

**Accuracy of Multi-Slice Computed Tomography  
Angiography in The Diagnosis of In-Stent Restenosis  
in Proximal Left Anterior Descending Coronary Artery  
Stent Compared to Conventional Coronary  
Angiography**

Thesis

*Submitted for Partial Fulfillment of Master Degree in  
Cardiology*

By

**Mina Aziz Maurice Tewfik**

*M.B., B.Ch, Ain Shams University*

Under Supervision of

**Prof. Dr. Yasser Gomaa El-Kashlan**

*Assistant Professor of Cardiology, Ain Shams University*

**Prof. Dr. Ahmed Mohamed Onsy**

*Assistant Professor of Cardiology, Ain Shams University*

**Dr. Maged Tewfik Saad Fahim**

*Lecturer of Cardiology, Ain Shams University*

Faculty of Medicine  
Ain Shams University

**2017**

# Acknowledgment

*First thanks to **God** to whom I relate any success in achieving any work in my life.*

*I wish to express my deepest thanks, gratitude and appreciation to **Dr. Yasser Gomaa El-Kashlan**, Assistant Professor of Cardiology for her meticulous supervision, kind guidance, valuable instructions and generous help.*

*Special thanks are due to **Dr. Ahmed Mohamed Onsy**, Assistant Professor of Cardiology for her sincere efforts and fruitful encouragement.*

*I am deeply thankful to **Dr. Maged Tewfik Saad Fahim**, Lecturer of Cardiology for her great help, outstanding support, active participation and guidance.*

*I would like to express my hearty thanks to all my family for their support till this work was completed.*

*Last but not least my sincere thanks and appreciation to all patients participated in this study.*

**Mina Aziz**

# *List of Contents*

Title	Page No.
List of Tables .....	4
List of Figures .....	5
List of Abbreviations .....	6
Introduction .....	1
Aim of the Work.....	10
Review of Literature	
▪ Instent Restenosis .....	11
▪ Natural History of MSCT .....	16
▪ Development of CT Coronary Angiography in Assessment of Instent Restenosis .....	21
Patients and Methods .....	34
Results .....	41
Discussion .....	49
Limitations of the Study .....	53
Conclusion and Recommendation.....	54
Summary .....	55
References .....	57
Master Table .....	72
Arabic Summary	

## *List of Tables*

Table No.	Title	Page No.
<b>Table (1):</b>	Summary of studies on EBCT imaging for assessing coronary stent patency .....	23
<b>Table (2):</b>	Summary of studies on 4-, 16-, and 40- slice CT imaging for assessing coronary stent patency and rule out of ISR .....	25
<b>Table (3):</b>	Summary of studies on 64-slice CT angiography for assessing coronary stents.....	28
<b>Table (4):</b>	Summary of studies on 64-slice DSCT and 320-slice Ct for assessing coronary stents .....	29
<b>Table (5):</b>	Age and sex of the studied patients .....	41
<b>Table (6):</b>	Risk factors among the studied patients .....	42
<b>Table (7):</b>	Stent characteristics at coronary angiography .....	43

## *List of Figures*

Fig. No.	Title	Page No.
<b>Fig. (1):</b>	Schematic image of 4 patterns of introduced classification of ISR in relation to previous dichotomous description of focal vs diffuse ISR.....	15
<b>Fig. (2):</b>	Detail render of drug-eluting stents. Diverse drug-eluting stents are currently available, differing in the type of metal used, stent design, and drug coating.....	33
<b>Fig. (3):</b>	Risk factors among the studied patients. ....	42
<b>Fig. (4):</b>	Diameter of stents at studied patients.....	43
<b>Fig. (5):</b>	Percentage of in-stent restenosis by QCA. ....	44
<b>Fig. (6):</b>	Comparison between total ISR detected by CA and MSCT. ....	45
<b>Fig. (7):</b>	Correlation between CA and MSCT in <b>stent &gt;3mm</b> .....	45
<b>Fig. (8):</b>	Correlation between CA and MSCT in <b>stent &lt;3mm</b> .....	46
<b>Fig. (9):</b>	A 48 years old male patient who underwent PCI to proximal LAD by stent 3x24mm is complaining of symptoms of anigina. He underwent MSCT and CA. they demonstrated a patent stent. ....	47
<b>Fig. (10):</b>	A 65 years old male patient who underwent PCI to proximal LAD by stent 2.75x18 mm is complaining of symptoms of anigina. He underwent MSCT and CA.....	48

## *List of Abbreviations*

<b>Abb.</b>	<b>Full term</b>
<i>AUC</i> .....	<i>Area under the curve</i>
<i>BMS</i> .....	<i>Bare metal stents</i>
<i>CA</i> .....	<i>Coronal angiography</i>
<i>DES</i> .....	<i>Drug eluting stents</i>
<i>EBCT</i> .....	<i>Electron beam CT</i>
<i>HU</i> .....	<i>Hounsfield units</i>
<i>IH</i> .....	<i>Intimal hyperplasia</i>
<i>ISR</i> .....	<i>Instant restenosis</i>
<i>IVUS</i> .....	<i>Intravascular ultrasonography</i>
<i>MDCT</i> .....	<i>Multidetector row computed tomography</i>
<i>MSA</i> .....	<i>Minimal stent area</i>
<i>NPV</i> .....	<i>Negative predictive value</i>
<i>PCI</i> .....	<i>Percutaneous coronary intervention</i>
<i>PES</i> .....	<i>Paclitaxel eluting stent</i>
<i>PPV</i> .....	<i>positive predictive value</i>
<i>SES</i> .....	<i>Sirolimus eluting stent</i>
<i>SPECT</i> .....	<i>Single photon emission tomographic</i>

## Abstract

Our study included 50 stents placed in the proximal segment of the left anterior descending coronary artery regardless their diameters and strut thickness assessed by MSCT and CA to assess the accuracy of MSCT in the diagnosis of significant in stent restenosis.

Our results showed that MSCT is an accurate and reliable tool in diagnosis of in stent restenosis in stents more than 3mm in diameter with 100% sensitivity and specificity. However this accuracy drops markedly in the assessment of stents less than 3mm in diameter due to an artificial narrowing of the stent diameter caused by the blooming effect.

Although MSCT can't be reliable in diagnosis of ISR in stents less than 3mm in diameter, it still can be used as a tool to rule out the presence of ISR in these stents with a negative predictive value almost 100%.

**Keywords:** Minimal stent area - Minimal stent area - Minimal stent area  
Intravascular ultrasonography - Sirolimus eluting stent

## Introduction

In the last years, coronary artery disease has been increasingly treated by coronary stent placement. Although stent implantation has been shown to greatly reduce restenosis after balloon angioplasty (*Schoenhagen et al., 2004*) in-stent restenosis can occur in 20-35% of patients for bare metal stents (*Morice et al., 2006*) and 5-10% for drug-eluting stents, as demonstrated by intravascular ultrasound.

Invasive coronary angiography is the gold standard technique for detection of in-stent restenosis. However, coronary angiography has limitations due to its invasiveness and association with potential risks of morbidity and mortality.

Given the high number of patients who receive coronary stents yearly, a non-invasive imaging technique for detection of in-stent restenosis will be clinically important and beneficial.

Since the introduction of multi-slice computed tomography (CT), CT angiography has emerged as a new tool in the diagnosis and monitoring of coronary heart disease. Additionally, noninvasive assessment of coronary stents is an attractive potential application of multi-slice CT technology (*Schoepf et al., 2004*).

Multi-slice computed tomography (MSCT) is increasingly used for non-invasive imaging of coronary artery disease and has

been reported to have a high diagnostic accuracy in the detection of coronary artery stenosis (*Fine et al., 2006*).

However, imaging of coronary stents by MSCT is more difficult than native coronary artery. This is due to the presence of artifacts interfering with the interpretation of lumen patency.

Although several reports have shown that MSCT may be used to evaluate stent patency, more precise evaluation of the lumen within stent is markedly affected by the blooming artifacts that can cause an appearance of artificial enlargement of the metallic stent struts (*Kruger et al., 2003*).

With increasing number of detector rows, promising results of MSCT in coronary artery disease have been reported with improved spatial and temporal resolution. However, it is unclear whether this also applies to the assessment of coronary stent implantation.

## **AIM OF THE WORK**

**T**o assess the accuracy of multi-slice Computed Tomography angiography in the diagnosis of in-stent restenosis in proximal Left anterior descending coronary artery stent compared to conventional coronary angiography.

## Chapter 1

# INSTENT RESTENOSIS

**P**ercutaneous coronary intervention (PCI) has seen an enormous increase and tends to be the preferred method for myocardial revascularization (*Mack et al., 2004*).

Nowadays, the routine use of balloon angioplasty is very limited due the recurrence of luminal narrowing due to recoil, the vessel wall remodeling or intimal hyperplasia induced by arterial wall injury and atherosclerosis progression. Because of the continuous evolution of stents and the introduction of newer drug eluting stents, interventional cardiologists have now a broad therapeutic spectrum. Thus the use of stents during PCI has achieved a significant decrease in complications and an improvement in all the patients' outcomes (*Serruys et al., 1994*).

The basic idea was for the implanted stent to serve as a scaffold that would maintain the artery's patency permanently. But the major drawback about implanted stents is still the chance of in-stent restenosis (*Kasaoka et al., 1998; Akiyama et al., 1998*).

Although the introduction of drug-eluting stents (DES) has made a huge decrease in the percentage of in-stent restenosis compared to bar-metal stents, but it still remains the Achilles' heel of PCI.

## **Mechanism and factors contributing to in-stent restenosis**

The rate of in-stent restenosis in BMS is very high reaching around 25% (*Serruys et al., 1994; Fischman et al., 1994*). Compared to a rate of less than 10% using DES (*Moses et al., 2003; Stone et al., 2004*).

However, its prevalence will be greater in view of the fact that, in the real world, DES are being increasingly used in complex lesions such as those in the left main artery, bifurcations, small vessels, vein grafts, chronic total occlusions, acute coronary syndromes and diabetic patients.

In these patient populations, “off-label” use has led to an ISR rate exceeding 10% (*Stone et al., 2005; Tanabe et al., 2004*).

Another fearful issue is that DES restenosis does not always present benignly, with myocardial infarction being the initial clinical manifestation in up to 10% of patients (*Abizaid et al., 1998*).

A number of predisposing factors have been associated with in-stent restenosis and can be divided into *vessel and lesion-related, procedure-related and patient-related*.

### **1- Vessel and lesion characteristics**

The vessel nature is a main indicator of the probability of ISR such as the vessel size, tortuosity, calcification or totally occluded vessels.

Another indicator is the lesion characteristics and sites especially lesions located in the left anterior descending coronary artery (LAD).

### **2- Technical failures of the implantation**

Improper implantation technique is still a risk factor for the occurrence of ISR such as small post-procedural minimum lumen diameter, underexpansion, overexpansion, stent fracture, non-uniform distribution of stent struts and malapposition.

### **3- Patient related factors**

Patients having diabetes mellitus have a higher probability for ISR (*Kip et al., 1996*). Genetic factors, such as the PIA polymorphism of glycoprotein IIIa (*Kastrati et al., 1999*), the insertion/deletion polymorphism and the plasma activity of angiotensin I-converting enzyme (*Ribichini et al., 1998*) have been reported to be important patient-related risk factors of ISR.

## Classification of ISR

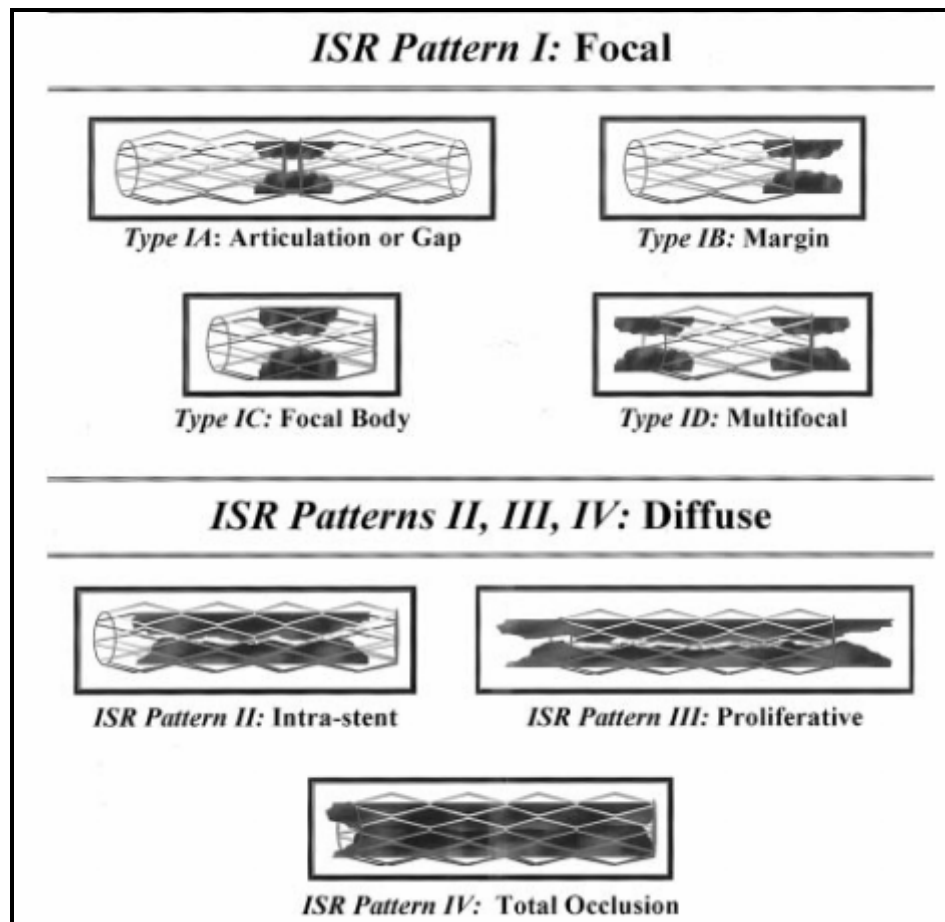
An earlier classification of lesions into either diffuse (lesion length >10 mm) or focal (<10 mm) has proved inadequate to predict the rate of target vessel revascularization (TVR). Nowadays, the angiographic pattern of restenosis based on Mehran's classification for ISR seems to have important prognostic value and may be used for further clinical assessment (*Mehran et al., 1999*).

- **Mehran classification (Figure 1)**

- **Class I:** Focal ISR group. Lesions are <10 mm in length and are positioned at the unscaffolded segment (ie, articulation or gap), the body of the stent, the proximal or distal margin (but not both), or a combination of these sites (multifocal ISR).
- **Class II:** “Diffuse intrastent” ISR. Lesions are 10 mm in length and are confined to the stent(s), without extending outside the margins of the stent(s)
- **Class III:** “Diffuse proliferative” ISR. Lesions are 10 mm in length and extend beyond the margin(s) of the stent(s).
- **Class IV:** ISR with “total occlusion.” Lesions have a TIMI flow grade of 0.

Recurrent ISR was more frequent with increasing grades of classification, as with diabetes.

Target lesion revascularization (TLR) increased according to ISR class, ranging from 19% to 83% for classes I to IV, respectively ( $p < 0.001$ ).



**Fig. (1):** Schematic image of 4 patterns of introduced classification of ISR in relation to previous dichotomous description of focal vs diffuse ISR. Pattern I contains 4 types (A-D). Patterns II through IV are defined according to geographic position of ISR in relation to previously implanted stent.