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## **Research Article**

# Sustained Ventricular Tachycardia in Acute Coronary Syndrome with ST Segment Elevation: Incidence, Predictive Factors and Related Mortality.

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# **Abstract:**

**Background:** Sustained Ventricular Tachycardia is the most serious complication; in acute coronary syndrome with ST segment elevation, several international studies have reported incidence and predictive factors of severe ventricular arrhythmias, without identifying sustained ventricular tachycardia separately; also, its epidemiological data is lacking in Algeria.

**Aims:** The main objective of our study is the determination of the frequency of sustained ventricular tachycardia in acute coronary syndrome with ST segment elevation, the secondary objective was the analysis of predictive factors of this arrhythmia, and related mortality.

**Methods and materials:** In this prospective study, conducted in the cardiology department of Hussein Dey hospital (Algiers-Algeria), 467 patients with acute coronary syndrome with elevated ST segment (87 women and 380 men) were enrolled between 28 February 2014 and 16 July 2015. The average age is  $60 \pm 13$  years; at admission, a Holter recorder was attached for continuous ECG monitoring during 48 hours

Kruskal's ANNOVA or H tests were used for comparison of quantitative variables,  $\chi 2$  test or Fisher's exact test, were used for qualitative variables, all tests were performed with 1<sup>st</sup> species risk of 5%.

**Results:** The frequency of Sustained Ventricular Tachycardia is 3.6 % (17 patients), CI 95%: [1.9%-5.3%], multivariate analysis identified the following independent predictors: low diastolic blood pressure, severe coronary artery lesions, Elevated high sensitive troponin concentrations more than 5ng/ml.

The risk of mortality expressed by Hazard Ration (HR) is 6. (CI95%: [1.7-21], p = 0.005); the predictors of mortality are: occurrence of ventricular fibrillation and low diastolic blood pressure.

Conclusion: Sustained Ventricular tachycardia is the most serious complication during acute coronary syndrome with elevated ST segment, its predictive factors according to our study are: low diastolic blood pressure, severe coronary lesions, Elevated high sensitive troponin concentrations more than 5ng/ml, its occurrence increases the risk of hospital mortality.

# Keywords: Acute Coronary Syndrome, Sustained Ventricular Tachycardia, Severe Coronary Artery Lesions.

## **Introduction:**

Sustained ventricular tachycardia (SVT) defined as ventricular tachycardia lasting at least 30 seconds, is one of the most serious ventricular arrhythmias, rarely observed early in acute coronary syndrome with ST segment elevation, because of its frequent degeneration into ventricular fibrillation.

Ventricular tachycardia may induce hemodynamic instability, and promotes the onset of cardiogenic shock during hospitalization.

Its mechanisms are complex and multifactorial; several electrophysiological modifications occur just after coronary artery occlusion, and lead to reentry phenomena, abnormal automatism, and triggered activity.

Several international studies have reported incidence and predictive factors of severe ventricular arrhythmias, without identifying sustained ventricular tachycardia separately; also its epidemiological data is lacking in Algeria.

The main objective of our study is to determine the frequency of sustained ventricular tachycardia in acute coronary syndrome with ST segment elevation, during the first 48 hours of hospitalization, while the secondary objective is the analysis of its predictive factors and the related mortality.

# Methods and materials:

We prospectively studied a group of 467 consecutive patients (380 men and 87 women; mean age  $60 \pm 13$  years) who presented acute coronary syndrome with ST segment elevation and admitted in cardiology department of Hussein-Dey hospital (Algiers, Algeria), between 28th February 2014 and 16th August 2015.

At emergency department admission, an ECG Holter recorder was attached for continuous ECG monitoring during 48 hours, the 17-leads surface ECG recorded at admission and repeated during hospitalization, Doppler Echocardiography, coronary

angiography, and biological assessment were performed in the majority of patients.

The most important rhythm and conduction disorders were identified, the patients with the same type of disorder are grouped together, and the name assigned to each group is that of the disorder that characterizes it; there are overlaps between the groups, so that several disorders may exist in the same patient.

The constitution of each group of the rhythm disorder implies the constitution of the opposite group without the corresponding disorder, the latter group is used for the comparative study; each group is therefore described and then compared to the corresponding opposite group.

In this sub study, the group of patients with sustained ventricular tachycardia was compared to the rest of patients without sustained ventricular tachycardia.

# **Statistical analysis:**

Data are presented as mean  $\pm$  SD, median, or frequency (percentage) where appropriate. Continuous variables were compared using the ANNOVA test, or H Kruskal Wallis test.  $\chi 2$  tests and Fisher's exact test were performed to distinguish differences between categorical variables. Statistical significance was defined as p < 0.05. In this first step, we used EPI-info version 6.0. A multivariate Binary regression was performed to determine the predictor factors of arrhythmias, and Cox regression was performed to identify the predictor factors of mortality.

The magnitude of the relationship between sustained ventricular tachycardia and their predictive factors is estimated by the Cramer V coefficient, a coefficient lower than 0.2 is in favor of a weak link, between 0.2 and 0.5: moderate link, greater than 0.5: strong link.

The statistical analysis was performed using SPSS Statistics (release 17).

## Results

#### **Incidence:**

The characteristics of the 467 patients included in our study are shown in Table 1; seventeen patients had presented sustained ventricular tachycardia at admission or during hospitalization, so its frequency in this present study is 3.6 % (17 patients), CI 95% [1.9%-5.3%].

This group of patients included 2 women and 15 men. The mean age was  $59.70 \pm 12$  years; the extreme age was 37 and 74 years. Six patients had presented SVT at admission, and eleven patients had SVT during their hospitalization.

The rate of SVT varied between 110 and 300 beats/min, the mean rate was 169 beats/min;

Monomorphic ventricular tachycardia with aspect of left bundle branch block was present in 15 patients, and with aspect of right bundle branch block in 2 patients.

Cardiovascular risk factors, clinical characteristics, medical history, treatment and evolution are shown in Table 1.

The Surface ECG had shown, anterior ACS in three patients, circumferential in two patients, inferior in five patients, right ventricular in one patient.

The mean heart rate at admission was  $91.52 \pm 30.59$  beats/min, the mean PR interval was  $143.33 \pm 29.19$  msec, the mean duration of the QRS complex was  $80 \pm 21.21$  msec, the mean amplitude of the ST segment elevation was  $4.41 \pm 2.62$  mm, the mean amplitude of the ST segment depression was  $1.58 \pm 3.06$  mm, the mean amplitude of the T wave was  $7.76 \pm 3.27$  mm and the mean corrected QT was  $439.86 \pm 46.28$  msec.

Five patients had persistence of the segment ST elevation during hospitalization.

	Patients with SVT (n = 17)	Patients without SVT (n = 450)	P-value	
Mean age	59.706	60.198	0.874	NS
Females	2	85	0.672	NS
Early consultation (within 6 hours)	12	331	0.455	NS
Hypertension	5/17	204/450	0.294	NS
Diabetes	7/27	149/450	0.667	NS
Current smoking	10/17	225/450	0.640	NS
hyperlipidemia	3/17	64/447	0.455	NS
GRACE score ≥ 155	13/17	154/449	0.001	S
Cardiogenic shock	5/17	13/450	0.000	S
Left ventricular heart failure	6/17	57/450	0.017	S
Right ventricular heart failure	1/17	10/450	0.337	NS
Persistence of chest pain	4/17	24/450	0.014	S
Mean SBP	91.235	131.867	0.000	S
SBP ≤100 mmHg	8/17	54/450	0.000	S
Mean DBP	56.000	78.311	0.000	S
DBP ≤ 60 mmHg	10/17	72/450	0.000	S

Hospital mortality (first 48 hours)	3/17	14/450	0.019	S
Previous myocardial infraction	0/17	21/450	0.451	NS
Electrocardiogram			l l	1
Right ventricular ACS	1/17	41/450	0.537	NS
Circumferential ACS	2/17	22/450	0.215	NS
Anterior ACS	3/17	41/450	0.209	NS
Inferior ACS	3/17	79/450	0.173	NS
QTc ≥ 440 msec	10/17	121/450	0.006	S
Persistence of ST segment elevation	5/17	25/448	0.002	S
Other associated arrhythmias				
Polymorphic PVC	5/14	46/434	0.014	S
PVC with R on T phenomenon	3/14	15/434	0.015	S
Medication before ACS	l		l .	I.
Beta blockers	2/17	34/449	0.383	NS
ARB	2/17	65/449	0.547	NS
ACE-inhibitor	3/17	32/449	0.3305	NS
Lipid-lowering drugs	2/17	31/449	0.2970	NS
Antiplatelet agents	4/17	36/449	0.048	S
Treatment at admission	l		<b> </b>	
Thrombolysis	15/17	391/450	0.612	NS
Primary or rescue percutaneous coronary intervention	0/11	14/318	0.615	NS
Beta blockers	6/17	180/450	0.891	NS
ACE-inhibitor	7/17	254/450	0.319	NS
Echocardiography				
Ejection fraction of left ventricle < 40	5/14	58/433	0.034	S
Mean Left ventricular diastolic diameter	59.643	53.954	0.032	S
Left ventricular diastolic diameter ≥ 59 mm	8/14	92/433	0.004	S
Coronary angiography		·		
Severe coronary artery lesions	8/11	94/318	0.004	S
Left main coronary artery severe lesion	0/11	12/318	0.660	NS
Left anterior descending artery lesion	9/11	200/318	0.168	NS
Left circumflex coronary artery lesion	4/11	114/318	0.603	NS
Right coronary artery lesion	9/11	130/380	0.008	S
Two-vessel coronary artery disease	8/11	120/318	0.022	S
Multi-vessel coronary artery disease	4/11	57/318	0.126	NS
TIMI flow grade 0	2/11	63/318	0.625	NS

ACS: Acute Coronary Syndrome, ACE inhibitors: Angiotensin-converting enzyme inhibitors ARB: Angiotensin receptor-blocker, DBP Diastolic blood pressure, LBBB: left bundle brunch block, PVC: Premature ventricular complexes, QTc: Corrected QT interval, RBBB: Right bundle brunch block, SBP: Systolic blood pressure.

**Treatment at admission and during hospitalization**: Metalyse (Tenecteplase) as fibrinolytics treatment were administered in 15 patients (88.23 %), 6 patients had presented SVT at admission before any therapy, 9 patients had presented SVT after thrombolysis. (Figure 1)

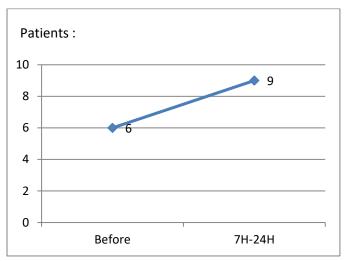


Figure 1: Sustained Ventricular Tachycardia onset delay as a Function of fibrinolytics treatment

Aspirin, Clopidogrel and Anticoagulants were administered in 17 patients, beta blockers in 6 patients (35.29 %), ACE inhibitors in 7 patients (41.17 %), sympathomimetic agents in 4 patients (23.52 %), diuretics in 2 patients (11.76 %), external electric shock in 13 patients (76.47 %), Amiodarone in 8 patients (47.05 %), Magnesium and Potassium supplementation at admission in 12 patients 70.58 %), Atropine in 1 patient (5.88 %), Insulin in 7 patients (41.17 %).

**Thrombolysis failure:** the persistence of chest pain after thrombolysis was observed in 3 patients, the persistence of ST segment elevation in 4 patients; for patients who had experienced SVT during hospitalization, the persistence of chest pain and ST segment elevation was observed in 3 patients.

**Doppler echocardiography** was performed in 14 patients, the left ventricular fraction less than 40 % was found in 5 patients (35.71%), left ventricular hypertrophy in 2 patients (14.28%), the mean area of the left atrium:  $18.64 \pm 5.31$  cm², that of the right atrium:  $12.84 \pm 5.16$  cm², the mean diastolic diameter of the left ventricle:  $59.64 \pm 10.28$  mm, diastolic diameter of left ventricle above or equal 59 mm in 8 patients (57.14 %), the mean diastolic diameter of the right ventricle was  $26.27 \pm 3.19$  mm, systolic pulmonary blood pressure:  $28.66 \pm 7.77$  mm Hg, wall akinesia in 10 patients (71.42%), apical thrombus in 1 patient (7.14%), and significant mitral insufficiency in 2 patients (14.28%).

**Holter ECG** was performed in 14 patients; this exam had participated in the recording of SVT, and also showed its duration, and frequency. (Figure 2)

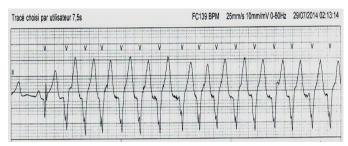


Figure 2: Initiation of Sustained Ventricular Tachycardia

Several arrhythmias were associated with SVT, ventricular fibrillation had occurred in 10 patients (58.82%), non-sustained ventricular tachycardia (NSVT) in 7 patients (41.17%), atrial fibrillation in 2 patients 11 %), complete left bundle branch block in 1 patients (5.8%) and complete right bundle branch block in 3 patients (17.64 %), bursts of atrial premature beats were found in 7 patients (4117 %), bursts of ventricular premature beats in 6 patients (35.29%), accelerated idioventricular rhythm in 2 patients (11.76 %), polymorphic ventricular premature beats in 5 patients (29.41 %), PVC with R on T phenomenon in 3 patients (17.64%).

**Evolution and complications:** stroke in 1 patient and aneurysm in 1 patient.

Coronary angiography was performed in 11 patients, severe coronary artery lesions were found in 8 patients (72.72%), severe stenosis of the left anterior descending artery in 9 patients (81.81%), circumflex artery in 4 patients (36.36%), right coronary artery in 9 patients (81.81%), two-vessel coronary artery lesions in 8 patients (72.72%), three vessel coronary artery lesions in 4 patients (36.36%), TIMI flow grade 0 in 2 patients (18.18%).

**Mortality:** three patients died during the first 48 hours of their hospitalization; 2 patients died during the first 2 hours and one patient after 12 hours.

**Biology:** the average blood glucose:  $2.31\pm1.27$  g/l, blood glucose level above or equal 2g/l in 9 patients (52.94%), average serum potassium  $3.98\pm0.64$  mmol/l, average blood urea:  $0.42\pm0.24$  g/l, blood creatinine:  $14.56\pm10.43$  mg/l; High-Sensitivity Troponin (hs-Trop) above or equal 5 ng/ml in 8 patients (66.66%).

**Predictive factors :** According to the univariate study, several variables had a statistically significant association with the occurrence of SVT: Cardiogenic shock, left heart failure, persistence of chest pain, GRACE Score  $\geq 155$ , left ventricular ejection fraction < 40 %, Mean diastolic diameter of the left ventricle  $\geq 59$  mm, mean corrected QT interval  $\geq 440$  mse Persistence of ST segment elevation after thrombolysis, Mean systolic blood pressure  $\leq 100$  mm Hg, Mean diastolic blood pressure  $\leq 60$  mm Hg, severe coronary artery lesions, right coronary artery stenosis two-vessel coronary artery lesions, blood glucose level above or equal 2g/l, High-Sensitivity Troponin (hs-Trop) above or equal 5 ng/ml etc. (Table 2)

But after the multivariate analysis using binary logistic regression, three predictive factors were identified: diastolic blood pressure  $\leq 60$  mm Hg, severe coronary artery lesions, High-Sensitivity Troponin (hs-Trop)  $\geq 5$  ng/ml. (Table 3) (Figure 3)

Variables	RR	CI 95%	P
Cardiogenic shock	10.98	4.36-27.67	0.0001
Left heart failure	3.48	1.33-9.08	0.01
Mean systolic blood pressure ≤ 100 mmHg	5.81	2.33-14.48	0.0005
Mean diastolic blood pressure ≤ 60 mmHg	6.71	2.63-17.10	0.0001
Persistence of Chet pain	4.82	1.68-13.03	0.01
GRACE Score ≥ 155	5.89	1.95-17.77	0.0008
Mean systolic blood pressure ≤ 100 mm Hg	3.84	1.84-8.01	0.001
Mean diastolic blood pressure ≤ 60 mm Hg	3.23	1.56-6.70	0.002
Polymorphic premature ventricular beat	4.32	1.51-12.40	0.01
PVC with R on T phenomenon	6.52	1.99-21.34	0.01
Persistence of ST segment elevation	6.04	2.28-16.03	0.002
Mean left ventricular ejection fraction < 40 %	3.39	1.17-9.78	0.03
Mean diastolic diameter of the left ventricle ≥ 59 mm	4.63	1.64-13.02	0.004
Severe coronary artery lesions	5.93	1.61-21.91	0.004
Right coronary artery stenosis	6.12	1.34-27.88	0.008
Two-vessel coronary artery lesions	4.19	1.13-15.49	0.02
High-Sensitivity Troponin (hs-Trop) ≥ 5 ng/ml	3.26	1-10.64	0.04
blood glucose level ≥ 2g/l	2.98	1.17-7.55	0.02
Antiplatelet agents (ongoing medication)	3.28	1.12-9.58	0.04
Persistence of ST segment elevation after thrombolysis	5.08	1.73-14.89	0.01
Persistence of chest pain after thrombolysis in patients who had experienced SVT during hospitalization	6.26	1.72-22.82	0.02
Persistence of ST segment elevation after thrombolysis in patients who had experienced VF during hospitalization	5.98	1.64-21.85	0.023

Table 3: Predictive factors of sustained ventricular tachycardia			
Predictive factors	OR	CI 95%	P
Diastolic blood pressure ≤ 60 mm Hg	8.923	2.0-39.10	0.004
Severe coronary artery lesions	8.544	1.951-37.421	0.004
High-Sensitivity Troponin (hs-Trop) ≥ 5 ng/ml	5.831	1.342-25.338	0.019

The magnitude of the relationship between SVT and low diastolic blood pressure is moderate, the Cramer V coefficient greater than 0.2, but low for the two others, the Cramer V coefficient does not exceed 0.2. (Table 4)

Table 4: Magnitude of the relationship between Sustained Ventricular Tachycardia and its predictive factors			
Predictive factors of sustained ventricular tachycardia	Cramer V coefficient	P	
Diastolic blood pressure ≤ 60 mm Hg	0.211	0.000	
Severe coronary artery lesions	0.101	0.038	
High-Sensitivity Troponin (hs-Trop) ≥ 5 ng/ml	0.168	0.002	

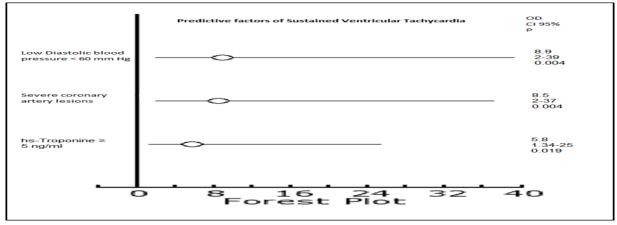


Figure 3: Predictive factors of Sustained Ventricular Tachycardia

# Mortality:

Hospital mortality (first 48 hours), in the SVT group is 17.64% while it does not exceed 3.11% in the group without SVT, p = 0.019. (HR at 6, CI 95% [1.7-21], p = 0.005. (Figure 4)

Cox regression was used for univariate and multivariate studies of mortality predictors.

According to the univariate study, some factors have a statistically significant association with the occurrence of mortality in the SVT group. (Table 5)

According to multivariate analysis, two predictive factors of mortality were identified: low diastolic blood pressure and occurrence of ventricular fibrillation. (Table 6)

The magnitude of the relationship between mortality and its predictive factors was moderate in the SVT group, but not significant. (Table 7)

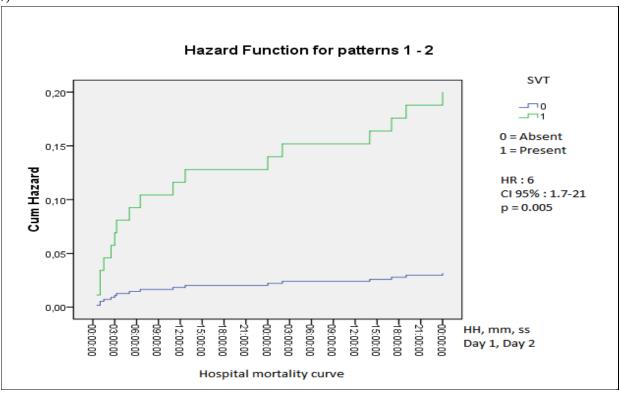


Figure 4: Hospital mortality curve (48h) in Sustained Ventricular Tachycardia (SVT) group versus group without SVT

Table 5: Univariate analysis, Factors related to mortality in group of Sustained Ventricular Tachycardia			
Factors	OR	IC95%	P
Feminine sex	4.5	1.7-12	0.002
Age ≥ 65 years	8.5	2.4-29.8	0.001
Right heart failure	65	23.4-181	0.000
Left heart failure	5.1	1.9-13.6	0.001
Cardiogenic shock	231	53-1003	0.000
Right ventricle acute coronary syndrome	6.1	2.2-16.7	0.000
Persistence of ST segment elevation	39.6	13.4-117.4	0.000
Persistence of chest pain	27.3	10.2-73	0.000
Diastolic blood pressure ≤ 60 mm Hg	3.28	1.1-9.1	0.023
Systolic blood pressure ≤ 100 mm Hg	8.8	3.2-24	0.000
Blood creatinine ≥ 17 mg/l	7.6	2.4-23.5	0.000
GRACE Score	6.9	1.9-24.9	0.003
Occurrence of ventricular fibrillation	12.4	4-37.8	0.000

Table 6: Predictive factors of mortality in Sustained Ventricular Tachycardia group				
Predictive factors of mortality in Sustained Ventricular Tachycardia group	OR	CI95%	P	
Occurrence of Ventricular Fibrillation	8.8	2.9-26.9	0.000	
Diastolic blood pressure ≤ 60 mm Hg	4.9	1.8-13.7	0.002	

The magnitude of the relationship between mortality and its predictive factors was moderate in the SVT group, but not significant. (Table 7)

Table 7: The magnitude of the relationship between mortality and its predictive factors			
Predictive factors of mortality in Sustained Ventricular Tachycardia group	Cramer V Coefficient	P	
Occurrence of Ventricular Fibrillation	0.387	0.11	
Diastolic blood pressure ≤ 60 mm Hg	0.240	0.32	

#### Discussion

Sustained ventricular tachycardia (SVT) is one of the most serious ventricular arrhythmias, rarely observed early in acute coronary syndrome with ST segment elevation, because of its frequent degeneration into ventricular fibrillation.

Sustained Ventricular Tachycardia may induce hemodynamic instability, and promotes the onset of cardiogenic shock during hospitalization.

Its mechanisms are complex and multifactorial; several electrophysiological modifications occur just after coronary artery occlusion, and lead to reentry phenomena, abnormal automatism, and triggered activity. [1]

In the literature the frequency of SVT is reported with that of Ventricular Fibrillation; in the GRACE register, the frequency of both arrhythmias is 10%, in the Thai register: 19.4%, in the Swedish register: 7%, in the APEX-AMI study: 5.7%, in the PAMI study: 4.3%, in the Swiss study: 8.7%, and in the Saudi Arabia register: 2.45%. [2-8]

The incidence of SVT in our study was 3.6 % (17 patients), CI 95% [1.9%-5.3%]. This incidence is within the range of that reported in the literature.

According to the Thai registry, predictive factors of ventricular arrhythmias are cardiogenic shock, tobacco, and elevated troponin levels [3]

For the PAMI study, the predictors of severe ventricular arrhythmias are tobacco, TIMI flow 0, the right territory of the ACS, early consultation time, and non-administration of beta blockers early on admission [6]

In another study published in 2012, the predictive factors of severe ventricular arrhythmias are the following: tobacco, taking beta blockers, digitalis and significant left main coronary artery disease [9]

In our study, predictive factors of sustained ventricular tachycardia were studied separately.

After multivariate analysis using binary logistic regression, the following predictive factors of Sustained ventricular tachycardia are: Low diastolic blood pressure, severe coronary artery lesions, High-Sensitivity Troponin (hs-Trop)  $\geq 5$  ng/ml. The low diastolic blood pressure reflects all other factors that reflect an altered hemodynamic status: cardiogenic shock, left or right ventricular failure, low systolic blood pressure, and high GRACE Score.

Severe coronary artery lesions reflect the presence of previous fibrosis and chronic ischemia which lead to constitution of arrhythmias substrate.

The high level of troponin released reflects the importance of tissue damage.

In the literature, hospital mortality related to SVT and ventricular fibrillation were regrouped together and reported as

hospital mortality related to severe ventricular arrhythmias.

According to the GRACE registry, the hospital mortality rate in patients with ventricular arrhythmias is 52% while it does not exceed 1.6% in patients without rhythmic disorders. [10]

In the Thai Registry (TACSR), the hospital mortality rate in patients with ventricular arrhythmia is 48%. [3]

In the VALIANT register, sustained ventricular tachycardia and ventricular fibrillation represent one of major predictors of hospital mortality (Relative Risk: 4.18). [11]

According to the Swedish primary angioplasty register published in 2012, hospital mortality rate is 16.5% in patients with ventricular tachycardia or fibrillation, while it is 1.5% in patients without ventricular arrhythmias; in the same register, Hospital mortality rate is higher and reaches 17.6% if ventricular arrhythmia (VF/VT) occurs after angioplasty, while it is 11.3% if this arrhythmia occurs before angioplasty. [4]

In our study, hospital mortality (first 48 hours), in the SVT group is about 17.64 %, this rate is lower than that reported by the GRACE and Thai registry, but similar to that of the Swedish registry, so our results were closer to those reported in study that exclusively uses primary angioplasty, which proves that fibrinolytics are as effective as primary angioplasty in reducing mortality in patients with severe ventricular arrhythmias.

Two predictive factors of mortality were identified: low diastolic blood pressure and occurrence of ventricular fibrillation, hospital mortality is therefore linked to hemodynamic instability and its electrophysiological consequences.

#### **Conclusion:**

Sustained ventricular tachycardia (SVT) is one of the most serious ventricular arrhythmias, rarely observed early in acute coronary syndrome with ST segment elevation, because of its frequent degeneration into ventricular fibrillation.

In our study we reported SVT as a separate entity, while in the literature; it's reported in the group of severe ventricular arrhythmias.

Its predictive factors according to our study are: low diastolic blood pressure, severe coronary artery lesions and High-Sensitivity Troponin (hs-Trop)  $\geq 5$  ng/ml.

The occurrence of SVT increases the risk of hospital mortality, related to hemodynamic instability and its electrophysiological consequences.

To our knowledge the low diastolic blood pressure and severe coronary artery lesions as predictors of sustained Ventricular Tachycardia were reported for the first time.

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