Ultrasound morphology of carotid lesions in retinal ischaemia

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Abstract
The extracranial carotid arteries of 165 patients with retinal ischaemic symptoms were examined with duplex ultrasound. Both the degree of stenosis and the morphological appearance of the lesions were examined. Ipsilateral carotid artery disease was found in 88% of patients. Degrees of stenosis of between 50–99% were found in 33% of patients, the majority of these lesions being complex heterogeneous in nature. Stenosis of <50% was found in 40% of arteries. The majority of lesions causing <20% stenosis were homogeneous in nature. However, in stenosis of 20–49%, 63% of lesions were found to be complex heterogeneous. A similar distribution of lesions was found in the contralateral carotid arteries. It is important to examine both ipsilateral and contralateral arteries, and to evaluate the morphological characteristics of all lesions causing >20% stenosis.

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A relationship between retinal ischaemia and carotid artery disease was first established in 1856, following autopsy findings of carotid artery thrombosis ipsilateral to retinal ischaemia. This relationship has been confirmed subsequently. Diagnostic studies of carotid arterial disease traditionally concentrate on identifying the degree of stenosis caused by atherosclerotic lesions, considering only those which significantly reduce blood flow or perfusion pressure as being potentially clinically important.

It has been shown in patients with amaurosis fugax that there is an increased risk of stroke, blindness and early death in those found to have carotid artery lesions causing even less than 50% stenosis compared with those patients with no evidence of carotid disease on ultrasound examination, indicating that retinal and cerebral ischaemic symptoms can be caused by carotid lesions previously considered too small to be of clinical significance. There is evidence that more attention should be given to the ultrasound morphological appearance of atherosclerotic lesions in order to determine their potential clinical significance. This factor may be as important in producing symptoms as is the amount of stenosis caused by the disease.

The purpose of this report is to give the results of analysis of carotid arterial lesions examined by duplex ultrasound in 165 patients presenting with symptoms of retinal ischaemia. Both the ipsilateral and contralateral carotid arteries have been examined. The atherosclerotic morphological characteristics and degree of stenosis found have been classified in order to try and identify clinically significant disease.

Material and methods
Between 1985 and 1989, 165 patients (101 male, 64 female, mean age 58 years) were referred to the Diagnostic Blood Flow Laboratory of the Vascular Medicine Unit from an ophthalmic centre for duplex ultrasound examination of the extracranial carotid system. Prior to referral definite retinal ischaemic disease was established on the basis of their ophthalmic assessment, which included funduscopic examination, retinal photography, or fluorescein angiography.

Amaurosis fugax was diagnosed in 79 patients, and 38 patients with permanent unilateral vision loss were found to have central retinal artery occlusion. Branch retinal artery occlusion was found in 48 patients, 17 of whom had partial field defects. Cholesterol emboli were identified in 31 of these patients.

The associated clinical conditions of these patients were: stroke, with a permanent neurological deficit, seven; cerebral transitory ischaemic attack five; peripheral vascular arterial disease 17; ischaemic heart disease 36. In addition, 62 patients were being treated with anti-hypertensive medication and six patients were on medication for control of diabetes mellitus.

An ATL MK5 duplex ultrasound system with a 7.5 MHz imaging facility combined with a 5 MHz Doppler system was used to examine the extracranial carotid arteries. The examinations were recorded on VHS video tape and hard copies made on a Mitsubishi Printer.

The presence or absence of atherosclerotic lesions was established, and where disease was found the degree of stenosis caused was noted together with a description of the morphological appearance of the lesion. This appearance was described as being either homogeneous or heterogeneous.

Homogeneous lesions had an even echo pattern

Figure 1 Complex heterogeneous lesion containing a low echo pool (p), without surface irregularity (s). (v) indicates vessel wall and arrowhead direction of blood flow.
throughout the lesion, and heterogeneous lesions had a mixture of low level and high level echo patterns within the lesion (Figs 1, 2, 3). Heterogeneous structures were further subdivided into simple and complex groups. Simple heterogeneous structures had a smooth surface and a more or less equal distribution of low level and high level echoes. Complex structures displayed low echo pools within the lesions and may also have had an irregular or pitted surface at the interface to flowing blood. The carotid arterial atherosclerosis was classified in terms of degree of stenosis caused and by the morphological characterisation of the lesions.

STATISTICAL ANALYSIS
The chi² method of statistical analysis was used throughout.

Results
Atheromatous lesions were found to be present in the ipsilateral arteries of 88% of the patients presenting with retinal ischaemia (Table 1). Degrees of stenosis of between 50–99% were found in 33% of patients, and a further 5% of patients presented with total occlusion of the internal carotid artery. Lesions causing less than 50% stenosis were found in 40% of ipsilateral carotid arteries.

The distribution of homogeneous and heterogeneous lesions in ipsilateral and contralateral arteries is shown in Table 2.

There was a significantly greater incidence of heterogeneous than of homogeneous lesions in both ipsilateral and contralateral arteries (p<0.001). The incidence of complex heterogeneous lesions was significantly greater than simple heterogeneous lesions (p<0.001); however, there was no difference in incidence of lesions between the ipsilateral and contralateral arteries.

Table 3 shows the relationship between degrees of stenosis and morphological structure. A homogeneous structure was found predominantly in lesions of less than 20% stenosis.

In stenosis of more than 50% all lesions were found to be heterogeneous and of these 82% were complex. In stenoses of 20–49% the majority of lesions were heterogeneous and, of these, 63% were complex in nature.

Discussion
The association between severe carotid artery disease and retinal ischaemic events has been well documented. In this study, the relationship between lesion morphological ultrasound characteristics and patient symptoms was investigated. Lesions were classified as being homogeneous or heterogeneous according to well established criteria.

Heterogeneous lesions were further subdivided into simple or complex appearances, the latter either containing low echo pools or having an irregular surface pattern. The degree of stenosis caused by the lesion was also established. Image quality and reproducibility are dependent on machine characteristics. In this study the same duplex ultrasound system was used for all scans.

A significant number of patients presenting
with retinal ischaemia, whose carotid artery disease might not have been considered clinically significant in terms of degree of stenosis, were found to have complex heterogeneous lesions. This suggests that complex heterogeneous lesions may be potential sources of emboli even when occupying less than 50% of the arterial lumen.

Several studies have compared the ultrasound characterisation of atherosclerotic plaques with their histological examination. Homogeneous plaques have been shown to consist of predominantly fibrous material, and have a poor association with symptom production. Heterogeneous plaques have been found to contain either intraplaque haemorrhage or lipid products, and while ultrasound cannot distinguish between these materials, both have been strongly associated with ischaemic symptoms.28, 29

Considerable variation has been reported in the accuracy of ultrasonic diagnosis of ulceration.30 However, as it is the acoustic surface of the lesion that is demonstrated by ultrasound, it is more accurate to describe the characteristics of the plaque surface rather than use the term ulceration, which has a specific pathological definition.

Both ulcerative plaques and lesions with surface irregularities have a reported association with ischaemic symptoms.31, 32

In a recent study of 108 patients with suspected carotid atherosclerosis, it was found that it is the heterogeneous nature of the lesion, rather than its size, that is associated with clinical symptoms. Duplex ultrasound of the carotid arteries did not reveal a correlation between degree of stenosis and hemispheric symptoms. However, significant correlation was found between heterogeneous plaques and hemispheric symptoms. The authors noted that the majority of plaques causing >50% stenosis were heterogeneous in nature.

A strong association between complex heterogeneous lesions and retinal ischaemic symptoms has been found in this study. There is a known association between severe stenosis and the production of symptoms, and lesions causing severe stenosis were usually found to be complex and heterogeneous in nature.

In this study of patients presenting with symptomatic retinal ischaemia, 40% of patients were found to have relatively small degrees of stenosis, but quite a high proportion of these smaller lesions were also complex and heterogeneous in structure. This emphasises the clinical importance of the ultrasound morphology of atherosclerotic lesions.

Clinical management of carotid artery disease is still controversial. The recent MRC European Carotid Surgery Trial report indicates benefit from surgery in those patients presenting with 70-99% stenosis, and does not recommend surgery for less than 30% stenosis. The findings for those with stenoses greater than 30% and less than 70% are still awaited.33 It should be noted that this trial did not take account of lesion morphology.

The role of anti-platelet therapy is widely established, and has been shown to reduce vascular mortality by 15% and non-fatal vascular events by 30% in patients with a history of cerebrovascular or cardiovascular disease.34

The non-invasive nature of ultrasound allows investigation of the contralateral asymptomatic arteries. This study shows an unexpectedly similar distribution of severity of lesions in both the ipsilateral and contralateral carotid arteries. There is a trend towards a higher incidence of more severe lesions on the ipsilateral side, a finding which has been observed by two previous reports.27, 28

A similar morphological pattern of atherosclerotic lesions was also noted on both arterial sides. It has been suggested that the type of carotid disease which gives rise to retinal ischaemic symptoms and that which results in hemispheric transient ischaemic attacks may be different, the former producing smaller emboli which also cause multiple silent infarcts confirmed by computed tomographic scan, with an associated increased incidence of cerebral atrophy. The similarity of distribution of lesions in the ipsilateral and contralateral sides in this study also raises the possibility that silent cerebral events may be occurring without involving the contralateral retina.

For this reason it is recommended that both carotid vessels be examined routinely as there is a potential danger of emboli from the contralateral side affecting either the retina or causing silent cerebral infarction.

All the patients in this study were referred by an ophthalmic centre specifically for carotid ultrasound studies. The results contain valuable information for the ophthalmic management of these patients. As a result the ophthalmologist may be the first professional to receive information about potential risk not only to the eye but also to the heart and brain.

14 Goes E, Janssens W, Maillet B, Freson M, Steyert L, Oestrey M. Tissue characterisation of atheromatous plaques:
25 European Carotid Surgery Trialist Collaborative Group. MRC European Carotid Surgery Trial: interim results for symptomatic patients with severe (70-99%) or with mild (0-29%) carotid stenosis. Lancet 1991; 337: 1235-43.