# **BMJ Open** Simple risk score based on the China Acute Myocardial Infarction registry for predicting in-hospital mortality among patients with non-ST-segment elevation myocardial infarction: results of a prospective observational cohort study

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# ABSTRACT

**Objectives** To simplify our previous risk score for predicting the in-hospital mortality risk in patients with non-ST-segment elevation myocardial infarction (NSTEMI) by dropping laboratory data.

Design Prospective cohort.

**Setting** Multicentre, 108 hospitals across three levels in China.

**Participants** A total of 5775 patients with NSTEMI enrolled in the China Acute Myocardial Infarction (CAMI) registry.

**Primary outcome measures** In-hospital mortality. **Results** The simplified CAMI-NSTEMI (SCAMI-NSTEMI) score includes the following nine variables: age, body mass index, systolic blood pressure, Killip classification, cardiac arrest, ST-segment depression on ECG, smoking status, previous angina and previous percutaneous coronary intervention. Within both the derivation and validation cohorts, the SCAMI-NSTEMI score showed a good discrimination ability (C-statistics: 0.76 and 0.83, respectively); further, the SCAMI-NSTEMI score had a diagnostic performance superior to that of the Global Registry of Acute Coronary Events risk score (C-statistics: 0.78 and 0.73, respectively; p<0.0001 for comparison). The in-hospital mortality increased significantly across the different risk groups.

**Conclusions** The SCAMI-NSTEMI score can serve as a useful tool facilitating rapid risk assessment among a broader spectrum of patients admitted owing to NSTEMI. **Trial registration number** NCT01874691.

#### INTRODUCTION

Acute myocardial infarction (AMI) is a leading cause of mortality worldwide, accounting for 2.4 million mortalities in the USA and more than 4 million mortalities in Europe and northern Asia.<sup>1</sup> AMI is commonly divided into ST-segment elevation myocardial infarction (STEMI) and non-STEMI (NSTEMI) based on the presence or absence of ST-segment

# Strengths and limitations of this study

- We developed and validated a simple risk score for predicting in-hospital mortality in Asian patients with non-ST-segment elevation myocardial infarction (NSTEMI).
- The simplified China Acute Myocardial Infarction (SCAMI)-NSTEMI score can be used at the first medical contact.
- The SCAMI-NSTEMI score requires external validation in a large independent cohort.
- The diagnostic performance between the SCAMI-NSTEMI and existing risk scores should be compared in future studies.

elevation on ECG. NSTEMI compromises approximately 70% of all myocardial infarction cases.<sup>2</sup> Patients with NSTEMI have varying prognoses. Accurate risk stratification of patients with NSTEMI is important, as it can not only help identify high-risk patients who will benefit from prompt revascularisation but also avoid inappropriate use of aggressive treatment among low-risk patients.

Many risk scores have been developed for estimating mortality risk in patients with acute coronary syndrome (ACS), including the Global Registry of Acute Coronary Events (GRACE) risk score,<sup>3</sup> Acute Coronary Treatment and Intervention Outcomes Network (ACTION) risk score,<sup>4</sup> Canadian ACS risk score,<sup>5</sup> and ProACS risk score.<sup>6</sup> However, these scores included only a small number of Chinese patients. Additionally, to our knowledge, no risk scores have been focused on patients with NSTEMI. To bridge the knowledge gap, our team previously developed and validated a novel risk score for predicting the

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in-hospital mortality risk among patients with NSTEMI based on the China Acute Myocardial Infarction (CAMI) registry (ie, CAMI-NSTEMI score).<sup>7</sup>

Early risk assessment is of clinical significance among patients with NSTEMI. A large-scale meta-analysis found that among high-risk patients with NSTEMI, an early invasive strategy was associated with a lower in-hospital mortality.<sup>8</sup> In addition, there is evidence indicating that among high-risk patients with NSTEMI, coronary angiography early within the initial 12 hours was associated with better outcomes.<sup>9</sup>

However, our original CAMI-NSTEMI score includes the white blood cell (WBC) count and creatinine level which might restrict its use at the time of the first medical contact before obtaining laboratory test results. Delayed risk stratification may have an adverse impact on patient outcomes, especially for these high-risk patients.

The objective of our study was to collect laboratory data from the previous CAMI-NSTEMI risk score and consequently develop and validate a simplified risk score which can save time in terms of score calculation and allow for early risk assessment.

### METHODS CAMI registry

The CAMI registry was designed as an integrated research and educational platform to reflect patients with AMI in China. The detailed description of the CAMI study design has been published previously.<sup>10</sup> In brief, the CAMI registry was a prospective multicentre registry that aimed to reflect the patient characteristics, medical care and management of Chinese patients with AMI. Multiteams with various roles cooperated to ensure smooth execution of the project. A total of 108 hospitals across three levels (provincial, prefectural and county) from 27 provinces and 4 municipalities participated in the project which assures a good representation of the contemporary cohort of AMI in China. The CAMI registry enrolled patients within 7 days of ischaemia symptoms, who had a diagnosis of AMI according to the Third Universal Definition of Myocardial Infarction.<sup>11</sup> Patient characteristics, physical examination results and laboratory test results were collected and submitted to an electronic web-based system.

#### **CAMI-NSTEMI score**

We previously developed and validated a novel risk score, that is, CAMI-NSTEMI risk score, to predict the in-hospital mortality risk among patients with NSTEMI.<sup>7</sup> Briefly, the CAMI-NSTEMI score was derived from a cohort of patients with NSTEMI registered in the CAMI registry between January 2013 and September 2014. Data were extracted by trained researchers using standard definitions to reduce measure and report bias. We excluded those with missing or invalid data on age, body mass index (BMI), admission diagnosis and in-hospital outcome and those diagnosed with left bundle branch

block (LBBB) and finally included 5775 patients to develop and validate the risk score. The primary outcome of our study was in-hospital mortality, which was evaluated by trained cardiologists during hospitalisation. We did not take actions to blind assessment and predictors of the outcome because of the hard outcome (in-hospital mortality) measure. Using a multivariable logistic regression model, we identified 11 independent predictors of in-hospital mortality: age, BMI, systolic blood pressure, Killip classification, cardiac arrest, ST-segment depression on ECG, serum creatinine level, WBC count, smoking status, previous angina and previous percutaneous coronary intervention. Although cardiac biomarkers are more available than the serum creatinine level and WBC count, the CAMI-NSTEMI risk score did not include cardiac biomarkers because the CAMI registry was a multicentre registry including 108 participating hospitals. The type of cardiac enzymes and the corresponding normal range differed across hospitals. Including cardiac biomarkers may reduce the diagnostic performance of the risk score. Therefore, based on these 11 predictors, we developed and validated the CAMI-NSTEMI risk score to predict the in-hospital mortality risk among the patients with NSTEMI. The detailed definition regarding each variable was described in the protocol.<sup>10</sup>

#### **Statistical analysis**

Continuous data were presented as mean ± SD and were compared using the Student's t-test. Categorical variables were summarised as counts and percentages and were compared using the  $\chi^2$  test or Fisher's exact test, as appropriate. All analyses were performed using the SAS V.9.4 system (SAS Institute). All p values were two-tailed, and a p value of <0.05 was considered statistically significant. We did not calculate the sample size, as this was a registry-based retrospective study, and we wanted to enrol as many patients as possible. We used simple imputation methods to deal with missing data, which were imputed with the median or mode values of the available cases. The methods for developing and validating simplified CAMI-NSTEMI (SCAMI-NSTEMI) risk score were the same as those for the original CAMI-NSTEMI score, which have been reported previously.<sup>7</sup> Briefly, the entire cohort was divided into the derivation (n=4332) cohort and the validation (n=1443) cohort chronologically. To develop the simplified risk score, we first fitted variables with p values of < 0.25 in the univariable selection into the logistic multivariable regression model. The WBC count and creatinine level were not included in the multivariable model. Thereafter, the multivariable model was constructed using stepwise variable selection with entry and exit criteria (p<0.05). We attributed integer numbers to each variable according to the coefficient. The area under the receiver-operating characteristic curve (AUC) and Hosmer-Lemeshow (HL) goodness-of-fit test were used to assess discrimination and calibration of the risk score. The scoring system was divided into three risk groups (low, intermediate and high risks) according to tertiles. We compared the diagnostic performance between the CAMI-NSTEMI score and GRACE score by calculating the AUC, net reclassification improvement (NRI) and integrated discriminatory index (IDI).<sup>12</sup>

#### Patient and public involvement

We did not involve patients or the public directly in our work.

### RESULTS

## **Patient characteristics**

From January 2013 to September 2014, a total of 6209 patients diagnosed with NSTEMI were registered in the CAMI registry. We excluded 393 patients owing to incomplete or invalid data on age, BMI, admission diagnosis and in-hospital outcome. We also excluded 41 patients with LBBB and finally included 5775 patients. A total of 342 patients (5.9%) died during hospitalisation. As shown in table 1, most patient characteristics were imbalanced between the groups. The patients who died during hospitalisation were older and more likely to present with diabetes mellitus or hypertension than the in-hospital survivors. Further, they had a lower BMI, and a lower proportion of male and current smokers.

Regarding clinical presentation, the patients who died during hospitalisation had higher heart rate, lower systolic blood pressure and higher Killip classification than the in-hospital survivors. A higher proportion of patients who died during hospitalisation presented with ST-segment depression on ECG and cardiac arrest. The laboratory test showed that the patients who died during hospitalisation had higher platelet count, serum creatinine level, WBC count and serum potassium level but lower haemoglobin level than the in-hospital survivors (table 1).

#### **SCAMI-NSTEMI score**

A univariable analysis was first performed to explore the unadjusted association between each baseline predictor and outcome (online supplementary table 1). Variables with p values of <0.25 were selected to entered into the multivariable logistic regression model. In the multivariable analysis, a total of nine variables were identified as independent risk factors of in-hospital mortality and used to develop the SCAMI-NSTEMI risk model: age, BMI, systolic blood pressure, Killip classification, cardiac arrest, ST-segment depression on ECG, smoking status, previous angina and previous percutaneous coronary intervention (table 2). We attributed integer numbers to each variable based on the coefficient in the multivariable logistic regression model (table 3) and established the SCAMI-NSTEMI risk score. The in-hospital mortality risk corresponding to each point is shown in online supplementary table 2.

Within the derivation cohort (n=4332), 248 patients died during hospitalisation. The AUC for the SCAMI-NSTEMI model was 0.7771 (95% CI: 0.7472 to 0.8071) which was slightly higher than that for the SCAMI-NSTEMI

 Table 1
 Baseline characteristics of patients who died vs

 survived

	In-hospital survivors (n=5433)	In-hospital deaths (n=342)	P value	
Age (years)	64.92±11.98	72.13±11.16	<0.01	
Male	3754/5433 (69.1)	187/342 (54.7)	<0.01	
BMI (kg/m <sup>2</sup> )	24.09±3.05	23.02±3.11	<0.01	
DM	1249/5418 (23.1)	98/342 (28.7)	0.02	
Hypertension	3154/5423 (58.2)	209/342 (61.1)	0.28	
Hyperlipidaemia	456/5421 (8.4)	14/342 (4.1)	<0.01	
LVEF (%)	55.10±11.76	46.75±13.01	<0.01	
Previous angina	2074/5408 (38.4)	144/342 (42.1)	0.17	
Previous MI (>1 month)	585/5411 (10.8)	63/342 (18.4)	<0.01	
Previous heart failure	263/5412 (4.9)	48/342 (14.0)	<0.01	
Previous PCI	358/5400 (6.6)	12/342 (3.5)	<0.01	
Previous CABG	48/5411 (0.9)	4/342 (0)	0.55	
Previous stroke	542/5410 (10.0)	46/342 (13.5)	0.05	
Previous renal dysfunction	137/5400 (2.5)	15/341 (4.4)	0.06	
Previous COPD	131/5382 (2.4)	16/340 (4.7)	0.02	
Family history of premature CAD	168/5417 (3.1)	6/342 (1.8)	0.13	
Previous peripheral vascular disease	64/5406 (1.2)	5/342 (1.5)	0.60	
Smoking status			<0.01	
Current smoker	1967/5406 (36.4)	60/339 (17.7)		
Previous smoker	721/5406 (13.3)	46/339 (13.6)		
Non-smoker	2718/5406 (50.3)	233/339 (68.7)		
Prior use of medication	on (within 1 week)			
Aspirin	1003/5405 (18.6)	69/339 (20.4)	0.42	
Thienopyridines	338/5386 (6.3)	29/339 (8.6)	0.11	
Statins	764/5327 (14.3)	54/334 (16.2)	0.36	
HR (beats/min)	79.19±19.90	89.97±25.39	<0.01	
SBP (mm Hg)	134.64±25.67	121.87±28.71	<0.01	
Killip classification			<0.01	
I	3873/5393 (71.8)	127/339 (37.5)		
II	989/5393 (18.3)	93/339 (27.4)		
III	382/5393 (7.1)	52/339 (15.3)		
IV	149/5393 (2.8)	67/339 (20.3)		
ST segment depression	2917/5340 (54.6)	223/335 (66.6)	<0.01	
Heart arrest	29/5404 (0.5)	14/340 (4.1)	<0.01	
Time to hospital			0.68	
1–7 days	2139/5345 (40.0)	140/334 (41.9)		
12–24 hours	758/5345 (14.2)	40/334 (12.0)		
6–12 hours	772/5345 (14.4)	54/334 (16.2)		
<6 hours	1676/5345 (31.4)	100/334 (30.5)		
PLT (10 <sup>9</sup> /L)	208.26±72.23	217.53±117.26	0.16	
Hb (g/L)	131.81±21.95	121.82±26.27	<0.01	

Continued

Table 1   Continued						
	In-hospital survivors (n=5433)	In-hospital deaths (n=342)	P value			
WBC (10 <sup>9</sup> / L)	9.06±3.43	12.00±5.84	<0.01			
Cr (µmol/L)	88.33±60.49	131.09±104.95	<0.01			
K⁺(mmol/L)	3.96±0.50	4.18±0.77	<0.01			

BMI, body mass index; CABG, coronary artery bypass grafting; CAD, coronary artery disease; COPD, chronic obstructive pulmonary disease; Cr, creatinine; DM, diabetes mellitus; Hb, haemoglobin; HR, heart rate; LVEF, left ventricular ejection fraction; MI, myocardial infarction; PCI, percutaneous coronary intervention; PLT, platelet count; SBP, systolic blood pressure; WBC, white blood cell count.

score (AUC: 0.7643; 95% CI: 0.7343 to 0.7943, p=0.0123 for comparison; figure 1A). Within the validation cohort (n=1443), 94 died during hospitalisation. The AUC value for SCAMI-NSTEMI model was 0.8614 (95% CI: 0.8173 to 0.9055), which was slightly higher than that for the SCAMI-NSTEMI score (AUC: 0.8286; 95% CI: 0.8286 to 0.8748, p=0.003 for comparison; figure 1B). Within the entire cohort, the AUC for the SCAMI-NSTEMI model and score was 0.7992 (95% CI: 0.7742 to 0.8243) and 0.7819 (95% CI: 0.7567 to 0.8072, p<0.001 for comparison), respectively (figure 1C). Good calibration was observed for both the risk model and score (HL test p value: 0.670 and 0.465, respectively).

The SCAMI-NSTEMI score ranged from 0 to 36. The in-hospital mortality risk associated with each point is shown in online supplementary table 2. To determine the risk categories of the SCAMI-NSTEMI risk score, we divided the patients into three groups according to guideline recommendations and the GRACE risk score.<sup>13</sup> The intertertile score range and event rate within each tertile are shown in

Table 2         Independent predictors of in-hospital death					
Predictors	OR	95% CI	P value		
Age (per 1-year increase)	1.027	1.014 to 1.041	< 0.0001		
BMI (per one 1 kg/m <sup>2</sup> increase)	0.946	0.903 to 0.990	0.0170		
SBP (per 1 mm Hg increase)	0.983	0.977 to 0.988	< 0.0001		
Killip classification	1.707	1.492 to 1.953	< 0.0001		
ST-segment depression	1.516	1.134 to 2.207	0.0049		
Heart arrest	3.103	1.270 to 7.578	0.0129		
Non-smoker vs current smoker	1.900	1.338 to 2.698	0.0003		
Ex-smoker vs current smoker	1.393	0.858 to 2.261	0.1804		
Previous MI	1.719	1.167 to 2.533	0.0061		
Previous PCI	0.334	0.148 to 0.753	0.0082		

BMI, body mass index; SBP, systolic blood pressure; MI, myocardial infarction; PCI, percutaneous coronary intervention

Table 3         Scores attributed to each variable						
Predictor	Categories	Score	Predictor	Categories	Score	
Age (years)	< 57	0	Killip classification	I	0	
	(57–66)	2		II	3	
	(66–75)	3		III	6	
	≥75	4		IV	9	
BMI (kg/ m²)	< 20.04	2	Heart arrest	No	0	
	(20.04–23.88)	1		Yes	6	
	(23.88–25.86)	1	Smoking status	Non-smoker	4	
	≥25.86	0		Ex-smoker	2	
SBP (mm Hg)	< 118.5	5		Current- smoker	0	
	(118.5–130)	4	Prior MI	No	0	
	(130–150)	2		Yes	3	
	≥150	0	Prior PCI	No	6	
ST segment depression	No Yes	0 2		Yes	0	

BMI, body mass index; MI, myocardial infarction; PCI, percutaneous coronary intervention; SBP, systolic blood pressure.

online supplementary table 3. Within the derivation cohort, the in-hospital mortality increased across the tertiles: 1.28% in tertile I (score range: 0–14), 3.33% in tertile II (score range: 15–18), and 10.53% in tertile III (score range:  $\geq$ 19) (p=0.001 for tertile II vs tertile I; p<0.001 for tertile III vs tertile I; p<0.001 for tertile III vs tertile I). Therefore, these three tertiles were defined as the low-risk, intermediate-risk and high-risk groups. Within the validation cohort, a similar trend was observed. The event rate was 1.04%, 2.65% and 15.21% in the low-risk, intermediate-risk groups, respectively.

# Comparison between the CAMI-NSTEMI score and GRACE risk score

We first compared the diagnostic performance between the SCAMI-NSTEMI score and the original CAMI-NSTEMI score. The AUC for the CAMI-NSTEMI score was higher than that for the SCAMI-NSTEMI score within the entire cohort (0.8080 vs 0.7819, p<0.0001 for comparison; online supplementary figure 1).

We then compared the diagnostic performance between the SCAMI-NSTEMI score and GRACE risk score within the entire cohort. The AUC for the SCAMI-NSTEMI score and GRACE risk score was 0.7819 (95% CI: 0.7567 to 0.8072) and 0.7272 (95% CI: 0.6995 to 0.7548), respectively, and the difference reached statistical significance (p<0.0001; online supplementary figure 2). The NRI and IDI for the SCAMI-NSTEMI score in relation to the GRACE score were 38.9% (p<0.0001) and 5.78% (p<0.0001), respectively.

# DISCUSSION

We developed and validated a simplified risk score to assess the in-hospital mortality risk of patients with NSTEMI.



Figure 1 ROC curves of SCAMI risk model and SCAMI risk score. (A) Within the derivation cohort, the C-statistic was 0.7771 (95% CI: 0.7472 to 0.8071) for SCAMI risk model and 0.7643 (95% CI: 0.7343 to 0.7943) for the SCAMI risk score. (B) Within the validation cohort, the C-statistic was 0.8614 (95% CI: 0.8173 to 0.9055) for SCAMI risk model and 0.8286 (95% CI: 0.7825 to 0.8748) for the SCAMI risk score. (C) Within the entire cohort, the C-statistic was 0.7992 (95% CI: 0.7742 to 0.8243) for SCAMI risk model and 0.7992 (95% CI: 0.7742 to 0.8243) for the SCAMI risk score. ROC, receiver operating characteristic curve; SCAMI, simplified China Acute Myocardial Infarction registry.

The simplified risk score incorporated nine variables, which are easily obtained during routine medical history-taking and bedside examination without the need to wait for laboratory test results. This allows timely risk stratification and treatment strategy selection, as well as risk assessment among a broader spectrum of patients with AMI, especially for those with missing variables during the first presentation. The SCAMI-NSTEMI score demonstrated good discrimination and calibration abilities, as well as better diagnostic performance, compared with the GRACE risk score.

#### Comparison with previous risk scores

Many risk scores have been developed to predict the short-term and long-term outcomes of patients with AMI, and the GRACE risk score is the most validated and commonly used risk prediction parameter by clinicians.<sup>3 14</sup> In addition, the GRACE risk score performed better than the Thrombolysis in Myocardial Infarction risk score in Chinese patients with NSTEMI.<sup>15</sup> Therefore, we compared the SCAMI-CAMI risk score with the GRACE risk score. These two scores differ in the following two aspects.

First, the SCAMI-NSTEMI risk score was derived from a more recent cohort. The original GRACE risk score was developed from a cohort of patients with ACS enrolled in the GRACE registry from 1 April 1999 to 31 March 2001. Patient profile and AMI management have evolved significantly over time, and it is reasonable to update the risk scores periodically.

Second, the SCAMI-NSTEMI score better represented unique prognostic factors among patients from Asia. While the GRACE registry was a multicentre registry covering the USA, Europe and Australia, only a few participants from Asia were enrolled. The risk factors of in-hospital mortality may differ across various ethnic groups. As Asia is the most populous continent worldwide, it is appropriate to develop a risk score more suitable for Asian patients. This is of clinical significance because NSTEMI affects a broad spectrum of patients with various prognoses, and a risk prediction parameter with high accuracy is important for the triage and management of patients with NSTEMI.

#### **Obesity and smoker's paradox**

Although obesity and smoking are well-established risk factors of coronary artery disease, our study found that the patients with a higher BMI had a lower in-hospital mortality than those with a normal BMI, and the current smokers had a lower in-hospital mortality than the current non-smokers. These phenomena are referred to as 'obesity paradox'<sup>9</sup> and 'smoker's paradox,'<sup>10</sup> respectively. The possible explanations for obesity paradox include the following: patients with obesity are younger than patients with normal weight and more likely to receive aggressive treatment.<sup>11</sup> In addition, when patients develop AMI, their metabolic demands increase sharply, and body fat may serve as nutritional reserves.<sup>12</sup>

Regarding smoker's paradox, the observed association may be subject to selection bias. The distribution of the risk factors was significantly different between the smokers and non-smokers. It is likely that we did not adjust for some unmeasured variables which may lead to selection bias. Conversely, the CAMI registry did not collect data on patients who died before hospitalisation. Failing to account for pre-hospital mortalities may also lead to selection bias.<sup>13</sup> In addition to selection bias, smoker's paradox may be explained by the biological effect of smoking. Smoking could lead to a chronic ischaemic state (ischaemic preconditioning)<sup>14</sup>; therefore, smokers may have better tolerance for an acute ischaemic event, such as AMI.

## **Clinical implications**

First, the simplified score can help save time in risk estimation at the first medical contact, that is, time of the first contact of the patient with the physician before obtaining laboratory test results. This may be beneficial particularly for high-risk patients with NSTEMI. A large-scale meta-analysis included 5324 patients from eight trials and found that an early invasive strategy was associated with a lower mortality among high-risk patients, including those with elevated cardiac biomarkers at baseline, diabetes mellitus, a GRACE risk score more than 140 and aged 75 years.<sup>8</sup> Consistently, a recent randomised trial showed that coronary angiography within the initial 12 hours was related to a lower risk of ischaemic outcomes among high-risk patients.<sup>9</sup> Therefore, early risk assessment enables high-risk patients with NSTEMI to receive revascularisation as soon as possible and may help improve their outcomes. Second, the SCAMI-NSTEMI score may help in better identification of high-risk patients. The current guidelines recommend prompt revascularisation in very high-risk patients with characteristics including cardiogenic shock, severe left ventricular dysfunction and haemodynamic instability.<sup>16</sup> However, many other baseline characteristics affect the mortality risk, and a patient may still be at a high risk even without the above-mentioned clinical presentation. Our study first identified independent risk factors based on the variables that can be easily obtained in clinical practice and then integrated these risk factors to establish the risk score system. Therefore, our score may help in better identifying patients at a high risk for in-hospital mortality with the absence of severe clinical presentation.

In addition, the SCAMI-NSTEMI score excluded laboratory test variables; this may broaden the applicability of the risk score in the management of AMI, especially in developing countries. Compared with the GRACE risk score, the SCAMI-NSTEMI score does not include the troponin level. Approximately 18.6% of hospitals do not have the capability to examine the troponin level in China. Even among hospitals with troponin-level testing capabilities, only 55.9% use the assay for >80% of patients with AMI.<sup>17</sup> Similarly, many patients may not have available data on the creatinine level at initial presentation. In a large-scale AMI cohort in the ACTION registry, approximately 11.6% of patients did not have available data on the initial or peak creatinine level.<sup>18</sup> Our SCAMI-NSTEMI score did not include the laboratory test variable and therefore enabled the providers to risk-stratify more patients admitted owing to suspected or confirmed AMI.

#### Limitations

Our study has several limitations. First, external validation of the SCAMI-NSTEMI score in a larger independent cohort from China and other countries is required in future studies. Second, we only assessed the in-hospital outcome; future studies are needed to examine whether the SCAMI-NSTEMI score is suitable for long-term risk prediction. Third, the CAMI registry did not collect data on the specific type and amount of tobacco products the patients used. Finally, all participants were from China. Our score requires further validation in another ethnic group.

#### **CONCLUSIONS**

The SCAMI-NSTEMI score, which was easier to calculate than the original score, showed good diagnostic performance and may aid in rapid risk stratification in more patients admitted owing to NSTEMI.

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#### REFERENCES

- Reed GW, Rossi JE, Cannon CP. Acute myocardial infarction. Lancet 2017;389:197–210.
- O'Gara PT, Kushner FG, Ascheim DD, et al. 2013 ACCF/AHA guideline for the management of ST-elevation myocardial infarction: a report of the American College of cardiology Foundation/American heart association task force on practice guidelines. *Circulation* 2013;127:e362–425.
- Granger CB, Goldberg RJ, Dabbous O, et al. Predictors of hospital mortality in the global registry of acute coronary events. Arch Intern Med 2003;163:2345–53.
- McNamara RL, Kennedy KF, Cohen DJ, et al. predicting in-hospital mortality in patients with acute myocardial infarction. J Am Coll Cardiol 2016;68:626–35.

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- Huynh T, Kouz S, Yan AT, et al. Canada acute coronary syndrome risk score: a new risk score for early prognostication in acute coronary syndromes. Am Heart J 2013;166:58–63.
- Timóteo AT, Aguiar Rosa S, Afonso Nogueira M, et al. ProACS risk score: an early and simple score for risk stratification of patients with acute coronary syndromes. *Rev Port Cardiol* 2017;36:77–83.
- Fu R, Song C, Yang J, et al. CAMI-NSTEMI score-China acute myocardial infarction registry-derived novel tool to predict in-hospital death in non-ST segment elevation myocardial infarction patients. *Circ J* 2018;82:1884–91.
- Jobs A, Mehta SR, Montalescot G, et al. Optimal timing of an invasive strategy in patients with non-ST-elevation acute coronary syndrome: a meta-analysis of randomised trials. *Lancet* 2017;390:737–46.
- Deharo P, Ducrocq G, Bode C, et al. Timing of angiography and outcomes in high-risk patients with Non-ST-Segment-Elevation myocardial infarction managed Invasively: insights from the TAO trial (treatment of acute coronary syndrome with Otamixaban). *Circulation* 2017;136:1895–907.
- Xu H, Li W, Yang J, et al. The China acute myocardial infarction (CAMI) registry: a national long-term registry-research-education integrated platform for exploring acute myocardial infarction in China. Am Heart J 2016;175:193–201.
- Thygesen K, Alpert JS, Jaffe AS, et al. Third universal definition of myocardial infarction. *Circulation* 2012;126:2020–35.
- 12. Steyerberg EW, Vickers AJ, Cook NR, *et al.* Assessing the performance of prediction models: a framework for traditional and novel measures. *Epidemiology* 2010;21:128–38.

- National Clinical Guideline C, National Institute for Health and Clinical Excellence: Guidance. Unstable angina and NSTEMI: the early management of unstable angina and non-ST-segment-elevation myocardial infarction. London: Royal College of Physicians (UK) National Clinical Guidelines Centre., 2010.
- 14. D'Ascenzo F, Biondi-Zoccai G, Moretti C, et al. Timi, grace and alternative risk scores in acute coronary syndromes: a metaanalysis of 40 derivation studies on 216,552 patients and of 42 validation studies on 31,625 patients. *Contemp Clin Trials* 2012;33:507–14.
- Wu C, Gao XJ, Zhao YY, et al. [Prognostic value of TIMI and GRACE risk scores for in-hospital mortality in Chinese patients with non-STsegment elevation myocardial infarction]. Zhonghua Xin Xue Guan Bing Za Zhi 2019;47:297–304.
- Roffi M, Patrono C, Collet J-P, et al. 2015 ESC guidelines for the management of acute coronary syndromes in patients presenting without persistent ST-segment elevation: Task force for the management of acute coronary syndromes in patients presenting without persistent ST-segment elevation of the European Society of cardiology (ESC). *Eur Heart J* 2016;37:267–315.
   Zhan L, Masoudi FA, Li X, et al. Trends in cardiac biomarker
- Zhan L, Masoudi FA, Li X, *et al.* Trends in cardiac biomarker testing in China for patients with acute myocardial infarction, 2001 to 2011: China PEACE-retrospective AMI study. *PLoS One* 2015;10:e0122237.
- Fox CS, Muntner P, Chen AY, *et al.* Short-term outcomes of acute myocardial infarction in patients with acute kidney injury: a report from the National cardiovascular data registry. *Circulation* 2012;125:497–504.