Heterogeneous Trends in Burden of Heart Disease Mortality by Subtypes in the United States, 1999-2018: Observational Analysis of Vital Statistics

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| Author: |  | 09-Mar-2020 $\quad$| Complete List of Authors: | Shah, Nilay; Northwestern University Feinberg School of Medicine, <br> Preventive Medicine <br> Molsberry, Rebecca; Northwestern University Feinberg School of <br> Medicine <br> Rana, Jamal; Kaiser Permanente Northern California <br> Sidney, Steve; Kaiser Permanente Northern California <br> Capewell, Simon; University of Liverpool, Department of Public Health <br>  <br> Policy <br> Carnethon, Mercedes; Northwestern University, Department of <br> Preventive Medicine <br> Lloyd-Jones, Donald; Northwestern University, Department of Preventive <br> Medicine <br> Khan, Sadiya; Northwestern University Feinberg School of Medicine |
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# Heterogeneous Trends in Burden of Heart Disease Mortality by Subtypes in the United States, 1999-2018: Observational Analysis of Vital Statistics 

Nilay S. Shah, MD, MPH, Postdoctoral Fellow; ${ }^{1,2}$ Rebecca Molsberry, MPH, Statistician; ${ }^{1}$ Jamal S. Rana MD, PhD, Chief of Cardiology; ${ }^{3}$ Stephen Sidney MD, MPH, Senior Research Scientist, ${ }^{3}$ Simon Capewell, DSc, MD, MBBS, Professor; ${ }^{4}$ Martin O’Flaherty MD, MSc, PhD, Hon(FPH), Professor; ${ }^{4}$ Mercedes Carnethon, PhD, Professor; ${ }^{1}$ Donald M. Lloyd-Jones, MD, ScM, Professor; ${ }^{1,2}$ Sadiya S. Khan, MD, MSc, Assistant Professor ${ }^{1,2}$

## Author Affiliations:

${ }^{1}$ Department of Preventive Medicine, Northwestern University Feinberg School of Medicine, 680 N. Lake Shore Drive, Suite 1400, Chicago, Illinois, 60611 USA
${ }^{2}$ Division of Cardiology, Department of Medicine, Northwestern University Feinberg School of Medicine, 676 N. St. Clair Street, Suite 600, Chicago, Illinois, 60611 USA
${ }^{3}$ Kaiser Permanente Northern California, 2000 Broadway, Oakland, California, 94612 USA
${ }^{4}$ Institute of Population Health Sciences, University of Liverpool, Waterhouse Building, Block B, Brownlow Street, Liverpool, L69 3GL United Kingdom

Author contributions: All authors contributed substantially to this manuscript and meet criteria for authorship, as outlined in CrediT Contribution designations provided upon submission.

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## Address for Correspondence:

Sadiya S. Khan, MD, MS, FACC, Assistant Professor of Medicine, Northwestern University
680 N. Lake Shore Drive, 14-002, Chicago, IL 60611
Tel: +1-312-503-2515, Fax: +1-312-908-9588
s-khan-1@northwestern.edu


#### Abstract

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Objective: To describe trends in burden of subtypes of heart disease (HD) mortality from 1999-2018, in order to inform targeted prevention strategies and reduce disparities.

Design: Serial cross-sectional analysis of cause-specific HD mortality rates using national death certificate data.

Setting: United States, 1999-2018
Participants: 12.9 million decedents from HD ( $49 \%$ women, $12 \%$ black, and $19 \%<65$ years old).
Main Outcome Measures: Age-adjusted mortality rates (AAMR) and years of potential life lost (YPLL) for each HD subtype as underlying cause of death, and mean annual percentage change (APC).

Results: Total HD deaths fell from 752,192 to 596,577 from 1999-2011, then increased to 655,381 in 2018. Proportions of deaths from each of the top 3 subtypes of HD deaths changed during the study period, including decrease in ischemic HD (IHD, $73 \%$ to $56 \%$ ) compared with increases in heart failure (HF, $8 \%$ to $13 \%$ ) and hypertensive HD (HTN-HD, $4 \%$ to $9 \%$ ). AAMR was highest from IHD, but APC from HF and HTN-HD was highest after 2011. The steepest increases in HF were in black men (APC $+4.9 \%[+4.0$ to +5.8$])$ whereas fastest increases in HTN-HD occurred in white men $(+6.3 \%[+4.9$ to $+9.4]$ ). The most rapid increases in midlife mortality (45-64 years) from HD occurred in black women due to HF $(+7.0 \%[+5.1$ to +8.9$])$, and in white women due to HTN HD $(+4.1 \%[+3.7$ to +4.5$])$. The burden of YPLL was greatest for IHD, but black-white disparities in YPLL were driven by HF and HTNHD.

Conclusions: AAMR from almost all subtypes of HD have become stagnant or increased since 2011. Increases in HD AAMR are attributed to HF and HTN-HD, which also account for the widest black-white disparities in YPLL. Early and targeted prevention and control of HTN is needed with a focus on highrisk groups to promote health equity.

## Summary Box

## Section 1 (What is already known):

- Surveillance data from national death certificates show that declines in heart disease mortality have slowed since 2011 in the United States.
- Advances in medical and surgical therapies may have led to heterogeneous patterns in death rates for specific subtypes of heart disease.
- A detailed understanding of how mortality from subtypes of heart disease contributes to overall heart disease mortality patterns will inform implementation of multi-level prevention interventions.


## Section 2 (What this study adds):

- In the United States, increases in heart failure and hypertensive heart disease mortality occurred parallel to declines in ischemic heart disease death rates, resulting in 3.5 million years of potential life lost from total heart disease in 2018 and warranting focus on reducing the growing burden of hypertension and other heart failure risk factors.
- Black men had the highest burden of mortality from leading subtypes of heart disease, but the greatest increases midlife mortality due to heart disease subtypes were observed in black women (heart failure) and white women (hypertensive heart disease), indicating the need to enhance prevention in high risk populations to reduce disparities and promote health equity.


## Introduction

Age-adjusted mortality rates (AAMR) due to heart diseases (HD) declined by more than $50 \%$ in the United States (US) in the second half of the $20^{\text {th }}$ century from a peak of 307 per 100,000 in 1950 . This fall was predominantly driven by rapid declines in mortality from ischemic heart disease (IHD), as a result of progress in prevention (e.g. decline in smoking rates) and evidence-based treatments (e.g. statins, anti-platelet agents, and anti-hypertensives). ${ }^{1-3}$ While mortality rates continued to decline after 2000, ${ }^{4}$ recent data demonstrated that HD AAMR plauteaued in 2011. ${ }^{256}$ Furthermore, some indicators now suggest a trend reversal with increasing HD death rates in certain population subgroups, such as middleaged Americans, ${ }^{78}$ thus undoing decades of progress in HD prevention and treatment and contributing to decreases in overall US life expectancy. ${ }^{9}$

However, subtypes of HD are heterogeneous in their pathophysiology and contribution toward preventable deaths. Recent analysis indicates that the stagnation in HD mortality rates in 2011 was driven by a slowing in the decline of IHD deaths, and increases in heart failure (HF) mortality rates. ${ }^{1011}$ Further detailed characterization of patterns for each subtype of HD across race-sex and age subgroups that are responsible for the observed changes in HD mortality rates would inform targeted prevention and treatment efforts. We therefore investigated trends across race-sex and age groups in cause-specific HD deaths (IHD, HF, hypertensive HD [HTN HD], valvular HD, arrhythmia, pulmonary HD, and other HD) using national death certificate data from 1999 to 2018 . We calculated complementary metrics to assess burden due to subtypes of HD , including age-adjusted mortality rates (AAMR), years of potential life lost (YPLL), and mean annual percent changes (APC) in AAMR and YPLL from HD subtypes to better evaluate the burden of avoidable HD mortality overall and by race-sex groups.

## Methods

Using mortality data from the Centers for Disease Control and Prevention's Wide-Ranging Online Data for Epidemiologic Research (WONDER) database of death certificates, we identified decedents from 1999 to 2018 by International Classification of Diseases (ICD)-10 code with an
underlying cause of death of HD (total), which comprised of HD subtypes classified as: IHD (ICD-10 code I20-I25), HF (I50), HTN HD (I11, I13), valvular HD (I34-I38), arrhythmia (I47-I49), pulmonary HD (I26-I28), and other HD (I00-I09, I30-I33, I40-I46, I51, which includes acute and chronic rheumatic heart disease, pericardial diseases, acute myocarditis, and unspecified cardiomyopathy and cardiac arrest not otherwise defined). The percentage of each HD subtype's contribution to total HD was calculated overall and by race-sex and age subgroups.

Trends were examined in the overall population (all race groups), separately by race-sex groups (black women, white women, black men, white men), and stratified by age at death ( $<45$ years, 45-64 years, $\geq 65$ years). The race-sex analysis was limited to black and white individuals because other race/ethnic groups (Asian Americans, Native Americans, Hispanic Americans) were either too small (e.g., Native Americans) or less reliably identified (e.g., Hispanic American, Asian Americans ${ }^{12}{ }^{13}$ ) in CDC

## WONDER.

We calculated AAMR per 100,000 population adjusted using the 2000 US standard population overall and stratified by race-sex and age subgroups. ${ }^{14}$ YPLL was calculated using standard methods previously employed. ${ }^{1315}$ An average life expectancy between 1999-2018 for the total US population and individually for race-sex subgroups was obtained from US vital statistics reports, and was used as the reference age for YPLL calculation. ${ }^{16}$ For each year of analysis, YPLL was calculated by multiplying number of deaths from HD subtypes in each of a series of 5-year age group decrements from the reference age, by the difference between the reference age and the midpoint age at death within each 5-year age group. This result was divided by the total 5-year age group population, then multiplied by 100,000 to obtain YPLL per 100,000 population. After age standardizing YPLL with the 2000 US standard population, the sum of YPLL in each 5-year age group provided cause-specific YPLL per 100,000 in the total population and race-sex subgroups, for each year of analysis. YPLL was calculated for total HD, IHD, HF, and HTN HD. Because of relatively few deaths due to other HD subtypes, YPLL was also calculated for an All Other HD category that comprised the remainder of the underlying cause of death codes.

Temporal trends in AAMR and YPLL were characterized by fitting log-linear regression models using Joinpoint Regression Program (National Cancer Institute). ${ }^{18}$ Log-transformed data were used as certain subtypes of HD had relatively few deaths. We used Joinpoint to identify an inflection point in the overall trend where significant temporal variation existed in the trend ( $\mathrm{p}<0.05$ ), the temporal intervals from which were applied to all subsequent analyses to calculate mean annual percent change (APC). Within each individual model, the slopes of trend intervals between up to three inflection points were used to calculate the annual percent change for each trend segment, and the weighted average of the annual percent change was reported as the mean annual percent change (APC) of AAMR and YPLL before and after 2011 (which was the inflection point identified for total HD during the study period, consistent with previously published trends identified with Joinpoint Regression ${ }^{5618}$ ), with weights equal to the length of the inter-inflection point annual percent change for each model. APCs based on the linear trend segments identified are dependent on the model-determined number of inflection points (i.e., if the trend is optimized with 0 inflection points, indicating 1 linear trend across the study period, APC before and after 2011 would be the same).

As a secondary analysis, AAMR and YPLL from HF as a contributing cause of cardiovascular death was evaluated using the CDC WONDER multiple cause of death files, in order to more broadly characterize quantify the burden of HF-related mortality. These metrics were calculated as previously described, for deaths defined as any mention of HF (ICD-10 code I50) on death certificates with any cardiovascular disease (I00-I78) listed as underlying cause of death.

This study was determined exempt from review by the Northwestern University institutional review board. As the results are based on publicly available national vital statistics, neither patients nor the general public were directly involved in the conduct of this analysis and direct dissemination is not applicable.

## Results

Study Population

Between 1999 and 2018 there were 12.9 million deaths from HD, of whom $49.1 \%$ were women, $11.6 \%$ were black, and $86.2 \%$ were white. From 1999 to 2018, IHD accounted for a declining proportion of total HD deaths (from $73.0 \%$ to $55.8 \%$ ), alongside increasing proportion of total HD from HF ( $7.6 \%$ to $12.8 \%$ ), HTN HD ( $3.6 \%$ to $9.2 \%$ ), valvular HD ( $2.7 \%$ to $3.7 \%$ ), arrhythmia ( $2.5 \%$ to $5.6 \%$ ), pulmonary HD ( $1.8 \%$ to $2.8 \%$ ), and other HD ( $8.7 \%$ to $10.3 \%$ ); see Figure 1. Overall, $2.6 \%$ of decedents were $<45$ years old, $16.3 \%$ were $45-64$ years old, and $81.2 \%$ were $\geq 65$ years old. The proportion of deaths among middle-aged decedents (45-64 years) increased from 13.7\% in 1999 to 17.3\% in 2018.

Total HD AAMR declined from 267 per 100,000 in 1999, to 164 per 100,000 in 2018 (Table 1). APC of overall HD AAMR declined, from -3.7\% per year ( $95 \%$ confidence interval -3.8 to -3.5 ) before 2011 to $-0.7 \%$ per year ( -1.1 to -0.3 ) after 2011 for all decedents. AAMR from IHD declined from 195 per 100,000 in 1999 , to 109 per 100,000 in 2011 , but further declined only to 91 per 100,000 in 2018 . The rate of AAMR declines (i.e., APC) from IHD-mortality slowed from $-4.7 \%$ per year ( -5.1 to -4.2 ) before 2011 to $-2.6 \%$ per year ( -3.3 to -2.0 ) after 2011. Trends in HF mortality rates reversed and increased (overall APC $-1.7 \%$ per year [ -2.5 to -0.8 ] before 2011, versus $+3.5 \%$ per year $[+2.6$ to +4.5$]$ after 2011), reflecting a decline in AAMR from 20 per 100,000 in 1999, to 17 per 100,000 in 2011, which rose to 21 per 100,000 again by 2018. Overall, APC of HTN HD mortality rates accelerated from $+1.2 \%$ per year $(+0.7$ to +1.7$)$ before 2011 to $+4.8 \%$ per year ( +3.3 to +6.3 ) after 2011. Absolute number of deaths from and AAMR of valvular HD, arrhythmia, and pulmonary HD were relatively low in comparison to IHD, HF, and HTN HD. While AAMR from valvular HD decreased from 2011 to 2018 (from 7 to 6 per 100,000 , APC $-1.6 \%$ per year [-2.8 to -0.4]), AAMR from arrhythmia and pulmonary HD increased to 9 per 100,000 and 5 per 100,000, respectively.

## Trends in HD Mortality Rates by Race-Sex

Black men had the highest AAMR from total HD (260 per 100,000 in 2018), which was reflected primarily in high AAMR from IHD (138 per 100,000 in 201), HF (29 per 100,000), and HTN HD (35 per $100,000)$. Trends in AAMR from total HD and IHD slowed similarly after 2011 in black and white
women and men (Figure 2 and Supplemental Table 1). All race-sex groups experienced a reversal of HF mortality trends after 2011; notably, after 2011 APC of HF mortality rates was highest in black men $(+4.9 \%$ per year $[+4.0$ to +5.8$])$ compared to black women, white men, and white women. Acceleration of HTN HD mortality rates was highest in white women (APC $+5.6 \%$ per year $[+4.3$ to +7.0$]$ ) and white men (APC $+6.3 \%$ per year $[+4.9$ to 9.4$]$ ) after 2011. Although APC of HTN HD mortality rates was lower in black women and black men, absolute AAMR was higher in black compared to white decedents.

## Trends in HD Mortality Rates by Age

AAMR was lowest for decedents $<45$ years old at time of death (Table 2). In this youngest age stratification, AAMR from any HD was 9 per 100,000 in 2018, predominantly attributable to IHD (AAMR 3 per 100,000 in 2018) and HTN HD (AAMR 1 per 100,000). APC of HTN HD mortality rates declined from $+4.8 \%$ per year $(+3.7$ to +5.9$)$ before 2011 to $+2.3 \%$ per year $(+1.9$ to +2.8$)$ after 2011 . However, APC of HF mortality rates accelerated after 2011, and APC of valvular HD reversed from decreasing to increasing.

For decedents age 45-64 years at time of death (Table 2), AAMR from all HD declined from 1999 to 2011 (from 164 to 121 per 100,000; APC $-2.5 \%$ per year [-2.8 to -2.2 ]), but remained generally unchanged between 2011 to 2018 (to 122 per 100,000; APC $+0.2 \% /$ year [ -0.1 to +0.4$]$ ). AAMR was highest for IHD ( 73 per 100,000 in 2018), HTN HD ( 15 per 100,000 in 2018) , and HF (7 per 100,000 in 2018). APC of IHD mortality rates declined after 2011, but accelerated for HTN HD mortality rates (APC $+3.0 \%$ per year $[+2.6$ to +3.5$]$ ) and HF mortality rates $(+5.7 \%$ per year [ +4.4 to 7.0$]$ ).

Decedents aged $\geq 65$ years had the highest AAMR for all types of HD compared to younger decedents (Table 2). AAMR for overall HD declined from 1767 per 100,000 in 1999, to 1116 per 100,000 in 2011 (APC -3.7\% per year [-4.3 to -3.1]), but further declined only to 1035 per 100,000 between 2011 to 2018 (APC $-1.1 \%$ per year [-1.6 to -0.6$]$ ). Similar to mortality patterns at younger ages, HTN HD mortality rate trends increased after 2011, and HF mortality rate trends reversed to increasing.

## Trends in Years of Potential Life Lost due to Subtypes of HD

Estimated YPLL per 100,000 population is shown in Table 3 and the Supplemental Figure. Total YPLL in the US population was estimated to be 1068 years per 100,000 population in 2018 . Of all the HD subtypes, YPLL in 2018 was highest for IHD (578 years per 100,000), followed by HTN HD (132 years per 100,000 ) then HF (64 years per 100,000). YPLL due to IHD was similar in white and black men, but 1.3-fold higher in black women compared with white women. For IHD, APC of YPLL declined from $-3.8 \%$ per year ( -3.9 to -3.7 ) overall before 2011 to $-1.6 \%$ per year ( -1.8 to -1.4 ) after 2011 with similar trends in all race-sex groups. YPLL from HTN HD was 2.9-fold higher in black women (219 years per 100,000 ) and 2.3-fold higher in black men ( 305 per 100,000 ) compared with white women and men in 2018 , respectively. However, greater increases in YPLL over the study period was observed in white women (APC $+4.2 \%$ per year [0.9 to 7.7 ] after 2011 ) and white men $(+3.3 \%$ per year [3.0 to 3.5 ] after 2011). For HF, YPLL changed from either decreasing or stagnant prior to 2011, to increasing after 2011. Black women and men had a 2-fold higher YPLL due to HF compared with white women and men, with highest APC of YPLL in black women $(+5.8 \%$ per year [4.5 to 7.2$])$ and black men $(+6.6 \%$ per year [5.0 to 8.3]).

## Discussion

Changing mortality patterns in HD subtypes in the US are reflected overall by a plateau in total HD AAMR since 2011, within which slowing in declines in IHD mortality rates are coupled with considerable increases in HF and HTN HD deaths in all sex-race groups. In particular, black men had both the highest AAMR and fastest increase in mortality rates due to HF. While HTN HD mortality rates increased for all race-sex groups, fastest increases in HTN HD AAMR were observed in white women and men between 2011-2018. However, AAMR for HTN HD remained about 2 times higher in black compared with white populations by 2018 . Our comprehensive estimates of the burden of HD leveraging YPLL translate into approximately 3.5 million potential years of life lost in 2018 due to total HD, but this burden was borne disproportionately in different race-sex groups and attributed to different HD subtypes.

Substantial burden of premature mortality increased from 2 of the top 3 subtypes of HD deaths: HTN HD and HF, reflected by an $80 \%$ and a $31 \%$ increase in YPLL from HTN HD and HF, respectively, from 1999-2018. While HF AAMR was higher than for HTN HD, YPLL from HTN HD was higher indicating that HTN HD-related deaths are occurring at younger ages. Patterns of AAMR from less common causes of HD (arrhythmia, pulmonary HD, and other HD) were either stagnant or increased from 2011 to 2018. Valvular HD was the only subtype where mortality rates declines accelerated after 2011, although relative burden of deaths from valvular HD remained low.

These findings align with recent data showing contemporaneous slowing in the decline of cardiometabolic disease mortality rates, and recent surveillance of HD mortality that showed that the growth in the population of older Americans age $\geq 65$ years during this time was associated with an increase in the number of HD deaths, despite a stagnant or slowly declining AAMR. ${ }^{611}$ Our results build on these reports by providing broader and detailed mortality patterns of the range of HD subtypes as underlying causes of death, particularly highlighting the rapid growth and high burden of death overall from HTN HD, increasing HF and HTN HD mortality rates at younger ages reflected in a high magnitude of YPLL from these causes, and worrisome patterns in mortality from arrhythmia and pulmonary HD.

The observed patterns may be due to the growing burden of cardiometabolic risk factors for HD. Obesity, diabetes, and hypertension likely play a sizeable role in the observed changes in recent HD mortality rates in the US, where the prevalence of obesity now exceeds $42 \%,{ }^{19}$ the prevalence of both prediabetes and diabetes approaches $50 \%,{ }^{20}$ and the prevalence of hypertension is approximately $30 \% .{ }^{21}$ Recent trends in assessment of cardiovascular health in the US project continued worsening in prevalences of poor levels of diet quality, physical inactivity, obesity, and diabetes in the coming decades. ${ }^{22-25}$ Such trends will likely result in continued increases in HF and HTN HD mortality given their direct association with these cardiometabolic risk factors if current trends continue without intervention. ${ }^{26-}$ 28

The observed differences in HD mortality by race may partly reflect underlying and pervasive disparities in cardiovascular health and burden of cardiovascular disease. ${ }^{22}$ Specifically, obesity, diabetes,
and hypertension in the US remain most prevalent in black Americans, and rates of control remain remain lower in black compared with white Americans. ${ }^{29}$ Differences in patterns of guideline-recommended medication use, including medication prescription, optimization, and adherence, especially for HTN, likely contribute to racial disparities in HD morbidity and mortality. ${ }^{30-33} \mathrm{~A}$ growing body of evidence supports that racial differences in cardiovascular health and HD mortality are in large part, representative of disparities in many other factors, including a range of social determinants of health and structural and systemic racism that require individual and policy-level changes to address and reduce persistent health inequities. ${ }^{3435}$

The principal strength of our analysis is investigation of contemporary nationwide mortality data attributed to leading subtypes of HD and differences among race-sex and age subgroups. These data allow for evaluating changes and disparities in cause-specific cardiovascular mortality patterns, which contribute to the observed increases in total HD deaths in the past decade and disproportionally burden black Americans. Limitations of our analysis include incomplete race/ethnic data as our investigation focus on the most reliable data available to evaluate trends in white and black Americans. However, data for disaggregated Asian American subgroups, Hispanic Americans, and Native Americans were either not available or unreliable. Additionally, death certificate data are subject to potential miscoding and misclassification. As the clinical syndrome of HF represents a mode of death frequently attributable to HTN, IHD, valvular HD, or diabetes as the underlying cause, ${ }^{36}$ our primary analysis may underestimate the burden of HF-related mortality. Notably, similar patterns were seen in our secondary analysis evaluating trends in mortality rates from HF as a contributing cause of cardiovascular death. Finally, the 'Other HD' category reflects a heterogenous mix of HD that may have dissimilar pathophysiology.

Future research and public health strategies should focus on dissemination and implementation of interventions that focus on prevention of HD in the highest risk populations in order to promote equity, clinical quality improvement to optimize primary and secondary prevention, and policies to improve diet quality and facilitate physical activity. These strategies may help reduce obesity, diabetes, and hypertension prevalence and improve control of these risk factors, and subsequently reduce deaths due to
the HD subtypes that account for the majority of HD mortality (i.e., IHD, HF, HTN HD). ${ }^{37}$ Ultimately, a comprehensive multi-level, multi-stakeholder approach focused on prevention of modifiable risk factors is urgently needed to curb worrisome trends in HD deaths in the US.

## References

1. CDC. Decline in deaths from heart disease and stroke--United States, 1900-1999. MMWR Morb Mortal Wkly Rep 1999;48(30):649-56. [published Online First: 1999/09/17]
2. Ma J, Ward EM, Siegel RL, et al. Temporal Trends in Mortality in the United States, 1969-2013. JAMA 2015;314(16):1731-9. doi: 10.1001/jama.2015.12319 [published Online First: 2015/10/28]
3. Ford ES, Ajani UA, Croft JB, et al. Explaining the decrease in U.S. deaths from coronary disease, 1980-2000. N Engl J Med 2007;356(23):2388-98. doi: 10.1056/NEJMsa053935 [published Online First: 2007/06/08]
4. Ritchey MD, Loustalot F, Bowman BA, et al. Trends in mortality rates by subtypes of heart disease in the United States, 2000-2010. JAMA 2014;312(19):2037-9. doi: 10.1001/jama.2014.11344 [published Online First: 2014/11/17]
5. Sidney S, Quesenberry CP, Jr., Jaffe MG, et al. Recent Trends in Cardiovascular Mortality in the United States and Public Health Goals. JAMA Cardiol 2016;1(5):594-9. doi: 10.1001/jamacardio.2016.1326 [published Online First: 2016/07/21]
6. Shah NS, Lloyd-Jones DM, O'Flaherty M, et al. Trends in Cardiometabolic Mortality in the United States, 1999-2017. JAMA 2019;322(8):780-2.
7. Curtin SC. Trends in cancer and heart disease death rates among adults aged 45-64: United States, 1999-2017. National Vital Statistics Reports 2019;68(5)
8. Woolf SH, Chapman DA, Buchanich JM, et al. Changes in midlife death rates across racial and ethnic groups in the United States: systematic analysis of vital statistics. BMJ 2018;362:k3096. doi: 10.1136/bmj.k3096 [published Online First: 2018/08/17]
9. Woolf SH, Schoomaker H. Life Expectancy and Mortality Rates in the United States, 1959-2017. JAMA 2019;322(20):1996-2016. doi: 10.1001/jama.2019.16932 [published Online First: 2019/11/27]
10. Sidney S, Quesenberry CP, Jr., Jaffe MG, et al. Heterogeneity in national U.S. mortality trends within heart disease subgroups, 2000-2015. BMC Cardiovasc Disord 2017;17(1):192. doi: 10.1186/s12872-017-0630-2 [published Online First: 2017/07/20]
11. Sidney S, Go AS, Jaffe MG, et al. Association Between Aging of the US Population and Heart Disease Mortality From 2011 to 2017. JAMA Cardiol 2019 doi: 10.1001/jamacardio.2019.4187 [published Online First: 2019/10/31]
12. Jose PO, Frank AT, Kapphahn KI, et al. Cardiovascular disease mortality in Asian Americans. J Am Coll Cardiol 2014;64(23):2486-94. doi: 10.1016/j.jacc.2014.08.048 [published Online First: 2014/12/17]
13. Iyer DG, Shah NS, Hastings KG, et al. Years of Potential Life Lost Because of Cardiovascular Disease in Asian-American Subgroups, 2003-2012. J Am Heart Assoc 2019;8(7):e010744. doi: 10.1161/JAHA.118.010744 [published Online First: 2019/03/21]
14. Wide-ranging Online Data for Epidemiologic Research (WONDER). https://wonder.cdc.gov/: Centers for Disease Control and Prevention, 2020.
15. Gardner JW, Sanborn JS. Years of potential life lost (YPLL)--what does it measure? Epidemiology 1990;1(4):322-9. [published Online First: 1990/07/01]
16. Arias E, Xu J, Kochanek KD. United States Life Tables, 2016. Natl Vital Stat Rep 2019;68(4):1-66. [published Online First: 2019/05/22]
17. Arias E, Xu J. United States Life Tables, 2017. Natl Vital Stat Rep 2019;68(7):1-65.
18. Joinpoint Regression Program, Version 4.7.0.0 [program]: National Cancer Institute, 2019.
19. Hales CM, Carroll MD, Fryar CD, et al. Prevalence of obesity and severe obesity among adults: United States, 2017-2018. NCHS Data Brief 2020;360
20. Virani SS, Alonso A, Benjamin EJ, et al. Heart Disease and Stroke Statistics-2020 Update: A Report From the American Heart Association. Circulation 2020;141(9):e139-e596. doi: 10.1161/CIR. 0000000000000757 [published Online First: 2020/01/30]
21. Fryar CD, Ostchega Y, Hales CM, et al. Hypertension prevalence and control among adults: United States, 2015-2016. NCHS Data Brief 2017(289)
22. Khan SS, Shah NS, Ning H, et al. Cardiovascular health behavior and health factor trends (19992014) and projections to 2050: Results from the National Health and Nutrition Examination Surveys. Circulation 2019;139(Supplement 1):AMP01.
23. Huffman MD, Capewell S, Ning H, et al. Cardiovascular health behavior and health factor changes (1988-2008) and projections to 2020: results from the National Health and Nutrition Examination Surveys. Circulation 2012;125(21):2595-602. doi: 10.1161/CIRCULATIONAHA.111.070722 [published Online First: 2012/05/02]
24. Hales CM, Fryar CD, Carroll MD, et al. Trends in Obesity and Severe Obesity Prevalence in US Youth and Adults by Sex and Age, 2007-2008 to 2015-2016. JAMA 2018;319(16):1723-25. doi: 10.1001/jama.2018.3060 [published Online First: 2018/03/24]
25. Lin J, Thompson TJ, Cheng YJ, et al. Projection of the future diabetes burden in the United States through 2060. Popul Health Metr 2018;16(1):9. doi: 10.1186/s12963-018-0166-4 [published Online First: 2018/06/16]
26. Dhingra R, Vasan RS. Diabetes and the risk of heart failure. Heart Fail Clin 2012;8(1):125-33. doi: 10.1016/j.hfc.2011.08.008 [published Online First: 2011/11/24]
27. Kenchaiah S, Evans JC, Levy D, et al. Obesity and the risk of heart failure. N Engl J Med 2002;347(5):305-13. doi: 10.1056/NEJMoa020245 [published Online First: 2002/08/02]
28. Uijl A, Koudstaal S, Vaartjes I, et al. Risk for Heart Failure: The Opportunity for Prevention With the American Heart Association's Life's Simple 7. JACC Heart Fail 2019;7(8):637-47. doi: 10.1016/j.jchf.2019.03.009 [published Online First: 2019/07/16]
29. Pool LR, Ning H, Lloyd-Jones DM, et al. Trends in Racial/Ethnic Disparities in Cardiovascular Health Among US Adults From 1999-2012. J Am Heart Assoc 2017;6(9) doi: 10.1161/JAHA.117.006027 [published Online First: 2017/09/25]
30. Ferdinand KC, Yadav K, Nasser SA, et al. Disparities in hypertension and cardiovascular disease in blacks: The critical role of medication adherence. J Clin Hypertens (Greenwich) 2017;19(10):1015-24. doi: 10.1111/jch. 13089 [published Online First: 2017/09/01]
31. Charles H, Good CB, Hanusa BH, et al. Racial differences in adherence to cardiac medications. J Natl Med Assoc 2003;95(1):17-27. [published Online First: 2003/03/27]
32. Zhang Y, Baik SH. Race/Ethnicity, disability, and medication adherence among medicare beneficiaries with heart failure. $J$ Gen Intern Med 2014;29(4):602-7. doi: 10.1007/s11606-013-2692-x [published Online First: 2013/12/25]
33. Ferdinand KC, Senatore FF, Clayton-Jeter H, et al. Improving Medication Adherence in Cardiometabolic Disease: Practical and Regulatory Implications. J Am Coll Cardiol 2017;69(4):437-51. doi: 10.1016/j.jacc.2016.11.034 [published Online First: 2017/01/28]
34. Mensah GA. Black and Minority Health 2019: More Progress is Needed. J Am Coll Cardiol 2019;74(9)
35. Havranek EP, Mujahid MS, Barr DA, et al. Social Determinants of Risk and Outcomes for Cardiovascular Disease: A Scientific Statement From the American Heart Association. Circulation 2015;132(9):873-98. doi: 10.1161/CIR.0000000000000228 [published Online First: 2015/08/05]
36. Snyder ML, Love SA, Sorlie PD, et al. Redistribution of heart failure as the cause of death: the Atherosclerosis Risk in Communities Study. Popul Health Metr 2014;12(1):10. doi: 10.1186/1478-7954-12-10 [published Online First: 2014/04/11]
37. Kontis V, Cobb LK, Mathers CD, et al. Three Public Health Interventions Could Save 94 Million Lives in 25 Years. Circulation 2019;140(9):715-25. doi: 10.1161/CIRCULATIONAHA. 118.038160 [published Online First: 2019/06/11]

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## Transparency Statement:

The lead author Dr. Nilay Shah and senior author Dr. Sadiya Khan affirm that the manuscript is an honest, accurate, and transparent account of the study being reported; that no important aspects of the study have been omitted; and that any discrepancies from the study as originally planned have been explained.

## Figure Titles and Legends

Figure 1. Proportion of total heart disease deaths attributable to each heart disease subtype category in the US, 1999-2018. Proportion of total heart disease deaths due to each individual subtype or cateogry. Proportion of HD deaths due to IHD decreased, while proportion of HD deaths due to all other causes increased, predominantly from HF and HTN HD, which were the second and third leading cause of HD death. HF: heart failure, HTN HD: hypertensive heart disease, IHD: ischemic heart disease, Other HD: other heart disease, Pulmonary HD: pulmonary heart disease, Valvular HD: valvular heart disease.

Figure 2. Trends in age-adjusted mortality rate due to leading heart disease subtypes in black and white women and men in the US, 1999-2018. Age-adjusted mortality rates shown for (A) total heart disease, (B) ischemic heart disease, (C) heart failure, and (D) hypertensive heart disease in black and white women and men. Dashed line represents temporal inflection point identified by Joinpoint regression. Average annual percentage change of AAMR before and after 2011 are listed in Supplemental Table 1. AAMR from total HD and IHD plateaued after 2011. AAMR from HF increased after 2011, particularly in black men. Absolute AAMR from HTN HD was highest in black men and women from 1999-2018, but increases in AAMR during this time was fastest in white men and women.

## TABLES

Table 1. Trends in age-adjusted mortality rate attributed to each heart disease subtype in the total US population, 1999-2018

|  | Age-adjusted mortality rate <br> (per 100,000) |  |  | Mean Annual Percent Change <br> $(\mathbf{9 5 \%}$ CI) |  |
| :--- | :---: | :---: | :---: | :---: | :---: |
|  | $\mathbf{1 9 9 9}$ <br> $(\mathrm{N}=725,192)$ | $\mathbf{2 0 1 1}$ <br> $(\mathrm{N}=596,577)$ | $\mathbf{2 0 1 8}$ <br> $(\mathrm{N}=655,381)$ | $\mathbf{1 9 9 9 - 2 0 1 1}$ | $\mathbf{2 0 1 1 - 2 0 1 8}$ |
| Total Heart <br> Disease | 266.5 | 173.7 | 163.6 | $-3.7(-3.8,-3.5)^{*}$ | $-0.7(-1.1,-0.3)^{*}$ |
| Ischemic Heart <br> Disease | 194.6 | 109.2 | 90.9 | $-4.7(-5.1,-4.2)^{*}$ | $-2.6(-3.3,-2.0)^{*}$ |
| Heart Failure | 20.3 | 16.9 | 20.8 | $-1.7(-2.5,-0.8)^{*}$ | $3.5(2.6,4.5)^{*}$ |
| Hypertensive <br> Heart Disease | 9.6 | 10.8 | 15.1 | $1.2(0.7,1.7)^{*}$ | $4.8(3.3,6.3)^{*}$ |
| Valvular Heart <br> Disease | 7.2 | 6.7 | 6.1 | $-0.4(-0.6,-0.1)^{*}$ | $-1.6(-2.8,-0.4)^{*}$ |
| Arrhythmia | 6.7 | 8.2 | 9.1 | $1.7(1.4,2.0)^{*}$ | $1.7(1.4,2.0)^{*}$ |
| Pulmonary Heart <br> Disease | 4.8 | 4.2 | 4.6 | $-0.9(-2.1,0.2)$ | $1.5(1.0,1.9)^{*}$ |
| Other Heart <br> Disease | 23.2 | 17.8 | 17.1 | $-2.0(-2.3,-1.6)^{*}$ | $-0.7(-1.0,-0.4)^{*}$ |

CI: Confidence interval.
*Indicates that the mean annual percent change is significantly different from zero, $\mathrm{p}<0.05$. Mean annual percentage change shown for AAMR for the specified time range.

Table 2. Trends in age-adjusted mortality rate attributed to each heart disease subtype by age at death in the total US population, 1999-2018

| Heart disease subtype | Age at Death | Age-adjusted mortality rates <br> per 100,000 |  |  | Mean Annual Percent Change (95\% CI) |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  |  | 1999 | 2011 | 2018 | 1999-2011 | 2011-2018 |
| Total Heart Disease | <45 | 10.3 | 9.0 | 8.7 | -1.0 (-1.5, -0.5)* | -0.3 (-0.7, 0.1) |
|  | 45-64 | 164.1 | 121.0 | 122.3 | -2.5 (-2.8, -2.2)* | 0.2 (-0.1, 0.4) |
|  | $\geq 65$ | 1767.0 | 1115.6 | 1034.6 | -3.7 (-4.3, -3.1)* | -1.1 (-1.6, -0.6)* |
| Ischemic <br> Heart <br> Disease | <45 | 4.9 | 3.8 | 3.2 | -1.9 (-2.6, -1.3)* | -2.3 (-2.7, -1.9)* |
|  | 45-64 | 121.1 | 79.3 | 73.1 | -3.4 (-3.6, -3.3)* | -1.4 (-1.6, -1.2)* |
|  | $\geq 65$ | 1301.6 | 704.7 | 573.8 | -4.9 (-5.4, -4.4)* | -3.0 (-3.4, -2.3)* |
| Heart Failure | <45 | 0.2 | 0.3 | 0.4 | 2.8 (0.5, 5.0)* | 5.1 (2.7, 7.6)* |
|  | 45-64 | 4.8 | 4.5 | 6.8 | -0.6 (-1.2, -0.1)* | 5.7 (4.4, 7.0)* |
|  | $\geq 65$ | 150.8 | 124.5 | 150.4 | -1.7 (-2.6, -0.9)* | 3.3 (2.4, 4.3)* |
| Hypertensive Heart Disease | <45 | 0.7 | 1.2 | 1.4 | 4.8 (3.7, 5.9)* | 2.3 (1.9, 2.8)* |
|  | 45-64 | 8.1 | 12.1 | 15.2 | 3.9 (2.1, 5.6)* | 3.0 (2.6, 3.5)* |
|  | $\geq 65$ | 57.6 | 58.0 | 85.5 | 0.3 (-0.2, 0.8) | 5.4 (3.6, 7.2)* |
| Valvular <br> Heart <br> Disease | <45 | 0.3 | 0.2 | 0.3 | -3.9 (-6.0, -1.8)* | 7.1 (1.4, 13.2)* |
|  | 45-64 | 2.5 | 2.1 | 2.2 | -1.6 (-3.2, 0.1) | 1.0 (-2.0, 4.0) |
|  | $\geq 65$ | 51.0 | 48.1 | 42.7 | -0.3 (-0.5, 0.0)* | -1.9 (-3.2, -0.6)* |
| Arrhythmia | <45 | 0.4 | 0.4 | 0.4 | -0.3 (-1.0, 0.4) | -0.3 (-1.0, 0.4) |
|  | 45-64 | 3.2 | 3.4 | 3.9 | 0.5 (-0.9, 2.0) | 2.2 (1.0, 3.4)* |
|  | $\geq 65$ | 45.7 | 56.6 | 63.1 | 1.8 (1.5, 2.1)* | 1.8 (1.5, 2.1)* |
| Pulmonary <br> Heart <br> Disease | $<45$ | 0.9 | 0.7 | 0.7 | -1.8 (-3.0, -0.6)* | 0.3 (-0.9, 1.5) |
|  | 45-64 | 4.6 | 4.0 | 4.3 | $-1.8(-2.5,-1.0)^{*}$ | 1.4 (0.4, 2.4)* |
|  | $\geq 65$ | 25.7 | 23.0 | 25.3 | -0.7 (-2.1, 0.7) | 1.7 (1.1, 2.2)* |
| Other Heart Disease | <45 | 2.8 | 2.4 | 2.4 | -1.3 (-1.7, -1.0)* | -0.1 (-1.0, 0.8) |
|  | 45-64 | 19.8 | 15.6 | 16.6 | -1.9 (-2.6, -1.2)* | $0.9(0.5,1.4)^{*}$ |
|  | $\geq 65$ | 134.5 | 100.7 | 93.8 | -2.2 (-2.6, -1.7)* | $-1.0(-1.3,-0.6)^{*}$ |

CI: Confidence interval. *Indicates that the mean annual percent change is significantly different from zero, $\mathrm{p}<0.05$. Mean annual percentage change shown for AAMR for the specified time range.

Table 3. Trends in age-standardized years of potential life lost per 100,000 population due to each HD category in the total population and race-sex subgroups in the US, 1999-2018

| Heart disease subtype | Race-Sex Group | Years of Potential Life Lost (per 100,000 population) |  |  | Mean Annual Percent Change (95\% CI) |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  |  | 1999 | 2011 | 2018 | 1999-2011 | 2011-2018 |
| Total Heart Disease | Total | 1494 | 1080 | 1068 | -2.7 (-2.9, -2.6)* | $0.1(-0.2,0.4)$ |
|  | Black women | 1824 | 1215 | 1210 | -3.5 (-3.8, -3.3)* | -0.1 (-0.7, 0.6) |
|  | White women | 1064 | 754 | 744 | -2.8 (-3.0, -2.7)* | 0.1 (-0.3, 0.4) |
|  | Black men | 2183 | 1549 | 1602 | -2.7 (-3.3, -2.2)* | 0.3 (-0.3, 0.9) |
|  | White men | 1677 | 1244 | 1205 | -2.5 (-2.6, -2.3)* | -0.5 (-0.7, -0.2)* |
| Ischemic Heart Disease | Total | 1025 | 645 | 578 | -3.8 (-3.9, -3.7)* | -1.6 (-1.8, -1.4)* |
|  | Black women | 983 | 543 | 479 | -5.0 (-5.5, -4.4)* | -1.6 (-2.0, -1.1)* |
|  | White women | 696 | 418 | 373 | -4.1 (-4.3, -3.9)* | -1.5 (-1.9, -1.1)* |
|  | Black men | 1254 | 770 | 716 | -3.8 (-4.3, -3.3)* | -1.1 (-1.7, -0.5)* |
|  | White men | 1247 | 816 | 723 | -3.5 (-3.7, -3.4)* | -1.7 (-2.0, -1.5)* |
| Heart Failure | Total | 49 | 45 | 64 | -1.0 (-1.7, -0.3)* | 5.5 (4.8, 6.1)* |
|  | Black women | 79 | 70 | 103 | -1.2 (-2.4, 0.1) | 5.8 (4.5, 7.2)* |
|  | White women | 50 | 43 | 53 | -1.4 (-2.9, 0.1) | 3.9 (2.6, 5.2)* |
|  | Black men | 71 | 74 | 116 | $0.8(-0.8,2.5)$ | 6.6 (5.0, 8.3)* |
|  | White men | 43 | 39 | 57 | $-0.9(-1.4,-0.5)^{*}$ | 5.5 (4.5, 6.6)* |
| Hypertensive Heart Disease | Total | 73 | 107 | 132 | 3.4 (2.6, 4.2)* | 3.2 (2.4, 4.1)* |
|  | Black women | 210 | 207 | 219 | 0.0 (-0.4, 0.3) | 0.0 (-0.4, 0.3) |
|  | White women | 39 | 57 | 76 | 3.4 (2.5, 4.3)* | 4.2 (0.9, 7.7)* |
|  | Black men | 237 | 271 | 305 | 1.6 (0.7, 2.5)* | 0.6 (0.2, 0.9)* |
|  | White men | 57 | 105 | 134 | 5.4 (4.7, 6.0)* | 3.3 (3.0, 3.5)* |
| All Other Heart Disease | Total | 346 | 283 | 293 | -1.6 (-2.0, -1.3)* | 0.5 (-0.7, 1.7) |
|  | Black women | 551 | 394 | 409 | -2.7 (-3.1, -2.4)* | 0.4 (-0.4, 1.2) |
|  | White women | 279 | 236 | 241 | -1.5 (-1.8, -1.1)* | 0.8 (0.2, 1.3)* |
|  | Black men | 620 | 435 | 465 | -2.7 (-3.9, -1.5)* | $0.7(0.1,1.4)^{*}$ |
|  | White men | 331 | 284 | 291 | -1.1 (-1.4, -0.8)* | 0.3 (0.0, 0.7) |

CI: Confidence interval. Mean annual percentage change shown for YPLL for the specified time range.
*Indicates that the mean annual percent change is significantly different from zero, $\mathrm{p}<0.05$.

## FIGURES

Figure 1. Proportion of total heart disease deaths attributable to each heart disease subtype category in the US, 1999-2018


Figure 2. Trends in age-adjusted mortality rate due to leading heart disease subtypes in black and white women and men in the US, 1999-2018.

B. Ischemic heart disease


D. Hypertensive heart disease


Heterogeneous Trends in Burden of Heart Disease Mortality by Subtypes in the United States, 1999-2018: Observational Analysis of Vital Statistics

Supplementary Material

Supplemental Table 1. Cause-specific age-adjusted heart disease mortality rate trends in race-sex subgroups of all ages, 1999-2018

| Heart disease subtype | Race-Sex Group | AAMR per 100,000 |  |  | Mean AnnualPercent Change (95\% CI) |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  |  | 1999 | 2011 | 2018 | 1999-2011 | 2011-2018 |
| Total Heart Disease | Black women | 283.7 | 176.2 | 162.2 | -3.9 (-4.3, -3.5)* | -1.2 (-1.6, -0.8)* |
|  | White women | 212.8 | 136.5 | 126.6 | -3.8 (-4.1, -3.6)* | -0.9 (-1.3, -0.4)* |
|  | Black men | 407.2 | 266.1 | 259.5 | -3.4 (-3.6, -3.2)* | -0.5 (-1.0, 0.0) |
|  | White men | 327.1 | 216.9 | 206.3 | -3.4 (-3.9, -2.9)* | $-0.6(-1.0,-0.3)^{*}$ |
| Ischemic Heart Disease | Black women | 188.0 | 99.7 | 77.9 | -5.3 (-5.6, -4.9)* | -3.3 (-3.7, -3.0)* |
|  | White women | 150.5 | 80.1 | 63.7 | $-5.1(-5.6,-4.6)^{*}$ | -3.3 (-3.7, -2.9)* |
|  | Black men | 278.7 | 161.5 | 138.1 | -4.4 (-4.7, -4.1)* | $-2.3(-2.9,-1.7)^{*}$ |
|  | White men | 251.2 | 146.5 | 125.6 | -4.4 (-4.8, -3.9)* | -2.2 (-2.6, -1.9)* |
| Heart Failure | Black women | 20.9 | 17.0 | 21.5 | -1.6 (-2.5, -0.7)* | 3.5 (2.8, 4.2)* |
|  | White women | 19.3 | 15.8 | 18.7 | $-1.7(-2.5,-0.9)^{*}$ | 2.6 (2.1, 3.2)* |
|  | Black men | 25.3 | 20.7 | 28.9 | -1.3 (-2.2, -0.4)* | 4.9 (4.0, 5.8)* |
|  | White men | 21.4 | 18.8 | 23.8 | -1.3 (-1.7, -0.9)* | 3.8 (2.8, 4.8)* |
| Hypertensive Heart Disease | Black women | 24.4 | 20.2 | 22.3 | -1.4 (-1.8, -1.0)* | $0.7(-1.0,2.3)$ |
|  | White women | 7.4 | 8.1 | 11.8 | 0.9 (0.5, 1.3)* | 5.6 (4.3, 7.0)* |
|  | Black men | 30.2 | 29.7 | 35.3 | $0.3(-0.5,1.1)$ | 1.8 (0.9, 2.6)* |
|  | White men | 7.8 | 10.4 | 15.7 | 2.5 (2.0, 3.0)* | 6.3 (4.9, 9.4)* |
| Valvular <br> Heart Disease | Black women | 5.1 | 4.0 | 3.8 | -1.8 (-2.1, -1.5)* | -1.8 (-2.1, -1.5)* |
|  | White women | 7.1 | 6.7 | 6.1 | -0.4 (-0.6, -0.2)* | $-1.5(-2.5,-0.4)^{*}$ |
|  | Black men | 5.4 | 4.2 | 4.2 | -1.5 (-2.0, -0.9)* | -1.5 (-2.0, -0.9)* |
|  | White men | 7.8 | 7.6 | 6.9 | $0.1(-0.2,0.4)$ | -1.6 (-3.1, 0.0) |
| Arrhythmia | Black women | 6.5 | 6.5 | 7.0 | 0.4 (-1.4, 2.3) | 0.9 (-0.1, 1.9) |
|  | White women | 6.2 | 8.1 | 8.8 | 2.0 (1.7, 2.3)* | 2.0 (1.7, 2.3)* |
|  | Black men | 8.9 | 8.2 | 9.3 | $0.1(-0.3,0.5)$ | $0.1(-0.3,0.5)$ |
|  | White men | 7.4 | 8.5 | 10.1 | 1.0 (0.5, 1.6)* | 2.7 (2.3, 3.2)* |
| Pulmonary <br> Heart Disease | Black women | 8.9 | 7.0 | 8.3 | -1.5 (-2.2, -0.8)* | 1.1 (0.4, 1.8)* |
|  | White women | 4.6 | 4.3 | 4.6 | $-0.7(-1.5,-0.2)^{*}$ | 1.4 (1.1, 1.8)* |
|  | Black men | 8.4 | 6.4 | 7.3 | -2.1 (-3.0, -1.2)* | 1.3 (0.2, 2.4)* |
|  | White men | 4.5 | 3.7 | 4.1 | -1.8 (-2.5, -1.0)* | 2.1 (1.1, 3.0)* |
| Other Heart Disease | Black women | 29.9 | 21.7 | 21.3 | -2.4 (-2.7, -2.1)* | -0.7 (-1.5, 0.1) |
|  | White women | 17.6 | 13.4 | 12.8 | -2.2 (-2.7, -1.7)* | -0.6 (-2.6, 1.6) |
|  | Black men | 50.2 | 35.4 | 36.4 | $-2.5(-2.9,-2.2)^{*}$ | $0.1(-0.8,0.9)$ |
|  | White men | 27.1 | 21.4 | 20.2 | -1.8 (-2.2, -1.4)* | -0.7 (-1.0, -0.4)* |

CI: Confidence interval. AAMR: Age-adjusted mortality rate per 100,000 population. Mean annual percentage change shown for AAMR for the specified time range. *Indicates that the mean annual percent change is significantly different from zero, $\mathrm{p}<0.05$.

Supplemental Table 2. Cause-specific age-adjusted heart disease mortality rate trends in race-sex subgroups, in decedents $<45$ years of age, 1999-2018

| Heart disease subtype |  | AAMR per 100,000 |  |  | Mean Annual Percent Change (95\% CI) |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  |  | 1999 | 2011 | 2018 | 1999-2011 | 2011-2018 |
| Total Heart Disease | Black women | 15.6 | 12.2 | 12.1 | -2.2 (-2.7, -1.8)* | 0.0 (-1.2, 1.3) |
|  | White women | 5.2 | 4.9 | 4.9 | -0.7 (-1.2, -0.3)* | 0.5 (-0.3, 1.3) |
|  | Black men | 27.4 | 22.6 | 23.4 | -1.0 (-1.2, -0.7)* | -1.0 (-1.2, -0.7)* |
|  | White men | 12.5 | 11.1 | 10.2 | -1.0 (-1.5, -0.4)* | -0.9 (-1.6, -0.1)* |
| Ischemic Heart Disease | Black women | 4.9 | 3.2 | 3.0 | $-2.8(-3.3,-2.3)^{*}$ | -2.8 (-3.3, -2.3)* |
|  | White women | 2.0 | 1.7 | 1.5 | -0.8(-2.1, 0.4) | -2.2 (-4.6, 0.2) |
|  | Black men | 11.3 | 8.4 | 7.8 | -2.0 (-2.3, -1.7)* | -2.0 (-2.3, -1.7)* |
|  | White men | 7.2 | 5.6 | 4.4 | -2.1 (-2.6, -1.5)* | -3.2 (-3.4, -3.0)* |
| Heart Failure | Black women | 0.5 | 0.7 | 0.8 | $2.1(-5.0,9.6)$ | $1.7(-1.9,5.5)$ |
|  | White women | 0.1 | 0.1 | 0.2 | $0.2(-2.8,3.2)$ | $0.2(-2.8,3.2)$ |
|  | Black men | 0.7 | 1.0 | 1.7 | 2.8 (0.0, 5.6)* | 8.8 (5.7, 12.0)* |
|  | White men | 0.2 | 0.2 | 0.4 | 2.8 (1.4, 4.2)* | 2.8 (1.4, 4.2)* |
| Hypertensive Heart Disease | Black women | 2.2 | 2.6 | 2.6 | 0.6 (0.1, 1.1)* | 0.6 (0.1, 1.1)* |
|  | White women | 0.2 | 0.4 | 0.6 | 4.7 (3.6, 5.8)* | 4.7 (3.6, 5.8)* |
|  | Black men | 3.6 | 4.6 | 5.1 | 1.8 (1.3, 2.3)* | 1.8 (1.3, 2.3)* |
|  | White men | 0.7 | 1.2 | 1.5 | 5.2 (3.7, 6.7)* | 2.7 (2.0, 3.3)* |
| Valvular <br> Heart Disease | Black women | 0.5 | 0.4 | 0.3 | -4.5 (-6.1, -2.8)* | -4.5 (-6.1, -2.8)* |
|  | White women | 0.3 | 0.2 | 0.2 | -0.2 (-1.9, 1.5) | -0.2 (-1.9, 1.5) |
|  | Black men | 0.6 | 0.3 | 0.5 | -4.1 (-5.6, -2.5)* | 3.6 (-4.9, 12.8) |
|  | White men | 0.3 | 0.3 | 0.3 | -0.1 (-0.1, -0.1)* | 0.8 (0.8, 0.8)* |
| Arrhythmia | Black women | 0.6 | 0.4 | 0.4 | -1.6 (-2.8, -0.4)* | -1.6 (-2.8, -0.4)* |
|  | White women | 0.3 | 0.3 | 0.3 | -0.4 (-1.6, 0.8) | $-0.4(-1.6,0.8)$ |
|  | Black men | 0.8 | 0.9 | 0.8 | $0.2(-0.6,1.1)$ | $0.2(-0.6,1.1)$ |
|  | White men | 0.4 | 0.4 | 0.4 | -0.5 (-4.6, 3.9) | $0.1(-1.1,1.3)$ |
| Pulmonary Heart Disease | Black women | 2.2 | 1.7 | 1.8 | -1.0 (-1.6, -0.3)* | -1.0 (-1.6, -0.3)* |
|  | White women | 0.8 | 0.7 | 0.7 | -2.1 (-3.6, -0.7)* | $0.5(-1.2,2.3)$ |
|  | Black men | 1.5 | 1.3 | 1.4 | -0.8(-1.8, 0.2) | -0.8 (-1.8, 0.2) |
|  | White men | 0.7 | 0.5 | 0.6 | $-1.7(-2.7,-0.8) *$ | 3.2 (1.3, 5.1)* |
| Other Heart Disease | Black women | 4.7 | 3.3 | 3.2 | -3.1 (-3.7, -2.5)* | -0.7 (-2.2, 0.9) |
|  | White women | 1.5 | 1.4 | 1.4 | $0.1(-0.5,0.3)$ | $0.1(-0.5,0.3)$ |
|  | Black men | 8.8 | 6.0 | 6.1 | $-2.6(-5.1,0.0)^{*}$ | $-0.4(-0.4,1.2)$ |
|  | White men | 3.0 | 2.9 | 2.6 | -0.8 (-1.0, -0.5)* | -0.8 (-1.0, -0.5)* |

CI: Confidence interval. AAMR: Age-adjusted mortality rate per 100,000 population. Mean annual percentage change shown for AAMR for the specified time range. Total includes all decedents of all race/ethnicity groups. *Indicates that the mean annual percent change is significantly different from zero, $\mathrm{p}<0.05$.

Supplemental Table 3. Cause-specific age-adjusted heart disease mortality rate trends in race-sex subgroups, in decedents age 45-64 years, 1999-2018

| Heart disease subtype |  | AAMR per 100,000 |  |  | Mean Annual Percent Change (95\% CI) |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  |  | 1999 | 2011 | 2018 | 1999-2011 | 2011-2018 |
| Total Heart Disease | Black women | 209.1 | 137.8 | 140.9 | -3.4 (-4.2, -2.6)* | 0.3 (-0.4, 1.0) |
|  | White women | 83.7 | 63.1 | 64.5 | -2.2 (-2.5, -2.0)* | 0.5 (0.1, 0.9)* |
|  | Black men | 414.1 | 278.8 | 288.1 | -3.3 (-3.6, -3.1)* | 0.3 (-0.3, 0.9) |
|  | White men | 222.2 | 166.2 | 165.3 | -2.4 (-2.7, -2.1)* | -0.1 (-0.4, 0.2) |
| Ischemic Heart Disease | Black women | 124.6 | 70.3 | 63.9 | -4.9 (-5.6, -4.1)* | -1.2 (-1.8, -0.6)* |
|  | White women | 57.7 | 37.9 | 35.6 | -3.3 (-3.5, -3.0)* | -0.8 (-1.2, -0.3)* |
|  | Black men | 273.2 | 160.4 | 149.7 | -4.2 (-4.6, -3.8)* | -1.1 (-1.6, -0.5)* |
|  | White men | 176.0 | 118.3 | 108.3 | -3.2 (-3.4, -3.0)* | -1.4 (-1.7, -1.2)* |
| Heart Failure | Black women | 9.0 | 7.9 | 12.3 | -1.2 (-3.0, 0.6) | 7.0 (5.1, 8.9)* |
|  | White women | 3.2 | 3.0 | 4.0 | -1.5 (-2.2, -0.7)* | 4.5 (2.7, 6.4)* |
|  | Black men | 14.1 | 13.4 | 21.0 | $0.4(-1.1,2.0)$ | 6.5 (4.9, 8.2)* |
|  | White men | 5.1 | 4.8 | 7.3 | -0.6 (-1.3, 0.1) | 5.8 (4.2, 7.5)* |
| Hypertensive Heart Disease | Black women | 24.7 | 23.5 | 26.1 | 0.0 (-1.4, 0.5) | 0.0 (-1.4, 0.5) |
|  | White women | 3.5 | 5.5 | 7.2 | 4.1 (3.7, 4.5)* | 4.1 (3.7, 4.5)* |
|  | Black men | 41.0 | 45.8 | 51.4 | $1.3(-0.4,2.9)$ | 1.0 (0.4, 1.6)* |
|  | White men | 7.3 | 13.6 | 17.6 | 5.6 (4.9, 6.3)* | 3.6 (3.3, 3.9)* |
| Valvular Heart Disease | Black women | 4.1 | 2.7 | 1.7 | -2.4 (-3.1, -1.6)* | -2.4 (-3.1, -1.6)* |
|  | White women | 1.9 | 1.5 | 1.6 | $-1.6(-2.2,-1.1)^{*}$ | $0.7(-0.6,1.9)$ |
|  | Black men | 5.2 | 3.7 | 4.0 | -2.4 (-3.2, -1.6)* | $-2.4(-3.2,-1.6) *$ |
|  | White men | 2.7 | 2.7 | 2.7 | $0.1(-0.3,0.6)$ | $0.1(-0.3,0.6)$ |
| Arrhythmia | Black women | 4.5 | 4.1 | 4.6 | -0.1 (-0.7, 0.6) | -0.1 (-0.7, 0.6) |
|  | White women | 2.1 | 2.5 | 2.5 | 1.2 (0.7, 1.7)* | 1.2 (0.7, 1.7)* |
|  | Black men | 7.9 | 6.7 | 8.4 | $-1.8(-2.9,-0.7)^{*}$ | 3.2 (1.5, 5.0)* |
|  | White men | 3.9 | 4.1 | 5.1 | $0.5(-0.3,1.3)$ | 3.4 (2.6, 4.1)* |
| Pulmonary <br> Heart Disease | Black women | 11.3 | 8.4 | 8.7 | -2.0 (-3.0, -0.9)* | 0.8 (-0.3, 1.8) |
|  | White women | 4.2 | 3.6 | 3.7 | -2.1 (-2.8, -1.4)* | $1.6(-0.3,3.6)$ |
|  | Black men | 9.9 | 8.1 | 8.3 | -2.1 (-3.1, -1.2)* | $1.9(0.7,3.1)^{*}$ |
|  | White men | 4.4 | 3.8 | 4.1 | -1.4 (-2.2, -0.6)* | 1.9 (1.1, 2.7)* |
| Other Heart Disease | Black women | 31.2 | 21.1 | 22.7 | -2.7 (-3.2, -2.1)* | 0.5 (-0.9, 1.9) |
|  | White women | 11.2 | 9.2 | 9.6 | $-1.3(-1.8,-0.8)^{*}$ | 1.0 (0.5, 1.5)* |
|  | Black men | 63.5 | 41.2 | 45.2 | $-3.0(-3.4,-2.6) *$ | 1.0 (0.0, 2.0)* |
|  | White men | 23.1 | 19.1 | 20.1 | -1.3 (-1.7, -0.9)* | $0.4(0.0,0.7)^{*}$ |

CI: Confidence interval. AAMR: Age-adjusted mortality rate per 100,000 population. Mean annual percentage change shown for AAMR for the specified time range. *Indicates that the mean annual percent change is significantly different from zero, $\mathrm{p}<0.05$.

Supplemental Table 4. Cause-specific age-adjusted heart disease mortality rate trends in race-sex subgroups, in decedents $\geq 65$ years of age, 1999-2018

| HD subtype |  | AAMR per 100,000 |  |  | Mean Annual Percent Change (95\% CI) |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  |  | 1999 | 2011 | 2018 | 1999-2011 | 2011-2018 |
| Total Heart Disease | Black women | 1797.1 | 1089.0 | 973.7 | -4.1 (-4.5, -3.7)* | -1.6 (-2.0, -1.2)* |
|  | White women | 1509.8 | 943.7 | 863.1 | -4.1 (-4.3, -3.8)* | -1.1 (-1.7, -0.6)* |
|  | Black men | 2353.2 | 1499.5 | 1426.5 | -3.6 (-3.9, -3.4)* | -0.9 (-1.2, -0.6)* |
|  | White men | 2133.2 | 1366.5 | 1289.3 | -4.2 (-4.2, -3.1)* | -0.8 (-1.1, -0.4)* |
| Ischemic Heart Disease | Black women | 1243.0 | 649.2 | 489.0 | -5.2 (-5.6, -4.9)* | -4.0 (-4.5, -3.5)* |
|  | White women | 1079.3 | 558.0 | 434.1 | -5.5 (-5.9, -5.0)* | -3.4 (-3.9, -2.9)* |
|  | Black men | 1667.2 | 952.4 | 789.1 | -4.7 (-5.1, -4.2)* | -2.7 (-3.2, -2.3)* |
|  | White men | 1641.3 | 922.5 | 780.7 | -4.7 (-5.2, -4.2)* | -2.4 (-2.8, -2.0)* |
| Heart Failure | Black women | 146.6 | 117.3 | 144.3 | -1.8 (-2.7, -0.8)* | 3.2 (2.4, 4.0)* |
|  | White women | 146.3 | 118.7 | 140.0 | -1.8 (-2.6, -0.9)* | 2.6 (2.0, 3.1)* |
|  | Black men | 171.7 | 135.2 | 183.1 | $-1.7(-2.2,-1.3) *$ | 4.1 (3.1, 5.2)* |
|  | White men | 159.0 | 139.1 | 173.6 | -1.3 (-1.8, -0.9)* | 3.6 (2.6, 4.7)* |
| Hypertensive Heart Disease | Black women | 138.5 | 105.7 | 117.6 | -2.0 (-2.4, -1.6)* | 0.6 (-1.0, 2.3) |
|  | White women | 51.6 | 52.2 | 77.8 | $0.3(0.0,0.6)$ | 5.8 (4.3, 7.3)* |
|  | Black men | 148.5 | 130.4 | 162.9 | -0.6 (-2.4, 1.2) | 2.5 (1.4, 3.6)* |
|  | White men | 45.3 | 51.8 | 85.1 | $1.2(0.6,1.9)^{*}$ | $7.4(5.7,9.1)^{*}$ |
| Valvular <br> Heart <br> Disease | Black women | 30.7 | 24.8 | 24.3 | -1.5 (-1.9, -1.2)* | -1.5 (-1.9, -1.2)* |
|  | White women | 51.5 | 49.2 | 44.1 | -0.3 (-0.5, 0.1)* | $-1.7(-2.9,-0.5)^{*}$ |
|  | Black men | 30.6 | 25.0 | 23.7 | -1.1 (-1.7, -0.4)* | -1.1 (-1.7, -0.4)* |
|  | White men | 55.3 | 54.1 | 47.8 | $0.1(-0.2,0.5)$ | -1.9 (-3.6, -0.1)* |
| Arrhythmia | Black women | 40.1 | 42.2 | 45.5 | 0.7 (-1.3, 2.6) | 1.0 (-0.1, 2.0) |
|  | White women | 44.0 | 58.7 | 64.1 | 2.1 (1.8, 2.4)* | 2.1 (1.8, 2.4)* |
|  | Black men | 52.5 | 48.5 | 54.4 | $0.1(-0.4,0.6)$ | $0.1(-0.4,0.6)$ |
|  | White men | 49.2 | 57.8 | 68.9 | 1.2 (0.6, 1.8)* | $2.8(2.3,3.3) *$ |
| Pulmonary <br> Heart <br> Disease | Black women | 42.4 | 33.5 | 40.9 | -1.3 (-2.1, -0.5)* | $2.2(0.7,3.8)^{*}$ |
|  | White women | 25.9 | 25.3 | 26.2 | 0.0 (-1.3, 1.3) | 1.5 (1.2, 1.8)* |
|  | Black men | 45.1 | 32.6 | 35.9 | -2.1 (-3.4, -0.9)* | $0.9(-0.6,2.5)$ |
|  | White men | 26.6 | 21.7 | 22.3 | -1.8 (-2.7, -0.9)* | $2.1(1.0,3.2)^{*}$ |
| Other Heart Disease | Black women | 157.7 | 117.3 | 112.1 | -2.3 (-2.6, -1.9)* | -1.1 (-1.9, -0.3)* |
|  | White women | 112.3 | 82.6 | 76.9 | -2.5 (-3.0, -1.9)* | -1.0 (-3.2, 1.1) |
|  | Black men | 240.5 | 176.8 | 177.3 | -2.4 (-2.8, -1.9)* | -0.1 (-1.2, 1.0) |
|  | White men | 158.5 | 120.9 | 111.0 | -2.1 (-2.6, -1.6)* | -1.0 (-1.4, -0.6)* |

CI: Confidence interval. AAMR: Age-adjusted mortality rate per 100,000 population. Mean annual percentage change shown for AAMR for the specified time range. *Indicates that the mean annual percent change is significantly different from zero, $\mathrm{p}<0.05$.

Supplemental Table 5. Heart failure-related cardiovascular mortality trends by race, sex, and age, 1999-2018

|  | Race-Sex Group | AAMR or YPLL per 100,000 |  |  | Mean Annual Percent Change (95\% CI) |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  |  | 1999 | 2011 | 2018 | 1999-2011 | 2011-2018 |
| AAMR <br> (All Ages) | Total | 74.6 | 53.3 | 58.1 | -2.8 (-3.5, -2.1)* | 1.3 (0.6, 1.9)* |
|  | Black women | 67.6 | 48.5 | 53.3 | -2.9 (-3.7, -2.2)* | 1.7 (0.8, 2.6)* |
|  | White women | 67.5 | 47.3 | 50.3 | -3.0 (-4.0, -2.1)* | 0.9 (0.1, 1.7)* |
|  | Black men | 82.1 | 61.3 | 73.6 | $-2.4(-2.8,-1.9)^{*}$ | 2.7 (2.2, 3.2)* |
|  | White men | 86.3 | 63.2 | 70.4 | -2.5 (-3.2, -1.9)* | 1.5 (1.2, 1.9)* |
| $\begin{aligned} & \text { AAMR } \\ & (<45) \end{aligned}$ | Total | 0.8 | 0.9 | 1.2 | 0.0 (-1.4, 1.4) | 4.9 (3.1, 6.7)* |
|  | Black women | 1.9 | 1.7 | 2 | -0.6 (-1.2, 0.0)* | 2.0 (0.9, 3.2)* |
|  | White women | 0.5 | 0.5 | 0.6 | $-1.5(-2.9,-0.2)^{*}$ | 4.2 (1.5, 6.9)* |
|  | Black men | 2.6 | 3.1 | 4.3 | $1.1(-1.4,3.7)$ | 4.8 (2.8, 6.9)* |
|  | White men | 0.8 | 0.8 | 1.1 | -0.1 (-1.9, 1.8) | 4.5 (2.8, 6.3)* |
| $\begin{aligned} & \text { AAMR } \\ & (45-64) \end{aligned}$ | Total | 20.1 | 15.1 | 20.2 | -2.5 (-2.8, -2.3)* | 4.3 (3.7, 5.0)* |
|  | Black women | 32 | 25.1 | 32 | -2.1 (-3.1, -1.0)* | 3.8 (2.7, 4.9)* |
|  | White women | 12.3 | 8.9 | 11.1 | -3.2 (-3.7, -2.8)* | 3.7 (2.6, 4.8)* |
|  | Black men | 52.8 | 41.4 | 58 | $-1.6(-2.5,-0.7)^{*}$ | 4.8 (3.8, 5.9)* |
|  | White men | 24.1 | 17.6 | 23.6 | -2.7 (-3.0, -2.5)* | 4.5 (3.9, 5.1)* |
| $\begin{aligned} & \text { AAMR } \\ & (\geq 65) \end{aligned}$ | Total | 550.5 | 390.6 | 418.4 | $-2.9(-3.6,-2.1)^{*}$ | 1.0 (0.4, 1.6)* |
|  | Black women | 468.7 | 330.8 | 354.8 | -3.1 (-3.9, -2.3)* | $1.4(0.5,2.3)^{*}$ |
|  | White women | 510 | 356.2 | 375.4 | -3.0 (-4.0, -2.1)* | 0.8 (0.0, 1.6)* |
|  | Black men | 543.7 | 396.4 | 458.6 | $-2.6(-3.1,-2.1)^{*}$ | 2.2 (1.6, 2.8)* |
|  | White men | 636 | 465.3 | 509.9 | $-2.6(-3.3,-1.8)^{*}$ | 1.3 (0.9, 1.7)* |
| YPLL | Total | 203 | 150 | 189 | -2.8 (-3.1, -2.5)* | 3.7 (3.3, 4.1)* |
|  | Black women | 284 | 218 | 268 | -2.3 (-3.1, 1.6)* | 3.3 (2.5, 4.1)* |
|  | White women | 189 | 131 | 152 | -3.2 (-4.3, -2.1)* | 2.3 (1.4, 3.2)* |
|  | Black men | 257 | 226 | 312 | -1.0 (-1.8, -0.2)* | 4.8 (3.9, 5.7)* |
|  | White men | 195 | 140 | 178 | -2.9 (-3.1, -2.7)* | 3.9 (3.4, 4.3)* |

CI: Confidence interval. AAMR: Age-adjusted mortality rate per 100,000 population. YPLL: Years of potential life lost per 100,000 population. Mean annual percentage change shown for AAMR or YPLL for the specified time range. *Indicates that the mean annual percent change is significantly different from zero, $\mathrm{p}<0.05$.

Supplemental Figure. Cause-specific age-standardized years of potential life lost per 100,000 population in the total and race-sex subgroup populations in the US, 1999-2018


