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Long-Term Trends in Short-Term Outcomes in Acute Myocardial Infarction

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Abstract

Background—The objectives of this study were to examine the magnitude of, and 20-year trends in, age differences in short-term outcomes among men and women hospitalized with acute myocardial infarction (AMI) in central Massachusetts.

Methods—The study population consisted of 5,907 male and 4,406 female residents of the Worcester, MA, metropolitan area hospitalized at all greater Worcester medical centers with AMI between 1986 and 2005.

Results—Overall, among both men and women, older patients were significantly more likely to have developed atrial fibrillation, heart failure, and to have died during hospitalization and within 30 days after admission compared to patients <65 years. Among men, age differences in the risk of developing atrial fibrillation have widened over the past 2 decades, while differences in the risk of developing cardiogenic shock have narrowed for men 75 years and older as compared with those <65 years. Among women, age differences in the risk of developing these major complications of AMI have not changed significantly over time. Age differences in short-term mortality have remained relatively unchanged over the past 20 years in both sexes, though individuals of all ages have experienced declines in short-term death rates over this period.

Conclusions—Elderly men and women are more likely to experience adverse short-term outcomes after AMI and age differences in short-term mortality rates have remained relatively unchanged in both sexes over the past 20 years. More targeted treatment approaches during

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hospitalization for AMI and thereafter are needed for older patients to improve their prognosis. Word count: 248

Keywords

Acute myocardial infarction; hospital complications; hospital mortality; age and sex differences

Introduction

Over the past several decades, dramatic advances in the medical management of patients hospitalized with acute myocardial infarction (AMI) have been accompanied by reductions in in-hospital clinical complications and short-term death rates.^{1,2} Despite encouraging declines in population death rates from coronary heart disease (CHD) and hospital mortality from AMI in the U.S. since the late 1960s,³ several groups, including older individuals, women, and patients with multiple comorbidities remain at increased risk for adverse outcomes after hospitalization for AMI.

Previous studies examining age and sex differences in the development of hospital complications and risk of dying in the setting of AMI have shown inconsistent results.^{4–9} While some studies found that older persons had higher hospital complication and mortality rates compared with younger individuals,^{3–6,10,11} other studies have not.^{7–9,12} Despite national interest, few studies have examined age-specific differences in important in-hospital complications and short-term death rates separately for men and for women. Further, there is a lack of data from a broad community-wide perspective that has examined changing, and contemporary, associations between age and short-term outcomes in men and women hospitalized with AMI.^{3,13}

The objectives of this study were to examine relatively contemporary age differences, and 20-year trends (1986–2005) therein, in the development of important in-hospital clinical complications and short-term death rates among residents of central Massachusetts hospitalized with AMI, separately for men and women. Data from the population-based Worcester Heart Attack Study were used for this investigation.^{14–16}

Methods

The Worcester Heart Attack Study is an ongoing clinical/epidemiologic investigation that is examining long-term trends in the incidence, hospital, and post-discharge case-fatality rates of AMI among residents of the Worcester metropolitan area hospitalized at all 16 greater Worcester medical centers in 15 biennial periods between 1975 and 2005.^{14–16} Fewer hospitals (n = 11) have been included during recent study years due to hospital closures, mergers, and conversion to chronic care facilities. In brief, computerized printouts of patients discharged from all greater Worcester hospitals with possible AMI were obtained and several International Classification of Disease (ICD) codes in which cases of AMI may have been diagnosed were reviewed. Cases of possible AMI were independently validated according to predefined criteria for AMI which included a suggestive clinical history, serum enzyme elevations, and serial electrocardiographic findings during hospitalization consistent with the presence of AMI; at least 2 of these 3 criteria needed to be present for an AMI to have occurred. Patients with an initial AMI (incident event), as well as those with a prior history of AMI, based on the review of information contained in hospital medical records, were included in our study population. Residents of the Worcester metropolitan area who satisfied these criteria were included in the present investigation.

Data Collection

The hospital medical records of greater Worcester residents with confirmed AMI were reviewed by trained study physicians and nurses who abstracted information about patient's demographic characteristics, medical history, clinical presentation, hospital treatment approaches, and hospital discharge status. Age was categorized into 3 strata of <65 years, 65–74 years, and ≥75 years. The principal study outcomes included the development of important in-hospital clinical complications and total hospital and 30 day mortality. Atrial fibrillation (AF) included the documentation of new onset AF in the hospital medical record or occurrence of typical electrocardiographic changes consistent with this diagnosis.¹⁷ Heart failure was indicated by clinical or radiographic evidence of pulmonary edema or bilateral basilar rales with an S3 gallop¹⁸ while cardiogenic shock was defined according to previously described criteria.¹⁹ Since the average length of stay for patients hospitalized with AMI has declined during the years under study,²⁰ we also examined 30-day postadmission death rates as a secondary study outcome. All patients discharged from greater Worcester hospitals after AMI were followed through a search of death certificates to determine their vital status.

Data Analysis

All analyses were performed separately for men and for women. Differences in baseline demographic, clinical characteristics, and hospital therapies in relation to patient age were compared using the chi-square test for categorical variables and Kruskal-Wallis test for continuous variables.

Using logistic regression models, we estimated odds ratios (ORs) and 95% confidence intervals (95% CIs) for the association between age and various hospital outcomes. Patients with prevalent disease, namely those who had the condition being examined previously diagnosed based on the review of information contained in hospital medical records since patients were never directly contacted as part of this study, were excluded from models examining the development of incident (initial events) cases of several hospital clinical complications (508 men and 471 women with prior AF, and 950 men and 1,155 women with prior heart failure). Potential confounding factors included in our regression models were selected on the basis of the findings from prior studies and on their clinical importance; these variables included race, marital status, comorbidities, AMI order (initial (incident) vs. prior (recurrent) event), type (Q wave vs. non-Q wave), duration of pre-hospital delay following the onset of acute coronary symptoms, and length of hospital stay. Because information on body mass index was not collected until 1995, and information on acute symptoms and whether the AMI was a non-ST-segment elevation myocardial infarction (NSTEMI) or an ST-segment elevation myocardial infarction (STEMI) was not recorded until 1997, these variables were not included in our regression models. Study year was grouped into 5 two-year periods (1986/1988, 1990/1993, 1995/1997, 1999/2001, and 2003/2005) for ease of analysis. In all regression models, patients <65 years served as the reference category. Interaction terms between age and study period were used to examine whether age differences in the principal study outcomes changed significantly over time. Likelihood ratio tests were used to compare models with and without interaction terms. For 30-day death rates after hospital admission, similar multivariable adjusted Cox proportional hazard models yielded estimated hazards ratios (HRs) and 95% CIs. The Institutional Review Board at the University of Massachusetts Medical School approved this study.

Results

Patient Characteristics

The study sample consisted of 10,313 residents of the Worcester metropolitan area (5,907 men and 4,406 women) hospitalized with validated AMI at all greater Worcester medical centers in 11 study years between 1986 and 2005. Overall, women were considerably older, on average, than men (75 years vs. 66 years). While 43% of men were aged <65 years, only 19% of women were in this age category; in contrast, 56% of women were 75 years and older compared with 31% of hospitalized men; relatively similar differences between the sexes according to age were observed in analyzing data from patients hospitalized in the 2 most recent periods under study (2003/2005).

During the past 20 years, among both men and women, age was strongly associated with several demographic and clinical characteristics (Table 1). In men and in women, the proportion of patients with a history of AF, heart failure, hypertension, diabetes, and stroke was higher in older patients. The proportion of patients with an initial, Q-wave MI, and STEMI was lower in older patients in both sexes. Heart rate, serum glucose, and creatinine levels were higher, on average, in older patients in both sexes; diastolic blood pressure, serum levels of total and LDL cholesterol, and patient's body mass index were, however, lower in older women and men (Table 1). A relatively similar distribution of patient characteristics, stratified according to age, was observed when we examined these characteristics among patients included in the 2 most recent cohorts (2003/2005).

Age Differences in Hospital Clinical Complications and Death Rates, and Trends over Time, among Men

Between 1986/1988 and 2003/2005, the likelihood of developing AF increased slightly in men aged 65–74 years (16% vs. 19%), whereas the odds of developing AF decreased slightly in younger (10% vs. 8%) and older men (26% vs. 24%) (Figure 1) (p <05). On the other hand, the incidence rates of new onset heart failure declined significantly over time in all age groups (19% vs. 12% for men<65 years; 34% vs. 27% for men 65–74 years; 48% vs. 40% for men \geq 75 years) (p<05). The frequency of cardiogenic shock decreased over time in men \geq 75 years (15% vs. 5%) (p<05), but did not significantly change among younger patients. Declining in-hospital death rates were observed for men of all ages over the past two decades: \geq 75 years (30% in 1986/88 vs. 13% in 2003/05), 65–74 years (14% vs. 8%); and <65 years (7% vs. 2%) (p<05) (Figure 1).

The significant interaction between age and study period (p = .04) indicated that age differences in the risk of developing AF have changed significantly over time (Table 2); in 1986/1988, the adjusted ORs for AF were 1.61 and 2.83 for men 65–74 years and men ≥ 75 years, respectively; in 2003/2005 these multivariable adjusted ORs were 3.73, and 4.60, respectively, compared with men <65 years. In examining 20 year trends in the risk of developing cardiogenic shock, the significant interaction between age and study period (p = .02) suggest that age differences in the risk of developing this clinical complication have changed significantly over time (Table 2); in 1986/1988, the adjusted ORs for cardiogenic shock were 0.75 and 2.19 for men 65–74 years and men ≥ 75 years, respectively; in 2003/2005 these ORs were 1.86, and 1.10, respectively, compared with men<65 years.

On the other hand, age differences in the risk of developing heart failure, and in-hospital and 30-day mortality, have not changed significantly over time (Table 2). Overall, older men were more likely to have developed heart failure during hospitalization than men <65 years. Similarly, older men of all ages were significantly more likely to have died during hospitalization and during the first 30 days after hospital admission compared with men <65 years (Table 2).

Age Differences in Hospital Clinical Complications and Death Rates, and Trends over Time, among Women

Between 1986/1988 and 2003/2005, the incidence rates of AF decreased in women <65 years (10% vs. 7%) and women \geq 75 years (33% vs. 22%), but remained relatively unchanged in women 65–74 years (Figure 2). The incidence rates of new onset heart failure and cardiogenic shock decreased in all age groups over time (Figure 2). Declining inhospital death rates were observed for women of all ages over the past two decades with the extent of decline ranging from 41% to 57% in the 3 age strata examined (Figure 2).

Age differences in hospital clinical complications and short-term mortality have not, however, changed significantly among women over the past 20 years (Table 2). Overall, older women were significantly more likely to have developed AF during hospitalization for AMI than women < 65 years (adjusted ORs were 1.74 and 3.09 for women 65–74 years and \geq 75 years, respectively). Older women were significantly more likely to have developed heart failure after AMI compared with younger women (adjusted ORs were 1.61 and 2.26 for women 65–74 years and women \geq 75 years, respectively). Compared with women <65 years, older women were more likely to have died during hospitalization as well as during the 30 days following hospital admission (Table 2).

Discussion

In this study of more than 10,000 residents of a large central New England metropolitan area hospitalized with AMI, older men and women were more likely to have developed AF and heart failure, and were more likely to have died during hospitalization and during the first 30-days after admission, compared with patients <65 years. Older men were also more likely to have developed cardiogenic shock compared with younger men. Among men, age differences in the risk of developing AF and cardiogenic shock have widened over time. Encouragingly, we noted a steady improvement in the majority of hospital outcomes examined in most age groups over the 20 year period under study, with a particularly marked improvement observed in the risk of developing cardiogenic shock in elderly but not in younger patients. While unknown, these latter findings may be due to the changing characteristics of patients hospitalized with AMI over time and/or to the more frequent use of cardiac catheterization and PCI in elderly patients.

Our results are consistent with the findings from previous studies which have shown that older patients hospitalized with AMI have a worse prognosis than younger patients.^{1,2,5,15} Older patients are more likely to have additional comorbidities present at the time of hospitalization for AMI which may increase their risk of developing clinically significant hospital complications and dying. Previous studies have shown that older patients are less likely to be treated with evidence-based cardiac medications and interventional procedures,^{4,21} which may have contributed to their greater risk of dying in the short-term. Other factors such as prolonged delay in seeking medical care,^{5,6} limited health care access, cognitive impairment, and frailty may also have played a role in the less favorable prognosis observed in older patients.

We found that age differences in the risk of developing new onset AF during hospitalization for AMI have widened during the past 20 years for men. On the other hand, differences in the risk of developing cardiogenic shock between men 65–74 years and men <65 years have widened over time but have narrowed for men \geq 75 years. Our findings also showed that, despite the fact that the overall in-hospital death rates among patients with AMI have decreased from 17% in 1986/1988 to 9% in 2003/2005, age differences in short-term mortality have remained relatively unchanged over time among both men and women; the elderly remain at higher risk for dying than younger patients.

The present findings may be partially explained by the fact that while the use of effective treatment modalities have increased in all age groups over time,^{22,23} the prevalence of clinically significant comorbidities have increased over time³, especially in older patients. These latter trends make the management of hospitalized patients all the more challenging and increase the risk for adverse outcomes. Inasmuch, physicians need to consider the greater use of these treatment modalities in older patients to improve their short-term outcomes. Indeed, it is possible that the more aggressive management of elderly patients with coronary interventional procedures led to their declining risk of cardiogenic shock, and improving hospital survival, during the period under study. The enhanced use of these treatment regimens may also result in greater quality of life in patients of all ages and improvements in long-term prognosis.

We also observed that the short-term death rates were much higher in younger women than in younger men, with these differences persisting in the most recently hospitalized study cohorts; there were no sex differences in the crude short-term death rates among older patients. This finding is consistent with the results of previous studies.^{6,11,24–26}

The reasons for worse short-term outcomes in younger women hospitalized with AMI are unclear but may be partially explained by the fact that women have a higher prevalence of comorbid conditions than men, and differences in these and other important prognostic factors are likely to be more pronounced in younger than in older individuals.²⁵ In addition, younger women have been shown to be less likely to be treated with effective cardiac medications.^{13,26} which can contribute to the worse outcomes noted in younger women. However, a previous study of patients enrolled in the National Registry of Myocardial Infarction suggested that differences in medical history, clinical severity of the infarction, and early management accounted only for about one third of the differences in early mortality observed between men and women hospitalized with AMI.⁶ The fact that men may be more likely to die out-of-hospital from coronary disease than women, and that this sex difference may be larger in younger than in older individuals,²⁵ could contribute to higher in-hospital death rates in younger women hospitalized with AMI. Additional prospective studies need to be carried out to understand the reasons behind the greater risk of adverse outcomes noted in younger women and older individuals hospitalized with acute coronary disease.

Study Strengths and Limitations

The strengths of this study include its population-based design that captured all validated cases of AMI occurring among residents of the Worcester metropolitan area hospitalized at all Central Massachusetts medical centers over a 20-year period. On the other hand, the study population was predominantly white and the generalizability of our findings to other race/ethnic groups may be limited. We did not have information available on several patient-associated characteristics (e.g., socioeconomic status, psychological factors) which may have confounded some of the observed associations. Because patients who died before hospitalization for AMI were not included, our findings are only generalizable to patients hospitalized with AMI.

In conclusion, while encouraging declines in hospital death rates and in the occurrence of several important clinical complications have declined in men and women of all ages during the past 20 years, older men and women were more likely to experience adverse short-term outcomes after hospitalization for AMI than patients <65 years. More targeted treatment approaches during hospitalization for AMI for older patients are needed to improve their short-term prognosis.

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Appendix. Characteristics of Patients Hospitalized with Acute Myocardial Infarction by Sex and by Age (2003/2005)

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Characteristics	<65 y (n = 466)	65-74y (n = 244)	≥75y (n = 408)	p- value	<65 y (n = 189)	65-74y (n = 166)	≥75y (n = 586)	p- value
Age (mean \pm SD, years)	53.4 (7.83)	69.9 (2.99)	82.8 (5.19)		55.4 (7.12)	70.0 (2.68)	84.3 (5.89)	
White (%)	86.6	83.3	92.6	0.001	95.1	94.6	98.4	0.005
Marital status (%)								
Single	19.7	11.6	7.6	<0.001	16.4	13.3	11.1	<0.001
Married	62.2	69.4	64.1		53.9	43.6	25.6	
Divorced	16.1	10.3	5.2		15.3	7.9	2.6	
Widowed	1.3	7.9	21.9		12.7	34.6	59.7	
Medical History (%)								
Atrial fibrillation	2.6	12.7	27.2	<0.001	3.2	10.8	18.9	<0.001
Hypertension	56.2	78.7	7.67	<0.001	69.8	85.5	83.3	<0.001
Heart failure	7.94	20.1	37.9	<0.001	17.5	25.9	36.9	<0.001
Diabetes	21.0	45.1	37.3	<0.001	37.0	52.4	32.6	<0.001
Stroke	4.51	10.7	20.6	<0.001	8.5	12.1	13.7	0.17
AMI characteristics (%)								
Initial	72.3	63.9	56.1	<0.001	71.9	66.3	62.3	0.05
Q-wave	34.9	20.5	9.3	<0.001	21.7	12.1	15.2	0.03
STEMI	50.9	40.6	17.7	<0.001	33.9	24.1	24.7	0.04
Pre-hospital delay (hr), median								
(IQR)	1.78(1.00-4.42)	2.00(1.00 - 3.50)	2.00(1.00 - 4.33)	<0.001	1.83(1.00-4.00)	2.00 (1.17-4.25)	2.08(1.20 - 4.43)	<0.001
Clinical parameters on admission, median (IQR)								
Heart rate (beats/min)	80 (68 – 92)	80 (68 - 100)	80 (68 - 100)	<0.001	85.5 (72 – 98)	84(69.5 - 106)	88 (75 – 104)	<0.001
Systolic BP, mmHg	140(120 - 162)	143(119-162)	138 (117–157)	0.203	142 (120 – 167)	144 (123 – 174)	139(118 - 162)	0.20
Diastolic BP, mmHg	83 (71–97)	78 (66 – 92)	72 (60 – 84)	<0.001	76 (63 – 90)	75 (62 – 89)	69 (56 – 83)	<0.001
BMI (kg/m ²)	28.9 (25.8–32.5)	27.8(24.4–31.1)	25.8 (23.1–28.3)	<0.001	28.3(24. 9–32.9)	28.9(23.4–32.6)	25.4(22.3-28.9)	<0.001
Laboratory findings on admission, median (IQR)								
Cholesterol (mg/dl)	179(154 –208)	158(137-193)	152(126 - 180)	<0.001	180(157 – 216)	164(136 -201)	168(142-201)	<0.001
LDL (mg/dl)	115(89-140)	96 (74 – 119)	91 (67 – 116)	<0.001	106(83.5 - 134)	96 (70-128)	98 (75 – 127)	<0.001
Glucose (mg/dl)	130 (112 – 170)	157(117-214)	150(119-207)	<0.001	149(113 – 219)	167(125 - 242)	153 (122 – 212)	<0.001
Creatinine (mg/dl)	1.1(0.9 - 1.2)	1.2(1.0-1.5)	1.4 (1.1 – 1.8)	<0.001	0.9 (0.8 – 1.2)	1.1 (0.9 – 1.3)	1.2 (1.0 – 1.6)	<0.001

		Men				Women		
Characteristics	<65 y (n = 466)	65-74y (n = 244)	≥75y (n = 408)	p- value	<65 y (n = 189)	65-74y (n = 166)	≥75y (n = 586)	p- value
Length of stay (day), median								
(IQR)	3.0(2.0-4.0)	4.0 (3.0 – 7.0)	$4.0(\ 3.0 - 7.0)$	<0.001	4.0 (2.5 – 5.0)	$3.0(2.0-4.0) 4.0(3.0-7.0) 4.0(3.0-7.0) <0.001 4.0(2.5-5.0) 5.0(3.0-8.0) 5.0(3.0-7.0) <0.001 <0.001 \\ -0.001 $	5.0(3.0-7.0)	<0.001
	I, body mass index; EF,	LDL, low-density l	ipoprotein choleste	rol				

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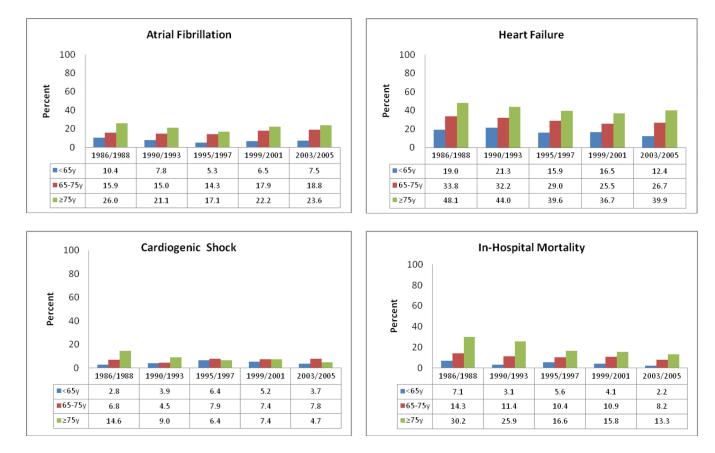


Figure 1.

Clinical Complications and In-Hospital Mortality According to Age and Study Period Among Men

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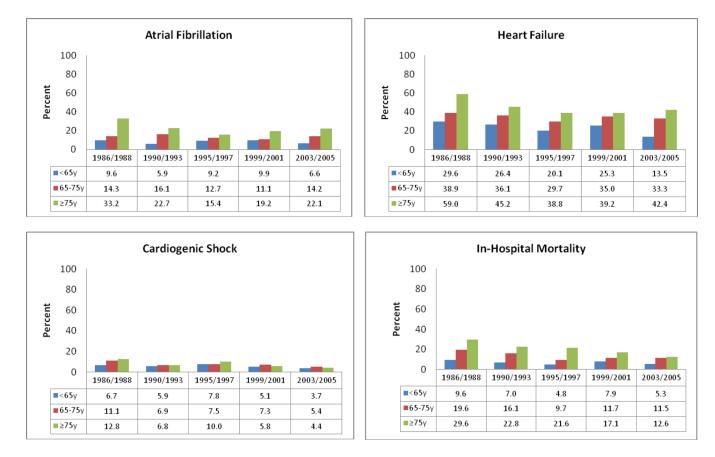


Figure 2.

Clinical Complications and In-Hospital Mortality According to Age and Study Period Among Women

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Table 1

Characteristics of Patients Hospitalized with Acute Myocardial Infarction (AMI) According to Age and Sex (1986-2005)

<			Men				Women		
$69.7(\pm 2.9)$ $81.8(\pm 5.1)$ $55.2(\pm 7.6)$ $70.0(\pm 2.8)$ $83.3(\pm 5.7)$ 95.0 97.6 0.001 90.0 93.2 96.4 9.6 7.2 0.001 11.4 10.4 10.6 76.9 67.3 60.01 11.4 10.4 10.6 76.9 67.3 20001 11.5 37.0 60.6 7.5 22.5 11.5 37.0 60.6 72.8 7.4 23.3 2001 17.5 37.0 60.6 7.5 22.5 0.001 55.0 69.9 72.8 17.2 2300 11.5 37.0 60.6 62.3 33.4 61.001 35.6 60.01 37.6 62.6 33.4 61.3 60.01 37.6 62.6 62.6 33.4 61.0 52.6 60.01 52.6 60.01 52.6 60.6	Characteristics	<65 y (n=2,554)	65–74y (n=1,545)	≥75y (n=1,808)	P- value	<65y (n=855)	65-74y (n=1,092)	≥75y (n=2,459)	P- value
95.0 97.6 <0001 90.0 93.2 96.4 76.9 7.2 <0001 11.4 10.4 10.6 76.9 <7.3 <0.001 11.4 10.4 10.6 75.9 <3.11 <1.43 <7.4 <2.8 7.5 $<2.2.5$ $<1.1.5$ <37.0 <60.6 <2.8 <7.5 $<2.2.5$ <0.001 <2.4 <2.8 <7.4 <2.8 <7.5 $<2.2.5$ <0.001 <2.4 <2.8 <7.5 <7.5 <7.7 <0.001 <2.4 <2.8 <7.6 <7.2 <7.6 <7.5 <7.7 <0.001 <7.6 <7.9 <7.2 <7.6 <7.2 <7.7 <7.6 <7.7 <7.6 <7.2 <7.2 <7.6 <7.6 <7.6 <7.6 <7.6 <7.6 <7.6 <7.6 <7.6 <7.6 <7.6 <7.6 <7.6 <7.6	Age (mean \pm SD, years)	53.4 (±8.0)	69.7 (±2.9)	81.8 (±5.1)		55.2 (±7.6)	70.0 (±2.8)	83.3 (±5.7)	
9.67.2 < 0001 11.410.410.676.9 67.3 < 0.01 45.3 26.0 5.9 3.1 1.4 $1.4,3$ 7.4 2.8 7.5 2.25 $1.1,5$ 37.0 60.6 7.5 2.25 $1.1,5$ 37.0 60.6 7.5 2.25 $1.1,5$ 37.0 60.6 7.5 $2.2,6$ 6.9 $7.2,8$ 7.12 29.0 < 0.001 35.6 69.9 72.8 7.12 29.0 < 0.001 35.8 39.8 30.0 7.12 29.0 < 0.001 35.8 39.8 30.0 7.12 29.0 < 0.001 35.8 39.8 30.0 7.12 29.0 < 0.001 37.2 31.5 27.1 33.4 31.0 < 0.001 37.2 31.5 27.1 33.4 31.0 60.01 37.2 31.5 27.1 33.4 21.0 60.01 37.2 31.5 27.1 33.4 20.01 37.2 31.5 27.1 35.9 28.1 < 0.001 37.2 31.5 27.1 43.0 26.9 20.01 37.2 31.5 $27.12-5.0$ 43.0 26.0 20.01 37.0 31.6 $27.12-5.0$ 44.0 $8.77-102$ $8.77-102$ $8.77-102$ $8.77-102$ $8.167-903$ $25.62.31-28.0$ 0.001 $28.10-9.23.1$ $27.12-5.02$ $8.167-903$ 2	White (%)	91.9	95.0	97.6	<0.001	90.0	93.2	96.4	<0.001
967.2 (001) 11.4 10.4 10.6 76.9 67.3 (011) 45.3 26.0 5.9 3.1 $1.4.3$ 7.4 2.8 7.5 22.5 11.5 37.0 60.6 7.5 22.5 11.5 37.0 60.6 7.5 22.5 11.5 37.0 60.6 7.5 22.6 60.1 5.6 69.9 72.8 7.12 29.0 <0.001 5.6 69.9 72.8 33.4 31.0 <0.001 5.0 69.9 72.8 33.4 31.0 <0.001 5.0 69.9 72.8 33.4 31.0 <0.001 5.0 99.9 33.1 33.4 31.0 <0.001 5.7 10.7 13.7 33.4 31.0 <0.001 35.8 39.8 30.0 11.7 16.8 <0.001 37.2 31.5 27.1 35.6 5.0 31.7 21.0 34.6 30.7 35.9 52.8 30.0 37.2 31.5 $27.1.2$ 43.0 26.9 <0.001 37.2 31.6 30.7 35.9 50.7 37.6 31.6 30.7 36.6 50.7 50.01 37.2 31.6 52.3 37.9 $20.1-4.5$ $21.1-4.5$ $21.1-5.5$ $27.1-2.5$ 37.6 37.6 37.6 37.6 34.6 30.7 31.6 50.7 37.6 3	Marital status (%)								
769673601453260593114.37.42.87522.511.537.060.67522.511.537.060.611.219.7 <001 2.4 8.8 17.5 59.4 64.3 <0001 2.4 8.8 17.5 59.4 64.3 <0001 2.4 8.8 17.5 53.4 31.0 <0001 3.0 21.0 33.1 33.4 31.0 <0001 3.7 21.0 33.1 33.4 31.0 <0001 37.2 39.8 30.0 11.7 16.8 <0001 37.2 31.6 27.1 33.4 31.0 <0001 37.2 31.5 27.1 33.4 25.9 <0001 37.2 31.5 27.1 33.4 25.1 <0001 37.2 31.5 27.1 43.0 26.9 <0001 37.2 31.5 27.1 43.0 26.9 <0001 37.2 31.5 27.1 $20(1-4.5)$ $25(12-5.1)$ <0001 $20(1.0-4.7)$ $24(1.3-5.5)$ $23(1.2-5.0)$ $81(67-100)$ $87(7-102)$ $8472-102$ $140(17-160)$ $140(17-160)$ $87(7-102)$ $80(67-91)$ $73(61-80)$ 0001 $80(64-92)$ $77(64-90)$ $70(57.5-80)$ $80(57-91)$ $256(23.1-280)$ 2001 $281(24-231)$ $2162(1.3-281)$ $2162(1.3-283)$ $80(58-23)$ $172(42-205)$	Single	14.3	9.6	7.2	< 0.001	11.4	10.4	10.6	<0.001
59 3.1 14.3 7.4 2.8 7.5 22.5 11.5 37.0 60.6 7.5 22.5 11.5 37.0 60.6 11.2 19.7 6001 2.4 8.8 17.5 59.4 64.3 6001 56.0 69.9 72.8 17.2 29.0 6001 13.0 21.0 33.1 33.4 31.0 6001 35.8 39.8 30.0 11.7 16.8 6001 37.2 31.6 30.7 33.4 31.0 6001 37.2 31.5 27.1 33.4 31.0 6001 37.2 31.5 27.1 35.9 58.1 6001 73.0 66.9 62.3 35.9 22.8 6001 37.2 31.5 27.1 43.0 26.9 6001 37.2 31.5 27.1 43.0 26.9 6001 37.2 31.5 27.1 43.0 26.9 6001 37.2 31.5 $27.12-5.0$ 47.0 $57.1-6.9$ 37.2 31.5 $27.12-5.0$ $81(67-100)$ $87(7-102)$ $83(70-99)$ $84(72-102)$ $8(74-105)$ $112(121-160)$ $140(117-160)$ 0001 $83(70-99)$ $84(72-102)$ $8(74-105)$ $80(67-91)$ $73(61-80)$ 0001 $80(64-92)$ $77(64-90)$ $70(57.5-80)$ $80(67-91)$ $73(61-80)$ 26.01 $20(110-47)$ $21.2(23.0-32.1)$ $24(21.3-28.3)$ <tr<< td=""><td>Married</td><td>73.8</td><td>76.9</td><td>67.3</td><td></td><td>60.1</td><td>45.3</td><td>26.0</td><td></td></tr<<>	Married	73.8	76.9	67.3		60.1	45.3	26.0	
7.5 22.5 11.5 37.0 60.6 11.2 19.7 0.001 2.4 8.8 17.5 59.4 64.3 0.001 56.0 69.9 72.8 33.4 31.0 0.001 35.8 39.8 30.0 33.4 31.0 0.001 35.8 39.8 30.0 11.7 16.8 0.001 35.8 39.8 30.0 33.4 31.0 0.001 35.8 39.8 30.0 33.4 31.0 6.001 35.8 39.8 30.0 35.9 58.1 0.001 73.0 66.9 62.3 35.9 22.8 0.001 37.2 31.5 27.1 43.0 26.9 6.001 37.2 31.5 27.1 43.0 26.9 0.001 37.2 31.6 32.6 43.0 26.9 0.001 37.2 31.5 $23(12-5.0)$ $210(1-4.5)$ $2.5(12-5.1)$ 0.001 337.0 $94.72-102$ $88.74-105$ $81(67-100)$ $8770-999$ $84.72-102$ $88.74-105$ $81(67-91)$ $73(61-86)$ 0.001 $83.70-999$ $84.72-102$ $88.74-105$ $80(67-91)$ $73(61-86)$ 0.001 $80(64-92)$ $77(64-90)$ $70(57.5-80)$ $80(67-91)$ $73(61-86)$ 0.001 $28.1(24.5-33.1)$ $21.3(14-251)$ $24.6(21.3-23.8)$ $80(67-91)$ $25.6(23.1-28.0)$ 0.001 $209(174-247)$ $21.5(174-251)$ $1999(1-235)$ <	Divorced	9.6	5.9	3.1		14.3	7.4	2.8	
11.2 19.7 6001 2.4 8.8 17.5 59.4 64.3 0001 56.0 69.9 72.8 17.2 29.0 0.001 13.0 21.0 33.1 33.4 31.0 0001 35.8 39.8 30.0 11.7 16.8 0.001 35.8 39.8 30.0 11.7 16.8 0.001 37.2 31.5 22.0 59.6 58.1 0.001 73.0 66.9 62.3 59.6 58.1 0.001 73.0 66.9 62.3 35.9 22.8 0.001 73.0 66.9 62.3 35.9 22.8 0.001 73.0 54.6 30.7 43.0 26.9 6.001 37.2 31.5 27.1 43.0 26.9 6.001 37.2 31.6 30.7 43.0 26.9 6.001 37.2 31.6 30.7 43.0 76.7 10.7 13.7 31.6 30.7 43.0 76.7 37.2 31.6 31.6 30.7 $2.0(1-4.5)$ $2.5(1.2-5.1)$ <0.01 $8370-99$ $8472-102$ $8771-02$ $81(67-100)$ $8771-102$ $8877-102$ $8874-105$ $1422(121-164)$ $143(120-170)$ $81(67-90)$ $736623.1-28.0$ 0.001 $8064-92$ $7764-90$ $70675-86$ $8067-91)$ $7361-860$ 2001 $20174-247$ $215(174-251)$ $19991-235$ $8061-823$ $172(122-2$	Widowed	2.2	7.5	22.5		11.5	37.0	60.6	
11.219.7 <0.001 2.4 8.8 17.5 59.4 64.3 <0.001 56.0 69.9 72.8 17.2 29.0 <0.001 13.0 21.0 33.1 33.4 31.0 <0.001 35.8 39.8 30.0 11.7 16.8 <0.001 35.8 39.8 30.0 33.4 31.0 <0.001 35.8 39.8 30.0 11.7 16.8 <0.001 73.0 66.9 62.3 59.6 58.1 <0.001 73.0 66.9 62.3 35.9 22.8 <0.001 73.0 66.9 62.3 35.9 22.8 <0.001 37.2 31.5 27.1 43.0 26.9 <0.001 37.2 31.5 27.1 43.0 26.9 <0.001 37.2 31.5 27.1 43.0 26.9 <0.001 37.2 31.5 27.1 43.0 26.9 <0.001 $837.0-99$ $8472-102$ $8874-105$ $81(67-100)$ $8771-102$ $8370-99$ $8472-102$ $8874-105$ $81(67-100)$ $8771-102$ $80.64-92$ $7764-90$ $70.675-80$ $8067-91$ $73(61-86)$ <0.001 $80(64-92)$ $7764-90$ $70675-86$ $8067-91$ $7361-86$ $273.1-28.3$ $273.32.3.1$ $273.32-32.1$ $276.21.3-28.3$ $8067-91$ $772(42-205)$ <0.001 $209(174-247)$ $215(174-251)$ $19991-235$ $186(158-223$	Medical History (%)								
59.4 64.3 <0.001 56.0 69.9 72.8 17.2 29.0 <0.001 13.0 21.0 33.1 33.4 31.0 <0.001 35.8 39.8 30.0 33.4 31.0 <0.001 35.8 39.8 30.0 33.4 31.0 <0.001 35.8 39.8 30.0 11.7 16.8 <0.001 5.7 10.7 13.7 59.6 58.1 <0.001 73.0 66.9 62.3 53.9 22.8 <0.001 77.0 34.6 30.7 43.0 26.9 <0.001 37.2 31.5 27.1 43.0 26.9 <0.001 47.0 34.6 30.7 $2.0(1-4.5)$ $2.5(1.2-5.1)$ <0.001 $2.0(1.0-4.7)$ $2.4(1.3-5.5)$ $2.3(1.2-5.0)$ $81(67-100)$ $8771-102)$ $83770-99)$ $84772-102)$ $88774-105)$ $80(67-91)$ $7361-860$ 0.001 $8370-99)$ $84772-102)$ $88(74-105)$ $80(67-91)$ $7361-860$ 0.001 $8370-99)$ $8472-102)$ $88(74-105)$ $80(67-91)$ $7361-860$ 20021 $8370-99)$ $8772-923.1)$ $27.3(23.0-32.1)$ $27.3(24.4-30.3)$ $25.6(23.1-28.0)$ 2001 $209(174-247)$ $215(174-251)$ $199(91-235)$ $186(158-223)$ $172(142-205)$ <0.001 $209(174-247)$ $215(174-251)$ $199(91-235)$	Atrial fibrillation	2.7	11.2	19.7	< 0.001	2.4	8.8	17.5	<0.001
17.2 29.0 <0.001 13.0 21.0 33.1 33.4 31.0 <0.001 35.8 39.8 30.0 33.4 31.0 <0.001 35.8 39.8 30.0 11.7 16.8 <0.001 5.7 10.7 13.7 59.6 58.1 <0.001 73.0 66.9 62.3 59.6 58.1 <0.001 73.0 66.9 62.3 35.9 22.8 <0.001 37.2 31.5 27.1 43.0 26.9 <0.001 37.2 31.6 30.7 43.0 26.9 <0.001 37.2 31.6 30.7 $2.0(1-4.5)$ $2.5(1.2-5.1)$ <0.001 $2.0(1.0-4.7)$ $2.4(1.3-5.5)$ $2.3(1.2-5.0)$ $81(67-100)$ $87771-102$ $8370-99$ $8477-102$ $8877-102$ $81(67-100)$ $87771-102$ $8370-99$ $84772-102$ $8877-105$ $80(67-91)$ $736(1-86)$ 0.001 $80(64-92)$ $77(64-90)$ $70(57.5-86)$ $80(67-91)$ $736(1-86)$ <0.001 $80(64-92)$ $77(64-90)$ $70(57.5-86)$ $80(67-91)$ $25.6(23.1-28.0)$ <0.001 $28.1(24.5-33.1)$ $27.3(23.0-32.1)$ $24.6(21.3-28.3)$ $186(158-223)$ $172(142-205)$ <0.001 $209(174-247)$ $215(174-251)$ $199(91-235)$	Hypertension	48.0	59.4	64.3	<0.001	56.0	6.69	72.8	<0.001
33.4 31.0 <0.001 35.8 39.8 30.0 11.7 16.8 <0.001 6.7 10.7 13.7 59.6 58.1 <0.001 73.0 66.9 62.3 59.6 58.1 <0.001 73.0 66.9 62.3 35.9 22.8 <0.001 73.0 54.6 62.3 35.9 22.8 <0.001 37.2 31.5 27.1 43.0 26.9 <0.001 47.0 34.6 30.7 $2.0(1-4.5)$ $2.5(1.2-5.1)$ <0.001 $2.0(1.0-4.7)$ $2.4(1.3-5.5)$ $2.3(1.2-5.0)$ $81(67-100)$ $877(71-102)$ $837(70-99)$ $8477-102)$ $8874-105$ $81(67-100)$ $877(71-102)$ (0.001) $837(70-99)$ $8477-102)$ $8874-105$ $80(67-91)$ $73(61-86)$ 0.001 $80(64-92)$ $77(64-90)$ $70(57.5-86)$ $80(67-91)$ $73(61-86)$ <0.001 $80(64-92)$ $77(64-90)$ $70(57.5-86)$ $27.3(24.4-30.3)$ $2.5.6(23.1-28.0)$ <0.001 $28.1(24.5-33.1)$ $27.3(23.0-32.1)$ $24.6(21.3-28.3)$ $186(158-223)$ $172(142-205)$ <0.001 $209(174-247)$ $215(174-251)$ $199(91-235)$	Heart failure	6.2	17.2	29.0	<0.001	13.0	21.0	33.1	<0.001
11.7 16.8 <0.001 6.7 10.7 13.7 59.6 58.1 <0.001 73.0 66.9 62.3 35.9 22.8 <0.001 37.2 31.5 27.1 43.0 26.9 <0.001 37.2 31.5 27.1 43.0 26.9 <0.001 47.0 34.6 30.7 $2.0(1-4.5)$ $2.5(1.2-5.1)$ <0.001 $2.0(1.0-4.7)$ $2.4(1.3-5.5)$ $2.3(1.2-5.0)$ $2.0(1-4.5)$ $2.5(1.2-5.1)$ <0.001 $8370-99$ $8472-102$ $8874-105$ $81(67-100)$ $87771-102$ <0.001 $8370-99$ $8472-102$ $8874-105$ $81(67-100)$ $87771-102$ <0.001 $8370-99$ $8472-102$ $8874-105$ $80(67-91)$ $73(61-86)$ 0.001 $80(64-92)$ $77(64-90)$ $70(57.5-86)$ $80(67-91)$ $73(61-86)$ <0.001 $80(64-92)$ $77(64-90)$ $70(57.5-86)$ $80(67-91)$ $73(61-86)$ <0.001 $80(64-92)$ $77(64-90)$ $70(57.5-86)$ $80(67-91)$ $73(61-86)$ <0.001 $80(64-92)$ $77(64-90)$ $70(57.5-86)$ $80(67-91)$ $73(61-280)$ <0.001 $80(64-92)$ $71(64-90)$ $70(57.5-86)$ $80(67-91)$ $73(61-280)$ <0.001 $28.1(24.5-33.1)$ $27.3(23.0-32.1)$ $24.6(21.3-283)$ $186(158-223)$ $172(142-205)$ <0.001 $209(174-247)$ $215(174-251)$ $199(91-235)$	Diabetes	20.9	33.4	31.0	<0.001	35.8	39.8	30.0	<0.001
59.6 58.1 <0.001 73.0 66.9 62.3 35.9 22.8 <0.001 37.2 31.5 27.1 43.0 26.9 <0.001 37.2 31.5 27.1 43.0 26.9 <0.001 47.0 34.6 30.7 $2.0(1-4.5)$ $2.5(1.2-5.1)$ <0.001 $2.0(1.0-4.7)$ $2.4(1.3-5.5)$ $2.3(1.2-5.0)$ $81(67-100)$ $87771-102)$ <0.001 $8370-99)$ $84772-102)$ $8874-105)$ $81(67-100)$ $87771-160)$ 0.05 $142(121-164)$ $143(120-170)$ $141(120-165)$ $80(67-91)$ $73(61-86)$ <0.001 $80(64-92)$ $77(64-90)$ $70(57.5-86)$ $27.3(24.4-30.3)$ $25.6(23.1-28.0)$ <0.001 $28.1(24.5-33.1)$ $27.3(23.0-32.1)$ $24.6(21.3-28.3)$ $186(158-223)$ $172(142-205)$ <0.001 $209(174-247)$ $215(174-251)$ $199(91-235)$	Stroke	4.2	11.7	16.8	<0.001	6.7	10.7	13.7	<0.001
59.6 58.1 <0.001 73.0 66.9 62.3 35.9 22.8 <0.001 37.2 31.5 27.1 43.0 26.9 <0.001 47.0 34.6 30.7 $2.0(1-4.5)$ $2.5(1.2-5.1)$ <0.001 47.0 34.6 30.7 $2.0(1-4.5)$ $2.5(1.2-5.1)$ <0.001 $2.0(1.0-4.7)$ $2.4(1.3-5.5)$ $2.3(1.2-5.0)$ $81(67-100)$ $87771-102)$ $8370-99)$ $8477-102)$ $8874-105)$ $81(67-100)$ $87771-160)$ 0.001 $8370-99)$ $8477-102)$ $8874-105)$ $80(67-91)$ $73(61-86)$ 0.001 $80(64-92)$ $77(64-90)$ $70(57.5-86)$ $80(67-91)$ $73(61-86)$ <0.001 $80(64-92)$ $77(64-90)$ $70(57.5-86)$ $27.3(24.4-30.3)$ $25.6(23.1-28.0)$ <0.001 $28.1(24.5-33.1)$ $27.3(23.0-32.1)$ $24.6(21.3-28.3)$ $186(158-223)$ $172(142-205)$ <0.001 $209(174-247)$ $215(174-251)$ $199(91-235)$	AMI characteristics (%)								
35.9 22.8 <0.001 37.2 31.5 27.1 43.0 26.9 <0.001 47.0 34.6 30.7 $2.0(1-4.5)$ $2.5(1.2-5.1)$ <0.001 $2.0(1.0-4.7)$ $2.4(1.3-5.5)$ $2.3(1.2-5.0)$ $81(67-100)$ $87771-102)$ <0.001 $8370-99)$ $8472-102)$ $8874-105)$ $81(67-100)$ $87771-160)$ 0.05 $142(121-164)$ $143(120-170)$ $141(120-165)$ $80(67-91)$ $73(61-86)$ <0.001 $80(64-92)$ $77(64-90)$ $70(57.5-86)$ $27.3(24.4-30.3)$ $25.6(23.1-28.0)$ <0.001 $28.1(24.5-33.1)$ $27.3(23.0-32.1)$ $24.6(21.3-28.3)$ $186(158-223)$ $172(142-205)$ <0.001 $209(174-247)$ $215(174-251)$ $199(91-235)$	Initial	72.7	59.6	58.1	<0.001	73.0	60.9	62.3	<0.001
$ \begin{array}{cccccccccccccccccccccccccccccccccccc$	Q-wave	46.1	35.9	22.8	<0.001	37.2	31.5	27.1	<0.001
$ \begin{array}{c ccccccccccccccccccccccccccccccccccc$	STEMI*	53.1	43.0	26.9	<0.001	47.0	34.6	30.7	<0.001
$\begin{array}{c ccccccccccccccccccccccccccccccccccc$	Pre-hospital delay (hours)*,								
$ \begin{array}{c ccccccccccccccccccccccccccccccccccc$	median (IQR)	1.9(1.0-4.2)	2.0(1-4.5)	2.5(1.2-5.1)	< 0.001	2.0(1.0-4.7)	2.4(1.3–5.5)	2.3(1.2-5.0)	0.05
81 67-100 87(71-102) <0.001 83(70-99) 84(72-102) 88(74-105) 142(121-160) 140(117-160) 0.05 142(121-164) 143(120-170) 141(120-165) 80(67-91) 73(61-86) <0.001	Clinical parameters on admis	ssion, median (IQR)							
$ \begin{array}{cccccccccccccccccccccccccccccccccccc$	Heart rate(beats/min)	80(67–94)	81(67–100)	87(71-102)	<0.001	83(70–99)	84(72–102)	88(74–105)	<0.001
80(67-91) 73(61-86) <0.001 80(64-92) 77(64-90) 70(57.5-86) 27.3(24.4-30.3) 25.6(23.1-28.0) <0.001	Systolic BP (mmHg)	141(122–162)	142(121–160)	140(117–160)	0.05	142(121–164)	143(120–170)	141(120–165)	0.83
$27.3(24.4-30.3) 25.6(23.1-28.0) <0.001 28.1(24.5-33.1) 27.3(23.0-32.1) 24.6(21.3-28.3) \\ 186(158-223) 172(142-205) <0.001 209(174-247) 215(174-251) 199(91-235) \\ \end{array}$	Diastolic BP (mmHg)	84(70–98)	80(67–91)	73(61–86)	<0.001	80(64–92)	77(64–90)	70(57.5–86)	<0.001
186(158–223) 172(142–205) <0.001 209(174–247) 215(174–251) 199(91–235)	BMI(kg/m2)*	28.6(25.7–32.3)	27.3(24.4-30.3)	25.6(23.1–28.0)	<0.001	28.1(24.5 - 33.1)	27.3(23.0-32.1)	24.6(21.3-28.3)	<0.001
210(171–236) 186(158–223) 172(142–205) <0.001 209(174–247) 215(174–251) 199(91–235)	Laboratory findings on admi	ssion, median (IQR)							
	Cholesterol(mg/dl)	210(171–236)	186(158–223)	172(142–205)	$<\!0.001$	209(174–247)	215(174–251)	199(91–235)	$<\!0.001$

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		Men				Women		
Characteristics	<65 y (n=2,554)	65–74y (n=1,545)	≥75y (n=1,808)	P- value	<65y (n=855)	65–74y (n=1,092)	≥75y (n=2,459)	P- value
LDL(mg/dl)	115(94–142)	105(82–124)	97(75-121)	<0.001	97(75-121) < 0.001 111(87-142)	107(84–141)	102(78–133)	<0.004
Glucose(mg/dl)	134(112–178)	151(119–209)	150(120-209)	<0.001	146(116–238)	163(125–247)	161(126-228)	0.02
Creatinine (mg/dl)	1.1(0.9 - 1.2)	1.2(1.0-1.5)	1.4(1.1-1.9) < 0.001	<0.001	0.9(0.8 - 1.2)	1.0(0.9 - 1.3)	1.2(0.9 - 1.6)	<0.001
Length of stay (days),								
median (IQR)	6(3–9)	7(4–10.5)	6(4-10)	< 0.001	6(4-10) < 0.001 6(4-10)	7(4–11)	6(4-10)	< 0.001
IOR: Inter quartile range: LDL, low-density lipoprotein cholesterol:	L. low-density lipoprot	cein cholesterol:						

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VK: Inter quartile range; LDL, low-density inpoprotein cholesterol;

* Information on pre-hospital delay, body mass index (BMI), acute presenting symptoms, and STEMI was recorded beginning in 1991, 1995, and 1997, respectively.

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Table 2

Age Differences in Clinical Complications and Short-term Mortality in Patients Hospitalized with Acute Myocardial Infarction

		Men				Women		
	<65 y	65–74 y	≥75 y	P value [†]	<65 y	65–74 y	≥75 y	P value ^ŕ
Atrial fibrillation								
Events, n (%)	185(7.4)	226(16.1)	325(21.7)		68(8.1)	141(13.9)	451(21.7)	
Crude OR (95% CI)	Referent	NA^*	NA	0.12	Referent	1.79(1.31–2.42)	3.17(2.42-4.15)	09.0
Adjusted OR (95% CI)	Referent	NA	NA	0.04	Referent	1.74(1.25–2.43)	3.09(2.27-4.21)	0.40
Heart failure								
Events, n (%)	414(17.3)	381(29.8)	526(41.0)		169(22.7)	299(34.7)	715(43.5)	
Crude OR (95% CI)	Referent	2.00(1.71 - 2.35)	3.41 (2.92–3.98)	0.76	Referent	1.75(1.40-2.18)	2.68(2.19–3.26)	0.28
Adjusted OR (95% CI)	Referent	1.68(1.38 - 2.05)	2.88(2.36–3.52)	0.23	Referent	1.61(1.22–2.11)	2.26(1.75–2.92)	0.24
Cardiogenic shock								
Events, n (%)	113(4.4)	102(6.6)	139(7.7)		58(5.7)	151(7.6)	479(7.2)	
Crude OR (95% CI)	Referent	NA	NA	<0.001	Referent	1.30(0.90 - 1.87)	1.33(0.96 - 1.84)	0.60
Adjusted OR (95% CI)	Referent	NA	NA	0.02	Referent	1.28(0.82 - 1.99)	1.38(0.91 - 2.10)	0.42
In-hospital mortality								
Events, n (%)	108(4.2)	171(11.1)	345(19.1)		58(6.8)	151(13.8)	479(19.5)	
Crude OR (95% CI)	Referent	2.77(2.15–3.56)	5.76(4.59–7.23)	0.15	Referent	2.11(1.54-2.90)	3.49(2.62-4.65)	0.41
Adjusted OR (95% CI)	Referent	2.83(2.09 - 3.84)	5.21(3.90–6.98)	0.13	Referent	2.10(1.43 - 3.08)	3.32(2.32-4.75)	0.22
30-hospital mortality								
Events, n (%)	129(5.1)	213(13.8)	427(23.8)		64(7.5)	166(15.2)	590(24.1)	
Crude HR (95% CI)	Referent	2.94(2.33–3.71)	5.78(4.68–7.14)	0.91	Referent	1.95(1.44-2.66)	3.74(2.85-4.92)	0.80
Adjusted HR (95% CI)	Referent	2.90(2.19–3.85)	5.10(3.90 - 6.66)	0.80	Referent	1.91(1.33–2.76)	3.44(2.46-4.80)	0.42

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* NA: Not applicable. Among men, since age differences in AF and cardiogenic shock have changed significantly over time, adjusted ORs for each study period were calculated separately and are presented in the text; there were no overall ORs for these outcomes.

p values for interaction terms between age and study period

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