

Presentation of acute coronary syndromes during covid-19 pandemic: attitude of patients and health care professionals

Covid-19 pandemisi sırasında akut koroner sendromların başvuru özellikleri: hastaların ve sağlık çalışanlarının tutumu

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SUMMARY

Objective: COVID-19 pandemic resulted in substantial alterations in health care demand and delivery. The aim of this study is to investigate presentations and treatment strategies of acute coronary syndromes during COVID-19 pandemic.

Method: The patients presented with acute coronary syndrome between March 10 2020-June 1 2020 and in the same timeframe in 2019 were analysed retrospectively. The patients in 2020 constituted the Pandemia group and the patients in 2019 were accepted to be controls. Presentation types, demographical and clinical characteristics as well as treatment strategies were compared between two groups.

Results: There was a 15.2% reduction in the number of acute coronary syndromes during the COVID-19 pandemic lockdown when compared to previous year. The number of patients presenting with STEMI was higher in Pandemia group when compared to controls ($p=0.033$). Less patients underwent coronary angiography and percutaneous coronary intervention in Pandemia group ($p=0.012$ and $p=0.033$, respectively). Antiagregant choice of physicians between clopidogrel and ticagrelol was similar ($p=0.227$). Aspartate aminotransferase and lactate dehydrogenase levels were significantly higher in Pandemia group compared to controls ($p=0.030$ and $p=0.003$, respectively). Hospitalization duration was similar between two groups while serum levels of troponin on the discharge day was significantly higher in Pandemia group ($p=0.512$ and $p=0.001$, respectively).

Conclusions: COVID-19 pandemic resulted in a decline in the number of acute coronary syndromes and a trend towards conservative approach in patients with NSTEMI. It is controversial whether mortality benefit driven from COVID-19 prevention is greater than the consequences these presentation and treatment alterations in heart attack patients.

Keywords: COVID-19, Pandemia, Acute coronary syndrome

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ÖZET

Amaç: COVID-19 pandemisi, sağlık hizmeti talep ve sunumunda önemli değişikliklere neden oldu. Bu çalışmanın amacı, COVID-19 pandemisine bağlı kısıtlamalar sırasında akut koroner sendromların başvuru şekillerini ve tedavi stratejilerini araştırmaktır.

Yöntem: 10 Mart 2020 ve 1 Haziran 2020 tarihleri arasında ve 2019' un aynı zaman diliminde akut koroner sendrom ile başvuran hastalar retrospektif olarak incelendi. 2020'deki hastalar Pandemi grubunu oluştururken, 2019' daki hastalar ise Kontrol grubu olarak kabul edildi. Başvuru şekilleri, demografik ve klinik özellikler ve tedavi stratejileri iki grup arasında karşılaştırıldı.

Bulgular: COVID-19 pandemisi sırasında akut koroner sendrom sayısında bir önceki yıla göre %15,2' lik bir azalma gözlemlendi. Pandemi grubunda STEMI ile başvuran hasta sayısı kontrol grubuna göre daha yüksekti ($p = 0.033$). Pandemi grubunda daha az sayıda hastaya koroner anjiyografi ve perkütan koroner girişim uygulandı (sırasıyla $p = 0.012$ ve $p = 0.033$). Klopidoğrel ve tikagrelol arasındaki antiagregan seçimi benzerdi ($p = 0.227$). Aspartat aminotransferaz ve laktat dehidrogenaz düzeyleri, Pandemi grubunda Kontrol grubuna göre anlamlı derecede yüksekti (sırasıyla $p = 0.030$ ve $p = 0.003$). Hastanede yatış süresi iki grup arasında benzer iken taburculuk gününde serum troponin düzeyleri Pandemi grubunda anlamlı olarak daha yüksek saptandı (sırasıyla $p = 0.512$ ve $p = 0.001$).

Sonuç: COVID-19 pandemisi, akut koroner sendromların sayısında azalma ve NSTEMI hastalarında konservatif yaklaşım yönünde bir eğilim ile sonuçlanmıştır. COVID-19 önlenmesinden kaynaklanan mortalite faydasının, kalp krizi hastalarındaki bu başvuru ve tedavi değişikliklerinin ortaya çıkardığı sonuçlardan daha fazla olup olmadığı tartışmalıdır.

Anahtar sözcükler: COVID-19, pandemi, akut koroner sendrom

INTRODUCTION

The Globe has faced with an extraordinary infectious disease recently. The first cases were reported in Wuhan-China in December 2019. The accumulation of serious pneumonia cases in hospitals and inability to isolate well-known pathogens raised suspicion of a possible unknown pathogen which is identified and named as SARS-CoV-2 afterwards. This new virus (SARS-CoV-2) belongs to the same severe acute respiratory syndrome-coronavirus (SARS-CoV) and Middle East respiratory syndrome-coronavirus (MERS-CoV) family. Coronavirus disease 2019 (COVID-19) has rapidly spread worldwide and adopted to be a pandemic by World Health Organization (WHO) in March 11 2020. Many countries have announced lockdown since then and most of the hospitals were reorganized as pandemic centers.

Beyond COVID-19 disease, acute coronary syndromes (ACS) continue to be the leading cause of mortality. While pandemic hospitals halted elective interventional procedures during lockdown, catheter laboratories have continued to admit patients presenting with ACS. On the other hand, since hospitals were considered to be the most hazardous places in terms of being infected, most patients hesitated to apply to hospitals even in the presence of severe symptoms. Chest pain is one of the most worrying symptoms in patients and the leading cause of emergency department admissions. Could the fear of being infected prevent hospital admission even in patients with chest pain? This study aimed to investigate the presentation of acute coronary syndromes and attitude of patients as well as the health care

professionals during COVID-19 pandemic lockdown.

MATERIAL AND METHODS

A total of 146 patients were included in the study. The study had a retrospective design. Sixty seven patients who presented with acute coronary syndrome between March 10 2020 and June 1 2020 constituted the Pandemic group. March 10 2020 was the day when first COVID-19 case was reported in Turkey, and June 1 2020 was the day when end of lockdown and beginning of new-normal life was announced by the government. The patients were analysed retrospectively from the hospital recordings. Seventy nine patients presented with acute coronary syndrome between March 10 2019 and June 1 2019, briefly in the same timeframe of the previous year were recruited as the Control group. Demographical characteristics, presentation types, laboratory findings, hospitalization durations and treatment approaches including choice of conservative or invasive strategy were compared between two groups.

Definitions

Diagnosis of acute coronary syndrome was established under the guidance of Expert Consensus Document on Fourth universal definition of myocardial infarction published in 2018¹. Patients with persistent chest discomfort or other symptoms suggestive of ischaemia and ST-segment elevation in at least two contiguous leads or new bundle branch blocks with ischaemic repolarization patterns along with evidence of myocardial injury (defined as an elevation of cardiac troponin values with at least one value

above the 99th percentile upper reference limit) were accepted as ST elevation myocardial infarction (STEMI). ST elevation on electrocardiogram was defined as new ST-elevation at the J-point in two contiguous leads with the cut-point: ≥ 1 mm in all leads other than leads V2–V3 where the following cut-points apply: ≥ 2 mm in men ≥ 40 years; ≥ 2.5 mm in men < 40 years, or ≥ 1.5 mm in women regardless of age. Patients with acute chest pain but no persistent ST-segment elevation (ECG changes may include transient ST-segment elevation, persistent or transient ST-segment depression, T-wave inversion, flat T waves or pseudo-normalization of T waves or the ECG may be normal) with evidence of myocardial injury (defined as an elevation of cardiac troponin values with at least one value above the 99th percentile upper reference limit) were accepted as non ST elevation myocardial infarction (NSTEMI). Unstable angina pectoris (USAP) was defined as myocardial ischaemia at rest or minimal exertion in the absence of cardiomyocyte necrosis demonstrated by normal cardiac troponin values.

Statistical analysis

Statistical analysis was performed using SPSS version 11.0 (SPSS Inc, Chicago, IL, USA) pocket program. The data were expressed as the mean \pm standard deviation (SD) and were tested for normal distribution using the Kolmogorov-Smirnov test. Comparisons between patients were made by using Student's independent t-test for normally distributed data and Mann-Whitney U test for non-normal distributed data. The chi-square test was used to compare the categorical variables. A p value of <0.05 was considered statistically significant.

RESULTS

The number of patients presented with acute coronary syndrome was 67 during lockdown and they were defined as Pandemia group. Control group consisted of 79 patients who presented with acute coronary syndrome during the same timeframe in 2019. There was a 15.2% reduction in the number of acute coronary syndromes during the COVID-19 pandemic lockdown when compared to previous year. The number of patients presenting with STEMI was higher in Pandemia group when compared to controls ($p=0.033$) while the number of patients with NSTEMI was similar ($p=0.033$ and $p=0.054$, respectively). There was no statistically significant difference between two groups in terms of age, gender and presence of hypertension, diabetes mellitus and chronic renal failure. The number of patients who underwent coronary angiography was lower in Pandemia group ($p=0.012$). Moreover percutaneous coronary intervention rates were also lower in Pandemia group ($p=0.033$). Coronary artery bypass grafting rates were similar between two groups ($p=0.165$). Antiagregant choice of physicians between clopidogrel and ticagrelol was similar ($p=0.227$). There was no statistically significant difference between Pandemia and Control groups in terms of serum sodium, potassium, LDL cholesterol, total cholesterol, alanine aminotransferase, hemoglobin A1C, glucose and creatinine levels. Serum HDL cholesterol levels were lower in Pandemia group ($p=0.049$) while aspartate aminotransferase and lactate dehydrogenase levels were significantly higher. ($p=0.030$ and $p=0.003$, respectively). Hospitalization duration was similar between two groups while serum levels of troponin on the discharge day was significantly higher in Pandemia group ($p=0.512$, and $p=0.001$, respectively).

Table 1: Comparison of demographical and clinical characteristics between Pandemia and Control groups

	Pandemia group (n:67)	Control group (n:79)	P
Age	65.5 (±13.6)	66.6 (±11.5)	0.601
STEMI (n, %)	47 (70.1 %)	67 (84.8 %)	0.033
Gender (Female, %)	22 (32.8 %)	19 (24.1 %)	0.344
NSTEMI (n, %)	20 (29.9 %)	13 (16.5 %)	0.054
Daibetes Mellitus	18 (26.9 %)	26 (32.9 %)	0.398
Hypertension	60 (89.6%)	63 (79.7%)	0.105
CRF	2 (3%)	5 (6.3%)	0.337
Na	138.9 (±3.2)	139.7 (±3.2)	0.151
K	4.4 (±0.5)	4.5 (±0.7)	0.167
HDL Cholesterol	38 (23-73)	42 (22-103)	0.049
LDL Cholesterol	108.5 (37-203)	102.5 (14-195)	0.083
Total Cholesterol	172.0 (109-310)	174.5 (94-318)	0.380
Triglyceride	137.5 (46-767)	141.5 (46-575)	0.695
AST	27.0 (11-431)	21.5 (8-327)	0.030
ALT	24.5 (7-154)	20.0 (7-368)	0.796
LDH	247.0 (110-993)	226.0 (112-691)	0.048
HbA1C	5.9 (4.8-13.8)	5.8 (5.0-11-7)	0.911
Glucose	109.0 (75-426)	112 (57-414)	0.523
BUN	42 (15-133)	35 (20-236)	0.933
Creatinine	0.9 (0.3-3.0)	0.9 (0.2-3.7)	0.710
WBC	8.3 (4.1-19.4)	8.1 (4.5-29.6)	0.978
Hb	13.8 (9.9-17.4)	13.4 (8.7-17.1)	0.246
Platletet	239 (142-422)	233 (109-470)	0.450

STEMI: St elevation myocardial infarction, **NSTEMI:** Non St elevation myocardial infarction, **CRF:** Chronic renal failure, **LDH:** Lactate dehydrogenase, **Na:** Sodium, **K:** Potassium, **HDL:** High density lipoprotein, **LDL:** Low density lipoprotein, **AST:** Aspartate aminotransferase, **ALT:** Alanine aminotransferase, **LDH:** Lactate dehydrogenase, **Hb:** Haemoglobin, **BUN:** Blood urea nitrogen, **WBC:** White blood cell

Table 2: Comparison of treatment strategies between Pandemia and Control groups

	Pandemia group (n:67)	Control group (n:79)	P
CAG	45 (67.2%)	67 (84.8%)	0.012
PCI	37 (55.2%)	57 (72.2%)	0.033
CABG	5 (7.5%)	2 (2.5%)	0.165
Ticagrelol	20 (29.8%)	31 (39.2%)	0.227
Hospitalization duration	3 (1-15)	3 (1-24)	0.894
Troponin on discharge day	0.34 (0.01-8.00)	0.03 (0.02-6.00)	0.001

CAG: Coronary angiography, **PCI:** Percutaneous coronary intervention, **CABG:** Coronary artery bypass grafting

DISCUSSION

The number of acute coronary syndromes diagnosed during COVID-19 pandemic lockdown were lower when compared with the same timeframe in 2019. On the other hand, the number of STEMI cases were higher during lockdown. There was a tendency for NSTEMI patients to be treated conservatively since the number of CAG and PCI were significantly lower in Pandemia group. Duration of hospitalization was similar between two groups but serum levels of troponin on discharge day was significantly higher in Pandemia group.

With the world health organization declaring COVID-19 disease as a pandemic, a climate of fear prevailed all over the world. Most of the countries announced intermittent or continuous lockdown. Elective outpatient clinic visits were suspended and the public was warned to apply to the hospitals only in emergency conditions. All these seemed to effect the hospital admission behaviours of the patients even in the presence of chest pain, the most worrying symptom for the patients. Recent reports have presented that the lockdown imposed due to the spread of COVID-19 infection has led to a change in the number and type of cardiology admissions in Italy²⁻⁴. Similar findings were

reported from United States⁵⁻⁷. Tan W. et al reported single center trends in acute coronary syndromes⁸. Reduction in the number of cases were presented from United Kingdom by Wilson SJ et al^{9,10}. Decline in the number of cases was also noticeable in Austria, France, Germany, Switzerland, Hong Kong and Egypt¹¹⁻¹⁸. These reports have demonstrated single or multi-center experiences about acute coronary syndrome admissions during COVID-19 pandemic. As it is seen the pandemic effected hospital admission for acute coronary syndromes all over the world. Our study results were also consistent with the previous reports. Although the total number of acute coronary syndromes including STEMI and NSTEMI was lower during pandemic, the ratio of STEMI was higher. This can be interpreted as increased chest pain threshold for patients to admit emergency departments. Significant amount of NSTEMI cases probably remained undiagnosed due to the fear of being infected.

Beside number of acute coronary syndromes, we also aimed to investigate frequency of interventional procedures. The number of patients who underwent coronary angiography was significantly lower in Pandemia group. In our center, all patients with STEMI underwent urgent coronary angiography and none received fibrinolytic therapy in accordance with the position statement of EAPCI on invasive management of acute coronary syndromes during the COVID-19 pandemic [19]. On the other hand, it seems that here was a tendency for conservative approach in patients with NSTEMI when decline in the number of interventional procedures is considered.

The major symptoms of COVID-19 disease were identified to be fewer and dyspnea. Since dyspnea is the most frequently seen side effect of ticagrelor, antiaggregant choice of the physicians was also compared between two groups. Did the concern that ticagrelor may mask COVID-19 disease have any effect on choice of antiaggregant therapy? This probably made no sense among physicians since rates of clopidogrel and ticagrelor use were similar. Another significant point is higher levels of lactate dehydrogenase and aspartate aminotransferase in the Pandemia group which demonstrates more myocardial damage. This may arise from longer delay in hospital admission. Moreover higher levels of troponin on discharge day in the Pandemia group was also an indicator of more damaged myocardium even in treated patients.

The outcomes of our study demonstrated a downwards trend in the hospital admissions of acute coronary syndromes. These findings were in line with the previous reports that have shown a

magnificent decline in the number of acute coronary syndromes from various countries. To the best of our knowledge this is the first scientific document to report the decrease in the number of acute coronary syndromes in the COVID-19 pandemic in Turkey. Although reduction in the number of overall hospital admissions was an expected finding, neglect of chest pain, the most worrying symptom for patients, was surprising. The fear of being infected probably raised the threshold for the patients to apply emergency departments.

Lack of some other demographical data such as family history and smoking status as well as clinical data such as onset to door time and door to balloon time can be considered as a limitation. However our study has a retrospective design and all the data were collected following announcement of the end of the lockdown after the patients were discharged. Unfortunately the above mentioned demographical and clinical data were not available in our recordings. Moreover this study reveals the outcomes from a single center. A multicenter design would probably better reflect the overall trend in our country. On the other hand the majority of the previous studies were reported from single centers.

Consequently, similar reports from all over the world proves that decline in the number of acute coronary syndromes was not occasional. COVID-19 fear resulted in "at home" heart attacks. It is controversial whether staying at home in order to prevent COVID-19 disease had advantage over the poor outcomes associated with neglected acute coronary syndromes in terms of mortality.

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