and in our study hypothermia clearly delayed increased intracranial pressure and thereby prolonged a higher cerebral perfusion pressure ($P=0.01$).

We propose that even in a well-controlled experimental study, and despite prolonged postinstitute control of extracerebral variables, with an adequately severe cerebral focal ischemic insult small animal-to-animal variations will lead to the inevitable death of a certain percentage of the animals. Present treatment modalities are inadequate in preventing this mortality from diffuse brain swelling and herniation. Prolonged moderate hypothermia (31°C) but not mild hypothermia (35°C) postpones intracranial hypertension while the hypothermia lasts, but it does not prevent the delayed intracranial hypertension that leads to herniation and brain death during rewarming in some of the animals.23 The modalities that are effective decrease the cerebral ischemic damage of surviving animals, as expressed by the brain infarct size and cerebral perfusion.

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Statistical Significance Versus Clinical Importance
The article by Walzl et al describes the results of heparin-induced extracorporeal low-density lipoprotein precipitation (HELP) treatment in patients with acute embolic stroke and multi-infarct dementia. The authors conclude that "the apheresis causes an immediate and significant improvement of the hemorheologic pattern, obviously followed by an improvement of clinical symptoms, which has so far not been achieved by any hemorheologically active substance to a comparable degree and time." I seriously doubt the validity of the conclusion on several grounds.

1. It is not clear how bias was controlled in selecting the control group of 30 patients from the 96 who were assigned to the non-HELP group by the computer. More importantly, no attempt was made to blind the outcome evaluators. This is particularly important in assessing disability (as opposed to death), where measurement is more prone to bias. Thus, a possibility exists that the results are influenced by bias in the selection and/or the measurement processes of the study.

2. Statistical significance does not equal clinical importance. It only tells us that the observed differences in the outcome are unlikely to be due to "chance alone." This may be a useful first step in evaluating the importance of the observed differences, but it is important to examine the clinical importance of the observed differences. We should keep in mind that there may be differences between groups that are "statistically significant" but "clinically meaningless." The treatment resulted in a 3.6- to 6.3-point difference in the Barthel Activities of Daily Living Scale score and a 1.8- to 2.9-point difference in the Mini-Mental State Examination (MMSE). There is no evidence to suggest that these represent clinically important differences. According to the Barthel Index scale, the minimum change that can be recorded is 5 points, and this represents just one level of change in only one of the items. In view of this, most clinicians would not consider a 6-point difference as clinically important. Similarly, in the MMSE, a 2- to 3-point difference may not mean improvement in even one item for several of the items. Therefore, it is doubtful that this will be considered clinically important.

To summarize, the study demonstrates a statistically significant but clinically insignificant difference. This is not to say that the scales are meaningless but to emphasize that their use requires careful consideration of their properties. One good example of its proper use is the multicenter United Kingdom trial of nimodipine in acute stroke, in which the Barthel Index score of 60 or more was used to define "independence," which was their primary outcome. This cutoff score was obviously irrelevant in the study by Walzl et al, because they included only cases with a score of 84 or more. One wonders what the authors expected to achieve in patients with a score of 84 or more, but a cutoff to meet their expectation could have been chosen. In the jargon of measurement theory, the scale has well-established "discriminative" but less validated "evaluative" properties. Some authors consider simple methods of assessing outcome as equally valid and reprimand the use of stroke scales. Even though this position seems extreme, it serves as a useful reminder to select outcome measures that are (1) clinically relevant and meaningful; and (2) reliable, valid, and sensitive for the purpose of the study.

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Association of Abdominal Aortic Aneurysm and Carotid Artery Stenosis
We read with interest the article by Karanja et al that identified the association of aortic aneurysms with carotid artery disease. We have also identified this association in a previously published article that to our knowledge is the only other article to do so. The authors identified aortic aneurysms in 20% of patients with carotid stenosis greater than 30% as measured by duplex ultrasound. The criteria that the authors used, however, defines an aortic aneurysm as any focal dilation of the vessel greater than 2.5 cm. Although a measurement of this diameter may be considered atheromsyral if a saccular dilatation is identified, it does not appear evident from the article. The subcommittee on reporting standards for arterial aneurysms has defined an aneurysm as a permanent
localized (i.e., focal) dilation of an artery having at least a 50% increase in diameter compared with the expected normal diameter of the artery in question. A recent article has further clarified the size of the normal infrarenal aorta based on size and age. The average size of the infrarenal aorta for an average-size man aged 65 to 75 years is 1.87 to 1.95 cm. The normal aortic size is approximately 2 mm smaller for women.

Based on this we reported our data, using a diameter of 3.0 cm for the definition of an aneurysm. We identified aneurysms in 8.4% of patients presenting to us for carotid duplex evaluation, and this increased to 11% in patients with high-grade stenosis of the internal carotid artery. If the authors had used this size for an aneurysm, the incidence of associated aneurysms would have been 13.5%, which is similar to our own results.

Our study was performed to determine the value of screening a cohort of patients with carotid disease for the presence of an aortic aneurysm. A screening examination (i.e., mammography) is performed to identify an occult, unsuspected abnormality. If one is trying to determine the efficacy of a screening exam for aortic aneurysms, patients with palpable or suspected aneurysms based on a physical examination should be excluded from the analysis. We eliminated patients in our study who had palpable aneurysms on physical examination. If we had included these patients, our incidence of associated aneurysms would have been higher and similar to that of Karanjia et al. They report that eleven of their patients had palpable masses, which if pulsatile would surely have been expected to be an aneurysm.

We believe this study confirms an association between carotid artery disease and abdominal aortic aneurysms. We advocate the use of ultrasound to screen asymptomatic patients for the presence of an aneurysm in those with elevated systolic velocities in the internal carotid artery.

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Response
We thank Drs Bays and Carty for their comments regarding our recent article in respect to their own investigation. Independent studies reporting generally similar data are always rather gratifying phenomena. As pointed out by these investigators, the difference between the two studies relates primarily to the size criteria used for abdominal aneurysm. This is an area of some controversy. In our study, we defined the size of an aneurysm as 2.5 cm, with clear focal dilatation present as judged by the radiographer. The correspondents imply that we were vague on the latter point, but we would refer them again to our "Subjects and Methods" section. Although many centers consider 3.0 cm the lower limit for the definition of aneurysm, other investigators argue for aneurysmal criteria even smaller than ours. Indeed, there is evidence that even the smallest aneurysms increase in size, possibly at a rate faster than that of larger aneurysms. We would also point out that some published criteria for aneurysm size restrict this term only for focal dilatation greater than twice the expected size in that individual, a definition that would exclude the 3.0- to 3.8-cm aneurysms of the correspondents' study. We believe, and we suspect the correspondents would agree, that the presence of even small aortic dilatations should prompt some clinical concern and should be followed at the least with serial ultrasounds.

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World Wide Web at:
http://stroke.ahajournals.org/content/25/7/1527.1.citation

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