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Everyday Discrimination and Mental Stress–Induced Myocardial Ischemia

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Abstract

Objectives: Mental stress-induced myocardial ischemia (MSIMI), a transient myocardial ischemic response to mental stress, is associated with poorer outcomes among patients with coronary heart disease (CHD) and is more likely to occur among women. Yet, predictors of MSIMI are not well explored. The current study investigated the association between experiences of everyday discrimination and MSIMI among patients with recent MI and contrasted the results with conventional stress-induced myocardial ischemia (CSIMI). We examined sex differences in associations.

Methods: We studied 295 post-MI patients (145 women, 150 men). Provocation of myocardial ischemia with mental stress (speech task) and conventional stress (exercise or pharmacologic) was assessed by myocardial perfusion imaging. Frequency of exposure to everyday discrimination was assessed via questionnaire using the Everyday Discrimination Scale (EDS).

Results: The mean age was 51 years in both women and men, and the EDS score ranged from 10 to 38 (mean 17; SD 6). After multivariable analysis, each standard deviation increase in EDS score (more frequent exposure) was associated with an increased odds of MSIMI (OR=1.57 [1.10 - 2.23]). EDS score was not associated with CSIMI (OR=0.86 [0.64 - 1.17]). Women demonstrated a two-fold increase (OR=1.96 [1.13 - 3.38], p=0.02) in the adjusted odds of MSIMI with each

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standard deviation increase in EDS score compared to a 1.4-fold increase (OR=1.40 [0.80 - 2.44], p=0.24) among men; however, interaction was not statistically significant.

Conclusions: Among post-MI patients, everyday discrimination was positively associated with occurrence of MSIMI, but not with CSIMI; associations were more pronounced among women.

Keywords

Coronary artery disease; Mental-stress induced myocardial ischemia (MSIMI); myocardial infarction (MI); Social Determinants of Health; Social Discrimination; Women's Health

Introduction

Mental stress-induced myocardial ischemia (MSIMI) is a transient myocardial ischemic response to mental stress. MSIMI has been observed in anywhere from 15% to 70% of patients with pre-existing coronary artery disease (CAD) (1). Similar to ischemia induced by conventional (exercise) stress tests, MSIMI is associated with a doubling of recurrent events and mortality among patients with CAD; however, unlike conventional (exercise) stress-induced myocardial ischemia (CSIMI), MSIMI occurs at a lower workload (i.e., lower heart rate, lower stroke volume), is largely independent of underlying CAD severity, and is usually silent (1–4).

Previous studies among patients with recent myocardial infarction (MI) and stable coronary heart disease (CHD) have shown that women are more likely to develop MSIMI compared to men, with only a modestly higher risk of CSIMI among women (5, 6). In fact, it has been suggested that MSIMI may be a potential mechanism explaining worse morbidity and mortality after MI among women, particularly young women (2, 7). There is growing evidence that the dominant mechanism driving MSIMI among women is not underlying CAD severity but microvascular dysfunction and peripheral vasoconstriction, for which women are particularly vulnerable, however other factors likely play a role (2, 8, 9). Yet, there has been limited research on the predictors of MSIMI overall and among young women in particular.

A number of psychosocial factors related to stress – including depression and state and traitanger – have been associated with higher risk for MSIMI (10–12). Not only are women more likely to experience psychosocial risk factors, they are potentially at higher risk of adverse cardiovascular consequences as a result of these factors (13). One psychosocial factor that may have particular relevance for women with MI is discrimination. Discrimination – unfair treatment based on group identity – is disproportionately experienced by members of groups that have historically been oppressed, marginalized, and disenfranchised, and young women with MI are more likely to be low income and Black than their male counterparts (14–16). There is an increasing body of evidence linking exposure to discrimination, conceptualized as a form of psychosocial stress, with physical health outcomes (17, 18). Several studies have identified associations between self-reported discrimination and cardiovascular disease (CVD) risk factors (i.e., hypertension, visceral fat, and atherosclerosis), inflammatory markers (i.e., C-reactive protein, interleukin-6), and incident CVD (19–23). However, most of these studies have focused on healthy populations,

and, to our knowledge, no studies have examined the association between discrimination and

MSIMI. Investigating discrimination in patients with recent MI is particularly important as this is a high-risk group that may be subject to discrimination not just in everyday settings, but in their more frequent interactions with the healthcare system.

The current study seeks to investigate the association between self-reported discrimination and MSIMI in a population of young and middle-aged survivors of MI. This is an important group to study as previous studies have found that young and middle-aged, but not older, women have higher mortality and complication rates after an MI compared with men of similar age (24, 25). Additionally, disparities in MSIMI between women and men with CHD were only found in younger patients (26). We used CSIMI as a control condition. We also examined differences in these associations by sex. Our hypothesis was that there would be an association between discrimination and MSIMI but not with CSIMI due to the hypothesized mechanisms through which emotional stressors trigger ischemia. Additionally, we hypothesized that discrimination will be more strongly associated with MSIMI among women compared to men, independent of sociodemographic and clinical risk factors.

Methods

Study Design

Study participants were patients with a previous MI who participated in the Myocardial Infarction and Mental Stress 2 (MIMS2) study. The protocol has been described before in detail (2). Participants with a documented history of MI in the previous eight months were recruited from the pool of patients admitted at Emory-affiliated hospitals in Atlanta, Georgia. MI case diagnosis was verified with medical record review based on standard criteria of troponin level increase, symptoms of ischemia, and changes in the electrocardiogram (ECG) or other evidence of myocardial necrosis. Inclusion criteria included being 18 to 60 years of age at the time of recruitment; presence of obstructive CAD was not required for inclusion. Each participant underwent a two-day protocol that included a blood draw, measured height and weight, and clinic tests of myocardial perfusion imaging following rest and mental and conventional stress procedures. A research nurse obtained sociodemographic, medical history and medication information, and participants completed questionnaires on behavioral, social, and health status information. Data for this study were collected between June 2011 and March 2015. The Emory University Institutional Review Board approved the protocol and all participants provided written informed consent.

Study Participants

Of the 313 MI cases, 14 were missing information on everyday discrimination and seven were missing information on MSIMI due to technical difficulties with testing. Thus, 295 participants were included in the analytic dataset. Those excluded from analyses had a mean single-photon emission computed tomography (SPECT) summed rest score of 7.8 (SD=7.6) compared to 3.7 (SD=6.3) among those included in analyses, indicating more resting perfusion defects among those excluded (p=0.0301). Also, those excluded were more likely to be in a higher income category compared to those included in analyses (p=0.0195).

Study Procedures

Mental Stress Procedure—Participants underwent a standardized mental stress test (7). After 30 minutes of resting in a quiet, dimly lit, temperature-controlled room, participants were given two minutes to make up a realistic story in which a close relative had been mistreated in a nursing home and three minutes to tell the story in front of a video camera and audience wearing white coats. Participants were told laboratory staff would be evaluating this presentation for content, quality, and duration. Blood pressure and heart rate were taken every five minutes during the 30-minute resting phase and every minute during the five-minute mental stress phase. The radioisotope for nuclear imaging, Technetium-99 (^{99m}Tc) sestamibi, was injected one minute after the start of the public speaking task.

Conventional Stress Procedure—Conventional stress testing took place on a separate day up to one week apart from the mental stress testing. This consisted of a standard Bruce protocol in which participants underwent treadmill exercise testing. The ECG, blood pressure and heart rate were continuously monitored. Once the participant attained 85% of their target heart rate, they were injected with a dose of ^{99m}Tc-sestamibi intravenously. Pharmacological stress testing with regadenoson was conducted among participants who were unable to exercise and ^{99m}Tc sestamibi was injected immediately after regadenoson injection. In our sample, 76% of men (N=112) and 66% of women (N=95) underwent exercise testing, and the remainder pharmacological stress testing. A physician was present during the study and the ECG and vital signs were continuously monitored.

Myocardial Perfusion Imaging—Three SPECT myocardial perfusion imaging scans were obtained: one at rest, one after mental stress, and one after conventional (primarily exercise) stress. Participants were scanned following injection of ^{99m}Tc sestamibi as described above, with a dose of 10 to 14 mCi for rest imaging and 30 to 40 mCi for stress imaging based on weight. Forty to 60 minutes after isotope injection, stress SPECT imaging was performed. Following standard nuclear cardiology protocols, antiischemic medications were withheld 24 hours before testing.

An experienced nuclear cardiologist performed visual interpretation of the imaging scans blinded to stress test type and other patient information. Each myocardial segment was scored from zero to four (zero = normal, one = possibly normal, two = definitely abnormal, three = severely abnormal, and four = no perfusion) which was used to create a summed stress score, summed rest score, and summed difference score (27). Presence of ischemia was defined as a summed difference score 3 for mental stress and 4 for exercise and pharmacological stress.

Measurement of Discrimination

The 10-item version of the Everyday Discrimination Scale (EDS) was used to assess exposure to everyday discrimination, or everyday occurrences of unfair treatment (28, 29). Sample items include being treated with less courtesy than other people, receiving poorer

service than other people at restaurants or stores, etc... without reference to race, gender or any other characteristic. For each item participants expressed frequency of occurrence using a 4-point Likert scale (one = "often", two = "sometimes", three = "rarely", four = "never"). Responses were summed and reverse coded for higher scores to represent more exposure to experiences of everyday discrimination, the resulting scores ranging from 10 to 40. The EDS has been widely used with good reliability and validity across racial/ethnic groups (30).

Other Study Measures

Demographic information included self-reported sex, race/ethnicity, age, educational attainment, and income. Educational attainment was assessed as years of education (continuous) and dichotomized by <12 years or 12 years. Household income was categorized (<\$35K, \$35-<\$75K, \$75K). Body mass index (BMI) was calculated as measured weight divided by the square of measured height (kg/m²). Previous medical history information included history of clinical cardiovascular risk factors (smoking, diabetes, hypertension, high blood cholesterol), type of previous MI (ST-Elevation Myocardial Infarction [STEMI] v. Non-ST-Elevation Myocardial Infarction [NSTEMI]), and left ventricular ejection fraction.

Depression was also measured using a validated self-report instrument. The Beck Depression Inventory-II (BDI-II), a 21-item inventory widely used in cardiac and noncardiac populations, was used to measure severity of depression symptoms.

Statistical Analysis

We calculated descriptive statistics for the sample by sex. We tested differences in these statistics by sex using chi-squared tests for categorical variables and t-tests for continuous variables. In order to examine the relationship between everyday discrimination and MSIMI (defined as the difference between summed rest and mental stress scores 3), and everyday discrimination and CSIMI (defined as the difference between summed rest and conventional stress scores 4), we ran generalized linear models with a logit link to derive odds ratios (OR) and 95% confidence intervals (CI) for MSIMI for each standard deviation increase in the EDS score. Covariates to be included in the models were considered a priori. Model 1 was adjusted for sociodemographic factors (race, sex, age, education, and income). Model 2 was further adjusted for cardiovascular risk factors (history of smoking, diabetes, hypertension, high blood cholesterol, and BMI) and indicators of previous MI severity (type of MI, ejection fraction, and SPECT imaging perfusion defects at rest). Model 3 further adjusted for depression symptom severity as this is a psychosocial factor often linked to discrimination and found to be a risk factor for MSIMI (12, 23, 31). While we considered a range of other potential mediators along with depression - including perceived stress, posttraumatic stress, and state and trait anger – depressive symptom severity was the most highly correlated with reports of everyday discrimination in a prior analysis from the MIMS2 cohort (32). Furthermore, we assessed sex differences with two methods: 1) we ran models 1–3 separately for men and women, and 2) examined interactions on the multiplicative scale by adding a product term to models with a logit link and on the additive scale by adding a product term to models with an identity link (33). All statistical analyses were conducted using SAS version 9.4.

Results

Table 1 shows descriptive characteristics of the analytic sample by sex. A greater proportion of women were Black (74.5%) compared to men (57.3%); the cohort primarily selfidentified as either Black or White, with only 17 identifying as other (7% of men (N=11))and 4% of women (N=6)). Women also had lower income than men. Additionally, women had a higher prevalence of diabetes, a higher BMI, and a higher left ventricular ejection fraction. Women and men were comparable on reports of everyday discrimination (women: mean 16.6 (SD 6.1); men: mean 17.2 (SD 6.0)), but women had higher scores on depression symptom severity. During rest, women had lower summed perfusion defect scores than men (women: mean 2.9 (range 0–32); men: mean 4.4 (range 0–30)), denoting fewer perfusion defects at rest compared with men. However, women had similar though slightly lower summed perfusion defect scores during mental stress (women: mean 4.1 (range 0-38); men: mean 4.9 (range 0-32)) and conventional stress (women: mean 4.9 (range 0-38); men: mean 5.3 (range 0-30)) compared to men. Therefore, women had higher summed difference scores for both MSIMI (mean 1.2 (range 0–13)) and CSIMI (mean 2.1 (range 0–16)) compared to men (mean 0.59 (range 0-10), mean 1.2 (range 0-14), respectively), indicating that women experience a more severe change in perfusion defects resulting from mental and conventional stress compared to at rest than men.

Of the 295 participants, 16.6% experienced MSIMI (N=49), with 18.2% of women (N=27) experiencing MSIMI and 14.7% of men (N=22). Although modeled continuously in all analyses, for descriptive purposes only we present the association between reports of everyday discrimination and the prevalence of MSIMI using approximate tertiles in Figure 1. From Figure 1 we see that as the everyday discrimination score increased, there was an increase in the proportion of participants experiencing MSIMI, showing a graded response. While approximately 11% of those in the lowest tertile of discrimination scores experienced MSIMI (N=11/104), 25% in the highest tertile experienced MSIMI (N=23/92).

Table 2 presents the results of the overall and sex-stratified models assessing the relationship between everyday discrimination and MSIMI, as well as results of interaction testing by gender. In the model adjusting for sociodemographic factors, the odds of MSIMI increased 1.4-fold (OR: 1.40, 95% CI: 1.07 - 1.83) for each standard deviation increase in the EDS score. This result was not fundamentally changed, after adjusting for cardiovascular risk factors and MI severity indicators, as well as depression symptom severity. Therefore, neither cardiovascular risk variables nor depression explained the association between discrimination and MSIMI.

In sex-stratified models, the association was more robust in women compared to men. In fully adjusted models, the OR of MSIMI in women was 1.96 (95% CI: 1.13 - 3.38) for each standard deviation increase in the EDS score; for men, the corresponding OR was 1.40 (95% CI: 0.80 - 2.44). However, formal testing for multiplicative and additive interactions indicated that the joint effect of everyday discrimination and sex was not greater than either the product or the sum of the individual effects, respectively.

Table 3 presents the results of the overall and sex-stratified models assessing the relationship between everyday discrimination and CSIMI, as well as results of interaction assessments. Both overall and sex-specific models did not provide evidence of a relationship between everyday discrimination and CSIMI.

Exploratory Analyses

Because the associations between everyday discrimination and MSIMI appear more robust in fully adjusted models with women relative to men, and the majority of women in our cohort were Black, we ran exploratory models examining the relationship between everyday discrimination and MSIMI among Black women in order to describe this association for this subgroup. Black women are the largest sub-group in this study and experience the most outcomes, and therefore we are able to appropriately examine the association of interest in this sub-group (unlike for non-Black women of which there were 37 in this sample, four of which experienced MSIMI). Of the 108 Black women included in analyses, 21.3% experienced MSIMI (N=23). After adjustment for sociodemographic and clinical risk factors, as well as depression symptom severity, the odds of MSIMI increased 2.5-fold (OR: 2.45, 95% CI: 1.27 - 4.71) for each standard deviation increase in the discrimination score among Black women.

Discussion

In this sample of men and women with recent MI, we found more frequent experiences of everyday discrimination were associated with an elevated risk for MSIMI after controlling for sociodemographic and clinical risk factors along with indicators of MI severity. Associations were especially robust among women. Even after adjustment for depression symptom severity, one standard deviation increase in the EDS score was associated with an OR for MSIMI of 1.96 (95% CI: 1.13 - 3.38) among women, nearly a two-fold increase, and 1.40 (95% CI: 0.80 - 2.44) among men. However, the interaction between everyday discrimination and sex was not statistically significant.

We found that experiences of discrimination were related to MSIMI and not to CSIMI. Findings from laboratory and human cohorts suggest that different mechanisms are involved in myocardial ischemia induced by mental stress compared to other forms of stress like exercise. Both mental stress and physical stress trigger an increase in myocardial workload (increased heart rate and blood pressure) and an increase in oxygen demand [1, 3]. However, while with mental stress there is also an increase in systemic vascular resistance and microvascular constriction, with exercise and pharmacological stress there is systemic vasodilation [1, 3, 31-33]. By these mechanisms, mental stress can impair coronary perfusion, potentially inducing myocardial ischemia even in the absence of coronary stenoses. In previous studies we have shown that these pathways for MSIMI are especially relevant for women (8, 34).

Previous studies have noted that reports of discrimination are related to mechanistic pathways that could explain vulnerability to MSIMI, including attenuated recovery of endothelial function in response to acute stress, increased visceral fat and increased inflammatory markers [18, 22, 23]. These studies also identified sex differences, with

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associations generally stronger among women than among men. Men in our study reported a similar exposure to everyday discrimination compared to women. However, women could be more vulnerable to the physiological effects of psychosocial stressors, as suggested by previous literature (13, 35, 36). Rodent and human studies have shown greater and prolonged hypothalamic-pituitary-adrenal (HPA) axis activation as well as greater potential for hyperarousal in response to emotional stimuli in females, which may reflect sex-related differences in signaling, structure, and function related to stress response (37). There is also evidence that women sustain higher levels of interleukin-6 before and after mental stress (38). Such dysregulation of neuroendocrine and immune responses to stress can facilitate microvascular dysfunction and peripheral vasoconstriction, which are emerging mechanisms of MSIMI, and more prevalent in women (2, 8, 39). Moreover, rumination over stressful events, to which women have a greater tendency, can result in prolonged duration of cortisol activation, hyperarousal and greater HPA axis response to stress (37, 40, 41). Women may cope in ways that make them especially sensitive to interpersonal stressors (42).

It should also be noted that women in this study were more likely to be Black compared to men, and therefore potentially exposed to social disadvantage and discrimination both for sex and for race (43). In exploratory models, adjusted associations between discrimination and MSIMI were very pronounced among Black women. We found it important to examine CVD risk among Black women in particular since recent research shows the prevalence of CVD among black women aged 35–44 years exceeded that of White women and Black and White men of the same age (44). Future research with larger subgroups of women and men identifying as White and other races are needed to further explore race-sex differences in MSIMI prevalence and associations between discrimination and MSIMI within these subgroups.

There are limitations to this study worth noting. The prevalence of MSIMI in this study is lower than that observed in others, such as the REMIT study, which used echocardiography with ischemia defined as left ventricular wall motion abnormalities, which can give a MSIMI prevalence upwards of 60% (6). Conversely, myocardial perfusion imaging was used in the current study as this is the gold standard for ischemia assessment, and yields a consistently lower prevalence of MSIMI (45-47). It is also possible that the older age of participants in the REMIT study (mean=63.81), or the lower proportion of women in the REMIT study (17%) may somewhat account for the MSIMI prevalence differences between studies. Another potential limitation worth discussing is the specific mental stress laboratory task used in the MIMS2 study, which while related to mistreatment, is fairly generic and designed to evoke a response from a wide range of sociodemographic groups. It is possible that a stressor focused more explicitly on discrimination or oppression on the basis of sociodemographic factors (i.e., sex, race/ethnicity, social class), might have evoked a stronger physiological stress response among those with more experiences of discrimination. Previous studies have shown that the prevalence of MSIMI varies depending on the type and duration of mental tasks being performed (48). Therefore, we may have seen stronger associations between discrimination and MSIMI with a more discrimination or oppressionbased stressor task, the more generic stressor used in MIMS2 potentially biasing our results toward the null.

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Further, although consistent with those reported in other studies, everyday discrimination scores in our cohort are fairly low (29, 49–51). This provides evidence in support of arguments that self-reported measures of interpersonal discrimination alone may not capture the full extent to which discrimination can affect health outcomes. For instance, institutional discrimination – systemic policies, practices, and norms that promote unequal access to opportunities such as employment, education, housing, and health care – may sustain persistent disparities in exposure to many stressors which may affect health (including work stress, violence, financial instability, and relationship stress) (31). Moreover, vigilance or anticipating discrimination is an everyday stressor that may better capture the worry and rumination related to discrimination. The latter can in turn lend to hyperarousal or dysregulation of physiological systems (52). Future studies should consider multiple domains of discrimination to more fully capture the effect of everyday discrimination on MSIMI and to inform future interventions that address the burden of MSIMI.

There are also important strengths to this study. To our knowledge, this is the first study to examine the relationship between discrimination and MSIMI. We used well-validated questionnaires and the gold-standard assessment of MSIMI, and controlled for a range of relevant psychological, behavioral and clinical confounders. By additionally exploring sex differences and focusing on a population with recent MI, this study offers important exploration of potential mechanisms explaining worse morbidity and mortality after MI among women (53). Also, the large number of Black patients and the nearly equal numbers of men and women makes this study well-suited to explore these questions.

In sum, in a sample of young and middle-aged men and women with a recent MI, experiences of everyday discrimination were associated with risk of MSIMI. There were no observed associations with conventional stress ischemia. The risk of MSIMI tended to be greater among women, suggesting that this emotional stressor may trigger a response to which women are particularly vulnerable, however we did not find a statistically significant interaction between everyday discrimination and sex. These findings further implicate the role of psychosocial exposures explaining worse morbidity and mortality after MI. This study adds important new data to a growing field pointing to discrimination as a key chronic stressor linked to CVD risk. Our results highlight the importance of assessing discrimination in post-MI patients, potentially for women in particular, and support the incorporation of this measure in future intervention efforts to ameliorate risk in this population.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

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Abbreviations

BDI-II	Beck Depression Inventory-II				
BMI	Body mass index				
CAD	Coronary artery disease				
CHD	Coronary heart disease				
CI	Confidence interval				
CSIMI	Conventional stress-induced myocardial ischemia				
CVD	Cardiovascular disease				
ECG	Electrocardiogram				
EDS	Everyday Discrimination Scale				
MI	Myocardial infarction				
MIMS2	Myocardial Infarction and Mental Stress 2				
MSIMI	Mental stress-induced myocardial ischemia				
NSTEMI	Non-ST-Elevation Myocardial Infarction				
OR	Odds ratio				
SPECT	Single-photon emission computed tomography				
STEMI	ST-Elevation Myocardial Infarction				
^{99m} Tc	Technetium-99				

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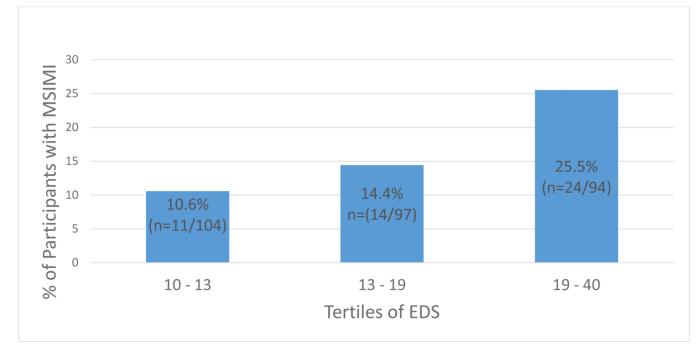


Figure 1.

Percent of Participants with Mental Stress-Induced Myocardial Ischemia (MSIMI) by Tertiles of EDS with outliers removed (n=2); Abbreviations: MSIMI: Mental Stress-Induced Ischemia, EDS: Everyday Discrimination Score

Table 1.

Descriptive characteristics of participants stratified by gender among post-MI participants (N= 295) in the Myocardial Infarction and Mental Stress 2 Study (MIMS2).

	Women (n=145)	Men (n=150)	p-value
Sociodemographic Factors			
Black race, n (%)	108 (74.5%)	86 (57.3%)	0.002
Age, years, mean (SD)	50.6 (7.3)	51.1 (6.0)	0.56
Education 12 Years, n (%)	85 (58.6%)	89 (59.3%)	0.90
Income			
<\$35K	78 (53.8%)	61 (40.7%)	
\$35 - <\$75K	35 (24.1%)	38 (25.3%)	
\$75K	23 (15.9%)	37 (24.7%)	0.02
Clinical Risk Factors & MI Severity			
History of Smoking, n (%)	81 (55.9%)	81 (54.0%)	0.75
History of Diabetes, n (%)	55 (37.9%)	40 (26.7%)	0.04
History of Hypertension, n (%)	120 (82.8%)	119 (79.3%)	0.45
History of High Blood Cholesterol, n (%)	114 (78.6%)	112 (81.3%)	0.56
BMI, kg/m2, mean (SD)	32.3 (8.0)	30.0 (5.8)	0.004
STEMI, N (%)	38 (26.2%)	48 (32.0%)	0.27
Ejection Fraction, %, mean (SD)	53.0 (11.1)	48.6 (12.5)	0.002
SPECT Summed Rest Score, mean (SD)	2.9 (5.7)	4.4 (6.8)	0.04
SPECT Summed Mental Stress Score, mean (SD)	4.1 (6.7)	4.9 (7.0)	0.33
SPECT Summed Physical Stress Score, mean (SD)	4.9 (6.7)	5.3 (7.2)	0.57
Psychosocial Factors			
BDI Score, mean (SD)	14.1 (11.1)	10.6 (9.7)	0.005
Everyday Discrimination Score, mean (SD)	16.6 (6.1)	17.2 (6.0)	0.88
Myocardial Ischemia Severity Score			
Summed Difference Score for Mental Stress, mean (SD)	1.2 (2.4)	0.59 (1.5)	0.01
Summed Difference Score for Exercise or Pharmacological Stress, mean (SD)	2.1 (3.3)	1.2 (2.6)	0.006

Abbreviations: SD: Standard deviation; BMI: Body mass index; BDI: Beck Depression Inventory.

Table 2.

Adjusted Association between Experiences of Everyday Discrimination (per 1-Standard Deviation in Total Score Increment) and Mental Stress Induced Myocardial Ischemia (N=295) in the Myocardial Infarction and Mental Stress 2 Study (MIMS2) Overall and By Sex.

				Interaction Assessment		
Model	Overall OR (95% CI)	Women OR (95% CI)	Men OR (95% CI)	Multiplicative p- value	Additive p- value	
Adjusted for sociodemographic factors *	1.40 (1.07 – 1.83)	1.43 (1.00 – 2.04)	1.36 (0.89 – 2.08)	0.82	0.92	
Adjusted for the above plus cardiovascular risk factors and MI severity indicators **	1.46 (1.08 – 1.98)	1.70 (1.08 – 2.66)	1.38 (0.85 – 2.25)	0.56	0.62	
Adjusted for the above plus depression symptom severity ***	1.57 (1.10 – 2.23)	1.96 (1.13 – 3.38)	1.40 (0.80 – 2.44)	0.51	0.60	

*Race, age, educational attainment, income

** History of smoking, diabetes, hypertension, and high blood cholesterol, BMI, type of MI, ejection fraction, SPECT summed rest score

*** Beck Depression Inventory score

Table 3.

Adjusted Association between Experiences of Everyday Discrimination (per 1-Standard Deviation in Total Score Increment) and Exercise or Pharmacologically-Induced Myocardial Ischemia (N=295) in the Myocardial Infarction and Mental Stress 2 Study (MIMS2) Overall and By Sex.

	Interaction Assessment				
Model	Overall OR (95% CI)	Women OR (95% CI)	Men OR (95% CI)	Multiplicative p- value	Additive p- value
Adjusted for sociodemographic factors *	1.01 (0.79 – 1.28)	1.03 (0.76 – 1.41)	0.99 (0.66 – 1.48)	0.62	0.51
Adjusted for the above plus cardiovascular risk factors and MI severity indicators	0.95 (0.74 – 1.22)	1.01 (0.72 – 1.42)	0.91 (0.59 – 1.40)	0.37	
Adjusted for the above plus depression symptom severity ***	0.86 (0.64 - 1.17)	0.76 (0.49 – 1.20)	1.02 (0.62 – 1.66)	0.45	0.64

Race, age, educational attainment, income

** History of smoking, diabetes, hypertension, and high blood cholesterol, BMI, type of MI, ejection fraction, SPECT summed rest score

*** Beck Depression Inventory score