

Survival curve identifies critical period for postoperative mortality in cardiac patients undergoing emergency general surgery

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Research article

Keywords: cardiac patients, emergency general surgery, survival curve, mortality

Posted Date: February 24th, 2020

DOI: <https://doi.org/10.21203/rs.2.24285/v1>

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Version of Record: A version of this preprint was published at Scientific Reports on September 23rd, 2020.
See the published version at <https://doi.org/10.1038/s41598-020-72647-7>.

Abstract

Introduction: The number of non-cardiac major surgeries carried out has significantly increased in recent years to around 200 million procedures carried out annually. Approximately 30% of patients submitted to non-cardiac surgery present some form of cardiovascular comorbidity. In emergency situations, with less surgery planning time and greater clinical severity, the risks become even more significant. The aim of this study is to determine the incidence and clinical outcomes in patients with cardiovascular disease submitted to non-cardiac surgical procedures in a single cardiovascular referral center.

Methods: This is a prospective cohort study of patients with cardiovascular disease submitted to non-cardiovascular surgery. All procedures were carried out by the same surgeon, between January 2006 and January 2018.

Results: 240 patients included were elderly, 154 were male (64%), 8 patients presented two diagnoses. Of the resulting 248 procedures carried out, 230 were emergency (92.8%). From the data obtained it was possible to estimate the day from which the occurrence of mortality is less probable in the postoperative phase.

Conclusion: Our research evaluated the epidemiological profile of the surgeries and we were able to estimate the survival and delimit the period of greatest risk of mortality in these patients. The high rate of acute mesenteric ischemia was notable, a serious and frequently fatal condition.

Introduction

The number of non-cardiac major surgeries carried out has significantly increased in recent years to around 200 million procedures carried out annually¹. This type of surgery exposes patients to risks such as hemorrhage, inflammatory systemic responses, infection, pulmonary embolism and acute myocardial infarction, which can lead to acute overload of the cardiovascular system. According to European Society of Cardiology and European Society of Anaesthesiology data, approximately 30% of patients submitted to non-cardiac surgery present some form of cardiovascular comorbidity², with a resulting greater risk of complications or a fatal cardiovascular event in the postoperative period.

In emergency situations, with less surgery planning time and greater clinical severity, the risks become even more significant, multiplying the incidence of myocardial infarction or mortality because of postoperative cardiovascular conditions by between 2.5 and 4 times³. This increased risk can be identified by the American Society of Anesthesiologists (ASA)⁴ score, among others.

In 2014, the American College of Cardiology/American Heart Association (ACC/AHA) Task Force for Practice Guidelines published a review of its recommendations for evaluation of cardiovascular risk in patients submitted to non-cardiac procedures⁵. As more surgical procedures take place in elderly patients, anesthesiologists will more often face patients with diagnosed or even suspected chronic heart failure in the perioperative period⁶. Patients with chronic heart failure form a distinct risk population, which is

associated with increased risk in the perioperative period. These initiatives show the relevance of the clinical conditions found in cardiopathic patients and their evolution in the postoperative period.

The aim of this study is to determine the incidence and clinical outcomes in patients with cardiovascular disease submitted to non-cardiac surgical procedures in a single cardiovascular referral center.

Methods

This is a prospective cohort study of patients with cardiovascular disease submitted to non-cardiovascular surgery. All procedures were carried out by the same surgeon, between January 2006 and January 2018, in the General Surgery Service at the Dante Pazzanese Institute of Cardiology (DPIC). The information related to the surgical procedures was taken from a database and complemented with an analysis of medical records.

The variables analyzed were sex, age, clinical diagnosis, surgical indication, surgical procedures carried out, short term mortality (up to 30 days postoperative) and long-term mortality.

Inclusion criteria

The patients included in this study were over the age of 18, with a cardiovascular disease referred for surgical non-cardiac treatment at the institution, either in emergency or for elective surgery.

Exclusion Criteria

There were no specific exclusion criteria, given that the aim was to report all adult patients that had a documented diagnosis and indication for a non-cardiovascular surgical procedure.

Statistical Analysis

A descriptive exploration is presented in frequency tables for qualitative variables, including the confidence intervals of the proportions.

The survival curves were estimated by the Kaplan-Meier method (observed in figure 1). For inter-group comparison the Log-Rank (Mantel-Cox), Breslow (Generalized Wilcoxon) and Tarone-Ware were used. All significance tests were two-tailed and with a confidence interval of 95%. The calculations were carried out using the statistical package IBM SPSS 25 (Statistical Package for the Social Sciences).

Ethical Considerations

This study was approved by the Ethics Committee of the Dante Pazzanese Institute of Cardiology.

Results

All the 240 patients included were elderly, 154 were male (64%), 8 patients presented two diagnoses. Of the resulting 248 procedures carried out, 230 were emergency (92.8%). The most common diagnoses were: acute mesenteric ischemia (53), acute cholecystitis (46), acute appendicitis (25), Fournier syndrome (24), emergency hernia (19), perforated peptic ulcer (9), colonic neoplasia (8). Other diagnoses comprised 64 procedures (25.8%) (Table 1).

We considered short term mortality as less than 30 days after surgical intervention. As an overall result, postoperative mortality was 30% (72 cases). The diagnoses with the highest mortalities were acute mesenteric ischemia (71.7%), perforated peptic ulcer (66.6%) and colonic neoplasia (37.5%). Other diagnoses also present significant mortality (table 2), except for elective hernias (no mortalities).

For all the procedures observed, there were no significant differences found in postoperative mortality between the sexes.

From the data obtained it was possible to estimate the day from which the occurrence of mortality is less probable in the postoperative phase (Table 3) and the necessary days to stabilize the survival curve (figure 1).

Discussion

In our study, an analysis of surgical procedures was carried out and the mortality was estimated in procedures of varying complexity, from exploratory laparotomy to elective hernioplasty. Many of the interventions (92.8%) were emergency surgery, which represents a risk factor for 30-day mortality.

The emergency procedures present significant numbers. In the United States, for example, there is an annual mean of around 3 million emergency surgeries carried out⁷. In this context, Reinke et al., carried out a study assessing hospitalization for non-cardiac general surgery⁸. The most common surgical procedures were cholecystectomy, wound debridement, partial resection of the small intestine and appendectomy, while in our study the most prevalent were exploratory laparotomy, cholecystectomy, appendectomy and wound debridement of perineal abscesses. A correspondence was verified between the cases in our study and those reported in others with similar risk factors and comorbidities^{8,9}.

Despite the higher number of patients with high cardiovascular risk, the mortality obtained in the American study in patients submitted to emergency surgery was 3.8%⁹, a much lower number than that found in our study. This contrast could be due to the severity of the conditions the patients presented in our center and the elevated rate of acute mesenteric ischemia (AMI). With 53 occurrences, this diagnosis was responsible for 52.7% of the mortality in our study.

AMI is caused by a sudden interruption of the blood supply for a segment of the small intestine, causing ischemia, cellular damage and intestinal necrosis. Most cases are caused by embolic occlusion (40 to 50%) or by thrombotic occlusion (25 to 35%). Mesenteric embolisms are associated with cardiac complications such as atrial fibrillation, left ventricle myocardial dysfunction and reduced ejection

fraction or cardiac valves damaged by endocarditis. AMI is an uncommon cause of abdominal pain, corresponding to less than 0.1% of hospitalizations in a general hospital, but its mortality is between 60 to 80%¹⁰.

In a recently published article, Gilshtein et al. identified 63 patients with ischemic colitis, the most common form of AMI, in one tertiary center¹¹. Most patients were treated clinically, and only the 20% most serious cases, with complications such as peritonitis, hemodynamic instability and untreatable disease, received surgical intervention. These patients had a mortality of 50%, lower than our study (71.7%), which reinforces the gravity of the condition of the patients referred to the center in our study.

From our statistical analysis it was possible to estimate that for AMI, mortality occurs until a mean of 11 days, which represents more than 90% of the deaths from this condition. This information could affect the management of the patient, as it defines the phase of greatest risk.

Another diagnosis that was relevant to our study was the perforated peptic ulcer. Complications of the ulcer, as well as perforation, include bleeding and obstruction. When perforated, the ulcer must be referred to emergency surgical treatment, and is associated with a 30% mortality rate in 30 days^{12,13}. Among our patients, a significantly higher mortality rate of 66.7% was observed, probably due to the severity of the condition of patients referred to the center. Furthermore, it was possible to identify that patients tended to die up to the 12th postoperative day. The large CI of 95% (between 5 and 20 days) was a result of our small sample with just 9 patients.

Acute cholecystitis and acute appendicitis were also highly prevalent in our study, but the 30-day mortality of both was low, which resulted in a survival curve with little predictive value.

Similarly, the other diagnoses did not collect enough cases, which shows a need for other studies with a larger sample of patients to obtain significant results. This would allow the period of greatest risk to patients to be identified, and allow care to be tailored accordingly, as with AMI.

Conclusion

Our research evaluated the epidemiological profile of the surgeries carried out in DPIC and we were able to estimate the survival and delimit the period of greatest risk of mortality in these patients. The high rate of AMI was notable, a serious and frequently fatal condition.

List Of Abbreviations

ASA - American Society of Anesthesiologists

ACC/AHA - American College of Cardiology/American Heart Association

DPIC - Dante Pazzanese Institute of Cardiology

Declarations

- Ethical Approval and Consent to participate

This study was approved by the Ethics Committee of the Dante Pazzanese Institute of Cardiology and all patients consented to participate

- Consent for publication

All authors consented for publication

- Availability of supporting data

Not applicable

- Competing interests

Not applicable

- Funding

No funding

- Authors' contributions

Diego Ramos Martines: Conceptualization; Data curation; Formal analysis; Investigation; Methodology; Roles/Writing - original draft; Writing - review & editing.

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Gustavo Oliveira: Supervision; Validation; Visualization; Writing - review & editing.

Antonio C Mugayar: Supervision; Validation.

Alberto Meyer: Supervision; Validation; Visualization; Writing - review & editing.

- Acknowledgements

The authors are thankful to Justin Axel-Berg for the English corrections.

References

1. Devereaux PJ, Sessler DI. Cardiac Complications in Patients Undergoing Major Noncardiac Surgery. *N Engl J Med.* 2015;373(23):2258-69.
2. Kristensen SD, Knuuti J, Saraste A, Anker S, Bøtker HE, De Hert S, Ford I, Gonzalez Juanatey JR, Gorenek B, Heyndrickx GR, Hoeft A, Huber K, Iung B, Kjeldsen KP, Longrois D, Luescher TF, Pierard L, Pocock S, Price S, Roffi M, Sirnes PA, Uva MS, Voudris V, Funck-Brentano C; Authors/Task Force Members. 2014 ESC/ESA Guidelines on non-cardiac surgery: cardiovascular assessment and management: The Joint Task Force on non-cardiac surgery: cardiovascular assessment and management of the European Society of Cardiology (ESC) and the European Society of Anaesthesiology (ESA). *Eur J Anaesthesiol.* 2014;31(10):517-73.
3. Smilowitz NR, Gupta N, Guo Y, Berger JS, Bangalore S. Perioperative acute myocardial infarction associated with non-cardiac surgery. *Eur Heart J.* 2017;38(31):2409-2417.
4. de Nadal M, Pérez-Hoyos S, Montejo-González JC, Pearse R, Aldecoa C; European Surgical Outcomes Study (EuSOS) in Spain. Intensive care admission and hospital mortality in the elderly after non-cardiac surgery. *Med Intensiva.* 2018;42(8):463-472.
5. Kaw R, Nagarajan V, Jaikumar L, Halkar M, Mohananey D, Hernandez AV, Ramakrishna H, Wijeyesundara D. Predictive Value of Stress Testing, Revised Cardiac Risk Index, and Functional Status in Patients Undergoing Noncardiac Surgery. *J Cardiothorac Vasc Anesth.* 2019;33(4):927-932.
6. Smit-Fun V, Buhre WF. The patient with chronic heart failure undergoing surgery. *Curr Opin Anaesthesiol.* 2016;29(3):391-6.
7. Gale SC, Shafi S, Dombrovskiy VY, Arumugam D, Crystal JS. The public health burden of emergency general surgery in the United States: A 10-year analysis of the Nationwide Inpatient Sample–2001 to 2010. *J Trauma Acute Care Surg.* 2014;77(2):202-8.
8. Reinke CE, Thomason M, Paton L, Schiffenbauer L, Rozario N, Matthews BD. Emergency general surgery transfers in the United States: a 10-year analysis. *J Surg Res.* 2017;219:128-135.
9. Ingraham A, Wang X, Havlena J, Hanlon B, Saucke M, Schumacher J, Fernandes-Taylor S, Greenberg C. Factors Associated With the Interhospital Transfer of Emergency General Surgery Patients. *J Surg Res.* 2019;240:191-200.
10. Clair DG, Beach JM. Mesenteric Ischemia. *N Engl J Med.* 2016;374(10):959-68.

11. Gilshtein H, Hallon K, Kluger Y. Ischemic colitis caused increased early and delayed mortality. *World J Emerg Surg.* 2018;13:31.
12. Søreide K, Thorsen K, Harrison EM, Bingener J, Møller MH, Ohene-Yeboah M, Søreide JA. Perforated peptic ulcer. *Lancet.* 2015;386(10000):1288-1298.
13. Møller MH, Engebjerg MC, Adamsen S, Bendix J, Thomsen RW. The Peptic Ulcer Perforation (PULP) score: a predictor of mortality following peptic ulcer perforation. A cohort study. *Acta Anaesthesiol Scand.* 2012;56(5):655-62.

Tables

Table 1: Other diagnosis (number of procedures)

Elective hernia (18)	Enteric fistula (2)	Ovarian cyst (1)
Infected hematoma (5)	Splenic trauma (1)	Ovarian neoplasia (1)
Gastric neoplasia (4)	Intestinal perforation (1)	Esophageal obstruction (1)
Choledocholithiasis (3)	Biliary neoplasia (1)	Liver failure (1)
Active obstructive abdomen (3)	Lymphoma (1)	Hemo retroperitonitis (1)
Splenic aneurysm (1)	Meckel's diverticulum (1)	Hepatic neoplasia (1)

Table 2: Distribution of mortality by diagnosis

	Live		Death within 30 days		Death after 30 days	
	N	%	N	%	N	%
ACUTE MESENTERIC ISCHEMIA	14	26,4%	38	71,7%	1	1,9%
ACUTE CHOLECYSTITIS	40	87%	3	6,5%	3	6,5%
ACUTE APPENDICITIS	20	80%	3	12%	2	8%
FOURNIER	15	62,5%	4	16,7%	5	20,8%
EMERGENCY HERNIA	11	57,9%	6	31,6%	2	10,5%
PERFORATED PEPTIC ULCER	3	33,3%	6	66,7%	0	0,0%
COLON NEOPLASIA	5	62,5%	3	37,5%	0	0,0%
OTHER DIAGNOSES	49	76,6%	9	14%	6	9,3%

Table 3: Estimate in days of fatal events during the postoperative period

Diagnosis	Estimate	Standard Error	95% Confidence interval	
			Lower limit	Upper Limit
ACUTE MESENTERIC ISCHEMIA	11.3	1.764	7.9	14.8
ACUTE CHOLECYSTITIS	28.3	0.961	26.4	30.2
ACUTE APPENDICITIS	27.6	1.466	24.8	30.5
FOURNIER	26.3	1.797	22.7	29.8
EMERGENCY HERNIA	23.8	2.919	18.1	29.5
PERFORATED PEPTIC ULCER	12.6	3.869	5	20.1
COLON NEOPLASIA	23.3	3.220	16.9	29.6

Figures

Overall patients 30 day survival

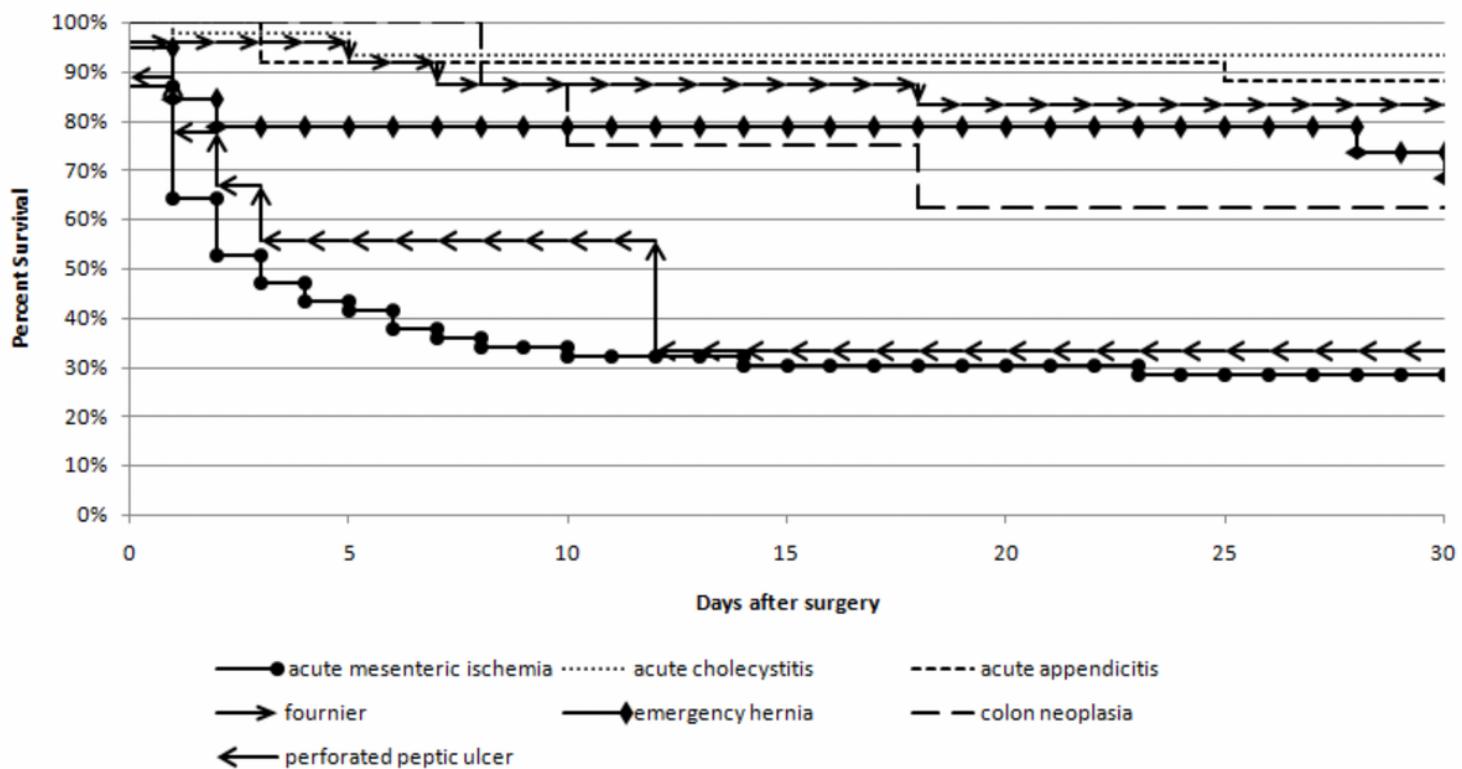


Figure 1

30 day survival curve in patients with the most common diagnoses.