Demographic description and outcomes of a metropolitan network for myocardial infarction treatment

Descripción demográfica y desenlaces de una red metropolitana de atención para el infarto agudo de miocardio

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Abstract

Objective: The objective of the study was to describe the myocardial infarction treatment network and compare in-hospital mortality in patients undergoing either primary angioplasty or pharmacoinvasive strategy in Mexico City and a broad metropolitan area. Methods: Cohort study including patients with ST-elevation myocardial infarction. We recorded demographic and clinical data, laboratory tests and in-hospital mortality in patients that underwent primary angioplasty and pharmacoinvasive strategy. Kaplan-Meier analysis was used to assess mortality and Cox-regression assessed mortality risk factors. Results: Three hundred forty patients from a network of 60 hospitals and 9 states were analyzed. Of the total population, 166 were treated with pharmacoinvasive strategy and 174 with primary angioplasty. Door to thrombolytic time was 54 min and door to wire crossing time was 72.5 min; no differences in total ischemia time were demonstrated. No differences for in-hospital mortality (6.3% vs. 5.4%, p = 0.49) were found when comparing pharmacoinvasive and primary angioplasty groups. The main predictors for in-hospital mortality were: glucose > 180 mg/dl (HR 3.73), total ischemia time > 420 min (HR 3.18), heart rate > 90 bpm (HR 5.46), Killip and Kimball > II (HR 11.03), and left ventricle ejection fraction < 40% (HR 3.21). Conclusions: This myocardial infarction network covers a large area and constitutes one of the biggest in the world. There were no differences regarding in-hospital mortality between pharmacoinvasive strategy and primary angioplasty. Pharmacoinvasive strategy is an effective and safe option for prompt reperfusion in Mexico.

Key words: Myocardial infarction. Thrombolytic therapy. Angioplasty. Mortality.

Resumen

Objetivo: Describir la red de atención de infarto agudo de miocardio y comparar los desenlaces intrahospitalarios en pacientes tratados con angioplastia coronaria o estrategia farmacoinvasiva en la Ciudad de México y su área metropolitana. Métodos: Estudio de cohorte que incluyó pacientes con infarto agudo de miocardio con elevación del segmento ST. Se recabaron datos demográficos y clínicos, así como estudios de laboratorio y mortalidad intrahospitalaria en los pacientes que fueron tratados...
con angioplastía coronaria o estrategia farmacoinvasiva. Se realizó un análisis de Kaplan-Meier para describir la mortalidad y un modelo de regresión de Cox para evaluar los factores asociados a mortalidad. Resultados: Se analizaron 340 pacientes provenientes de una red compuesta por 60 hospitales. Del total de la población, 166 fueron tratados con estrategia farmacoinvasiva y 174 con angioplastía primaria. El tiempo puerta-aguja fue 54 min. y el tiempo puerta-dispositivo de 72.5 min.; no se encontraron diferencias en el tiempo total de isquemia. Además, no existieron diferencias en la mortalidad intrahospitalaria (6.3% vs. 5.4%, p = 0.49) al comparar la estrategia farmacoinvasiva y la angioplastía primaria. Los principales predictores de mortalidad intrahospitalaria fueron: glucosa > 180 mg/dl (HR 3.73), tiempo total de isquemia > 420 min. (HR 3.18), frecuencia cardiaca > 90 lpm (HR 5.46), Killip and Kimball > II (HR 11.03) y fracción de eyecación < 40% (HR 3.21).

Conclusiones: En esta red de atención al infarto agudo de miocardio no se encontraron diferencias en la mortalidad intrahospitalaria entre la estrategia farmacoinvasiva y la angioplastia primaria. La estrategia farmacoinvasiva puede ser una alternativa efectiva y segura para lograr reperfusión adecuada en México.

Palabras clave: Infarto de miocardio. Terapia trombolítica. Angioplastia. Mortalidad.

Introduction

Optimal timing for reperfusion in ST-segment elevation myocardial infarction (STEMI) reduces infarct size, prevents, and delays ventricular remodeling and increases survival. In spite of this, a big proportion of patients, especially in low-income countries do not receive reperfusion promptly or receive it lately.

Clinical practice guidelines recommend two main strategies for reperfusion in STEMI: primary percutaneous coronary intervention (PPCI) and pharmacoinvasive strategy (PS). PPCI consists in an urgent recanalization of the obstruction in the lumen of the affected coronary artery by means of a balloon or stent, without previous administration of a fibrinolytic agent. It has been demonstrated that coronary flow can return to normal in 90% of the cases, while for fibrinolysis flow is only restored in about 50-60%. The success of reperfusion depends on many factors such as the time of onset of symptoms, the ability and experience of the operator and resource availability for the procedure.

There is a problem with the availability of PCI centers in many parts of the world, as it has been stated in other studies. To overcome these difficulties, PS consists in the administration of a fibrinolytic agent at first moment after diagnosis of STEMI, followed by a coronary intervention in the next 3-24 h. This practice has reduced reinfarction and recurrent ischemia compared to medical treatment alone. The STREAM trial demonstrated that the PS, together with contemporary antithrombotic therapy (clopidogrel, aspirin, and enoxaparin), has the same efficacy and safety than PPCI. Other trials stand out the ability to overcome the social and geographic limitations.

Mexico City and its metropolitan area constitute one of the most populated urban areas in the world. The peculiar social and economic problematics of our country, together with a high population density, makes it difficult to treat STEMI in optimal timing so to accomplish guidelines goals represents a big challenge. Therefore, the means of this study were to compare in-hospital mortality in patients taken to PPCI against those taken to PS in Mexico City and a large metropolitan area with the goal to establish the efficacy of PS in a real life setting.

Methods

A cohort from the PHASE-MX trial was taken, which included all patients from both genders, between 18 and 80 years of age, with a diagnosis of STEMI, that were admitted to the Emergency Department and Coronary Care Unit of the Instituto Nacional de Cardiología, from April 1, 2018, to March 31, 2019. The myocardial infarction definition used in this study was the one proposed by the European Society of Cardiology and Thygesen et al.

At admission, the following data were collected: age, gender, date of admission, the presence of diabetes mellitus, systemic arterial hypertension, smoking, chronic kidney disease, obesity, previous history of myocardial infarction, previous revascularization, vital signs, TIMI, GRACE, and CRUSADE scores, blood biomarkers, blood glucose, troponin, NT proBNP, total ischemic time, first medical contact time, door-to-needle time, door-to-wire crossing or device time, medical treatment before reperfusion, time to PS, and treatment success. In this study, all the patients from the PS group received fibrinolysis at their first medical contact center and angiography was done at Instituto Nacional de Cardiología, both decisions were made according to the medical staff’s discretion. Afterward, we made an in-hospital follow-up where mortality and date of home discharge were registered.
Statistical analysis: all tests were done in STATA v13 (StataCorp LP, College Station, Tx). Quantitative variables were analyzed with descriptive methods depending on their distribution, corroborated by the Shapiro–Wilk test. Variables with a normal distribution were described with mean value and standard deviation. Otherwise, median and interquartile ranges were used. Taking into consideration the normality of each quantitative variable, an analysis with Student’s t and U Mann–Whitney tests was performed. Qualitative variables were described through frequencies and percentages, while for the bivariate analysis \( \chi^2 \) or Fisher’s test were performed depending on the number of events. For the survival analysis, tables and Kaplan–Meier curves were made to describe mortality in both groups. Differences between survival times for both treatment groups were compared with log rank test. Cox regression models, adjusted by sex and age, were built to determine the main predictors of in-hospital mortality in patients treated with both strategies. A \( p < 0.05 \) was considered as statistically significant.

Furthermore, for the descriptive analysis, we created a map of the hospitals that are part of our STEMI network. All hospitals were geocoded by finding their latitude and longitude using Google Maps. The coordinates were recorded in a separate datasheet. We used QGIS 3.10 (2019, QGIS Geographic Information System. Open Source Geospatial Foundation. URL: http://qgis.org) to create our maps. From the University of California at Berkeley library for GeoData (https://geodata.lib.berkeley.edu/), we downloaded polygon shapefiles of the states and municipalities where the hospitals are located and added the hospitals (points) as another layer. Finally, we showed our center with a different mark to highlight the distances between the different hospitals in the network and our center.

### Results

A total of 340 patients were included, 166 for PS and 174 for PPCI. The mean age was 59 ± 10.8 years, 87.1% were male and 12.9% were female. Regarding medical history, 35% had diabetes mellitus, 46.8% hypertension, 17.1% dislipidemia, 46.2% were current smokers, and 9.7% had a previous myocardial infarction (Table 1).

As to the place of residency, 55.29% came from Mexico City, 29.41% came from Estado de México, and 5.29% came from Morelos. We received patients from 60 different hospitals, with a mean distance of 25.2 km, the smallest distance was 1.3 km and the longest was 312 km. The mean estimated transfer time was 53 min, being the minimum and maximum of 8 and 263 min, respectively. Moreover, 23.35% of the patients had their first medical contact at the Instituto Nacional de Cardiología and hospitals who referred the majority of patients to our center were Hospital General “Dr. Manuel Gea González,” Hospital General Balbuena, Hospital General “La Perla”, Hospital General de Cuernavaca, and Cruz Roja Mexicana (Table 2). Figures 1 and 2 show the geographic distribution of the PS network in the Instituto Nacional de Cardiología.

Regarding the clinical characteristics at admission, no overt differences were found between PS and PPCI groups—even though PPCI presented significantly higher blood pressure levels, these were not clinically relevant. Furthermore, a greater proportion of patients in the PPCI group presented a better Killip–Kimball score compared to the PS group (\( p < 0.001 \) (Table 3). Likewise, as shown in table 4, laboratory profiles between both groups were similar and most of the tests were among reference values, still some differences were found. For instance, C reactive protein was higher among PPCI patients, while cardiac dysfunction (NT-PROBNP) and damage (troponin I) markers were significantly higher among PS patients (\( p < 0.001 \)).

In relation to the time of first medical contact, it was 120 min (IQR: 60-225) for PS and 150 min for PPCI (IQR: 60-270), without significant differences (\( p = 0.11 \)). Moreover, the mean total ischemic time was 347.5 min (IQR: 200-600) for PS versus 310 min (IQR: 205-557) for PPCI (\( p = 0.52 \)). Furthermore, for PS patients the mean door-to-needle time was 54 min (IQR: 30-103), and the mean time for pharmacoinvasion was 1440 min (IQR: 600-2880), while among PPCI patients the time to door-to-device was 72.5 min (IQR: 60-95). Finally, hospital stay was similar with both strategies, with a mean time of 6 days, and an interquartile range of 3-9 days.

With regard to in-hospital mortality, a total of 20 patients died during the follow-up – 11 patients (6.3%) who underwent PCI and nine (5.4%) who underwent PS (\( p = 0.82 \)). A further subanalysis was made according to the time taken to pharmacoinvasion where the cutoff point was set according to the median value of 1440 min, without differences in mortality (6.56 vs. 5.73 %, \( p = 0.49 \)). Moreover, 96% of patients from this cohort survived after 6 days of follow-up (Fig. 3). As to the type of post-infarction treatment, mean survival after 6 days was essentially the same between PPCI and PS patients (95% vs. 96%; \( p = 0.54 \)) as shown in the Kaplan–Meier curves from figure 4.
Table 1. Demographic characteristics of patients with STEMI

| Variable                        | Total (n = 340) | PS (n = 166) | PPCI (n = 174) | p   |
|---------------------------------|-----------------|--------------|---------------|-----|
|                                 | n   | %   | n   | %   | n   | %   |     |
| Male                            | 296 | 87.1| 148 | 89.2| 148 | 85.1| 0.26|
| Female                          | 44  | 12.9| 18  | 10.8| 26  | 14.9|     |
| Diabetes                        | 119 | 35  | 58  | 34.9| 61  | 35.1| 0.98|
| Hypertension                    | 159 | 46.8| 72  | 43.4| 87  | 50  | 0.22|
| Dyslipidemia                    | 58  | 17.1| 21  | 12.7| 37  | 21.3| 0.03|
| Current smoking                 | 157 | 46.2| 88  | 53  | 69  | 39.7| 0.01|
| Previous smoking                | 56  | 16.5| 23  | 13.9| 33  | 18.97| 0.2 |
| Chronic kidney disease          | 7   | 2.1 | 4   | 2.4 | 3   | 1.72| 0.47|
| Obesity                         | 77  | 22.7| 35  | 21.1| 42  | 24.1| 0.50|
| Previous myocardial infarction  | 33  | 9.7 | 14  | 8.4 | 19  | 10.9| 0.43|
| Previous PCI                    | 23  | 6.8 | 7   | 4.3 | 16  | 9.2 | 0.05|
| Previous CABG                   | 5   | 1.5 | 1   | 0.6 | 4   | 2.3 | 0.20|
| Heart failure                   | 3   | 0.9 | 0   | 0   | 3   | 1.7 | 0.08|
| Valvular heart disease          | 2   | 0.6 | 0   | 0   | 2   | 1.2 | 0.26|
| Atrial fibrillation             | 1   | 0.3 | 0   | 0   | 1   | 0.6 | 0.32|
|                                 | n   | Mean ± SD | n   | Mean ± SD | n   | Mean ± SD | p   |
| Age (years)                     | 340 | 59 ± 10.8 | 166 | 58.5 ± 10.9 | 60 | 61 | 0.08 |

PS: pharmacoinvasive strategy; PPCI: primary percutaneous coronary intervention; CABG: coronary artery bypass grafting; SD: standard deviation.

Figure 1. Regional map of places of origin of patients with STEMI.
### Table 2. Description of the place of first medical contact in patients with STEMI

| Place of first medical contact                          | n | %     | Distance to NHI (km) | Estimated time to NHI (hh:mm) |
|----------------------------------------------------------|---|--------|----------------------|-------------------------------|
| Instituto Nacional de Cardiología                        | 76| 22.35  | -                    | -                            |
| Médico particular                                        | 55| 16.17  | -                    | -                            |
| Hospital General “Dr. Manuel Gea González”               | 42| 12.35  | 1.3                  | 0:08                         |
| Hospital General Balbuena                               | 18| 5.29   | 19.7                 | 0:41                         |
| Hospital General “La Perla”                             | 17| 5      | 25.1                 | 0:53                         |
| Hospital General de Cuernavaca “Dr. José G. Parres”      | 13| 3.82   | 67.7                 | 0:55                         |
| Cruz Roja de México                                      | 9 | 2.65   | 21.9                 | 1:40                         |
| Hospital General “La Villa”                             | 9 | 2.65   | 28.3                 | 0:55                         |
| Instituto Nacional de Ciencias Médicas y Nutrición       | 6 | 1.76   | 0.85                 | 0:06                         |
| Hospital de Especialidades “Dr. Belisario Domínguez”     | 6 | 1.76   | 12                   | 0:50                         |
| Instituto Nacional de Enfermedades Respiratorias         | 5 | 1.47   | 1.4                  | 0:08                         |
| Hospital General de Texcoco “Guadalupe Victoria”        | 5 | 1.47   | 46.5                 | 1:06                         |
| Hospital General de Tulancingo                           | 4 | 1.18   | 131                  | 2:07                         |
| Hospital General “Dr. Enrique Cabrera”                   | 4 | 1.18   | 20.8                 | 0:39                         |
| Hospital General de Naucalpan “Dr. Maximiliano Ruiz Castañeda” | 4 | 1.18   | 28.8                 | 1:10                         |
| Centro Médico de Toluca “Adolfo López Mateos”            | 4 | 1.17   | 67.6                 | 1:12                         |
| Hospital General “Dr. Gustavo Baz Prada”                 | 3 | 0.88   | 25.3                 | 0:53                         |
| Hospital General de Cuautitlán “José Vicente Villada”    | 3 | 0.88   | 56.2                 | 1:22                         |
| Hospital ISSEMYM Tlalnepantla                            | 3 | 0.88   | 39.6                 | 1:10                         |
| Hospital General “Dr. Nicolás San Juan”                  | 2 | 0.59   | 68.1                 | 1:13                         |
| Clínica 25, IMSS                                         | 2 | 0.59   | 22.4                 | 0:44                         |
| Hospital General de Huichapan                            | 2 | 0.59   | 183                  | 2:41                         |
| Hospital General de Chilpancingo “Dr. Raymundo Abarca Alarcón” | 2 | 0.59   | 251                  | 2:36                         |
| Clínica Médica Mardán                                     | 2 | 0.59   | 16.6                 | 0:36                         |
| Hospital General de Jilotepec                            | 2 | 0.59   | 114                  | 2:07                         |
| Hospital General Gregorio Salas                          | 2 | 0.59   | 19.2                 | 0:45                         |
| Hospital General de México                               | 2 | 0.59   | 17.3                 | 0:36                         |
| Hospital General del Valle del Mezquital                  | 2 | 0.59   | 168                  | 2:50                         |
| Hospital General de Tláhuac “Dr. Miguel Lima Ramírez”    | 2 | 0.59   | 120                  | 1:55                         |
| Hospital General de Taxco                                 | 2 | 0.59   | 161                  | 2:06                         |
| Hospital General de Valle de Bravo                       | 2 | 0.59   | 144                  | 2:01                         |
| Hospital General de Tlaxcala                              | 1 | 0.29   | 120                  | 1:55                         |
| Hospital General de Milpa Alta                           | 1 | 0.29   | 22.6                 | 0:54                         |
| Instituto Nacional de Cancerología                        | 1 | 0.29   | 1.2                  | 0:06                         |
| Hospital General de Cuautla “Dr. Mauro Belauzarán Tapia”  | 1 | 0.29   | 90.2                 | 1:16                         |

(Continues)
Finally, variables associated with in-hospital mortality for both strategies were assessed through a Cox regression model, wherein blood glucose > 180 mg/dl (HR 3.73, IC 95% 1.02-13.56), total ischemic time > 420 min (HR 3.18, IC 95% 1.01-10.2), heart rate > 90 bpm (HR 5.46, IC 95% 1.69-17.59), Killip and Kimball > II (HR 11.03, IC 95% 1.42-85.15), and left ventricular ejection fraction < 40% (HR 3.21, IC 95% 1.03-10.01) determined a greater mortality risk in the whole cohort (Table 5).

### Discussion

Optimal treatment for STEMI is timely reperfusion. The European Society of Cardiology (ESC) proposes that PCI is the treatment of choice; however, if the catheterization laboratory is beyond 2 h, the recommended treatment is PS\(^2\). Unfortunately, Mexico has many limitations, not only in urban infrastructure but also in social and economic development which keeps us away from achieving the proper timing for PCI. As an example of this, we received patients that came from 9

| Place of first medical contact                                      | n  | %  | Distance to NHI (km) | Estimated time to NHI (hh:mm) |
|---------------------------------------------------------------------|----|----|----------------------|------------------------------|
| Hospital General de Huitzuco                                       | 1  | 0.29 | 180                  | 2:07                         |
| Hospital General de Jojutla “Dr. Ernesto Meana San Román”          | 1  | 0.29 | 128                  | 1:37                         |
| Hospital Escandón                                                  | 1  | 0.29 | 21.2                 | 0:38                         |
| Hospital General de Ixtapan de la Sal                              | 1  | 0.29 | 122                  | 1:46                         |
| Hospital Regional “1° de Octubre”                                  | 1  | 0.29 | 24.9                 | 1:02                         |
| Hospital General de Tenancingo ”Miguel Hidalgo y Costilla”         | 1  | 0.29 | 96.3                 | 1:49                         |
| Hospital General “Darío Fernández Fierro”, ISSSTE                  | 1  | 0.29 | 17.5                 | 0:30                         |
| Clínica Materno Infantil Sagrada Familia                           | 1  | 0.29 | 30                   | 0:55                         |
| Hospital General del Altiplano                                    | 1  | 0.29 | 104                  | 2:10                         |
| Instituto Nacional de Neurología y Neurocirugía                    | 1  | 0.29 | 6.8                  | 0:16                         |
| Hospital General de San Felipe del Progreso                        | 1  | 0.29 | 147                  | 2:05                         |
| Hospital General de Ticomán                                       | 1  | 0.29 | 29.1                 | 1:20                         |
| Hospital General de Zona #8                                       | 1  | 0.29 | 11.3                 | 0:30                         |
| Hospital General de Atizapán                                      | 1  | 0.29 | 40.5                 | 1:06                         |
| Hospital General “Ajusco Medio”                                   | 1  | 0.29 | 8.9                  | 0:29                         |
| Hospital General de Las Américas                                  | 1  | 0.29 | 43.1                 | 1:08                         |
| Hospital General de Temixco                                       | 1  | 0.29 | 87.6                 | 1:09                         |
| Hospital General “Rubén Leñero”                                   | 1  | 0.29 | 28.8                 | 0:48                         |
| Hospital General de Querétaro                                     | 1  | 0.29 | 231                  | 2:59                         |
| Hospital General de Tacuba, ISSSTE                                  | 1  | 0.29 | 27.7                 | 0:49                         |
| Hospital Municipal de Temascaltepec                                | 1  | 0.29 | 130                  | 2:13                         |
| Hospital General de Coatepec                                       | 1  | 0.29 | 312                  | 4:23                         |
| Hospital General de Pachuca                                       | 1  | 0.29 | 114                  | 2:08                         |
| Hospital General de Ecatepec                                       | 1  | 0.29 | 34.9                 | 0:57                         |
| Hospital Ángeles del Pedregal                                     | 1  | 0.29 | 14.9                 | 0:25                         |
| Hospital Durango                                                   | 1  | 0.29 | 25                   | 0:43                         |
Table 3. Clinical characteristics in patients with STEMI

| Variable          | Total (n = 340) | PS (n = 166) | PPCI (n = 174) | p   |
|-------------------|-----------------|--------------|----------------|-----|
| Killip-Kimball I  | 181 (54.4)      | 74 (45.1)    | 108 (63.5)     | 0.00|
| Killip-Kimball II | 132 (39.6)      | 78 (47.6)    | 54 (31.8)      |     |
| Killip-Kimball III| 10 (3)          | 5 (3.1)      | 5 (2.9)        |     |
| Killip-Kimball IV | 10 (3)          | 4.3          | 3 (1.8)        |     |

| Variable          | Median (IQR)    | Median (IQR)  | Median (IQR)  | p   |
|-------------------|-----------------|---------------|---------------|-----|
| Heart rate (bpm)  | 75.5 (68.5-90)  | 75.5 (70-90)  | 75.5 (68-90)  | 0.59|
| Respiratory rate (bpm) | 18 (16-19)  | 18 (16-19)    | 18 (16-20)    | 0.87|
| Systolic pressure (mmHg) | 127 (114-147) | 126 (112-140)| 130 (117-150)| 0.01|
| Diastolic pressure (mmHg) | 80 (70-90)   | 76.5 (70-86) | 80 (70-93)    | 0.00|
| Pulse oximetry (%) | 92 (90-95)     | 92 (90-95)    | 92 (90-95)    | 0.94|
| TIMI score        | 4 (2-5)         | 4 (2-5)       | 3.5 (2-5)     | 0.36|
| GRACE score       | 125 (101-150)   | 126 (106-153)| 123 (99-147)  | 0.19|
| CRUSADE score     | 26 (18-35)      | 27 (19-35)    | 26 (18-37)    | 0.72|

PS: pharmacoinvasive strategy; PPCI: primary percutaneous coronary intervention; CABG: coronary artery bypass grafting; SD: standard deviation; IQR: interquartile range.

Figure 2. Regional map with a close-up to Mexico City and its metropolitan area indicating the places of first medical contact in patients with STEMI.
### Table 4. Laboratory tests in patients with STEMI

|                        | Total (n = 340) | PS (n = 166) | PPCI (n = 174) | p  |
|------------------------|----------------|--------------|---------------|----|
| Hemoglobin (g/L)       | 15.6 (14.4-16.7) | 15.35 (14.4-16.3) | 15.8 (14.5-16.9) | 0.08 |
| Creatinine (mg/dL)     | 1 (0.8-1.2)     | 1 (0.8-1.2)   | 1 (0.8-1.1)   | 0.99 |
| Urea nitrogen (mg/dL)  | 17 (14-23)      | 18.1 (15-25)  | 16.6 (14-21)  | 0.04 |
| Na (mEq/L)             | 136 (134-138)   | 136 (134-138) | 136 (134-137) | 0.30 |
| C reactive protein (mg/L) | 6.9 (2.7-28.7) | 13.06 (4.26-46) | 4.5 (2-19) | 0.00 |
| Leukocytes (10^3/µL)   | 11.7 (9.3-14.4) | 11.4 (9.3-14.7) | 11.9 (9.2-14.2) | 0.82 |
| NT-PROBNP (pg/mL)      | 793.5 (222.5-3284.5) | 1445 (421-3643) | 389 (100.5-2644) | 0.00 |
| Troponin I (ng/mL)     | 12.7 (0.9-52.8) | 35 (12-80)    | 1.85 (0.3-14.9) | 0.00 |
| Maximum Troponin I (ng/mL) | 64 (24-80)     | 67.9 (23.8-80) | 62.9 (26-80) | 0.46 |
| Glucose (mg/dL)        | 162.5 (1278-238.5) | 150 (115-230) | 174 (136-246.8) | 0.00 |
| K (mEq/L)              | 4.1 (3.8-4.4)   | 4.1 (3.86-4.5) | 4.1 (3.8-4.4) | 0.07 |
| Cl (mEq/L)             | 103 (100-105.52) | 103 (101-107) | 103 (100-105) | 0.30 |
| Glycated hemoglobin (%)| 6.1 (5.65-8.2)  | 6.1 (5.6-7.6) | 6.1 (5.7-8.3) | 0.41 |
| Albumin (g/dL)         | 3.6 (3.3-3.9)   | 3.6 (3.4-4)   | 3.7 (3.4-3.9) | 0.88 |
| Uric acid (mg/dL)      | 6.7 (5.6-7.86)  | 6.8 (5.8-8.1) | 6.4 (5.4-7.8) | 0.03 |
| Platelets (10^3/µL)    | 217 (183-259)   | 209.5 (177-257) | 221.5 (192-263) | 0.10 |
| Cholesterol (mg/dL)    | 154.9 (130-188.9) | 153 (129.5-186) | 157 (131-189) | 0.36 |
| LDL Cholesterol (mg/dL)| 98.3 (75-121.8) | 99.9 (74.4-121) | 97.2 (75.8-122.6) | 0.72 |
| HDL Cholesterol (mg/dL)| 34.4 (29.7-40)  | 34.3 (29.2-40.3) | 34.5 (30.7-40) | 0.33 |
| STH (mIU/L)            | 1.4 (0.7-2.8)   | 1.35 (0.9-2.8) | 1.4 (0.7-2.9) | 0.88 |

PS: pharmacoinvasive strategy; PPCI: primary percutaneous coronary intervention; SD: standard deviation; IQR: interquartile range; LDL: low density lipoprotein; HDL: high density lipoprotein; STH: stimulant thyroid hormone.

**Figure 3.** General survival of patients with STEMI.
states and 60 different health facilities, being the most distant the Coatepec General Hospital (312 km and a mean estimated transfer time of 4 h and 23 min); moreover, it must be acknowledge that this network covers a big area with a median radius of 25.2 km. This pharmacoinvasive network covers a big area of the Mexican territory and constitutes one of the biggest in the world.

There was a higher proportion of men, diabetes mellitus, hypertension, and active smoking that seemed to surpass the national statistics, and it is striking that there was a lower prevalence of obesity compared to Mexico’s National Health and Nutrition Survey (ENSA-NUT in Spanish) 2012\textsuperscript{25}.

The time of the first medical contact was higher for PCI than for PS, however, there were no differences among the total ischemic time even though there was a slight increase in the latter one probably explained by the time of transfer.

The door-to-needle time was found 5 times higher than the time proposed by the ESC guidelines, which emphasizes the fact that training of the medical staff is necessary, because the lower the total ischemic time, the higher the proportion of rescued myocardial tissue\textsuperscript{2}.

On the other side, the door-to-device time was 72.5 min, which goes hand in hand with the established international guidelines. Furthermore, the time to pharmacoinvasion calculated in our study was 1440 min, equivalent to 1 day, the same time according to the recommendations of the ESC guidelines\textsuperscript{2,26}.

### Table 5. Cox regression model for prediction of in-hospital mortality in STEMI

| Variable                        | HR   | SE  | p     | 95% CI          |
|---------------------------------|------|-----|-------|-----------------|
| Male gender                     | 2.03 | 2.11| 0.49  | 0.26-15.55      |
| Diabetes                        | 1.75 | 0.94| 0.29  | 0.61-5.02       |
| Hypertension                    | 1.03 | 0.55| 0.95  | 0.36-2.94       |
| Chronic kidney disease          | 4.26 | 4.48| 0.16  | 0.54-33.4       |
| CRP > 5 mg/L                    | 1.57 | 0.93| 0.44  | 0.49-5.02       |
| Glucose >180 mg/dl              | 3.73 | 2.45| 0.04  | 0.18-13.55      |
| Total ischemia time > 420 min   | 3.18 | 1.89| 0.04  | 1.01-10.20      |
| First medical contact > 50 min  | 1.28 | 0.98| 0.74  | 0.28-5.75       |
| Heart rate > 90 bpm             | 5.46 | 3.26| 0.04  | 1.69-17.59      |
| Systolic pressure < 90 mmHg     | 6.75 | 7.12| 0.07  | 0.85-53.41      |
| Killip & Kimball > II           | 11.03| 11.5 |0.02  | 1.42-85.15      |
| GRACE score > 140               | 3.04 | 1.84| 0.06  | 0.93-9.98       |
| LVEF < 40%                      | 3.21 | 1.86| 0.04  | 1.03-10.01      |

CRP: C reactive protein; LVEF: left ventricular ejection fraction; HR: hazard ratio; SE: standard error; 95% CI: 95% confidence interval.

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Figure 4. Survival according to the reperfusion strategy in patients with STEMI. PS: pharmacoinvasive strategy; PPCI: primary percutaneous coronary intervention.
It is important to acknowledge that our patients have longer hospital stays than usual, approximately 6 days, independently of the strategy. The ESC STEMI guidelines recommend a rapid home discharge, especially between the next 48-72 h, in low risk infarctions and when the assurance of a rehabilitation program and follow-up is reliable².

Within the most relevant data of this study is the in-hospital mortality, represented by 11 patients taken to PCI and 9 patients taken to PS, 6.3% and 5.4% (p = 0.82), respectively. The percentage of survival after STEMI was 94.1%, similar to what Sierra-Fragoso et al. reported in their study, where authors reported in-hospital mortality of 5.1% in PS and 5.3% in PCI²². On the other hand, the RENASCA registry reported a cardiovascular mortality of 14.9% higher than what we described²⁷. Meanwhile, in other international registries, similar mortality has been reported among both strategies with a follow-up to 1 year after the index event, demonstrating that PS is a safe and effective method¹⁹,¹⁵,¹⁸,²⁸,⁰.

Finally, the data obtained by the Cox regression model for predictors associated with in-hospital mortality were blood glucose >180 mg/dl, total ischemic time > 420 min, heart rate > 90 bpm, Killip-Kimball > II, and left ventricular ejection faction < 40%. This is relevant since we do not have tools to predict in-hospital mortality in our population.

Epidemiological transition has left its mark. Nowadays chronic diseases have a high prevalence and most of them will develop an acute myocardial infarction as a final outcome. Mexico has up to 3 times more in-hospital mortality than the rest of the countries belonging to the Organization for Economic Cooperation and Development (OECD) due to the lack of strategies that does not allow optimal access to medical services and proper timing for treatment. In the national context, the PS headed by the National Heart Institute has proven to be equally effective than PCI at least for in-hospital mortality, which by now will improve cardiovascular outcomes in the future. This model should keep growing and spreading to the health care centers that have the capacity to perform PCI.

Conclusions

There were no differences in survival and mortality in STEMI patients treated by means of PCI or PS. PS is a viable, effective, and safe option for optimal reperfusion in Mexican population according to its social and economic limitations.

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Conflicts of interest

None.

Ethical disclosures

Protection of human and animal subjects. The authors declare that no experiments were performed on humans or animals for this study.

Confidentiality of data. The authors declare that they have followed the protocols of their work center on the publication of patient data.

Right to privacy and informed consent. The authors declare that no patient data appear in this article.

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