

ACUTE MYOCARDIAL INFARCTION IN OLDER ADULT PATIENTS

Infarto agudo do miocárdio em pacientes idosos

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ABSTRACT

OBJECTIVES: To assess clinical and coronary angiographic characteristics, previous medical history, and clinical course, by age group, in older adults after myocardial infarction who underwent primary percutaneous coronary intervention (pPCI). **METHODS:** Single-center, cohort study that enrolled all patients with ST-segment elevation myocardial infarction who underwent pPCI at a specialized cardiology reference center in the South of Brazil. Older adults were defined as age ≥ 60 years, as set out in Brazilian legislation. Patients in the following age groups were compared: 60 to 64 years, 65 to 69 years, 70 to 74 years, 75 to 79 years, and ≥ 80 years. Patients' clinical course was assessed at initial hospital admissions and after 2 years of clinical follow-up. Data were analyzed using SPSS 19, and significance was established at $p < 0.05$. **RESULTS:** From December 2015 to December 2018, a total of 636 patients were enrolled consecutively. Angiographic success rates were around 90% in all age groups. There were no differences in medications used, except for glycoprotein IIb/IIIa inhibitors, which were more frequently used in patients of lower age groups. Older patients had more in-hospital acute renal failure and higher in-hospital mortality. Predictors of mortality were age over 75, chronic renal failure, need for ventilatory support, severe arrhythmia, and sepsis. **CONCLUSIONS:** pPCI in older adult patients is a safe procedure with a high success rate. **KEYWORDS:** myocardial infarction; angioplasty; critical care; health services for the aged; aged.

RESUMO

OBJETIVOS: Avaliar características clínicas e angiográficas, história clínica pregressa e evolução clínica, por faixa etária, em idosos submetidos a intervenção coronária percutânea primária (ICPp) após infarto do miocárdio. **METODOLOGIA:** Estudo de coorte, de centro único, que incluiu todos os pacientes com infarto do miocárdio com supradesnívelamento do segmento ST submetidos ICPp em um centro de referência especializado em cardiologia no sul do Brasil. Os idosos foram definidos como aqueles com idade ≥ 60 anos, conforme estabelecido na legislação brasileira. Os pacientes nas seguintes faixas etárias foram comparados: 60 a 64 anos, 65 a 69 anos, 70 a 74 anos, 75 a 79 anos e ≥ 80 anos. O curso clínico dos pacientes foi avaliado nas admissões hospitalares iniciais e após 2 anos de acompanhamento clínico. Os dados foram analisados usando o SPSS 19, e $p < 0,05$ foi considerado significativo. **RESULTADOS:** De dezembro de 2015 a dezembro de 2018, 636 pacientes foram incluídos consecutivamente. As taxas de sucesso angiográfico foram de cerca de 90% em todas as faixas etárias. Não houve diferenças nos medicamentos utilizados, com exceção dos inibidores da glicoproteína IIb/IIIa, que foram mais frequentemente utilizados em pacientes em faixas etárias mais baixas. Pacientes mais velhos apresentaram mais insuficiência renal aguda intra-hospitalar e maior mortalidade hospitalar. Os preditores de mortalidade foram: idade superior a 75 anos, insuficiência renal crônica, necessidade de suporte ventilatório, arritmia grave e sepse. **CONCLUSÕES:** O ICPp em pacientes idosos é um procedimento seguro e com alta taxa de sucesso. **PALAVRAS-CHAVE:** infarto do miocárdio; angioplastia; cuidados críticos; serviços de saúde para idosos; idoso.

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INTRODUCTION

Brazil is going through a demographic transition process. The population ageing increases the rates of chronic diseases and comorbidities, increasing, as a consequence, the demand for health services and requiring the recognition of the needs of the population of this age group.¹ Cardiovascular diseases are among the main causes of morbidity and mortality worldwide, with rates in Brazil and the United States of 28% and 31%, respectively.^{2,3} Percutaneous coronary revascularization is increasingly indicated for older adults, both for stable coronary artery disease and for acute coronary syndromes.⁴

It is estimated that 60 to 65% of ST-segment elevation myocardial infarctions (STEMI) occur in patients over the age of 60.⁵ A small proportion of the participants in large clinical trials are older adults and these patients have atypical presentations, including silent or unrecognized acute myocardial infarction and left bundle branch block.⁵ Compared with the younger population, older adult patients submitted to percutaneous angioplasty present more frequently with diffuse disease, ventricular dysfunction, acute coronary syndromes, and comorbidities such as renal failure, strokes, and peripheral arterial disease, making their clinical status less favorable. These characteristics make it more difficult to assess the results of interventions in this age group.⁶⁻⁸

There are no contemporary studies conducted in Brazil that analyze the clinical and angiographic characteristics and clinical outcomes of older adults who underwent primary percutaneous coronary intervention. The objective of the present study was to record, analyze, and compare these characteristics as age increases in patients who had a recent infarction and were treated with primary percutaneous coronary intervention (pPCI) at a specialized cardiology reference center in the South of Brazil.

METHODS

Cross-sectional, single-center cohort study that enrolled all patients with ST-segment elevation myocardial infarction (STEMI) who underwent pPCI at a specialized cardiology reference center in Southern Brazil from December 2015 to December 2018. The study was approved by the institutional Research Ethics Committee and all patients signed an informed consent form. Patients were followed for 30 days, 1 year and 2 years by medical records and telephone calls to determine their health status.

Patients were considered for inclusion in the study if they were admitted via the emergency department at our institution by STEMI. Older adults were defined as ≥ 60 years, as set out in the Brazilian legislation.⁹

According to the V Brazilian Cardiology Society Guidelines on Treatment of ST-Segment Elevation Acute Myocardial Infarction,¹⁰ diagnosis of STEMI was defined as raised levels of cardiac markers (with at least one result over the 99th percentile) followed by a drop off, constituting an enzyme curve. Additionally, at least one of five criteria had to be present to confirm a diagnosis of STEMI:

- symptoms of myocardial ischemia;
- abnormalities of the ST segment/T wave or new left bundle branch block;
- development of pathological Q waves on the ECG;
- loss of viable myocardial muscle or abnormalities of segmental motion seen on an imaging examination;
- identification of an intracoronary thrombus by angiography or autopsy.

Exclusion criteria were chest pain lasting more than 12 hours or patient refusal to participate.

Primary percutaneous coronary intervention (pPCI) was performed as recommended in the literature. All patients were medicated at admission with 300 mg acetylsalicylic acid and 300 to 600 mg of clopidogrel or 180 mg of ticagrelor. Unfractionated heparin (60 to 100 U/kg) was administered before pPCI in the emergency room or the Cath Lab. Technical aspects of the procedure, such as stents characteristics and number, use of devices and use of glycoprotein IIb/IIIa inhibitors were decided by the interventional physician responsible for PCI.¹¹ Patients were interviewed during their hospital stay in order to obtain information on previous history and risk factors. Clinical evolution were obtained from the hospital system.

Criteria used to define risk factors were as follows: smoking: regular cigarette smoking; hypertension (H): prior diagnosis of hypertension and/or use of antihypertensive drugs; diabetes mellitus (DM): prior diagnosis of DM and/or use of drugs to treat diabetes, fasting Glycemia ≥ 126 mg/dL, Glycemia 2 hours after a Glucose challenge ≥ 200 mg/dL, or Glycated Hemoglobin (HbA1c) $\geq 6.5\%$; dyslipidemia (DSLIP): fasting serum cholesterol exceeding 240 mg/dL, prior diagnosis of DSLIP, and/or use of antihyperlipidemic medication; family history for coronary artery disease: AMI or sudden death among first-degree relatives.

Data were analyzed using SPSS for Windows. Significance was established at $p < 0.05$. Medians and interquartile ranges were calculated for ages. The study sample was classified into five age groups: 60 to 64; 65 to 69; 70 to 74, 75 to 79, and 80 or older. Quantitative variables were expressed as means and standard deviations and compared with one-way ANOVA or the Kruskal-Wallis test. Categorical variables

were expressed as absolute and relative frequencies and analyzed with the χ^2 test. Multiple logistic regression was used to identify predictors of mortality and Kaplan Meier curves were plotted to illustrate event-free survival by age group.

RESULTS

From December 2015 to December 2018, 1274 patients underwent primary percutaneous coronary intervention, of which 636 were over 60 years of age and were included in the present analysis. Regarding the age groups analyzed, 202 patients were aged from 60 to 64 years old, 157 patients from 65 to 69 years, 125 patients from 70 to 74 years, 81, from 75 to 79, and 71 patients were 80 years or older. With regard to clinical characteristics, we observed that younger patients were smokers more frequently and had a family history of coronary artery disease, while the oldest patients had hypertension and had left bundle branch block (LBBB) more frequently than other patients. Considering the medical history, the oldest patients had more strokes, heart failure, and chronic renal failure. There was no difference in the prevalence of previous AMI, previous percutaneous coronary interventions, or coronary artery bypass graft.

The angiographic profiles of patients were very similar in all groups, with no difference in the percentages of patients with multiarterial involvement. Older patients had coronary calcification more frequently. There was no significant difference between the study groups in culprit vessel IAM, percentage of left main disease, or bifurcations lesions. Need for an intra-aortic balloon or pacemaker did not differ according to age. Angiographic success rates were similar in all age groups. The door-to-balloon time, however, was longer among patients over the age of 80 (Table 1). It can be observed in Table 2 that medications used were also similar in all age groups, with the exception of Glycoproteins IIb/IIIa inhibitors, which were more frequently used in younger patients.

During hospital stay (Table 3), there were no significant differences in percentages of severe arrhythmia or sudden death averted, bleeding, sepsis, or need for ventilatory support. However, acute kidney injury (AKI) was more frequent among the older patients. In-hospital mortality was higher among the older patients.

The patients were followed for 328 ± 280 days. We found 6% and 8% losses in 1 and 2 years of follow-up, respectively. The overall mortality was 10.2% and higher among the patients in the 75- to 79-year-old group (25%).

Table 1 Coronary angiographic characteristics of 636 older adults by age group.

| Characteristics | 60–64 years n = 202 | 65–69 years n = 157 | 70–74 years n = 125 | 75–79 years n = 81 | ≥ 80 years n = 71 | p |
|------------------------------|------------------------|------------------------|------------------------|-----------------------|----------------------|---------|
| Triple vessel disease, n (%) | 47 (23.40) | 44 (28.60) | 32 (25.80) | 23 (28.70) | 24 (34.80) | 0.13 |
| Culprit vessel | | | | | | |
| AD, n (%) | 89 (44.90) | 61 (39.60) | 49 (40.20) | 27 (34.20) | 32 (46.40) | 0.37 |
| CX, n (%) | 19 (9.60) | 21 (13.60) | 17 (13.90) | 9 (11.40) | 4 (5.80) | |
| RC, n (%) | 85 (42.90) | 71 (46.10) | 49 (40.20) | 39 (49.40) | 33 (47.80) | |
| LCA, n (%) | 3 (1.50) | 0 | 2 (1.60) | 1 (1.30) | 0 | |
| Mammary, n (%) | - | - | - | 1 (1.30) | 0 | |
| Saphenous Bypass, n (%) | 2 (1.00) | 1 (0.60) | 5 (4.10) | 2 (2.50) | 0 | |
| LMD, n (%) | 6 (3.00) | 4 (2.60) | 10 (8.10) | 4 (5.00) | 3 (4.30) | 0.22 |
| Bifurcation, n (%) | 8 (4.00) | 6 (3.90) | 8 (6.60) | 2 (2.50) | 3 (4.40) | 0.92 |
| Calcification, n (%) | 6 (3.00) | 11 (7.10) | 9 (7.40) | 11 (13.80) | 16 (23.50) | < 0.001 |
| Vessel diameter, mm | 3.13 ± 0.44 | 3.09 ± 0.59 | 3.12 ± 0.40 | 3.12 ± 0.40 | 3.16 ± 0.60 | 0.85 |
| Length of lesion, mm | 24.85 ± 12.61 | 26.00 ± 11.90 | 24.36 ± 11.12 | 24.56 ± 11.12 | 25.68 ± 11.48 | 0.51 |
| Femoral access, n (%) | 44 (22.30) | 33 (21.40) | 34 (28.10) | 24 (30.80) | 23 (33.30) | 0.03 |
| Pre-dilatation, n (%) | 146 (75.60) | 106 (70.70) | 85 (70.20) | 53 (74.60) | 54 (77.10) | 0.92 |
| Stenting, n (%) | 188 (94.50) | 139 (92.10) | 105 (86.80) | 67 (91.80) | 63 (88.70) | 0.07 |
| Bare-metal Stent, n (%) | 115 (61.20) | 76 (55.10) | 59 (57.80) | 40 (60.60) | 35 (56.50) | 0.67 |
| Post dilatation, n (%) | 71 (36.40) | 54 (36.20) | 36 (29.80) | 27 (37.00) | 22 (31.40) | 0.43 |

Continue...

Table 1 Continuation.

| Characteristics | 60–64 years n = 202 | 65–69 years n = 157 | 70–74 years n = 125 | 75–79 years n = 81 | ≥ 80 years n = 71 | p |
|-----------------------------|------------------------|------------------------|------------------------|-----------------------|----------------------|-------|
| IAB, n (%) | 13 (6.40) | 5 (3.20) | 6 (4.80) | 2 (2.60) | 5 (7.40) | 0.78 |
| Pacemaker, n (%) | 10 (5.00) | 12 (7.80) | 9 (7.50) | 5 (6.40) | 5 (7.10) | 0.54 |
| TIMI 3 Post, n (%) | 184 (93.40) | 132 (89.80) | 109 (91.60) | 62 (87.30) | 60 (92.30) | 0.71 |
| Angiographic Success, n (%) | 183 (93.40) | 131 (89.70) | 105 (89.70) | 60 (87.00) | 60 (92.30) | 0.50 |
| Complications, n (%) | 25 (12.40) | 18 (11.50) | 19 (22.90) | 12 (14.80) | 9 (12.70) | 0.59 |
| Door-to-Balloon, minutes | 0:44 [0:33–0:72] | 0:52 [0:37–0:65] | 0:48 [0:39–0:73] | 0:40 [0:37–0:61] | 0:56 [0:43–0:87] | 0.007 |
| EF% | 51.30 ± 12.40 | 51.00 ± 15.60 | 52.70 ± 13.60 | 53.80 ± 13.10 | 49.00 ± 14.00 | 0.84 |

AD: anterior descending artery; CX: circumflex; RC: right coronary; LCA: Left main coronary artery; LMD: left main disease; IAB: intra-aortic balloon; TIMI: thrombolysis in myocardial infarction grade = myocardial perfusion; EF: left ventricle ejection fraction.

Table 2 Medications used by 636 older adults.

| Medication | 60–64 years n = 202 | 65–69 years n = 157 | 70–74 years n = 125 | 75–79 years n = 81 | ≥ 80 years n = 71 | p |
|---|------------------------|------------------------|------------------------|-----------------------|----------------------|------|
| Clopidogrel 600 mg, n (%) | 186 (93.00) | 146 (94.20) | 112 (91.10) | 73 (91.30) | 65 (95.60) | 0.88 |
| Ticagrelor 180 g, n (%) | 7 (3.50) | 6 (3.90) | 2 (1.60) | 2 (2.50) | 2 (2.90) | 0.52 |
| Heparin Bolus, n (%) | 175 (87.50) | 136 (88.30) | 104 (86.00) | 70 (89.70) | 58 (84.10) | 0.59 |
| ASA, n (%) | 198 (99.00) | 153 (98.70) | 121 (98.40) | 78 (97.50) | 67 (98.50) | 0.41 |
| Glycoprotein IIb/IIIa Inhibitors, n (%) | 36 (17.80) | 24 (15.30) | 21 (16.80) | 7 (8.60) | 5 (7.00) | 0.01 |

ASA: acetylsalicylic acid.

Table 3 Mortality and in-hospital events in 636 older adults.

| Events | 60–64 years n = 202 | 65–69 years n = 157 | 70–74 years n = 125 | 75–79 years n = 81 | ≥ 80 years n = 71 | p |
|----------------------------|------------------------|------------------------|------------------------|-----------------------|----------------------|-------|
| In hospital | | | | | | |
| Severe arrhythmia, n (%) | 15 (7.40) | 11 (7.10) | 10 (8.30) | 9 (11.30) | 6 (9.00) | 0.37 |
| Bleeding, n (%) | 3 (1.50) | 2 (1.30) | 3 (2.40) | 3 (3.80) | 0 | 0.23 |
| Sepsis, n (%) | 10 (5.00) | 4 (2.50) | 9 (7.40) | 7 (8.80) | 4 (6.00) | 0.20 |
| Ventilatory support, n (%) | 23 (11.40) | 18 (11.50) | 21 (17.10) | 12 (15.00) | 8 (11.90) | 0.41 |
| Acute kidney injury, n (%) | 13 (6.50) | 8 (5.10) | 9 (7.50) | 13 (16.30) | 6 (9.00) | 0.04 |
| Death, n (%) | 14 (6.90) | 9 (5.70) | 13 (10.70) | 14 (17.50) | 11 (16.20) | 0.001 |

Table 4 shows the independent predictors of mortality in this sample, specifically: age ≥75 years, female, chronic renal failure, need for ventilatory support, and occurrence of severe arrhythmia and sepsis on admission. Chronic kidney failure (CKF) and severe arrhythmia led to a four-fold increase in the risk of death and sepsis increased risk by 13 times.

In Figure 1, it can be observed that although mortality was higher among patients over 75, event-free survival was still around 80% during late clinical follow-up.

DISCUSSION

In this report, we analyzed a consecutive cohort of older adult patients with STEMI submitted to pPCI in a high-volume, tertiary cardiology center. The main findings were that older adults had similar rates of success in the pPCI procedure when compared to younger individuals, but the older patients had acute kidney injury more frequently and had higher mortality in the follow-up. We believe that this is an important piece of information, because it confirms that older STEMI patients have a higher risk

of adverse clinical events, but it shows that contemporary pPCI treatment has similar results in these patients when compared to younger individuals.

Within this perspective, a sub-analysis from 5475 STEMI patients in the APEX TRIAL study¹² showed that patients over 75 years had less angiographic success than their younger counterparts. Although this analysis was derived from a large sample, its results may not be representative of the current daily practice, as it was published in 2011. Since then, several significant developments have occurred in the field, such as newer and more potent antiplatelet drugs, the widespread

adoption of the radial approach,¹³ stent implantation techniques and improvements in drug-eluting stents technology.¹⁴ Recently, in a sub-study of the AIDA STEMI TRIAL,¹⁵ there were no differences in angiographic success between patients with multivessel coronary disease vs. single-vessel disease in both groups. Thus, technical and personnel improvement in these 10 years may contribute to better angiographic success of older patients.

The AIDA STEMI study¹⁵ also demonstrated that patients over 75 who underwent primary percutaneous coronary intervention had the same frequency of renal failure as those who did not undergo pPCI. Thus, we can consider that this complication is probably not only due to contrast-induced nephropathy (CIN). Ischemia of renal tissues and reduced glomerular filtration may be the primary pathophysiologic components, and hydration and reduced contrast volumes are considered the most effective methods for prevention of CIN.¹⁶ Regarding in-hospital evolution, older patients had more AKI, which increased mortality. However, when AKI was added to the multivariate model, the 1-year increase in mortality was not significant.¹⁷

In this study the multivariate analysis showed that besides age ≥ 75 years, the independent predictors of mortality also included chronic renal failure, severe arrhythmia, sepsis, and need for ventilatory support. We emphasize that these predictors are related to prior histories of patients, since, for example, the most common causes of arrhythmia are coronary disease, valve disease, and heart failure. Considering sepsis, in a previous study, we demonstrated that among the 3.9% of patients who had infections, 2% were community acquired — i.e., contracted before hospital admission.¹⁸

Door-to-balloon time was longer for patients over 80 years, but within the limit set out in the guidelines. It should be pointed out that the care teams and operation teams had the same technical training, so this longer time may be due to anatomic conditions associated with more difficult procedures, such as calcification, more diffuse disease, and tortuosity.

Similarly, there was no difference in relation to medications used, with the exception of glycoprotein IIb/IIIa inhibitors, which were less frequently used among older patients. However, as previously reported, the use of IIb/IIIa inhibitors increases the rates of hemorrhagic complications. Nevertheless, older adults present a slightly elevated risk of these complications.¹⁹ Therefore, the potential benefits of treating this population should be considered.

Our study has certain limitations that merit mention. Although our institution performs a very high volume of

Table 4 Multiple logistic regression for predictors of mortality.

| Characteristics | Odds | 95%CI | p |
|--------------------------|-------|------------|---------|
| Age ≥ 75 | 4.47 | 1.19–16.74 | 0.03 |
| Male | 0.36 | 0.14–0.96 | 0.04 |
| CHF | 2.44 | 0.79–7.61 | 0.12 |
| CKF | 4.74 | 1.22–18.30 | 0.02 |
| Killip 3 or 4 | 2.59 | 0.61–10.96 | 0.20 |
| Ventilatory support (MV) | 3.64 | 1.07–12.37 | 0.04 |
| Arrhythmia severe | 4.64 | 1.26–17.03 | < 0.001 |
| Bleeding | 1.43 | 0.16–12.49 | 0.74 |
| Sepsis | 13.26 | 1.32–12.45 | < 0.001 |

CHF: congestive heart failure; CKF: chronic kidney failure; MV: mechanical ventilation; 95%CI: confidence interval of 95%.

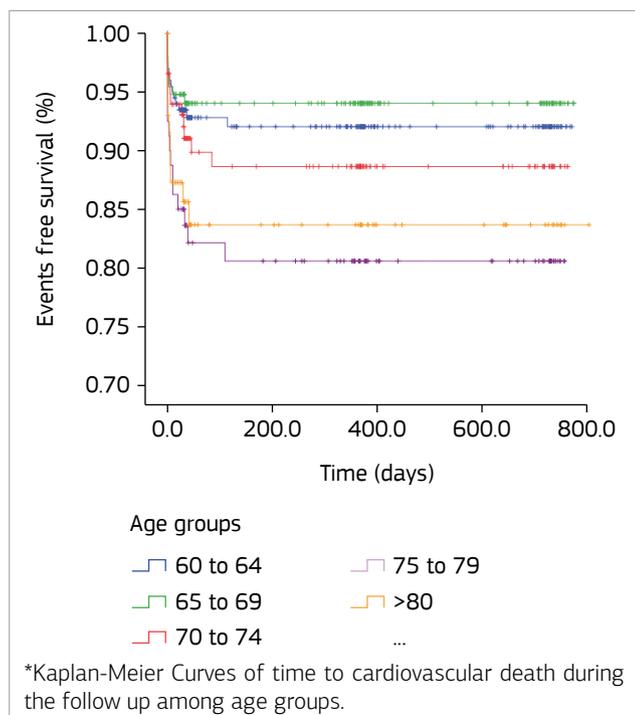


Figure 1 Survival curves by age group*.

primary percutaneous coronary intervention procedures, this is still a single-center study that enrolled a modest number of patients. Patients' prior histories were taken by interviewing the patients themselves, so there may be information bias. Information on late clinical follow-up was collected both by electronic medical records and telephone interviews with patients and family members, which could also generate information bias.

CONCLUSIONS

In conclusion, we consider that primary percutaneous coronary intervention in patients over the age of 60 is a safe procedure providing high success rates. The independent predictors of mortality in this population were chronic

kidney injury, need for ventilatory support, sepsis, and severe arrhythmia while still in hospital.

CONFLICT OF INTERESTS

There are no conflicts of interests to declare.

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