SUPPLEMENTAL MATERIALS TABLE OF CONTENTS

Supplemental Table 1: P-values for interaction by the following terms on associations between symptoms and subsequent myocardial infarction

	Prior myocardial infarction or revascularization	eGFR (continuous)	eGFR (categorical)	Proteinuria (continuous)	Sex
Chest pain	0.18	0.92	0.97	0.85	0.13
Shortness of breath	0.69	0.46	0.70	0.15	0.10
Inability to climb stairs	0.18	0.96	0.44	0.48	0.95

Adjusted for sex, self-reported race/ethnicity, time-updated age, time-updated prior myocardial infarction or revascularization, prior stroke, diabetes, systolic and diastolic blood pressure, heart failure, atrial fibrillation, BMI, COPD, eGFR, current smoking status, log-transformed proteinuria, use of coronary vasodilators, calcium channel blockers, and beta blockers.

Supplemental Table 2: Additive interactions between absence or presence of chest pain, shortness of breath, and inability to climb stairs with Relative Excess Risk due to Interaction

Symptoms	Relative excess risk due to interaction (95% CI)
Chest Pain (any vs none) and Shortness of Breath (any vs none)	-0.00 (-0.84, 0.83)
Chest Pain (any vs none) and Inability to Climb Stairs (any vs none)	-0.18 (-1.64, 1.29)
Shortness of Breath (any vs none) and Inability to Climb Stairs (any vs none)	-0.80 (-2.05, 0.44)

Supplemental Table 3: Associations between categorical time-updated symptoms of atherosclerosis and myocardial infarction, stratified by diabetes status

	Hazard Ratio (95% Confidence Interval)	
	Adjusted, without diabetes	Adjusted, with diabetes
Chest Pain		
No bother	1.0 (Referent)	1.0 (Referent)
Mild bother	1.51 (0.95, 2.39)	1.22 (0.91, 1.64)
Moderate or worse bother	**2.48 (1.50, 4.11)**	**1.46 (1.03, 2.07)**
P-value for trend across symptom severity	< 0.001	0.02
Shortness of Breath		
No bother	1.0 (Referent)	1.0 (Referent)
Mild bother	**1.55 (1.01, 2.36)**	**1.3 (1.01, 1.67)**
Moderate or worse bother	**2.68 (1.79, 4.03)**	1.00 (0.75, 1.33)
P-value for trend across symptom severity	< 0.001	0.72
Inability to climb stairs		
No limitation	1.0 (Referent)	1.0 (Referent)
Mild limitation	1.49 (0.97, 2.27)	**1.43 (1.01, 2.02)**
Severe limitation	**1.77 (1.13, 2.77)**	**1.94 (1.38, 2.72)**
P-value for trend across symptom severity	0.01	< 0.001

Adjusted for sex, self-reported race/ethnicity, time-updated age, prior myocardial infarction or revascularization, prior stroke, diabetes, systolic and diastolic blood pressure, heart failure, atrial fibrillation, BMI, COPD, eGFR, current smoking status, log-transformed proteinuria, use of coronary vasodilators, calcium channel blockers, and beta blockers.
Bold font indicates statistically significant confidence interval

Supplemental Table 4: Associations between categorical time-updated symptoms of atherosclerosis and myocardial infarction, additionally adjusted for changes in symptoms since the last visit (N=3909)

	Hazard Ratio (95% Confidence Interval)			
	Model 1	**Model 2**	**Model 3**	**Model 4: additionally adjusted for preceding symptoms**
Chest Pain				
No bother	1.0 (Referent)	1.0 (Referent)	1.0 (Referent)	1.0 (Referent)
Mild bother	**1.63 (1.28, 2.08)**	**1.36 (1.06, 1.74)**	**1.30 (1.01, 1.67)**	**1.37 (1.06, 1.78)**
Moderate or worse bother	**2.22 (1.67, 2.94)**	**1.80 (1.35, 2.40)**	**1.70 (1.27, 2.27)**	**1.97 (1.40, 2.76)**
Shortness of Breath				
No bother	1.0 (Referent)	1.0 (Referent)	1.0 (Referent)	1.0 (Referent)
Mild bother	**1.62 (1.31, 2.00)**	**1.38 (1.11, 1.71)**	**1.37 (1.10, 1.70)**	**1.45 (1.16, 1.82)**
Moderate or worse bother	**1.80 (1.44, 2.26)**	**1.37 (1.08, 1.74)**	**1.33 (1.05, 1.69)**	**1.55 (1.18, 2.04)**
Inability to climb stairs				
No limitations	1.0 (Referent)	1.0 (Referent)	1.0 (Referent)	1.0 (Referent)
Mild limitations	**1.86 (1.43, 2.43)**	**1.47 (1.12, 1.92)**	**1.44 (1.10, 1.89)**	**1.51 (1.15, 1.99)**
Severe limitations	**3.00 (2.32, 3.87)**	**1.95 (1.48, 2.56)**	**1.89 (1.44, 2.49)**	**2.08 (1.54, 2.82)**

Model 1: Adjusted for sex, self-reported race/ethnicity, and time-updated age
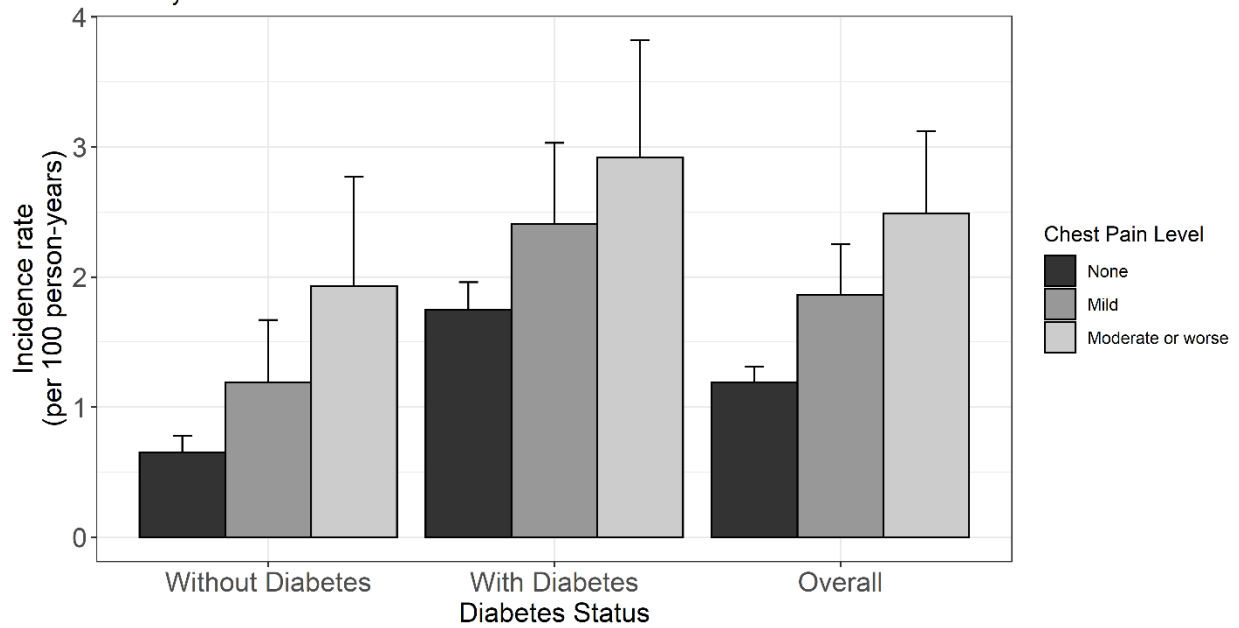
Model 2: Adjusted for the components of Model 1 plus time-updated prior myocardial infarction or revascularization, prior stroke, diabetes, systolic and diastolic blood pressure, heart failure, atrial fibrillation, BMI, COPD, eGFR, current smoking status, and log-transformed proteinuria

Model 3: Adjusted for the components of Model 2 plus time-updated use of coronary vasodilators, calcium channel blockers, and beta blockers

Model 4: Adjusted for the components of Model 3 plus changes in symptoms from the visit preceding the analyzed visit
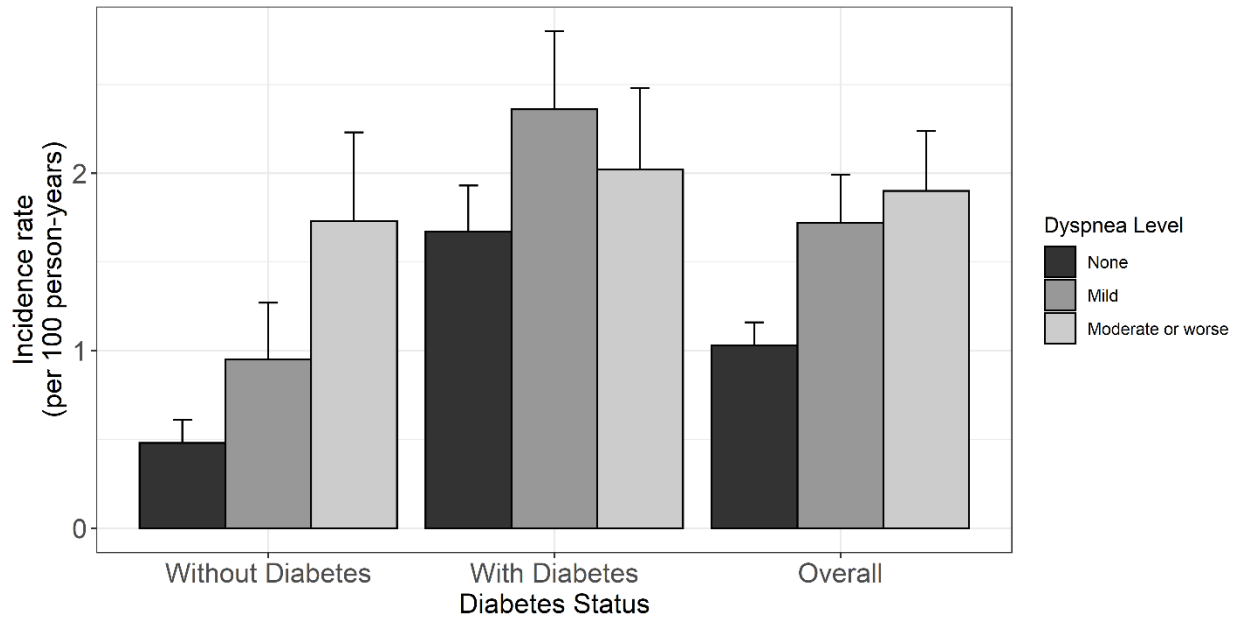
Bold font indicates statistically significant confidence interval

Supplemental Figure 1: Incidence rate of myocardial infarction by chest pain severity and diabetes status
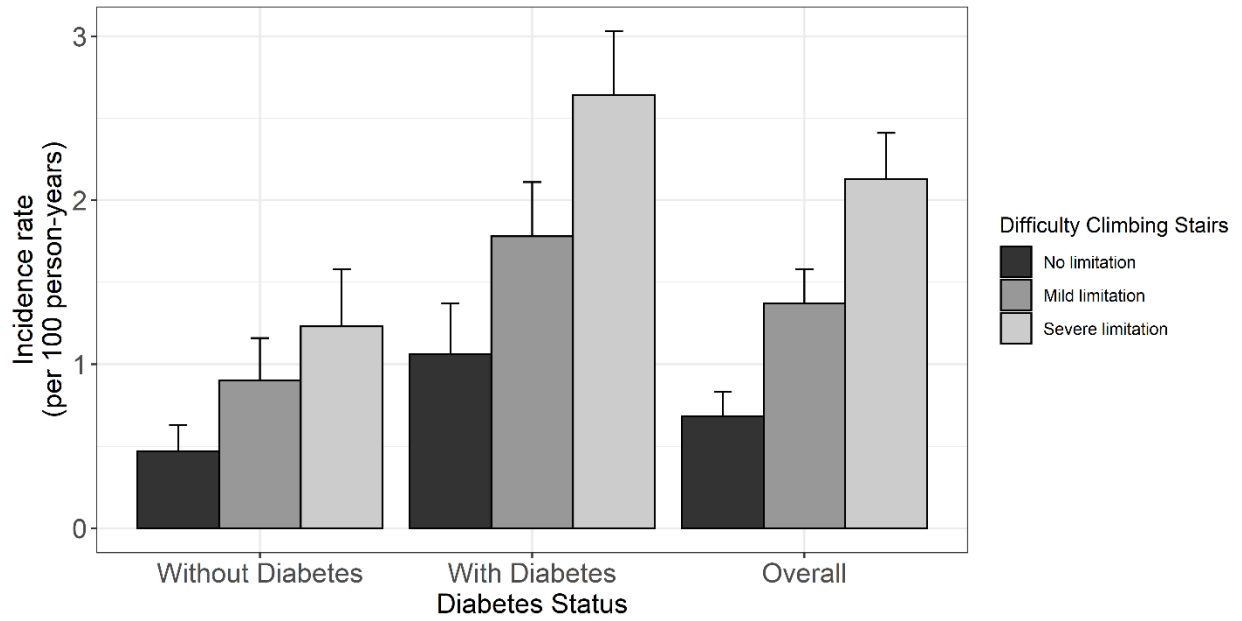

Error Bars represent the upper bounds of the 95% confidence interval.

Supplemental Figure 2: Incidence rate of myocardial infarction by shortness of breath severity and diabetes status

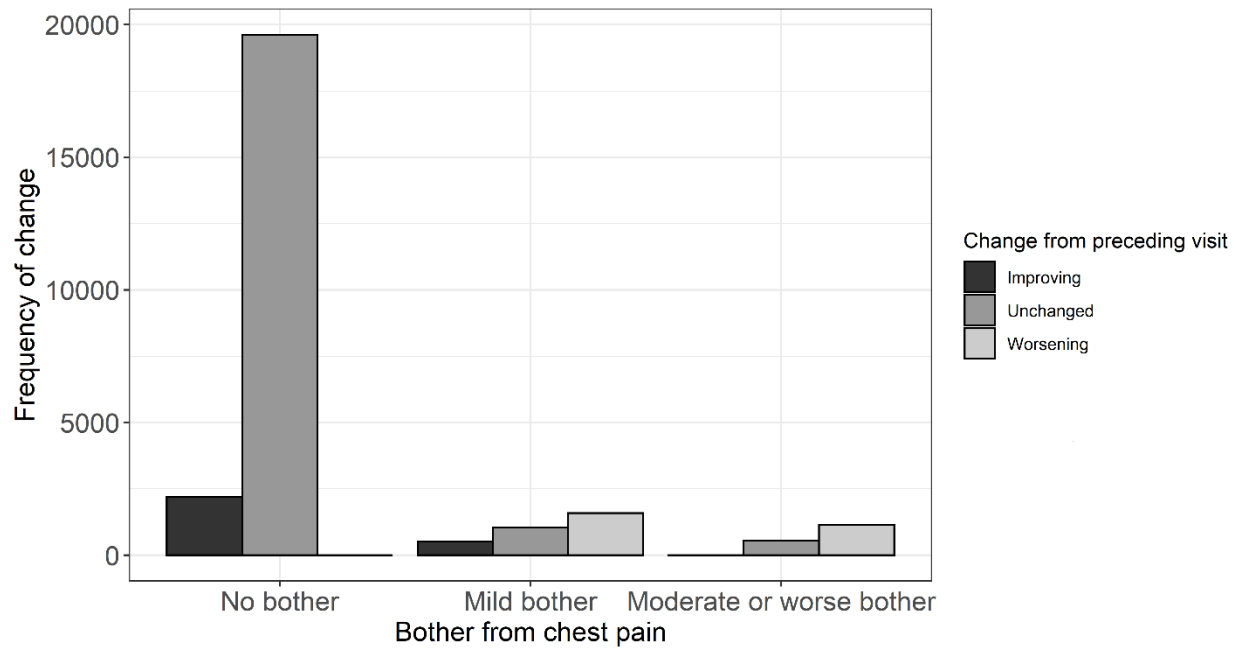Error Bars represent the upper bounds of the 95% confidence interval.

Supplemental Figure 3: Incidence rate of myocardial infarction by limitation climbing stairs and diabetes status
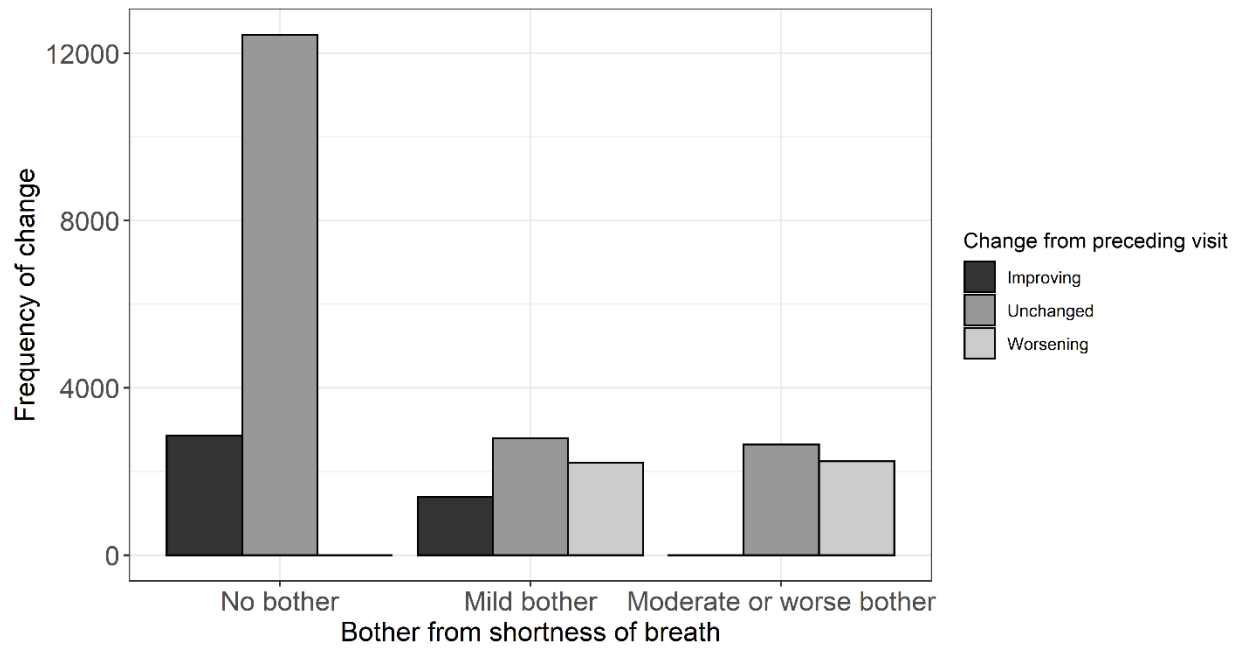
Error Bars represent the upper bounds of the 95% confidence interval.

Supplemental Figure 4: Frequency of changes in chest pain from preceding annual visit

Supplemental Figure 5: Frequency of changes in shortness of breath from preceding annual visit

Supplemental Figure 6: Frequency of changes in difficulty climbing stairs from preceding annual visit