Patterns of Transcranial Doppler Flow Velocities in Recent Ischemic Stroke Patients

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Abstract

Context: Intracranial atherosclerosis is a common cause of stroke in India. Transcranial Doppler (TCD) provides a noninvasive way to study basal intracranial blood vessels. The Oxfordshire Community Stroke Project (OCSP) classification is a simple clinical stroke classification system that has prognostic significance and has been associated with size and location of the infarct. **Aim:** This study was undertaken to identify patterns of TCD abnormalities in our stroke population particularly in relation to the OCSP classification. **Setting and Design:** A cross-sectional study was conducted at a tertiary care center in South India. **Methods:** Recent nondisabling ischemic stroke patients were studied. TCD was used to insonate bilateral middle cerebral, bilateral anterior cerebral, bilateral vertebral, and basilar arteries. Mean flow velocity was used to define normal or abnormal flow as per standard criteria. **Statistical Methods:** Association between abnormal flow velocities and OCSP classification was studied using Chi-square tests. Univariate and multivariate analysis was performed to determine factors associated with abnormal flow velocities. **Results:** Of the 59 participants studied, 42 (71%; 95% confidence interval [CI]: 57.3-84.7%) had abnormal flow velocities were blunted flow velocities. Anterior circulation flow velocity abnormalities were seen among all OCSP stroke subtypes, but posterior circulation flow abnormalities were associated with posterior circulation infarcts (P = 0.03). **Conclusion:** Intracranial flow velocity abnormalities are frequent among Indian stroke population. Further studies are needed to characterize these abnormalities fully.

Keywords: Blunted flow velocities, Oxfordshire Community Stroke Project classification, transcranial Doppler

INTRODUCTION

Stroke subtypes in the East have been shown to be different from the West. A review of South Asian strokes by Wasay *et al.* reported that among Indians, the most common etiology of stroke was large vessel stroke accounting for 41%.^[1] However, unlike western populations where extracranial atherosclerosis is common, among South Asians, intracranial atherosclerosis is more common, with reported prevalence of 30%.^[1,2]

Transcranial Doppler (TCD) is an important and noninvasive tool to study the flow characteristics of the intracranial basal cerebral blood vessels. It has a sensitivity of 70%–90% and specificity of 90%–95% for detection of intracranial stenosis in the anterior circulation and sensitivity of 50%–80% and specificity of 80%–96% for posterior circulation.^[3]

The Oxfordshire Community Stroke Project (OCSP) classification is a clinical stroke classification system which was derived by Bamford *et al.*^[4] This uses clinical

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information to categorize computed tomography-proven ischemic strokes as total anterior circulation infarcts (TACIs) (hemiparesis/hemisensory loss with higher cortical disturbances and hemianopia), partial anterior circulation infarcts (PACIs) (any two of the three or pure higher cortical disturbance), lacunar infarcts (LACIs) (pure motor or pure sensory or ataxic hemiparesis or clumsy hand-dysarthria), and posterior circulation infarcts (POCIs) (bilateral signs, crossed signs, and isolated homonymous hemianopia). It has been shown to be associated with size and location of the infarct. It has also been used to prognosticate.^[5] Previous studies have shown that distinct vascular abnormalities maybe associated with different stroke subtypes.^[6] If these findings

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are true, then OCSP classification can be used to identify patients for specific therapies or investigations for underlying vascular pathology.

We undertook this study to identify the common patterns of intracranial flow abnormalities in our stroke patients and their association with different stroke subtypes.

METHODS

The study was a retrospective analysis of data collected for a trial of yoga in stroke between December 2012 and September 2014. It was conducted in the stroke outpatient department of a tertiary care institute in the south of India. Patients with recent ischemic stroke (between 1 and 6 months of the ictus) with modified Rankin score ≤ 3 and insonable temporal and occipital windows were included in this study. Patients with serious comorbid cardiovascular or respiratory illness or aphasia were excluded from the study. Baseline data including demographic information, details of comorbid illness, and investigations performed as part of stroke workup including echo and extracranial duplex studies were collected. OCSP classification was performed using clinical and imaging data by a neurologist. TCD studies were performed on all the included patients using 2MHz probe on Rimed, Israeli, four-view software by a trained neurovascular technician using standard techniques. Bilateral middle cerebral arteries (MCAs), anterior cerebral arteries (ACAs), vertebral arteries (VAs), and the basilar artery (BA) were insonated and mean flow velocities documented. Assuming a prevalence of 30%,^[1] estimating the proportion of those with intracranial atherosclerosis using 59 participants and p = 0.05, we were able to achieve a precision of 12%.

The Stroke Outcomes and Neuroimaging of Intracranial Atherosclerosis (SONIA) trial laid down criteria for detection of >50% intracranial stenosis of MCA and vertebrobasilar arteries.^[7] Zhao *et al.* further refined these criteria.^[8] These have been used to classify velocities [Table 1] of participants in our trial as normal or abnormal.^[9,10] Association between abnormal flow velocities and OCSP classification was studied using Chi-square tests. Univariate and multivariate analysis was performed to determine factors associated with abnormal flow velocities. All statistical analyses were carried out using STATA ver. 14.2, StataCorp, Texas, USA.

RESULTS

A total of 59 participants were part of the study. Most (80%) were male. Anterior circulation strokes (OCSP classification of PACI (47%) and LACI (35%)) were the most common type of stroke occurring in 50 participants (85%). Other baseline characteristics are in Table 2.

The flow velocities in both MCA and BA were classified as abnormal according to criteria by Zhao *et al*. Forty-two participants (71%; 95% confidence interval [CI]: 57.3%–84.7%) had abnormal flow velocity in one or more vessels while 10 (17%) participants had abnormal flows in all three vessels. All abnormal velocities were blunted. Multivariate analysis using logistic regression [Table 3] revealed that only smoking was significantly associated with abnormal flow velocities.

Comparisons of the mean velocity of MCA, BA, and VA between the different types of strokes according to OCSP yielded significantly lower mean velocity in BA in posterior circulation strokes as compared to anterior circulation strokes (P = 0.03) [Figure 1]. The other velocities were not significantly different between the groups.

Table 1: Criteria for abnormal velocity				
Vessel	Lower limit of normal velocity (cm/s)	Upper limit of normal velocity (cm/s)		
Middle cerebral artery	30	80		
Vertebral or basilar artery	20	60		

Table 2: Baseline characteristics

Factor	Mean (SD) or <i>n</i> (%) or median (IQR)			
Age (years)	51.2 (10.8)			
Sex (male)	48 (80)			
Hypertension	40 (66)			
Diabetes	13 (21)			
Dyslipidemia	7 (11)			
Smoking	24 (40)			
Previous stroke	1 (2)			
Ischemic heart disease	3 (5)			
Abnormal duplex@	12 (20)			
Abnormal echo#	9 (15)			
Blood pressure				
Systolic	143.3 (23.3)			
Diastolic	87.6 (12.1)			
Fasting blood sugar (mg/dL)	108.8 (34.6)			
Total cholesterol	146.1 (32.6)			
LDL	83.9 (24.3)			
HDL (mg/dL)	36.3 (13.3)			
TOAST classification				
Large artery atherosclerosis	39 (66)			
Small vessel disease	20 (34)			
OCSP classification				
Total anterior circulations infarct	1 (1)			
Partial anterior circulation infarct	28 (47)			
Lacunar infarct	21 (35)			
Posterior circulation infarct	9 (15)			
NIHSS 5 (3, 7)				
mRS	1 (1, 2)			

[#]Left ventricular hypertrophy 6, sclerotic aortic valve 2, anterior wall hypokinesia 1, [@]None of the vessels, namely, bilateral external and internal carotids and bilateral vertebral arteries, insonated had >50% stenosis. LDL=Low-density lipoprotein, HDL=High-density lipoprotein, OCSP=Oxfordshire Community Stroke Project, NIHSS=National Institutes of Health Stroke Scale, SD=Standard deviation, IQR=Interquartile range, TOAST=Trial of ORG 10172 in acute stroke treatment

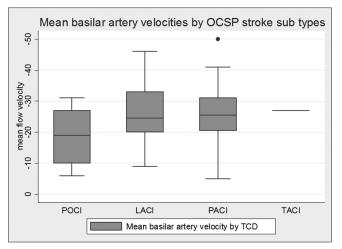


Figure 1: Mean basilar artery flow velocities[#]. [#]*P* value for mean flow velocity in posterior circulation infarcts versus others = 0.03

Table 3: Multivariate analysis for factors associated with abnormal flow velocities

Factor	Adjusted OR	CI
Male sex	0.9	0.2-3.9
Diabetes	0.6	0.1-2.7
Hypertension	1.1	0.3-3.9
Smoking	5	1.2-21.8
Abnormal extracranial duplex	0.8	0.2-3.5

CI=Confidence interval, OR=Odds ratio

DISCUSSION

In our study of recent ischemic strokes, we found that TCD velocity of basal cerebral vessels were abnormal in 72% (95% CI: 57.3%–84.7%) of participants tested and all abnormal velocities were blunted velocities. These blunted flow velocities were significantly associated with smoking. While anterior circulation flow abnormalities had no specific distribution, posterior circulation flow abnormalities were more frequent among patients with posterior circulation strokes.

Sada et al. studied 374 asymptomatic participants with vascular risk factors using TCD. They defined middle cerebral stenosis by velocity criteria given by Feldman et al. from SONIA.^[7] They found a prevalence of 7.2% for intracranial MCA stenosis in the study population.^[11] They, however, did not comment on blunted flow velocities. Our study shows that using velocity criteria, 71% (95% CI: 57.3-84.7) of our cohort of recent nondisabling ischemic strokes had evidence of abnormal flow in at least one of the vessels studied. Among patients with stroke and extracranial carotid disease, Ghosh et al. reported a prevalence of 52.7% for intracranial atherosclerosis using magnetic resonance angiogram (MRA).^[12] Although we found a comparable prevalence of intracranial atherosclerosis, none of our patients had significant extracranial carotid or vertebral stenosis. All our abnormal flow velocities were blunted velocities. The significance of blunted flow velocities is not completely understood. Blunted velocities could signify diffuse atherosclerosis, long-segment stenosis, or multiple distal obstructions. Wijnhoud *et al.* studied 489 strokes or TIA patients followed over 2 years. They found that compared to the second decile of mean flow velocities, the first decile (mean flow velocity of 33.7 cm/s) was associated with a hazard ratio of 10.9 (95% CI: 1.3–88) for recurrent strokes or TIA.^[13] This suggests that blunted flow velocities are not benign.

We found the abnormal velocities to be significantly associated with smoking. Icikay *et al.* showed that mean flow velocity falls with hematocrit and smoking produces rise in hematocrit.^[14] This may explain the association. Another possibility is smoking which is often associated with a diffuse vascular process^[15] and this may also result in blunted flow velocities.

Some previous studies have attempted to explore the relationship between OCSP subtypes and TCD abnormalities. Among 30 patients with stroke, Naylor et al. found MCA occlusion or severe stenosis in all 14 with TACI and 2 with PACI. Distal branch occlusions were seen in 6 of 13 patients with PACI while no abnormalities were seen in patients with LACI.^[16] Mead et al. conducted a study to explore patterns of vascular abnormalities by OCSP subtypes in Edinburgh, UK. They studied 476 patients of whom 36% had PACI while LACI and POCI accounted for 25% and 24%, respectively. They found anterior circulation abnormalities (MCA and ACA) to be significantly more common in patients with TACI and PACI as against POCI and LACI. Similarly, posterior circulation abnormalities were more frequently associated with POCI. They concluded that intracranial abnormalities were significantly related to OCSP subtypes.^[6] Among 699 Chinese patients, Li et al. found abnormalities in MCA and vertebrobasilar velocities among all four OCSP subtypes, with no significant differences between the groups.^[17] In a slightly more diverse South Asian population from Singapore, TCD and MRA were used by De Silva et al. to find that intracranial arterial flow abnormalities were equally distributed among all the OCSP subtypes.^[18]

Our study of recent nondisabling ischemic strokes had a distribution of stroke patients more like Mead *et al.* with 47% of participants with PACI, rather than the Asian studies both of which had predominant LACI subtype. We did not find anterior circulation abnormalities to be associated with any stroke subtype, like the two Asian studies. However, posterior circulation abnormalities were associated with POCI like the study by Mead *et al.* and unlike the Asian studies. We used only mean flow velocities to determine normal or abnormal blood flows while other studies have variably included other parameters including pulsatility index and spectral patterns. While these differences may contribute to these disparate findings, we postulate that the pattern of abnormalities we found reflects the unique Asian Indian stroke phenotype.

Our study has certain limitations. We only included patients if they had nondisabling strokes. Hence, we could include only one TACI. Most of our participants were male probably reflecting the referral bias of a hospital-based population. TCD is a subjective measure, but all the velocities were recorded by a single neurosonographer minimizing interobserver error. We did not look at other TCD parameters such as pulsatility index and spectral patterns, in this study.

CONCLUSION

TCD velocity abnormalities are frequent among Asian Indian stroke patients, with blunted flow velocities being the most common abnormality. More studies are needed with inclusion of wider variety of patients, including other TCD parameters to fully characterize the nature of intracranial vascular disease among Indian stroke patients.

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Conflicts of interest

There are no conflicts of interest.

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