

## CT ANGIOGRAPHY

# Computed Tomography of the Coronary Arteries

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**ABSTRACT** MSCT Coronary Angiography is a fast developing non-invasive diagnostic technique that can detect a coronary stenosis. The detection of a coronary stenosis is hampered by limited image quality and by motion artefacts and extensive calcifications. However, MSCT-coronary angiography is highly reliable to rule out coronary stenosis. The role of MSCT-coronary angiography in the diagnostic work-up of coronary artery disease needs further research.

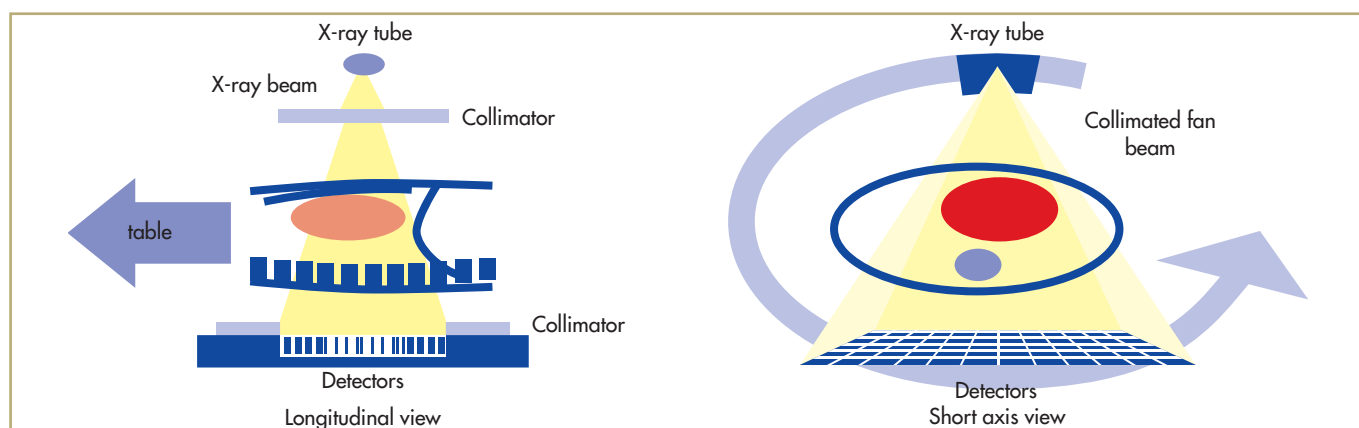
## INTRODUCTION

CT-coronary imaging is a robust technology which has a high temporal and spatial resolution allowing non-invasive visualization of the coronary plaque burden (coronary calcified and non-calcified plaques) and coronary lumen obstructions<sup>(1-3)</sup>. CT-coronary imaging has developed rapidly and is now poised to become a clinically reliable non-invasive diagnostic modality to assess the coronary lumen and coronary plaques. However, CT-coronary imaging is still fraught with limitations.

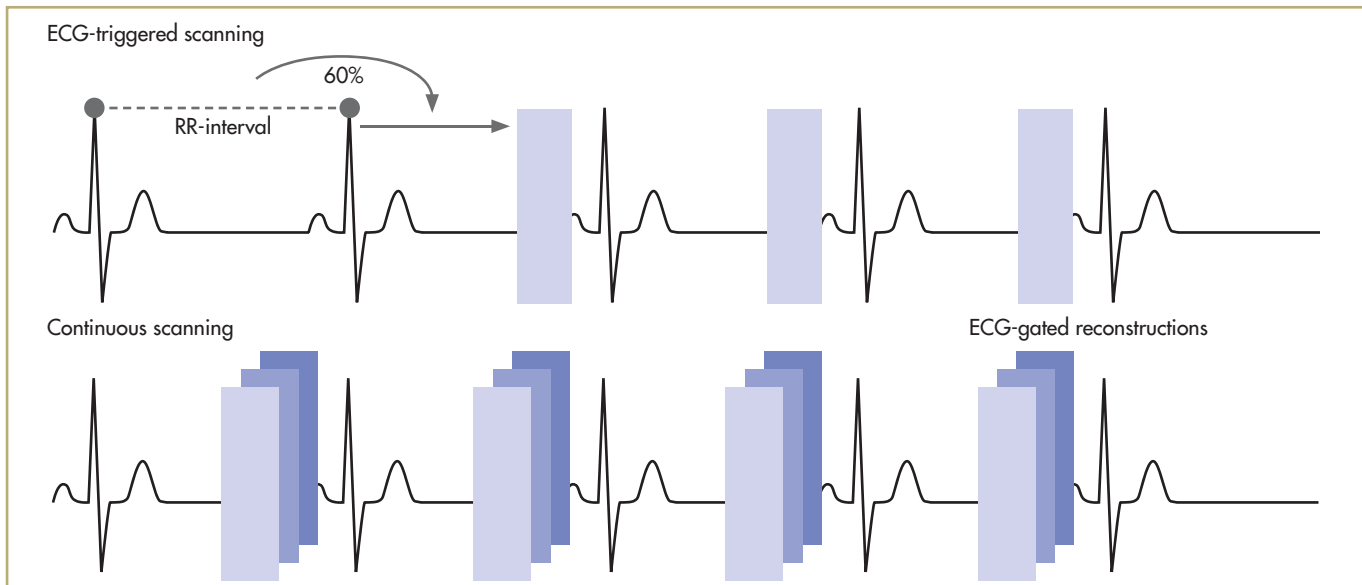
This article reviews the diagnostic utility and limitations of CT-coronary imaging.

## BASIC COMPUTED TOMOGRAPHY

The gantry and patient couch are referred to as the scanner. The gantry is a mounted framework that surrounds the patients in a vertical plane<sup>(1)</sup>. It contains an X-ray tube and a row of detectors which rotate around the patient, who is positioned in the center of the gantry (Figure 1). The X-ray tube produces a fan-shaped X-ray beam which is collimated and passes through the patient and is collected on the detectors at the opposite side. The CT-uses X-ray absorption to create images. Image contrast depends on the physical density and the number of atoms with differing atomic numbers. Iodine and calcium have high atomic numbers, which is associated with high absorption and will appear white in CT-images. Hydrogen, which is abundant in fat, has a low atomic number and low absorption and appears dark grey. Blood and soft tissue containing hydrogen, oxygen and carbon appear light grey. Lung tissue contains air and appears black. Each individual tomogram can be seen as a slice, which is composed of voxels, each of which contains a specific CT-number. Each CT-number is proportional to the absorption of X-rays of tissue in that voxel. An image is reconstructed from the various CT-numbers within the slice. The CT-numbers are established on a relative basis, using the attenuation of water as a reference. The CT-number for water is 0, for air -1000 and for bone +1000. A CT-image is reconstructed from CT-numbers obtained during a full rotation (360°) of the X-ray tube around the patient. Because the first and second halves of the X-ray tube rotation provide comparable data partial scan (180°) reconstruction data, can be used that reduce the temporal resolution (see later) to half the X-ray tube rotation time.



**FIGURE 1:** The X-ray tube and the detectors rotate in an opposing position of the gantry around the patient. A collimated X-ray beam is passed through the patient and the attenuated X-ray beam is collected on the detectors while the patient on the couch is continuously advanced through the gantry. 64 parallel detector rows acquire the data in a very short scan time.



**FIGURE 2:** ECG-synchronized image reconstruction with sequential scan protocols using prospective ECG triggering to synchronize the data acquisition and reconstruction window to the motion of the heart. Spiral CT scanning protocols acquire data continuously and record the ECG during the scan. Isocardiophasic images are reconstructed using retrospective ECG-gating. The reconstruction window can be positioned anywhere within the R-R-interval.

Multislice CT-imaging is carried out either in the spiral mode or in the sequential mode. Multi-slice spiral CT is performed with the patient on the couch, continuously moving at a predefined speed through the scanner while the X-ray tube and detectors rotate continuously around the patient. Thus a volumetric dataset is acquired from which cross-sectional images can be reconstructed. In the sequential mode ("slice by slice") the table, and thus the patient, is moved incrementally between successive rotations of the X-ray tube ("step and shoot" approach).

Imaging of the heart requires acquisition or image reconstruction that is synchronized to the motion of the heart, to be able to reduce cardiac motion artefacts. This is achieved by simultaneously recording the ECG, which is used to synchronize prospective data acquisition or retrospective data reconstruction. In the sequential CT-mode the reconstruction window is determined by prospective triggering of the patient's ECG (Figure 2). In the spiral mode, the ECG signal is used to retrospectively, after data acquisition, select a cardiac motion-free dataset which mostly occurs in the diastolic phase of the cardiac cycle, but sometimes also in the systolic phase. Because spiral CT coronary angiography is associated with a rather high radiation exposure, the radiation exposure can be reduced by prospectively ECG-triggered X-ray tube modulation (see radiation exposure).

## CT IMAGE QUALITY

Cardiac imaging suffers from cardiac motion-artefacts and, therefore, the data used for reconstruction of the coronary images are preferentially obtained from a relative cardiac motion-free period, which occurs mostly during the diastolic phase of the cardiac cycle.

Because the diastolic phase is longer during low heart rates (<60 bpm), the optimal motion-free images are obtained during low heart rates, and therefore  $\beta$ -blockers are usually given to reduce the higher heart rates (>65 bpm)<sup>(4,5)</sup>.

Although the technical developments of CT have been spectacular, compared to the almost unbeatable high quality images of conventional invasive coronary angiography, the spatial resolution of 64-CT is still limited, but allows near isotropic resolution with voxel size of  $0.4 \times 0.4 \times 0.4 \text{ mm}^3$  (Table I).

**TABLE I:** Significant parameters of CT-coronary angiography compared to invasive coronary angiography

	16-slice CT	64-slice CT	Invasive Coronary Angiography
Spatial resolution	0.8 mm	0.4 mm	0.2 mm
Temporal resolution	180 – 210 ms	165 ms	20 ms
Contrast volume	120 ml	80 ml	60 ml
Radiation exposure	5 – 10 mSv	6 – 12 mSv	3 – 4 mSv

The temporal resolution is of utmost importance when imaging the heart and plays a major role in the creation of motion-free cardiac images. The temporal resolution is defined as the duration of the reconstruction window during which time a slice is reconstructed (Figure 2).

The duration of the reconstruction window is related to the tube-rotation speed and is half the time needed for the X-ray tube to rotate  $360^\circ$  around the patient. Thus the faster the rotation speed the better the temporal resolution.

**X-RAY RADIATION EXPOSURE**

CT-coronary imaging is associated with X-radiation exposure, which is expressed as the effective dose. The effective dose reflects the potential biological effect of the radiation and is given in mSv (Sievert). The effective dose for various CT-examinations and scan protocols is presented in Table 2<sup>(6-9)</sup>.

**TABLE 2:** Radiation exposure associated with cardiac CT-examination

	Total – body effective dose mSv	
	male	female
EBCT coronary calcium	1.0	1.3
MDCT coronary calcium	2.3 – 2.9	3.2 – 3.6
16-slice CT coronary angiogram (no pulsing)	7.9 – 11.8	11.1 – 16.3
16-slice CT coronary angiogram (pulsing)	4.0 – 6.2	5.6 – 8.7
64-slice CT coronary angiogram (no pulsing)	9.6 – 15.2	13.5 – 21.4
64-slice CT coronary angiogram (pulsing)	4.8 - 10	6.8 - 14

Helical scanning is associated with higher radiation exposure than sequential scanning. Prospectively ECG triggered X-ray tube modulation does reduce the radiation exposure by about 40%. Radiation exposure is reduced because the full tube current is only generated during a short time interval during a predefined period in diastole and reduced tube current during systole and early diastole. However, this mode is not very robust and relies on a stable heart rhythm, and the occurrence of an extrasystole during scanning may ruin the image quality. Furthermore, use of the tube modulation mode prevents data reconstruction during systole which, in approximately 30% of cases produces better image quality.

**CT-CORONARY CALCIFICATION IN ASYMPTOMATIC INDIVIDUALS**

CT-coronary calcification is imaged in the sequential mode using a low-dose radiation technique without contrast enhancement. The calcified coronary artery plaque is traditionally quantitated using the Agatston score in which the area of a calcific plaque with a CT-attenuation of more than 130HU is multiplied by a weighting factor that is based on the peak density of the calcified lesion<sup>(11)</sup>. A factor of 1,2,3, or 4 has been assigned to calcium attenuation of 130-199HU, 200-299HU, 300-399HU, or >400HU respectively.

**TABLE 3:** Predictive value of CT-calcium of adverse cardiac events. Coronary Calcium Score

Author	N subjects	Calcium score	Prevalence (%)	F-up months	Annual event rate % (death, nonfatal/MI)
Detrano	1196	>0	67	41	1.1
Raggi	676	>0	53	32	1.7
Kondos	5635	>0	60	37	0.3
Shaw	10,377	≥11	43	60	0.5 (all-cause death)
Vliegenthart	1795	>0	63	40	0.2
Arad	4903	>0	49	52	0.2
Lamonte	10,746	>100	50	42	0.7

The presence of coronary calcium is directly associated with the presence of coronary atherosclerosis and it has been shown that the presence of coronary calcium can predict adverse cardiac events, which is independent of and incremental to the presence of traditional risk factors<sup>(11-17)</sup> (Table 3).

Currently, the definitive role of CT-calcium scoring in asymptomatic individuals is not yet established. We will need more evidence from well-designed studies, such as the RECALL (Risk-factors Evaluation of Coronary calcium and Life style), The Rotterdam Study and MESA-study (Multi-Ethnic Study of Atherosclerosis), and CARDIA (Coronary Artery Risk Development in Young Adults) to better understand the additional role of calcium scoring for risk factor management in asymptomatic persons.

**MSCT CORONARY ANGIOGRAPHY: 16-SLICE CT-SCANNER**

The 16-slice MSCT scanner has a temporal resolution of 200 msec and an in-plane resolution of 0.4 x 0.4mm with a slice thickness of 0.75mm. Comparative invasive angiographic studies have shown that the diagnostic performance of 16-slice MSCT coronary angiography to detect significant coronary stenosis is high, with sensitivities ranging from 67% to 98% and specificities from 79% to 98%<sup>(18-31)</sup> (Table 4). Pooling the results of these studies reveals an average sensitivity of 87% and a specificity of 96%. This is achieved in approximately 92% of all coronary segments, while around 8% of the coronary segments were not analysable due to motion-artifacts or severe calcification. These limitations render the 16-slice CT-technique not reliable for routine clinical use and it is of note that the reported sensitivities and specificities were obtained in the larger proximal, mid and distal coronary segments that had lumen diameters of at least 1.5 to 2 mm.

**TABLE 4:** Diagnostic performance of 16 CT-coronary angiography for detection of significant coronary stenosis (luminal diameter > 50%): a coronary segmental analysis.

Author	Number patients	Excl. Segments %	Sensitivity %	Specificity %	PPV %	NPV %
Nieman	58	0	95	86	80	97
Mollet ('04)	127	7	92	95	79	98
Martuscelli	72	16	89	98	90	98
Kuettner ('04)	58	21	72	97	72	97
Leta	31	12	75	91	65	94
Hoffmann** ('04)	33	17	67	95	64	96
Mollet ('05)	51	0	95	98	87	99
Kuettner	72	7	82	98	86	97
Cademartiri	40	0	96	96	86	99
Schuijf	45	6	98	97	89	99
Kefer	52	0	82	79	46	95
Hoffmann	103	7	95	98	95	99
Kuettner	120	7	85	98	93	95
Achenbach	50	4	94	96	68	99
(weighted)	912	7.8	87	96	81	97.5

\*\* stenosis > 70% PPV = positive predictive value NPV = negative predictive value

FIGURE 3

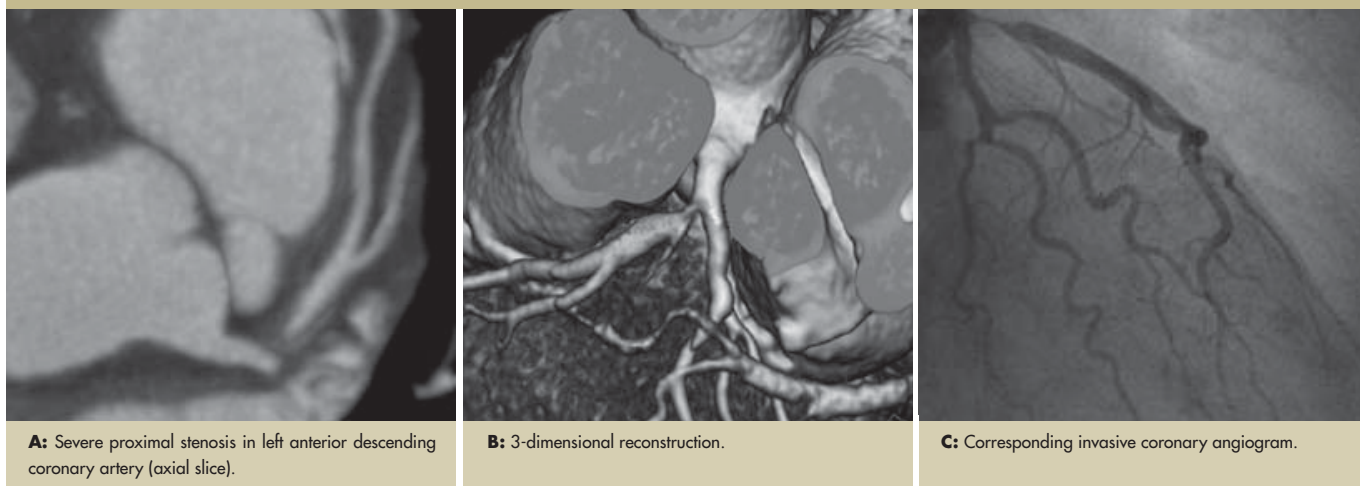


FIGURE 4



**MSCT CORONARY ANGIOGRAPHY:  
64-SLICE CT-SCANNER**

Significant technical advances in CT-technology have been introduced with almost unprecedented speed during the recent few years, which has culminated in the production of a 64-slice CT scanner which nowadays is available for clinical use and offered by the various vendors (Siemens, Philips, General Electronic and Toshiba). The 64-CT scanners from the various vendors differ slightly in design and configuration but all scanners feature a high temporal and spatial resolution and fast coverage of the entire heart with an acquisition time ranging from as low as 6 seconds to as high as 12 seconds. The diagnostic performance of the 64-slice CT-scanner when compared to quantitative invasive coronary angiography, to detect (Figure 3) or rule out (Figure 4) a significant coronary stenosis is shown in Tables 5 and 6 (32-40). The 64-slice CT-scanner still is not perfect but the high negative predictive value of 98% underscores the reliability of CT to exclude the presence of

significant coronary stenosis. Unfortunately, the 64-CT scanner still was not always able to analyse all coronary segments, and around 6% of coronary segments were excluded from analysis.

**TABLE 5:** Diagnostic performance of 64-slice CT coronary angiography to detect coronary stenosis (>50% luminal diameter: a segmental analysis)

Author	Number patients	Excl. Segments %	Sensitivity %	Specificity %	PPV %	NPV %
Leschka	67	0	94	95	88	98
Leber	55	0	76	97	75	97
Raff	70	12	86	95	66	99
Mollet	51	0	99	95	76	99
Ropers	81	4	93	97	56	100
Schuijff	60	1.4	85	98	82	96
Ong	134	9.7	82	96	83	96
Ehara	69	8	90	94	89	95
Nikolaou	72	10	82	95	72	97
Total Weighted	659	5.6	90	96	79	98

PPV = positive predictive value NPV = negative predictive value

**TABLE 6:** Diagnostic performance of 64-slice CT-coronary angiography to detect a coronary stenosis (>50% luminal diameter): a patient-based analysis

Author	Number patients	Sensitivity %	Specificity %	PPV %	NPV %
Leschka	67	100	100	88	98
Leber	45	88	85	75	97
Raff	70	95	90	66	99
Mollet	51	100	92	76	99
Ropers	81	96	91	56	100
Schuijff	60	94	97	97	93
Ehara	67	98	86	98	86
Nikolaou	68	97	79	86	96
Total (weighted)	509	96	90	94	95

PPV = positive predictive value NPV = negative predictive value

### **CT-CORONARY ANGIOGRAPHY FOR ASSESSMENT OF CORONARY STENTS**

Assessment of coronary stents is extremely challenging. The stent struts create a blooming artefact which makes the stent struts much larger, thereby obscuring part of the in-stent lumen. Stent evaluation is usually limited to the assessment of stent patency which is quite reliable, but in particular, the smaller sized stents ( $\leq 3\text{mm}$  in diameter) do not permit evaluation of the presence of neo-intimal hyperplasia or in-stent restenosis<sup>(41-45)</sup>.

However, the use of 64-slice CT-scanners has improved the visualization of stents, and now allows in-stent restenosis evaluation in larger sized stents ( $> 4\text{mm}$  in diameter) (Figure 5). A recent study demonstrated in 70 patients who underwent a follow-up study of patients who underwent left main stent implantation that a negative 64-slice CT-scan could reliably exclude the presence of in-stent restenosis<sup>(46)</sup>. This study suggested that 64-slice CT-ca could serve as an initial diagnostic modality to rule out the presence of in-stent restenosis in patients with left main stem stent implantation, while in case of a positive 64 CT-scan a confirmative invasive coronary angiogram is recommended because of the rather low positive predictive value of the scan.

**TABLE 7:** Diagnostic performance of 16 and 64-slice CT for the detection of bypass graft occlusion or non-occlusive obstruction and obstructions in native coronary arteries

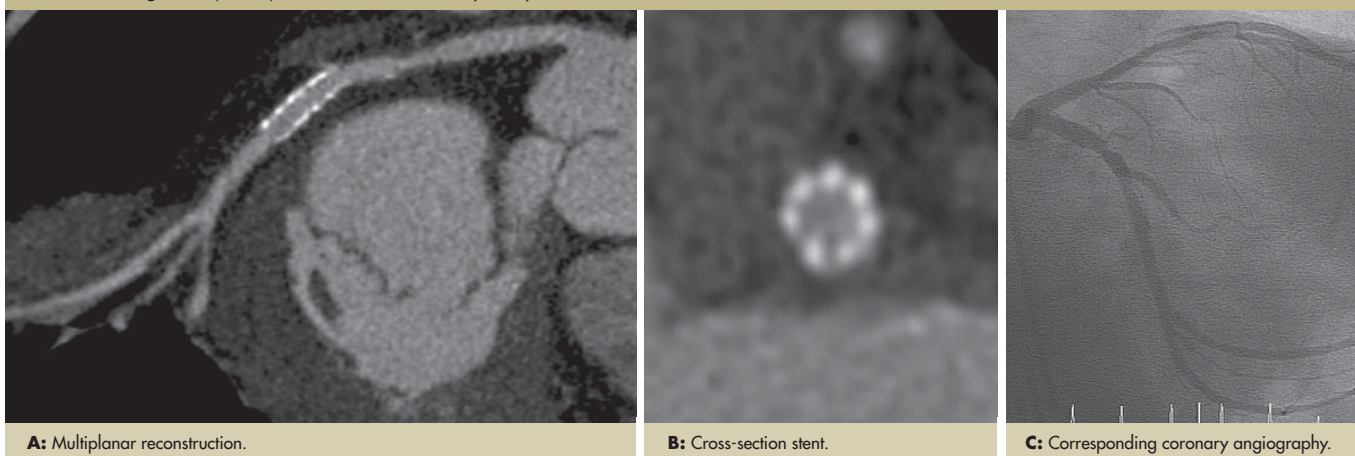
Author	Number	Endpoint patients	Sensitivity %	Specificity %
Martuscelli	96	Occlusion	100	100
		Stenosis	90	100
Schlosser	51	Occlusion	100	100
		Stenosis	90	100
Chiurlia	51	Occlusion	100	100
		Stenosis	96	100
Moore	50	Occlusion	100	100
		Stenosis	100	99
Salm	25	Occlusion	100	100
		Stenosis	100	94
		Coronary stenosis	100	89
Nieman	24	Occlusion	100	98
		Stenosis	70	88
		Coronary stenosis	85	73
Malagutti*	52	Occlusion	99	96
		Stenosis	89	93
		Coronary stenosis		

\* 64-slice CT-scanner

### **CT-CA FOR ASSESSMENT OF BYPASS PATIENTS**

The imaging of venous bypass grafts is less challenging because the grafts usually have a large diameter, are relative immobile and less calcified, but surgical metal clips causing significant CT-artefacts may cause problems with accurate assessment of adjacent disease. The detection of graft-occlusion using the 16-slice CT-scanner is quite reliable (Table 7) while the detection of non-occlusive stenoses is more problematic<sup>(47-53)</sup>. The 64-slice scanners perform slightly better, but evaluation of a post-bypass patient should include not only the bypass grafts but also the run-off native coronary segments distal to the graft anastomosis and non-grafted native coronary segments (Figure 6). Imaging of these coronaries is challenging because they usually are rather small, diffusely diseased and often heavily calcified.

**FIGURE 5:** Large stent (3.5mm) in left circumflex coronary artery without in-stent restenosis.



**A:** Multiplanar reconstruction.

**B:** Cross-section stent.

**C:** Corresponding coronary angiography.

**FIGURE 6**



**A:** LIMA with anastomosis on left anterior descending coronary artery.  
Note: bright dots are surgical clip artefacts.



**B:** RIMA with anastomosis on the right coronary artery.



**C:** RIMA anastomosis and proximal occlusion of right coronary artery.

$\beta$ -blockade remedies this problem in the majority of the cases. The temporal resolution can be further reduced by use of bisegmental or multisegmental reconstruction algorithms that combine data from 2 or more cardiac cycles and thus prevent the occurrence of cardiac motion blurred images. However, these algorithms are most optimal when the heart rate is constant, the ECG signal is reliable and no arrhythmia occurs, since they rely on an identical cardiac contraction pattern with time-consistent positioning of the cardiac structures during each consecutive heart cycle.

### **NEWEST CT-SCANNER: THE DUAL – SOURCE 64-CT**

The temporal resolution of CT-scanners is still limited and the cause of motion artefact of fast moving coronary segments. The dual-source CT scanner features a gantry composed of two X-ray tubes placed at an orthogonal angle and each has an opposite detector row. This new CT-configuration has improved the temporal resolution twofold as compared to a conventional scanner<sup>(54)</sup>. The initial results are impressive and demonstrate that, also at high heart rates, motion-free images can be obtained. Further studies are needed to evaluate its diagnostic performance.

### **CARDIAC CT: EVOLVING INDICATIONS**

64-Slice CT-ca has now sufficient diagnostic accuracy to exclude significant coronary stenosis, suggesting that the technique is now ready to establish its role in the clinical work-up of patients presenting with chest pain. However, CT-ca is still limited and the technique fails in cases of persistent arrhythmia, frequent extra-systoles and severe segmental calcification, which may occur quite frequently alone or in combination in elderly populations (>70 years) where the incidence of symptomatic coronary artery disease is most frequent. Furthermore, a new technique would only be useful as a reliable alternative diagnostic technique if it is able to assess all relevant coronary segments and is not limited by the exclusion of non-evaluable segments due to motion artefacts or significant coronary calcifications. Currently CT-ca may be a viable technique in a limited number of clinical situations (Table 8).

**TABLE 8:** CT-ca as an alternative for invasive coronary angiography

1. coronary anomalies
2. patency bypass grafts
3. exclusion of significant CAD in patients referred for cardiac valve surgery
4. exclusion of significant CAD in patients with a low or intermediate pretest-risk
5. exclusion of in-stent restenosis in stents implanted in left main stem

### **ABBREVIATIONS**

CT	Computed Tomography
HU	Hounsfield Units
MSCT	Multi-Slice Computed Tomography
CT-ca	Computed Tomography Coronary Angiography

### **LIMITATIONS OF MSCT CORONARY ANGIOGRAPHY**

Despite significant technological progress and increasing experience with the evaluation of MSCT coronary angiography, a number of shortcomings remain. CT-coronary angiography cannot be used in patients with persistent irregular heart rhythm (atrial fibrillation) or frequent extrasystoles which preclude reconstruction of coronary images. Severe calcification poses two problems: 1) it obscures the underlying lumen and thus detection of a significant stenosis and 2) due to its blooming effects the severity of an adjacent stenosis may be overestimated. Fast heart rates (>70 bpm) are associated with cardiac motion-artefacts or unsharp images because the end-diastolic reconstruction time required for motion-free coronary reconstruction imaging is reduced. Reduction of the heart rate with oral or intravenous

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