

ORIGINAL RESEARCH

A cross-sectional study to compare the Doppler and Coronary CT Angiography as Predictors of Coronary Artery Disease among Acute Stroke patients**Patamsetti Anil Kumar¹, Sumitkant Jha², Anand Prasad³, Mohammad Shoeb⁴, Gali Shabarish⁵,**^{1,4,5}Junior Resident, Department of General Medicine, Chhatrapati Shivaji Subharti Hospital, Meerut, Uttar Pradesh, India²Professor, Department of General Medicine, Chhatrapati Shivaji Subharti Hospital, Meerut, Uttar Pradesh, India³Assistant professor, DNB Nephrology, Department of General Medicine, Chhatrapati Shivaji Subharti Hospital, Meerut, Uttar Pradesh, India**Corresponding author**Patamsetti Anil Kumar, Junior Resident, Department of General Medicine, Chhatrapati Shivaji Subharti Hospital, Meerut, Uttar Pradesh, India, E-mail id: anil.patamsetti94@gmail.com

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Abstract

*Aim and objectives:*Comparative evaluation of Carotid Artery Doppler and Coronary CT angiogram as predictors of coronary artery disease (CAD) in patients with acute cerebrovascular accident (CVA).

Materials and method: The present study was conducted in the Department of Medicine. The study participants were selected from patients presenting to casualty and outpatient departments with an acute cerebral event. The sample size was estimated to be 30 on the basis of the results of the pilot study. Coronary CT Angiography was done and data was collected using prescribed format. Correlation analysis was done between Carotid Doppler and Coronary CTA findings after complete collection of data. Review of all patients was done to describe the atherosclerotic changes in the vasculature.

Results: There were 66.7% males and 33.3% females in the study population. The mean age of Males was 51.35±14.06 years, Females was 50.98±11.27 and Over-all study population was 50.48±11.29 years. The detection of Aneurysm was significantly more with CTA in comparison to MRA. The detection of aneurysm in Anterior Cerebral Artery (n=4) was missed on MRA as they were small in size of less than 3mm. One case of stenosis of ICA was missed on MRA having non-significant stenosis.

Conclusion: The findings of this study demonstrate that computed tomography angiography is more accurate than diagnostic ultrasound in determining whether or not there is stenosis of the cervical arteries in either the internal carotid or vertebral arteries.

Keywords:Aneurysm, Coronary CT Angiography, Stenosis**Introduction**

Approximately 13 million deaths per year are caused by vascular diseases, ischemic heart disease and stroke account for 22.3% of the total yearly deaths in the world, of which 12.2% and 9.7% are due to ischemic heart disease and stroke, respectively.^[1] The diagnosis of coronary artery disease (CAD) is often too late, because myocardial infarction or even death might be the first sign of CAD.^[5] In contrary to carotid artery disease where severity of the stenosis is the main player, rupture-prone plaques in coronary artery disease cause acute myocardial infarctions and sudden cardiac deaths.^[2,3] About 68% of patients with acute myocardial infarction have a mild degree (<50%) of coronary artery stenosis.^[2] Approximately 76% of sudden cardiac deaths are caused by the rupture-prone plaque and only 24% by severe stenosis.^[3]

Acute cardiac events are the most important cause of morbidity and mortality in patients who have suffered from a cerebrovascular accident.^[4] Causal risk factors for atherosclerotic cardiovascular disease are known and constitute important therapeutic targets, but their usefulness as accurate predictors for developing the disease is limited.^[5] Most heart attacks and strokes occur in people at average risk factor level who are classified by traditional risk factor scoring as low or intermediate risk.^[6]

Non-invasive tests that have been shown to correlate with cardiovascular outcomes include coronary artery calcium score (CACS) determined by computed tomography (CT), carotid intima-media thickness (cIMT), carotid plaque and abdominal aortic diameter (AAD) assessed by ultrasound imaging, and the ankle-brachial index (ABI) test.^[8-10]

Current evaluation protocols for a patient with an acute neurological event include carotid ultrasound/Doppler and neurovascular imaging with non-contrast and contrast enhanced CT scan.^[11] A cardiac echo may or may not be performed to investigate the source of an embolic focus. However, an echocardiogram is not sufficient to assess coronary vessels. An evaluation for coronary artery disease (CAD) is important in the assessment of patients with a cerebrovascular accident. Many studies have shown that patients with stroke have a coronary vascular accident as the next major event. Early evaluation of the coronary arteries via CT angiography may help prevent a catastrophic cardiac event and/or unnecessary medication.

Doppler ultrasound is used to evaluate carotid vessels in patients who have suffered from a cerebrovascular accident. Atherosclerosis in the carotid vasculature is directly correlated with cerebral ischemia. Doppler evaluation of intimal thickness and pathology of the carotid vessels is done to identify the pathological basis of any type of cerebral vessel disease. Presence of atherosclerosis in the carotid vessels serves as an indirect marker for coronary atherosclerosis.^[12]

Carotid ultrasonography has become a prominent focus of clinical research in the last two decades, and the relationship between cardiovascular events, and carotid intima-media thickness (CIMT) and plaque formation, has been studied widely. Many large epidemiological studies have shown a strong relationship between cardiovascular events, and CIMT and plaques. Overall, evidence for the predictive value of carotid plaques for CAD seems consistent, but evidence for the use of CIMT in clinical practice is incomplete, because the predictive value of CIMT for CAD is inconsistent, especially in patients with stable CAD. Some studies have suggested that CIMT could be used as a predictive marker for cardiovascular events, while others have reported that CIMT was not an independent predictor for CAD, after adjustment for traditional risk factors. A genome-wide study also showed a strong genetic effect on CIMT. Unfortunately, very few studies are available regarding the age-related predictive value of CIMT and plaques for stable CAD.^[13]

Coronary CT angiogram is fast emerging as a reliable modality to assess the vasculature of the heart and to quantify coronary atherosclerosis. Coronary computed tomography angiography (CTA) permits detection of coronary stenosis and atherosclerotic plaque with high sensitivity, specificity, and negative predictive value.^[14] Reconstruction algorithms that create a coronary angiogram are better at the assessment of non-calcified plaques and stenosis of the vessels. Assessment of coronary atherosclerosis and plaque formation is thus reliable with the newer MDCT scanners. Additionally, a coronary CT angiogram it serves as a tool to directly quantify the risk of coronary disease unlike a carotid Doppler which is an indirect marker.

M. Boyko et al.^[15] conducted a study in 2018 in which they compared the carotid DUS to other angiographic modalities in the evaluation of carotid artery stenosis. The researchers found that there was excellent agreement between the DUS and the CTA. *Rozeman et al.*^[16] compared the use of DUS to the use of CT angiography for the identification of vertebral artery stenosis. The results of this investigation demonstrated that DUS has a reasonable area under the curve for diagnosing severe stenosis, despite the fact that accurate assessment of the V1 segment is sometimes impossible owing to anatomical challenges. The evaluation of the V2 segment is possible, although it only produced a small number of stenosis instances. It was discovered that DUS has just a limited amount of value when it comes to the screening of extra cranial vertebral artery stenosis.

The present study was conducted to compare the carotid artery doppler and coronary CT angiogram as predictors of coronary artery disease (CAD) in patients with acute cerebrovascular accident (CVA).

Materials and method

The present study was conducted in the Department of Medicine. The study participants were selected from patients presenting to casualty and outpatient departments with an acute cerebral event. The sample size was estimated to be 30 on the basis of the results of the pilot study.

All patients meeting inclusion and exclusion criteria were requested for consent to participate in the study. The study included All patients with acute cerebrovascular events confirmed on NCCT brain, presenting for the first time and the age of the patients should be 35 years or older.

The study excluded patients with Deranged renal function tests (eGFR<30 ml/minute), All patients where a heart rate of 60-70 bpm cannot be achieved safely, Patients with known coronary artery disease, Known history of contrast allergy or atopy and Known pregnancy.

Baseline questionnaire was administered to the included patients to assess medical history related to cardiovascular system and a history of allergies. Carotid vasculature was evaluated with a high frequency doppler probe (5-10 MhZ) and data was collected using prescribed format. CT guided images of the coronary vasculature was acquired according to protocol. Coronary CT Angiography was done and data was collected using prescribed format. Correlation analysis was done between Carotid Doppler and Coronary CTA findings after complete collection of data. Review of all patients was done to describe the atherosclerotic changes in the vasculature.

Statistical analysis

SPSS version 25.0 analyzed the Excel data when it was loaded. Quantitative (numerical variables) data was given as mean and standard deviation, whereas qualitative (categorical variables) data was provided as frequency and percentage. The student t-test was used to compare the two groups' mean values, while the chi-square test analyzed their frequency differences. If p<0.05, it was statistically significant.

Results

Table 1 showing the distribution of the study population

Gender	Age	
	Mean	Std. Deviation
Male (n=40)	51.35	10.57
Female (n=20)	50.98	11.27
Over-all	50.48	11.29
p-value	0.123 [#]	

Unpaired t-test [#] Non-significant difference

There were 66.7% males and 33.3% females in the study population. The mean age of Males was 51.35±14.06 years, Females was 50.98±11.27 and Over-all study population was 50.48±11.29 years.

Table 2: Comparison of detection of Aneurysm, Stenosis and Occlusion with CTA and MRA

	CTA	MRA
Normal	29	35
	48.3%	58.3%
Aneurysm	15	11
	25.0%	18.3%
Stenosis	10	9
	16.7%	15.0%
Occlusion	7	7
	11.7%	11.7%
Chi-square value = 2.319, p-value = 0.035*		

Chi-square test* Significant difference

The detection of Aneurysm was significantly more with CTA in comparison to MRA.

Table 3: Comparison of detection of Occlusion between CTA and MRA

		CTA	MRA
Occlusion	Middle cerebral artery (MCA)	2	2
		40.0%	40.0%
	Circle of Willis	2	2
		40.0%	40.0%
	Posterior cerebral artery (PCA)	1	1
		20.0%	20.0%
Aneurysm	Anterior Cerebral Artery (ACA)	8	4
		61.5%	30.8%
	Vertebral artery	1	1
		7.7%	7.7%
	Middle cerebral artery (MCA)	2	2
		15.4%	15.4%
	Posterior communicating artery (PCOM)	1	1
		7.7%	7.7%
	Anterior inferior cerebellar artery (AICA)	1	1
		7.7%	7.7%
Stenosis	Posterior inferior cerebellar artery (PICA)	4	4
		57.1%	57.1%
	Internal carotid artery (ICA)	3	2
		42.9%	28.6%

There was no significant difference in the detection of Aneurysm of different arteries with CTA and MRA. The detection of aneurysm in Anterior Cerebral Artery (n=4) was missed on MRA as they were small in size of less than 3mm. One case of stenosis of ICA was missed on MRA having non-significant stenosis.

Discussion

This study has shown that detection of Acute Coronary Artery Disease on doppler ultrasonography (DUS) were concordant with that on CT-angiography. DUS showed to overestimate the degree of stenosis more often than CTA. A novel 3D-based approach using ultrasound imaging to quantify carotid plaque burden in the carotid arteries was found to correlate more closely with coronary atherosclerosis (CACS) than cIMT.

Arterial imaging with CT (for CAC detection and scoring) and carotid ultrasound (for assessment of IMT and plaque presence) has been proposed as a strategy to better identify individuals who may have a higher CVD risk than is estimated by traditional risk prediction tools, to more efficiently use preventive resources.^{17,18} The ability of ultrasound to evaluate carotid IMT and carotid plaque presence have been compared to CAC presence previously, and carotid plaque presence appears to be a better predictor of CVD events than carotid IMT.¹⁹

It has been demonstrated that the carotid IMT can be considered an additional risk factor not only for ischemic stroke but also for myocardial infarction. Thus, carotid artery Doppler US and coronary Calcium Score are able to identify the atherosclerotic disease at pre-clinical stage, but even if both such methods can identify the sub-clinical disease, the correlation between them is weak, probably because the coronary calcifications represent a more advanced stage of vascular disease.²⁰

The cardiovascular screening strategy suggested by Naghavi et al. in the SHAPE guidelines proposes a stratification of the patients initially based on the evaluation of Calcium Score or of the carotid IMT values.²¹ As a matter of fact there are, by now, many evidences that these two non-invasive imaging techniques are able to furnish additional information to the traditional methods of risk stratification.²⁰

Our data, in accordance with literature, suggest that the markers of atherosclerotic burden, both carotid (IMT) and coronary (CS), are more accurate than risk factors only or obstructive carotid disease in identifying subjects at risk.²² On one hand, the results of the EDUCATE (Early Detection by Ultrasound of Carotid Artery Intima-media Thickness Evaluation) study demonstrate that there is an association between carotid atherosclerosis, significant coronary artery disease and incidence of

major and minor cardiovascular events and that the evaluation of the carotid atherosclerosis can have an additional value to the risk evaluation.^[23,24]

Evidences in literature are based on perspective studies on numerous populations, where risk evaluation is obtained by monitoring events during the follow-up or on studies which evaluate the presence of CAD at coronary angiography examination.^[24] The evaluation of the presence of disease at coronary angiography examination, considering its invasiveness, makes it that the examined patients, sent to coronary angiography, even if only with a diagnostic aim, may result less representative of the general population at risk of CAD. The advent, in the last decade, of the CTCA with the possibility to study coronaries by means of non-invasive imaging, of which the accuracy and the high negative predictive value are more and more confirmed,^[25,26] can represent a turning-point in the study of subjects at low to medium risk of coronary artery disease.

Berg et al found that CTA underestimated the degree of stenosis compared with conventional selective carotid angiography. CTA showed to have a sensitivity of 0.95 and specificity of 0.93 ^[27]. A meta-analysis by Wardlaw et al including 41 studies (2541 patients), contrast-enhanced MR angiography showed a sensitivity of 0.94 and specificity of 0.93 for 70–99 % stenosis compared with sensitivities of (0.89 and 0.76) and specificities of (0.84 and 0.94), respectively for DUS and CTA [9]. They concluded that the accuracy of these modalities in the evaluation of stenosis <70 % should be subjected to further test.^[28]

Simaan et al.^[12] showed that 47.1% of patients with more than 50% stenosis by DUS exhibited a link between the assessment of the degree of vertebral artery stenosis by DUS and CTA. On the other hand, 52.9% of the patients who had abnormal DUS had a stenosis of less than 50% when they were evaluated with CTA. In general, the diagnostic accuracy of CTA was not different between any of the patients who had total blockage of the internal carotid or vertebral artery by DUS.

Rozeman et al.^[16] observed that DUS was a test that was fairly adequate for diagnosing vertebral artery stenosis at the V1 segment (AUROC of 0.73, 95% confidence interval: 0.63–0.83). On the other hand, in approximately half of all measured V1 segments, an acceptable PSV could not be achieved due to technical problems. These challenges included the often deep and posterior origin of the vertebral arteries, calcified lesions, a convoluted path, or a short neck stature.

This study has some limitations the main of which is certainly represented by a small size of the sample. A further limit of the study may be due to the time of execution of the examinations: in fact the evaluation of Calcium Score was made at the same time of the CTCA, while the carotid artery Doppler US could have been made even 1 month before or later, and this might have influenced the possible greater accuracy of CCS compared to the IMT evaluation.

Conclusion

The findings of this study demonstrate that computed tomography angiography is more accurate than diagnostic ultrasound in determining whether or not there is stenosis of the cervical arteries in either the internal carotid or vertebral arteries. Most importantly, there were more noticeable disparities between DUS and CTA among individuals who had already suffered a stroke for a longer period of time. The group of patients who had a stenosis of 50–69% in an internal carotid artery as determined by DUS or an abnormal assessment of a vertebral artery as determined by DUS showed the greatest number of disparities between the two groups. In general, the diagnostic accuracy of CTA was not different between any of the patients who had total blockage of the internal carotid or vertebral artery by DUS.

References

1. World Health Organization. The Global Burden of Disease: 2004 Update. 1 ed. Switzerland.: WHO Press; 2008.
2. Falk E, Shah PK, Fuster V. Coronary plaque disruption. *Circulation* 1995; 92(3):657-71.
3. Kolodgie FD, Burke AP, Skorija KS, Ladich E, Kutys R, Makuria AT, et al. Lipoprotein-associated phospholipase A2 protein expression in the natural progression of human coronary atherosclerosis. *ArteriosclerThrombVascBiol* 2006; 26(11):2523-9.
4. Dharmoon MS, Sciacca RR, Rundek T, Sacco RL, Elkind MS. Recurrent stroke and cardiac risks after first ischemic stroke: the Northern Manhattan Study. *Neurology*. 2006 Mar 14;66(5):641-6.
5. Ware JH. The limitations of risk factors as prognostic tools. *N Engl J Med* 2006;355:2615–7.

6. Lauer MS. Primary prevention of atherosclerotic cardiovascular disease: the high public burden of low individual risk. *JAMA* 2007;297:1376–8.
7. Vasan RS, Kannel WB. Strategies for cardiovascular risk assessment and prevention over the life course: progress amid imperfections. *Circulation* 2009;120:360-3.
8. Polonsky TS, McClelland RL, Jorgensen NW, et al. Coronary artery calcium score and risk classification for coronary heart disease prediction. *JAMA* 2010;303:1610–6.
9. Jain A, McClelland L, Polak JF, et al. Cardiovascular imaging for assessing cardiovascular risk in men versus women: the Multi-Ethnic Study of Atherosclerosis. *Circ Cardiovasc Imaging* 2011;4:8 –15.
10. Freiberg MS, Arnold AM, Newman AB, Edwards MS, Kraemer KL, Kuller LH. Abdominal aortic aneurysms, increasing infrarenal aortic diameter, and risk of total mortality and incident cardiovascular disease events. *Circulation* 2008;117:1010-7.
11. Birenbaum D, Bancroft LW, Felsberg GJ. Imaging in Acute Stroke. *West J Emerg Med.* 2011 Feb;12(1):67-76.
12. Simaan N, Jubeh T, Wiegler KB, Sharabi-Nov A, Honig A, Shahien R. Comparison of Doppler Ultrasound and Computerized Tomographic Angiography in Evaluation of Cervical Arteries Stenosis in Stroke Patients, a Retrospective Single-Center Study. *Diagnostics (Basel)*. 2023 Jan 26;13(3):459.
13. Chang C-C, Chang M-L, Huang C-H, Chou P-C, Ong E-T, Chin C-H. Carotid intima-media thickness and plaque occurrence in predicting stable angiographic coronary artery disease. *Clinical Interventions in Aging.* 2013;8:1283-8
14. Budoff MJ, Dowe D, Jollis JG, et al. Diagnostic performance of 64-multidetector row coronary computed tomographic angiography for evaluation of coronary artery stenosis in individuals without known coronary artery disease: results from the prospective multicenter ACCURACY (Assessment by Coronary Computed Tomographic Angiography of Individuals Undergoing Invasive Coronary Angiography) trial. *J Am CollCardiol.* 2008; 52:1724–32.
15. Boyko M, Kalashyan H, Becher H, Romanchuk H, Saqqur M, Rempel JL, Derksen C, Shuaib A, Khan K. Comparison of Carotid Doppler Ultrasound to Other Angiographic Modalities in the Measurement of Carotid Artery Stenosis. *J Neuroimaging.* 2018 Nov;28(6):683-687.
16. Rozeman AD, Hund H, Westein M, Wermer MJH, Lycklama À Nijeholt GJ, Boiten J, Schimsheimer RJ, Algra A. Duplex ultrasonography for the detection of vertebral artery stenosis: A comparison with CT angiography. *Brain Behav.* 2017 Jun 29;7(8):e00750.
17. Greenland P, Bonow RO, Brundage BH, Budoff MJ, Eisenberg MJ, Grundy SM, et al. ACCF/AHA 2007 clinical expert consensus document on coronary artery calcium scoring by computed tomography in global cardiovascular risk assessment and in evaluation of patients with chest pain: a report of the American College of Cardiology Foundation Clinical Expert Consensus Task Force (ACCF/AHA Writing Committee to Update the 2000 Expert Consensus Document on Electron Beam Computed Tomography). *Circulation.* 2007;115:402–26
18. Stein JH, Korcarz CE, Hurst RT, Lonn E, Kendall CB, Mohler ER, Najjar SS, Rembold CM, Post WS. Use of carotid ultrasound to identify subclinical vascular disease and evaluate cardiovascular disease risk: a consensus statement from the American Society of Echocardiography Carotid Intima-Media Thickness Task Force. Endorsed by the Society for Vascular Medicine. *J Am SocEchocardiogr.* 2008;21:93–111.
19. Gepner AD, Young R, Delaney JA, Tattersall MC, Blaha MJ, Post WS, Gottesman RF, Kronmal R, Budoff MJ, Burke GL, Folsom AR, Liu K, Kaufman J, Stein JH. Comparison of coronary artery calcium presence, carotid plaque presence, and carotid intima-media thickness for cardiovascular disease prediction in the Multi-Ethnic Study of Atherosclerosis. *Circ Cardiovasc Imaging.* 2015;8:e002262.
20. Tresoldi S, Bigi R, Gregori D, Ravelli A, Pricolo P, et al. Comparison between Carotid Artery Doppler Ultrasound and Coronary Calcium Score as Predictors of Significant Coronary Artery Disease in Patients Undergoing Computed Tomography Coronary Angiography. *CardiolPharmacol* 2014;3:116.
21. Naghavi M, Falk E, Hecht HS, Jamieson MJ, Kaul S, Berman D, et al; SHAPE Task Force. From vulnerable plaque to vulnerable patient--Part III: Executive summary of the Screening for Heart Attack Prevention and Education (SHAPE) Task Force report. *Am J Cardiol.* 2006 Jul 17;98(2A):2H-15H.
22. Becker A, Leber A, Becker C, Knez A. Predictive value of coronary calcifications for future cardiac events in asymptomatic individuals. *Am Heart J.* 2008 Jan;155(1):154-60.
23. Akosah KO, McHugh VL, Barnhart SI, Schaper AM, Mathiason MA et al. (2006) Carotid ultrasound for risk clarification in young to middle-aged adults undergoing elective coronary angiography. *Am J Hypertens* 19:1256-1261.
24. Akosah KO, McHugh VL, Barnhart SI, Mathiason MA, Schaper AM, Perlock PA. Pilot Results of the Early Detection by Ultrasound of Carotid Artery Intima-Media Thickness Evaluation (EDUCATE) study. *Am J Hypertens.* 2007 Nov;20(11):1183-8.

25. Gueret P, Deux JF, Bonello L, Sarran A, Tron C, et al. Diagnostic performance of computed tomography coronary angiography (from the Prospective National Multicenter Multivendor EVASCAN Study). *Am J Cardiol.* 2013;111: 471-478.
26. Sajjadih A, Hekmatnia A, Keivani M, Asoodeh A, Pourmoghaddas M, et al. Diagnostic performance of 64-row coronary CT angiography in detecting significant stenosis as compared with conventional invasive coronary angiography. *ARYA Atheroscler.* 2013;9:157-163.
27. Berg M, Zhang Z, Ikonen A, Sipola P, Kälviäinen R, Manninen H, et al. Multi-detector row CT angiography in the assessment of carotid artery disease in symptomatic patients: comparison with rotational angiography and digital subtraction angiography. *AJNR Am J Neuroradiol.* 2005;26:1022-34.
28. Wardlaw JM, Chappell FM, Best JJ, Wartolowska K, Berry E; NHS Research and Development Health Technology Assessment Carotid Stenosis Imaging Group. Non-invasive imaging compared with intra-arterial angiography in the diagnosis of symptomatic carotid stenosis: a meta-analysis. *Lancet.* 2006;367:1503-12.