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Authors

Powell-Wiley, Tiffany M Ngwa, Julius Kebede, Selomie <u>et al.</u>

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Impact of BMI on Heart Failure by Race/Ethnicity from Get With The Guidelines-Heart Failure Registry

Tiffany M. Powell-Wiley, MD, MPH¹, Julius Ngwa, PhD², Selomie Kebede, MD², Di Lu, MS³, Phillip J. Schulte, PhD⁴, Deepak L. Bhatt, MD, MPH⁵, Clyde Yancy, MD⁶, Gregg C. Fonarow, MD⁷, and Michelle A. Albert, MD, MPH⁸

¹Cardiovascular and Pulmonary Branch, National Heart, Lung, and Blood Institute, NIH, Bethesda, MD, USA

²Division of Cardiovascular Medicine, Howard University Hospital and School of Medicine, Washington, D.C., USA

³Duke Clinical Research Institute, Duke University School of Medicine, Durham, NC, USA

⁴Department of Health Sciences Research, Mayo Clinic, Rochester, MN, USA

⁵Brigham and Women's Hospital Heart & Vascular Center, Harvard Medical School, Boston, MA, USA

⁶Cardiovascular Division, Feinberg School of Medicine, Northwestern University, Chicago, IL, USA

⁷Ronald Reagan-University of California at Los Angeles Medical Center, Los Angeles, CA, USA

⁸CeNter for the StUdy of AdveRsiTy and CardiovascUlaR DiseasE (NURTURE Center), Cardiology Division, University of California at San Francisco Medical Center, San Francisco, CA, USA

Address for Correspondence to: Tiffany Powell-Wiley, MD, MPH, FAHA, Cardiovascular and Pulmonary Branch, National Heart, Lung, and Blood Institute, National Institutes of Health, Bethesda, MD, USA, 10 Center Drive, Room 5E-3340, Bethesda, MD, USA, 20892, USA, Tel/Fax: 301-594-3735/301-402-0888, tiffany.powell-wiley@nih.gov.

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Abstract

Objective—To evaluate race/ethnicity influence on BMI and mortality relationship among HFpEF and HFrEF patients.

Background—Prior studies demonstrate an "obesity paradox" among overweight/obese patients, where they have a better HF prognosis compared to normal-weight patients. Less is known about BMI and mortality relationship among diverse HF patients, particularly given disparities in obesity and HF prevalence.

Methods—Utilizing GWTG-HF data, we assessed BMI and in-hospital mortality relationship using logistic regression modeling. We assessed 30-day and 1-year all-cause mortality following discharge using Cox regression modeling.

Results—39,647 HF patients were included[white=32,434(81.8%); black=3,809(9.6%); Hispanic=1,928(4.9%); Asian=544(1.4%); other=932(2.3%)] with 59.7% HFpEF and 30.7% obese. More black and Hispanic patients had class I obese or higher (BMI 30 kg/m²) than white, Asian or patients of other racial/ethnic groups(P <0.0001). Among HFpEF, higher BMI was associated with lower 30-day mortality, up to 30 kg/m² with a small risk increase above 30 kg/ m²[BMI=30 vs 18.5 kg/m² hazard ratio(HR)=0.63, 95% confidence interval(CI) 0.54–0.73]. A modest relationship was observed in HFrEF(BMI=30 vs BMI=18.5 kg/m² HR=0.73, 95% CI 0.60–0.89), with no risk increase above 30 kg/m². There were no significant BMI by race/ethnicity interactions related to 30-day mortality(p>0.05).

Conclusions—Our work is one of the first suggesting the obesity paradox for 30-day mortality exists at all BMI levels in HFrEF, not HFpEF. Higher BMI was associated with lower 30-day mortality across racial/ethnic groups in a manner inconsistent with the J-shaped relationship noted for CAD. The differential slope of obesity and mortality among HFpEF and HFrEF patients potentially suggests differing mechanistic factors requiring further exploration.

Keywords

heart failure; obesity; race/ethnicity; mortality; GWTG-Heart Failure

INTRODUCTION

Both obesity and heart failure (HF) are unremitting in their rise; in the United States, 35% of Americans are obese (1) and HF afflicts 5.7 million individuals (2). Consequently, these conditions combined contribute to an estimated 246 billion dollars in healthcare expenditure (3,4). Although evidence independently implicates overweight (body mass index [BMI] 25 kg/m²) and obesity (BMI 30 kg/m²) with increased HF risk (1), most studies have suggested that increased BMI is associated with lower mortality in HF patients (6,7). However, whether this relationship differs between HF with preserved (HFpEF) and reduced ejection fraction (HFrEF) patients and whether there are important differences among HF patients of different racial/ethnic groups is less clear.

Considering the diverse racial composition of the U.S., the association of race/ethnicity on any BMI-HF mortality relationship becomes increasingly important. For example, African Americans (blacks) have the highest rates of both overweight/obesity and HF compared to

other racial/ethnic groups in the U.S. (8) as well as increased HF and HF hospitalization rates (9) - factors likely significantly contributing to gaps in mortality and longevity by race/ ethnicity. Additionally, as hospital readmissions are higher in blacks and Hispanics (10–13) and readmissions are closely linked to morbidity and mortality, it is imperative that BMI's association with HF be characterized both in general and according to race/ethnicity to understand its effect on HF outcomes, and to potentially inform therapeutic interventions. Utilizing Get With The Guidelines Heart Failure (GWTG-HF) registry data for both HFpEF and HFrEF patients, we sought to 1) assess the association between BMI and in-hospital mortality according to race/ethnicity; and 2) determine associations between BMI and 30-day and 1-year all-cause mortality following discharge alive according to race/ethnicity.

METHODS

Data

Data Source—The GWTG-HF is a registry and performance improvement initiative started in 2005 to enhance adherence to practice guidelines for hospitalized HF patients. This voluntary American Heart Association (AHA) program collects data on patient characteristics using web-based information systems. The program's methods, design, and validity have been published previously (14–17). Hospitals participating in the registry submit clinical information regarding medical history, laboratories, diagnostic testing, hospital care and outcomes of patients hospitalized for HF using an online, interactive case report form and patient management tool (Quintiles, Cambridge MA). To be eligible for GWTG-HF, patients had to be adults hospitalized for a HF episode as the primary cause of admission or with significant HF symptoms that developed during hospitalization with a primary discharge diagnosis of HF. Race/ethnicity data were collected for evaluating subgroup differences in outcomes. Patients, based on self-reported race/ethnicity, were assigned to race/ethnicity categories using options defined by the case report form as follows: race - American Indian or Alaska Native, Asian, Black or African-American, Native Hawaiian or Pacific Islander, White or unable to be determined; ethnicity, Hispanic- yes, no or unable to be determined.

Data Collection Protocols—HF status, including HFpEF vs. HFrEF diagnosis, was determined by point-of-care providers, based on American Heart Association guidelines. HFrEF patients had ejection fraction (EF) 40% and HFpEF patients had EF 40% (18). Patient height and weight were collected at time of admission and BMI was imputed by the GWTG Patient Management Tool (PMT) data collection form. Covariates were collected from historical records or during admission, depending on time of presentation. For example, HF clinical characteristics were recorded based on current hospital admission or when the condition was first recognized. Quintiles is the GWTG data collection (through their Patient Management Tool – PMT) and coordination center. Duke Clinical Research Institute (DCRI) serves as the data analysis center and has an agreement to analyze aggregate de-identified data for research purposes. Participating institutions were required to comply with local regulatory guidelines and the local institutional review board.

Study Population

The starting population for this study was 65,037 GWTG-HF patients linked to CMS from 292 sites. The study period was from January 2005 through December 2011. We sequentially excluded patients who were not enrolled in fee-for-service Medicare at discharge (n= 2,473), patients without race/ethnicity data (n= 1,768), BMI missing (n= 14,661), ejection fraction missing (n= 5,416), transfers out or discharge information missing or not documented or left against medical advice (n= 1,072) (Figure 1). A total of 39,647 patients having HFpEF or HFrEF, with BMI and race/ethnicity data were documented. For post discharge outcomes, we excluded the study population patients that died in hospital or discharged to hospice (n=2,086).

Outcome Measures

The study objective was to assess the association between BMI and in-hospital mortality according to race/ethnicity and associations between BMI and 30-day or 1-year all-cause mortality following discharge alive, according to race/ethnicity. The outcomes were in-hospital mortality, 30-day and 1-year all-cause mortality. The 30-day and 1-year mortality were evaluated from discharge date to 30 days and 1 year afterwards.

Statistical Analysis

The study population was stratified by HFpEF and HFrEF and analyses were performed separately for each cohort. For descriptive analyses, baseline patient and hospital characteristics were stratified among Whites, Blacks, Hispanic, Asian and other race/ ethnicity groups. For categorical variables, proportions were used and differences assessed by Chi-square test. For continuous/ordinal variables, means and standard deviations were presented across race/ethnicity groups assessed by Kruskal-Wallis tests.

The relationship between BMI, race/ethnicity, and in-hospital mortality was evaluated using logistic regression. We tested the interaction between BMI and race/ethnicity to assess whether the relationship between BMI and odds of mortality were similar among racial/ ethnic groups. A multivariable model adjusted for patient and hospital characteristics, including age, sex, medical history (anemia, ischemic history, CVA/TIA, diabetes, hyperlipidemia, hypertension, COPD or asthma, PVD, renal insufficiency, smoking), admission vitals and labs (systolic blood pressure, heart rate, sodium, and blood urea nitrogen), hospital region, academic status and number of beds. Post-discharge outcomes of 30-day and 1-year mortality were similarly evaluated using Cox proportional hazards regression.

For missing adjustment variables, medical history variables were imputed to "no" as data abstractors were likely to skip this section of the data collection form when none applied; multiple imputation with 25 imputations was used for other patient covariates. Hospital characteristics were not imputed. Adjustment covariates were assessed for linearity and proportional hazards assumptions as needed and transformations applied when appropriate. Restricted cubic spline transformations flexibly illustrate relationships between BMI and mortality. To interpret results numerically, we also fit linear splines of BMI. A spline knot was chosen which balanced model-fit by maximizing model likelihood and interpretation of

results. In HFrEF, we report results for BMI per 1 kg/m² increase up to 25 kg/m² and per 1 kg/m² increase above 25 kg/m² for in-hospital mortality. Among HFpEF, the knot point for in-hospital mortality was 30 kg/m². For 30-day and 1-year mortality, the BMI knot point was 30 kg/m². Restricted cubic spline relationships are plotted for 30-day mortality.

All tests were two-tailed and a p-value<0.05 was considered statistically significant. All analyses were performed using SAS version 9.3 (SAS Institute, Cary, NC). The authors had full access to all study data and take responsibility for its integrity and the data analysis.

RESULTS

Baseline Characteristics

Of the 39,647 patients in the study population, 23,653(59.7%) had HFpEF and 15,994(40.3%) had HFrEF. Baseline patient and hospital characteristics for those with HFpEF are summarized in Table 1 and e-Table 1 in Supplement, while those of patients with HFrEF are summarized in Table 2 and e-Table 2 in Supplement. Figure 2 demonstrates the mean BMI distribution by race and HF status.

HFpEF Patients—Blacks with HFpEF were significantly younger than other racial/ethnic groups with HFpEF [mean age [(standard deviation(SD)]=77(8.3) years](p < 0.0001) (Table 1). Across all racial/ethnic groups, the majority of HFpEF patients were women. The mean BMI for the overall cohort was 28.6 (7.9) kg/m²; blacks with HFpEF had the highest BMI [mean BMI=30.9(9.3) kg/m²] and highest likelihood of Class III obesity (14.3%, p < 0.0001). Insulin-dependent diabetes was more common among Hispanics (26.2%) with HFpEF compared with other racial/ethnic groups (p< 0.0001). Additionally, hypertension, prior stroke, and renal insufficiency were more prevalent among blacks with HFpEF as compared to other racial/ethnic groups (89.0%, 20.4% and 26.2% respectively, p < 0.0001). Whites (55.7%) were more likely to have an ischemic etiology for HFpEF compared with Hispanics (54.7%), blacks (46.9%), Asians (50.3%) and others (54.7%) (p < 0.0001).

HFrEF Patients—Among HFrEF patients, blacks were younger [mean age 76(7.8) years] and more likely women (46.4%) compared with other racial/ethnic groups with HFrEF (Table 2). Mean BMI for the overall HFrEF population was 26.7(6.6) kg/m², and blacks with HFrEF had a higher mean BMI than other racial/ethnic groups (p<0.0001); 3.7% of the overall HFrEF population had Class III obesity with blacks (5.5%) having a higher likelihood of Class III obesity as compared to whites (3.6%), Hispanics (3.1%), Asians (1.1%), and others (4.1%) (p<0.0001). Hispanics (23.5%) had the highest rates of insulindependent diabetes (p<0.0001), while Asians (34.3%) had the highest rates of non-insulin dependent diabetes (p<0.0001). Hispanics with HFrEF had the highest rates of hypertension (p<0.0001), while blacks had the highest rates of prior CVA (p=0.0114). Asians with HFrEF (28.0%) had the highest rates of renal insufficiency as compared to whites (19.6%), blacks (24.1%), Hispanics (19.8%) and others (20.4%) (p<0.0001). As with HFpEF patients, whites with HFrEF were more likely to have an ischemic etiology compared with other racial/ethnic groups (p<0.0001).

Patient Mortality by BMI Category

HFpEF Patients—Among HFpEF patients, higher BMI was associated with lower 30-day all-cause mortality up to 30 kg/m² with a slight increase in risk above 30 kg/m² (Figure 3; BMI=30 kg/m² vs. BMI=18.5 kg/m²: hazard ratio (HR) 0.63, 95% CI=0.54 – 0.73). Additionally, hazard of 1-year, all-cause mortality was 4% lower for one-unit BMI increase up to 30 kg/m² [HR=0.96 (95% CI=0.95–0.96)] (Table 3). There was no significant relationship between BMI and 1-year mortality among those with HFpEF above BMI of 30 kg/m². With regard to in-hospital mortality, there was no significant relationship between BMI and in-hospital mortality among HFpEF patients up to BMI of 30 kg/m². For BMI above 30 kg/m², each one-unit increase in BMI was associated with a 2% greater odds of inhospital death for HFpEF patients[odds ratio (OR) = 1.02(95% confidence interval (CI)=1.01–1.04)] (e-Table 3 in supplement).

HFrEF Patients—For HFrEF patients, up to a BMI of 30 kg/m², 30-day all-cause mortality decreased with every BMI unit increase (BMI=30 kg/m² vs BMI=18.5 kg/m² HR 0.73, 95% CI:0.60–0.89) (Figure 4). A smaller relationship between BMI and lower 30-day all-cause mortality was observed in HFrEF patients, as shown in Figure 4. Thirty-day mortality rates were similar across obesity classes. Above BMI of 30 kg/m², each one-unit BMI increase was associated with a 1% higher hazards of 1-year all-cause mortality [HR=1.01(95% CI=1.00–1.02)] (Table 3). Among patients with HFrEF, each BMI unit increase up to 25 kg/m² was associated with a 5% lower odds of in-hospital death [OR = 0.95(95% CI= 0.91–0.99)]. Above a BMI of 25 kg/m², each one unit BMI increase was associated with a 4% higher odds of in-hospital death [OR = 1.04(95% CI=1.03–1.05)] (e-Table 3 in supplement).

Interactions Between BMI and Race/Ethnicity

No significant BMI by race/ethnicity interactions related to 30-day mortality were seen among HFpEF or HFrEF patients (all p>0.05, Table 3). The relationship between BMI below 30 kg/m² and hazards of 1-year all-cause mortality was significantly different among racial/ethnic groups with HFrEF. BMI by race/ethnicity interactions were not statistically significant for 1-year, all-cause mortality among HFpEF and HFrEF patients with a BMI > 30 kg/m². There were no significant BMI by race/ethnicity interactions related to in-hospital mortality for patients with HFpEF or HFrEF (all p>0.05, e-Table 3 in supplement).

DISCUSSION

Our findings demonstrate that in a nationally representative, racially and ethnically diverse cohort, high BMI was both common (30.7%) and associated with lower 30-day all-cause mortality for older (>65 years) HFpEF and HFrEF patients. While blacks and Hispanics had higher obesity rates than whites, Asians, and other racial/ethnic groups, there was no significant interaction between BMI and race in the relationship between HF and mortality. Unlike the relationship between BMI and mortality for CAD patients, the relationship between BMI and 30-day mortality was not J-shaped for HFpEF and HFrEF patients and 30-day mortality rates remained relatively constant in Class II and Class III obesity in HFrEF patients. However, there was a small, but statistically significant increase in 30-day mortality

rates for HFpEF patients with a BMI above 30 kg/m². Differential slopes in the association between BMI below 30 kg/m² and 30-day mortality among HFpEF and HFrEF patients likely suggests differences in mechanistic factors that promote early mortality between the two HF phenotypes.

Our findings contribute to the literature on BMI and mortality in HF patients in several important ways. First, this is one of few studies to compare the relationship between BMI and 30-day all-cause mortality across racial/ethnic groups with HFpEF or HFrEF (19), as highlighted in Table 4. Prior GWTG-HF studies have been limited to evaluating in-hospital mortality among those with HFpEF and HFrEF in one racial/ethnic group (20) or have not distinguished outcomes between HFpEF and HFrEF patients when comparing in-hospital or 30-day mortality across racial/ethnic groups (13,21). In GWTG-HF, Hispanics with HFpEF were less likely to have ischemic heart disease compared with Hispanics with HFrEF. Additionally, Hispanics with HFpEF - but not HFrEF - had lower in-hospital mortality compared with whites (21). Thomas and colleagues demonstrated that blacks and Hispanics in the GWTG-HF registry had a lower likelihood of in-hospital death as compared to white patients. One potential explanation for the differences was that HF with a non-ischemic etiology was more common among blacks and Hispanics (20). More recent data suggested that 30-day survival after index admission is greater among blacks compared with whites, even after controlling for co-morbidities, hospital characteristics, and socioeconomic status. Our work extends the findings from these prior GWTG-HF studies to demonstrate that BMI differences do not appear to explain differences in 30-day all-cause mortality across racial and ethnic groups. This conclusion is supported by a recent meta-analysis which demonstrated that greater BMI was associated lower one-year all-cause mortality in racially/ ethnically diverse cohorts with HF from different parts of the world (i.e., Asia, North/South America, Europe) (19).

Second, our findings enhance the body of knowledge about the perplexing repercussions of increasing BMI on HF outcomes. Many prior studies have demonstrated the presence of an "obesity paradox" among overweight and Class I and II obese patients, or that these patients have a better prognosis with HF as compared to patients with a normal BMI (22-25. Our work is one of the first studies to suggest that the BMI paradox for 30-day mortality exists at all BMI levels, including Class II and Class III obesity, among HFrEF patients; our findings appear consistent with a recent meta-analysis of six studies and over 22,000 patients (26). Other cohorts in which the relationship between Class II or III obesity and HF mortality has been investigated have been limited in size (27), in the numbers of those with Class III obesity (28), or in diversity (29). Padwal and colleagues, for example, did not look across race/ethnic groups when evaluating the obesity paradox in HF patients (29), while Ziaeian and colleagues looked across race/ethnic groups without discussing the obesity paradox (30). Differences in the proportion of study participants with HFpEF compared with HFrEF, in the racial/ethnic composition of the populations, and prevalent co-morbidities among study participants may explain the differences between our findings and those of other studies looking at Class II/III obesity and mortality. In particular, further work is needed to differentiate how Class III obesity impacts HF mortality because Class III obesity is strongly associated with adverse cardiac remodeling and the subsequent HF development (31).

Finally, this is one of the first studies to distinguish between HFpEF and HFrEF when evaluating the association of BMI and mortality among HF patients. Unlike prior studies, which have only looked at HFrEF or HFpEF populations (32,33), our findings suggest heterogeneity in mortality risk at lower BMI between the two HF phenotypes, with a greater 30-day mortality risk at higher BMI among HFpEF patients. From a pathophysiologic standpoint, this may represent differences in co-morbidities between those with HFrEF and HFpEF who survive hospitalization. For example, compared with HFpEF patients, HFrEF patients who survive a hospitalization may be more likely to have reductions in ejection fraction due to mechanisms unrelated to cardiovascular risk factors or cardiovascular disease and which may carry a better prognosis (27). Additionally, HFpEF patients at a given BMI likely have higher blood pressures and tolerate higher doses of cardioprotective medications compared with those with HFrEF, leading to differences in mortality risk (34). Recent work suggests that patient characteristics, including age, left ventricular function, and HF chronicity, impact the prognostic association between BMI and all-cause mortality. As compared with our findings, BMI was more strongly associated with all-cause mortality for HF patients with a left ventricular ejection fraction less than 50% as compared to those with an ejection fraction greater than or equal to 50% in this meta-analysis of international cohorts (19). Our study findings highlight the importance of differentiating between HFpEF and HFrEF when evaluating BMI and mortality in HF patients.

The strengths of the current study include data from a nationally representative, multiethnic cohort with well-established protocols for data collection and analyses. The GWTG-HF data was also linked to high-quality, standardized data from the Centers for Medicare and Medicaid Services to determine 30-day mortality for study participants, further enhancing the reliability of the study data. However, limitations of the study must be acknowledged. There are significant differences in demographics, clinical characteristics and treatments by BMI categories and we cannot exclude residual measured and unmeasured confounding factors contributing to these findings. We were unable to compare mortality across measures of body fat distribution or more accurate measures of adiposity, such as waist circumference in this cohort. Alternative measures of body fat distribution instead of BMI may be of particular importance when attempting to differentiate cardiovascular risk relative to adiposity among racially and ethnically diverse populations (35). Patients' weights for determining BMI were obtained during hospitalization for HF and thus may not represent the patients' weight at a time they are well compensated. However, relatively few patients would be expected to change BMI categories on the basis of wet versus dry weight. Another potential limitation is that GWTG-HF program hospitals are voluntary participants and may be more motivated for quality improvement, which may lead to better patient outcomes as compared to other hospitals around the country, limiting study generalizability. Additionally, the proportion of racial/ethnic minority patients seen in GWTG-HF hospitals may differ from the proportion in those hospitals not represented in the program. Hospitals outside of the GWTG-HF program may disproportionately care of racial/ethnic minority patients and may have differ from GWTG-HF hospitals in quality of care provided. Finally, method of recording race and ethnicity by patient self-designation as recorded by administrative staff or admitting providers is likely not as reliable as direct patient reporting.

In conclusion, black and Hispanic patients in the GWTG-HF registry were more likely to be obese than white, Asian or other patients. Higher BMI was associated with lower 30-day all-cause mortality in each racial/ethnic group in a manner not consistent with the J-shaped obesity paradox noted in prior studies. Additionally, the differential slope of the BMI and 30-day all-cause mortality association below BMI of 30 kg/m² in HFpEF and HFrEF and higher mortality risk above BMI of 30 kg/m² in HFpEF possibly suggests differing mechanistic factors which require further exploration.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

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Abbreviations

HF	heart failure
HFpEF	heart failure with preserved ejection fraction
HFrEF	heart failure with reduced ejection fraction
CAD	coronary artery disease
GWTG-HF	Get With The Guidelines-Heart Failure
BMI	body mass index
HR	hazard ratio
OR	odds ratio
SD	standard deviation
CI	confidence interval

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CLINICAL PERSPECTIVE

Race/ethnicity does not appear to modify the relationship between BMI and mortality among HF patients, and the obesity paradox appears to exist across BMI levels for HF patients with preserved or reduced ejection fraction.

TRANSLATIONAL OUTLOOK

Further research is needed to investigate mechanisms by which the obesity paradox exists across BMI levels for patients with preserved or reduced ejection fraction HF.



Figure 1. Flow diagram of the study population selection process

Participants were excluded systematically from the original GWTG-HF cohort.



Figure 2. Mean BMI by race/ethnicity among HFpEF and HFrEF patients, GWTG-HF, 2005–2011

Gray bars represent patients with HFpEF, while blue bars represent patients with HFrEF. Error bars represent standard deviation.



Figure 3. Adjusted Association between Body Mass Index and 30-day Mortality for HFpEF Patients, GWTG-HF, 2005–2011

Among HFpEF patients, higher BMI was associated with lower 30-day all-cause mortality up to 30 kg/m^2 with little change in risk above 30 kg/m^2 .



Figure 4. Adjusted Association between Body Mass Index and 30-day Mortality for HFrEF Patients, GWTG-HF, 2005–2011

Up to a BMI of 30 kg/m², 30-day all-cause mortality decreased with every BMI unit increase (BMI=30 kg/m² vs BMI=18.5 kg/m² HR 0.73, 95% CI:0.60–0.89).

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Table 1

Baseline Patient and Hospital Characteristics of Patients with HFpEF, GWTG-HF, 2005-2011

Variable	Overall (N=23653)	White (N=19575)	Black (N=2085)	Hispanic (N=1088)	Asian (N=368)	Other (N=537)	P-value
Demographics							
Age, yrs, mean(SD *)	80.8 (8.1)	81.35 (8.0)	77.2 (8.3)	79.1 (8.3)	80.4 (7.9)	79.6 (7.9)	<0.0001
Female, No. (%)	14872 (62.7)	12188 (62.3)	1411 (67.7)	675 (62.0)	214 (58.2)	339 (63.1)	<0.0001
Insurance Status, No. (%)							<0.0001
Medicare	16806 (74.4)	14078 (75.2)	1342 (68.9)	741 (70.6)	267 (75.0)	378 (72.7)	
Medicaid	1301 (5.8)	688 (3.7)	326 (16.7)	186 (17.7)	55 (15.5)	46 (8.9)	
Other	4496 (19.9)	3963 (21.2)	281 (14.4)	122 (11.6)	34 (9.6)	96 (18.5)	
BMI, kg/m ² , mean(SD *)	28.6 (7.9)	28.4 (7.8)	30.9 (9.3)	29.5 (7.8)	24.8 (6.2)	28.1 (7.3)	<0.0001
BMI, categories, No. (%)							<0.0001
Underweight (18.5 kg/m^2)	1076 (4.6)	911 (4.7)	66 (3.2)	33 (3.0)	41 (11.1)	25 (4.7)	
Normal $(18.5 - 24.9 \text{ kg/m}^2)$	7602 (32.1)	6438 (32.9)	508 (24.4)	292 (26.8)	182 (49.5)	182 (33.9)	
Overweight $(25 - 29.9 \text{ kg/m}^2)$	6646 (28.1)	5518 (28.2)	555 (26.6)	332 (30.5)	89 (24.2)	152 (28.3)	
Obese I $(30 - 34.9 \text{ kg/m}^2)$	4147 (17.5)	3406 (17.4)	404 (19.4)	213 (19.6)	38 (10.3)	86 (16.0)	
Obese II $(35 - 39.9 \text{ kg/m}^2)$	2178 (9.2)	1754 (9.0)	253 (12.1)	110 (10.1)	9 (2.5)	52 (9.7)	
Obese III (40 kg/m^2)	2004 (8.5)	1548 (7.9)	299 (14.3)	108 (9.9)	9 (2.5)	40 (7.5)	
Comorbidities							
Diabetes-Insulin Treated, No. (%)	4087 (17.4)	3111 (16.0)	527 (25.4)	283 (26.2)	63 (17.2)	103 (19.4)	<0.0001
Diabetes-Non-insulin Treated, No. (%)	5511 (23.5)	4380 (22.6)	571 (27.6)	330 (30.5)	103 (28.1)	127 (24.0)	<0.0001
Hypertension, No. (%)	18930 (80.6)	15436 (79.5)	1844 (89.0)	922 (85.3)	318 (86.9)	410 (77.4)	<0.0001
Ischemic Etiology, No. (%)	12858 (54.8)	10821 (55.7)	972 (46.9)	591 (54.7)	184 (50.3)	290 (54.7)	<0.0001
CAD, No. (%)	11434 (48.7)	9634 (49.6)	880 (42.5)	514 (47.6)	160 (43.7)	246 (46.4)	<0.0001
CVA/TIA, No. (%)	4064 (17.3)	3355 (17.3)	422 (20.4)	149 (13.8)	67 (18.3)	71 (13.4)	<0.0001
Anemia, No. (%)	5246 (22.4)	4287 (22.1)	587 (28.3)	200 (18.5)	65 (17.8)	107 (20.2)	<0.0001
Smoking, No. (%)	1827 (7.8)	1451 (7.5)	243 (11.7)	76 (7.0)	17 (4.7)	40 (7.5)	<0.0001
Renal Insufficiency, No. (%)	4451 (19.0)	3493 (18.0)	543 (26.2)	210 (19.4)	85 (23.2)	120 (22.6)	<0.0001
Hospital Characteristics							
Number of Beds, mean (SD^*)	410.0 (235.6)	409.2(236.9)	456.9(234.9)	358.3(198.4)	362.0(222.2)	392.3(234.7)	<0.0001

* SD - standard deviation.

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Table 2

Baseline Patient and Hospital Characteristics of Patients with HFrEF, GWTG-HF 2005–2011

Variable	Overall (N=15994)	White (N=12859)	Black (N=1724)	Hispanic (N=840)	Asian (N=176)	Other (N=395)	P-value
Demographics							
Age, yrs, mean (SD^*)	78.6 (7.9)	79.2 (7.8)	75.7 (7.8)	76.7 (7.7)	79.0 (7.6)	77.3 (8.3)	<0.0001
Female, No. (%)	6297 (39.4)	4943 (38.4)	800 (46.4)	326 (38.8)	72 (40.9)	156 (39.5)	<0.0001
Insurance Status, No. (%)							<0.0001
Medicare	11454 (75.9)	9307 (76.3)	1161 (72.8)	570 (74.0)	132 (79.5)	156 (39.5)	
Medicaid	728 (4.8)	387 (3.2)	208 (13.0)	106 (13.8)	16 (9.6)	284 (77.4)	
Other	2910 (19.3)	2500 (20.5)	226 (14.2)	94 (12.2)	18 (10.8)	11 (3.0)	
BMI, kg/m ² , mean(SD *)	26.7 (6.6)	26.6 (6.6)	27.6 (7.4)	27.0 (6.0)	23.4 (5.5)	26.6 (6.4)	<0.0001
BMI, categories, No. (%)							<0.0001
Underweight (18.5 kg/m^2)	903 (5.7)	734 (5.7)	89 (5.2)	37 (4.4)	24 (13.6)	19 (4.8)	
Normal $(18.5 - 24.9 \text{ kg/m}^2)$	6363 (39.8)	5156 (40.1)	628 (36.4)	314 (37.4)	100 (56.8)	165 (41.8)	
Overweight $(25 - 29.9 \text{ kg/m}^2)$	4890 (30.6)	3972 (30.9)	489 (28.4)	275 (32.7)	40 (22.7)	114 (28.9)	
Obese I $(30 - 34.9 \text{ kg/m}^2)$	2344 (14.7)	1831 (14.2)	304 (17.6)	135 (16.1)	8 (4.6)	66 (16.7)	
Obese II $(35 - 39.9 \text{ kg/m}^2)$	899 (5.6)	709 (5.5)	120 (7.0)	53 (6.3)	2 (1.1)	15 (3.8)	
Obese III (40 kg/m^2)	595 (3.7)	457 (3.6)	94 (5.5)	26 (3.1)	2 (1.1)	16 (4.1)	
Comorbidities							
Diabetes-Insulin Treated, No. (%)	2510 (15.8)	1899 (14.9)	329 (19.2)	196 (23.5)	23 (13.1)	63 (16.3)	<0.0001
Diabetes-Non-insulin Treated, No. (%)	3706 (23.4)	2894 (22.7)	426 (24.8)	245 (29.3)	60 (34.3)	81 (20.9)	<0.0001
Hypertension, No. (%)	11633 (73.4)	9108 (71.5)	1423 (83.0)	695 (83.2)	132 (75.4)	275 (71.1)	<0.0001
Ischemic Etiology, No. (%)	10807 (68.2)	8981 (70.5)	915 (53.4)	553 (66.2)	114 (65.1)	244 (63.1)	<0.0001
CAD, No. (%)	9369 (59.1)	7812 (61.3)	783 (45.7)	482 (57.7)	89 (50.9)	203 (52.5)	<0.0001
CVA/TIA, No. (%)	2487 (15.7)	2002 (15.7)	298 (17.4)	99 (11.9)	27 (15.4)	61 (15.8)	0.0114
Anemia, No. (%)	2535 (16.0)	2024 (15.9)	325 (19.0)	113 (13.5)	25 (14.3)	48 (12.4)	0.0007
Smoking, No. (%)	1775 (11.2)	1345 (10.6)	285 (16.6)	86 (10.3)	6 (3.4)	53 (13.6)	<0.0001
Renal Insufficiency, No. (%)	3204 (20.2)	2497 (19.6)	414 (24.1)	165 (19.8)	49 (28.0)	79 (20.4)	<0.0001
Hospital Characteristics							
Number of Reds mean (SD *)	429.9(233.6)	430.3 (236.2)	465.0 (233.8)	386.9(180.6)	350,5(2,12,4)	393 3(229.1)	<0.0001

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* SD - standard deviation.

Table 3

GWTG-HF 2005-2011
IFpEF and HFrEF Patients,
ise Mortality, H
1-year All-Cau
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between BMI i
Association

Outcome	Variable	$\mathrm{HR}^{*, \dagger}$	Lower 95% CI	Upper 95% CI	P-value
	Overall BMI up to 30 kg/m ² *Race				0.92
20 Jan All Carnes Meadeline for UBABE metioner	BMI per 1 unit increase up to 30 kg/m ²	0.96	0.94	16.0	<0.0001
20-day All-Cause Moltanty for ALPEF pauents	Overall BMI above 30 kg/m ² *Race				0.96
	BMI per 1 unit increase above 30 kg/m ²	1.01	1.00	1.03	0.0477
	Overall BMI up to 30 kg/m ² *Race				0.08
20 Jan All Carros Montellin for UEADE antionts	BMI per 1 unit increase up to 30 kg/m ²	0.97	0.95	66.0	0.0016
outage All-Cause Moltanty for ALFLEF patents	Overall BMI above $30 \text{ kg/m}^2 \text{ *}$ Race				0.92
	BMI per 1 unit increase above 30 kg/m ²	1.01	66.0	1.03	0.51
	Overall BMI up to 30 kg/m ² * Race				0.27
1 All Course Manufity for UB-DE actionts	BMI per 1 unit increase up to 30 kg/m^2	0.96	0.95	0.96	<0.0001
1-year Alt-Cause Moriality for the percents	Overall BMI above 30 kg/m ² *Race				0.65
	BMI per 1 unit increase above 30 kg/m ²	1.00	1.00	1.01	0.67
	Overall BMI up to 30 kg/m ² * Race				0.0246
1 room All Connor Montality for UD-DD mations	BMI per 1 unit increase up to 30 kg/m ²	0.96	0.95	0.97	< 0.0001
1-year Alt-Cause Morianty for HITLY parents	Overall BMI above $30 \text{ kg/m}^2 \text{ *}$ Race				0.96
	BMI per 1 unit increase above 30 kg/m ²	1.01	1.00	1.02	0.0438

JACC Heart Fail. Author manuscript; available in PMC 2019 March 01.

HR reported per 1 kg/m² BMI increase

 $\dot{ au}_{30}$ -day and 1-year all-cause mortality were evaluated from the discharge date to 30 days and 1 year afterward

Table 4

Comparison of Studies Examining the Obesity Paradox in Cardiovascular Disease, 2014 to Present

Reference	Main Findings
Powell-Wiley TM, Ngwa J, Kedebe S et al. 2017. Impact of BMI on Heart Failure with Preserved or Reduced Ejection Fraction: Data by Race/Ethnicity from Get With The Guidelines- Heart Failure Registry.	Race/ethnicity does not appear to modify the relationship between BMI and mortality among HF patients, and the obesity paradox appears to exist across BMI levels for HF patients with preserved or reduced ejection fraction.
Wang ZJ, Zhou YJ, Galper BZ, et al. Association of body mass index with mortality and cardiovascular events for patients with coronary artery disease: a systematic review and meta- analysis. <i>Heart.</i> 2015;101(20):1631–8.	There is a J-shaped relationship between BMI categories and risk of mortality among patients with CAD, with underweight patients having highest risk. Survival benefit of obesity was attenuated by 5-year follow up.
Vivo RP, Krim SR, Liang L, et al. Short- and long-term rehospitalization and mortality for heart failure in 4 racial/ethnic populations. <i>J Am Heart Assoc.</i> 2014;3:e001134.	Black and Hispanic HF patients had higher 30-day and 1-year hospital readmission rates, but lower 30-day and 1-year mortality compared to white patients, after adjustment for socioeconomic status and patient/hospital characteristics.
Shah R, Gayat E, Januzzi JL, Jr., et al. Body mass index and mortality in acutely decompensated heart failure across the world: a global obesity paradox. <i>J Am Coll Cardiol.</i> 2014;63:778–85.	There is an obesity paradox in 1-year mortality of acute decompensated HF patients globally, after adjustment for patient characteristics. However, the association of BMI and outcome is restricted to specific groups, such as non-diabetic, older patients with de novo HF diagnosis or HF with reduced LVEF.
Padwal R, McAlister FA, McMurray JJV et al. The obesity paradox in heart failure patients with preserved versus reduced ejection fraction: a meta-analysis of individual patient data. <i>Int J</i> <i>Obesity</i> . 2014;38:1110–1114.	There is a U-shaped mortality curve among HF patients with preserved or reduced LVEF. Lowest mortality was seen in patients with BMI between 30 and 30.4 kg/m ² . After adjustment for confounders, high BMI status remained significantly associated with lower mortality risk.
Ziaeian B, Heidenreich PA, Xu H, et al. Race/ethnic differences in outcomes among hospitalized Medicare patients with heart failure and preserved ejection fraction. <i>J Am Coll Cardiol HF</i> . 2017;5:483–493.	Lower mortality risk after hospitalization for HFpEF was found in black, Hispanic, and Asian patients after adjusting for SES and patient/hospital characteristics. Black and Hispanic HFpEF patients had higher readmission rates, but lower short- and long-term mortality.