918 Letters to the Editor

JACC Vol. 39, No. 5, 2002 March 6, 2002:916-21

## REPLY

We appreciate the interest of Dr. Karamanoukian and colleagues in our study about the relation between postoperative stroke in cardiac surgery and presence, location and extent of atherosclerosis in the ascending aorta (1). Our study showed a 3.5% incidence of stroke due to atherosclerosis of the ascending aorta as detected by epiaortic ultrasound (in 26% of the patients) despite minor surgical modifications. Furthermore, a multivariate analysis showed that the two best predictors for perioperative stroke were atherosclerosis of the ascending aorta and diabetes mellitus. Age is usually associated with atherosclerosis in the ascending aorta. Consequently, atherosclerosis of the ascending aorta (detected by intraoperative ultrasound) is a better predictor than age. Therefore, we recommended a more radical change in surgical strategy in the presence of atherosclerosis of the ascending aorta, and especially when the disease of the aortic wall is extended.

One interesting option in this situation is off-pump coronary artery bypass (OPCAB). Still, partial clamping of the ascending aorta is usually used in OPCAB. Clamping of the ascending aorta has been shown to generate cerebral emboli (2–5). Thus, with OPCAB cross-clamping can be avoided, but usually partial clamping is conducted, unless an anastomotic device is employed or the ascending aorta is totally avoided. A further possibility is the use of intraoperative intraaortic filters, if clamping is necessary (5).

We congratulate Dr. Karamanoukian and colleagues for their successful results with OPCAB in octogenarians. Notably, approximately 12% of their patient cohort had atherosclerosis of the ascending aorta, presumably detected by palpation. It is unclear how these patients were handled in terms of proximal anastomosis on the ascending aorta. Considering that their patients received only on average 1.8 grafts, we assume that their conclusion should be that OPCAB incomplete revascularization without touching the ascending aorta may be the preferred operative technique in high-risk patients (i.e., octogenerians).

Recently, we employed OPCAB to achieve complete revascularization in 20 patients with extensive disease of the ascending aorta according to intraoperative ultrasound and totally avoided the ascending aorta in the majority of the patients. None of the 20 patients suffered a perioperative stroke.

In conclusion, we believe that OPCAB techniques may have a justified place in high-risk patients (i.e., with extended atherosclerosis the ascending aorta according to epiaortic ultrasound) in order to prevent perioperative stroke. However, well-designed randomized studies have to be conducted to prove the superiority of OPCAB over conventional coronary surgery.

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## **REFERENCES**

- van der Linden J, Hadjinikolaou L, Bergman P, Lindblom D. Postoperative stroke in cardiac surgery is related to the location and extent of atherosclerotic disease in the ascending aorta. J Am Coll Cardiol 2001;38:131–5.
- Barbut D, Yau FS, Hager DN, et al. Comparison of transcranial Doppler ultrasound and transesophageal echocardiography to monitor emboli during coronary artery bypass surgery. Stroke 1996;27:87–90.
- 3. Barbut D, Lo YW, Gold JP, et al. Impact of embolization during coronary bypass grafting on outcome and length of stay. Ann Thorac Surg 1997;63:998–1002.
- Barbut D, Yao FS, Lo YW, et al. Determination of size of aortic emboli and embolic load during coronary artery bypass grafting. Ann Thorac Surg 1997;63:1262–7.
- Van Boven NW on behalf of the International Council of Emboli Management (ICEM) Study Group. The role of aortic clamp manipulation as a source of particulate emboli generation: risk factors and outcomes of the ICEM study group (abstr). Ann Thorac Surg 2000; 70:1790.

## Cytokine Gene Polymorphisms and Development of CAD Associated With CP Infection

We read with great interest the study by Momiyama et al. (1) in a recent issue of the *Journal*. The study reports that cytokine gene polymorphisms, in particular interleukin-1 (IL-1) polymorphisms, play a role in the development of coronary artery disease (CAD) in patients with *Chlamydia pneumoniae* (CP) infection. Moreover, they found that CP seropositivity was not significantly different between patients with and without CAD. Momiyama et al. (1) suggested that "triggers" such as IL-1 gene polymorphisms could influence the effect of such infectious agents as CP on development of CAD.

Increasing evidence exists indicating that inflammation plays an important role in atherogenesis (2). The hypothesis of infectious agents that might play an important role in the atherogenesis is supported by results of several epidemiologic studies suggesting possible atherogenic potential not only from CP (3) but also from such pathogens as cyotmegalovirus (4), herpes simplex virus (HSV) (5) and *Helicobacter pylori* (6).

However, existing epidemiologic data about the association of some of these pathogens and atherosclerosis are conflicting (7). We support the finding of Momiyama et al. (1) that that CP seropositivity is not associated with CAD by data from 218 consecutive patients undergoing coronary angiography. Blood of all subjects was tested for serum IgG antibodies to CP and for seromarkers of five other pathogens (hepatitis A-virus, *Helicobacter pylori*, HSV, influenza type A and influenza type B). Of the 218 patients, 88 (40.4%) had anti-CP IgG antibodies. The CAD prevalence was 61.4% in CP-seropositive and 66.9% in CP-seronegative patients (p = 0.49). Moreover, seropositivity for each other pathogen (tested in our study) was not associated with CAD.

In contrast, the number of infectious pathogens to which an individual has been exposed (8) ("infectious burden") correlates with prevalence of CAD. Four or more of the six seromarkers tested for particular infections were positive in 48.8% of patients with CAD and in 31.2% of patients in patients without CAD (p = 0.02). Therefore, our data support the results from Momiyama et al. (1) that seropositivity for a particular infectious agent, like CP, represents no predictor of risk for CAD. However, some "triggers," such as cytokine gene polymorphisms or additional exposure to