Heart Failure Incidence and Mortality in the Southern Community Cohort Study

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Background—There is a paucity of data on heart failure (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 United States.

Methods and Results—Participants were 27078 white and black men and women enrolled during 2002 to 2009 in the SCCS (Southern Community Cohort Study) who had no history of HF and were receiving Centers for Medicare and Medicaid Services. Incident HF diagnoses through December 31, 2010 were ascertained using International Classification of Diseases 9th Revision codes 428.x via linkage with Centers for Medicare and Medicaid Services research files. Most participants were black (68.8%), women (62.6%), and earned <$15000/y (69.7%); mean age was 55.5 (10.4) years. Risk factors for HF were common: hypertension (62.5%), diabetes mellitus (26.5%), myocardial infarction (8.6%), and obesity (44.8%). Over a median follow-up of 5.2 years, 4341 participants were diagnosed with HF. The age-standardized incidence rates were 34.8, 37.3, 34.9, and 35.6 /1000 person-years 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 of 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 women compared with white women.

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. (Circ Heart Fail. 2017;10:e003553. DOI: 10.1161/CIRCHEARTFAILURE.116.003553.)

Key Words: epidemiology ■ heart failure ■ incidence ■ population ■ survival
patterns by race and sex, in a low-income underinsured population underrepresented in previous studies. Between 2002 and 2009, the SCCS enrolled ≈66000 adults (≈two-thirds black) aged 40 to 79 years living in 12 southeastern states to investigate various chronic disease outcomes. Approximately 96% of participants were recruited at community health centers, which provide primary health and preventive care services for low-income populations, so that the cohort is made up of sizeable numbers in previous cohort studies, particularly those investigating CVD. 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 computer-assisted personal interviews for community health center participants, and via self-administered mailed questionnaire for the general population participants. Detailed description of SCCS methods has been previously published.

SCCS participants (n=27078) included in the current analyses were individuals aged ≥65 years (n=77001) at cohort enrollment, or individuals aged <65 years (n=20077) at enrollment who (1) reported being covered by Medicare (which provides medical benefits to low-income adults and uninsured individuals) on the baseline questionnaire, (2) reported being covered by Medicare (the primary health insurance program for individuals aged ≥65 years) on the baseline questionnaire, or (3) 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 or Medicaid from the time of SCCS enrollment to the end of the follow-up period (December 31, 2010), for the ascertainment of incident HF events. Analyses were restricted to self-reported African American or black and non-Hispanic white SCCS participants, as too few individuals in other racial groups were available for stable statistical analysis.

**Outcome Ascertainment**

HF events were ascertained via linkage of the SCCS cohort with CMS Research Identifiable Files (which include Medicare institutional and noninstitutional 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 (ICD-9), discharge code of 428.x (428.0–428.9) within the Medicare institutional (Medicare Provider Analysis and Review [MEDPAR], which includes inpatient, outpatient, and skilled nursing facility base files), Part B carrier (includes noninstitutional physician services and durable medical equipment), or outpatient-based claims files or the Medicaid Analytic Extract Inpatient and Other Services claims files, from the date of SCCS enrollment through December 31, 2010. Detailed description of the CMS research files are published elsewhere.

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 epidemiological researchers and the National Death Index through December 31, 2010. Both National Death Index and SSA are well-established and reliable means of identifying deaths in the United States, and are expected to capture nearly all deaths.

**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 the date of entry into the SCCS until the date of the first diagnosis of HF, date of death, or December 31, 2010, whichever occurred first. IRs of HF 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 1000 person-years (PY). The 95% confidence intervals (CIs) were calculated using the quadratic approximation to the Poisson log likelihood for the log-rate parameter. 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 used 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 (BMI; restricted cubic splines with 4 knots), and history of diabetes mellitus, hypertension, high cholesterol, myocardial infarction (MI)/coronary artery bypass graft or stroke (all yes/no); and model 3 additionally adjusted for the following covariates: annual household income (<$15000, $15000–$24999, and ≥$25000), education (chips school, high school/vocational training/junior college, and college degree or higher), smoking (never, former, current <19.5 pack-years, current ≥19.5 pack-years, and 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, and single/never married), and enrollment source (community health centers versus general population). Knots were placed at quantiles of covariate distributions, equally spaced in sample size. 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, 2010, whichever occurred first. When the date of death was coincident with the date of HF diagnosis, follow-up time was set to 0.5 days. We computed cumulative mortality for both HF cases and noncases 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 of <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, TX, USA) and the “rms” package for R version 3.1.1 (R Core Team 2014).

**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 27078 SCCS participants included in this study, 68.8% were black, 62.6% were women, 69.7% had annual household income <$15000, 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 mellitus (26.5%); MI (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 mellitus and hypertension.

**HF Incidence**

Over a median (25th, 75th percentile) follow-up time of 5.2 (3.1, 6.7) years, 4341 participants (16%) developed incident HF (IR, 32.8/1000 PY; 95% CI, 31.8–33.8). White men had the highest age-standardized IR, 37.3/1000 PY, compared
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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 (hazard ratio, 0.89; 95% CI, 0.82–0.98). The risk of HF was similar among white men (hazard ratio, 1.09; 95% CI, 0.97–1.23) and black men (hazard ratio, 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 4341 individuals who developed incident HF, 952 died (cumulative mortality=21.9%) over a median (25th, 75th percentile) post-HF follow-up time of 2.3 (0.9, 4.2) years (Table 3). Men had higher percent mortality than women (29% versus 18%), with little difference by race. In individuals without HF (n=22 737), there were 1929 deaths, corresponding to a percent mortality of 8.5%.

Figure 1 shows age-adjusted survival curves for individuals 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 mellitus, high cholesterol, history of MI 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 nonsignificant relative differences in risk by race in all models. The race–sex interaction term was not statistically significant (P=0.92).

Discussion

We investigated HF incidence and post-HF mortality in a large multi-ethnic low-income sample from the southeastern United States. Our principal findings are as follows: (1) the IR for HF was remarkably high across all race and sex groups in the SCCS, (2) there was no significant difference in age-adjusted IRs 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) men had higher post-HF mortality with no significant racial differences.

The IRs for HF in the SCCS exceeded those previously reported from established CVD cohorts (Table 4). In CHS (n=5888; age range: ≥65 years), for example, the age-standardized HF IRs after 10 years of follow-up among white men, black men, white women, and black women were 30.2, 19.2, 19.2, and 22.6/1000 PY, respectively. In ARIC (n=14,933; age range: 45–64 years), Loehr et al found IRs of 6.0, 9.1, 3.4 and 3.8/1000 PY, respectively, in these race-sex groups. The FHS and MESA reported even lower IRs (7.2 and 4.2/1000 PY in men and women in FHS and 3.1/1000 PY overall in men and women in MESA).

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Table 2. Risk of Incident HF Among Participants in the Southern Community Cohort Study, Overall and Stratified by Race and Sex

<table>
<thead>
<tr>
<th></th>
<th>Overall, n=27,078</th>
<th>White Women, n=5,252</th>
<th>White Men, n=3,202</th>
<th>Black Women, n=11,688</th>
<th>Black Men, n=6,936</th>
</tr>
</thead>
<tbody>
<tr>
<td>Incident HF cases, n</td>
<td>4,341</td>
<td>801</td>
<td>511</td>
<td>1,940</td>
<td>1,089</td>
</tr>
<tr>
<td>Person-years (PY)</td>
<td>132,500</td>
<td>23,339</td>
<td>13,934</td>
<td>60,639</td>
<td>34,589</td>
</tr>
<tr>
<td>Cumulative incidence, %</td>
<td>16.0</td>
<td>15.3</td>
<td>16.0</td>
<td>16.6</td>
<td>15.7</td>
</tr>
<tr>
<td>Incidence rate/1000 PY (95% CI)</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Crude</td>
<td>32.8 (31.8–33.8)</td>
<td>34.3 (32.0–36.8)</td>
<td>36.7 (33.6–40.0)</td>
<td>32.0 (30.6–33.4)</td>
<td>31.5 (29.7–33.4)</td>
</tr>
<tr>
<td>Age-adjusted</td>
<td>35.1 (34.1–36.2)</td>
<td>34.8 (32.4–37.2)</td>
<td>37.3 (34.0–40.6)</td>
<td>35.6 (33.9–37.2)</td>
<td>34.9 (32.7–37.1)</td>
</tr>
<tr>
<td>Hazard Ratio (95% CI)</td>
<td>1.00</td>
<td>1.02 (0.91–1.14)</td>
<td>0.91 (0.83–0.99)</td>
<td>1.06 (0.97–1.17)</td>
<td>1.04 (0.94–1.15)</td>
</tr>
</tbody>
</table>

CI indicates confidence interval; HF, heart failure; and MI, myocardial infarction. Model 1: Includes age (restricted cubic splines with 4 knots), race, and sex. Model 2: Model 1+body mass index (restricted cubic splines with 4 knots), history of diabetes mellitus, hypertension, high cholesterol, MI, and stroke (all yes/no). Model 3: Model 2+annual household income (<$15,000, $15,000–$24,999, and ≥$25,000), education (<high school, high school/vocational training/junior college, and college degree or higher), smoking (never, former, current <19.5 pack-years, and current ≥19.5 pack-years) and alcohol intake (linear and quadratic term), marital status (married/living as married with partner, separated/divorced, widowed, and single/never married), and enrollment source (community health centers vs general population). P value for race×sex interaction=0.22.

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Table 3. Percent Mortality of SCCS Participants According to Heart Failure Status, Overall and Stratified by Race and Sex

<table>
<thead>
<tr>
<th></th>
<th>Overall, n=4,341</th>
<th>White Women, n=801</th>
<th>White Men, n=511</th>
<th>Black Women, n=1,940</th>
<th>Black Men, n=1,089</th>
</tr>
</thead>
<tbody>
<tr>
<td>Deaths, n</td>
<td>952</td>
<td>144</td>
<td>152</td>
<td>343</td>
<td>313</td>
</tr>
<tr>
<td>Mortality, %</td>
<td>21.9</td>
<td>18.0</td>
<td>29.7</td>
<td>17.7</td>
<td>28.7</td>
</tr>
<tr>
<td>Risk of death: Hazard ratio (95% CI)</td>
<td>1.00 (ref)</td>
<td>1.73 (1.37–2.17)</td>
<td>0.91 (0.75–1.10)</td>
<td>1.61 (1.32–1.96)</td>
<td></td>
</tr>
</tbody>
</table>

CI indicates confidence interval; and SCCS, Southern Community Cohort Study. Model 1: Includes age (restricted cubic splines with 4 knots), race, and sex. Model 2: Model 1+body mass index (restricted cubic splines with 4 knots), history of diabetes mellitus, hypertension, high cholesterol, myocardial infarction, and stroke (all yes/no). Model 3: Model 2+annual household income (<$15,000, $15,000–$24,999, and ≥$25,000), education (<high school, high school/vocational training/junior college, and college degree or higher), smoking (never, former, current <19.5 pack-years, and current ≥19.5 pack-years) and alcohol intake (linear and quadratic term), marital status (married/living as married with partner, separated/divorced, widowed, and single/never married), and enrollment source (community health centers vs general population). P value for race×sex interaction=0.92.
The higher SCCS HF incidence could be explained, in part, by notably higher prevalence of CVD risk factors (in particular hypertension, diabetes mellitus, obesity, 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 of <$15,000. Previous evidence suggests a strong independent association between socioeconomic status and HF risk; 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 IR of HF in the SCCS, consistent with findings from the CHS. However, minimal differences in age-adjusted IRs and HF risk between groups after adjustment for CVD risk factors (except for black women who had significantly lower risk) suggest the homogeneity of HF risk profile. Similarly, in ARIC, crude racial and sex differences in incidence density were attenuated by adjustment for CVD risk factors.

Overall, the 5-year post-HF survival in SCCS was higher than the 52% previously reported (data from the Olmsted county study). This may be due, in part, to the fact that SCCS participants had shorter post-HF follow-up time and were younger at baseline (55.5 versus 74 years), and to temporal trends suggesting improved post-HF survival 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, which has a less favorable prognosis compared with HF with preserved ejection fraction.

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 after admission for HF.

### Table 4. Comparison of Heart Failure Incidence Between SCCS, ARIC, MESA, and CHS Cohorts

<table>
<thead>
<tr>
<th></th>
<th>SCCS,6 45–64 y</th>
<th>ARIC,6 45–64 y</th>
<th>SCCS,6 65–79 y</th>
<th>CHS,5† ≥65 y</th>
<th>SCCS,6 40–79 y</th>
<th>MESA,6 45–84 y</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>n=15,321</td>
<td>n=14,933</td>
<td>n=7,001</td>
<td>n=5,888</td>
<td>n=27,078</td>
<td>n=68,145</td>
</tr>
<tr>
<td>Overall</td>
<td>35.8</td>
<td>5.7</td>
<td>39.5</td>
<td>19.3</td>
<td>24.0</td>
<td>33.7</td>
</tr>
<tr>
<td>White women</td>
<td>38.3</td>
<td>3.4</td>
<td>34.6</td>
<td>14.5</td>
<td>19.2</td>
<td>34.3</td>
</tr>
<tr>
<td>White men</td>
<td>39.7</td>
<td>6.0</td>
<td>38.3</td>
<td>24.9</td>
<td>30.2</td>
<td>34.3</td>
</tr>
<tr>
<td>Black women</td>
<td>35.1</td>
<td>8.1</td>
<td>42.4</td>
<td>19.6</td>
<td>22.6</td>
<td>33.4</td>
</tr>
<tr>
<td>Black men</td>
<td>34.0</td>
<td>9.1</td>
<td>41.9</td>
<td>23.5</td>
<td>27.5</td>
<td>4.6</td>
</tr>
</tbody>
</table>

ARIC indicates Atherosclerosis Risk in Communities; CHS, Cardiovascular Health Study; HF, heart failure; MESA, Multi-Ethnic Study of Atherosclerosis; and SCCS, Southern Community Cohort Study.

Incidence rates computed for SCCS participants aged 45–64, 65–79, and 40–79 years 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.

†The tabulated values are computed from values presented in Arnold et al and standardized to the age distribution of CHS participants.
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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 after admission, but the differences by sex were nonsignificant. In CHS, the mortality rate in white women, white men, black women, and black men was 35.5, 40.5, 33.6, and 44.4/1000 PY, respectively. After full adjustment for covariates, there were no significant racial differences, but women had a 15% lower risk of all-cause mortality (hazard ratio, 0.85; 95% CI, 0.73–0.99). In SCCS, white men had the lowest 5-year survival post-HF diagnosis; but after full adjustment, there were mainly sex differences in post-HF mortality with higher risk of death among men and no significant racial 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 with reduced ejection fraction and worse prognosis compared with HF with preserved ejection fraction. 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 used by the SCCS was tailored toward low-income, rural, and underinsured 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 used in other cohorts. A review of the detection of HF in administrative claims data that included eight studies conducted among Medicare beneficiaries reported positive predictive values between 76% and 99%, with the majority of the studies reporting positive predictive values over 90%. These codes have also been used with high specificity in several studies, although no independent validation was conducted by the SCCS investigators. An over-representation of groups with elevated HF risk (individuals > 65 years and individuals <65 years receiving Medicare) in our SCCS subcohort compared with the SCCS base population may have contributed to higher HF IRs than would be expected for the total SCCS cohort. However with the mean age of the total cohort being 52.6 years versus 55.5 years for our subcohort, 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%, it is plausible that we may have underestimated the IR of HF in our subcohort. Also, when contrasting the IRs between our study and previous CVD cohorts (such as 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 about the
continuous coverage in CMS of individuals younger 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 individuals aged <65 years 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 the use of medications (diabetes mellitus and hypertension). Although self-report may be susceptible to recall and misclassification bias, these methods have been successfully used and validated in large epidemiological 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 mellitus, height, and weight.9

The SCCS cohort comprised a substantial number of individuals from minority and low-income populations who are traditionally under-represented in most studies investigating CVD and HF in particular. The IRs 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 National Death Index and SSA allowed for robust ascertainment of all-cause mortality.

In conclusion, in this low-income multiethnic population, we found higher IRs 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.

Acknowledgments

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Disclosures

None.

References


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