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# **Research Article**

# The Value of Longitudinal Strain versus Coronary Angiography in Detection of Coronary Artery Disease

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# ABSTRACT

**Aims:** The aim of this study was to evaluate the value and accuracy of longitudinal strain in detection of coronary artery disease compared to coronary angiography.

**Results:** The left ventricular longitudinal strain-speckle tracking showed evidence of stenosis of left anterior descending artery, circumflex artery and right coronary artery in (86.1%), (76.4%), and (84.7%) respectively. For the stenosis in left anterior descending artery, the current study showed that the longitudinal strain was a good predictor for presence of significant stenosis with a sensitivity of (93.8%), specificity (75%) and accuracy (91.7%) compared with coronary angiography. For the stenosis in right coronary artery, the left ventricular longitudinal strain had a sensitivity of (93.5%), specificity of (70.0%) and accuracy of (90.3%) compared with coronary angiography.

**Conclusion:** Speckle-tracking echocardiography technique has a good performance and validity in detection of coronary artery stenosis with good agreement with angiography.

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# Introduction

Coronary artery disease is characterized by atherosclerosis in the epicardial coronary arteries. The reduction in coronary artery flow may be symptomatic or asymptomatic, occurs with exertion or at rest, and culminate in myocardial infarction or angina, depending on obstruction severity and the rapidity of development [1].

Recent studies have suggested that decreased left ventricular (LV) compliance accompanies both coronary artery disease and acute myocardial infarction. Many facilities measure only left ventricular ejection fraction (LVEF) and left ventricular end-diastolic pressure

(LVEDP) when assessing left ventricular dysfunction in patients with coronary artery disease (CAD) [2, 3].

Speckle-tracking echocardiography is a new noninvasive ultrasound imaging technique that allows for an objective and quantitative evaluation of global and regional myocardial function independently from the angle of insonation and from cardiac translational movements [4-7].

**Patients and Methods** 

Study design

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A cross sectional study was conducted in Ibn Al-Bittar Cardiac Surgical Center/ Baghdad. It included a sample of 72 patients with coronary artery disease proved by coronary angiography and then echocardiography for each patient was done. Also, evaluation of risk factors of CAD like smoking, diabetes mellitus, hypertension, lipid profile along with body mass index were taken in consideration.

The study was conducted in compliance with the medical ethics rules and all participants have given their consent. The study protocol was approved by the ethical committee of the department of medicine/ Baghdad medical school.

#### Types of investigations used

All the study subjects were evaluated by treadmill test TMT (which is considered positive TMT if there is ST segment deviation one mm or more in one or more ECG leads and/or if the patient develops ischemic chest pain). Lipid profile, conventional echocardiography & 2D LV Longitudinal strain using speckle tracking analysis by the available 2D strain software (GE Vivid E9) within 1wk of the scheduled coronary angiography.

#### Conventional echocardiography

Conventional echocardiography was done using echocardiography machine (GE Vivid E-9) and phased-array probe with a frequency of 2-4 MHz.

Evaluation of LV systolic function (LV ejection fraction) was done using linear M-mode echocardiography with M-mode cursor at the tips of mitral valve leaflets in parasternal long axis view. Regional wall motion abnormality was assessed using apical long axis view for assessment of anterio-septal and posterior-lateral walls at basal, mid, apical segments; LV 4-chamber view for septal and lateral walls at basal, mid, apical segments; LV 2-chamber view for anterior and inferior walls at basal mid apical segments, correlated with LV short axis view at basal, mid, and apical levels.

#### Two-dimensional speckle tracking analysis

Using available 2D strain software (GE E-9), acquisition of 3 apical views (2-chamber, 3-chamber and 4- chamber) was done. All images were taken at end expiration to avoid LV apical foreshortening and reducing sector width and depth was used to increase frame rate for better image resolution. All images were digitally stored at a high frame rate (>63 frames/s, mean  $70 \pm 12$ ).

Region of interest was localized at LV endocardial border 2 at base, and one apical to include the entire myocardium at end systolic frame. The software algorithm automatically segmented the LV into six equidistant segments and selected suitable speckles in the myocardium for tracking. The software algorithm then tracked the speckle patterns on a frame-byframe basis using the sum of absolute difference algorithm. Finally, the software automatically generated LV strain profiles for each of the six segments of each view, from which end-systolic strain was measured. The average value of strain at each level (basal, middle, and apical) and All measurements were preformed off-line on a dedicated workstation using available 2D strain software (GE E9). The examiner was blinded to patient's results of coronary angiography. Longitudinal strain was computed using 2D speckle-tracking analysis by automated function imaging (AFI). For strain processing, the peak of the R-wave on the electrocardiogram was used as the reference time point for end-diastole and segments with poor-quality tracking were manually discarded. GLS was only computed from patients with >92% of segments adequately tracked ( $\geq$ 15 segments for a 17-segment model). Given the average value of ( $-17 \pm 0.1\%$ ) for normal GLS, any value assessed by AFI less than (-16%) was considered as impaired GLS.

Thorough correlation of LV regional longitudinal impairment with coronary territories was evaluated in all patients, value of regional longitudinal strain less than (-16%) were abnormal and diagnostic of CAD if it involved more than 3 segments in same coronary territories.

#### **Coronary angiography**

Coronary angiography was done for all patient using standard protocol and the diagnosis of CAD was defined if there is stenosis more than or equal to 50% in left main coronary artery (LMS) and/ or if the stenosis was more than or equal to 70% in the right coronary artery (RCA), left anterior descending coronary artery (LAD) and/ or circumflex coronary artery (CX).

#### Statistical analysis

Data of the patients were entered and analyzed using the statistical package for social sciences (SPSS) version 22, IBM Inc., Chicago, USA, 2013. Descriptive statistics were presented as means, standard deviations, ranges, frequencies and proportions according to the variable types. Cross-tabulation and Receiver operating characteristics (ROC) curve analysis was used to assess the validity of the 2D LV Longitudinal strain -speckle tracking in detection of stenosis in different coronary arteries, kappa statistics used to assess the performance and the percent agreement of the 2D LV Longitudinal strain -speckle tracking in detection in comparison with angiography. Level of significance was set at 0.05 to be significant difference or correlation. Finally results and findings were presented in tables, figures and explanatory paragraphs using Microsoft office (Word), 2013 software for windows.

#### Result

The baseline demographic characteristics of the patients are shown in (**Table 1**). Regarding age, the mean age of patients was  $(56.4 \pm 10.1 \text{ yrs})$ , and the mean body mass index (BMI) was  $(27.2 \pm 3.2 \text{ kg/m}^2)$ . Only 20 patients (27.8%) had BMI within normal levels (BMI < 25 kg/m<sup>2</sup>). The smokers represented about half of the patients (51.4%), 53 patients (73.6%) were diabetic, 24 (33.3%) were hypertensive, 29 (40.3%) had pericardial fat and 42 patients (58.3%) had hyperlipidemia.

Table 1: Demographic characteristics of the study population

Characteristic			N= 72
Age (year)		Mean± SD	$56.4 \pm 10.1$
BMI category		Mean± SD	$27.2\pm3.2$
DIG	(Normal)		20 (27.8%)
BMI	(Overweight)		35 (48.6%)
	(Obese)		17 (23.6%)
	(Yes)		37 (51.4%)
Smoking			
	(No)		35 (48.6%)
Diabetes mellitus	(Yes)		53 (73.6%)
	(No)		19 (26.4%)
Hypertension	(Yes)		24 (33.3%)
	(No)		48 (66.7%)
Pericardial Fat	Pericardial Fat (Yes)		29 (40.3%)
(No)			43 (59.7%)
Hyperlipidemia	(Yes)		42 (58.3%)
	(No)		30 (41.7%)

According to the angiographic findings, it had been found that left anterior descending artery (LAD) stenosis was present in 64 patients (88.9%), the circumflex coronary artery (LCX artery) stenosis in 56 patients (77.8%), and the right coronary artery (RCA) stenosis in 62 patients (86.1%) (Table 2).

According to the findings of 2-D LV longitudinal strain-speckle tracking, positive finding of left anterior descending artery (LADA) stenosis was present in 62 patients (86.1%), the circumflex (CX) artery stenosis was found in 55 patients (76.4%) and the right coronary artery (RCA) stenosis was found in 61 patients (84.7%) (Table 3).

**Table 2:** Angiographic findings of the study group (N = 72)

	Stenosis			
Coronary artery	Present		Absent	
	No	%	No	%
Left anterior	64	88.9	8	11.1
descending				
Circumflex	56	77.8	16	22.2
Right	62	86.1	10	13.9

 Table 3: Findings of 2-D LV Longitudinal strain –speckle tracking in study patients

	Stend	Stenosis				
Coronary artery	Present		Absent			
	No	%	No	%		
Left anterior	62	86.1	10	13.9		
descending						
Circumflex	55	76.4	17	23.6		
Right	61	84.7	11	15.3		

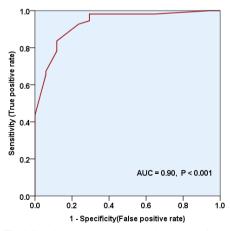
Table 4: Shows the number of stenosed arteries detected by both methods.

Method	Number of diseased arteries	No.	%
Angiography	One vessel	9	12.5
	Two vessels	16	22.2
	Three vessels	47	65.3
2D LV Longitudinal	One vessel	10	13.9
strain	Two vessels	18	25.0
	Three vessels	44	61.1

Table 5: shows the cross-tabulation of the findings of longitudinal strain against angiography regarding the stenosis in left anterior descending artery (LADA); this revealed that longitudinal strain was a good predictor for LADA stenosis with a sensitivity of (93.8%), specificity of (75%) and accuracy of (91.7%). On the other hand, kappa value was 0.82 which indicated good performance of the test and a percent agreement of 91.8% between both tests (P-value < 0.001). Furthermore, ROC curve revealed an area under the curve (AUC) of 0.90 which indicated good predictive value and accuracy of the test (P < 0.001) (Fig. 1).

 
 Table 5: Validity and performance of 2-D LV longitudinal strainspeckle tracking in detection of left anterior descending artery stenosis.

Longitudinal strain	Angio	ography	Total				
	Positi	ive	Negat	Negative			
	No.	%	No.	%	No.	%	
Positive	60	93.8	2	25.0	62	86.1	
Negative	4	6.3	6	75.0	10	13.9	
Total	64	100.0	8	100.0	72	100.0	
Validity test							
Sensitivity			93.	8%			
Specificity			75.	0%			
Accuracy	Accuracy 91.7%						
<i>PPV (positive predictive value)</i> 96.8%							
<i>NPV (negative predictive value)</i> 60.0%							
<i>Kappa</i> = 0.82, H	Percent a	greement	= 91.8%	6, P-value	< 0.001		

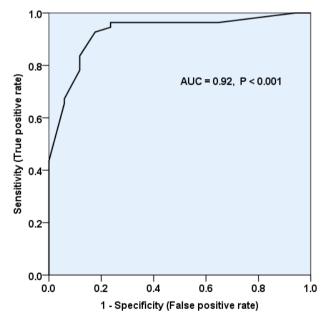


**Fig 1:** Receiver operating characteristics curve for the validity of 2D LV Longitudinal strain -speckle tracking in detection of LADA stenosis.

In (table 6), the longitudinal strain showed good performance in detection of stenosis in LCX artery with a sensitivity of (92.9%), specificity of (81.3%) and accuracy of (90.3%), (*kappa* = 0.82, percent agreement = 90.2%, P< 0.001), (Fig. 2).

**Table 6:** Validity and performance of 2-D LV Longitudinal strainspeckle tracking in detection of left circumflex coronary artery stenosis.

Longitudinal strain	Angi	ography	Total					
	Posit	ive	Nega	tive				
	No.	%	No.	%	No.	%		
Positive	52	92.9	3	18.8	55	76.4		
Negative	4	7.1	13	81.3	17	23.6		
Total	56	100.0	16	100.0	72	100.0		
Validity test								
Sensitivity			92.	.9%				
Specificity			81	.3%				
Accuracy 90.3%								
<i>PPV (positive predictive value)</i> 94.5%								
NPV (negative p	NPV (negative predictive value) 76.5%							
Kappa = 0.82, H	Kappa = 0.82, Percent agreement = 90.2%, P-value < 0.001							



**Fig 2:** Receiver operating characteristics curve for the validity of 2D LV Longitudinal strain -speckle tracking in detection of CXA Stenosis.

Regarding the performance and validity of 2-D LV longitudinal strain in detection of RCA stenosis in comparison with angiography, it had a sensitivity of (93.5%), specificity of (70.0%) and accuracy of (90.3%), (*kappa* = 0.88, percent agreement = 90.2%, P< 0.001), (Table 7), (Fig. 3).

 Table 7: Validity and performance of 2D LV Longitudinal strain –

 speckle tracking in detection of RCA stenosis.

Longitudinal strain	Angio	ography	Total				
	Posit	ive	Nega	Negative			
	No.	%	No.	%	No.	%	
Positive	58	93.5	3	30.0	61	84.7	
Negative	4	6.5	7	70.0	11	15.3	
Total	62	100.0	10	100.0	72	100.0	
Validity test							
Sensitivity			93.	.5%			
Specificity			70	.0%			
Accuracy	90.3%						
PPV (positive predictive value) 95.1%							
<i>NPV (negative predictive value)</i> 63.6%							
Kappa = 0.88, H	Percent a	greement	= 90.29	%, P-value	e < 0.00	1	

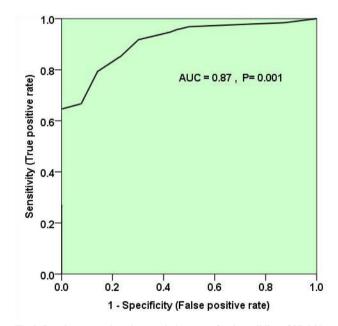


Fig 3: Receiver operating characteristics curve for the validity of 2D LV Longitudinal strain -speckle tracking in detection of CXA stenosis

Additionally, the validity of longitudinal strain in detection of stenosis in different numbers of coronary arteries are shown in (**Table 8**), (*Kappa* = 0.86, Percent agreement = 84.7%, P-value < 0.001), hence, the validity and performance in detection of multiple vessel stenosis were good and the best values reported in detection of three-vessels disease.

Table 8: Validity and performance of 2-D LV Longitudinal strainspeckle tracking in detection of different numbers of stenosed coronary arteries

	Angio	graphy					Tota	ıl
Longitu dinal strain	One v	essel	Two v	Two vessels		5		
Strum	No.	%	No.	%	No.	%	Ν	%
							0.	
One vessel	7	77.8	3	18.8	0	0.0	10	13.9
Two vessels	2	22.2	11	68.8	4	8.5	17	23.6
Three	0	0.0	2	12.5	43	91.5	45	62.5
vessels								
Total	9	100	16	100	47	100	72	100
Validity	One		Two		Thre			
test	vess		vess		e			
	el		els		vesse			
					1s			
Sensitivity	77	.8%	68.89	%	91.5	%		
Specificity	95	5.2%	88.59	%	92.0	)%		
Accuracy	93	.1%	79.2% 91.7%					
PPV(positi	ve		_					
predictive	value) 7(	).0%	64.79	6	95.6	%		
NPV (nega	tive		_					
predictive	value) 90	5.8%	83.69	6	85.2	2%		
Kappa = 0.	86, Perc	ent agre	ement =	84.7%,	P-value	< 0.001		

## Discussion

Speckle Tracking Echocardiography (STE) is an echocardiographic imaging technique that analyzes the motion of tissues in the heart by using the naturally occurring speckle pattern in the myocardium when imaged by ultrasound. This novel method of documentation of myocardial motion represents a noninvasive method of definition of vectors and velocity. When compared to other technologies seeking noninvasive definition of ischemia, speckle tracking seems a valuable endeavor. This speckle pattern is a mixture of interference patterns and natural acoustic reflections [8]. These reflections are also described as speckles or markers. The pattern is being random, each region of the myocardium, has a unique speckle pattern (also called patterns, features, or fingerprints), that allows the region to be traced from one frame to the next, and this speckle pattern is relatively stable, at least from one frame to next [9, 10].

The current study shows that the main age group of patients was between 50-59 years with mean age  $56.4 \pm 10.1$  years, which is the same that was found by Cimino et al., and by Montgomery et al [11, 12]

Smoking is a major risk factor for coronary heart disease which is also found in the present study that shows that more than half of the studied group were smokers; this is consistent with that found by Cimino et al., where about 59% of patients were smokers [11].

In asymptomatic diabetic patients with a preserved LVEF, it has been shown that speckle-tracking echocardiography has the potential for detecting subclinical LV systolic dysfunction, which is unmasked by the alteration of longitudinal strain. In this view, speckle-tracking echocardiography might provide useful information about the development of subclinical myocardial dysfunction in the diabetic setting before the overt appearance of diabetic cardiomyopathy. This evidence confirms previous experiences using either color tissue velocity imaging or doppler-derived strain rate imaging [13]. The current study shows that about two thirds of the participants had diabetes mellitus which is the same finding that was reported by Tsutomu Takagi [14].

The present study shows that about one third of the patients were suffering from hypertension, which in accordance with that revealed by Ji Qiang et al [15].

Regarding the hyperlipidemia, it was found that more than half of the patients had hyperlipidemia, which is less than found by Cimino et al. where more than two thirds of the patients suffer from it. This may be due to that the patients in the present study were diagnosed previously and were on treatment which includes cholesterol lowering medications [11].

For the body mass index (BMI), the current study shows that about half of the patients were overweight and the mean BMI was  $(27.2 \text{ kg/m}^2)$  which is concordant with that revealed by Lipinski, Michael et al., where it was (28.5%) [16].

According to the angiographic findings, it was found that left anterior descending coronary artery (LAD) stenosis was reported in (88.9%), the left circumflex coronary artery (LCX) stenosis reported in (77.8%) and the right coronary artery (RCA) stenosis was found in (86.1%) of the patients while in a study done in USA by Montgomery, David et al <sup>12</sup> the results were: the stenosis in LAD only (21%), in LCX only (9%) and the stenosis in RCA only (7%). Additionally, it was (25%) in stenosis of LAD + LCX, (11%) in LAD + RCA and it was (4%) for the stenosis in LCX + RCA.

In the present study, 2-D LV Longitudinal strain-speckle tracking showed evidence of stenosis of LAD (left anterior descending artery) in (86.1%), LCX artery stenosis in (76.4%) and the RCA stenosis was found in (84.7%) of the patients.

As for the stenosis in LAD, the current study reported that the longitudinal strain was a good predictor with a sensitivity of (93.8%), specificity (75%) and accuracy (91.7%), with good performance of the test between tests, good predictive value and accuracy of the test.

The longitudinal strain showed that there was a good performance in detection of stenosis in LCX with high sensitivity, specificity and accuracy. Regarding stenosis of RCA, 2-D LV longitudinal strain had a sensitivity of (93.5%), specificity of (70.0%) and accuracy of (90.3%). Additionally, this study showed the validity of longitudinal strain in detection of stenosis in different number of coronary arteries, hence, the validity and performance in detection of multiple vessel stenosis were good and the best values were reported in detection of three vessels disease.

Aggeli Constantina et al. reported that the sensitivity for detection of coronary artery disease was 38%, 29% and 28% in LAD, LCX and RCA respectively [17].

The average value of global longitudinal strain (GLS) in this study was  $(-17 \pm 0.1\%)$  for normal GLS, and any value assessed by AFI less than (-16%) was considered as impaired GLS, while the normal GLS was  $(-14 \pm 3.3\%)$  in Cimino et al. study [11].

In a study of 2-D strain imaging, a peak systolic longitudinal strain rate (SR) of (-0.83s<sup>-1</sup>) and an early diastolic SR of (0.96 s<sup>-1</sup>) predicted significant (>70%) stenosis with a sensitivity and specificity of 85% and 64% and 77% and 93%, respectively suggesting the potential for early diastolic deformation to improve diagnostic accuracy [18]. In another study of 108 patients undergoing coronary angiography, a 2-D longitudinal strain of (-17.9%) discriminated severe 3-vessel or left main disease from lesser coronary artery disease with a sensitivity and specificity of 79% and 79%, respectively [19]. Finally, in a cohort of patients with normal ejection fraction at increased atherosclerotic risk and/or with stable chest pain, a progressive impairment of 2-D global strain and SR (the former with lower variability and higher reproducibility than the latter) was directly related to increasing severity of coronary disease as determined from multislice computed tomography. A global longitudinal strain (the average of segmental longitudinal strains) less than (-17.4%) provided high sensitivity and specificity (83% and 77%, respectively) in identifying patients with obstructive coronary disease [20].

According to the angiographic findings, the present study revealed that (12.5%) of the patients had stenosis in one vessel, (22.2%) in two vessels and the main percentage was found in three vessels (65.3%) of patients. This is not consistent with that mentioned by Aggeli et al. where the main percentage was found in one vessel (46%), 14% in two vessels and 7% in three vessels [17]. This may be attributed to the difference in sample selection as in the last study only 67% of the patients had coronary artery disease. On the other hand, the results of the current study are in accordance to that reported by Jiangping et al. in their study to assess the coronary artery stenosis by coronary angiography; it revealed that the main artery stenosis was found in three vessels of coronary arteries (50%), (29.5%) had 1 artery stenosis and (20.4%) had 2 arteries stenosis [21].

In a study carried by Brian D. LV strain has also been studied as an adjunct to dobutamine stress echocardiography (DSE) [22]. In response to a flow-limiting stenosis at rest and a non-flow limiting stenosis during dobutamine infusion in open-chest pigs, longitudinal and circumferential (but not radial) strain was reduced. Radial strain was decreased only in the presence of a flow-limiting stenosis during dobutamine, suggesting that longitudinal and circumferential mechanics are altered earlier than radial mechanics in the ischemic cascade. The incremental value of post-processed 2-D and tissue Doppler longitudinal strain to the stress-echocardiographic diagnosis of coronary artery disease was compared in 150 consecutive patients undergoing dobutamine infusion. Diagnostic accuracy of strain (and wall motion analysis) was similar for ischemia in the left anterior descending territory and similar (but not as good as wall motion analysis) for the left circumflex territory; however, tissue

Doppler strain was more sensitive than either 2-D strain or wall motion analysis in the right coronary territory [22].

In another study, the optimal cutoff values for longitudinal, circumferential, and radial strains at peak dobutamine stress derived from 62 patients with a coronary angiographic reference to detect significant coronary stenoses were 20%, 26%, and 50%, respectively; the diagnostic accuracy derived from an additional 40 patient validation group was 85%, 76%, 70%, and 82% for longitudinal, circumferential, radial strains, and wall motion score index, respectively, and the combination of longitudinal strain and wall motion scoring yielded diagnostic accuracy that was incremental to either alone. In addition, 2-D strain measured in early diastole may increase the accuracy of detecting coronary disease during stress. Delayed LV relaxation detected by radial strain from apical views (transverse strain) during the first third of diastole 5 and 10 minutes after exercise ("diastolic stunning") identified significant coronary stenosis with a sensitivity of 97% and specificity of 93%. These encouraging data suggest that quantitative strain data are sensitive and provide incremental diagnostic information in DSE [22].

#### Limitation of the study

- Selection bias is present in the current study in which only patients with coronary artery disease were included.
- 2. Only one type of the echocardiography equipment (2D strain software GE E9) equipment was used.
- 3. Small sample size with single center study.

# Conclusions

- Speckle-tracking echocardiography is a sophisticated new echocardiographic technique that, working with standard 2dimensional images and devoid of the limitations of Doppler techniques, provides a comprehensive analysis of global and regional myocardial deformation evaluated in all spatial directions.
- 2. Speckle-tracking echocardiography technique shows a good performance and validity in detection of coronary artery stenosis with very good agreement with angiography.

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