Prevalence of the Metabolic Syndrome in Patients With Acute Coronary Syndrome in Six Middle Eastern Countries

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The objective of this study was to evaluate the prevalence and effect of the metabolic syndrome (MetS) on patients with acute coronary syndrome (ACS) in six Middle Eastern countries using the new definition of MetS. Analysis of the Gulf Registry of Acute Coronary Events (Gulf RACE), which included 8716 consecutive patients hospitalized with ACS, was conducted and patients were divided into two groups: patients with and patients without the MetS. Overall,

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46% of patients had MetS. Patients with MetS were more likely to be female and less likely to be smokers. In-hospital mortality and cardiogenic shock were comparable between the two groups, although MetS patients were more likely to have congestive heart failure and recurrent ischemia. In ST-elevation myocardial infarction, MetS was also associated with increased risk of recurrent myocardial infarction and stroke. Using the recent MetS definition, MetS is highly prevalent among Middle Eastern patients presenting with ACS. MetS is associated with higher-risk profile characteristics and increased risk for development of heart failure and recurrent myocardial ischemia without an increase in hospital mortality. J Clin Hypertens (Greenwich). 2010;12:890-899. ©2010 Wiley Periodicals, Inc.

Worldwide, the metabolic syndrome (MetS) is a major health problem associated with increased morbidity and mortality. The syndrome was first described in Western societies, but it is now emerging in developing countries. MetS represents a constellation of cardiovascular risk factors, including elevated blood pressure (BP), hyperglycemia, elevated triglyceride levels, central obesity, and decreased high-density lipoprotein cholesterol.¹ The prevalence of MetS among patients with acute coronary syndrome (ACS) is limited by small registries and post hoc

analyses of randomized trials that use three different guideline definitions for MetS (Third Report of the Adult Treatment Panel National Cholesterol Education Program [NCEP-ATP III] definition; National Heart, Lung and Blood Institute/American Heart Association [NHLBI/ AHA] definition; and International Diabetes Federation [IDF] definition), making consensus about their findings difficult.²⁻¹⁵ More recently, a joint statement of the IDF Task Force on Epidemiology and Prevention, NHLBI, AHA, the World Heart Federation, the International Atherosclerosis Society, and the International Association for the Study of Obesity unified the diagnostic criteria of MetS.¹⁶ In this study we evaluate the prevalence of MetS and patient outcomes based on the new definition among a large cohort of patients presenting with ACS in six Middle Eastern countries.

METHODS

The Gulf Registry of Acute Coronary Events (Gulf RACE) prospectively enrolled 8716 patients (of which 6071 patients had data on weight and abdominal waist circumference) with ACS from 65 centers in six adjacent Middle Eastern gulf countries (Bahrain, Kuwait, Qatar, Oman, United Arab Emirates, and Yemen) during a six-month period. The diagnosis of ACS included unstable angina (UA) and non-ST-segment myocardial infarction (NSTEMI), collectively referred to here as non-STsegment elevation ACS (NSTEACS) or ST-segment elevation myocardial infarction (STEMI). The study received ethical approval from the institutional ethical bodies in all participating countries. There were no exclusion criteria. Full details of the methods have been previously published.^{17,18} Diagnosis of the different types of ACS and definitions of data variables were based on the American College of Cardiology clinical data standards.¹⁹

Patients were evaluated for the presence of MetS based on the most recent definition that was outlined in a joint statement of the IDF Task Force on Epidemiology and Prevention, NHLBI, AHA, World Heart Federation, International Atherosclerosis Society, and International Association for the Study of Obesity. Patients were classified as having MetS when at least three of the following were present: (1) abdominal obesity (waist circumference in Middle Eastern and Mediterranean populations: men \geq 94 cm and women \geq 80 cm); (2) high triglyceride level (\geq 150 mg/dL [\geq 1.7 mmol/L]); (3) low highdensity lipoprotein cholesterol level (<40 mg/dL [<1.04 mmol/L] for men and <50 mg/dL [<1.30 mmol/L] for women); (4) high BP (systolic \geq 130 mm Hg or diastolic \geq 85 mm Hg or taking antihypertensive medication); and (5) high fasting plasma glucose concentration (\geq 100 mg/dL [\geq 6.10 mmol/L]) or a clinical history of diabetes mellitus.¹⁶

Statistical Analysis

Baseline and clinical characteristics of patients are presented as median and 25th and 75th percentiles for continuous variables, whereas frequency distribution and percentages are presented for categoric variables. Independent t tests were used for comparing continuous variables, and chi-square tests for categoric variables have been used for comparing MetS vs non-MetS cases. A univariate logistic regression was performed for all important variables, and important variables were considered multivariate logistic regression analysis to examine the importance of MetS in heart failure (HF) patients. Chi-square test was performed. A $P \le .05$ was considered statistically significant. All data analyses were carried out using the Statistical Package for Social Sciences version 14 (SPSS Inc, Chicago, IL).

RESULTS

Study Population Characteristics

Of the 6701 patients studied, 3108 patients (46%) had MetS. Patients with MetS were more likely to present with NSTEACS, NSTEMI (37% vs 28%), or UA (33% vs 25%) than STEMI (30% vs 47%) (all *P*=.001).

Presenting Symptoms and Baseline Clinical Characteristics

Overall, MetS patients with ACS were three years older than those without MetS, more likely to be female, and less likely to be smokers. Patients with MetS were more likely to have a history of previous coronary artery disease and prior coronary revascularization and were more likely to be taking aspirin therapy. Patients with MetS were also more likely to have renal impairment at presentation. The admission heart rate and systolic and diastolic BPs were significantly higher in MetS patients. MetS patients were more likely to have Killip class >1 on admission. While MetS patients were more likely to present with atypical presentations, they were more likely to present earlier than patients without MetS. MetS patients had higher first blood sugar level at admission, serum creatinine, and triglyceride levels when compared when patients without MetS. Subset analysis of patients with STEMI, NSTEMI, and UA demonstrated similar clinical

	No Metabolic	Metabolic	
	Syndrome	Syndrome	P VALUE
No. %	3593 (54)	3108 (46)	
Age, mean, y	55±13	58±11	<.001
Female, %	15	35	<.001
Diabetes mellitus, %	18	67	<.001
Hypertension, %	28	76	<.001
Dyslipidemia, %	20	46	<.001
Smoking, %	43	32	<.001
Prior aspirin use, %	32	53	<.001
Family history of CAD, %	13	14	.43
Prior CAD, %	38	55	<.001
Prior revascularization, %	11	21	<.001
Renal impairment, %	15	21	<.001
At presentation			
Heart rate, beats per min	84±22	87±32	<.001
Systolic BP, mm Hg	135±30	145 ± 31	<.001
Diastolic BP, mm Hg	83±19	85±18	<.001
Body mass index, mean	26±4	29±6	<.001
Killip class >1%	20	25	<.001
GRACE risk score			
Low, %	45	38	.001
High, %	27	33	
Laboratory results			
Total cholesterol	5±2	5±3	.33
Serum triglyceride	1.5 ± 1	2.3±4	.001
First blood sugar	9.4±8	12.5 ± 12	.001
First serum creatinine	103 ± 78	$114{\pm}110$.001

Table I. Clinical Characteristics, Management, and Hospital Outcomes in Patients With and Without the Metabolic Syndrome Presenting With Acute Coronary Syndrome

characteristics and risk factor findings regardless of the admission diagnosis (Table I).

A subset analysis of the interaction between age at presentation with ACS and sex was performed (Figure 1). The prevalence of MetS among the different age groups was consistently higher among women when compared with men and was >70%among women in the age group of 50 to 79 years.

Treatment

In-Hospital. Overall, aspirin, clopidogrel, and β -blockers use was high and glycoprotein IIb/IIIa inhibitor use was low and comparable between the two groups. Patients with MetS were more likely to receive low molecular weight heparin than unfractionated heparin; they were also more likely to receive angiotensin-converting enzyme (ACE) inhibitors, angiotensin receptor blockers (ARBs), and statins. Thrombolytic therapy was the primary reperfusion strategy in our registry, and no differences were observed in its use between the two groups (Table II).

At Discharge. Patients with MetS were more likely to be prescribed aspirin, clopidogrel, ACE inhibitors, ARBs, and statins at discharge. There was no difference in β -blocker prescription at discharge between the two groups.

In-Hospital Outcome

In all ACS patients, there were no differences in mortality, cardiogenic shock, major bleeding, or recurrent myocardial infarction (MI) between the two groups, although MetS patients were more likely to develop HF and recurrent myocardial ischemia. In NSTEMI, MetS patients had more HF and recurrent myocardial ischemia than non-MetS patients, with no differences in the incidence of other complications between the groups. In UA, MetS patients had comparable complication rates to non-MetS patients. In the STEMI group, there were no differences between the two groups regarding mortality, cardiogenic shock, or major bleeding complications, although MetS patients had a significantly higher incidence of recurrent MI, stroke, and HF (Table II).

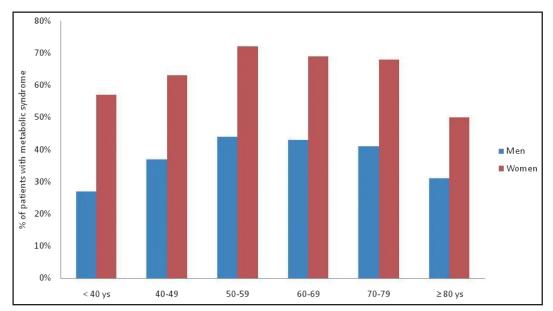


Figure 1. Prevalence of metabolic syndrome stratified according to age and gender.

Multivariate Predictors of HF

Multivariate regression demonstrated age, female sex, diabetes mellitus (odds ratio [OR], 1.6; 95% confidence interval [CI], 1.31–1.88; P<.001), and hypertension (OR, 1.4; 95% CI, 1.16–1.69; P<.001) to be independent predictors for HF complications. MetS was not an independent predictor of HF (OR, 0.082; P=.08) (Figure 2).

DISCUSSION

The present study is the largest registry to date that reports the prevalence of MetS among patients with ACS and is the first that applies the new criteria of MetS that was recently jointly approved by multiple international associations. MetS was highly prevalent among Middle Eastern patients presenting with ACS (46%). ACS patients with MetS were more likely to present with atypical presentations and had higher Killip class. Although MetS in ACS patients was not associated with increased in-hospital mortality, it was associated with increased risk of congestive HF. This increased risk was mainly attributed to diabetes mellitus and hypertension, as MetS was not an independent predictor of HF. In STEMI, patients with MetS had double the risk of developing stroke and recurrent MI.

Prevalence and Characteristics of MetS Among Patients With ACS

Few studies reported the prevalence of MetS, as defined by the NCEP-ATP III, NHLBI/AHA, and/ or IDF criteria in patients with ACS,^{2–15} (Table III),

which were mostly registry data conducted in North America, Europe, Japan, and Australia and included a small number of patients (107-633 patients) with an MetS prevalence ranging from 20.8% to 79.1%. Our study confirms this high prevalence of MetS (46%) using the new MetS definition in a larger unselected population (6701 patients). Patients with MetS were consistently younger (age range, 67.8 to 70 years) when compared with non-MetS patients in almost all studies except two (Yasar¹⁰ and Dohi¹² and colleagues). The current study extends these observations in an overall younger population with a mean age of 56 years (55 years with MetS vs 58 years without MetS). Moreover, it extends the observations of Zeller and colleagues³ of higher female and lower smoking prevalence among MetS patients. Our study, based on an unselected large population of ACS patients in the Middle East reports the highest prevalence of diabetes mellitus (40%) when compared with previous reports, which ranged from 19% (in Australia)⁸ to 34% (in Japan).⁹ In a post hoc analysis of the Gruppo Italiano per lo Studio della Sopravvivenza nell'Infarto Miocardico (GISSI-Prevenzione) trial, which included 10,384 patients who had recent MI (within 3 months), Levantesi and colleagues¹³ reported a 29.3% and 21% prevalence of MetS and diabetes mellitus, respectively. To further expand on the current observations, additional large-scale multi-ethnic studies using the new definition of MetS are urgently needed.

	No Metabolic	Metabolic	
	Syndrome	Syndrome	P VALU.
Presentation >12 h, %	33	28	.007
Atypical presentation, %	18	23	<.001
Final diagnosis			
STEMI, %	47	30	.001
Non-STEMI, %	28	37	
Unstable angina, %	25	33	
Coronary angiogram, %	18	19.5	.09
In-hospital therapy			
Thrombolytic therapy, % ^a	57	59	.35
Primary PCI ^a	5	3	.006
Aspirin, %	97	98	.62
Clopidogrel, %	54	53	.95
Unfractionated heparin, %	52	42	.001
Low molecular weight heparin, %	42	53	<.001
Glycoprotein IIb/IIIa inhibitors, %	10	11	.07
Angiotensin-converting enzyme inhibitor, %	64	66	.03
Angiotensin receptor blocker, %	3	8	.001
β-blockers, %	64	66	.09
Therapy at discharge			,
Aspirin, %	93	95	.005
Clopidogril, %	48	51	.004
Angiotensin receptor blocker, %	4	10	.001
Angiotensin-converting enzyme inhibitor, %	70	69	.99
β-Blockers, %	75	76	.07
Statins, %	79	83	.001
Hospital outcomes			
Recurrent ischemia, %	8	10	.01
Reinfarction,%	2	2.4	.37
Heart failure, %	15	18	.002
Cardiogenic shock, %	5.5	5	.20
Major bleeding, %	0.8	0.7	.55
Stroke, %	0.6	0.9	.56
Mortality, %	3	2.4	.22
Hospital stay, d	5.5±4	5.7±5	.19

MetS and HF

One of the main findings of the current study is higher Killip class (Killip class >1, overall 25% vs 20% in non-MetS patients) and higher incidence of HF complications among patients with MetS (overall ACS group, 18% vs 15%). The increased HF risk mainly occurs in patients with MI only, STEMI (20% vs 14%), and NSTEMI (18% vs 14%). The incidence of HF among MetS patients with UA was not increased (15% vs 15.2%, *P*=.09). However, DM and hypertension (not MetS) were independent predictors of HF. Several investigators reported the prevalence of HF complications among MetS patients with ACS. The Observatoire des Infarctus de Cote-d'Or Survey³ reported higher Killip class (Killip class >1, 26% vs 16%) and HF complications among MetS patients. In that survey (n=633), the overall reperfusion therapy rate was low and was even lower among MetS patients (49% vs 38%). The investigators hypothesized that this increased risk may be related to fasting hyperglycemia, and this hypothesis was supported by earlier studies reporting increased risk of pump failure in patients with MI and hyperglycemia, even after adjustment for age.^{20–22} Another explanation however, may be lower reperfusion therapy in the MetS group. More recently, in a case-control study of 461 patients with MI, Takeno and colleagues⁹ also reported increased risk of HF and revascularization rates among MetS patients despite high reperfusion therapy rates >70% and lack of significant differences in left ventricular ejection fraction

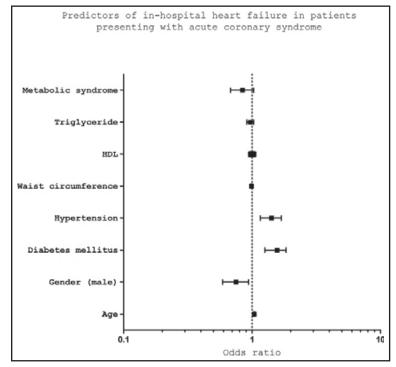


Figure 2. Predictors of in-hospital heart failure in patients presenting with acute coronary syndrome. HDL indicates high-density lipoprotein.

when compared with non-MetS patients. They attributed increased risk of these two complications (HF and revascularization) to the presumed development of new coronary lesions or restenosis during the follow-up period (19 months). On the other hand, in a 36-month follow-up study of patients with recent MI by Laventsei and colleagues,¹³ patients with diabetes mellitus and not MetS had increased risk of HF when compared with controls. Clavijo and colleagues⁴ reported increased infarct size in 405 patients with MetS and acute MI. They determined the infarct size by peak creatine kinase-MB (CK-MB) levels. However, Zeller³ and Yasar¹⁰ and colleagues did not observe higher peak CK and CK-MB levels or lower left ventricular ejection fraction in patients with MetS when compared with those without MetS. In conclusion, MetS patients presenting with MI (not UA) are at increased risk for HF, which appears to not be related to worse systolic function and appears to be related to the presence of diabetes and hypertension. Further studies specifically designed to elucidate the pathogenesis of this association are needed.

MetS and Prognosis

Several investigators (using either IDF or ATP III or both definitions of MS) reported in-hospital and fol-

low-up mortality rates among MetS patients. In a 17.6-month follow-up of 461 patients, Takeno and colleagues9 reported MetS (using the ATP III definition) no independent predictor of cardiovascular death (relative risk, 0.96; 95% CI, 0.3-3.30). Boulon and colleagues² reported an increase in total mortality in MetS patients at 19-month follow-up (using the IDF definition, among 480 ACS patients); however, this increase was in part due to an increase in noncardiovascular deaths. On the other hand, Zeller and colleagues³ reported an increased in-hospital mortality rate among MetS patients (10.7% vs 3.8%; P=.01). However, after adjustment for variables, MetS was no longer an independent predictor of mortality. On the other hand, in the Myocardial Ischemia Reduction With Aggressive Cholesterol Lowering (MIRACLE) trial,¹⁵ patients with ACS and MetS had increased all-cause mortality and an increased rate of reaching the primary end point (death, nonfatal MI, cardiac arrest, or recurrent unstable myocardial ischemia) during a 16-week follow-up. More recently, Dohi and colleagues¹² reported an increased risk of cardiovascular mortality among MetS patients in a 10-year follow-up of 383 Japanese patients.

The current study, which involved a much larger cohort of patients of different ethnicities than those previously reported, supports lack of increased

Table III. Reported Studies of the MetS and ACS	he MetS and ACS					
Author, Year (Origin)	Study Design	No. of Patients	Previous DM, %	Previous MetS Definition, %	Follow-Up	OUTCOME
Boulon, 2006 (France) ²	Registry	480	21.6	ATP III: 20.8; IDF: <i>27.7</i>	19 mo	Increase in total mortality in the MetS group (5.2% vs 1.4%, P<.01) due in part to an increase in noncardiovascular death. No increase in minor or major cardiovascular
Zeller, 2005 (France) ³	Registry	633	26	ATP III: 46	In-hospital	events. Independent predictor of heart failure (OR, 2.13; 95% CI, 1.28–3.57; P=.003)
Clavijo, 2006 (United States) ⁴	Registry	405 ^a	۵	ATP III: 41 ^b	In-hospital	Associated with larger infarct size determined by cardiac enzymes (P <.001), more complications (21.1% vs 9.2%, P =.003), and a marked increase of acute renal failure
Celik, 2006 (Turkey) ⁵	Registry	283 ^a	م	ATP III: 25	In-hospital	Independent predictor of poor myocardial perfusion grade post-primary PCI (OR, 254: 95% CI: 135-475: P=003)
Chung, 2007 (United States) ⁶	Registry	223°	13	ATP III: 47	Not provided	MetS highly prevalent in young (\leq 45 y)
Milionis, 2007 (Greece) ⁷	Registry	136°	2.2	ATP III: 40.4	In-hospital	With ML. MetS strongly associated with ACS in young persons, independent of other risk factors. Smoking and a positive family history of premature CAD in young persons with MetS had an incremental effect on the
Prasad, 2009 (Australia) ⁸	Registry	107ª	19	IDF: 54 ATP III: 49	In-hospital	odds of having AUS (UK, /.1.2; 95%) CI, 2.42–20.96; <i>P</i> <.001). Same extent of coronary atherosclerosis and coronary flow post-PCI. No differences in systolic function but greater diastolic dysfunction with
Tokeno, 2008 (Japan) ⁹	Registry	461 ^a	34	ATP III: 37	17.6 mo	Mets (P <.05). Independent predictor of heart failure (OR, 2.60; 95% CI, 1.01–6.66; P =.04) and repeat revascularization (OR, 2.1; 95% CI, 1.27–3.47; P <.01).

Table III. Reported Studies of the MetS and ACS (Continued)	f the MetS and ACS (Co	ntinued)				
Author, Year (Origin)	Study Design	No. of Patients	Previous DM, %	Previous MetS Definition, %	Follow-Up	OUTCOME
Yasar, 2009 (Turkey) ¹⁰	Registry	116	27	ATP III: 55.2	In-hospital	MetS patients have lower rates of TIMI grade 3 flow (OR, 3.545; 95% CI, 1.064–11.808; <i>P</i> =.03) and higher corrected TIMI frame counts after thrombolytic therapy for AMI.
Koutsovasilis, 2009 (Greece) ¹¹	Registry	211	30.3	ATP III: 72.5 NHLBI/AHA: 81.2 IIDF: 79.1	Compared with control	Only IDF-defined MetS was significantly associated with ACS (OR, 2.23; 95% CI, 1.3–3.82; P=.003). Of the MetS components, only waist circumference was independently associated with ACS (OR, 1.045; 95% CI, 1.014–1.078: P=.005).
Dohi, 2009 (Japan) ¹²	Registry	384 (with complete revascularization)		ATP III: 42.5	10.4±3.4 y	Independent predictor of long-term all-cause (OR, 1.62; 95% CI, 1.01–2.59; <i>P</i> =.046) and cardiovascular death (OR, 2.40; 95% CI, 1.16–4.94; <i>P</i> =.018).
Levantesi, 2005 (Italy) ^{d13}	RCT-post hoc (GISSI-Prevenzione)	10,384	20.6	ATP III: 29.3	3.5 y	Increased probability of death (OR, 1.29; 95% CI, 1.1–1.51; <i>P</i> =.002) and cardiovascular events (OR, 1.23; 95% CI, 1.06–1.42; <i>P</i> =.005) at follow-up; however, hospitalization for heart failure
Germain, 2004 (Europe-United States) ^{e14}	RCT-post hoc (4S and AFCAPS/ TexCAPS)	5179 (not all with prior ACS)	Ъ	ATP III: 20.6–46 5–5.4 y	5-5.4 y	48: Independent predictor of major adverse cardiac events (OR, 1.5; 95% CI, 1.2–1.8). AFCAPS/TexCAPS: Independent predictor of major adverse cardiac events (OR, 1.4; 95% CT 1.04–1.9)
Schwatrz, 2005 (United States) ¹⁵	RCT-Post hoc (MIRACL)	3083	23	ATP III: 38	16 wk	Independent predictor of major adverse cardiac events at 16-week follow-up (OR, 1.49; 95% CI, 1.24–1.79; <i>P</i> <.0001). Independent predictor of all-cause mortality (OR, 1.49; 95% CI, 1.04–2.14; <i>P</i> =.029).

Table III. Reported Studies of the MetS and ACS (Continued)	of the MetS and ACS	(Continued)				
Author, Year (Origin)	Study Design	No. of Patients	Previous DM, %	Previous MetS Definition, %	Follow-Up	OUTCOME
Current study (6 Middle Eastern countries)	Registry	6701	40	New unified: 46	In-hospital	Increased risk of heart failure and recurrent myocardial ischemia but not death. STEMI patients have double the risk of recurrent myocardial infarction and stroke.
Abbreviations: ACS, acute coronary syndrome; ATP III IDF, International Diabetes Federation; MIRACL, the intervention; RCT, randomized controlled trial; STEM patients only. ^b Nondiabetic with metabolic syndrome (Italiano per lo Studio della Sopravvivenza nell'Infarto N analysis of two randomized trials (the Scandinavian Sin	ronary syndrome; ATP ederation; MIRACL, 1 ed controlled trial; STJ ith metabolic syndrom opravvivenza nell'Infart ials (the Scandinavian	¹ III, National Chc the Myocardial Isc. EMI, ST-elevation ie (MetS). ^o The stu o Miocardico [GR Simvastatin Surviv	alesterol Educatic hemia Reduction myocardial infar ady included onl. SSI-Prevenzione] al Study [4S] an.	nn Program Adult Treat n With Aggressive Chole ction; TIMI, thromboly y patients younger than Trial), which included d the Air Force/Texas (nent Panel III; CI, c sterol Lowering trial; sis in myocardial inf 45 years. ^d Post hoc patients who had a r boronary Atheroscler	Abbreviations: ACS, acute coronary syndrome; ATP III, National Cholesterol Education Program Adult Treatment Panel III; CI, confidence interval; DM, diabetes mellitus; IDF, International Diabetes Federation; MIRACL, the Myocardial Ischemia Reduction With Aggressive Cholesterol Lowering trial; OR, odds ratio; PCI, percuraneous coronary intervention; RCT, randomized controlled trial; STEMI, ST-elevation myocardial infarction; TIMI, thrombolysis in myocardial infarction. ^a Acute myocardial infarction (AMI) patients only. ^b Nondiabetic with metabolic syndrome (MetS). ^c The study included only patients younger than 45 years. ^d Post hoc analysis of randomized trial (the Gruppo Italiano per lo Studio della Sopravvivenza nell'Infarto Miocardico [GISSI-Prevenzione] Trial), which included patients who had a recent MI (within 3 months). ^e Post hoc analysis of two randomized trials (the Scandinavian Sinwastatin Survival Study [4S] and the Air Force/Texas Coronary Atherosclerosis Prevention Study [AFCAPS/TexCAPS]).

hospital mortality risk among MetS patients. On the other hand, diabetes mellitus was an independent predictor of increased mortality. However, patients with MetS were at increased risk for recurrent myocardial ischemia in the overall ACS group (P=.01). Furthermore, it reports for the first time that STEMI patients with MetS have double the risk of stroke and recurrent MI. Increased risk of recurrent myocardial ischemia, MI, and stroke may be attributed to prothrombotic state associated with MetS.²³⁻²⁵ Increased levels of plasminogen activator inhibitor-1 (PAI-1) are reported in patients with MetS. PAI-1 inhibits spontaneous and pharmacologically induced fibrinolysis. In patients with MetS, fibrinolytic dysfunction together with increased platelet activation may lead to a prothrombotic state and impaired response to thrombolytic agents. Arenillas and colleagues²⁴ investigated the impact of MetS on the response to systemic thrombolysis in patients with acute middle cerebral artery ischemic stroke and reported that MetS was associated with a poor response to thrombolysis. They suggested that insulin resistance might exert independent deleterious actions that contribute to limit the efficacy of thrombolytic therapy. More recently, Mente and colleagues,²⁶ using the IDF and ATP III criteria among patients in the INTERHEART study, reported increased risk of MI among patients with MetS; however, this risk does not appear to be greater than the risk conferred by its component factors. It appears that the behavior of MetS as a cardiovascular risk factor is similar to that of hypertension and different from diabetes mellitus. MetS increases the risk of developing MI and, subsequently, HF and recurrent myocardial ischemia complications without increased in-hospital mortality. Further studies, are required to clarify the prognostic impact of MetS among MI patients.

Study Limitations

Our data were collected from an observational study, which is a limitation. The fundamental limitations of observational studies cannot be eliminated because of the nonrandomized nature and unmeasured confounding factors. However, welldesigned observational studies provide valid results and do not systematically overestimate the results compared with the results of randomized controlled trials. Long-term follow-up is needed in both groups to consolidate our findings.

CONCLUSIONS

MetS is highly prevalent among patients presenting with ACS in six Middle Eastern countries. MetS is

associated with increased risk for development of HF and recurrent myocardial ischemia without an increase in hospital mortality. STEMI patients have double the risk of recurrent MI and stroke. To further expand on our understanding of the prevalence and outcome of MetS in patients with ACS, further large-scale studies in several ethnicities using the new definition of MetS are urgently needed.

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