

## Patient Care of STEMI In Algeria: The STAMI Study

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### Abstract

**Background:** ST-segment elevation myocardial infarction (STEMI) is associated with a significant short- and medium-term mortality rate despite progress at several levels of intervention worldwide. Few data are available in Algeria.

**Objectives:** To evaluate major cardiovascular events in STEMI patients in Algeria, during hospitalization, 1 month and 12 months after discharge.

**Materials and Methods:** STAMI (STEMI: Algerian national registry) is a descriptive, non-interventional, national, multicenter, prospective registry-based study that was conducted in Algerian university public hospitals and included adults hospitalized within 24 hours of STEMI symptoms' onset. Major in-hospital cardiovascular events and after hospital discharge were recorded at 1 and 12 months.

**Results:** Between November 2016 and January 2017, 329 patients were enrolled; 322(97.9%) patients constituted the full-analysis set. Median age was 61[53; 68] years, 79.5% were males, their median body mass index was 26.0[23.4; 28.4] kg/m<sup>2</sup>. Most of them presented with a Killip class I. The first 2 main risk factors were smoking (46.6%) and high blood pressure (40.1%). Median time between symptoms' onset and arrival to the emergency room was 240 minutes and median time between symptoms' onset and primary angioplasty was 330 minutes. Fibrinolysis was administered to 194 (60.2%) patients and primary angioplasty to 94 (29.2%) patients. During hospitalization, 7 (2.2%) patients had a bleeding, 3(0.9%) patients had a recurrent MI, 3 (0.9%) patients a stroke and overall, 12 (3.7%) patients experienced at least 1 major cardiovascular event. In-hospital death rate was 2.2%. At 1 month and 12 months, major cardiovascular events rates were: death 2.3% and 6.2%; MI 0.3% and 0.4%, and stroke 1.0% and 0.4%, respectively.

**Conclusion:** Coronary emergencies remain a major public health issue in Algeria; more effective strategies aimed at the prevention of cardiovascular disease need to be implemented in Algeria to reduce the delays in access to hospital and minimize morbidity and mortality.

**Keywords:** ST-segment elevation myocardial infarction (STEMI); Algeria; Registry; Outcomes; Mortality

### Introduction

In 2017, the Global Burden of Diseases, Injuries, and Risk Factors Study, estimated that the cardiovascular diseases (CVD) were responsible for the largest number of deaths within non-communicable diseases (17.8 million; 95% uncertainty interval [UI]: 17.5–18.0 deaths). On a global level, between 2007 and 2017, deaths from ischemic heart disease increased from 7.30 million (95% UI: 7.22–7.46) to 8.93 million (95% UI: 8.79–9.14) [1]. According to the World Health Organization (WHO), at least three-quarters of the world's deaths from CVDs occur in low- and middle-income countries. People living in low- and middle-income countries often do not have the benefit of primary health care programs for early detection and treatment of people with risk factors for CVDs [2]. According to the Algerian Society of

Cardiology, CVD, including myocardial infarction (MI), kill almost twice as many people as all cancers combined. In 2019, ischemic heart disease was regarded as responsible for 127 deaths per 100,000 population in Algeria [3].

Acute ST-segment elevation myocardial infarction (ST-elevation myocardial infarction (STEMI) is defined as a separate entity among acute coronary syndromes due to its to its own pathophysiological, prognostic and therapeutic characteristics [4]. Overall, 30% to 50% of patients die within one month after MI, half of them by cardiac death suffered during the first two hours after the onset of symptoms. In contrast to the pre-hospital prognosis, early mortality in the hospital phase over the last 40 years has been reduced to 6-8.4% thanks to the creation of specialized units for revascularization treatments and

secondary prophylactic drug treatments [5]. Factors of poor prognosis are age, high heart rate, diabetes, history of MI, coronary artery bypass grafting, Killip class at admission and previous location of necrosis [6]. The objectives of the STAMI study were to evaluate major cardiovascular events in STEMI patients in Algeria. Secondary objectives included evaluation of time to patient's care, hospitalization and mortality at 1 and 12 months.

## Materials and Methods

### Study Design

STAMI (ST elevation acute myocardial infarction: Algerian national registry) is a descriptive, non-interventional, national, multicenter, prospective and longitudinal registry-based study conducted among cardiologists in university hospitals, from the public sector in Algeria.

### Patient's Selection

Females and males  $\geq 18$  years diagnosed with ST-segment elevation myocardial infarction (STEMI), defined by a history of chest pain/discomfort and persistent ST-segment elevation ( $> 30$  min) of  $\geq 0.1$  mV in 2 or more contiguous leads on electrocardiogram (ECG) or presumed new left bundle branch block (LBBB) upon admission, were eligible to participate if they signed the informed consent and completed the Contact Order Form. They also had to be hospitalized within 24 hours of current episode symptoms onset. Patients were not eligible to participate if STEMI was precipitated by or as a complication of surgery, trauma or gastrointestinal bleeding or post-percutaneous coronary intervention (PCI) or occurred in patients already hospitalized for other reasons. Additional exclusion criteria were any condition which could significantly limit the complete follow-up of the patient (e.g., tourist, does not understand the local language), and presence of serious/severe comorbidities which may limit short-term (i.e., 12 months) life expectancy.

### Data Collection

During the hospitalization period, socio-demographics, medical history, patient care, evolution and hospital discharge, and treatment patterns of eligible patients were collected in an electronic case report form (eCRF). During the follow-

up, data on medical procedures/hospitalizations, and living status were recorded over the phone at one and 12 months. Site's characteristics were also recorded. Patients underwent clinical assessments and received the standard medical care as determined by the treating physician. Patients did not receive any experimental intervention or treatment as a consequence of their participation in STAMI.

### Ethical Considerations

Collection of data started only after the obtention of the patient's written informed consent and data were analyzed respecting their autonomy and anonymity. The study was conducted in accordance with the Declaration of Helsinki, the Good Clinical Practice guidelines, and the applicable local legislations on non-interventional studies.

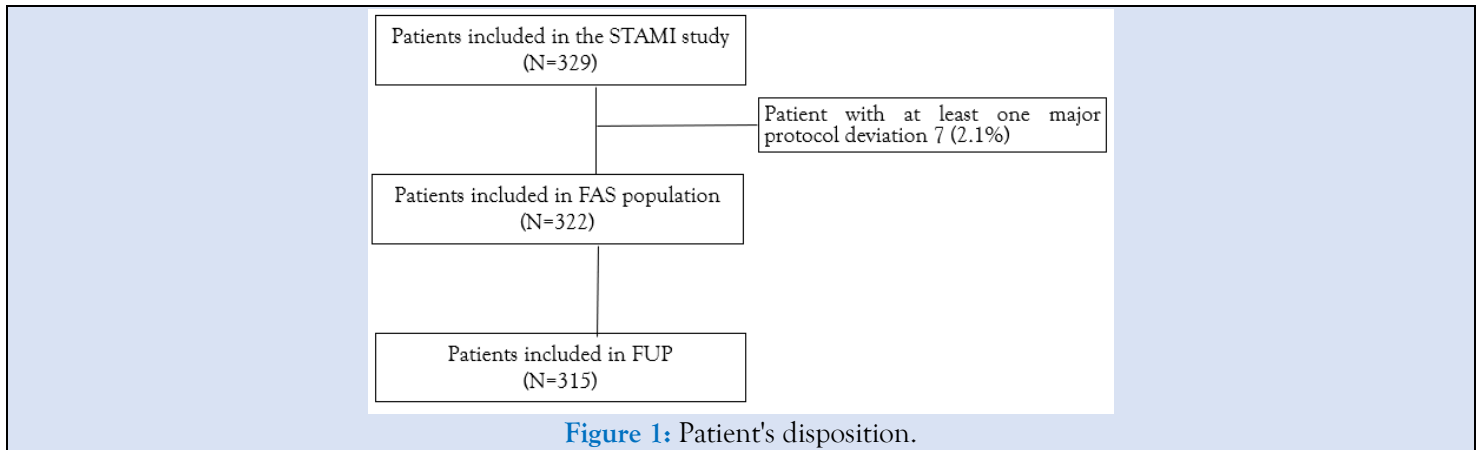
### Statistical Analysis

The primary endpoint was the percentage of major cardiovascular events (mortality, recurrence of MI and stroke). A total of 300 patients were to be recruited from the participating sites in order to have an acceptable precision.

Mean  $\pm$  Standard Deviation (SD) or median and interquartile ranges [IQR] for non-normal distributions are displayed. Missing data were reported but not included in the analyses. The full-analysis set (FAS) was defined as all enrolled patients with no major protocol deviation and the follow-up population (FUP) as all FAS patients discharged from hospital. Data are summarized using standard descriptive statistics. All statistical tests are two-sided and are displayed with their bilateral 95% confidence interval (95%CI). Cumulative mortality rate after discharge was estimated using the Kaplan-Meier method with its 95% CI at 30 days and 12 months. Statistical analysis was done using SAS/STAT software, version 14.3 for Windows.

## Results

Overall, 329 patients were enrolled between November 2016 and January 2017, of whom 322 (97.9%) patients constituted the FAS. The 315 (95.8%) patients who were discharged from the hospital constituted the FUP (Figure 1); 310 (98.4%) patients had at least one follow-up phone call (Figure 1).



**Figure 1:** Patient's disposition.

FAS: Full Analysis Set (patients with no major protocol deviation); FUP: Follow-up population (patients included in FAS population, not dead during hospitalization, and discharged)

### Baseline Characteristics, Initial Presentation, And Early Management

Patient's characteristics are displayed in Table 1. Patients were mostly male with a median age of 61 [53; 68] years old and a median body mass index (BMI) of 26.0 [23.4; 28.4] kg/m<sup>2</sup>. Most patients presented with a Killip class I.

**Table 1:** Patient's characteristics upon admission (FAS population).

	Overall n=322
Males	256 (79.5%)
Age (years)	61 [53; 68]
BMI (kg/m <sup>2</sup> )	26 [23.4; 28.4]
Heart rate (beat/min)	80 [70; 97]
SBP (mmHg)	130 [120; 150]
DBP (mmHg)	80 [70; 90]
Killip class	
I	299 (92.9%)
II	13 (4.0%)
III	9 (2.8%)
IV	1 (0.3%)

n (%) patients or median [Q1; Q3]. BMI: body mass index; DBP: diastolic blood pressure; SBP: systolic blood pressure.

Patient's risk factors are presented in Table 2.

**Table 2:** Patient's risk factors (FAS population).

	Overall n=322
Smoking	150 (46.6%)
Hypertension	129 (40.1%)
Dyslipidemia	38 (11.8%)
Family history of coronary heart disease	33 (10.2%)
Diabetes	
Type 1	1 (0.3%)
Type 2	108 (33.5%)
Unstable angina	25 (7.8%)
Renal failure	19 (5.9%)

n (%) patients

Initial patient management is presented Table 3. Median time between symptoms' onset and arrival to the emergency room was 240 [120; 420] minutes; 191 (59.3%) patients had previously been in another medical facility. At arrival, an ECG was performed for

all patients. ST+ elevation show that ischemia affected the anterior area for 181 (56.2%) patients, the inferior area for 130 (40.4%) patients, the basal area for 49 (15.2%) and the lateral area for 52 (16.1%) patients. Arrhythmia was present in 27 (8.4%) patients

including: ventricular tachycardia in 7 (25.9%) patients, ventricular fibrillation in 6 (22.2%) patients and atrial fibrillation in 9 (33.3%) patients. Overall, 22 (6.8%) patients presented conduction defects with

mostly 3rd degree atrioventricular block [12 (54.5%) patients] or right bundle branch block [7 (31.8%) patients].

**Table 3:** Initial presentation and management (FAS population).

Overall n=322	
Time delays (minutes)	
Time between symptoms' onset and arrival to the emergency room	240 [120; 420]
Time between symptoms' onset and primary angioplasty	330 [189; 600]
Procedures upon arrival	
ECG (at inclusion)	322 (100%)
ECG (90 min)	306 (95.0%)
Primary angioplasty	94 (29.2%)
Fibrinolysis:	194 (60.2%)
Tenecteplase	154 (81.9%)
Alteplase	34 (18.1%)
Medications	
Unfractionated Heparin	41 (13.6%)
LMWH	261 (86.4%)
Aspirin	321 (99.7%)
Beta-blockers	202 (62.7%)
Clopidogrel	308 (97.2%)
Ticagrelor	9 (2.8%)
Diuretic	21 (6.5%)
Statin	200 (62.1%)
Nitrates	30 (9.3%)
Morphine	48 (14.9%)
ACE inhibitor	200 (62.1%)
Others	60 (18.6%)

median [Q1; Q3] or n (%) patients. ACE: angiotensin-converting enzyme; ECG: electrocardiogram; LMWH: Low Molecular Weight Heparin.

### In-Hospital Complications

The median duration of hospitalization was 4 [3; 6] days. During hospitalization, a control angiography was performed in 129 (40.1%) patients. Recurrent MI occurred in 3 (0.9%) patients (95%CI: [0.0%; 2.0%]) and 17 (5.3%) experienced recurrent chest pain. Emergency revascularization by angioplasty was performed in 65 (20.2%) patients. A stroke occurred in 3 (0.9%) patients [0.0%; 2.0%]. Overall, 12 (3.7%)

patients experienced at least 1 major cardiovascular event during hospitalization (3.7% [1.7%; 5.8]). In-hospital death rate was 2.2% [0.6%; 3.8%]; in the 7 patients who died (1 stroke and 7 cardiac deaths), median time to death was 2 [1; 5] days.

### Discharge

Overall, 315 patients were discharged. Treatments prescribed at discharge are displayed in Table 4.

**Table 4:** Medications at discharge.

Overall n=315	
Aspirin	311 (98.7%)
Clopidogrel	296 (94.9%)
Ticagrelor	16 (5.1%)
Oral Anticoagulant	21 (6.7%)
Statin	308 (97.8%)
Beta-blockers	295 (93.7%)
ACE inhibitor	291 (92.4%)
Oral Antidiabetics	46 (14.6%)
Insulin	41 (13.0%)

n (%) patients. ACE: angiotensin-converting enzyme.

## Follow-up

The follow-up by phone was effective in 307 (97.5%) and 271 (86.0%) patients, at 1 and 12 months

respectively. Cardiovascular events reported during the follow-up are presented Table 5. Mortality rate was 2.3% [0.6%; 3.9%] at 30 days and 6.2% [3.4%; 9.0%] at 12 months.

**Table 5:** Follow-up.

	1 month n=307	Between 1- and 12-months n=271
Myocardial infarction	1 (0.3%)	1 (0.4%)
New hospitalization for a cardiac event	16 (5.2%)	29 (10.7%)
Stroke	3 (1.0%)	1 (0.4%)
Hemorrhage	3 (1.0%)	0

*n* (% column) patients.

## Discussion

The STAMI study was the first national registry describing patient management of acute MI and risk assessment after MI in Algeria. Patient's characteristics were similar to the STEMI patients from the French FAST-MI 2015 study in term of age (61 [53; 68] years vs 63 ± 14 years), sex (79 % vs 75 % males) and risk factors (smoking: 47% vs 42%; hypertension: 40% vs 45%) [7]. Previous registry studies conducted in the Middle East and North Africa (MENA) region reported a higher prevalence of cardiovascular risk factors especially tobacco which was recorded in 58% to 65 % in STEMI patients [8-10]. Indeed, a recent estimation by the WHO, indicated that prevalence of smoking in Algeria is decreasing [11], most likely as a result from the national preventive measures. One main difference between the STAMI patients and the FAST-MI 2015 patients was the percentage of Killip class I: in STAMI, most patients (93%) present a Killip class I when the FAST-MI 2015 proportion was slightly lower (88%). This suggests that the most serious cases of STEMI died before arrival at the hospital or were not able to give their informed consent.

The 240-minute median time between symptoms' onset and arrival to the emergency room was high whereas in another study performed in the MENA median was 178 minutes [12]. The median time between symptoms' onset and primary angioplasty was 330 minutes when the ESC guidelines recommend to initiate primary angioplasty within 120 minutes of STEMI diagnosis [13]. These differences may be due to variations in clinical management, but also to socioeconomic factors such as the type of hospital, and delays to presentation and treatment [10]. For hospitals with no available percutaneous coronary intervention, transfers remain the optimal strategy. For expected delays of greater than 120

minutes, a pharmaco-invasive strategy is recommended [14]. To minimize patient delay, guidelines recommend increasing public awareness of how to recognize common symptoms [13]. Indeed, in the STAMI study, more than half patients consulted in another facility before arriving to the emergency room.

The main objective of the STAMI study was to estimate the incidence of major cardiovascular events. In-hospital bleeding and stroke were higher than the respectively reported <1% and 0.5% in the FAST-MI 2015 study. MI recurrence was comparable in both studies. However, in-hospital death was slightly lower than the 2.8% in FAST-MI 2015 which could be linked to the lower proportion of Killip class >1. In-hospital mortality of unselected patients with STEMI in the national registries of the European Society of Cardiology countries, varies between 4 and 12% [13]. In a similar STAMI setting, a Tunisian study reported a higher in-hospital mortality in STEMI patients of 7.0 % [9].

The rates of major cardiovascular events at 1 month were lower in the STAMI study than in a large Indian study: death (2.3% vs 8.6%), reinfarction (0.3% vs 2.3%) and stroke (1.0% vs 0.7%) [15].

In the present study the acute reperfusion rate was high (fibrinolysis 60.2% and primary angioplasty 29.2%) compared to other developing countries. A prospective analysis of registry data of acute coronary syndromes in India, the CREATE registry, reported a high rate of reperfusion with fibrinolytic therapy for patients with STEMI (59%) but a very low rate of in-hospital percutaneous coronary intervention (PCI) (8.0%) [15]. In comparison with a similar study performed in Maghreb, only 30% of STEMI patients received a fibrinolytic therapy, and PCI was performed in 27% of STEMI patients, of whom 97% received a stent [10]. Additionally, the FAST-MI Tunisia registry (2014-2015) recorded a total rate of

reperfusion of 61.8% of STEMI patients: 30% with primary PCI and 31.8% with intravenous fibrinolysis, total reperfusion rate is still lower than the one reported in the present study [16].

More effective strategies aimed at the prevention of cardiovascular disease need to be implemented in Algeria to reduce the delays in access to hospital and minimize morbidity and mortality. Also, a longer door-to-balloon delay in primary PCI for STEMI is related to higher risk of adverse outcomes [17].

This observational study is subject to limitations and biases, as all observational studies, such as potential selection bias and missing or incomplete information. As with most registries, hospital and patient enrolments were voluntary; thus, the study results may not be representative of patients in all hospitals in the country.

## Conclusion

Coronary emergencies remain a major public health issue in Algeria during which saving time is a real challenge. The more systematic use of primary angioplasty and adjuvant therapies to angioplasty has allowed an improvement of the prognosis of infarction. Early diagnosis in the hospital setting makes it possible to offer appropriate therapeutic management.

## Declarations

### Patient Consent and Ethical Approval

This study was approved by the Algerian Ministry of Health and a central ethical committee on October 06, 2016. All patients provided written informed consent before any collection of study data, and data were analyzed respecting their autonomy and anonymity. The study was conducted in accordance with the Declaration of Helsinki, the Good Clinical Practice guidelines and all the applicable local legislations on non-interventional studies.

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### Competing Interests

All authors have no conflict of interest to declare.

### Author Contribution

All authors read, revised, and approved the final version of the article.

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