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Heart Failure Incidence and Mortality in the Southern Community Cohort Study

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Abstract

Background—There is a paucity of data regarding HF incidence among low-income and minority populations. Our objective was to investigate HF incidence and post-HF survival by race and sex among low-income adults in the southeastern US.

Methods and Results—Participants were 27,078 white and black men and women enrolled during 2002–2009 in the Southern Community Cohort Study (SCCS) who had no history of HF and were receiving Centers for Medicare or Medicaid services (CMS). Incident HF diagnoses through December 31, 2010 were ascertained using ICD-9 codes 428.x via linkage with CMS research files. Most participants were black (68.8%), women (62.6%) and earned < \$15,000/year (69.7%); mean age was 55.5 (10.4) years. Risk factors for HF were common: hypertension (62.5%), diabetes (26.5%), myocardial infarction (8.6%) and obesity (44.8%). Over a median follow-up of 5.2 years, 4,341 participants were diagnosed with HF. The age-standardized incidence rates were 34.8, 37.3, 34.9 and 35.6 PY/1000 in white women, white men, black men and black women, respectively, remarkably higher than previously reported. Among HF cases, 952 deaths occurred over a median follow-up 2.3 years. Men had lower survival; hazard ratios and 95% confidence intervals were 1.63 (1.27–2.08), 1.38 (1.11–1.72) and 0.90 (0.73–1.12) for white men, black men and black men and black women compared with white women.

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Conclusions—In this low-income population, HF incidence was higher for all race-sex groups than previously reported in other cohorts. The SCCS is a unique resource to investigate determinants of HF risk in a segment of the population underrepresented in other existing cohorts.

Keywords

heart failure; incidence; survival; race; epidemiology

Journal Subject Terms

Heart Failure; Epidemiology; Race and Ethnicity

There are over 26 million persons living with heart failure (HF) worldwide.¹ In the US, over 5.7 million adults ($\approx 2.5\%$ of the US adult population) are estimated to have HF.² About half of persons diagnosed with HF die within 5 years and the estimated total costs of HF in the US exceeded \$30 billion in 2012.^{3,4} Several established cardiovascular disease (CVD) cohorts have investigated HF incidence and mortality, including the Framingham Heart Study (FHS), Cardiovascular Health Study (CHS), Multi-ethnic Study of Atherosclerosis (MESA) and Atherosclerosis Risk in Communities (ARIC).^{5–8} The FHS included predominantly white individuals. Other cohorts, including CHS, MESA, and ARIC, enrolled multi-ethnic middle-class populations from select communities and their relatively small sample sizes limited assessment of differential risk patterns between demographic groups defined by both race and sex.

While data from these previous cohorts suggest differences in HF incidence rates and post-HF survival between population subgroups, knowledge gaps persist regarding the magnitude and direction of these differences, particularly in multi-ethnic low income populations with high burden of CVD risk factors. The prospective Southern Community Cohort Study (SCCS) provided a valuable opportunity to investigate differences in the incidence of HF as well as in post-HF survival between groups defined by race and sex: white women, black women, white men and black men.⁹

Methods

Study sample

The SCCS is a prospective cohort study designed to investigate the incidence of cancer and other chronic diseases, including differential patterns by race and sex, in a low-income under-insured population underrepresented in previous studies. Between 2002 and 2009, the SCCS enrolled approximately 86,000 adults (\approx two-thirds black) aged 40–79 living in 12 southeastern states to investigate various chronic disease outcomes.⁹ Approximately 86% of participants were recruited at community health centers (CHC), which provide primary health and preventive care services for low-income populations so that the cohort is made up of a segment of society (minority, poor, rural) seldom included in sizeable numbers in previous cohort studies; particularly those investigating CVD.^{9, 10} The remaining 14% were recruited via mail-based general population sampling. Data on socioeconomic, demographic (including self-reported race), lifestyle, and anthropometric characteristics, as well as

personal medical history, were ascertained at cohort enrollment via standardized computerassisted personal interviews for CHC participants, and via self-administered mailed questionnaire for general population participants. Detailed description of SCCS methods has been previously published.^{9, 10}

SCCS participants (n = 27,078) included in the current analyses were individuals aged 65 years (n = 7001) at cohort enrollment, or persons < 65 years (n = 20,077) at enrollment who: a) reported being covered by Medicaid (which provides medical benefits to low-income adults and uninsured persons) on the baseline questionnaire; or b) reported being covered by Medicare (the primary health insurance program for persons aged 65) on the baseline questionnaire; or c) did not report Medicare or Medicaid on the baseline questionnaire but had a Centers for Medicare and Medicaid Services (CMS) claim within 90 days of being enrolled in SCCS. The restriction to these groups ensures that participants would likely have continuous coverage in Medicare and/or Medicaid from the time of SCCS enrollment to the end of the follow-up period (December 31st, 2010), for the ascertainment of incident HF events. Analyses were restricted to self-reported African American or black and non-Hispanic white SCCS participants, since too few persons in other racial groups were available for stable statistical analysis.

Outcome ascertainment

Heart failure events were ascertained via linkage of the SCCS cohort with CMS Research Identifiable Files (which include Medicare institutional and non-institutional files, and the Medicaid Analytic Extract files). Incident HF was defined as the first occurrence of a medical claim with an International Classification of Diseases, 9th revision, discharge code of 428.x (428.0 to 428.9) within the Medicare institutional (Medicare Provider Analysis and Review, MEDPAR, which includes inpatient, outpatient and skilled nursing facility base files), Part B carrier (includes non-institutional physician services and durable medical equipment), or outpatient-based claims files or the Medicaid Analytic Extract (MAX) Inpatient and Other Services claims files, from the date of SCCS enrollment through December 31st, 2010. Detailed description of the CMS research files are published elsewhere. ¹¹

Deaths, including dates and causes of death, were ascertained via linkage of the SCCS cohort with both the Social Security Administration (SSA) vital status service for epidemiologic researchers and the National Death Index (NDI) through December 31st, 2010. Both NDI and SSA are well-established and reliable means of identifying deaths in the US, and are expected to capture nearly all deaths.^{12–14}

Statistical Analysis

Descriptive statistics (means and standard deviations for continuous variables and counts and percentages for categorical variables) were computed for all study participants by race and sex.

To investigate the incidence of HF, duration of follow-up was computed from date of entry into the SCCS until the date of the first diagnosis of HF, date of death, or December 31st, 2010, whichever occurred first. Incidence rates (IR) of heart failure were calculated for

white women, black women, white men and black men by dividing the number of HF cases by person-time of follow-up, presented per 1,000 person-years. The 95% confidence intervals (CI) were calculated using the quadratic approximation to the Poisson log likelihood for the log-rate parameter.¹⁵ To account for age differences between the demographic categories, age-standardized rates were computed using the overall age distribution of the SCCS participants.

Multivariable Cox models were utilized to test whether differences in crude IRs between categories defined by race and sex persisted after adjustment for baseline covariates. Three models were constructed, with white women as the referent category: model 1 included indicator variables for white men, black women and black men and age (restricted cubic splines with 4 knots); model 2 additionally adjusted for body mass index (restricted cubic splines with 4 knots), and history of diabetes, hypertension, high cholesterol, MI/CABG or stroke (all yes/no); model 3 additionally adjusted for the following covariates: annual household income (<\$15,000; \$15,000–\$24,999; \$25,000); education (< high school, high school/vocational training/junior college, college degree or higher), smoking (never, former, current <19.5 pack-years, current 19.5 pack-years, 19.5 being the median pack-years among current smokers), alcohol intake (linear and quadratic term), marital status (married/ living as married with partner, separated/divorced, widowed, single/never married) and enrollment source (community health centers vs general population). Knots were placed at quantiles of covariate distributions, equally spaced in sample size.¹⁶

For analyses of post-HF survival among those with a diagnosis of incident HF, follow-up time was defined as time from HF diagnosis to death or December 31^{st} 2010 whichever occurred first. When date of death was coincident with date of HF diagnosis, follow-up time was set to 0.5 days. We computed cumulative mortality for both HF cases and non-cases using contingency tables. Age-adjusted estimates of the survivor functions (adjusted to the mean age of SCCS participants diagnosed with HF) were obtained from a stratified Cox model fit and plotted for all race-sex groups. Cox models were used to investigate differences in cumulative hazard for death (all-cause mortality) using white women as the referent group. Model 1 comprised indicator variables for white men, black women and black men and age (restricted cubic splines with 4 knots). Variables included in models 2 and 3 are the same as described previously. P-values for race-by-sex interaction were computed in models for HF incidence and post-HF survival; and a p-value < 0.05 was considered statistically significant. Model assumptions were verified using Schoenfeld residuals and log (–log) plots.

All analyses were performed using STATA (version 12.1, Stata Corp, College Station, Texas, USA) and the 'rms' package for R version 3.1.1 (R Core Team 2014).^{16, 17}

Ethics statement

SCCS participants provided written informed consent, and protocols were approved by the Institutional Review Boards of Vanderbilt University Medical Center and Meharry Medical College.

Results

Among the 27,078 SCCS participants included in this study, 68.8% were black, 62.6% were women, 69.7% had annual household income < 15,000 and 38.4% had less than a high school education. The mean (SD) age at enrollment was 55.5 (10.4) years. At baseline, risk factors for HF were common: hypertension (62.5%); diabetes (26.5%); myocardial infarction (8.6%); and obesity, BMI 30 kg/m² (44.8%) (Table 1).

Overall, white men were older and had the highest prevalence of MI and stroke at baseline (Table 1). In contrast, black women were more likely to be obese at baseline and report a history of diabetes and hypertension.

HF incidence

Over a median (25th, 75th percentile) follow-up time of 5.2 (3.1, 6.7) years, 4,341 participants (16%) developed incident HF (IR: 32.8/1000 person-years; 95% CI: 31.8–33.8). White men had the highest age-standardized IR, 37.3/1000 PY, compared with 34.8, 34.9 and 35.6 in white women, black men and black women, respectively (Table 2).

In models adjusted for age and other risk factors for HF, black women had a significantly lower risk of HF when compared with white women [HR=0.89; 95% CI: 0.82–0.98]. The risk of HF was similar among white men (HR=1.09; 95% CI: 0.97–1.23) and black men (HR=1.04; 95% CI: 0.94–1.15) compared with white women (Table 2). There was no evidence of race-by-sex interaction [p = 0.22].

Post-HF survival

Among the 4,341 individuals who developed incident HF, 952 died (cumulative mortality = 21.9%) over a median (25^{th} , 75^{th} percentile) post-HF follow-up time of 2.3 (0.9, 4.2) years (Table 3). Men had higher percent mortality than women (29% vs. 18%), with little difference by race. In persons without HF (n= 22,737), there were 1,929 deaths, corresponding to a percent mortality of 8.5%.

Figure 1 shows age-adjusted survival curves for persons diagnosed with HF stratified by race and sex. The 5-year post-HF survival probability was significantly lower among white men (0.55; 95% CI: 0.49–0.61) and black men (0.64; 95% CI: 0.60–0.67) compared with white women (0.73; 95% CI: 0.69–0.78) and black women (0.77; 95% CI: 0.74–0.79), respectively [p < 0.0001]. Racial differences within sex groups were not statistically significant. Similar patterns were observed for 1-year and 3-year survival probabilities.

Compared with white women, the risk of death was 60% (95% CI: 27%–202%) higher in white men and 35% (95% CI: 9%–65%) higher in black men in analyses adjusted for age, BMI, hypertension, diabetes, high cholesterol, past history of MI/CABG and stroke (Table 3). These findings were robust to further adjustment for lifestyle factors and enrollment source. In contrast, comparisons between black women and white women suggested minimal non-significant relative differences in risk by race in all models. The race-sex interaction term was not statistically significant [p = 0.92].

Discussion

We investigated heart failure incidence and post-HF mortality in a large multi-ethnic lowincome sample from the southeastern United States. Our principal findings are: 1) the incidence rate for HF was remarkably high across all race and sex groups in the SCCS, 2) there was no significant difference in age-adjusted incidence rates across groups defined by race and sex but after full adjustment for socioeconomic status and traditional cardiovascular risk factors black women had the lowest risk of HF, and 3) higher post-HF mortality among men with no significant racial differences.

The incidence rates for heart failure in the SCCS exceeded those previously reported from established CVD cohorts (Table 4). In CHS (n = 5,888; age range: 65 years) for example, the age-standardized HF incidence rates after 10 years of follow-up among white men, black men, white women and black women were 30.2, 19.2, 27.5 and 22.6 per 1000 PY respectively.⁵ In ARIC (n = 14,933; age range: 45–64), Loehr et al found IRs of 6.0, 9.1, 3.4 and 3.8 per 1000 PY, respectively, in these race-sex groups.⁸ The FHS and MESA reported even lower IRs (7.2 and 4.2 per 1000 PY in men and women in FHS and 3.1 per 1000 PY overall in MESA).^{6, 7} The higher SCCS HF incidence could be explained in part by notably higher prevalence of CVD risk factors (in particular hypertension, diabetes, obesity, prior history of MI) in the SCCS study sample at baseline compared with ARIC, CHS, FHS and MESA (Table 5). In addition, SCCS participants were largely of low socioeconomic status, with over two-thirds having annual household income <\$15,000. Prior evidence suggests a strong independent association between socioeconomic status and HF risk; ^{18–21} thus participants in SCCS may be at higher risk of unfavorable societal stressors and an elevated risk of adverse cardiovascular outcomes including HF.

White men had the highest crude incidence rate of HF in the SCCS, consistent with findings from the CHS. However, minimal differences in age-adjusted incidence rates and HF risk between groups after adjustment for CVD risk factors (except for black women who had significantly lower risk) suggest homogeneity of HF risk profile. Similarly, in ARIC, crude racial and sex differences in incidence density were attenuated by adjustment for CVD risk factors.⁸

Overall, the 5-year post-HF survival in SCCS was higher than the 52% previously reported (data from the Olmsted county study).³ This may be due in part to the fact that SCCS participants had shorter post-HF follow-up time, were younger at baseline (55.5 vs 74 years) and temporal trends suggesting improved post-HF survival ³ related to recent improvements in therapeutic options. In addition, participants in the Olmsted county study were mostly non-Hispanic Whites who may be at higher risk of HF with reduced ejection fraction (HF*r*EF), which has a less favorable prognosis compared with heart failure with preserved ejection fraction (HF*r*EF).^{22, 23}

The relative patterns of post-HF survival for the four demographic subgroups in SCCS were substantially different from those seen in ARIC and CHS. In ARIC, compared with white men and women, black men and women had the lowest survival probability following admission for HF. The 5-year case fatality for white women, white men, black women and

black men were 35.8%, 41.2%, 46.1% and 51.8%, respectively. The racial differences were significant, with black men having the highest all-cause mortality following admission, but the differences by sex were non-significant. In CHS, the mortality rate in white women, white men, black women and black men were 35.5, 40.5, 33.6 and 44.4 per 100 PY respectively. After full adjustment for covariates there were no significant racial differences, but women had a 15% lower risk of all-cause mortality [HR: 0.85; 95% CI: 0.73, 0.99].¹³ In SCCS, white men had the lowest 5-year survival post-HF diagnosis; but after full adjustment, there were mainly sex-differences. This could be explained in part by the higher prevalence of MI among men. MI is associated with greater risk for the development of HF/EF which is known to have a worse prognosis compared with HF/pEF.^{22, 23} However, MI does not fully account for the higher risk of post HF mortality among men, as this risk persisted even after full adjustment for relevant baseline covariates (including history of MI).

Limitations of our study should be noted. Our study sample may not be representative of the background population of the Southeastern states as the recruitment and sampling scheme utilized by the SCCS was tailored towards low-income, rural and under-insured populations not often included in sizeable numbers in other cohorts investigating chronic disease outcomes. Also, HF was ascertained via linkage with CMS Research Identifiable Files using ICD-9 codes 428.x, rather than independent physician adjudication. However, the diagnosis codes (ICD-9 428.x) algorithm for identification of HF used in this study has been previously validated and utilized in other cohorts.²⁴⁻²⁶ A review of the detection of HF in administrative claims data that included eight studies conducted among Medicare beneficiaries reported positive predictive values (PPVs) between 76% and 99%, with the majority of the studies reporting PPVs over 90%.²⁴ These codes have also been used with high specificity in a number of studies ^{25, 26} even though no independent validation was conducted by the SCCS investigators. An over-representation of groups with elevated HF risk (persons > 65 and persons < 65 receiving Medicare) in our SCCS sub-cohort compared with the SCCS base population, may have contributed to higher HF incidence rates than would be expected for the total SCCS cohort. However with the mean age of the total cohort being ≈ 52.6 years²⁷ versus 55.5 years for our sub-cohort, the small age difference between both populations may have had less than dramatic effects on the HF incidence. In addition, with studies suggesting that the sensitivity of ICD-9 code 428.x for HF ascertainment varies between 62.8 and 89%^{24, 28}, it is plausible that we may have underestimated the incidence rate of HF in our sub-cohort. Also, when contrasting the incidence rates between our study and previous CVD cohorts (like ARIC and CHS) we used data for comparable age groups between studies (Table 4). However, the fact that HF represents a myriad of clinical conditions, the lack of universality in the definition of HF and the heterogeneity in the methods for HF ascertainment between studies makes head-to-head comparisons between studies difficult. Our analyses required assumptions regarding the continuous coverage in CMS of persons less than 65 years, raising the possibility of incomplete capturing of HF events in this age stratum of the SCCS cohort. However, we found that over 81.9% of persons aged < 65 who reported CMS coverage at baseline had a claim for any condition within 90 days of being enrolled in SCCS. This suggests that an even greater proportion of participants included in this study filed at least one claim at some point during follow-up

from 2002 to 2010 and thus any HF event would likely have been captured if it occurred. Data on baseline covariates (including anthropometric and cardiovascular risk factors) were based on self-report of a physician diagnosis and use of medications (diabetes and hypertension). While self-report may be susceptible to recall and misclassification bias, these methods have been successfully used and validated in large epidemiologic cohorts, including the SCCS. Many of the questions on the SCCS questionnaire were adapted from questionnaires used and validated in other settings; and a series of independent validation studies using biomarkers, repeat interviews or medical records have demonstrated the reliability of the questionnaire within the SCCS population for variables such as smoking status, self-reported diseases including diabetes, height and weight.⁹

The SCCS cohort is comprised of a substantial number of individuals from minority and low-income populations who are traditionally under-represented in most studies investigating CVD and heart failure in particular. The incidence rates for HF in the SCCS exceeded that of most existing cardiovascular cohorts. Therefore, the SCCS provides an unparalleled opportunity to investigate patterns in HF incidence and mortality among the highest risk individuals. In addition, both black and white participants included in this cohort had minor differences in income and education levels thereby curtailing confounding by socioeconomic differences. The availability of a large sample of participants and HF cases provided the opportunity to adequately explore differential patterns across sex and racial categories. Also, linkage with the NDI and SSA allowed for robust ascertainment of all-cause mortality.

In conclusion, in this low-income multiethnic population, we found higher incidence rates for HF in all race-sex groups than previously reported in other CVD cohorts which was paralleled by high prevalence of CVD risk factors at baseline. This suggests that SCCS can be a unique resource to investigate determinants of HF risk in a segment of the population underrepresented in other existing cohorts.

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Clinical Perspective

Heart failure (HF) is a major public health problem. Most of the epidemiologic data regarding the incidence, prevalence, and survival in HF is derived from cardiovascular disease (CVD) cohorts that enrolled mostly middle-class individuals and with relatively low representation of black individuals. However, the greatest burden of HF risk factors resides within individuals of lower socioeconomic status, particularly within the southeastern United States. Therefore, we investigated HF incidence and post-HF survival by race and sex among 27,078 low-income individuals enrolled in the Southern Community Cohort Study (SCCS). Over a median follow-up of 5.2 years, 4,341 participants were diagnosed with HF. The age-standardized incidence rates were 34.8, 37.3, 34.9 and 35.6 PY/1000 in white women, white men, black men and black women, respectively, which are approximately two to ten times higher than previously reported from cohorts such as the Cardiovascular Health Study (CHS), Atherosclerosis Risk in Communities (ARIC), and the Multi-Ethnic Study of Atherosclerosis (MESA). Among incident HF cases in the SCCS, 952 deaths occurred over a median follow-up of 2.3 years. Compared with women, men had lower post-HF survival; hazard ratios and 95% confidence intervals were 1.63 (1.27–2.08), 1.38 (1.11–1.72) and 0.90 (0.73–1.12) for white men, black men and black women compared with white women. Individuals of lower socioeconomic status in the southeastern United States are at particularly high risk for the development of HF. In order to curb the HF epidemic, population and individual level resources should be directed toward this high risk segment of the population.

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Figure 1.

Age-adjusted Survival curves for SCCS participants diagnosed with HF stratified by race and sex. The adjusted survival estimates were computed at the mean age (58 years) of all participants with HF (n = 4341). Compared with white women and black women, white men and black men had significantly lower survival (p<0.0001). The 1-year, 3-year and 5-year age-adjusted survival estimates were (0.91, 0.83, 0.77); (0.90, 0.82, 0.74); (0.84, 0.71, 0.64) and (0.85, 0.73, 0.58) for black women, white women, black men and white men respectively.

Table 1

Comparison^{*} of baseline characteristics of 27, 078 SCCS participants who were receiving Medicare or Medicaid during follow-up between 2002 and 2010, according to race and sex.

	Overall $N = 27,078$	White Women n = 5,252	White Men $n = 3,202$	Black Women n = 11,688	Black Men n = 6,936
Age (SD), years	55.5 (10.4)	57.7 (10.6)	58.7 (10.5)	54.4 (10.4)	54.3 (9.5)
Age Categories %					
40–54	51.2	41.9	38.0	56.1	56.1
55-64	22.9	23.6	22.7	21.5	24.9
65	25.9	34.5	39.3	22.4	18.9
BMI (kg/m ²) (SD)	30.4 (7.8)	30.7 (8.2)	28.7 (6.4)	32.4 (8.2)	27.7 (6.1)
BMI Categories %					
< 18.5	1.7	2.2	1.1	1.4	2.0
18.5 - < 25.0	24.0	23.6	28.2	16.5	35.0
25 - < 30.0	29.5	28.0	36.4	25.4	34.3
30.0	44.8	46.2	34.3	56.7	28.8
History of MI %	8.6	8.6	17.7	6.1	8.6
History of Stroke %	9.6	10.0	10.8	9.1	9.4
Diabetes %	26.5	24.3	23.8	29.8	23.8
Hypertension %	62.5	57.0	56.7	67.6	60.5
High Cholesterol %	39.5	49.5	47.1	38.0	31.0
Education %					
< High school (HS)	38.4	32.0	29.0	40.4	44.2
HS/Junior college/VT	53.1	58.7	53.1	52.1	49.0
College degree	8.5	9.4	17.9	6.5	6.8
Annual Income < \$15,000, %	69.7	65.9	53.4	74.5	71.8
Smoking %					
Never	34.7	37.0	21.7	45.1	21.3
Former	25.3	26.8	40.4	20.5	25.3
Current	40.1	36.2	37.9	34.5	53.4

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	Overall N = 27,078	White Women n = 5,252	White Men n = 3,202	Black Women n = 11,688	Black Men n = 6,936
Alcohol Intake %					
0 drink per day	54.9	66.8	48.8	61.0	38.5
>0-2 drinks per day	33.2	29.7	35.6	31.4	37.6
>2 drinks per day	11.9	3.5	15.6	7.5	23.9

 $^*_{\rm Ml}$ comparisons between demographic groups were significant (p=0.02 for stroke; p<0.0001 for all other baseline variables).

VT: Vocational Training; MI: Myocardial Infarction; SD: Standard Deviation

Table 2

Risk of incident heart failure among participants in the Southern Community Cohort Study, overall and stratified by race and sex.

Incident HF cases (n) 4,341 801 511 1,940 Person-Years (PY) 132,500 23,339 13,934 60,639 Person-Years (PY) 132,500 23,339 13,934 60,639 Cumulative Incidence (%) 16.0 15.3 16.0 16.6 Incidence Rate/1000PY (95% CI) 16.0 15.3 36.7 (33.6,40.0) 32.0 (30.6, 33.4) Incidence Rate/1000PY (95% CI) 32.8 (31.8, 33.8) 34.3 (32.0, 36.8) 36.7 (33.6, 40.0) 32.0 (30.6, 33.4) Incidence Rate/1000PY (95% CI) 35.8 (31.8, 33.8) 34.3 (32.4, 37.2) 37.3 (34.0, 40.6) 32.0 (30.6, 33.4) Age-adjusted 35.1 (34.1, 36.2) 34.8 (32.4, 37.2) 37.3 (34.0, 40.6) 35.6 (33.9, 37.2) Age-adjusted 35.1 (34.1, 36.2) 34.8 (32.4, 37.2) 37.3 (34.0, 40.6) 35.6 (33.9, 37.2) Age-adjusted 35.1 (34.1, 36.2) 34.8 (32.4, 37.2) 37.3 (34.0, 40.6) 35.6 (33.9, 37.2) Age-adjusted 35.1 (34.1, 36.2) 37.3 (34.0, 40.6) 35.6 (33.9, 37.2) 10.0 (0.94.1.11) Model 1 Model 2 Model 2 10		Overall $N = 27,078$	White Women n = 5,252	White Men n = 3,202	Black Women n = 11,688	Black Men n = 6,936
Person-Years (PY) 132,500 23,339 13,934 60,639 Cumulative Incidence (%) 16.0 15.3 16.0 16.6 Incidence (%) 16.0 15.3 16.0 16.6 Incidence (%) 16.0 15.3 16.0 16.6 Incidence Rate/1000PY (95% CI) 32.8 (31.8, 33.8) 34.3 (32.0, 36.8) 36.7 (33.6, 40.0) 32.0 (30.6, 33.4) Actual 32.8 (31.8, 33.8) 34.3 (32.0, 36.8) 36.7 (33.6, 40.0) 32.0 (30.6, 33.4) Actual 35.1 (34.1, 36.2) 34.8 (32.4, 37.2) 37.3 (34.0, 40.6) 32.6 (33.9, 37.2) Actual Ratio (95% CI) 35.1 (34.1, 36.2) 34.8 (32.4, 37.2) 37.3 (34.0, 40.6) 35.6 (33.9, 37.2) Actual Ratio (95% CI) 35.1 (34.1, 36.2) 34.8 (32.4, 37.2) 37.3 (34.0, 40.6) 10.0 (0.91, 11.1) Model 1 1.00 1.00 1.00 (0.91, 11.4) 0.91 (0.83, 0.99) 10.0 Model 2 100 1.00 1.00 (0.97, 1.23) 0.89 (0.82, 0.98) 10.0	Incident HF cases (n)	4,341	108	511	1,940	1,089
Cumulative Incidence (%) 16.0 15.3 16.0 16.6 Incidence Rate/1000PY (95% CI) 15.3 16.0 16.6 16.6 Incidence Rate/1000PY (95% CI) 32.8 (31.8, 33.8) 34.3 (32.0, 36.8) 36.7 (33.6, 40.0) 32.0 (30.6, 33.4) Crude 32.8 (31.8, 33.8) 34.3 (32.0, 36.8) 36.7 (33.6, 40.0) 32.0 (30.6, 33.4) Age-adjusted 35.1 (34.1, 36.2) 34.8 (32.4, 37.2) 37.3 (34.0, 40.6) 35.6 (33.9, 37.2) Age-adjusted 35.1 (34.1, 36.2) 34.8 (32.4, 37.2) 37.3 (34.0, 40.6) 35.6 (33.9, 37.2) Age-adjusted 35.1 (34.1, 36.2) 34.8 (32.4, 37.2) 37.3 (34.0, 40.6) 10.2 (0.94, 1.11) Model 1 1.00 1.00 1.02 (0.91, 1.14) 0.91 (0.83, 0.99) 100 Model 3 1.00 1.00 1.00 (0.97, 1.23) 0.89 (0.82, 0.98) 100	Person-Years (PY)	132,500	23,339	13,934	60,639	34,589
Incidence Rate/1000PY (95% CI) 32.8 (31.8, 33.8) 34.3 (32.0, 36.8) 36.7 (33.6, 40.0) 32.0 (30.6, 33.4) Crude 32.8 (31.8, 33.8) 34.3 (32.0, 36.8) 36.7 (33.6, 40.0) 32.0 (30.6, 33.4) Age-adjusted 35.1 (34.1, 36.2) 34.8 (32.4, 37.2) 37.3 (34.0, 40.6) 35.6 (33.9, 37.2) Hazard Ratio (95% CI) 35.1 (34.1, 36.2) 34.8 (32.4, 37.2) 37.3 (34.0, 40.6) 10.0 (0.91, 1.11) Model 1 1.00 1.00 1.02 (0.91, 1.14) 0.91 (0.83, 0.99) Model 2 1.00 1.00 (0.97, 1.23) 0.89 (0.82, 0.98) 089 (0.82, 0.98)	Cumulative Incidence (%)	16.0	15.3	16.0	16.6	15.7
Crude32.8 (31.8, 33.8)34.3 (32.0, 36.8)36.7 (33.6, 40.0)32.0 (30.6, 33.4)Age-adjusted35.1 (34.1, 36.2)34.8 (32.4, 37.2)37.3 (34.0, 40.6)35.6 (33.9, 37.2)Hazard Ratio (95% CI)35.1 (34.1, 36.2)34.8 (32.4, 37.2)37.3 (34.0, 40.6)1.02 (0.94, 1.11)Model 11.001.001.04 (0.93, 1.16)1.02 (0.94, 1.11)Model 201.001.00 (0.97, 1.13)0.91 (0.83, 0.99)Model 31.001.001.00 (0.97, 1.23)0.89 (0.82, 0.98)	Incidence Rate/1000PY (95)	% CI)				
Age-adjusted 35.1 (34.1, 36.2) 34.8 (32.4, 37.2) 37.3 (34.0, 40.6) 35.6 (33.9, 37.2) Hazard Ratio (95% CI)	Crude	32.8 (31.8, 33.8)	34.3 (32.0, 36.8)	36.7 (33.6, 40.0)	32.0 (30.6, 33.4)	31.5 (29.7, 33.4)
Hazard Ratio (95% CI) Model 1 1.00 1.04 (0.93, 1.16) 1.02 (0.94, 1.11) Model 2 1.00 1.00 1.02 (0.91, 1.14) 0.91 (0.83, 0.99) Model 3 1.00 1.00 1.09 (0.97, 1.23) 0.89 (0.82, 0.98)	Age-adjusted	35.1 (34.1, 36.2)	34.8 (32.4, 37.2)	37.3 (34.0, 40.6)	35.6 (33.9, 37.2)	34.9 (32.7, 37.1)
Model 1 1.00 1.04 (0.93, 1.16) 1.02 (0.94, 1.11) Model 2 1.00 1.02 (0.91, 1.14) 0.91 (0.83, 0.99) Model 3 1.00 1.09 (0.97, 1.23) 0.89 (0.82, 0.98)	Hazard Ratio (95% CI)					
Model 2 1.00 1.02 (0.91, 1.14) 0.91 (0.83, 0.99) Model 3 1.00 1.09 (0.97, 1.23) 0.89 (0.82, 0.98)	Model 1		1.00	1.04 (0.93, 1.16)	$1.02\ (0.94,1.11)$	$0.99\ (0.91,\ 1.09)$
Model 3 1.09 (0.97, 1.23) 0.89 (0.82, 0.98)	Model 2		1.00	1.02 (0.91, 1.14)	$0.91\ (0.83, 0.99)$	1.06 (0.97, 1.17)
	Model 3		1.00	1.09 (0.97, 1.23)	$0.89\ (0.82,0.98)$	1.04 (0.94, 1.15)

Model 1: Includes age (restricted cubic splines with 4 knots), race and sex. Model 2: Model 1 + BMI (restricted cubic splines with 4 knots), history of diabetes, hypertension, high cholesterol, MI and stroke (all yes/no). Model 3: Model 2 + annual household income (<\$15000, \$1500-\$24999 & \$25000), education (< high school, high school, view school, view school, sc smoking (never, former, current < 19.5 pack-years, current 19.5 pack-years) and alcohol intake (linear and quadratic term), marital status (married/living as married with partner, separated/divorced, widowed, single/never married) and enrollment source (community health centers vs general population). P-value for race×sex interaction = 0.22.

CI: Confidence Interval

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	Overall N = 4,341	White Women n = 801	White Men n = 511	Black Women n = 1,940	Black Men n = 1,089
Deaths (n)	952	144	152	343	313
Percent Mortality (%)	21.9	18.0	29.7	17.7	28.7
Risk of death: Hazard r	atio (95% CI)				
Model 1		1.00 (ref)	1.73 (1.37, 2.17)	0.91 (0.75, 1.10)	1.61 (1.32, 1.96)
Model 2		1.00 (ref)	1.60 (1.27, 2.02)	$0.89\ (0.73,\ 1.09)$	1.35 (1.09, 1.65)
Model 3		1.00 (ref)	1.63 (1.27, 2.08)	0.90 (0.73, 1.12)	1.38 (1.11, 1.72)

Model 1: Includes age (restricted cubic splines with 4 knots); race and sex. Model 2: Model 1 + BMI (restricted cubic splines with 4 knots), history of diabetes, hypertension, high cholesterol, MI and stroke (all yes/no). Model 3: Model 2 + annual household income (<\$15000, \$15000-\$24999 & \$25000); education (< high school, high school/vocational training/junior college, college degree or higher), smoking (never, former, current < 19.5 pack-years) and alcohol intake (linear and quadratic term), marital status (married/living as married with partner, separated/divorced, widowed, single/never married) and enrollment source (community health centers vs general population). P-value for race×sex interaction = 0.92.

CI: Confidence Interval

Table 4

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			HF Incidenc	e Rates	(IR) per	1000 PY		
	SCCS [*] 45–64 yrs	ARIC ⁸ 45–64 yrs	SCCS [*] 65–79 yrs	99 CH	IS ⁵ † 5 yrs		SCCS [*] 40–79 yrs	MESA ⁶ 45–84 yrs
	n = 15,321	n = 14,933	n = 7,001	= u	5,888		n = 27,078	n = 6,814
				5-yr	10-yr			
Overall	35.8	5.7	39.5	19.3	24.0	Overall	33.7	3.4
White Women	38.3	3.4	34.6	14.5	19.2			
White Men	39.7	0.0	38.3	24.9	30.2	Whites	34.3	2.4
Black Women	35.1	8.1	42.4	19.6	22.6			
Black Men	34.0	9.1	41.9	23.5	27.5	Blacks	33.4	4.6
*	c.							

Incidence rates computed for SCCS participants aged 45–64, 65–79 and 40–79 for comparability with the ARIC, CHS and MESA cohorts respectively. In addition, the rates are standardized to the age distribution of the SCCS study participants within these age ranges.

 $\dot{\tau}^{\prime}$ The tabulated values are computed from values presented in Arnold et al⁵ and standardized to the age distribution of CHS participants.

ARIC: Atherosclerosis Risk in Communities; CHS: Cardiovascular Health Study; MESA: Multi-ethnic Study of Atherosclerosis; SCCS: Southern Community Cohort Study.

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Table 5

ARIC and CHS participants	
MESA,	
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characteristics	
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Comparison (

Age (years)* $55.5 (10.4)$ $54 (6)$ Age (years)* 62.6 $54 (6)$ Women (%) 62.6 54 Blacks (%) 68.8 27 Education (<high (%)<="" school)="" td="">$38.4$$24$Bdm (kg/m²)*$30.4 (7.8)$$24$BMI (kg/m²)*$30.4 (7.8)$$27.3 (5.1)$Obese (BMI > 30) %$44.8$$26$Diabetes (%)$26.5$$11$Diabetes (%)$62.5$$33$Mypertension (%)$8.6$$4$Stroke (%)$9.6$$1.4$</high>	SCCS (n= $27,078$) ARIC ⁸ (n= $14,99$	$(3) CHS^5 (n=5,888)$	MESA ⁶ $(n = 5,923)^{\dagger}$	${ m FHS^7}$ (n=9,405) [‡]
Women (%) 62.6 54 Blacks (%) 68.8 27 Blacks (%) 68.8 27 Education (< high school) (%)	55.5 (10.4) 54 (6)	72.8 (5.6)	61.8 (10.3)	41 (10)
Blacks (%) 68.8 27 Blacks (%) 38.4 24 Education (<high (%)<="" school)="" td="">$38.4$$24$BMI (kg/m²)*$30.4$ (7.8)27.3 (5.1)BMI (kg/m²)*30.4 (7.8)27.3 (5.1)Dobese (BMI > 30) %$44.8$$26.5$Diabetes (%)$26.5$11Hypertension (%)$62.5$$33$Myocardial infraction (%)$8.6$$4$</high>	62.6 54	57.6	53	53
Education (<high (%)<="" school)="" td=""> 38.4 24 BMI (kg/m²)* 30.4 (7.8) 27.3 (5.1) Dhese (BMI > 30) % 44.8 26 Obese (BMI > 30) % 24.8 26 Diabetes (%) 26.5 11 Hypertension (%) 62.5 33 Myocardial infaction (%) 8.6 4 Stroke (%) 9.6 1.4</high>	68.8 27	15.7	26.1	$0 \approx$
$BMI (kg/m^2)^*$ $30.4 (7.8)$ $27.3 (5.1)$ $Dbese (BMI > 30) \%$ 44.8 26 $Diabetes (\%)$ 26.5 11 $Hypertension (\%)$ 62.5 33 $Myocardial infraction (\%)$ 8.6 4 Stroke (\%) 9.6 1.4	38.4 24	29.5	16	26 <i>§</i>
Obese (BMI > 30) % 44.8 26 Diabetes (%) 26.5 11 Hypertension (%) 62.5 33 Myocardial infraction (%) 8.6 4 Stroke (%) 9.6 1.4	30.4 (7.8) 27.3 (5.1)	26.7 (4.7)	28.0 (5.4)	24.9 (3.8)//
Diabetes (%) 26.5 11 Hypertension (%) 62.5 33 Myocardial infarction (%) 8.6 4 Stroke (%) 9.6 1.4	44.8 26	19	32	10.8//
Hypertension (%) 62.5 33 Myocardial infarction (%) 8.6 4 Stroke (%) 9.6 1.4	26.5 11	16.4	11.6	4.1
Myocardial infarction (%) 8.6 4 Stroke (%) 9.6 1.4	62.5 33	57.7	42	7
Stroke (%) 9.6 1.4	8.6 4	9.6	n/a	$1.6^{#}$
	9.6 1.4	4.2	n/a	0.5 **
Ever smoked cigarettes (%) 65.3 58.2	65.3 58.2	53.5	49	57.3 [§]

* Tabulated values are mean (SD). n/a = not applicable.

Circ Heart Fail. Author manuscript; available in PMC 2018 March 01.

 $\dot{\tau}^{t}$ By design, participants enrolled in MESA were free of CVD at baseline so the prevalence of MI and stroke at baseline in this cohort may be best described as not applicable.

tThe baseline data for the FHS pertains whenever available to both the original cohort and the offspring cohort given this larger sample was utilized to compute HF incidence rates referenced in the manuscript. In the absence of such data, we have presented data from the original cohort (or subsamples thereof) as a proxy and indicated so in each case.

 g Obtained from a subsample of the parental FHS cohort data (n=1037, 45–62yrs and CHD-free at baseline).

%Obtained from a subsample of the parental FHS cohort data (n=2922, 30–62yrs and CHD-free at baseline).

(Obtained from the parental FHS cohort data (n=5209) and pertains to the composite of MI, CHD-related sudden death and angina pectoris.

** Obtained from the parental FHS cohort data (n=5209). ARIC: Atherosclerosis Risk in Communities; CHS: Cardiovascular Health Study; SCCS: Southern Community Cohort Study; MESA: Multi-ethnic Study of Atherosclerosis; FHS: Framingham Heart Study.