# **BUKTI KORESPONDENSI**

Judul Artikel : The Impact of Classical Cardiovascular Risk Factors on Hospitalization and Mortality among Hajj Pilgrims

Jurnal : Scientific World Journal

Author : Meity Ardiana, Eka Rahayu Utami, Makhyan Jibril Al Farabi, and Yusuf Azmi

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No.	Perihal	Tanggal	Komentar Editor Jurnal	Komentar Penulis
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#### 28 March 2022

## Academic Editor, The Scientific World Journal

## Dear Professor/Doctor,

We wish to submit the manuscript entitled **"The impact of classical cardiovascular risk factors on hospitalization and mortality among Hajj pilgrims"** for consideration of publication in **The Scientific World Journal**.

Hajj, an annual pilgrimage in the Kingdom of Saudi Arabia (KSA), is the largest mass gathering event in the world, performed by more than 2 million Muslims from more than 183 countries. Indonesia is the largest single source of pilgrims, with more than 200,000 people contributing about 10% of total Hajj pilgrims. The increasing number of pilgrims combined with extreme physical stressors increases the health risks and exacerbation of underlying conditions. Cardiovascular disease (CVD) is the leading cause of morbidity and mortality during Hajj. Herein, we aim to identify the impact of classical factors of the cardiovascular event and impact on hospitalization requirement among Hajj.

This study showed that pilgrims with classical cardiovascular risk factors were associated with increased hospitalization and mortality during the Hajj period. Cardiovascular disease was accounted for about 13% of hospitalization and the majority (38.2%) of the causes of death. These findings indicate the need to identify classical cardiovascular risk factors before departure. Cardiac prevention approaches and monitoring are needed for pilgrims with CVD or those at risk of CVD during the Hajj period to reduce hospitalization and death associated with cardiovascular events.

We believe this study may fulfill scientific holes in the field and hopefully fit the scope of your journal. This study has not been published in part or whole or is not under consideration for publication elsewhere. All authors in this study have agreed to be listed and approved the manuscript.

Thank you for considering our work for publication in The Scientific World Journal.

Yours sincerely,

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# 1 Title:

- 2 The impact of classical cardiovascular risk factors on hospitalization and mortality among
- 3 Hajj pilgrims
- 4 Running title:
- 5 Classical cardiovascular risk factors in Hajj pilgrims
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## 38 Abstract

Background. Cardiovascular disease (CVD) is the leading cause of morbidity and mortality
during Hajj. This study aims to The objective of the present study was to -examine the effect
of classical cardiovascular disease risk factors on identify the impact of classical factors of the
cardiovascular event and impact on mortality and hospitalization requirement among Hajj
pilgrims from East Java, Indonesia during 2017, 2018, and 2019-Hajj.

Methods. This study was a retrospective cohort of Hajj pilgrims from East Java, Indonesia,
from 2017 to 2019. The data of risk factors were obtained from the pre-embarkation Hajj
screening records. The diagnosis of hospitalization and cause of death during the Hajj period
were obtained from the medical report and hospital/flight doctor death certificate.

48 **Results.** A total of 72078 eligible subjects were included in this study. 33807 (46.9%) were 49 men, and 38271 (53.1%) were women, and the majority (35%) were aged between 50 and 59 50 years old. A total of 42446 pilgrims (58.9%) were classified as high risk due to underlying 51 health conditions such as hypertension, diabetes, or if they were aged 60 years or older. The 52 overall hospitalization rate is 971 per 100,000 pilgrims and the overall death rate is 240 deaths 53 per 100,000 pilgrims. Multivariate analysis using logistic regression showed that male gender, 54 age > 50 years old, hypertension grade II-III, diabetes, overweight and obesity were associated 55 with higher risk of hospitalization. Moreover, male gender, diabetes, and overweight were 56 associated with higher risk of mortality. Of all hospitalized patients, 92 patients (13.1%) had 57 an initial diagnosis of CVD, and CVD is the main cause of mortality (38.2%) of pilgrims.

58 Conclusion. Pilgrims with classical cardiovascular risk factors were associated with increased
59 hospitalization and mortality.

60 **Keywords:** Hajj, pilgrims, classical cardiovascular risk factor, hospitalization, mortality

#### 61 Introduction

62 Hajj, an annual pilgrimage in the Kingdom of Saudi Arabia (KSA), is the largest annual global 63 temporary migration and largest mass gathering event in the world, performed by more than 64 2 million Muslims from more than 183 countries.(1) Indonesia, as the country with the largest 65 Muslim population, holds the largest Hajj visa quota and is the largest single source of 66 pilgrims, with more than 200,000 people contributing about 10% of total Hajj pilgrims.(2) The 67 increasing number of pilgrims combined with extreme physical stressors such as sun exposure, heat, thirst, traffic jams, and steep inclines for a long period increases the health 68 69 risks and exacerbation of underlying conditions.(3)

70 Cardiovascular disease (CVD), including coronary heart disease, hypertension, congestive 71 heart failure, and arrhythmias, is the leading cause of morbidity and mortality during Hajj.(4) 72 It accounts for about 25% of hospitalizations and 64% of ICU admissions among pilgrims in 73 Hajj.(5,6) Cardiovascular disease was reported as the most common cause of death during 74 Hajj, accounting for 66% of all deaths (446 deaths) of 206,831 Indonesian pilgrims in Hajj 75 2008.(2) Due to the high morbidity and mortality of CVD among Hajj pilgrims, it is important 76 to know the risk factors and their significance in increasing the need for hospitalization and 77 mortality. This study aims to identify the impact of classical cardiovascular risk factors on 78 hospitalization requirements among pilgrims from East Java during the 2017, 2018, and 2019 79 Hajj.

#### 80 Methods

#### 81 Data Collection

82 The design of this study was a retrospective cohort, non-intervention study. The sample 83 population was all Hajj pilgrims from East Java, Indonesia, from 2017 to 2019. The follow-up 84 period is 30-50 days (median 43 days) covering the entire Hajj period from departure to KSA to return to Indonesia (Figure 1). Data of risk factors were obtained from the Hajj medical 85 86 service records carried out by trained health workers during the pre-embarkation medical 87 assessment. During pre-departure, pilgrims undertake a medical test to receive a meningitis 88 vaccine and confirm travel fitness. Pilgrims aged 60 years old or older and had at least one preexisting medical condition (e.g., hypertension and diabetes mellitus) were classified as 89 90 high risk. Variables extracted from pre-departure screening records include demographic data 91 of name, age, gender, home address, and physical examination data, including blood pressure 92 measurement and body mass index (BMI) calculation. Hypertension was defined following 93 the guidelines from the European Society of Cardiology (ESC) 2018 with office systolic blood 94 pressure (SBP) values of 140 and/or diastolic blood pressure (DBP) of 90 mmHg. The 95 classification of hypertension is divided into grade 1 with SBP 140-159 mmHg and/or DBP 90-99 mmHg, grade 2 with SBP 160-179 mmHg and/or DBP 100-109 mmHg, and grade 3 with 96 97 SBP 180 mmHg and/or DBP 110 mmHg.(7) Obesity was defined based on BMI following the 98 World Health Organization (WHO) classification for overweight and obesity recommendation 99 for Asian population. Participants with BMIs ranging from 18.5–22.9 kg/m2 were categorized 100 as normal, BMIs less than 18.5 kg/m2 were categorized as underweight, BMIs ranging from 101 23.0–24.9 kg/m2 were categorized as overweight, BMIs ranging from 25–29.9 kg/m2 were 102 categorized as obesity class I, BMIs  $\geq$ 30 kg/m2 were categorized as obesity class II.(8) Postprandial blood sugar and the history of cigarette smoking were extracted from the medical report. Diabetes was defined according to the American Diabetes Association (ADA) criteria as a self-reported diagnosis determined previously by a healthcare professional or among participants without self-reported diabetes using an oral glucose tolerance test with a two-hour plasma glucose 200 mg/dL (11.1 mmol)/L).(9)

The diagnosis of hospitalization during the Hajj period from 2017 to 2019 was obtained from the medical report. Cause of death was obtained from hospital or flight doctor death certificate. Hospitalizations and deaths <u>during the Hajj period from 2017 to 2019</u> were documented by the Indonesian public health team based in Saudi Arabia. Diagnosis of hospitalization and mortality data were coded according to the International Classification of Diseases-10 (ICD-10) coding.

114 Classical cardiovascular risk factors

115 Classical cardiovascular risk factors were identified in the Framingham Heart Study as 116 conferring increased risk of CVD in the general population.(10) Classical cardiovascular risk 117 factors analyzed in this study were age, <u>mengender</u>, hypertension, diabetes, smoking, and 118 obesity. Blood pressure (BP) determination was made using a periodically calibrated mercury 119 sphygmomanometer. Diabetes was diagnosed at a two-hour blood sugar greater than or 120 equal to 200 mg/dl of oral glucose tolerance test (OGTT).(11)

#### 121 Statistical Analysis

All statistical analyses were performed using SPSS 20.0. The normal distribution of data was evaluated using the Kolmogorov-Smirnov test. Continuous variables were presented as median and 25th/75th percentiles, while categorical variables were presented as absolute frequencies and percentages.(12) Pearson's Chi-square or Fisher's exact tests were used to 126 compare univariate continuous variables, and independent T-tests were used to compare 127 univariate numeric categorical variables. A P-value of < 0.05 was considered statistically 128 significant. Furthermore, multivariate analysis was performed using logistic regression to to 129 identify which classical cardiovascular risk factors predict hospitalization and mortality among 130 Hajj pilgrims. The variables included in the logistic regression analysis were variables with P value < 0.25 in bivariate analysis. Receiver operating characteristic (ROC) curves were 131 constructed for classical cardiovascular risk factors and two outcomes of hospitalization and 132 133 mortality.

## 134 Ethics Approval

The clinical and epidemiological data collection was submitted for ethical review to the Ministry of Health of the Republic of Indonesia with approval number of HJ.01.03/I/2900/2020.

#### 138 Results

#### 139 Characteristics of patients

140 Of the 106680 subjects obtained from pilgrims attending for the Hajj period from 2017 to 141 2019, 34602 (32.43 %) were excluded from analysis due to incompleteness in demographic 142 data, physical examination, laboratory results, or patient outcomes. The final sample was 143 therefore composed of 72078 eligible subjects in the study (Table 1) with 21488, 23865 and 144 26725 pilgrims respectively from the 2017, 2018 and 2019 Hajj years. Of these, 33807 (46.9%) 145 were men and 38271 were (53.1%) women. All pilgrims were aged  $\geq$  18 years old with a 146 median age of 54 years old (47 - 61), and the majority (35%) were aged between 50 and 59 147 years old. From the three Hajj year periods from 2017 to 2019, there were significant 148 differences in age groups, prevalence of hypertension, BMI distribution, average fasting blood 149 sugar level, and smoking history. In addition, there were also differences in the hospitalization 150 rate and mortality of the Hajj participants from 2017 to 2019. The highest reported 151 hospitalization rate was in the 2019 Hajj year but with the highest mortality rate was in 2017. 152 A total of 42446 pilgrims (58.9%) were classified as high risk due to underlying health 153 conditions such as hypertension, diabetes, or if they were aged 60 years or older. An increase 154 in BMI was also common in pilgrims, with 13922 (19.3%) subjects being overweight and 36080 155 (50.1%) subjects being diagnosed as obese. In addition, smoking history was recorded in 156 10389 (14.4%) subjects, with the majority were men (10202 subjects; 98.2%)

157 Factors contributing to the hospitalization in pilgrims

During 2017-2019, 700 out of 72078 pilgrims were required hospitalization during Hajj in Saudi Arabia (Table 2). The overall hospitalization rate is 971 per 100,000 pilgrims. Men were associated with increased hospitalization than women (1.18% vs 0.80%, p<0.001). Pilgrims 161 aged 60 years or older (2.17%) was also required more hospitalization compared to younger 162 ones, and the rates significantly increased with increasing age (p<0.001). Hospitalization was 163 required in 4.44% of patients with hypertension (versus 0.79% of patients without hypertension; P < 0.001) and 2.28% in patients with diabetes (versus 0.82% of patients 164 165 without diabetes, P < 0.001). Obese (BMI ≥ 30) and underweight (BMI < 18.5) pilgrims had the 166 highest percentage of hospitalization (3.18% and 3.17%, respectively) compared to pilgrims 167 with normal BMI (BMI 18.5-24.9; 1.09%; p < 0.001). There was no association between 168 smoking history and hospitalization among pilgrims (p = 0.905).

#### 169 Factors contribute to mortality in pilgrims

170 During 2017-2019, 173 out of 72078 pilgrims died during Hajj in Saudi Arabia. The overall 171 death rate is 240 deaths per 100,000 pilgrims (Table 2). The mortality rate was higher in men 172 (144 deaths per 100,000 pilgrims) than in women (96 deaths per 100,000 pilgrims, p<0.001). 173 The mortality rate was highest in those aged 60 years or older (593 per 100,000 pilgrims), and 174 rates significantly increased with increasing age (p<0.001). Mortality was found in 0.29% of 175 patients with hypertension (versus 0.21% of patients without hypertension; P < 0.001) and 176 0.55% in patients with diabetes (versus 0.20% of patients without diabetes, P < 0.001). 177 Underweight pilgrims (BMI < 18.5) had the highest percentage of mortality (0.79%) followed 178 pilgrims with normal BMI (BMI 18.5-24.9; 0.25%) and overweight and obese pilgrims (BMI ≥ 179 25; 0.16%; p < 0.001). There was no association between smoking history and mortality 180 among pilgrims (p = 0.839). In addition, when combined, the cumulative total of classical 181 cardiovascular risk factors increases with the incidence of hospitalization and death (Figure 182 2).

Multivariate analysis using logistic regression showed that male gender, age > 50 years old, hypertension grade II-III, diabetes, overweight and obesity were associated with higher risk of hospitalization. Moreover, several variables such as male gender, diabetes, and overweight, but not obesity, were associated with higher risk of mortality (Table 3). Areas under the ROC curve (AUC) for male gender, age, hypertension, diabetes, BMI and risk of hospitalization was 0.762. While AUC for male gender, diabetes, BMI and mortality was 0.806 (Figure 3).

# 190 Diagnosis during hospitalization and cause of mortality

Of all hospitalized patients (n = 700 pilgrims), 92 patients (13.1%) had an initial diagnosis of CVD (Figure 4). The top three CVDs leading to hospitalization were congestive heart failure (24 pilgrims), acute myocardial infarction (12 pilgrims), and hypertensive heart disease (9 pilgrims). Meanwhile, CVD is the main cause of mortality for pilgrims (38.2%), followed by respiratory diseases (29.5%), circulatory diseases (9.8%), and infectious and parasitic diseases (7.5%) (Figure 5). 197 Discussion

198 Over the last few decades, CVD has emerged as an important cause of hospital admission and 199 death among Hajj pilgrims. Especially pilgrims with preexisting heart disease are at high risk of experiencing physical stress leading to ischemia.(4) A cross-sectional study analyzing 200 201 admissions to 1487 beds and 104 intensive care units (ICUs) from three hospitals in Arafat 202 and four hospitals in Mena was conducted during the Hajj 2004. This study showed that CVD 203 accounted for nearly 25% of admissions, including congestive heart failure, hypertension, ischemic heart disease, and arrhythmias. Furthermore, it also showed that about 64% of ICU 204 205 admissions were due to CVD, which was dominated by left ventricular failure and myocardial 206 infarction. In addition, more than 70% of these ICU patients had underlying diseases that 207 required medical attention, with more than half of the comorbid were CVDs such as 208 congestive heart failure, hypertension, and ischemic heart disease. (5,6) Another prospective 209 cohort study from four hospitals in Mina during the 2009 Hajj period showed that CVD (23.6%) 210 was the second most critically ill patient after severe infections (46.5%) with a diagnosis of 211 myocardial infarction, atrial fibrillation (4.7%), and cardiogenic shock (2.7%).(13) CVD was the 212 most common cause of morbidity (34.1%) in patients necessitating admission to a tertiary 213 care hospital in Makkah during the 2005 Hajj.(14) Previous study in Indonesia showed the 214 association between CVD and increased morbidity and hospital admissions among Hajj 215 pilgrims. Moreover, CVD the most common cause of death during Hajj, accounting for 66% of all deaths (446 deaths) of 206,831 Indonesian pilgrims in Hajj 2008.(2) Cardiovascular disease 216 217 during Hajj was also reported by Saudi Arabian Ministry of Health as the most common cause 218 of death compared to other medical diseases, both communicable and non-communicable 219 diseasesIn addition to increased morbidity and hospital admissions, CVD is the most common 220 cause of death during Hajj, accounting for 66% of all deaths (446 deaths) of 206,831

221 Indonesian pilgrims in Hajj 2008.(2) Cardiovascular diseases during the Pilgrimage cause more

222 death than other medical ailments, both communicable and non-communicable.-(15)\_(4)

223 Due to the high morbidity and mortality of CVD among Hajj pilgrims, it is important to know 224 the risk factors and their significance in increasing the need for hospitalization and mortality. 225 In the second half of the twentieth century, many epidemiological studies identified risk 226 factors for CVD that have contributed to primary prevention. One of the most widely used 227 risk prediction equations is the Framingham model to estimate the 10-year risk of coronary 228 heart disease (CHD). The Framingham model is based on classical risk factors, including age, 229 gender, blood pressure, low-density lipoprotein (LDL) and high-density lipoprotein (HDL) 230 cholesterol, smoking, and diabetes.(10) Although long-standing, recent evidence-based on 231 population risk calculations in the United States using the Framingham 10-year risk estimate 232 for CHD suggests that 75% to 85% of CHD can be prevented by avoiding classic risk factors. (16) 233 Identifying cardiovascular risk factors in Hajj participants is important because it is estimated 234 that the obligation to carry out the rites during Hajj increases the burden on cardiac function, 235 especially in pilgrims with underlying CVD. Sudden strenuous physical activity results in 236 stressful exercises and triggers several mechanisms, including the decreased venous return 237 and decreased cardiac output that may lead to an acute cardiovascular attack. This process is 238 counterproductive and can compromise disease management and, in extreme cases, can be 239 fatal.(4) This condition is exacerbated by the heat in which sweating causes fluid loss, leading 240 to dehydration and hypovolemia, decreased cardiac output, and loss of body fluids into the

beyond this compensatory stage, the central venous pressure falls sharply, causing a further

interstitial fluid spaces with subsequent cardiovascular collapse. If heat stress continues

241

rise in core body temperature with subsequent failure of thermoregulatory mechanisms andleading to heatstroke.(17)

245 In multivariate analysis, diabetes consistently and significantly increased the risk of 246 hospitalization and death of Hajj participants in this study. A systematic review showed that 247 the prevalence of diabetes is one of the most common comorbidities found in Hajj pilgrims 248 with a prevalence of 5%.(18) Diabetes and related complications have also been reported as 249 a very influential risk factor and is one of the main causes of hospitalization and mortality 250 among pilgrims.(19) In addition, the male gender is also more often found in hospitalized and 251 deceased pilgrims. This was also found in a report from Indian Hajj Pilgrims where of the 163 252 deaths, 68.7% were male and the most common terminal event was cardiorespiratory 253 arrest.(20) Interestingly, this research also showed that underweight pilgrims (BMI < 18.5) 254 had a higher rate of hospitalization (3.18%) and mortality (1.10%) compared to normal BMI, 255 although this risk factor failed to show significant impact on multivariate analysis. This 256 unexpected result of the increased mortality risk in underweight group can be associated with 257 various clinical factors, such as poor nutritional status, sarcopenia, and aging in underweight 258 population. Previous research also suggested that underweight may be a risk for CVD disease 259 with 19.7% greater risk than the normal-weight. (21)

A study in Iran provided intervention to prospective pilgrims with a history of CVD. These interventions include cancellation of prospective Hajj pilgrims with severe CVD (history of recent myocardial infarction, unstable angina, advanced heart failure, and uncontrolled hypertension) and provision of adequate therapy and monitoring for other patients with stable angina, mild heart failure, controlled hypertension, and other cardiovascular disorders. It showed that mortality and hospitalization rates were significantly reduced in the intervention population. Iranian study and the results of this study support the need for
health screening before departure, especially for classical cardiovascular risk factors. Pilgrims
with severe CVD should be excluded from Hajj. Other pilgrims with CVD or those at risk of
CVD need appropriate intervention and monitoring during Hajj to reduce hospitalization and
mortality rates in Hajj pilgrims.

271 When compared to studies with similar area and period to this study, the results were similar. 272 A study that screened cardiovascular risk factors in adults aged 40 years and in Malang, East Java, in 2016–2017 showed that out of 22,093 participants, 6,455 (29.2%) had high 273 274 cardiovascular risk with an estimated 10-year CVD risk of  $\geq$  30%. In this study, high 275 cardiovascular risk was defined as the presence of coronary heart disease, stroke or other 276 atherosclerotic disease.(22) Another study analyzing the report of Riskesdas 2013, a 277 nationally representative survey conducted by the Indonesian Ministry of Health in 2013, 278 estimated the coronary heart disease (CHD) burden caused by five major and modifiable 279 vascular risk factors: smoking, hypertension, diabetes, increased total cholesterol, and 280 overweight. The results of this study showed hypertension as a major vascular risk factor 281 (20%–25% of all CHD) and smoking in men explains most of the vascular events (25% of 282 <u>CHD).(23)</u>

## 283 Study limitation

Although this study uses large data of pilgrims from three time periods of Hajj (2017-2019), the sample size only represents a portion of the overall Hajj pilgrims in Indonesia. In addition, the retrospective cohort design using only univariate analysis limits the generalizability of the findings. Data on risk factors were collected from pre-departure screening reports so that the responses obtained were susceptible to information bias. The multivariate analysis showed that overweight, but not obesity, significantly contributed to mortality. This may be due to
the low mortality rate of 173 from the total 71904 (0.24%) hajj participants, thus the
distribution may not be representative.

## 292 Conclusion

Pilgrims with classical cardiovascular risk factors were associated with increased hospitalization and mortality. Cardiovascular disease was accounted for about 13% of hospitalization and the majority (38.2%) of the causes of death. These findings indicate the need to identify classical cardiovascular risk factors before departure. Cardiac prevention approaches and monitoring are needed for pilgrims with CVD or those at risk of CVD during the Hajj period to reduce hospitalization and death associated with cardiovascular events.

## 299 **Disclosure statement**

# **Declaration of Conflicting Interests**

301 The authors declare no Conflict of Interest for this article.

# 302 Ethics Approval

303 Not applicable.

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**Table 1.** Characteristic of the study population of Hajj pilgrims from East Java, Indonesia,from 2017 to 2019.

Variable	2017 (n = 21488)	2018 (n = 23865)	2019 (n= 26725)	P-value (χ2 test / Anova)	Post-hoc test	P value (Man- whitney)
Men	10018	11254	12535	0.521		
	(46.6)	(47.2)	(46.9)			
Women	11470	12611	14190			
	(53.4)	(52.8)	(53.1)			
Age (year)	54.0 (11.3)	53.8 (10.8)	54.14 (11.8)	0.145		
18-40 years old	2369 (11.0)	2473 (10.4)	3086 (11.5)	< 0.001		
41-49 years old	5082 (23.7)	5499 (23.0)	5902 (22.1)			
50-59 years old	7490 (34.9)	8860 (37.1)	9442 (35.3)			
≥ 60 years old	6547 (30.5)	7033 (29.5)	8295 (31.0)			
SBP (mmHg)	127.3	127.9 (20.5)	128.4 (21.0)	<0.001	2017 vs	0.001
	(20.4)				2018	
					2017 vs	<0.001
					2019	
					2018 vs	0.045
					2019	
DBP (mmHg)	80.4 (10.2)	80.5 (10.2)	80.3 (10.4)	0.006	2017 vs	0.232
					2018	
					2017 vs	0.660
					2019	
					2018 vs	0.002
					2019	
Non-	10000	15376	1282 (4.8)	< 0.001		
hypertension	(46.5)	(64.4)				
HT grade I	7612 (35.4)	5491 (23.0)	6936 (26.0)			
HT grade II	2785 (13.0)	2200 (9.2)	5129 (19.2)			
HT grade III	1091 (5.1)	798 (3.3)	3614 (13.5)			
BMI	25.3 (4.6)	25.3 (4.5)	25.3 (4.6)	0.349		
Underweight	974 (4.5)	953 (4.0)	1282 (4.8)	0.002		
Normal	5686 (26.5)	6245 (26.2)	6936 (26)			
Overweight	4130 (19.2)	4663 (19.5)	5129 (19.2)			
Obesity class I	7885 (36.7)	8872 (13.1)	9764 (36.5)			
Obesity class II	2813 (13.1)	3132 (13.1)	3614 (13.5)			
PPG levels	136.9	138.0 (62.4)	138.6 (63.8)	< 0.001	2017 vs	<0.001
(mg/dL)	(63.4)				2018	
					2017 vs	<0.001
					2019	
					2018 vs	0.776
Diabata	2240 (40.0)		2005 (44.2)	0.247	2019	
	2340 (10.9)	2556 (10.7)	2985 (11.2)	0.247		
Smoking history	2987 (13.9)	3420 (14.3)	3982 (14.9)	0.007		
Hospitalization	173 (0.8)	230 (1.0)	297 (1.1)	0.003		

	Mortality	77 (0.4)	42 (0.2)	54 (0.2)	< 0.001	
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Table 2. Univariate analysis for hospitalization and mortality of Hajj pilgrims from East Java,
 Indonesia, from 2017 to 2019.

	Hospitalization		P-value	Mortality		P-value
Variable	No	Yes		Alive	Death	
	n=71378	n=700		n=71904	n=173	
Men	33412	395	<0.001	33703	104	<0.001
Women	37966	305	-	38202	69	-
Age (year)	53.9	64.2	<0.001	54.0	66.8	<0.001
	(11.3)	(11.1)		(11.3)	(10.2)	
Age group						
18-40 years old	7914	14	<0.001	7928	0	<0.001
41-49 years old	16436	47	-	16474	9	-
50-59 years old	25618	174	-	25757	35	-
≥ 60 years old	21410	465	-	21746	129	-
SBP (mmHg)	127.8	136.1	<0.001	127.9	134.3	< 0.001
	(20.6)	(23)		(20.7)	(22.4)	
DBP (mmHg)	80.4	82.9	<0.001	80.4	82.5	0.005
	(10.3)	(11.1)		(10.3)	(11.1)	
Non-HTN	42351	333	<0.001	42597	87	0.001
HTN grade I	18897	198	_	19050	45	_
HTN grade II	7282	119	_	7375	26	_
HTN grade III	2848	50	_	2883	15	_
BMI	25.3	23.7	<0.001	25.3	22.9	<0.001
	(4.6)	(5.3)		(4.6)	(5.5)	
Underweight	3110	99	<0.001	3174	35	<0.001
Normal	18634	233	-	18810	57	_
Overweight	13803	119	-	13898	24	-
Obesity class I	26355	166	-	26482	39	-
Obesity class II	9476	83	_	9541	18	-
PPG levels (mg/dL)	137.6	169.3	<0.001	137.8	169.4	<0.001
	(62.8)	(90.6)		(63.1)	(97.6)	
Diabetes						
Yes	7705	176	<0.001	7838	43	<0.001

No	63672	524		64067	130	
Smoking history						
Yes	10287	102	0.905	10365	24	0.839
No	61091	598	_	61540	149	-

**Table 3.** Multivariate analysis using logistic regression for risk factors of hospitalization andmortality Hajj pilgrims from East Java, Indonesia, from 2017 to 2019.

Variable	Hospita	alization	
Vallable	Odds ratio	95% CI	P-value
Men	1.334	1.146-1.554	<0.001
Age group			
18-40 years old	Reference		
41-49 years old	1.546	0.850-2.812	0.153
50-59 years old	3.262	1.856-5.640	<0.001
≥ 60 years old	8.906	5.204-15.244	<0.001
Hypertension			
Non-HTN	References		
HTN grade I	1.055	0.882-1.263	0.559
HTN grade II	1.546	1.247-1.917	<0.001
HTN grade III	1.657	1.222-2.247	<0.001
Diabetes	2.552	2.099-3.102	<0.001
BMI			
Normal	Reference		
Underweight	1.031	0.796-1.336	0.816
Overweight	2.244	1.653-3.047	<0.001
Obesity class I	0.788	0.592-1.048	0.102
Obesity class II	0.615	0.471-0.803	<0.001
Variable	Hospita	alization	
Variable	Odds ratio	95% CI	P-value
Men	1.480	1.088-2.014	0.012
Diabetes	2.567	1.808-3.643	<0.001
BMI			
Normal	Reference		

Underweight	1.009	0.588-1.731	0.975
Overweight	2.985	1.660-5.369	<0.001
Obesity class I	0.664	0.358-1.231	0.193
Obesity class II	0.631	0.360-1.106	0.108





393

Figure 1. Flow chart of the study design.





hospitalization and mortality (in 1000 individuals).





399 Figure 3. Area under the curve for the association between classical cardiovascular (CV) risk



factors and hospitalization (A) and mortality (B).



401

Figure 4. Early diagnosis during hospitalization among Hajj pilgrims.



404

Figure 5. Cause of death according to hospital/flight doctor death certificate.



Figure 1. Flow chart of the study design.



**Figure 2.** The cumulative total of classical cardiovascular risk factors with the incidence of hospitalization and mortality (in 1000 individuals).



Figure 3. Area under the curve for the association between classical cardiovascular (CV) risk

factors and hospitalization (A) and mortality (B).





Figure 4. Early diagnosis during hospitalization among Hajj pilgrims.



Figure 5. Cause of death according to hospital/flight doctor death certificate.



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