

Research Article

· Open Access ·

Coronary artery bypass grafting in the octogenarians: should we intervene, or leave them be?

Anil Ozen, Ertekin Utku Unal, Murat Songur, Sinan Sabit Kocabeyoglu, Onur Hanedan, Metin Yilmaz, Basak Soran Turkcan, Ferit Cicekcioglu, Sadi Kaplan, Cemal Levent Birincioglu

Department of Cardiovascular Surgery, Turkiye Yuksek Ihtisas Training and Research Hospital, Kizilay Sok, No. 4 06100 Sihhiye Ankara, Turkey

Abstract

Objective Coronary artery bypass grafting (CABG) is gradually increasing in the elderly population. We aimed to investigate the risk factors and the results of CABG along with the long term survival in patients at an age of 80 and older. **Methods** Between January 2002 and December 2011, a total of 101 consecutive patients at an age of 80 and older who underwent CABG in our hospital were included in the study. The patients were followed and the long-term survival was estimated. **Results** The mean age of the patients was 82.98 ± 2.27 years. Sixty-four (63.4%) were males and 37 (36.6%) were females. Emergency surgery, duration of cardiopulmonary bypass, the intensive care unit (ICU) stay, inotropic support, intra aortic balloon pulsation application, amount of erythrocyte transfusion and fresh frozen plasma transfusion and ventilation period were significantly higher in the patients who died in the hospital. The duration of cardiopulmonary bypass (CPB) was found to be an independent predictor of mortality (OR: 1.18, 95% CI 1.01-1.38, P = 0.034). The in-hospital mortality was 16.8%. Kaplan-Meier analyses revealed a survival ratio of 91.3% at one year, 82.9% at three years and 69.0% at five years. **Conclusions** Patients at the age of 80 and older can be candidates for the CABG procedure bearing in mind that they may have a longer ventilation period and intensive care unit stay. The morbidity and mortality of this age group is considered within an acceptable range. Approaches to minimize CPB, or the choice of off-pump surgery, may be a preventive method to lower the incidence of mortality. Hence, CABG may be performed in this age group with a satisfactory survival ratio.

J Geriatr Cardiol 2015; 12: 147–152. doi:10.11909/j.issn.1671-5411.2015.02.005

Keywords: Cardiopulmonary bypass; Complications; Coronary artery bypass; Mortality; Octogenarians

1 Introduction

Avoiding coronary artery bypass grafting (CABG) in the elderly population is attributed to the increased morbidity and mortality of this age group. The main reasons to avoid surgery in this group are due to low life expectancy and the intolerance to cardiopulmonary bypass which is an important constituent of the operation. Furthermore, most of these patients possess multiple systemic disorders, such as diabetes, renal failure and neurological problems.^[1]

Coronary artery disease presents a rapid increase that is directly proportional to age after the fourth decade. Cardio-vascular disease is frequent over the age of 80.^[2,3] Edmunds, *et al.*^[1] revealed the prevalence of symptomatic cardiac disease at 40% in this age group. The population over the age

Correspondence to: Anıl Ozen, MD, Turkiye Yuksek Ihtisas Training and Research Hospital, Department of Cardiovascular Surgery, Kizilay Sok, No.4 06100, Sihhiye Ankara, Turkey. E-mail: anil_ozen@hotmail.com

 Telephone: +90-533-736-43-43
 Fax: +90-312-310-1791

 Received: November 9, 2014
 Revised: January 7, 2015

 Accepted: January 16, 2015
 Published online: March 12, 2015

of 80 reached 12 million in the USA in 2010, [4] and the expected increase in the aging population in Turkey between 2008 and 2040 is 201%. [5]

Akins, *et al.*^[6] displayed that 87% of the patients over the age of 80 benefited dramatically from CABG resulting in an increase in the quality of life. We aimed to investigate the morbidity and mortality of CABG, along with the long-term survival, in patients at age 80 and older.

2 Methods

The study was approved by the institutional board and the appropriate ethical committee and has been performed in accordance with the ethical standards detailed in the Declaration of Helsinki and its later amendments.

2.1 Patient profile

A total of 101 patients at age 80 and older, who underwent CABG in our hospital between 2002 January and 2011 December were included in the study. In this study, 17 potential confounding variables were studied: age, sex, diabe-

tes mellitus, hypertension, chronic obstructive pulmonary disease, hyperlipidemia, obesity, family history, increased levels of urea and creatinine (not more than 0.3 mg/dL from the basal value of creatinine), carotid artery stenosis, preoperative ejection fraction, emergency surgery, total number of grafts used, surgery type, duration of cross clamp and operation time. We were interested in the following seven outcomes: mortality, length of time in intensive care unit (ICU) and hospital stay, use of inotropics and intra aortic balloon pulsation (IABP), transfusion and need for ventilator support.

Patients were divided into two groups: Group 1 consisting of the patients who died in the hospital and Group 2 including the patients who were discharged from the hospital.

2.2 Surgical technique

Sternotomy was followed by aortocaval cannulation. The site of the aortic cannulation was decided by epiaortic ultrasonography. The plaque free sites were used for aortic cannulation. Cardiopulmonary bypass (CPB) was initiated by cooling the patient down to 32°C. After application of cross-clamp onto the aorta, the heart was arrested using antegrade and retrograde cardioplegia, and then the distal anastomoses were performed. The proximal anastomoses were performed using a site biting clamp following defibrillation of the heart. Fifteen (17.9%) patients underwent beating heart operation due to diffuse atherosclerotic disease of the aorta.

The CPB procedures were similar in most cases, except in the patients undergoing concomittant replacement of the ascending aorta. Axillary artery cannulation was applied for these patients and the distal anastomosis was held under low flow (10 mL/kg per minute) unilateral cerebral antegrade perfusion at 28°C.

Patients with high levels of urea and creatine were hydrated and dopamine was administered (2.5 mg/kg per minute). The perfusion pressure was kept around 70 mmHg and hemofiltration was applied during the surgery when required.

The patients regaining muscular power following surgery without severe left ventricular dysfunction and postoperative hemorrhage were extubated generally within the first eight hours. Early mobilization was enhanced the next day following surgery.

2.3 Statistical analysis

Kolmogorov Smirnov test was used for analyzing the normal distribution of the data. Normally distributed continuous variables were expressed as mean \pm SD, or median values if abnormally distributed. Categorical variables were

expressed as numbers and percentages. Demographic characteristics and perioperative variables were compared using "independent samples t-test" or "Mann-Whitney-U test" for continuous variables and "chi-square test" or "Fisher's exact test" for categorical variables. Potential predictive preoperative and intraoperative significant variables were further analyzed by logistic regression models. Survival analysis was performed by the Kaplan Meier method. SPSS pocket program, version 15.0, was used for statistical analysis. A P value < 0.05 was accepted as significant.

3 Results

The demographic characteristics of the patients are summarized in Table 1. The mean age of the patients was 82.98 ± 2.27 years. A total of 64 (63.4%) were male and 37 (36.6%) were female. Number of patients with a variable number of vessels bypassed are as follows: 17 (16.8%) patients with one vessel, 37 (36.6%) patients with two vessels, 23 (22.8%) patients with three vessels, 24 (23.8%) patients with five or more vessels.

Table 1. Demographic characteristics of the patients.

Age, years	82.98 ± 2.27
Male sex	64 (63.4%)
Diabetes mellitus	16 (15.8%)
Hypertension	44 (43.6%)
Chronic obstructive pulmonary disease	8 (7.9%)
Hyperlipidemia	21 (20.8%)
Obesity	5 (5.0%)
Positive family history	11 (10.9%)
Number of smokers	6 (5.9%)
Raised levels of urea and creatinine	28 (27.7%)
Carotid artery stenosis	8 (7.9%)
Preoperative ejection fraction, %	$50.25\% \pm 9.69\%$

Data are presented as n (%) or mean \pm SD.

The operative and postoperative variables of the patients are given in Table 2. The procedures in addition to CABG were: aortic valve replacement (AVR) in one patient (0.9%), AVR together with supra-coronary graft implantation in two patients (1.9%), carotid endarterectomy together with patch plasty in six patients (5.9%), plication of left ventricular aneurysm together with closure of atrial septal defect in one patient (0.9%), supra-coronary graft implantation in three patient (2.7%), post-infarct ventricular septal defect (VSD) repair (0.9%) in one patient, David I operation in one patient (0.9%), and mitral repair together with atrial septal defect and VSD closure in one patient (0.9%).

Table 2. Operative and postoperative variables of the patients.

Number of bypassed vessels	2.64 ± 1.24		
Cardiopulmonary bypass			
On-pump	84 (83.2%)		
Off-pump	17 (16.8%)		
Duration of cardiopulmonary bypass, min $(n = 84)$	96.39 ± 40.17		
Duration of cross-clamp, $min (n = 84)$	57.41 ± 29.30		
Duration of the operation, min	233.77 ± 77.18		
Intensive care unit stay, days	4.36 ± 11.42		
Inotropic support	40 (39.6%)		
Intra-aortic balloon pump	5 (5.0%)		
Erythrocyte suspension (units)	1.92 ± 1.96		
Fresh frozen plasma (units)	3.30 ± 3.47		
Ventilator support (hours)	43.37 ± 127.68		
Hospital stay (days)	10.80 ± 13.38		

Data are presented as n (%) or mean \pm SD.

Table 3. In-hospital mortality vs. patients discharged.

There were no differences concerning the preoperative data among the patients who died in the hospital and the patients discharged from the hospital (Table 3). However, the percentage of emergency surgery was predominantly higher in the mortality group (P = 0.003). Among the intraoperative factors, the duration of CPB was found to be significantly different between the groups (P = 0.022). There were no differences among the groups concerning the number of bypassed vessels, duration of cross-clamp, and the duration of the operation. The intensive care unit stay (P <0.001), inotropic support (P = 0.001), IABP application (P= 0.003), number of erythrocyte suspension (P < 0.001), and fresh frozen plasma (P = 0.002) transfused and ventilation periods (P < 0.001) were higher in the patients who died in the hospital. CABG with additional procedures compared to the isolated CABG group did not reveal a statistically significant difference in mortality (P = 0.206).

	Discharged $(n = 84)$		In-hospital mortality $(n = 17)$			
	Mean	n (%)	Mean	n (%)	– <i>P</i> -value	
Preoperative characteristics						
Age, yrs	83.05 ± 2.35		82.65 ± 1.84		0.515	
Male sex		55 (65.5%)		9 (52.9%)	0.328	
Diabetes mellitus		13 (15.5%)		3 (17.6%)	0.823	
Hypertension		36 (42.9%)		8 (47.1%)	0.750	
Chronic obstructive pulmonary disease		8 (9.5%)		0 (0.0%)	0.185	
Hyperlipidemia		21 (25.0%)		0 (0.0%)	0.021	
Obesity		4 (4.8%)		1 (5.9%)	1.000	
Positive family history		9 (10.7%)		2 (11.8%)	0.899	
Number of smokers		5 (6.0%)		1 (5.9%)	1.000	
Raised levels of urea and creatine		28 (33.3%)		0 (0.0%)	0.005	
Carotid Artery Stenosis		7 (8.3%)		1 (5.9%)	0.733	
Preoperative ejection fraction, %	50.08 ± 10.08		51.71 ± 5.53		0.984	
Emergency surgery		3 (3.6%)		4 (23.5%)	0.003	
Operative characteristics						
Number of vessels bypassed	2.55 ± 1.20		3.06 ± 1.34		0.126	
Cardiopulmonary bypass duration ($n = 84$)	89.85 ± 33.90		123.40 ± 52.75		0.022	
On-pump		69 (82.1%)		15 (88.2%)	15 (88.2%) 2 (11.8%) 0.540	
Off-pump		15 (17.9%)		2 (11.8%)		
Duration of cross-clamp, min	55.27 ± 28.45		66.86 ± 32.21		0.203	
Duration of the operation, min	225.90 ± 67.12		272.65 ± 109.19		0.111	
Postoperative characteristics						
Intensive care unit stay, days	2.10 ± 3.15		15.53 ± 24.59		< 0.001	
Inotropic support		27 (32.1%)		13 (76.5%)	0.001	
Intra-aortic balloon pump		1 (1.2%)		4 (23.5%)	0.003	
Erythrocyte suspension	1.54 ± 1.43		3.94 ± 2.98		< 0.001	
Fresh frozen plasma	2.55 ± 1.43		7.25 ± 6.93		0.002	
Ventilator support, h	18.33 ± 22.42		174.81 ± 287.89		< 0.001	
Hospital stay, days	8.70 ± 5.96		21.18 ± 28.23		0.147	

Data are presented as n (%) or mean \pm SD.

Further logistic regression analysis revealed that the duration of CPB is an independent predictor of mortality in our patient population (OR: 1.18, 95%CI 1.01-1.38, P = 0.034).

The in-hospital mortality was 16.8%. Postoperatively, twenty-one (21%) patients required prolonged ventilation, eight (8%) underwent surgical re-exploration due to bleeding or pericardial tamponade, eight (8%) developed atrial fibrillation following the operation, eight (8%) developed pulmonary or wound infection, five (5%) had low cardiac output, two (2%) had sternal dehiscence and one (1%) had a permanent pacemaker implantation due to complete AV block.

At follow-up, a total of 98 patients were contacted and three patients remained out of reach. The mean follow up period was 39.8 ± 23.9 months (range 2–108 months). Kaplan Meier analyses revealed a survival ratio of 91.3% in one year, 82.9% in three years and 69.0% in five years (Figure 1), while 74% of the patients were angina free (totally angina-free or Class I according to Canadian Cardiovascular Society grading of angina pectoris) among the survivors.

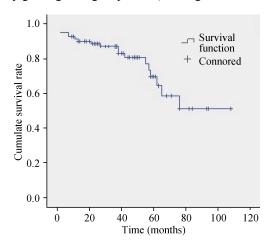


Figure 1. Kaplan-Meier analysis of the patients.

4 Discussion

In this study, we found that octogenarians continue to have higher morbidity and mortality after cardiac surgery than the younger population. Additionally, the duration of CPB was found to be an independent predictor for mortality in the elderly.

Cardiac surgery in octogenarians has risen steadily since the 1980s. The elderly are now the fastest growing population in Western countries, and the number of elderly patients with coronary artery disease potentially eligible for surgery is expected to increase.^[7] Is it cost-effective to operate on these patients? To answer this question, it is important to examine late postoperative survival and functional outcome in addition to morbidity and mortality.

There are three options for treatment of octogenarians with coronary artery disease: medical therapy, coronary intervention, and surgery. It was found that revascularization is superior to optimal medical treatment in those older than 75 years. At present, open heart surgery in octogenarians is more safe and efficient as a result of improvements in myocardial protection, surgical technique, extracorporeal perfusion, anesthetic management and postoperative care. [10,11]

In our study, we found that 86% of our patients required isolated CABG and 14% of patients required additional procedures. Although bypass surgery in elderly patients is associated with favorable outcomes, combined CABG with additional procedures is associated with significantly higher risks. [12] However, CABG with additional procedures compared to the isolated CABG group did not reveal a statistically significant difference in mortality in our study.

Off-pump CABG is known to be less invasive, reduces mortality, the postoperative complications and shortens the postoperative recovery period. [13,14] However, we found that there was no statistically significant difference in terms of mortality between these two groups. As a result of additional procedures and severe atherosclerosis of coronary arteries, our off-pump CABG rate was only 15%.

We used the left internal mammary artery (LIMA) in the majority of the patients. Harvesting the single LIMA in the octogenarians is safe, and the LIMA graft provides better patient outcomes, even in octogenarians.^[15,16]

Mortality in octogenarians after CABG has been reported to range from 8% to 24% and the in-hospital mortality was 16.8%. In our series, however, mortality is higher than what had been observed in the three major previous studies.^[12,17,18] Peterson, *et al.*^[17] reported a hospital mortality of 11.5%. Likosky, *et al.*^[18] and Alexander, *et al.*^[12] observed a mortality of 7.7% and 8.1%, respectively.

It should be emphasized that the long-term survival and functional improvement can be achieved in elderly patients despite the existence of a serious cardiovascular disease. In this study, Kaplan Meier analyses revealed a survival ratio of 91.3% in one year, 82.9% in three years, and 69.0 % in five years. The 3-year survival rates after CABG in other series were between 66.1%–84.5%. [19] Our results were similar to those reports. Salomon, *et al.* [20] demonstrated in a 6-year follow-up period a percentage of angina-free survival of 77% and Mullany, *et al.* [21] in a 5-year follow-up period with 159 patients over 80 years old, observed a percentage of 79. The percentage of angina-free survival in our study was also similar to the one of Solomon and colleagues.

Previous series have reported a high incidence of postop-

erative complications in elderly patients. [22] In this study, the most common complication was prolonged ventilation (21%) followed by arrhythmias (9%). Other complications included postoperative bleeding, infection, low cardiac output and sternal dehiscence. The incidences and nature of postoperative complications in our study are comparable to other reports. [7,23–27] It has been previously reported that octogenarians tend to have a longer postoperative length of stay than younger patients. [23] In this report, the mean intensive care unit length of stay was significantly higher in the in-hospital mortality group.

4.1 Clinical predictors of mortality

For patients of any age, the variables that were independent predictors of mortality in patients following CABG were age, gender, prior CABG, vascular disease, chronic obstructive pulmonary disease, history of congestive heart failure, shock or emergency procedure, ejection fraction, renal insufficiency or dialysis and preoperative myocardial infarct. Previous studies have reported different predictors of mortality in octogenarians undergoing open heart surgery. Akins, et al. [6] identified chronic lung disease and congestive heart failure as independent predictors of mortality, while Edmunds, et al. [1] found that preoperative variables predictive of early death, include New York Heart Association (NYHA) functional classification IV, previous myocardial infarction, and emergency operation. In our study, we found that CPB duration was the only independent predictor of mortality. Besides, in the mortality group, potential predictors such as intensive care unit stay, inotropic support, application of IABP, transfusion and ventilator support time were found statistically higher than the discharged group. However, these parameters should be addressed as a consequence of the procedure rather than as a predictor.

In our study, the hospital mortality was 17 (16.8%). It was cardiac related and associated with co-morbid conditions which were similar to those reported by others. [10,24-30] The main causes of death in our series were due to low cardiac output, respiratory failure and the consequent of infections.

4.2 Limitations

This report is retrospective and describes a series of cases transpiring or occurring operated at a single center. Outcome comparisons are made with literature and are also discussed, since a reliable statistical comparison could not be performed with reports by other authors. Meta-analysis regarding this subject may clarify this issue further. Comparative analyses with other forms of treatment were not carried out, such as drug, or percutaneous treatment. Our off-pump CABG rate was only 15%. Comparison of sur-

vival of surgically treated patients and percutaneously, or medically treated, patients would give more detailed information about the benefits of therapy strategy.

In conclusion, this clinical series provides a profile of octogenarians currently undergoing cardiac surgery and their associated outcomes. Octogenarians represent an operable cohort (or population of patients) despite a longer ventilation period, and prolonged ICU stay. Duration of CPB should be minimized for this age group of patients. The morbidity and mortality of this age group is within an acceptable range. Therefore, CABG can be performed in this age group with a reasonable survival ratio.

Acknowledgement

The authors declare that there is no conflict of interest and this work has not been published before and there is no consideration under publication elsewhere.

References

- Edmunds LH, Stephenson LW, Edie RN, *et al.* Open-heart surgery in octogenarians. *N Engl J Med* 1988; 319: 131–136.
- 2 Hancock EW. Aortic stenosis, angina pectoris, and coronary artery disease. *Am Heart J* 1977; 93: 382–393.
- Assey ME. Heart disease in the elderly. *Heart Dis Stroke* 1993;
 330–334.
- 4 United States Population Pyramid for 2010. Age and sex distribution for the year 2010. http://www.nationmaster.com/country-info/profiles/United-States/People
- 5 Kinsella K, Wan H. International population reports. An Aging World: 2008. http://www.census.gov/prod/2009pubs/p95-09-1.pdf. (Accessed on December 7, 2014).
- 6 Akins CW, Daggett WM, Vlahakes GJ, et al. Cardiac operations in patients 80 years old and older. Ann Thorac Surg 1997; 64: 606–614.
- 7 Speziale G, Nasso G, Barattoni MC, et al. Short-term and long-term results of cardiac surgery in elderly and very elderly patients. J Thorac Cardiovasc Surg 2011; 141: 725–731.
- 8 TIME Investigators. Trial of invasive versus medical therapy in elderly patients with chronic symptomatic coronary artery disease (TIME): a randomised trial. *Lancet* 2001; 358: 951–957.
- 9 Graham MM, Ghali WA, Faris PD, et al. Alberta provincial project for outcomes assessment in coronary heart disease (APPROACH) investigators: survival after coronary revascularization in the elderly. Circulation 2002; 105: 2378–2384.
- 10 Tsai TP, Nessim S, Kass RM, *et al.* Morbidity and mortality after coronary artery bypass in octogenarians. *Ann Thorac-Surg* 1991; 51: 983–986.
- 11 Merrill WH, Stewart JR, Frist WH, *et al.* Cardiac surgery in patients age 80 years or older. *Ann Surg* 1990; 211: 772–775.

- 12 Alexander KP, Anstrom KJ, Muhlbaier LH, *et al.* Outcomes of cardiac surgery in patients > or = 80 years: results from the national cardiovascular network. *J Am Coll Cardiol* 2000; 35: 731–738.
- Hirose H, Amano A, Yoshida S, et al. Off-pump coronary artery bypass: early results. Ann Thorac Cardiovasc Surg 2000; 6: 110–118.
- 14 Amano A, Hirose H, Takahashi A, et al. Off-pump coronary arterial bypass: mid-term results. Jpn J Thorac Cardiovasc Surg 2001; 49: 67–78.
- 15 Morris RJ, Strong MD, Grunewald KE, et al. Internal thoracic artery for coronary artery grafting in octogenarians. Ann Thorac Surg 1996; 62: 16–22.
- 16 Craver JM, Puskas JD, Weintraub WW, et al. 601 octogenarians undergoing cardiac surgery: outcome and comparison with younger age groups. Ann Thorac Surg 1999; 67: 1104–1110.
- 17 Peterson ED, Cowper PA, Jollis JG, *et al.* Outcomes of coronary artery bypass graft surgery in 24,461 patients aged 80 years or older. *Circulation* 1995; 92: 85–91.
- 18 Likosky DS, Dacey LJ, Baribeau YR, et al. Long-term survival of the very elderly undergoing coronary artery bypass grafting. Ann Thoracic Surg 2008; 85: 1233–1237.
- Hirose H, Amano A, Takahashi A. Coronary artery bypass grafting for octogenarians: Experience in a private hospital and review of the literature. *Ann Thorac Cardiovasc Surg* 2001; 7: 282–291.
- 20 Salomon NW, Page US, Bigelow JC, et al. Coronary artery bypass grafting in elderly patients. Comparative results in a consecutive series of 469 patients older than 75 years. J Thorac Cardiovasc Surg 1991; 101: 209–217.
- 21 Mullany CJ, Darling GE, Pluth JR, et al. Early and late results

- after isolated coronary artery bypass surgery in 159 patients aged 80 years and older. *Circulation* 1990; 82: 229–236.
- 22 Cane ME, Chen C, Bailey BM, et al. CABG in octagenarians: Early and late events and actuarial survival in comparison with matched population. Ann Thorac Surg 1995; 60: 1033–1037.
- 23 Fruitman DS, MacDougall CE, Ross DB. Cardiac surgery in octogenarians: Can elderly patients benefit? Quality of Life after Cardiac Surgery. Ann Thorac Surg 1999; 68: 2129–2135.
- 24 Ullery BW, Peterson JC, Milla F, *et al.* Cardiac surgery in select nonagenarians: should we or shouldn't we? *Ann Thorac Surg* 2008; 85: 854–860.
- 25 Williams DB, Carrillo RG, Traad EA, et al. Determinants of operative mortality in octogenarians undergoing coronary bypass. Ann Thorac Surg 1995; 60: 1038–1043.
- 26 Ko W, Krieger KH, Lazenby WD, et al. Isolated coronary artery bypass grafting in one hundred consecutive octogenarian patients: a multivariate analysis. J Thorac Cardiovasc Surg 1991; 102: 532–538.
- 27 Durukan AB, Gurbuz HA, Durukan E, et al. Propensity matched comparison of outcomes of cardiac surgery in octogenarians. *Turk J Geriatr* 2013; 16: 25–30.
- 28 Demir A, Pepeşengül E, Aydinli B, et al. Cardiac surgery and anesthesia in an elderly and very elderly patient population: a retrospective study. Turk Gogus Kalp Dama 2011; 19: 377–383.
- 29 Uncu H, Acipayam M, Altinay L, et al. Is the risk of isolated coronary artery bypass graft surgery in women aged above 75 years higher compared to men? *Turk Gogus Kalp Dama* 2014; 22: 247–252.
- 30 Bridges CR, Edwards FH, Peterson ED, et al. Cardiac surgery in nonagenarians and centenarians. J Am Coll Surg 2003; 197: 347–356.