

Investigation of the Postoperative Complications Rate and Predictors in Patients Undergoing Surgery due to Associated Carotid and Coronary Occlusive Disease

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ABSTRACT

Background: The aim of this study was to evaluate the frequency of postoperative complications in patients who underwent coronary artery bypass grafting (CABG) and simultaneous carotid endarterectomy (CEA) and find predictors of postoperative complications.

Methods: We retrospectively evaluated 86 patients after simultaneous CABG and CEA. Inclusion criteria were: patients with asymptomatic carotid stenosis with a reduction of the carotid lumen diameter of more than 70% detected with Doppler ultrasound and diagnosed with one, two, or three vessel coronary artery disease with coronary stenosis more than 75% and hemodynamic significant stenosis of the left main artery. Exclusion criteria were patients with urgent and previous cardiac surgery and patients with myocardial infarction and stroke in the past one month. We monitored preoperative (ejection fraction, coronarography status), operative (number of grafts, on-pump or off-pump technique) and postoperative (extubation, unit care and hospital stay, bleeding and reoperation) details and complications (myocardial infarction, neurological events, inotropic agents and transfusion require, infection, arrhythmic complication, renal failure, mortality).

Results: Postoperative complications were observed in 18 (20.9%) patients. Two patients (2.3%) had postoperative stroke and one patient (1.2%) had transient ischemic attack (TIA). Previous stroke was a predictor for increased postoperative neurological events ($P < .05$). Intra-hospital mortality was 8.1%.

Conclusion: Simultaneous CEA and CABG were performed with low rates of stroke and TIA. Previous stroke was identified as a predictor for increased postoperative neurological complications.

INTRODUCTION

Concomitant carotid and coronary disease is a common problem and an important risk factor of coronary artery

bypass grafting (CABG)-associated stroke. The risk of cerebrovascular accident has been reported to increase up to 14% in patients with severe carotid artery stenosis [Huh 2003; Khaitan 2000]. There is no consensus as to the optimal management of patients with concomitant carotid and coronary disease [Naylor 2010]. Also, treatment strategies are not supported by studies with level 1 evidence [Cywinski 2006]. One of the strategies includes simultaneous CABG and CEA that is associated with low rates of mortality and morbidity [Eren 2005; Levy 2012; Ren 2012; Yuan 2009]. On the other hand, a systematic review of 97 studies in patients treated with CEA and CABG showed that the highest rate of stroke and death was noticed in the group of patients where these procedures were performed simultaneously [Naylor 2003]. There is insufficient data in domestic literature regarding the outcome of simultaneous operative treatment of patients with associated carotid and coronary occlusive disease. Thus, the purpose of this study was to evaluate postoperative complications of patients who underwent CABG and simultaneous CEA, with particular reference to neurological events. Also we wanted to identify significant predictors that were associated with postoperative outcome.

METHODS

The retrospective cohort study with a short-term follow-up period was conducted in order to identify predictors of postoperative complications. The study included 86 patients who underwent simultaneous CABG and CEA during the same anesthetic setting in a 4-year period from 2012 to 2016 (Clinic for Cardiac Surgery, Clinical Centre of Serbia, Medical School, University of Belgrade). The observed follow-up period was duration of hospitalization after surgical treatment. The study was approved by the Medical School, University of Belgrade research ethics committee. The indications for simultaneous procedure were: patients with asymptomatic carotid stenosis with a reduction of the carotid lumen diameter of more than 70% detected with Doppler ultrasound and diagnosed with one, two, or three vessel coronary artery disease with coronary stenosis more than 75% and hemodynamic significant stenosis of the left main artery. Exclusion criteria were patients with urgent and previous cardiac surgery and patients with myocardial infarction and stroke in the

Received March 19, 2018; received in revised form November 8, 2018; accepted November 14, 2018.

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Table 1. Demographic, Preoperative, and Operative Characteristics

	n = 86
Age, mean \pm SD	67.2 \pm 7.2
Sex, n (%)	
Male	57 (66.3)
Female	29 (33.7)
Comorbidity, n (%)	
Hypertension	83 (96.5)
Hyperlipidemia	63 (73.2)
Diabetes mellitus	34 (39.5)
COPD	10 (11.9)
Renal failure	1 (1.2)
TIA	1 (1.2)
Stroke	5 (6)
EF (%), mean \pm SD	51.2 \pm 11.9
Coronary disease, n (%)	
One-vessel	3 (3.5)
Two-vessel	10 (11.6)
Three-vessel	42 (48.8)
Left main artery	31 (36)
Off-pump CABG, n (%)	12 (14)
On-pump CABG, n (%)	74 (86)
Number of grafts, med (IR)	3 (1)

COPD indicates chronic obstructive pulmonary disease; TIA, transient ischemic attack; EF, ejection fraction; med, median; IR, interquartile range.

past one month. Our strategy was that after the CAE (done by the vascular team), the wound was completely closed on the neck, and then the off-pump CABG procedure was done. If an extracorporeal circulation (ECC) was necessary for a given patient, it was advantageous for the bypass on a full to create proximal anastomosis. Regardless of the choice of CABG procedure, full heparinization was applied (400 IU/kg). Patients monitoring during hospitalization included demographic characteristics (age and sex), comorbidity, preoperative (ejection fraction [EF], coronarography status), operative (number of grafts, on-pump or off-pump technique) and postoperative (extubation, unit care stay, hospital stay, bleeding, and reoperation) details, complications (myocardial infarction [MI], neurological events, inotropic agents and transfusion require, infection, arrhythmic complication, renal failure, mortality). The multidisciplinary team participated in the treatment of the patients. On suspicion of a neurological event, a neurologist was consulted who, after a neurological examination, indicated CT scan of the endocranium based on which the diagnosis of stroke was made. Statistical analyses were performed using IBM SPSS version 24.0 packaged software and included

Table 2. Postoperative Characteristics and Complications

	n = 86
Extubation, days, median (IR)	1 (1)
Intensive unit stay, days, med (IR)	4 (2)
Hospital stay, days, mean \pm SD	11 \pm 6.2
Transfusion, n (%)	48 (55.8)
Complications, n (%)	18 (20.9)
Inotropic support, n (%)	20 (23.2)
Reoperation for bleeding, n (%)	5 (5.8)
Hemodialysis, n (%)	6 (7)
Stroke, n (%)	2 (2.3)
TIA, n (%)	1 (1.2)
Mediastinitis, n (%)	1 (1.2)
Arrhythmic complications, n (%)	7 (8.1)
Mortality, n (%)	7 (8.1)

IR indicates interquartile range; TIA, transient ischemic attack.

methods of descriptive statistics (measures of central tendency [mean and median], measures of variability [standard deviation], interquartile range [IQR], relative numbers and methods of analytic statistics [chi square test, Student t test, Pearson and Spearman correlations). Cox regression analysis was used to identify predictors of postoperative complications.

RESULTS

Demographic, preoperative, and operative data are summarized in Table 1. The mean age of patients was 67.2 \pm 7.2 years. Of the 86 patients, 57 (66.3%) were men, and 29 (33.7%) were women. A history of hypertension was present in 83 patients (96.5%). Hyperlipidemia and diabetes mellitus (DM) were present in 63 (73.2%) and 34 (39.5%) patients, respectively. Five patients (6%) had a history of previous stroke and one patient (1.2%) had a history of previous transient ischaemic attack (TIA). Ten patients (11.6%) reported chronic obstructive pulmonary disease (COPD) and one patient (1.2%) reported renal failure (Table 1). The mean ejection fraction (EF) was 51.2 \pm 11.9 (range: 21% to 75%). A large number of patients (48.8%) had three vessel coronary disease, 36% of patients had left main disease. 3.5% and 11.6% of patients had one and two vessel coronary disease, respectively (Table 1). The majority of patients (86%) underwent on-pump CABG and the median number of grafts was 3 (Table 1). Postoperative characteristics and complications are listed in Table 2. Eighteen patients (20.9%) developed complications including 2 strokes (2.3%), 1 TIA (1.2%), 7 arrhythmic complications (8.1%), 6 renal failure requiring hemodialysis (7%), 5 reoperations for bleeding (5.8%), 1 mediastinitis (1.2%). Twenty patients (23.2%) required the consumption of

Table 3. Variables Associated with Postoperative Stroke and Mortality

Variables	Postoperative stroke		Mortality	
	RR(CI)	P	RR (CI)	P
Previous stroke	15.8 (1.02-252.6)	.049	2.63 (0.32-21.88)	.370
Postoperative complications	–		9.44 (1.83-48.68)	.007

RR indicates relative risk; CI, confidence interval.

inotropic agents and 48 patients (55.8%) required transfusion. The average length of unit care and hospital stay was 4 and 11 ± 6.2 days, respectively (Table 2). In-hospital deaths occurred in 7 patients (8.1%) (Table 2). Of the 7 patients, 4 were men and 3 were women. Causes of deaths were neurologic events (strokes) in two patients, arrhythmic complications in one patient (atrial fibrillation in the early postoperative course), heart failure in one patient. The cause of death of three patients were not neurological and cardiac complications

Regression analysis revealed some predictors for mortality and postoperative strokes (Table 3). We found significant correlation between postoperative complications and mortality rate. Patients with complications had higher mortality rate ($\rho = -0.370$; $P < .001$) and presence of complications was significantly associated with in-hospital mortality ($P = .007$) (Table 3). Hospital stay correlated weakly with mortality ($\rho = -0.256$; $P = .041$). Patients with longer hospital stay had higher mortality rate but hospital stay was not significantly associated with mortality rate ($P = .088$). Patients with three or more associated diseases had a higher incidence of mortality compared to patients with fewer associated illnesses but without significant difference ($P = .095$). Individually, previous stroke was not significantly associated with mortality rate ($P > .05$) (Table 3). In the group of patients with previous stroke, we found higher prevalence of postoperative neurologic events (20%) in relation to group of patients without previous stroke (1.3%) and that was statistically significant ($\chi^2 = 7.101$; $P = .008$). Previous stroke was significantly associated with the prevalence of postoperative neurologic events ($P < .05$) (Table 3).

DISCUSSION

Concomitant carotid and coronary disease is a very frequent clinical problem. In the associated coronary and carotid disease, a multidisciplinary team composed of cardiac surgeons, cardiologists, anesthesiologists, neurologists, transfusionists, and vascular surgeons has established the protocol of the Clinical Center of Serbia according to which the primary coronary operation without previous carotid operation is carried out in the following conditions: Asymptomatic unilateral stenosis of the carotid arteries less than 80% or previous stroke with a carotid stenosis of less than 80%. Primary carotid operation without previous coronary operation-carotid stent is performed in patients with acute neurological symptoms with bilateral lesions of carotid arteries

greater than 50% with or without contralateral occlusion or in the case of a stroke linked to carotid stenosis greater than 80% in the case of stable chronic angina in the absence of a recent infarction. Simultaneous surgery in patients with both carotid and coronary disease is performed in patients with unstable angina pectoris, left main stenosis, and diffuse coronary disease without satisfactory collaterals. The basic controversy is about a combined procedure in patients with asymptomatic carotid artery stenosis greater than 80%, and in this case the team decision adjusts to individual patients characteristics. Postoperative stroke is one of the most concerning complications after CABG, with reported 24.8% mortality [John 2000]. A 9.2% rate of stroke or TIA in cardiac surgical patients with asymptomatic carotid stenosis is reported when compared with 1.3% of patients with no carotid stenosis [Brenner1987]. The possible mechanism is via embolization from unstable plaques or by decreased cerebral blood flow distal to critical stenosis. Considering the patient's risk profile, lesion morphology and symptom status, simultaneous surgical strategy (CEA before CABG) may be considered. The main goal is stroke prevention or reduction of the stroke rate. Thus, the purpose of this study was to provide a description of simultaneous CEA and CABG outcome and find possible factors that were associated with patient's postoperative status. The majority of studies reported postoperative complications in relation to stroke and mortality [Aydin 2014; Nwakanma 2006; Gansera 2012; Illuminati 2011]. We primarily examined the frequency of stroke and mortality, but we also investigated and described other complications that might be expected during the postoperative course. The first studies on combined surgical approach showed a reduced incidence of postoperative complications primarily in the sense of stroke and myocardial infarction in simultaneous CEA and CABG [Trachiotis 1997]. On the other hand, there are studies that report a significantly higher percentage of postoperative complications (stroke 4.48%, TIA 1.66%, and intrahospital mortality 3.56%) relative to reference groups [Gopaldas 2011; Prasad 2010]. The authors of these studies emphasized the importance of further detailed analysis and scientific research when it comes to the choice of treatment of patients with associated carotid occlusive and coronary disease. Generally speaking, the majority of patients in our study did not have complications during the early postoperative period (79.1%) and 20.9% of patients had complications. The most frequent complication was inotropic support (31.3%), while the least frequent complications were TIA (1.2%) and mediastinitis (1.2%). Similarly, the rate

of TIA (1.2%) was close to the results by Aydin et al (0.9%) [Aydin 2014]. A higher percent of patients with postoperative TIA (2%) was observed in the study by Kolh et al [Kolh 2006]. There were no patients with myocardial infarction in the early postoperative period and that is in accordance with other studies [Aydin 2014; Nwakanma 2006]. Arrhythmic complications in our study were higher (5.8%) in relation to the results by Aydin et al (1%) [Aydin 2014]. The highest rate of arrhythmic complication (18.5%) was reported in the study by Nwakanma et al [Nwakanma 2006]. 5.9% of patients were reoperated due to postoperative bleeding, 7% of patients required hemodialysis. In other studies, a number of reoperations were observed (8.04%) [Prasad 2010] and there were no patients with acute renal insufficiency [Nwakanma 2006]. Simultaneous CEA and CABG in our study population were performed with low rates of stroke (2.3%) and TIA (1.2%) and that is in accordance with other studies [Aydin 2014]. The authors of a previous study reported the postoperative stroke rate of 2% and postoperative TIA rate of 0.9% [Aydin 2014]. Similar results were observed in other research papers [Levy 2012; Kougiass 2007; Naylor 2011]. In the study by Levy et al [Levy 2012], the prevalence of postoperative stroke was 2.5% while the study by Naylor et al [Naylor 2011] and Kougiass et al [Kougiass 2007] reported the prevalence of postoperative stroke at 2.7%. A higher frequency of postoperative strokes (3.7%) was registered by Nwakanma et al [Nwakanma 2006], but the mean age of the patients in their study was higher in relation to our study and they reported a higher frequency of emergency elective surgery (40.7%). In our study, all patients underwent emergency surgery. Also, a higher percent of postoperative neurological events (stroke: 3.9%) was noticed in the study by Timaran et al [Timaran 2008]. In our study, both patients with postoperative stroke were from the “on-pump” group, which was expected considering systemic hypotension, and in combination with stenosis of intracranial arteries, it can lead to brain hypoperfusion. It has been shown that this phenomenon is responsible for 8.8% of strokes in patients undergoing CABG [Likosky 2003]. In our study, seven patients died (8.1%), neurological and cardiac complications were not the cause of the death of three patients, one patient died due to atrial fibrillation in the early postoperative course, one patient had heart failure, while the cause of death of two patients was related to neurological complications (stroke). There was a strong correlation between postoperative complications and mortality rate. Also, in the group of patients with complications, a higher incidence of mortality was observed, which was statistically significant. Similar results were observed by Kolh et al [Kolh 2006]. In our study, the regression analysis revealed that the presence of postoperative complications was the predictor of mortality ($P = .007$) (Table 3). Patients with three or more associated diseases had a higher incidence of mortality compared to patients with fewer associated illnesses, and this correlation was close to the statistical significance level, while in the regression model it did not distinguish itself as a significant predictor. This is comparable with results by Busch et al [Busch 1999]. The results of regression analyses in literature data indicate that socio-demographic characteristics (age), comorbidity (cardiac insufficiency), and

postoperative characteristics (reintubation) are predictors of mortality [Nwakanma 2006; Gansera 2012]. In our study, comorbidity (previous stroke) was a predictor of postoperative stroke ($P < .05$). The results of this study showed that simultaneous CEA and CABG were performed with low rates of stroke and TIA. On the other hand, the rate of mortality was high (8.1%). These results should have a direct clinical application and influence on the optimal decision-making and plan in terms of the details of surgical treatment of coronary disease in patients with simultaneous and associated carotid occlusive disease. The limiting factors of this study are the lack of a control group and evaluation of long-term results.

Conclusion

Simultaneous CEA and CABG were performed with low rates of stroke and TIA and previous stroke was identified as a predictor for increased postoperative neurological events. Further studies are needed to maintain the long-term follow-up.

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