among participants <65 years of age, the less healthy patterns (vs. healthy pattern) was associated with an increased hazard of death in diabetic individuals (hazard ratio [HR] = 2.17, 95% confidence interval [CI] = 1.09–4.29; and HR = 2.50, 95% CI = 1.39–4.50) and cardiovascular events among nondiabetic individuals (HR = 1.49, 95% CI = 1.04–2.43; and HR = 2.97, 95% CI = 1.49–5.90). Individuals with the more obese/ sedentary pattern had an increased risk of CKD progression in those who were diabetic (HR = 1.34, 95% CI = 1.13–1.59). Among older adults, the less healthy patterns were associated with increased risk of death (HR = 2.97, 95% CI = 1.43–6.19; and HR = 3.47, 95% CI = 1.48–8.11) in those who were nondiabetic. Potential barriers to recommended health behaviors include lower health literacy and self-efficacy.

**Conclusion**: Identifying health behavior patterns and barriers may help target high-risk groups for strategies to increase participation in health behaviors.

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KEYWORDS: all-cause death cardiovascular outcomes; chronic renal failure; chronic renal insufficiency; CKD progression; health behavior; self-management

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**Correspondence**: Sarah J. Schrauben, Founders 1 - Renal Division, 3400 Spruce Street, Philadelphia, Pennsylvania 19103, USA. E-mail: Sarah.Schrauben@uphs.upenn.edu

<sup>12</sup>The CRIC Study Investigators are Lawrence J. Appel, Harold I. Feldman, Alan S. Go, Jian He, John W. Kusek, James P. Lash, Panduranga S. Rao, Mahboob Rahman, and Raymond R. Townsend.

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T he number of adults 65 years and older in the United States is projected to be more than 20% of the population by the year 2030.<sup>1</sup> With each advancing decade of life, CKD incidence and prevalence increases.<sup>2</sup> CKD increases the risk of developing cardiovascular disease (CVD), progressing to end-stage renal disease (ESRD), and premature death.<sup>2,3</sup> Age is an important effect modifier of these risk associations, with older adults less frequently progressing to ESRD but experiencing more cardiovascular mortality than younger adults.<sup>4,5</sup>

Current CKD guidelines are not age specific and broadly recommend patients to actively participate in health behaviors to manage their disease and mitigate its sequelae; these behaviors include monitoring blood pressure, abstaining from smoking, maintaining a healthy weight, consuming a healthy diet, and participating in physical activity.<sup>6–8</sup> The effectiveness of these health behavior recommendations in CKD is relatively unknown, as they were informed by research in the general population.<sup>9</sup> In addition, to date, there are limited data examining the health behavior practices of adults with CKD.<sup>10</sup> In particular, even though older adults likely face unique challenges to health behavior engagement, such as higher rates of comorbid conditions, frailty, and decreased functional status, little consideration has been given to whether adoption of health behaviors differs for older (vs. younger) adults. Such information is important to understand gaps in care and to develop age-appropriate recommendations and interventions to help all patients with CKD. The primary goals of this study were to examine the level of engagement in recommended health behaviors, to identify potential barriers to this engagement, and to elucidate the association of clustering of health behaviors with important outcomes separately among younger and older adults with CKD.

# **METHODS**

This study utilized data from the Chronic Renal Insufficiency Cohort (CRIC) Study, a multicenter, prospective, observational cohort of 5,499 men and women with mild-to-moderate CKD in the United States.<sup>11,12</sup> Participants were recruited from 7 clinical centers during 2 phases of the study: (i) phase I, from 2003 to 2008 (3939 participants); and (ii) phase III, from 2013 to 2015 (1560 participants). The study protocol was approved by the institutional review boards of all participating centers and is in accordance with the Declaration of Helsinki. All participants provided informed consent. The full inclusion and exclusion criteria have been previously reported.<sup>11,12</sup> In brief, major eligibility criteria for phase I included adults aged 21 to 74 years with age-based estimated glomerular filtration rate (eGFR) of 20 to 70 ml/min per 1.73 m<sup>2</sup>; entry into phase III required eGFR 45 to 70 ml/min per 1.73 m<sup>2</sup> and age 45 to 79 years. Exclusion criteria included inability to provide consent and the presence of certain severe conditions.

Table 1. Measures of Health Behaviors in CKD

Health Behavior Measures	Recommended	Not recommended
Cigarette smoking	Never/former	Current
Body mass index (kg/m <sup>2</sup> )	20 to <25	${<}20 \text{ or }{\geq}25$
Physical activity (min/wk)	$\geq$ 150 moderate, $\geq$ 75 vigorous, or moderate + vigorous $\geq$ 150	Less than recommended
Healthy diet score <sup>a</sup>	3 or 4	≤ 2
Blood pressure (mm Hg)	≤140/90	>140/90
Blood glucose control	HbA1c ≤7%	$\rm HbA1c > 7\%$

<sup>a</sup>Healthy diet score: based on 12-month diet history of fruits, vegetables, fish, grains, and sweets/sweetened beverages. A diet score of 3 or 4 was "recommended," and  ${\leq}2$  was "not recommended."

The selection of recommended health behavior measures was guided by previous literature and guidelines (Table 1).<sup>8,13,14</sup> Briefly, body mass index (BMI) served as a proxy of weight maintenance, was dichotomized into "recommended" for a BMI of 20 to  $<25 \text{ kg/m}^2$  and "not recommended" if otherwise.<sup>8,9</sup> A "healthy" diet score (range 0-4) was constructed from a 12-month diet history<sup>15</sup> based on fruits, vegetables, fish, grains, and sweets/sweetened beverages.<sup>9</sup> A diet score >2 was "recommended" and  $\leq 2$  "not recommended." Reported weekly physical activity was categorized as "recommended" for moderate activity  $\geq 150$  minutes or vigorous  $\geq$ 75 minutes, or moderate plus vigorous  $\geq$ 150 minutes<sup>9</sup> and "not recommended" if otherwise. Past use or never use of tobacco was "recommended" and current use "not recommended." A mean blood pressure was a proxy for adherence to antihypertensive treatment, with  $\leq 140$  mm Hg systolic blood pressure and  $\leq$ 90 mm Hg diastolic blood pressure defined as "recommended" and >140/90 mm Hg as "not recommended."<sup>16</sup> A hemoglobin A1c (HbA1c) value among individuals with diabetes was a proxy for blood glucose management, with  $\leq$ 7.0% "recommended" and >7.0% "not recommended."<sup>17</sup>

## Other Study Variables

Age, gender, race/ethnicity, education, income, insurance, duration of CKD awareness, and medical history were collected through self-report. Comorbidities reported were used to construct an index using the Charlson Comorbidity Index scoring system.<sup>18</sup> Assessments of health literacy, physical health, general health status, depressive symptoms, and cognitive function were collected through validated questionnaires. Health literacy was determined by a score on the short version of the Test of Functional Health Literacy Assessment.<sup>19</sup> Reported health status (including mental and physical health) and health-related quality of life were collected using the Kidney Disease Quality of Life Short Form 36.<sup>20</sup> The Beck Depression Inventory, which is widely used in CKD, assessed symptoms of depression.<sup>21–23</sup> Global cognitive function was assessed with the Modified Mini-Mental State Examination.<sup>24</sup>

Several measures were available only in the phase III Cohort, including social support, which was collected using the Lubben Social Network Scale,<sup>25</sup> general disease management self-efficacy, and self-efficacy in the patient—provider interaction, which were both assessed using 5-item validated instruments.<sup>26,27</sup> Blood samples were used to measure HbA1c, and to estimate eGFR with a CRIC-specific equation using serum creatinine and cystatin C.<sup>28</sup> Cystatin C may be less related to muscle mass than creatinine, and therefore may have a particular advantage in older adults.<sup>29</sup> A 24-hour urine collection was used to assess proteinuria.

The following outcomes were evaluated: (i) CKD progression, defined as a 50% decrease in eGFR from baseline or occurrence of ESRD (incident dialysis or kidney transplantation); (ii) atherosclerotic cardiovascular events (myocardial infarction, stroke, or peripheral arterial disease); and (iii) death from any cause. Progression of CKD was defined by achieving the composite endpoint of development of ESRD or halving of eGFR from baseline. Ascertainment of time to eGFR halving was imputed assuming a linear decline in kidney function between annual visits.<sup>30</sup> End-stage renal disease was defined as the initiation of maintenance dialysis or kidney transplantation, and was ascertained through self-report or report from family members of deceased participants. Information collected on ESRD was supplemented by data from the United States Renal Data System. Cardiovascular events were adjudicated by 2 physician reviewers using hospital records.<sup>11</sup> Deaths were ascertained from reports by next of kin, death certificates, obituaries, reviews of hospital records, and the Social Security Death Master File. Participants were followed until the occurrence of death, withdrawal from the study, or the end of the follow-up period (mid-2014).

# **Statistical Analysis**

We described the study population using mean, SD, or median and interquartile range for continuous variables, and frequency and proportion for categorical variables. The Pearson  $\chi^2$  or Fisher exact test and analysis of variance or Kruskal-Wallis test were used to compare categorical and continuous variables, respectively. We examined for patterns of health behavior engagement, as many CKD patients engage only in certain behaviors. To account for clustering of behaviors, we used latent class analysis (LCA), a wellvalidated statistical technique that uses mixture modeling to identify unobserved clusters based on variables of interest (i.e., the measures of health behaviors) without mandating consideration of the outcome.<sup>31</sup> To determine the optimal number of clusters (or patterns), latent class models with successive numbers of patterns were compared, and the bestfitting model was chosen. Our previous publication describes this process in more detail.<sup>32</sup> A priori, we chose to stratify the LCA by age greater or less than 65 years, as many studies have demonstrated effect modification at 65 years of age in relation to the clinical outcomes of interest. Further information on the agestratified LCA is provided in the Supplementary Data. A sensitivity LCA stratified by age group was performed with the recommended blood pressure defined as  $\leq 130$  mm Hg systolic pressure and  $\leq 80$  mm Hg diastolic pressure.

To evaluate for potential barriers to health behavior engagement, we analyzed the association of psychosocial and clinical factors in multinomial logistic regression models with reported prevalence ratios to approximate relative risk. The covariates included in the regression models were thought to influence participation in health behaviors and/or have been associated with self-management behaviors in prior studies. Potential barriers previously identified in other chronic diseases include inadequate social support, low self-efficacy, physical limitations or low levels of physical functioning, presence of depression, low level of health literacy, lack of knowledge about one's medical conditions, and presence of comorbid diseases.<sup>33–40</sup> Subjective well-being has also been linked to self-management in individuals with ESRD.<sup>41</sup> In addition, social and systems-based characteristics, such as uninsured/underinsured, may play major roles in increasing complexity of patients' ability to adopt health behaviors.<sup>42,43</sup> Education, income, and insurance were included as key measures of social and systemsbased characteristics. Cognition was included, as it may influence the ability or motivation of individuals with CKD to participate in health behaviors.

Cox proportional hazards models were used to generate crude and adjusted HRs using the phase I cohort only, stratified by diabetes, given the observed differences of CKD progression within the CRIC population. Insufficient follow-up time did not allow for analyses using the phase III cohort. Sample sizes included in each analytical step are provided in Supplementary Figure S1. Secondary analyses were performed using Cox proportional hazards models to estimate the risk of outcomes with individual measures of health behaviors. For each outcome, hierarchical modeling was performed with sequential adjustment for socio-demographic covariates including gender, race/ethnicity, clinical center, education, eGFR, proteinuria, diabetes, and history of CVD. In the secondary analyses, a final adjustment was made for the individual measures of health behaviors. For analyses of cardiovascular events, death was a competing event. The assumptions of proportionality were tested in the fully adjusted model; if violated, the covariates were modified by an interaction with time, and for the health behavior patterns, HRs were reported in intervals of follow-up time. Statistical analyses were conducted using STATA version 14.2, using the LCA Stata Plugin, Version 1.2 and the LCA Bootstrap, Version 1.0 (Penn State Methodology Center, University Park, PA).<sup>44–46</sup>

# RESULTS

Characteristics of the study participants are listed in Tables 2 and 3. Participants <65 years of age had a higher level of education, income, cognition, and fewer comorbidities, compared to those  $\geq 65$  years. The younger age group reported lower disease self-efficacy and more depressive symptoms. Three similar patterns of health behavior engagement were identified among participants <65 and  $\geq 65$  years of age based on model fit indices (Supplementary Table S1). In both age groups, the 3 health behavior patterns were characterized as follows: 1 "healthy" pattern, and 2 "less healthy" patterns comprising 1 pattern with more obesity and sedentary activity (more obese/sedentary) and the other with more smoking and less obesity (less obese/smoker) (Supplementary Figure S2). In both age groups, individuals with the healthy pattern had high levels of engagement in recommended health behaviors; those with the more obese/sedentary pattern had low levels of recommended BMI and physical activity; and those with the less obese/smoker pattern had high levels of smoking and recommended BMI. Blood glucose control was similar across the patterns. Among individuals <65 years of age, the healthy pattern was the most prevalent (55.7%), the more obese/sedentary pattern the next most prevalent (40.1%), and the less obese/smoker pattern the least prevalent (4.2%). Among the older age group, the most prevalent was the more obese/sedentary pattern (47.9%), the healthy pattern was less prevalent (42.3%), and, the less obese/ smoker pattern was the least prevalent (9.8%).

In both age groups, individuals with the healthy pattern had the highest levels of education, income, self-efficacy, renal function, cognition, and physical functioning, and the lowest levels of depressive symptoms and CVD history (Tables 2 and 3). Comorbidity scores were similar across the health behavior patterns. In older adults, the less healthy patterns, compared to the healthy pattern, were associated with inadequate health literacy, less social support, as well lower physical functioning as mental and (Supplementary Table S2).

Overall, event rates were increased in individuals with the less healthy patterns, and among those with

diabetes (Supplementary Table S3). The proportional hazards assumption was violated for death among those <65 years of age; therefore the HR and 95% CI are reported in intervals of follow-up time in which the hazard appeared to change (before and after 3 years for those without diabetes and 5 years for those with diabetes). Results are reported for the final multivariable-adjusted model (Figures 1 and 2; Supplementary Table S4). Among individuals without diabetes, increased risk of death was seen in the more obese/sedentary pattern (HR = 2.17, 95% CI = 1.09-4.29, <3 years of follow-up), and of cardiovascular events in both the more obese/sedentary and less obese/ smoker patterns (HR = 1.59, 95% CI = 1.04-2.43, and HR = 2.97, 95% CI = 1.49–5.90, respectively). Among individuals with diabetes, increased risk for death was observed among the more obese/sedentary and less obese/smoker patterns (HR = 1.37, 95% CI = 1.01-1.86and HR = 2.50, 95% CI = 1.39–4.50, respectively for >5 years of follow-up) and CKD progression in the more obese/sedentary pattern (HR = 1.34, 95% CI = 1.13-1.59). The proportional hazards assumption was also violated for the outcome of death among older adults without diabetes, and the HR (95% CI) reported before and after 3 years of follow-up time. Increased rates of death were seen in the more obese/sedentary and less obese/smoker patterns (HR = 2.97, 95% CI = 1.43-6.19and HR = 3.47, 95% CI = 1.48-8.11, respectively for <3 years), and CKD progression for the less obese/ smoker pattern (HR = 1.85, 95% CI = 1.07-3.22).

#### Individual Health Behaviors

In individuals <65 years of age without diabetes, current smoking and uncontrolled blood pressure were associated with increased risk of death (HR = 2.62, 95% CI = 1.81–3.79, and HR = 1.53, 95% CI = 1.06– 2.19, respectively) as well as increased risk of atherosclerotic events (HR = 2.60, 95% CI = 1.57-4.29, and HR = 2.11, 95% CI = 1.33-3.35, respectively) (Figure 3, Supplementary Table S5). Among individuals <65 years of age with diabetes, current smoking and less physical activity were associated with increased risk of death (HR = 1.47, 95% CI = 1.10-1.97, and HR = 1.40, 95% CI = 1.09-1.80, respectively); uncontrolled blood pressure and less healthy diet were associated with increased risk of CKD progression (HR = 1.65, 95% CI = 1.34–2.04, and HR = 1.32, 95% CI = 1.08-1.62, respectively); and less physical activity and increased HbA1c were associated with increased risk of atherosclerotic events (HR =1.41, 95% CI = 1.06–1.87, and HR = 1.65, 95% CI = 1.21–2.23, respectively). Of note, a BMI  $<20 \text{ kg/m}^2$ or  $\geq 25 \text{ kg/m}^2$  was associated with decreased risk of CKD progression in those with and without diabetes

	Age <65 years					Age ≥65 years						
Characteristic	Overall (N = 3552)	Healthy pattern n = 1979 (55.7%)	More obese/sedentary pattern n = 1423 (40.1%)	Less obese/smoker pattern n = 150 (4.2%)	P value <sup>a</sup>	Overall (N = 1947)	Healthy pattern n = 823 (42.3%)	More obese/sedentary pattern $n = 933 (47.9\%)$	Less obese/smoker pattern N = 191 (9.8%)	P value		
Age, mean (SD)	54 (9)	54 (9)	53 (9)	53 (8)	< 0.001	70 (3)	70 (4)	70 (3)	69 (3)	0.01		
Gender, male, %	55	56	54	59	0.19	42	61	56	57	0.10		
Race, %					< 0.001					0.001		
Non-Hispanic white	39	41	37	30		45.9	50.3	43.6	37.0			
Non-Hispanic black	45	42	47	65		42	38	44	52			
Hispanic	12	13	12	3		9	8	10	7			
Other	4	4	4	2		3	4	3	3			
Education, %					< 0.001					< 0.001		
High school or less	36	33	39	50		41	33	46	49			
College or more	64	67	61	50		59	67	54	51			
Insurance, %					< 0.001					0.01		
Medicaid	24	20	29	42		13	10	15	15			
Medicare/VA	42	42	41	40		80	82	79	80			
Private/commercial	34	38	30	19		7	9	6	5			
Income, %					< 0.001					< 0.001		
≤\$20,000	37	33	40	52		33	26	36	50			
\$20–50,000	26	26	27	23		35	35	36	27			
>\$50,000	37	41	22	25		32	39	28	23			

Table 2. Demographic characteristics of latent class-defined health behavior engagement patterns in the CRIC phase I and III cohorts overall, and by age group (<65 years and ≥65 years)

Values for continuous data given as mean (SD); values for categorical data given as percentage.

<sup>a</sup>P value for test of the difference across behavior engagement patterns.

			Age <65 years					Age ≥65 years					
Characteristic	Overall (N = 3552)	Healthy pattern n = 1979 (55.7%)	More obese/sedentary pattern $n = 1423$ (40.1%)	Less obese/smoker pattern n = 150 (4.2%)	P value <sup>a</sup>	Overall (n = 1947)	Healthy pattern $n = 823 (42.3\%)$	More obese/sedentary pattern n = 933 (47.9%)	Less obese/smoker pattern n = 191 (9.8%)	P value			
Adequate Health Literacy <sup>b</sup> , %	87	87	88	84	0.29	83	83	81	75	0.08			
Duration of CKD awareness, %					0.15					0.45			
Unknown/≤1 yr	45	46	43	42		48	48	49	53				
>1 yr	56	54	57	58		52	52	51	47				
Disease self-efficacy score, mean (SD)	40 (9)	41 (9)	39 (10)	39 (10)	0.03	42 (8)	43 (7)	40 (9)	42 (7)	0.001			
Patient provider efficacy score, mean (SD)	21 (4)	21 (4)	21 (4)	21 (4)	0.73	21 (4)	22 (4)	21 (4)	21 (5)	0.09			
Social Support score <sup>c</sup> , mean (SD)	N/A	N/A	N/A	N/A		16 (6.)	17 (6)	16 (6)	15 (6)	0.02			
Reported general health excellent/good, %	57	62	51	57	< 0.001	64	73	58	59	< 0.001			
Cognition score, mean (SD)	92 (8)	92 (8)	92 (8)	91 (8)	0.08	89 (9)	91 (8)	89 (10)	88 (10)	< 0.001			
BDI depressive symptom score, mean (SD)	9 (9)	8 (8)	10 (9)	11 (9)	< 0.001	7 (7)	6 (6)	8 (7)	8 (6)	< 0.001			
BMI, kg/m <sup>2</sup> , mean (SD)	33 (8)	33 (8)	33 (9)	29 (7)	< 0.001	32 (7)	31 (6)	34 (7)	27 (6)	< 0.001			
Any CVD, %	29	28	31	35	< 0.001	42	38	44	48	0.01			
eGFR, ml/min per 1.73 m <sup>2</sup> , mean (SD)	50 (18)	51 (17)	48 (18)	46 (17)	< 0.001	47 (15)	49 (15)	46 (15)	46 (15)	< 0.001			
Proteinuria, g/24 h, mean (SD)	1.2 (2.5)	1.1 (2.3)	1.3 (2.6)	1.3 (2.2)	< 0.001	0.7 (1.5)	0.6 (1.1)	0.7 (1.6)	0.9 (1.1)	0.22			
Charlson Comorbidity Index score, mean (SD)	4.9 (2.1)	4.9 (2.1)	4.9 (2.1)	5.1 (2.0)	0.52	6.8 (1.7)	6.6 (1.7)	7.0 (1.7)	6.6 (1.7)	< 0.001			
KDQOL Mental Composite Score, mean (SD)	49.1 (10.9)	49.9 (10.4)	48.2 (11.3)	48.0 (11.2)	< 0.001	52.4 (9.5)	53.4 (8.6)	51.4 (10.1)	52.8 (9.5)	< 0.001			
KDQOL Physical Composite Score, mean (SD)	41.1 (11.7)	42.3 (11.3)	39.5 (12.1)	40.9 (11.3)	< 0.001	40.5 (11.1)	43.1 (10.4)	38.3 (11.1)	39.7 (11.4)	< 0.001			
Total number of medications, mean (SD)	8.8 (4.8)	8.9 (4.8)	8.8 (4.8)	7.8 (4.4)	0.02	10.3 (4.5)	10.1 (4.4)	10.6 (4.5)	9.8 (5.1)	0.003			
Health behavior measures, %													
Recommended body mass index	14	12	14	33	< 0.001	11	14	0	55	< 0.001			
Nonsmoking	85	97	77	0	< 0.001	92	94	100	42	< 0.001			
Controlled blood pressure	72	77	65	61	< 0.001	72	73	70	73	0.29			
Recommended diet	33	58	5	6	< 0.001	31	33	32	20	0.004			
Recommended physical activity	41	66	0	100	< 0.001	42	100	0	0	< 0.001			
HbA1c <7.0% (if diabetic)	36	37	37	30	0.52	51	53	49	57	0.29			

Table 3. Clinical characteristics of latent class-defined health behavior engagement patterns in the CRIC phase I and III cohorts overall, and by age group (<65 years and ≥65 years)

Values for continuous data given as mean (SD); values for categorical data given as percentage. BDI, Beck Depression Inventory; CVD, cardiovascular disease; eGFR, estimated glomerular filtration rate; KDQOL, Kidney Disease Quality of Life; NA, not available.

<sup>a</sup>*P* value for test of the difference across behavior engagement patterns.

<sup>b</sup>Adequate health literacy was defined by score >23 on the Short Test of Functional Health Literacy.<sup>19</sup>

<sup>c</sup>Assessed only in older adults.



**Figure 1.** Tree display of sample size by age group, diabetes status, and health behavior patterns with reported hazard ratios (HR) and *P* values for each clinical outcome. Reference group is the healthy pattern group. \*HR is reported from baseline to 3 years of follow-up and thereafter for participants without diabetes and from baseline to 5 years of follow-up for those with diabetes, as the proportionality of hazard assumption was violated, with the hazard of death appearing to change at selected intervals.

(HR = 0.66, 95% CI = 0.49–0.89, and HR = 0.49, 95% CI = 0.36–0.67).

In individuals  $\geq$ 65 years of age without diabetes, current smoking was associated with increased risk of death (HR = 1.88, 95% CI = 1.16-3.03), and uncontrolled blood pressure was associated with increased atherosclerotic events (HR = 1.85, 95% CI = 1.11-3.09) (Figure Supplementary Table S5). Among 3, individuals  $\geq 65$  years of age with diabetes, uncontrolled blood glucose and blood pressure were associated with increased risk of death (HR = 1.37, 95% CI = 1.03-1.83, and HR = 1.77, 95% CI = 1.28-2.45, respectively), and uncontrolled blood pressure was associated with increased CKD progression (HR = 2.41, 95% CI = 1.66–3.52). Notably, a BMI <20 kg/m<sup>2</sup> or  $\geq 25 \text{ kg/m}^2$  was associated with decreased risk of death (HR = 0.55, 95% CI = 0.33-0.92), and a less healthy diet was associated with decreased risk of CKD progression (HR = 0.65, 95% CI = 0.44-0.95).

In the sensitivity analyses in which recommended blood pressure was defined as  $\leq$ 130 mm Hg systolic and  $\leq$ 80 mm Hg diastolic, slightly different behavior patterns were identified from the primary analysis (Supplementary Tables S6–S11). Behavior patterns in individuals with a lower prevalence of recommended health behaviors were associated with poorer clinical characteristics, and the health behavior patterns with less obesity and more smoking similarly trended toward an increased hazard of poor clinical outcomes, with a significant finding for CKD progression for those <65 years of age.

# CONCLUSION

In a multicenter, diverse cohort of individuals with CKD, we identified 3 similar patterns of health behavior engagement among younger and older adults. The health behavior engagement patterns consisted of 1 "healthy" pattern, and 2 "less healthy" patterns comprising 1 pattern with more obesity and sedentary activity and another with more smoking and less obesity. Overall, the less healthy behavior patterns were associated with increased hazard of poor clinical outcomes. Potential barriers to recommended health

## (a) Less than 65 years of age

Outcome	Nu Behavior engagement pattern	umber of events (rate per 10 person- years)	s Hazard ratio (95% C	I) Hazard ratio (95% CI)
	More obese/sedentary – ND (<3 yr)	26 (1.5)	2.17 (1.09-4.29)	▶ <b>──</b>
	More obese/sedentary $-$ ND (>3 yr)	59 (1.8)	1.13 (0.78–1.62)	┝┼╋──┤
	More obese/sedentary – DM (<5 yr)	88 (3.8)	1.21 (0.89–1.64)	<b>⊢</b> ∎1
Death	More obese/sedentary – DM (>5 yr)	82 (6.0)	1.37 (1.01–1.86)	<b>⊢</b> ∎I
	Less Obese/Smoker – ND (<3 yr)	5 (2.4)	2.13 (0.75–6.02)	
	Less Obese/Smoker – ND (>3 yr)	14 (4.2)	1.73 (0.96–3.13)	HI
	Nonobese/Smoker – DM (<5 yr)	6 (3.1)	0.73 (0.27–2.01)	
	Nonobese/Smoker – DM (>5 yr)	14 (11.7)	2.50 (1.39-4.50)	F
	More obese/sedentary - ND	154 (4.43)	1.22 (0.95–1.55)	<b>⊢</b> ∎1
CKD	More obese/sedentary – DM	272 (12.7)	1.34 (1.13–1.59)	<b>⊢</b> ∎→1
Progression	Less Obese/Smoker – ND	21 (6.2)	1.12 (0.70–1.79)	F
	Less Obese/Smoker – DM	27 (14.7)	1.46 (0.96–2.21)	<b>⊢_</b> ∎i
	More obese/sedentary – ND	52 (1.17)	1.59 (1.04–2.43)	<b>⊢</b>
Atherosclerotic	More obese/sedentary – DM	117 (3.7)	1.26 (0.98–1.63)	<b>⊢</b> ∎1
events	Less Obese/Smoker – ND	11 (2.7)	2.97 (1.49–5.90)	<b>—</b>
	Less Obese/Smoker – DM	13 (4.7)	1.40 (0.75–2.61)	
			0	.5 1.0 1.5 2.0 2.5 3.0 3.5 4.0 4.5 5.

## (b) 65 Years and older

Outcome	Nur Behavior engagement pattern	mber of events (rate per 10 person- years)	Hazard ratio (95% (	CI)	1		Haza	rd ratio	o (95%	CI)			
	More obese/sedentary – ND (<3 yr)	26 (4.2)	2.97 (1.43–6.19)			<b>—</b>							7
	More obese/sedentary - ND (>3 yr)	48 (4.9)	0.95 (0.64-1.40)										
	More obese/sedentary – DM	137 (6.11)	0.89 (0.67–1.17)	-									
Death	Less Obese/Smoker – ND (<3 yr)	12 (5.5)	3.47 (1.48-8.11)			-							-
	Less Obese/Smoker – ND (>3 yr)	24 (7.4)	1.54 (0.94–2.54)		+	-		-					
	Less-Obese/Smoker – DM	24 (9.3)	1.18 (0.72–1.93)	-	-		-						
	More obese/sedentary - ND	49 (4.1)	1.06 (0.70–1.61)	<b>-</b>	-								
CKD	More obese/sedentary – DM	118 (8.1)	0.97 (0.69–1.34)	F	-	-							
Progression	Less Obese/Smoker – ND	21 (5.3)	1.85 (1.07–3.22)		<b>-</b>		•						
	Less Obese/Smoker – DM	76 (6.6)	0.62 (0.32-1.24)	-	+								
	More obese/sedentary - ND	29 (2.0)	0.61 (0.38–1.00)	-	-								
Atherosclerotic	More obese/sedentary - DM	76 (4.0)	1.12 (0.76–1.64)	H	-								
events	Less Obese/Smoker - ND	16 (3.4)	1.36 (0.74–2.50)	F		•							
	Less Obese/Smoker – DM	13 (5.9)	1.32 (0.67–2.59)	-	-	-							
	-		C	).5	1.0	1.5	2.0	2.5	3.0	3.5	4.0	4.5	

**Figure 2.** (a,b) Forest plot of hazard ratios (HRs) (95% confidence intervals [CIs]) of clinical outcomes by health behavior pattern for participants without diabetes (ND) (white rows) and those with diabetes (DM) (gray rows) for participants (a) <65 years of age and (b)  $\geq 65$  years of age. Reference group is the healthy pattern group for both DM and ND. HR is reported from baseline to 3 years of follow-up and after for ND and from baseline to 5 years of follow-up and thereafter for DM, as the proportionality of hazards assumption was violated, with the hazard of death appearing to change at selected intervals.

behavior engagement included social determinants of health, as well as depressive symptoms, inadequate health literacy, poor social support, and a higher burden of comorbidities. Previous studies regarding other chronic diseases have shown a link between health behaviors and poor outcomes.<sup>47,48</sup> This is the first study to examine the health behavior pattern–outcome link separately

Outcome	Individual Behavior Measures	Hazard Ratio (95% CI)	0
	Current smoking – ND	2.62 (1.81-3.79)	► <b>■</b>
	Current smoking DM	1.47 (1.10–1.97)	
	BMI <20 and >25 (kg/m <sup>2</sup> ) — ND	1.00 (0.66–1.52)	▶ <b>•</b>
	BMI <20 and >25 (kg/m <sup>2</sup> ) — DM	1.53 (1.06–2.19)	
Death	BP >140/90 mm Hg — ND	1.09 (0.84-1.42)	┝┼┳──┤
	BP >140/90 mm Hg — DM	1.09 (0.84–1.42)	
	Not recommended Diet - ND	1.13 (0.77–1.67)	
	Not recommended Diet — DM	1.28 (0.99–1.64)	┝─┲─┤
	Not Recommended Physical Activity - ND	1.01 (0.72–1.42)	
	Not Recommended Physical Activity - DM	1.40 (1.09–1.80)	<b>⊢</b>
	HbA1c > 7.0% – ND	n/a	
	HbA1c > 7.0% – DM	1.06 (0.82–1.37)	<b>⊢</b> ∎1
	Current smoking – ND	1.09 (0.79–1.50)	
	Current smoking DM	1.19 (0.93–1.54)	
	BMI <20 and >25 (kg/m <sup>2</sup> ) – ND	0.66 (0.49-0.89)	⊢∎→
	BMI <20 and >25 (kg/m <sup>2</sup> ) — DM	0.49 (0.36–0.67)	HEH
CKD	BP >140/90 mm Hg - ND	1.29 (0.96–1.72)	k <b>∎</b> _i
Progression	BP >140/90 mm Hg – DM	1.65 (1.34–2.04)	
	Not recommended Diet - ND	0.91 (0.68–1.21)	
	Not recommended Diet – DM	1.32 (1.08–1.62)	<b>⊢∎</b> →1
	Not Recommended Physical Activity – ND	1.08 (0.83–1.40)	⊢∎→
	Not Recommended Physical Activity – DM	1.13 (0.93–1.37)	H-₩-1
	HbA1c > 7.0% – ND	n/a	
	HbA1c > 7.0% – DM	1.07 (0.88–1.31)	⊢∎1
	Current smoking – ND	2.60 (1.57-4.29)	<b>₽</b>
	Current smoking DM	1.27 (0.89–1.82)	
	BMI <20 and >25 (kg/m <sup>2</sup> ) – ND	1.00 (0.57—1.76)	
	BMI <20 and >25 (kg/m <sup>2</sup> ) — DM	1.14 (0.68—1.91)	
Atherosclerotic	BP >140/90 mm Hg — ND	2.11 (1.33–3.35)	
events	BP >140/90 mm Hg – DM	1.17 (0.87–1.57)	
	Not recommended Diet – ND	1.52 (0.88–2.61)	F <b>■</b> 1
	Not recommended Diet – DM	1.02 (0.78–1.35)	
	Not Recommended Physical Activity - ND	1.22 (0.78–1.93)	
	Not Recommended Physical Activity – DM	1.41 (1.06–1.87)	<b>⊢_</b> ∎i
	HbA1c > 7.0% – ND	n/a	
	HbA1c > 7.0% – DM	1.65 (1.21–2.23)	
		0.0	

#### (a) Less than 65 years of age

**Figure 3.** (a,) Forest plot of hazard ratios (HRs) (95% confidence intervals [CIs]) of clinical outcomes by individual measures of health behaviors for participants without diabetes (ND) (white rows) and those with diabetes (DM) (gray rows) among participants (a) <65 years of age and (b)  $\geq 65$  years of age.

among older and younger adults with CKD. We explored for patterns of health behavior engagement, as we thought it imperative to recognize that participation in health behaviors cluster at the individual level. We found that clustering of certain health behaviors in less healthy behavior patterns was associated with increased risk of poor outcomes, but differentially, depending on diabetes status. Among younger adults, the less healthy behavior patterns were associated with increased risk of CKD progression in only those with diabetes and increased atherosclerotic events in those without diabetes. Among the older adults, less healthy behavior patterns were associated with increased risk of death and CKD progression in only those without diabetes. There were no statistically significant associations for older adults with concomitant diabetes. This finding suggests that the presence of

diabetes may largely explain the association in this age group. Alternatively, the lack of a significant finding could also be related to reduced power among this smaller sample of older adults.

In addition to exploring the health behavior pattern–outcome link, we investigated the association of individual health behaviors to further explore which health behaviors could potentially be driving the risk of poor outcomes. Previously, within the CKD population, several individual health behaviors, such as smoking, blood pressure, and physical activity, have been shown to be associated with outcomes, but these behaviors have not been specifically explored in different age groups.<sup>49,50</sup> In this study, the strongest associations between individual health behaviors and outcomes existed for current smoking and higher blood pressure. There were paradoxical findings with

Outcome	Individual behavior measures	Hazard ratio (95% CI)	Hazard ratio (95% CI)
	Current smoking - ND	1.88 (1.16-3.03)	· · · · ■ · · · · · · · · · · · · · · ·
	Current smoking DM	1.30 (0.73–2.32)	F ■ 1
	BMI <20 and >25 (kg/m <sup>2</sup> ) – ND	1.10 (0.78–1.72)	
	BMI <20 and >25 (kg/m <sup>2</sup> ) – DM	1.16 (0.78–1.72)	
Death	BP >140/90 mm Hg - ND	1.77 (1.2802.45)	⊢_∎
	BP >140/90 mm Hg – DM	1.77 (1.2802.45)	<b>⊢</b>
	Not recommended Diet - ND	0.93 (0.64-1.34)	
	Not recommended Diet – DM	0.97 (0.79–1.43)	
	Not Recommended Physical Activity - ND	1.22 (0.86-1.73)	
	Not Recommended Physical Activity – DM	1.06 (0.78–1.43)	
	HbA1c > 7.0% – ND	n/a	
	HbA1c > 7.0% – DM	1.37 (1.03–1.83)	<b>⊢</b> ∎−−-1
	Current smoking – ND	1.39 (0.71–2.73)	
	Current smoking DM	0.87 (0.44–1.76)	
	BMI <20 and >25 (kg/m <sup>2</sup> ) – ND	0.66 (0.40-1.11)	
	BMI <20 and >25 (kg/m <sup>2</sup> ) – DM	0.93 (0.46–1.90)	⊢ <b></b>
CKD	BP >140/90 mm Hg - ND	1.25 (0.79–1.99)	
Progression	BP >140/90 mm Hg - DM	2.41 (1.66–3.52)	· · · · · · · · · · · · · · · · · · ·
	Not recommended Diet – ND	0.65 (0.42-1.03)	<b>⊢∎</b>
	Not recommended Diet – DM	0.65 (0.44-0.95)	
	Not Recommended Physical Activity – ND	1.20 (0.78–1.84)	
	Not Recommended Physical Activity – DM	0.89 (0.62–1.27)	
	HbA1c > 7.0% — ND	n/a	
	HbA1c > 7.0% – DM	1.11 (0.79–1.56)	
	Current smoking – ND	1.08 (0.49–2.39)	⊢ <b>∎</b>
	Current smoking DM	0.64 (0.25–1.65)	
	BMI <20 and >25 (kg/m <sup>2</sup> ) - ND	1.07 (0.58–1.99)	
	BMI <20 and >25 (kg/m <sup>2</sup> ) – DM	0.66 (0.32–1.36)	
Atherosclerotic	BP >140/90 mm Hg — ND	1.85 (1.11–3.09)	<b>⊢∎</b>
events	BP >140/90 mm Hg – DM	0.79 (0.50–1.27)	
	Not recommended Diet – ND	0.97 (0.58–1.65)	
	Not recommended Diet – DM	0.94 (0.61-1.44)	<b>⊢_∎</b> I
	Not Recommended Physical Activity – ND	0.77 (0.48–1.23)	
	Not Recommended Physical Activity – DM	1.17 (0.77–1.77)	<b>⊢</b>
	HbA1c > 7.0% - ND	n/a	
	HbA1c > 7.0% – DM	1.34 (0.90–1.98)	
		0.0	05 10 15 20 25 30 35 4

#### (b) 65 years and older

Figure 3. Continued

the recommended BMI of  $<20 \text{ kg/m}^2$  or  $\geq 25 \text{ kg/m}^2$ , in that it was associated with a decreased risk of death among older adults with diabetes, which has not yet been reported,<sup>51</sup> and a decreased risk of CKD progression in younger adults; however, the published reports of BMI contributing to CKD progression have been conflicting.<sup>52,53</sup> Furthermore, a diet that did not fall in the recommended range was differentially associated with CKD progression between age groups with diabetes, as well as an increased risk among younger adults and a decreased risk in older adults. The available studies on the influence of diet have yet to demonstrate a consistent association with CKDrelated outcomes.<sup>54</sup>

Based on this study's results, and if future studies support the findings, health care providers caring for CKD patients could tailor their management recommendations. For example, ensuring that we prioritize

blood pressure control regardless of age, as it was a central component of the healthy behavior pattern in both age groups, and as an individual measure, it appears to be protective for death, CKD progression, and atherosclerotic events. In addition, a focus on increasing physical activity as an intervention target would be beneficial, since the health behavior patterns in both age groups with sedentary activity were associated with poorer outcomes across diabetes strata. In addition, even though the prevalence of smoking is quite low, smoking cessation is likely a prime intervention target for individuals <65 years of age in the health behavior pattern with less obesity and more smoking, as this group already participates in increased physical activity. At this time, weight management should likely not be a focus of CKD management recommendations, as it remains unclear which BMI is most appropriate in predialysis CKD.

Potential barriers identified in this study, such as less education, lower self-efficacy, and more depressive symptoms, have also been identified in other chronic disease populations, but had yet to be explored for differences between younger and older adults.<sup>34,38</sup> Age-specific barriers are important to consider, because certain clinical and psychosocial factors may affect older adults' ability to adopt health behaviors. In particular, older adults with CKD have higher rates of comorbid conditions, frailty, and decreased functional status, which could further challenge their ability to engage in the recommended health behaviors.<sup>29,55,56</sup> In this study, we demonstrate that a higher burden of comorbidities and medications, lower social support, inadequate health literacy, lower physical functioning, and lower efficacy in the patient-provider interaction are associated with less healthy behavior engagement patterns.

A logical first step to enhance the ability of individuals with CKD to participate in recommended health behaviors is to address the barriers to behavior engagement. The more readily modifiable barriers include physical functioning, self-efficacy, social support, health literacy, and depressive symptoms, which could serve as potential targets for intervention. Specifically addressing health literacy could be particularly beneficial as it has been associated with better outcomes in other chronic diseases,<sup>57</sup> can lead to greater self-efficacy, which in turn leads to improved self-management, which includes participation in recommended health behaviors.<sup>58</sup> In addition, future directions in CKD management could further explore the role of self-efficacy in the patient-provider relationship, as well as conducting randomized clinical trials of health behavior promotion programs, exploring the role of supporting health behaviors to address poor outcomes.

Unique to this study is the assessment of engagement in health behaviors, identification of potential barriers to health behavior engagement, and description of the association of health behaviors to important clinical outcomes separately for younger and older adults with CKD. Strengths of this study includes the wealth of data obtained from the CRIC Study, which represents the leading causes of CKD, a broad spectrum of ages, and diverse racial/ethnic groups. The data also allowed for the assessment of psychosocial, clinical and sociodemographic factors that have been minimally explored in relation to health behaviors and, more broadly, self-management in CKD.

Despite these strengths, there are limitations to discuss. This study used baseline measures and therefore could not assess whether the severity of kidney disease or other important covariates such as cognitive

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decline and depression mediate the association with clinical outcomes. It will be important to investigate how psychosocial characteristics and change in renal function influence health behavior engagement in a longitudinal manner in order to potentially identify the reasons for the excess mortality and the cognitive and functional decline unique to older adults with CKD. Furthermore, the generalization of our study results could be limited, as the CRIC Study population likely differs from the overall CKD population. The prevalence of adequate health literacy among older participants within the CRIC Study was somewhat higher (75.2%–83.2%) than national levels (71%).<sup>59</sup> However, the engagement in most of the health behaviors was similar to that in a national sample of older adults.<sup>60</sup>

In conclusion, we identified 3 similar patterns of engagement in health behaviors among younger and older adults with CKD that varied in association with clinical outcomes. In both age groups, individuals with less healthy behavior patterns had an increased hazard of poor clinical outcomes, but differed depending on diabetes status. Identification of the health behavior patterns and barriers to health behavior engagement may help target high-risk groups for strategies to increase participation in health behaviors.

# DISCLOSURE

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# SUPPLEMENTARY MATERIAL

Supplementary Data. Latent class analysis.

**Figure S1**. Analytical steps and associated sample sizes for each step.

Figure S2. Response probabilities (x-axis) of recommended health behaviors (y-axis), given behavior engagement pattern, (A) <65 years and (B) 65+ years. Pattern 1 (healthy pattern), pattern 2 (more obese/sedentary pattern), pattern 3 (less obese/smoker pattern).

**Table S1.** Model fit indices of successive latent-classmodels by age group.

**Table S2.** Multinomial logistic regression model results of the association of the 3 behavior engagement patterns with selected demographic, psychosocial, and clinical factors among age groups in the 5499 CRIC Study participants.

**Table S3.** Events, event rate (per 100 person-years) reported for the phase I CRIC population (N = 3939), stratified by diabetes and age (<65 years and  $\geq$ 65 years of age). **Table S4.** Associations of health behavior engagement patterns with clinical outcomes by <65 years and  $\geq$ 65 years of age in the phase I CRIC population (N = 3939), stratified by diabetes.

**Table S5.** Associations of individual health behavior measures among adults with CKD <65 years and  $\geq 65$  years, stratified by diabetes.

**Table S6.** Model fit indices of successive latent-class models by age group, in the sensitivity LCA with controlled BP  $\leq$ 130/80 mm Hg.

**Table S7.** Behaviors of latent class-defined health behaviorengagement patterns in the CRIC phase I and III cohortparticipants<65 years, in the sensitivity LCA with</td>controlled BP  $\leq$ 130/80 mm Hg.

**Table S8.** Health behaviors of latent class-defined health behavior engagement patterns in the CRIC phase I and III cohort participants  $\geq$ 65 years, in the sensitivity LCA with controlled BP  $\leq$ 130/80 mm Hg.

**Table S9.** Clinical characteristics of latent class-defined health behavior engagement patterns in the CRIC phase I and III cohorts by age group (<65 years and  $\geq$ 65 years), in the sensitivity LCA with controlled BP  $\leq$ 130/80 mm Hg.

Table S10. Associations of health behavior engagement patterns from the sensitivity LCA and clinical outcomes in the phase I CRIC population (N = 3939) <65 years, stratified by diabetes.

**Table S11.** Associations of health behavior engagement patterns from the sensitivity LCA and clinical outcomes in the phase I CRIC population (N = 3939)  $\geq$ 65 years, stratified by diabetes.

Supplementary material is linked to the online version of the paper at www.kireports.org.

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