# CORRELATION BETWEEN MYOCARDIAL PERFUSION IMAGING AND CAROTID ULTRA-SONOGRAPHY IN PATIENTS WITH CHEST PAIN

By

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# ABSTRACT

**Background:** Atherosclerosis is a major risk factor for coronary artery disease (CAD). Carotid ultrasound (U/S) may be a useful method for prediction of CAD through measurement of intimio-media thickness (IMT) of right and left common carotic arteries.

**Objective:** To emphasize the correlation between myocardial perfusion imaging using single photon emission computed tomography (SPECT) and carotid ultrasound through measuring IMT in early prediction and diagnosis of CAD.

**Patients and methods:** The study included 80 patients complaining from chest pain. Single photon emission computed tomography (SPECT) and carotid U/S were done to each of them. All Patients were recruited from El-Hussein university hospital outpatient clinic and Kobry El-Koba military hospital outpatient clinic between February 2020 and November 2020.

**Results:** The study included 80 patients, 57 of them were males while 23 of them were females. 52 patients were hypertension (HTN), 31 patients were diabetes mellitus (DM), 33 patients were smokers, 15 patients of them had positive family history. 52 patients had positive SPECT stress test revealing CAD "ischemic group", while 28 of them had negative SPECT stress test "Non-ischemic group". For the ischemic group, 43 patients (82.7%) had increased intimio-media thickness IMT. Right IMT mean  $\pm$  SD "0.69 $\pm$ 0.10", t-test "7.771", weighted Kappa test 0.655 substantial agreement and P-value "p <0.001". Lefy IMT mean  $\pm$  SD "0.77 $\pm$ 0.12", t-test "5.122", weighted Kappa test 0.614 substantial agreement and p-value "p <0.001". Only 9 patients (17.3%) from the ischemic group had below 75th percentile IMT. For Non-ischemic group, only 9 patients (32.1%) had abnormal IMT while the other 19 patients (67.9%) had normal IMT.

**Conclusion:** Carotid ultrasound and intimio-media thickness are good predictors and highly correlated to coronary artery disease.

Keywords: SPECT, Carotid U/S, Intimio-media thickness, CAD.

## **INTRODUCTION**

The worldwide health burden has shifted from communicable, maternal and perinatal causes to non-communicable diseases with cardiovascular diseases (CVDs) being the largest single contributor worldwide. CAD is the most common and hazardous cardiovascular disease. Its incidence and the resultant morbidity and mortality increase every year, especially in the elderly (*Rusnak et al., 2017*).

Coronary atherosclerosis can lead to different graded stenosis of coronary

arteries causing decreased blood flow to the myocardium and dysfunction of the affected myocardial segments. Early intervention and detection whether medically or invasively are of utmost value to preserve left ventricular (LV) functions (McPherson systolic and Tybjaerg-Hansen, 2016).

Noninvasive identification of patients with CAD remains a clinical challenge despite the widespread use of imaging and provocative testing; more than 50% of patients currently referred to coronary angiography show normal or nonobstructive CAD (*Ruigómez et al.*, 2010).

Myocardial perfusion imaging (MPI) is a non-invasive imaging test that shows how well blood flows through (perfuses) your heart muscle. It can show areas of the heart muscle that aren't getting enough blood flow. This test is often called a nuclear stress test. It can also show how well the heart muscle is pumping. There are 2 techniques for MPI: single photon emission computed tomography (SPECT) and positron emission tomography (PET). MPI is useful in patients with chest discomfort to see if the discomfort comes from lack of blood flow to the heart muscle caused by narrowed or blocked heart arteries (angina). Myocardial perfusion imaging doesn't show the heart arteries themselves, but can tell your doctor with good certainty if any heart arteries are blocked and how many. MPI can also show if you've previously had a heart attack. A radioactive tracer (called Myoview) is injected into the patient's bloodstream and is taken up by the heart photon tissue. А single emission computed tomography (SPECT) camera detects the radiation released by the tracer to produce images of the heart. Two sets of images are taken, one after an injection at rest and another after an injection during a stress test – either exercise stress on a treadmill or drug-induced stress with persantine (dipyridamole) medication, which simulates exercise (*Fihn et al.*, 2012).

Intima-media thickness (IMT), also called intimal medial thickness, is a measurement of the thickness of tunica intima and tunica media, the innermost two layers of the wall of an artery. The measurement is usually made by external ultrasound and occasionally by internal, invasive ultrasound catheters. Measurements of the total wall thickness of blood vessels can also be done using other imaging modalities (*Go et al., 2013*).

The carotid artery is the usual site of measurement of IMT and consensus statements for carotid IMT have been published for adults and children. Often, IMT is measured in three locations: in the common carotid artery (typically at one cm proximal to the flow divider), at the bifurcation, and in the internal carotid artery. IMT measurements of the far (deeper) wall, by ultrasound, are generally considered more reliable than measurements performed on the near superficial) wall: although (more measurement of both near and far wall IMT has also been advocated (Staub, 2018).

The aim of this study was to identify the correlation between myocardial perfusion imaging using SPECT., and measuring carotid ultra-sonography as

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predictors of presence of coronary artery disease in patients with chest pain.

### PATIENTS AND METHODS

This study included 80 patients of both genders "males and females" complaining from chest pain. The included cases were recruited from the Outpatient clinic and Inpatients at El-Hussein univesity hospital and Kobry El Kobba Military Hospital between February 2020 and November 2020.

**Inclusion criteria:** Adult patients between 18–80 years of age with chest pain.

#### **Exclusion criteria:**

Any patient with:

- 1. Multi-valvular Lesions.
- 2. Severe malignant arrhythmia.
- 3. End organ damage "Renal failure, Liver failure, Metastatic cancer"
- 4. Any contraindication to exercise test.
- 5. Any contraindication to MPI test.

#### All patients were subjected to:

History and physical examination: Upon enrollment all patients were have a review of their medical history, including demographic data (age, gender, body mass index [BMI]), and presence of risk factors for coronary atherosclerosis (smoking, diabetes, hypertension, dyslipidemia, family history of premature CAD) and associated co-morbidities.

#### **Clinical examination:**

**Vital signs:** e.g.: heart rate, blood pressure. Signs of heart failure or hemodynamic instability. Local cardiac examination.

**Twelve-lead ECG:** 12 leads surface electrocardiography to exclude any arrhythmia or the presence of Q waves or ST-T wave changes at rest.

**Laboratory investigation:** Lipid profile (LDL- HDL – TGDs – Cholesterol), kidney function tests.

# Transthoracic Echocardiography analysis:

All echocardiographic examinations were performed after 20-30 min of rest with the patient in quiet respiration in the partial left lateral decubitus position, using 2–4 MHz transducer. and was а accompanied by recording resting electrocardiography. All measurements were obtained online and echocardiographic parameters were measured according to the American Society of Echocardiography Values. For each parameter was obtained by averaging measurements from three successive cardiac cycles. Left ventricularenddiastolic and end-systolic diameters was measured and LV ejection fraction (EF) was calculated by Simpson's rule.

**Carotid arteries U/S Doppler study:** All patients were have carotid U/S Doppler study with measuring carotid intimamedia thickness at the nearest wall and stratified into two groups according to European society of cardiography guidelines.

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Figure (1): Carotid U/S measuring Intima media thickness.

# Myocardial perfusion study using SPECT:

The study was done at "two-day" protocol. In the first day all patients undergo exercise stress on treadmill according to Bruce protocol. Then patients were injected with radioactive tracer and

subjected to gamma camera for the stress film. The next day patients were subjected to gamma camera for the rest film and then the 2 films were analyzed by SPECT analysis system.



Figure (2): Real patient SPECT stress/rest film after analysis.

For each patient LV EF was calculated and ischemic areas either reversible or scared tissues was detected.

#### **Statistical Analysis:**

Recorded data were analyzed using the statistical package for social sciences, version 20.0 (SPSS Inc., Chicago, Illinois, USA). Quantitative data were expressed as mean± standard deviation (SD). Qualitative data were expressed as frequency and percentage. Independent-samples t-test of significance was used when comparing between two means. Chi-

square (x2) test of significance was used in order to compare proportions between qualitative parameters. Kappa measure of agreement between Technicium scan and Carotid duplex, a value of: 0 - 0.20indicates slight agreement; 0.21-0.40, fair agreement; 0.41 -0.60. moderate agreement; 0.61-0.80, substantial agreement; and 0.81-1.00, almost perfect agreement. The confidence interval was set to 95% and the margin of error accepted was set to 5%. So,

#### RESULTS

This was a prospective study conducted on 80 patients to demonstrate the correlation between ischemic heart disease and atherosclerotic changes at carotid arteries.

The mean age of the patients was  $57.50\pm8.77$  years (range 39-76 years). There were 23 (28.8%) female and 57 (71.3%) male of demographic data. This table shows that there were HTN 52 (65.0%), DM 31 (38.8%), Smoking 33 (41.3%), CA or PCI PCI 15 (18.8%), CA 5 (6.3%), CABG 2 (2.5%), PCI LAD 1 (1.3%), PCI, CABG 1 (1.3%), Post covid 1 (1.3%), Family Hx 15 (18.8%), Urea 38.28±12.05 and Creat 1.12±0.50 of history & Laboratory data (**Table 1**).

Table (1): Distribution of patients with chest pain according to their demographic data regarding gender, age, HTN, DM, CA or PCI, family history, urea and creat (n=80). (n=80)

Demographic data	Total (n=80)
Gender	
Female	23 (28.8%)
Male	57 (71.3%)
Age (years)	
Range	39-76
Mean±SD	57.50±8.77
HTN	52 (65.0%)
DM	31 (38.8%)
Smoking	33 (41.3%)
CA or PCI	
CA	5 (6.3%)
CABG	2 (2.5%)
No	55 (68.8%)
PCI	17 (21.2%)
Post covid	1 (1.3%)
Family Hx	15 (18.8%)
Urea	26-112 [38.28±12.05]
Creat	0.6-4.2 [1.12±0.50]

#### **Ischemic and Non-Ischemic patients:**

Comparison between Ischemic and non-ischemic patients "by SPECT results" according to demographic data, history, lab & Echo results.

This table shows statistically significant higher male in Ischemic group (80.8%) compared to non-Ischemic group (53.6%), with p-value <0.05 S; while age (years) insignificant difference. This table statistically shows significant higher Ischemic smoker in group (50%)compared to non-Ischemic group (25%), with p-value <0.05 S; while HTN, DM,

CA or PCI, family, urea and creat. insignificant difference.

This table shows statistically significant increase mean in Ischemic group compared to non-Ischemic according to TGDs and LDL, while HDL significant decrease mean in Ischemic group compared to non-Ischemic group.

This table shows highly statistically significant higher SWMA in Ischemic group (80.8%) compared to non-Ischemic group (10.7%), with p-value <0.001 HS; while EF% significant decrease mean in Ischemic group compared to non-Ischemic group (**Table 2**).

			1	
Demographic data	Ischemic	Non-Ischemic	p-value	
	(n=52)	(n=28)		
Gender				
Female	10 (19.2%)	13 (46.4%)	0.010*	
Male	42 (80.8%)	15 (53.6%)	0.010*	
Age (years)				
Mean±SD	56.65±9.00	59.07±8.25	0.242	
Range	39-76	43-76	0.242	
History & Lab data				
HTN				
No	20 (38.5%)	8 (28.6%)	0.27(	
Yes	32 (61.5%)	20 (71.4%)	0.376	
DM				
No	30 (57.7%)	19 (67.9%)	0.252	
Yes	22 (42.3%)	9 (32.1%)	0.373	
Smoking	×			
No	26 (50.0%)	21 (75.0%)	0.000	
Yes	26 (50.0%)	7 (25.0%)	0.029*	
CA or PCI				
CA	4 (7.7%)	1 (3.6%)		
CABG	3 (5.8%)	0(0.0%)		
PCI	12 (23.1%)	4 (14.3%)		
PCILAD		1 (3.6%)	0.235	
Post covid		1 (3.6%)		
No	34 (65 4%)	21 (75.0%)	-	
Family Hy	54 (05.470)	21 (73.070)	+	
No	40 (76 9%)	25 (80 3%)		
Vos			0.177	
Lizza	12 (23.170)	3 (10.7 /0)		
Moon+SD	20 71+14 12	25 61+6 12		
	26 112	35.01±0.12	0.147	
Croat	20-112	20-34		
Moon SD	1 19 0 50	1.02+0.21		
Mean±SD	1.18±0.59		0.166	
Kange	0.0-4.2	0.7-1.4		
		1		
Lipid profile				
Cholesterol				
Mean±SD	216.90±42.71	201.75±36.09	0.115	
Kange	113-301	154-275		
TGDs				
Mean±SD	240.12±56.21	209.36±42.18	0.013*	
Range	124-376	134-289		
LDL				
Mean±SD	117.75±19.49	107.68±17.66	0.026*	
Range	77-154	73-145		
HDL				
Mean±SD	34.62±4.99	36.89±5.27	0.041*	
Range	26-45	26-47		
Echo				
SWMA				
No	10 (19.2%)	<b>25 (89.3</b> %)	<0.001**	
Yes	42 (80.8%)	3 (10.7%)		
EF%				
Mean±SD	48.29±13.22	63.07±5.79	0.001 **	
Range	25-73	51-75	<0.001**	

# Table (2): Comparison between Ischemic and non-Ischemic according to<br/>demographic data, history & Lab data, lipid profile and echo

#### Comparison between Ischemic and nonischemic patients according to results of Carotid U/S:

This table shows highly statistically significant higher mean in Ischemic group compared to non-Ischemic group according to carotid duplex regarding RT IMT and Lt IMT. This table shows highly statistically significant higher in Ischemic group (82.7%) compared to non-Ischemic group (32.1%) according to RT IMT normality, weighted Kappa test 0.655 substantial agreement, while LT IMT normality in Ischemic group (82.7%) compared to non-Ischemic group (35.7%), weighted Kappa test 0.614 substantial agreement, about carotid duplex. This table shows statistically significant higher in Ischemic group (21.2%) compared to non-Ischemic group (3.6%) according to presence of atherosclerotic plaque (**Table 3**).

Table (3): Comparison between Ischemic and non-Ischemic groups according to<br/>carotid U/S results regarding RT IMT, Lt IMT, RT IMT normality, LT<br/>IMT normality and presence of a plaque

Carotid duplex	Ischemic	Non-Ischemic	p-value
	(n=52)	(n=28)	
RT IMT			
Mean±SD	0.69±0.10	0.58±0.06	<0.001**
Range	0.5-1.1	0.5-0.7	
Normal RT			
Mean±SD	0.60±0.04	0.60±0.02	0.799
Range	0.5-0.62	0.53-0.62	
Lt IMT			
Mean±SD	0.77±0.12	0.66±0.09	<0.001**
Range	0.56-1.1	0.5-0.9	
Normal Lt			
Mean±SD	0.67±0.05	0.61±0.25	0.134
Range	0.57-0.7	-0.64-0.7	
Carotid duplex			
<b>RT IMT normality</b>			
Abnormal	43 (82.7%)	9 (32.1%)	<0.001**
Normal	9 (17.3%)	19 (67.9%)	
LT IMT normality			
Abnormal	43 (82.7%)	10 (35.7%)	<0.001**
Normal	9 (17.3%)	18 (64.3%)	
Presence of a plaque			
No	41 (78.8%)	27 (96.4%)	0.036*
Yes	11 (21.2%)	1 (3.6%)	

#### DISCUSSION

This was a prospective study conducted on 80 subjects of both genders "males and females" complaining from chest pain. The included cases were recruited from the outpatient clinic and inpatients at El Hussein univesity hospital and Kobry El Kobba military hospital between february 2020 and november 2020.

Many studies evaluated the role of carotid U/S in detection of arterial

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atherosclerosis through measuring of intima media thickness and detection of any atheromatous plaques at carotid arteries; however a limited number of researchers in literature were discussing the correlation between stress cardiac imaging & viability tests and carotid U/S.

So, we aimed in this study to emphasize on the role of carotid U/S in early detection of ongoing atherosclerotic process and its correlation to diagnosis of coronary artery disease through myocardial perfusion imaging through SPECT.

There was a statistically significant higher number of males with positive stress imaging through SPECT "Ischemic group" (80.8%) compared to negative stress imaging "non-Ischemic group" (53.6%), with "p-value <0.0".

There was statistically significant higher number of smokers in Ischemic group (50%) compared to non-Ischemic group (25%), with "p-value <0.05".

Increased IMT was considered abnormal if more than "75th percentile per age and sex" of right and left carotid IMT by *Yano et al.* (2016).

There was statistically significant higher mean IMT in the "ischemic group" than mean IMT in the "non-ischemic group".

For Ischemic group, right IMT range was 0.5-1.1 with Mean  $\pm$  SD  $0.69\pm0.10$  while normal 75th percentile range was 0.5-0.62 with Mean  $\pm$  SD  $0.60\pm0.04$ .

For Ischemic group, left IMT range was 0.56-1.1 with Mean±SD  $0.77\pm0.12$  while normal 75th percentile range was 0.57-0.7with Mean±SD  $0.67\pm0.05$ .

For Non-ischemic group, right IMT range was 0.5-0.7 with Mean±SD 0.58±0.06 while normal 75th percentile range was 0.5-0.62 with Mean±SD 0.60±0.04.

For Non-ischemic group, left IMT range was 0.5-0.9 with Mean±SD 0.66±0.09 while normal 75th percentile range was 0.64-0.7with Mean±SD 0.61±0.25.

There was 43 patients with increased IMT out of 52 patients in the "ischemic group" with percentage 82.6% with (P-value <0.001\*\*) weighted Kappa test 0.614 substantial agreement.

There were only 9 patients with increased IMT out of 28 patients in the "non-ischemic group" with percentage 32.1%.

This agreed with Mitevska et al. (2017) who had 60 asymptomatic DMT2 patients (pts) without known coronary artery disease (CAD) underwent one day rest Dypiridamole stress Tc-99m sestamibi single photon emission computed perfusion tomography myocardial imaging (MPI). 18 (31%) patients had myocardial ischemia. Mild ischemia was found in 6 pts, moderate in 7 patients and severe ischemia in 5 patients. Carotid IMT was increased in 34 pts and 15 pts had carotid plaques. Mean c-IMT value in patients with normal MPS results was 0.7  $\pm$  0.1: in moderate ischemia 0.9  $\pm$  0.1 and in pts severe ischemia  $1.0 \pm 0.2$ .

This also agreed with *Djaberi et al.* (2010) who found that Increased CIMT was significantly related to the presence and extent of abnormal myocardial perfusion. Assessment of CIMT may be useful to identify asymptomatic patients

with type 2 diabetes at higher risk for CAD.

Overall, 34 patients (35%) showed abnormal perfusion (SSS  $\geq$  3), including severely abnormal perfusion (SSS  $\geq 8$ ) in 14 patients (14%). Average CIMT was  $0.68 \pm 0.12$  mm. Comparison with reference values revealed normal CIMT in 60 patients (61%), while in the remaining 38 patients (39%) an increased CIMT value was observed in at least one CCA. Abnormal perfusion was present in 9% of patients with normal CIMT versus 75% of patients with increased CIMT. Notably, prevalence of severely abnormal perfusion increased from 3% in patients with normal CIMT to 28% in those with increased CIMT with P value (P < 0.001).

It also agreed with *Boucher et al.* (2010) who conducted repeated Doppler measurements of the CIT of primitive carotid arteries in 52 diabetic patients aged 49 to 75. All these asymptomatic patients, without coronary antecedents, diabetic for more than 10 years and exhibiting at least one cardiovascular risk factor, had undergone myocardial scintigraphy.

He found that CIMT was greater in the group with positive scintigraphies and is the best predictive factor of the presence of ischemia. For a CIT value < 0.55 mm, the negative predictive value of the CIT was of 77% with 80% sensitivity; in non-smokers the predictive negative value increased to 92% with 95% sensitivity.

# CONCLUSION

Myocardial perfusion imaging through SPECT is a useful method of diagnosis of CAD. Carotid U/S is a good predictor of atherosclerosis. Carotid Intimio-media thickness is a good predictor and highly correlated to coronary atherosclerosis and CAD.

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## AHMED A. H. ABD EL-AZIZ et al.,

العلاقة بين تصوير نضح عضلة القلب والتصوير فوق الصوتي السباتي في المرضى الذين يعانون من ألم في الصدر أحمد عبد العزيز حسن عبد العزيز، عبد الحليم محمد أبو المجد، أيمن السعيد صادق، محمد طلعت محمد\*

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خلفية البحث: يعد تصلب الشرايين أحد عوامل الخطر الرئيسية لمرض الشريان التاجي، قد تكون الموجات فوق الصوتية السباتية طريقة مفيدة للتنبؤ بمرض الشريان التاجي من خللال قياس سمك البطانة الداخلية و الوسطى للشرايين السباتية اليمنى واليسرى.

الهدف من البحث: التأكيد على الارتباط بين تصوير نضح عضلة القلب باستخدام التصوير المقطعي المحوسب بإصبدار فوتون واحد والموجات فوق الصوتية السباتية من خلال قياس قياس سمك البطانة الداخلية و الوسطى في التنبؤ المبكر وتشخيص مرض الشريان التاجي.

المرضى وطرق الدراسة: شملت الدراسة 80 مريضاً يشكون من آلام في الصدر. تم عمل التصوير المقطعي المحوسب بانبعاث فوتون واحد والموجات فوق الصوتية السباتية لكل منهما. تم استقدام جميع المرضى من العيادات الخارجية بمستشفى الحسين الجامعي والعيادة الخارجية بمستشفى كوبري القبة العسكري بين فبراير 2020 ونوفمبر 2020.

النتائج: اشتملت الدراسة على 80 مريضا ، 57 منهم من الذكور و 23 من الإناث. كان 52 مريضاً يعانون من ارتفاع ضغط الدم، و 31 مريضاً من داء السكري، و 33 مريضاً من المدخنين، و 15 مريضاً لديهم تاريخ عائلي إيجابي. كان لدى 52 مريضاً اختبار التصوير المقطعي المحوسب بإصدار فوتون واحد بالمجهود إيجابي أظهر "مجموعة مرض القلب الإقفاري"، بينما كان لدى 28 منهم اختبار التصوير المقطعي المحوسب بإصدار فوتون واحد المحمومية المحمود سابي

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(82.7%) لحديهم زيادة في سمك البطانة الداخلية و الوسطي للشريان السباتي، سمك البطانة و الوسطي للشريان السباتي، سمك البطانة الداخلية و الوسطى التغيير المعياري" «معاملك البطانية الداخلية و الوسطى التغيير المعياري" معاملك المتريان المتحوازن 0.655 باتفاق جو هري و قيمة بي <0.001

سمك البطانة الداخلية و الوسطى للشريان الأيسر "متوسط ل التغيير المعياري" "0.12±0.77، اختبار تماير معامل كابسا المتوازن 0.614 باتفاق جو هري و قيمة بي <0.001.

فقط 9 مرضى (17.3%) من مجموعة مرض القلب الإقفاري كنان لديهم سمك البطانة الداخلية و الوسطى للشريان الثباتي أقل من الشريحة المئوية ال 75.

بالنسبة للمجموعة الغير الففارية 9 مرضى فقط (32.1%) لديهم سمك البطانة الداخلية و الوسطى للشريان السباتي غير طبيعية بينما البقية 19 مريضاً (67.1%) كان لديهم سمك البطانة طبيعي.

**الخلاصة:** الموجات فوق الصوتية للشريان السباتي وسماكة الوسائط الداخلية منبئات جيدة ومرتبطة بشكل كبير بمرض الشريان التاجي.

**الكلمات الدالة:** التصوير المقطعي المحوسب بإصدار فوتون واحد، والموجات فوق الصوتية السباتية، سماكة الوسائط الداخلية، لمرض الشريان التاجي.