

Reoperation for bleeding in cardiac surgery

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Abstract

At Odense University Hospital (OUH), 5–9% of all unselected cardiac surgical patients undergo reoperation due to excessive bleeding. The reoperated patients have an approximately three times greater mortality than non-reoperated. To reduce the rate of reoperations and mortality due to postoperative bleeding, we aim to identify risk factors that predict reoperation. A total of 1452 consecutive patients undergoing cardiac surgery using extracorporeal circulation (ECC) between November 2005 and December 2008 at OUH were analysed. Statistical tests were used to identify risk factors for reoperation. We performed a case-note review on propensity-matched patients to assess the outcome of reoperation for bleeding regarding morbidity and mortality. In total, 101 patients (7.0%) underwent surgical re-exploration due to excessive postoperative bleeding. Significant risk factors for reoperation for bleeding after cardiac surgery was low ejection fraction, high EuroSCORE, procedures other than isolated CABG, elongated time on ECC, low body mass index, diabetes mellitus and preoperatively elevated s-creatinine. Reoperated patients significantly had a greater increase in postoperative s-creatinine and higher mortality. Surviving reoperated patients significantly had a lower EuroSCORE and a shorter time on ECC compared with non-survivors. The average time to re-exploration was 155 min longer for non-survivors when compared with survivors.

Keywords: Cardiac surgery • Reoperation • Postoperative bleeding • Risk factors

INTRODUCTION

Risk of reoperation due to bleeding after cardiac surgery is shown to be 2.2–4.2% [1–6]. Risk factors are high age, low body mass index (BMI) or body surface area (BSA), time on extracorporeal circulation (ECC), five or more anastomosis or non-elective operation [2–4]. Reoperated patients have a two to six times greater mortality [1, 3–5] and a greater morbidity regarding renal and pulmonary function, sepsis and arrhythmia [1, 4, 5]. Postoperative bleeding can be due to surgical or coagulopathic factors [1]. Patient-related factors are also indicative for reoperation due to bleeding.

At Odense University Hospital (OUH), 5–9% of all unselected cardiac surgical patients undergo reoperation due to excessive bleeding. The reoperated patients had three times greater mortality and, to lower the rate of reoperation due to bleeding and improve the mortality, we have tried to identify risk factors in patients as well as in procedures.

MATERIALS AND METHODS

Patient population and data

Patient data were collected consecutively from The Western Denmark Heart Registry [7] (WDHReg) during 18 November 2005 until 31 December 2008 at OUH. During this period, we included all patients undergoing cardiac surgery using ECC. Of

1452 patients found, 101 patients (7.0%) underwent at least one reoperation within 24 h due to excessive postoperative bleeding. Pre-, peri- and postoperative characteristics were collected from WDHReg complemented by case records. Information included the patient's age, type of operation, complications, bleeding and possible causes of death. We qualified risk factors for reoperation by comparing the results of 101 reoperated patients with the remaining 1351 patients.

Using recorded operative details, we classified the bleeding as coagulopathic or surgical. Blood or haematoma without ongoing bleeding and/or diffuse oozing was defined as coagulopathic bleeding. Specific bleeding requiring clips or suture was defined as surgical bleeding.

Propensity-matched group

To assess the outcome of reoperation for bleeding regarding morbidity and mortality, we performed a case-note review on propensity-matched patients. Patients requiring re-exploration for bleeding were propensity matched with a unique patient not re-explored using the following criteria: priority of the operation, age, BMI, medicine (acetylsalicylic acid, clopidogrel, coumadine or heparin) within the last 5 days, EuroSCORE (European System for Cardiac Operative Risk Evaluation) and sex. When possible we also matched patients with regard to ejection fraction (EF), diabetes mellitus (DM) and the type of procedure. The mentioned criteria are the result of a literature study [2, 5]. Matching

the patients using these criteria, we minimized their influence as confounders when analysing the consequences of reoperations.

Mortality was defined as death within 30 days after the first operation. Complications such as perioperative myocardial infarction, transient cerebral ischaemic attack (TIA), stroke and sternal infection had to be diagnosed by a doctor within the same hospital admission as the index surgery. The morbidity definitions were according to the definitions of the protocol at WDHReg [7]. Stroke is defined as a neurological deficit of cerebrovascular cause that persists beyond 24 h or is interrupted by death within 24 h. TIAs share the same aetiology as strokes, but the symptoms resolve within 24 h. Sternal infections often appeared after the discharge, so we searched the case records after discharge.

The decision to perform reoperation was made by the surgeon on call and based on conventional guidelines: (i) drainage of >500 ml in the first hour, total drainage of >800 ml in the first 2 h, 900 ml in the first 3 h, >1000 ml in the first 4 h or 1200 ml in the first 5 h; (ii) sudden massive bleeding or cardiac tamponade.

Statistical analysis

Differences between the two groups were explored using an independent *t*-test for continuous variables and Fisher's exact test and Pearson's χ^2 test for categorical variables. Data were given as mean values, standard deviations of mean values and in numbers and percentages. In all cases, a *P*-value <0.05 was considered as significant.

Statistical analysis was performed with SPSS (SPSS Inc., PASW Statistics 18).

RESULTS

Causes of death were divided into circulatory, multi-organ failure, bowel ischaemia and unknown (see Fig. 1).

The total population

Of the 1452 patients undergoing on-pump cardiac surgery during the study period, 101 patients (7.0%) underwent a surgical re-exploration for postoperative bleeding (types of bleeding, Fig. 2).

Mortality in patients re-explored for surgical bleeding and coagulopathic bleeding was 19 and 10.5%, respectively.

Patients in the re-explored group had a higher mortality when compared with patients who were not re-explored (15.8 versus 5.7%, *P* < 0.001). Predictors for reoperation due to bleeding are listed in Table 1.

Matched group

The group of reoperated patients and propensity-matched group were homogeneous with respect to preoperative and perioperative variables. However, preoperative s-creatinine was significantly higher in the re-explored group when compared with the control group (Table 2).

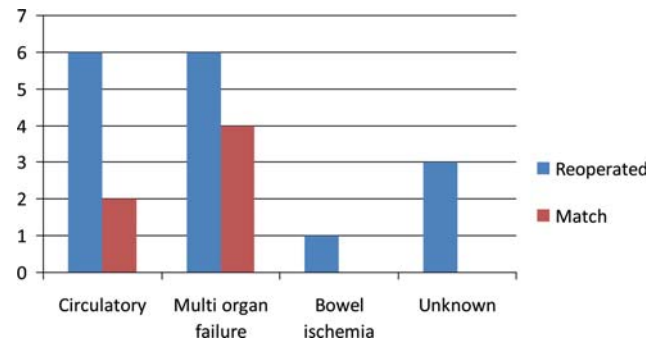


Figure 1: Causes of death among the 16 reoperated and 6 matched patients.

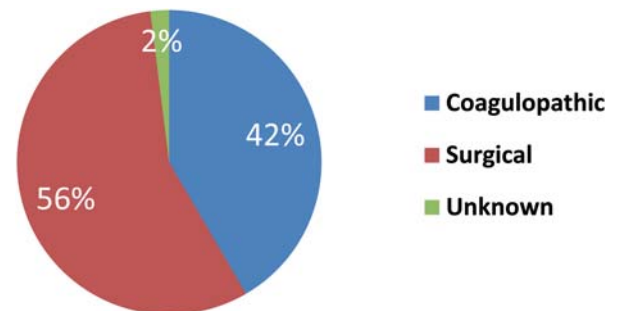


Figure 2: Types of bleeding found at reoperation. Forty-two patients had a coagulopathic bleeding, 57 patients had a surgical bleeding and 2 patients had an unknown type of bleeding as it was not described further in the operation description.

Table 1: Risk factors for reoperation

Variables	Reoperated (n = 101), mean \pm SD or number (%)	Not reoperated (n = 1351), mean \pm SD or number (%)	<i>P</i> -value
Age (year)	67.0 \pm 11.6	66.1 \pm 10.6	0.414
Sex (male)	73 (72.3)	984 (72.8)	0.908
Priority (acute)	15 (14.9)	125 ^a (9.3)	0.079
EuroSCORE	12.2 \pm 13.1	8.3 \pm 11.7	0.001
EF	48.5 \pm 12.9	51.3 \pm 11.8	0.022
Procedure: isolated CABG	25 (24.8)	564 (41.7)	0.001
Time on ECC (min)	158.3 \pm 83.9	126.7 \pm 57.6	0.000
BMI (kg/m ²)	25.4 \pm 4.0	27.1 \pm 4.4	0.000
DM	25 (25.8)	199 ^a (14.7)	0.010
Hypertension	51 (50.5)	596 ^a (44.4)	0.255
Previous AMI	31 (30.7)	395 ^a (29.3)	0.821
Previous PCI	16 (15.8)	176 ^a (13.1)	0.447
Medicine within 5 days	23 (22.8)	302 (22.4)	0.902
Creatinine >134 μ mol/l at admission	17 (16.8)	105 ^a (7.8)	0.004

^aNumber (n) different from 1351 due to missing data. Results are calculated with another 'n'. Priority, DM, creatinine and epidural, n = 1350, hypertension, n = 1341; previous AMI, n = 1346; previous PCI, n = 1340.

Table 2: Comparison of case and match group

Variables	Reoperated (n = 101), mean ± SD or number (%)	Match (n = 101), mean ± SD or number (%)	P-value
Priority (acute)	15 (14.9)	15 (14.9)	1.000
Age (year)	67.0 ± 11.6	67.0 ± 11.6	0.990
BMI (kg/m ²)	25.4 ± 4.0	25.6 ± 4.2	0.711
EuroSCORE	12.2 ± 13.1	13.2 ± 16.1	0.647
Medicine within 5 days	23 (22.8)	23 (22.8)	1.000
Sex (male)	73 (72.3)	72 (71.3)	1.000
EF	48.5 ± 12.9	49.1 ± 12.4	0.710
DM	25 (24.8)	17 (16.8)	0.225
CABG	59 (58.4)	58 (57.4)	1.000
Aortic valve	55 (54.5)	42 (41.6)	0.091
Mitral valve	11 (10.9)	17 (16.8)	0.309
Other procedure	33 (32.7)	22 (21.8)	0.114
S-creatinine (µmol/l)	109.8 ± 39.4	97.6 ± 24.2	0.008

Table 3: Comparing the outcome between reoperated and match group

Variables	Reoperated (n = 101), Mean ± SD or number (%)	Match (n = 101), Mean ± SD or number (%)	P-value
TIA ^a	2 (2.0)	0 (0.0)	0.129
Stroke ^a	0 (0.0)	3 (3.0)	
Lacunar infarct ^a	3 (3.0)	1 (1.0)	
Sternal infection	0 (0.0)	1 (1.0)	1.000
AMI			
Yes	4 (4.0)	4 (4.0)	0.929
Uncertain	4 (4.9)	3 (3.0)	
Mortality	16 (15.8)	6 (5.9)	0.040
Drainage during the admission at intensive care unit (ml) ^b	2905.3 ± 2332.9	829.9 ± 626.3	0.000

^aNumber different from 202; reoperated 99, match 100.

^bNumber different from 202, 91 in both groups.

Patients undergoing re-exploration had higher mediastinal drainage in the intensive care unit (2905.3 versus 829.9 ml). Furthermore, the mortality rate was higher in the re-explored group (15.8 versus 5.9%) (Table 3).

To analyse the impact of re-exploration on the renal function, we excluded patients with preoperatively elevated s-creatinine (s-creatinine > 134 µmol/l at admission). This reduced the group of re-explored patients from 101 to 82 patients and the control group was reduced to 97. The increase in postoperative s-creatinine is greater for the re-explored when compared with the control group ($P = 0.001$). The maximum value of s-creatinine is also different in the two groups (160.4 versus 126.1 µmol/l) (Table 4).

Table 4: Comparing s-creatinine between reoperated and match

Variables	Reoperated (n = 82), mean ± SD	Match (n = 97), mean ± SD	P-value
S-creatinine admission (µmol/l)	96.1 ± 16.8	94.3 ± 16.4	0.476
S-creatinine max	160.4 ± 89.4	126.1 ± 47.3	0.001
S-creatinine discharge/ death	107.5 ± 39.2	103.5 ± 32.8	0.458
Average increase Difference discharge-admission	64.3 ± 84.2 11.5 ± 33.1	31.8 ± 44.2 9.2 ± 28.0	0.001 0.626

Patients with creatinine above 134 at admission were excluded from this comparison. Furthermore, two patients are excluded due to missing data.

Table 5: Comparison of reoperated survivors and non-survivors

Variables	Non-survivors (n = 16), mean ± SD or number (%)	Survivors (n = 85), mean ± SD or number (%)	P-value
Sex (male)	12 (75.0)	61 (71.8)	1.000
Age (year)	70.8 ± 5.8	66.3 ± 12.3	0.151
EuroSCORE	19.9 ± 20.0	10.8 ± 10.9	0.010
BMI (kg/m ²)	26.7 ± 5.1	25.1 ± 3.9	0.166
EF	43.8 ± 12.1	49.3 ± 13.0	0.114
DM	5 (31.3)	20 (23.5)	0.535
Priority (acute)	3 (18.8)	12 (14.1)	0.702
Medicine <5 days	4 (20.0)	19 (22.4)	0.756
ECC-time (min)	216.7 ± 92.7	147.3 ± 77.9	0.002
S-creatinine admission (µmol/l)	127.4 ± 42.5	106.5 ± 38.2	0.052
S-creatinine max ^a	226.6 ± 84.1	172.4 ± 105.4	0.081
S-creatinine increase ^a	97.0 ± 66.8	68.6 ± 92.3	0.290
Type of bleeding (surgical)			0.256
Isolated CABG	3 (18.8)	22 (25.9)	1.000
Drainage ^b	2049.3 ± 1135.1	1783.5 ± 1049.6	0.376
Time to reoperation (min) ^c	561.4 ± 331.1	406.4 ± 275.4	0.061

^a97 patients; 13 non-survivors and 84 survivors.

^b95 patients; 15 non-survivors and 80 survivors, drainage until reoperation.

^c99 patients; 14 non-survivors and 85 survivors.

Surviving patients in the re-explored group had a lower EuroSCORE than non-survivors ($P = 0.010$). Additionally, a significant difference in the re-explored group is a longer ECC time. On average, the surviving patients were re-explored 155 min earlier than the others (561.4 versus 406.4 min) (Table 5).

One surgeon had a greater risk for re-exploration compared with the other surgeons (surgeon 12, $P = 0.003$) (Fig. 3), but no difference in mortality was seen among the surgeons.

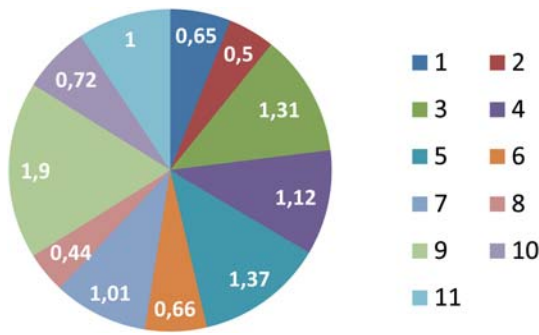


Figure 3: Relative risk for reoperation with respect to the surgeon. The surgeons' identification numbers are listed to the right. The relative risk is shown in the figure.

DISCUSSION

Of 1452 patients undergoing cardiac surgery, 101 patients (7.0%) required at least one reoperation within 24 h due to excessive bleeding. This rate is greater than previously reported rates of 2.2–4.2% [1–6] and also greater than the target rate of the surgical ward of 5.0%. It has not been possible for the surgeons to stay within the recommended rates over the examined 3-year period, but the rate is no greater than the national average rate for Denmark in the given period estimated by WDHReg [7].

It has been shown that reoperation is associated with an increased mortality [1, 3–6]. Our study confirmed this by showing an increase in mortality by three times (2.7–2.8) for re-explored patients compared with both all non-re-explored and matched control group.

We identified seven significant pre- and intraoperative variables that were predictive for re-exploration due to bleeding: low BMI; low EF; high EuroSCORE; preoperatively increased creatinine; DM; procedures other than coronary artery bypass graft surgery (CABG) and prolonged time on ECC. These variables are intertwined in various ways. Other studies have found age and priority associated with the risk of reoperation, but our study did not show this association [2–4]. We did not find any significant association between a medical history with hypertension or previous AMI/PCI (respectively, acute myocardial infarction and percutaneous coronary intervention) and risk for reoperation. Interestingly we found that patients with DM have a significantly greater risk for reoperation due to bleeding which is not seen in other studies [2–4]. Thus the risk for reoperation seems to depend on the type of preoperative morbidity the patients present themselves with.

Karthik *et al.* [2] studied 2898 patients undergoing CABG with a reoperation rate of 3.1% to identify risk factors for reoperation. Variables such as low BMI, non-elective surgery, increasing number of grafts and increasing age was identified as having a significantly greater risk for reoperation for bleeding. Moulton *et al.* [4] studied 6015 patients undergoing cardiac surgery with a reoperation rate of 4.2%. High age, preoperative renal insufficiency, procedures other than CABG and prolonged time on ECC were at greater risk of needing reoperation for bleeding. Dacey *et al.* [3] studied 8586 patients with a reoperation rate of 3.6%. In this substantial study, high rates of re-exploration for haemorrhage were observed in patients with increasing age, smaller BSA, prolonged time on ECC and number of grafts.

We differentiated the haemorrhage in coagulopathic and surgical to assess whether it had an influence on mortality or not. While 56.4% of the patients had surgical bleeding, the remaining 41.6% were coagulopathic bleeding. Mortality was 19.0% for patients with surgical bleeding, and 10.5% for patients with coagulopathic bleeding; this difference is not significant ($P = 0.256$). Hall *et al.* [1] differentiated the haemorrhage between surgical and coagulopathic to determine the differences in patients' outcome. Of the 2263 patients undergoing cardiac surgery, 3.6% were reoperated due to bleeding. While 66% had a surgical bleeding, the remaining 34% had coagulopathic bleeding. The mortality for reoperated group was 8.7 and 12.5%, respectively, and for non-reexplored it was 2%. Our study did not find the same tendency as Hall *et al.*'s study regarding mortality as our patients with surgical bleedings had a greater mortality than the coagulopathic.

Morbidity could be examined using the propensity-matched score. We did not find a significant difference in morbidity between the re-explored group compared with the propensity-matched group regarding AMI, sternal infection, stroke and the level of creatinine after discharge. This is similar to the findings from Karthik *et al.* [2]. It should be stressed that these postoperative complications are infrequent and a large number of patients are needed to show a difference.

Focusing on renal function we excluded patients in both the reoperated group and the propensity-matched group with preoperative elevated creatinine ($>134 \mu\text{mol/l}$). Having only patients with normal pre-operative creatinine in both groups, we evaluated the level of postoperative rise of s-creatinine, and patients reoperated for bleeding had a greater increase in s-creatinine both average and maximum which was significant. Surprisingly, there was no difference in the values at discharge/death between the two groups.

Among the patients undergoing reoperation the prolonged ECC time was associated with increased mortality. We did not find any significant difference regarding the time to re-exploration between the reoperated survivors and non-survivors, but one could consider that it might make a difference if the patients underwent earlier re-exploration. The average time to re-exploration in the two groups was 561 and 406 min, respectively. This is in conflict with the guidelines regarding postoperative bleeding and reoperation outlining the volume of bleeding up to 5 h (300 min) after the first operation.

CONCLUSIONS

Our aim was to find factors with significant influence on the risk for reoperation due to haemorrhage after cardiac surgery and evaluate the consequence of the reoperation. Like several other studies, we found a higher mortality for reoperated patients. We also found that low EF, high EuroSCORE, procedures other than CABG, the time on ECC, low BMI and preoperatively elevated s-creatinine had a significant influence on the risk for reoperation. These variables are intertwined in various ways. We found a significant association between DM and the risk for reoperation.

By comparing the reoperated patients with a propensity-matched group, we found that the reoperated patients had a significantly greater increase in postoperative s-creatinine.

By analysing the reoperated patients we found that the group of surviving patients had a lower EuroSCORE, a shorter time on the ECC and a shorter time to re-exploration.

The haemorrhage was differentiated in coagulopathic and surgical. In total, 56.4% of the patients had surgical bleeding.

Patients with high EuroSCORE, low EF, low BMI, DM, pre-operative s-creatinine >134 µmol/l and procedures other than CABG should have a very carefully planned operation. Preoperatively, discontinuation of pertinent medication and screening coagulation in blood samples could reduce coagulopathic bleeding. Initiatives such as checklists, action cards, guidelines and regular audits can help reduce surgical causes for reoperation due to bleeding. It is mandatory to strictly follow guidelines regarding reoperation for post-operative bleeding and thereby possibly reduce the amount of time and blood spent before performing a necessary reoperation.

Conflict of interest: none declared.

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- eComment. Postoperative bleeding in cardiac surgery: the issue is not resolved yet**
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- I read with great interest the article by Kristensen *et al.* [1]. They showed an incidence of reexploration for bleeding after cardiac surgery of 7%. Also, it was found that low ejection fraction, high EuroSCORE, procedures other than isolated coronary artery bypass graft surgery (CABG), prolonged time on extracorporeal circulation, low body mass, and others were significant risk factors for reoperation for bleeding. Indeed, operative mortality (15.8%) increased by three times for reexplored patients.
- It is noteworthy that not only the reoperation for excessive haemorrhage per se had a negative impact on operative mortality and morbidity. Christensen *et al.* [2] demonstrated that postoperative haemorrhage exceeding 200 ml/h in any single hour or part thereof, or 2 ml/kg/h for 2 consecutive hours in the first 6 hours after surgery, or > 495 ml in the first 24 hours was associated with a higher 30-day mortality and other major postoperative complications. In fact, death at 30-day after surgery increased from 5.5% to 22.4% in the postoperative haemorrhage group. When postoperative haemorrhage was present, reexploration for bleeding occurred in 50% of the cases. Postoperative haemorrhage was also associated with ICU stays > 72 h, and mechanical ventilation > 24 h. Possible explanations for these circumstances include more blood transfusions, a more hypovolemic status, and systemic hypotension resulting in secondary organ failure, such as prolonged respiratory support, the need for renal replacement therapy, and a higher incidence of systemic inflammatory response syndrome. Vivacqua *et al.* [3] have demonstrated that both greater blood transfusion and reoperation for excessive haemorrhage are independently associated with an elevated risk of mortality and major morbidity.
- Traditionally, the decision to perform re-sternotomy is based on conventional guidelines: drainage of > 500 ml in the first hour, > 800 ml in the first 2 h, 900 ml in the first 3 h, 1000 ml in the first 4 h, 1200 ml in the first 5 h, sudden massive bleeding, or cardiac tamponade. However, the important point here is that postoperative bleeding, according to data mentioned in [2], may carry a higher risk of major complications and early mortality, even when the patient does not require reoperation for bleeding.

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