Since January 2020 Elsevier has created a COVID-19 resource centre with free information in English and Mandarin on the novel coronavirus COVID-19. The COVID-19 resource centre is hosted on Elsevier Connect, the company's public news and information website.

Elsevier hereby grants permission to make all its COVID-19-related research that is available on the COVID-19 resource centre - including this research content - immediately available in PubMed Central and other publicly funded repositories, such as the WHO COVID database with rights for unrestricted research re-use and analyses in any form or by any means with acknowledgement of the original source. These permissions are granted for free by Elsevier for as long as the COVID-19 resource centre remains active.
ORIGINAL ARTICLE

The impact of the COVID-19 pandemic on acute coronary syndrome admissions to a tertiary care hospital in Portugal

João Calvão*, Ana Filipa Amador, Catarina Martins da Costa, Paulo Maia Araújo, Teresa Pinho, João Freitas, Sandra Amorim, Filipe Macedo

Serviço de Cardiologia, Hospital São João, Porto, Portugal

Received 20 October 2020; accepted 31 January 2021
Available online 20 March 2021

KEYWORDS
COVID-19;
Acute coronary syndrome;
Percutaneous coronary intervention;
Public health

Abstract
Introduction and objectives: The coronavirus SARS-CoV-2 (COVID-19) pandemic has been an unmatched challenge to global healthcare. Although the majority of patients admitted with acute coronary syndrome (ACS) may not be infected with COVID-19, the quarantine and public health emergency measures may have affected this particular high risk group. The objective of this study is to assess the impact of the early period of the COVID-19 pandemic on ACS admissions and clinical course in a tertiary care hospital in Portugal’s most affected region.

Methods: This retrospective, case-control study included patients admitted with a diagnosis of ACS during March and April 2020 (pandemic group) and in the same period in 2019 (control group). Clinical course and complications were also assessed.

Results: During the pandemic, there were fewer ACS admissions but presentation was more severe, with a larger proportion of acute ST-elevation myocardial infarctions (54.9% vs. 38.8%, p=0.047), higher maximum troponin levels and greater prevalence of left ventricular systolic dysfunction at discharge (58.0% vs. 35.0%, p=0.01). In this population, although not statistically significant, it was observed a delay between the onset of symptoms and percutaneous coronary intervention, which may traduce a deferred search for urgent medical care during the pandemic.

Conclusion: The lockdown phase of COVID-19 pandemic was associated with fewer and more severe ACS in a Tertiary Care Hospital in Portugal’s most affected region by the pandemic.

© 2021 Sociedade Portuguesa de Cardiologia. Published by Elsevier España, S.L.U. This is an open access article under the CC BY-NC-ND license (http://creativecommons.org/licenses/by-nc-nd/4.0/).

* Corresponding author.
E-mail address: joacalvao1@gmail.com (J. Calvão).

https://doi.org/10.1016/j.repc.2021.01.007
0870-2551/© 2021 Sociedade Portuguesa de Cardiologia. Published by Elsevier España, S.L.U. This is an open access article under the CC BY-NC-ND license (http://creativecommons.org/licenses/by-nc-nd/4.0/).
Introduction

COVID-19, caused by the severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2), is a highly contagious condition that may lead to serious respiratory illness and premature death; with up to 20% of hospitalized patients requiring mechanical ventilation and intensive care. The first cases were reported in December 2019 in Wuhan, Hubei, China but rapidly spread worldwide, being declared as a pandemic by the WHO on 11 March 2020. Currently, there are about 34 million cases, with more than 1 million deaths globally.

In order to contain the COVID-19 pandemic, there was a widespread lockdown worldwide, with restrictive home confinement and social distancing measures. Overwhelmed healthcare systems were also promptly restructured, expanding intensive care units and altering or cancelling non-essential procedures.

In the meantime, many countries reported a significant reduction in hospital admissions for several acute conditions that require life-saving evidence-based emergent treatment, such as acute coronary syndrome (ACS) and stroke. Moreover, patients with cardiovascular disease are at increased risk of severe COVID-19 illness, endorsing the direct and indirect consequences of delayed care of this particular group of patients. In Portugal, the first case was reported on 2 March 2020 in Porto. The mandatory national lockdown was imposed from 18 March to 2 May, amounting to 25 190 cases and 1023 deaths due to COVID-19 at the end of this period. The northern region of Portugal, mainly Porto’s metropolitan area, had the highest incidence of COVID-19 cases, reaching around 14 951 cases and 585 deaths.

The aim of this study was to assess the impact of the COVID-19 pandemic on ACS admissions to the Cardiology Department of the Centro Hospitalar Universitário de São João, a tertiary care hospital in Porto’s metropolitan area.

Methods

This retrospective, case-controlled study included all patients admitted to the Cardiology Department of Centro Hospitalar Universitário de São João with a diagnosis of ACS, during the months of March and April of 2020 (pandemic group) and in the same period of 2019 (control group). Patients who tested positive for SARS-CoV-2, and therefore admitted to COVID-19 exclusive units, were also included.

The database used in this study was completed retrospectively using the available clinical records. Patients were selected based on the final diagnosis at the time of discharge (ST-segment elevation myocardial infarction (STEMI), non-ST-segment elevation myocardial infarction (NSTEMI) or unstable angina (UA)). ACS was defined according to the Fourth Universal Definition of Myocardial Infarction.
Demographic characteristics (gender, age, presence of cardiovascular risk factors, chronic kidney disease, previous vascular events and LV dysfunction), in-hospital clinical course (length of stay, Killip class, maximum troponin value, complications, prevalence of left ventricular (LV) systolic dysfunction at discharge), and short term mortality were compared between groups. Continuous variables were summarized as mean ± standard deviation or median (25, 75 percentiles) and categorical variables as proportions. Groups were compared using Student’s t tests, Wilcoxon Rank Sum tests, and χ² tests, as appropriate. Values of p<0.05 were considered significant. All statistical analyses were performed using IBM SPSS statistics 25 (New York, United States).

Results

Baseline patient characteristics

Our study included 151 patients hospitalized with ACS in the two aforementioned periods. There were 71 ACS admissions between March and April, 2020 (n=71), representing a reduction of 11.3% relative to the same period in 2019 (n=80).

Overall, there were no significant differences in age, gender, and cardiovascular risk factors in both groups (Table 1). The pandemic group had a higher prevalence of atherosclerotic disease (defined as history of peripheral, coronary or cerebral arterial disease) (45.1% vs. 28.7%, p=0.04), although there was no difference considering previous ACS or revascularization procedures.

Three patients from the pandemic group tested positive for SARS-COV-2.

Initial presentation and management

The proportion of ACS presenting as STEMI was significantly higher in the pandemic group (54.9% vs. 38.8%, p=0.047). This was driven not only by a decrease in the number of NSTEMI and UA patients (49 vs. 32) but also by an increase in the absolute number of STEMIs (31 vs. 39).

Among STEMI admissions, although not statistically significant, there was a longer period between time of first symptom onset to first medical contact (FMC), during the pandemic (control vs. pandemic; median 145±178 min vs. 180±360 min, p=0.31) (Table 2). This was more pronounced considering NSTEMI and UA presentations (control vs. pandemic; median 180±330 min vs. 300±1170 min, p=0.38).

The proportion of STEMI patients who underwent emergent angiography was similar in both years (87.1% vs. 82.1%, p=0.41). Time from FMC to emergent percutaneous coronary intervention (PCI) was also similar (150+/−270 min vs. 120+/−492 min, p=0.56). Among patients admitted to hospitals without the ability to perform primary PCI, there was no significant difference in TMC to PCI times in both groups (190+/−632 vs. 210+/−600, p=0.57).

One of the patients from the pandemic group had known SARS-COV-2 infection at the time of presentation. He was admitted with inferior wall STEMI (less than one hour after symptom onset) and underwent fibrinolysis, followed by routine PCI. The decision to perform fibrinolysis was made based on a STEMI reperfusion algorithm published at the time.¹⁰

Clinical course

Both groups had a similar length of hospitalization (control vs. pandemic: 5±8 vs. 5±6 days, p=0.377) (Table 3). There was no difference regarding Killip class classification. Maximum measured high-sensitivity Troponin I was significantly higher in the pandemic group (19 439±70 265 ng/L vs. 10 222±36 870 ng/L, p=0.03). There were also significantly more patients with LV systolic dysfunction at discharge (58.0% vs. 35.0%, p=0.01).

Finally, there was a greater proportion of patients in the pandemic group with in-hospital complications (23.9% vs. 13.8%, p=0.11) and in-hospital (5.6% vs. 1.3%, p=0.15) and 30-day (7.0% vs. 2.5%, p=0.18) mortality, although these did not reach statistical significance (Table 3). At the time of discharge, there was no difference in the number of patients receiving angiotensin-converting-enzyme inhibitors and/or beta-blockers (control vs. pandemic; 96.8% vs. 94.6%, p=0.55). Almost all patients had undergone angio-

---

Table 1  Baseline characteristics of the participants.

|                      | Control group (N=80) | Pandemic group (N=71) | p value |
|----------------------|---------------------|-----------------------|---------|
| Age (mean, SD)       | 65.7 (±12.8)        | 63.3 (±12.7)          | 0.25    |
| Male (%)             | 60 (75.0)           | 56 (78.9)             | 0.35    |
| ≥1 CVRF (%)          | 78 (97.5)           | 64 (90.1)             | 0.06    |
| Hypertension (%)     | 48 (60.0)           | 49 (69.0)             | 0.25    |
| Dyslipidemia (%)     | 49 (61.3)           | 38 (53.5)             | 0.33    |
| Smoking history (%)  | 39 (51.2)           | 40 (57.1)             | 0.47    |
| Diabetes (%)         | 26 (32.5)           | 23 (32.4)             | 0.99    |
| Atherosclerotic disease (%) | 23 (28.7) | 32 (45.1) | 0.04*   |
| Previous ACS (%)     | 15 (18.8)           | 17 (23.9)             | 0.44    |
| Previous coronary revascularization (%) | 16 (20.0) | 17 (23.9) | 0.56    |
| CKD (%)              | 12 (15.0)           | 8 (11.3)              | 0.50    |
| Previous LV dysfunction (LVEF<50%) (%) | 7 (8.8)     | 6 (8.5)               | 0.95    |

ACS: acute coronary syndrome; CVRF: cardiovascular risk factors; CKD: chronic kidney disease; LV: left ventricular; LVEF: left ventricular ejection fraction; SD: standard deviation. *p<0.05.
Table 2  Initial presentation and management of ST-elevation myocardial infarction patients.

|                      | Control group (N=31) | Pandemic group (N=39) | p value |
|----------------------|----------------------|-----------------------|---------|
| Primary PCI strategy (%) | 27 (87.1)            | 32 (82.1)             | 0.41    |
| Fibrinolysis (%)      | 0 (0.0)              | 1 (1.4)               | 0.47    |
| Time from symptoms to FMC (min) (median, IQR) | 145 (178)           | 180 (360)             | 0.31    |
| Time from FMC to PCI (min) (median, IQR)      | 150 (388)            | 120 (426)             | 0.4     |
| Time from symptoms to PCI (min) (median, IQR) | 270 (750)            | 330 (804)             | 0.56    |

FMC: first medical contact; IQR: interquartile range; PCI: percutaneous coronary intervention.

Table 3  Clinical course and outcomes.

|                      | Control group (N=80) | Pandemic group (N=71) | p value |
|----------------------|----------------------|-----------------------|---------|
| Length of stay (days) (median, IQR) | 5 (8)                | 5 (6)                 | 0.38    |
| Coronary angiography (%) | 78 (97.5)            | 67 (94.4)             | 0.42    |
| Revascularization (%) | 56 (70.0)            | 58 (82.9)             | 0.66    |
| PCI                  | 45 (56.3)            | 53 (76.8)             | 0.01*   |
| CABG                 | 12 (15.0)            | 6 (8.7)               | 0.24    |
| Killip class ≥2 (%)  | 16 (22.5)            | 19 (28.8)             | 0.40    |
| Max troponin (ng/L) (median, IQR) | 10 222 (36 870) | 19 439 (70 265) | 0.03*   |
| LV dysfunction at discharge (EF<50%) (%) | 28 (35.0)          | 40 (58.0)             | 0.01*   |
| In-hospital complications (%) | 11 (13.8)         | 17 (23.9)             | 0.11    |
| Cardiac arrest (%)   | 2 (2.5)              | 8 (11.3)              |         |
| Pericarditis (%)     | 3 (3.8)              | 5 (7.0)               |         |
| 2nd/3rd degree AV block (%) | 1 (1.3)            | 5 (7.0)               |         |
| Sustained VT (%)     | 2 (2.5)              | 3 (4.2)               |         |
| VF<24 h (%)          | 1 (1.3)              | 3 (4.2)               |         |
| VF>24 h (%)          | 0 (0.0)              | 3 (4.2)               |         |
| LV thrombus (%)      | 4 (5.0)              | 2 (2.8)               |         |
| Mechanical complication (%) | 0 (0.0)           | 1 (1.4)               |         |
| Moderate-severe MR (%) | 0 (0.0)           | 6 (8.5)               |         |
| AF (%)               | 5 (6.3)              | 3 (4.2)               |         |
| Stroke (%)           | 2 (2.5)              | 6 (8.5)               |         |
| In-hospital mortality (%) | 1 (1.3)         | 4 (5.6)               | 0.15    |
| 30-day mortality (%) | 2 (2.5)              | 5 (7.0)               | 0.18    |

AF: atrial fibrillation; AV: atrioventricular; CABG: coronary artery bypass graft; EF: ejection fraction; IQR: interquartile range; LV: left ventricular; MR: mitral regurgitation; PCI: percutaneous coronary intervention; VF: ventricular fibrillation; VT: ventricular tachycardia. *p<0.05.

Globaly, there was a trend towards higher revascularization rates in the pandemic group (70.0% vs. 82.9%, p=0.66), with a significantly higher number of patients undergoing PCI (56.3% vs. 76.8%, p=0.01) and fewer patients undergoing coronary artery bypass grafting (15.0% vs. 8.7%, p=0.24).

Discussion

In the early COVID-19 pandemic, the number of patients requiring hospitalization and intensive care overwhelmed the healthcare facilities in numerous countries. Meanwhile, a major decline in hospital admissions with ACS in several medical centers was observed, possibly due to the quarantine and other large-scale public health measures. Some centers in highly affected regions, like Northern Italy and China, implemented early fibrinolysis protocols in STEMI treatment, with a direct decrease in the number of primary PCIs. Afterwards, position statements reinforced that during the COVID-19 pandemic, the standard of care for STEMI patients at PCI-capable hospitals remained primary PCI when it could be provided in a timely manner. During the first phase of the COVID-19 pandemic in Portugal, the population of Porto and the surrounding urban area was the most affected by SARS-COV-2 disease. Contrary to what was seen in other countries, we did not see a clear reduction in hospitalization but presentations were more severe, with a larger proportion of STEMs, higher maximum troponin values and greater prevalence of LV systolic dysfunction at discharge. The fear of becoming infected by SARS-Cov-2 may have discouraged patients that experienced milder symptoms to seek urgent medical care. In fact, previous studies in other countries noted a deferred search for urgent medical care during the pandemic. Although not statistically significant, our data suggests that there was a delay between the onset of symptoms and search for medical care during the pandemic. The non-significance of this finding may be related to the small sample size. This delay may explain
the larger number of STEMI patients admitted and consequently the higher prevalence of LV systolic dysfunction at the time of discharge. It is of relevance that during the pandemic there was a significantly higher prevalence of atherosclerosis among patients and this may have had some impact on the differences seen in the outcomes of the two groups.

In-hospital and 30-day mortality were numerically higher in patients admitted during the pandemic, similar to those observed in other studies.12 Our study was innovative since it also assessed complications related to ACS; the rate was overall low but a higher proportion of complications occurred during the pandemic. This may translate into higher mortality and higher morbidity in the near future. Our concern also extends to patients who did not reach out for medical care during the pandemic, which may negatively affect long-term cardiovascular health. This may partially explain the higher revascularization rates, since milder cases may not have come to medical attention. More studies should be designed to explore these events.

Importantly, there was no evidence of significant delay in the healthcare response to STEMI patients, including in cases initially admitted to hospitals without the ability to perform primary PCI. Revascularization rates were not precluded and fibrinolysis use was residual. These data may be explained since Portugal did not reach a situation in which the health care system was overwhelmed or began to collapse.34

This analysis has some limitations. Only patients admitted to one center were included. Additionally, acute morbidity and mortality may be underestimated, since patients who died in the pre-hospital phase/emergency room or were admitted at other intensive care units were not included.

Conclusion

During the lockdown phase of the COVID-19 pandemic, there were fewer admissions for ACS but more serious presentations in a tertiary hospital center in the region most affected by COVID-19 in Portugal, possibly due to a deferred search for healthcare by the population. As ACS requires prompt in-hospital assessment and treatment in order to prevent significant morbidity and mortality, the general population should be made aware of this problem. Also, a public healthcare safety message addressing non-COVID emergencies must be transmitted. The global healthcare policy focused on addressing the current pandemic must not disregard other serious medical emergencies. More studies are needed to assess the long-term repercussion of the COVID-19 pandemic in acute cardiovascular patients.

Conflicts of interest

The authors have no conflicts of interest to declare.

References

1. Guan W-J, Ni Z-Y, Hu Y, et al. Clinical characteristics of coronavirus disease 2019 in China. N Engl J Med. 2020;382:1708–20.
2. Rodríguez-Morales AJ, Cardona-Ospina JA, Gutiérrez-Ocampo E, et al. Clinical, laboratory and imaging features of COVID-19: a systematic review and meta-analysis. Travel Med Infect Dis. 2020;34:101623.
3. Timeline of WHO’s response to COVID-19. https://www.who.int/news-room/detail/29-06-2020-covidtimeline [accessed 12.08.20].
4. WHO. https://covid19.who.int/table [accessed 03.10.20].
5. World Economic Forum: ‘Nearly 3 billion people around the globe under COVID-19 lockdown—today’s coronavirus updates’; 26th March 2020. https://www.weforum.org/Agenda/2020/03/todays-coronavirus-updates/ [accessed 12.08.20].
6. BBC News: “Coronavirus: The world in lockdown in maps and charts”; 7th April 2020. https://www.bbc.com/news/world-52103747 [accessed 12.08.20].
7. Grasselli G, Pesenti A, Cecconi M. Critical care utilization for the COVID-19 outbreak in Lombardy Italy: early experience and forecast during an emergency response. JAMA. 2020;323:1545.
8. Saglietto A, D’Ascenzo F, Zoccai GB, et al. COVID-19 in Europe: the Italian lesson. Lancet. 2020;395:1110–1.
9. De Filippo O, D’Ascenzo F, Angelini F, et al. Reduced rate of hospital admissions for ACS during COVID-19 outbreak in Northern Italy. New Engl J Med. 2020, http://dx.doi.org/10.1056/NEJMct2009166.
10. Garcia S, Albaghadi MS, Meraj PM, et al. Reduction in ST-segment elevation cardiac catheterization laboratory activations in the United States during COVID-19 pandemic. J Am Coll Cardiol. 2020;75:2871–2.
11. Rodríguez-Leor O, Cid-Alvarez B, Ojeda S, et al. Impacto de la pandemia de COVID-19 sobre la actividad asistencial en cardiología intervencionista en España. REC Interv Cardiol. 2020;2:82–9.
12. Metzler B, Siostrzonek P, Binder RK, et al. Decline of acute coronary syndrome admissions in Austria since the outbreak of COVID-19: the pandemic response causes cardiac collateral damage. Eur Heart J. 2020;41:1852–3.
13. Tam CF, Cheung KS, Lam S, et al. Impact of coronavirus disease 2019 (COVID-19) outbreak on ST-segment-elevation myocardial infarctionare in Hong Kong, China. Circ Cardiovasc Qual Outcomes. 2020;13:e006261.
14. Wilson SJ, Conolly MJ, Elghamry Z, et al. Effect of the COVID-19 Pandemic on ST-segment–elevation myocardial infarction presentations and in-hospital outcomes. Circ Cardiovasc Interv. 2020:13.
15. Bhatt AS, Moscone A, McEllrath EE, et al. Declines in hospitalizations for acute cardiovascular conditions during the COVID-19 pandemic: a multicenter tertiary care experience. J Am Coll Cardiol. 2020;76:280–8.
16. Solomon MD, McNulty EJ, Rana JS, et al. The COVID-19 pandemic and the incidence of acute myocardial infarction. N Engl J Med. 2020.
17. De Rosa S, Spaccarotella C, Basso C, et al. Reduction of hospitalizations for myocardial infarction in Italy in the COVID-19 era. Eur Heart J. 2020.
18. Morelli N, Rota E, Terracciano C, et al. The Baffling case of ischemic stroke disappearance from the casualty department in the COVID-19 era. Eur Neurol. 2020;83:213–5, 10.
19. Zhao J, Li H, Kung D, et al. Impact of the COVID-19 epidemic on stroke care and potential solutions. Stroke. 2020.
20. Manhem MM, Spata E, Goldacre R. COVID-19 pandemic and admission rates for and management of acute coronary syndromes in England. Lancet. 2020;396:381–9.
21. Clerkin KJ, Fried JA, Raikhelkar J, et al. Coronavirus disease 2019 (COVID-19) and cardiovascular disease. Circulation. 2020;141:1648–55.
22. Yang J, Zheng Y, Gou X, et al. Prevalence of comorbidities in the novel Wuhan coronavirus (COVID-19) infection: a systematic review and meta-analysis. Int J Infect Dis. 2020;94:91–5.
23. Wang B, Li R, Lu Z, et al. Does comorbidity increase the risk of patients with COVID-19: evidence from meta-analysis. Aging (Albany NY). 2020;12:6049–57.
24. Relatório de Situação COVID-19 em Portugal; 3rd March 2020. https://covid19.min-saude.pt/wp-content/uploads/2020/03/Relatório-de-Situação-1.pdf
25. Relatório de Situação COVID-19 em Portugal; 2nd May 2020. https://covid19.min-saude.pt/wp-content/uploads/2020/05/61_DGS_boletim_20200502_V2.pdf
26. Decreto de lei n.º 2-A/2020 – início do estado de emergência. https://www.portugal.gov.pt/download-ficheiros/ficheiro.aspx?v=3f8e87a6-3cf1-4d0c-b5ee-72225a73cd4f
27. Decreto de lei n.º 20/2020 – fim do estado de emergência. https://dre.pt/application/conteudo/132883356
28. Thygesen K, Alpert JS, Jaffe AS, et al. Fourth universal definition of myocardial infarction (2018). J Am Coll Cardiol. 2018;72:2231–64.
29. Chor-Cheung FT, Kent-Shek C, Simon L, et al. Impact of coronavirus disease 2019 (COVID-19) outbreak on ST-segment – elevation myocardial infarction care in Hong Kong, China. Circ Cardiovasc Qual Outcomes. 2020;(March). CIRCOUT-COMES120006631.
30. Matthew JD, Mauricio GC, Anthony A, et al. Reperfusion of STEMI in the COVID-19 era – business as usual? Circulation. 2020;141:1948–50.
31. Tumminello G, Barbieri L, Lucreziotti S, et al. Impact of COVID-19 on STEMI: second youth for fibrinolysis or time to centralized approach? IJC Heart Vascul. 2020;30:100600.
32. Vecchio S, Fileti L. Reggi Almpatto della pandemia COVID-19 sui ricoveri per sindrome coronarica acuta: revisione della letteratura ed esperienza monocentrica. G Ital Cardiol. 2020;21:502–8. ID: covidwho-611788 [in italian].
33. Piccolo R, Bruzzese D, Mauro C, et al. Population trends in rates of percutaneous coronary revascularization for acute coronary syndromes associated with the COVID-19 outbreak. Circulation. 2020;141:2035–7.
34. Mahmud E, Dauerman HL, Welt FGP, et al. Management of acute myocardial infarction during the COVID-19 pandemic – a consensus statement from the Society for Cardiovascular Angiography and Interventions (SCAI), the American College of Cardiology (ACC), and the American College of Emergency Physicians (ACEP). Catheter Cardiovasc Interv. 2020;1–10.