



PLATFORM OF LABORATORIES FOR ADVANCES IN CARDIAC EXPERIENCE

**ROMA**

Centro Congressi  
di Confindustria

**Auditorium  
della Tecnica**

**9<sup>a</sup> Edizione**

**30 Settembre**

**1 Ottobre**

**2022**

**IMAGING E STRATIFICAZIONE DEL RISCHIO NELLA  
CARDIOPATIA ISCHEMICA**

**IMAGING IBRIDO ANATOMO-FUNZIONALE: PERCHÉ È  
LA DIREZIONE GIUSTA**

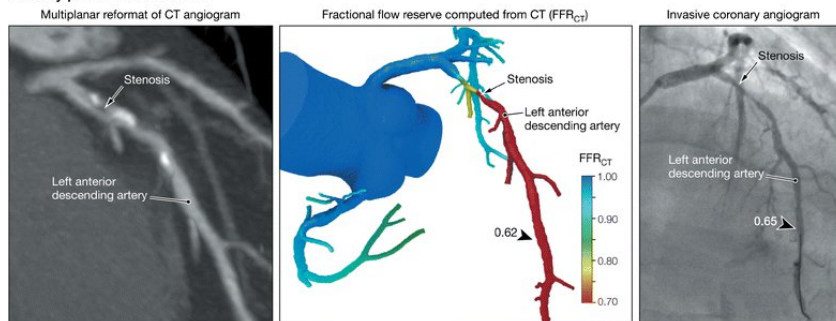
**Nudi Francesco**



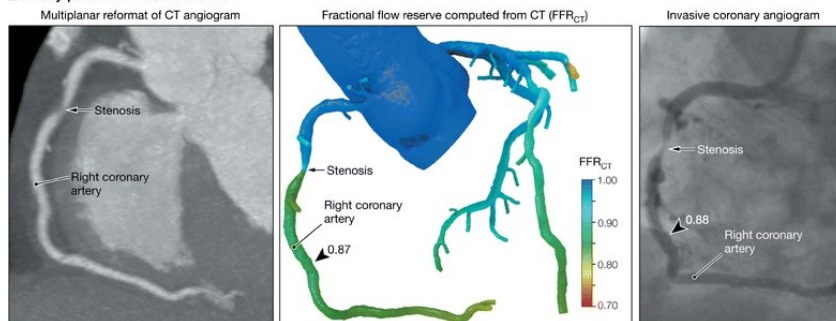


# Diagnostic accuracy of fractional flow reserve from anatomic CT angiography

## A Study patient with ischemia



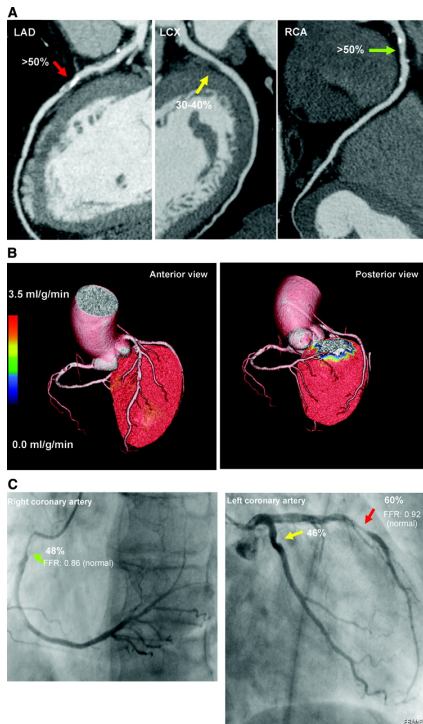
## B Study patient without ischemia



Non-invasive fractional flow reserve derived from conventional CCTA. Although both patients have obstructive coronary artery disease by computed tomographic angiography (CT), FFRCT identified myocardial ischemia in one patient (A), whereas FFRCT revealed no ischemia in the other patient (B). Multiplanar reformat of a CT angiogram demonstrating obstructive stenosis of the proximal portion of the left anterior descending artery (LAD) and an FFRCT value of **0.62**, indicating vessel ischemia. Invasive coronary angiogram demonstrates obstructive stenosis of the proximal portion of the LAD and measured FFR values of **0.65**, indicating vessel ischemia. (B) CT angiogram demonstrating obstructive stenosis of the mid portion of the right coronary artery (RCA) and an FFRCT value of **0.87**, indicating no vessel ischemia. Invasive coronary angiogram demonstrates obstructive stenosis of the mid portion of the RCA and a measured FFR value of **0.88**, indicating no vessel ischemia



## Cardiac Positron Emission Tomography/Computed Tomography Imaging Accurately Detects Anatomically and Functionally Significant Coronary Artery Disease



A 69-year-old man with attacks of atypical anginal pain. A, CT showed significant LAD and RCA stenoses with only mild stenosis in the LCx.

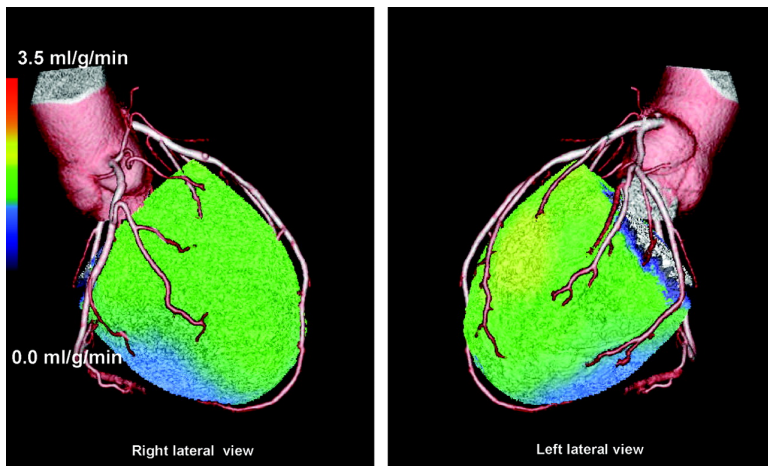
B, Hybrid images with normal stress PET perfusion (absolute scale, 0 to 3.5 mL g<sup>-1</sup> min<sup>-1</sup>). Normal perfusion is >2.5 mL g<sup>-1</sup> min<sup>-1</sup> (yellow or red). C, ICA with quantitative analysis and FFR.

Despite anatomically significant narrowing of the LAD and borderline changes in the RCA, FFR was normal in both vessels, indicating functionally nonsignificant disease.

Kajander S, et al. (Circulation. 2010;122:603-613)



## Cardiac Positron Emission Tomography/Computed Tomography Imaging Accurately Detects Anatomically and Functionally Significant Coronary Artery Disease

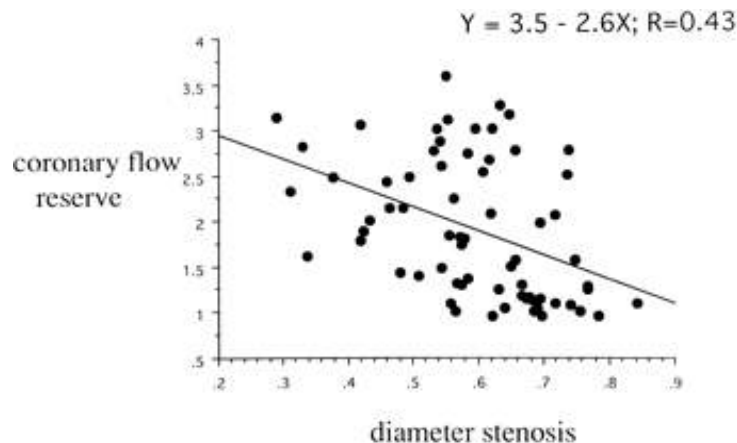


A 63-year-old man with positive family history. Good performance, atypical chest pain, and 2-mm ST depression in the ECG at the exercise test. In hybrid images, stress myocardial perfusion was reduced in most regions (green and blue). However, both CT and ICA showed normal coronary arteries, indicating possible microvascular disease.





## Intracoronary Doppler assessment of moderate coronary artery disease



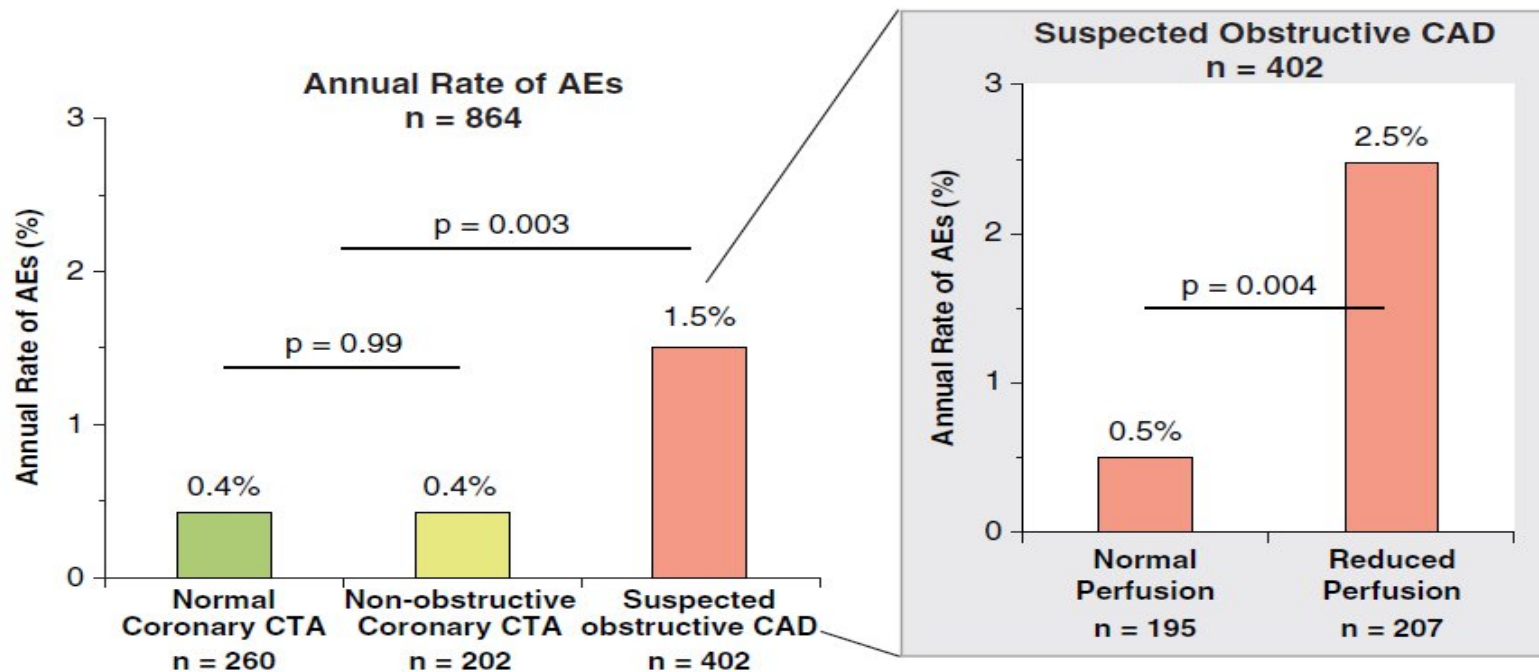
	Pos TI-201	Neg TI-201
CFVR < 1.7	87%	13%
CFVR > 1.7	11%	89%

**r = 88%**

Heller et al, Circulation 1997



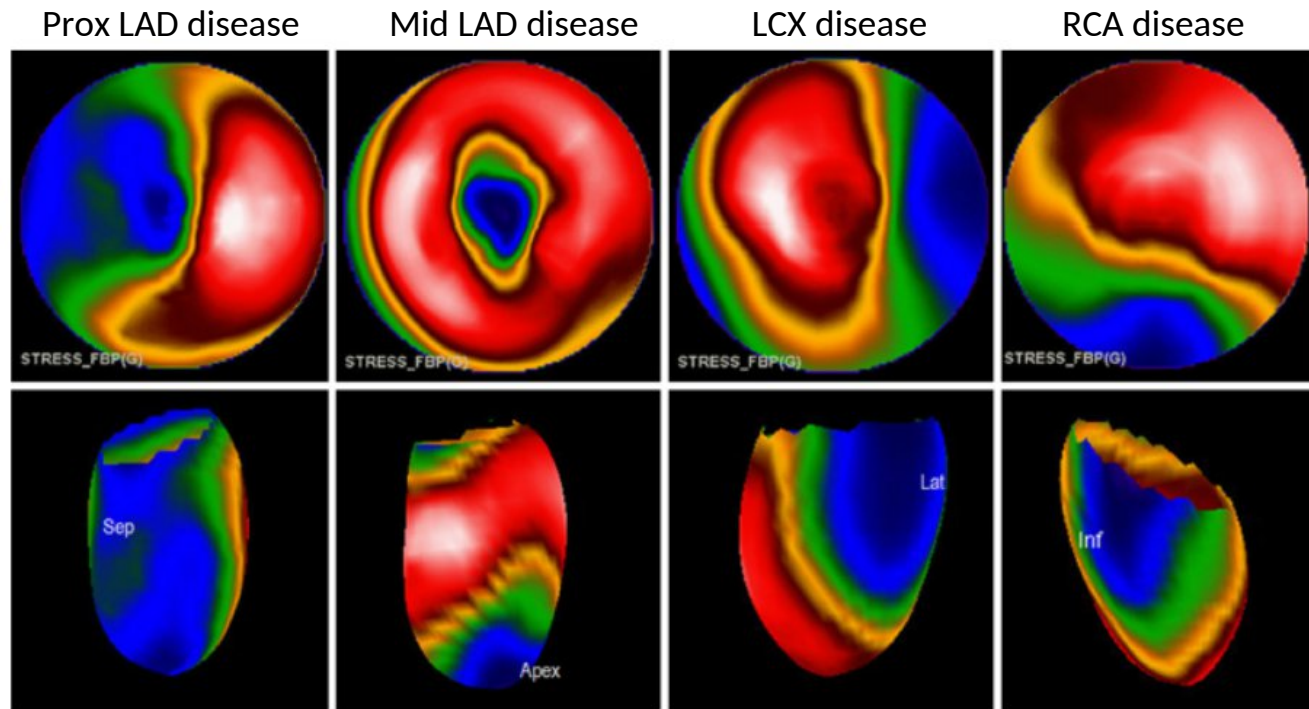
## CT followed by selective PET



Maanitty et al, J Am Coll Cardiol Img 2017



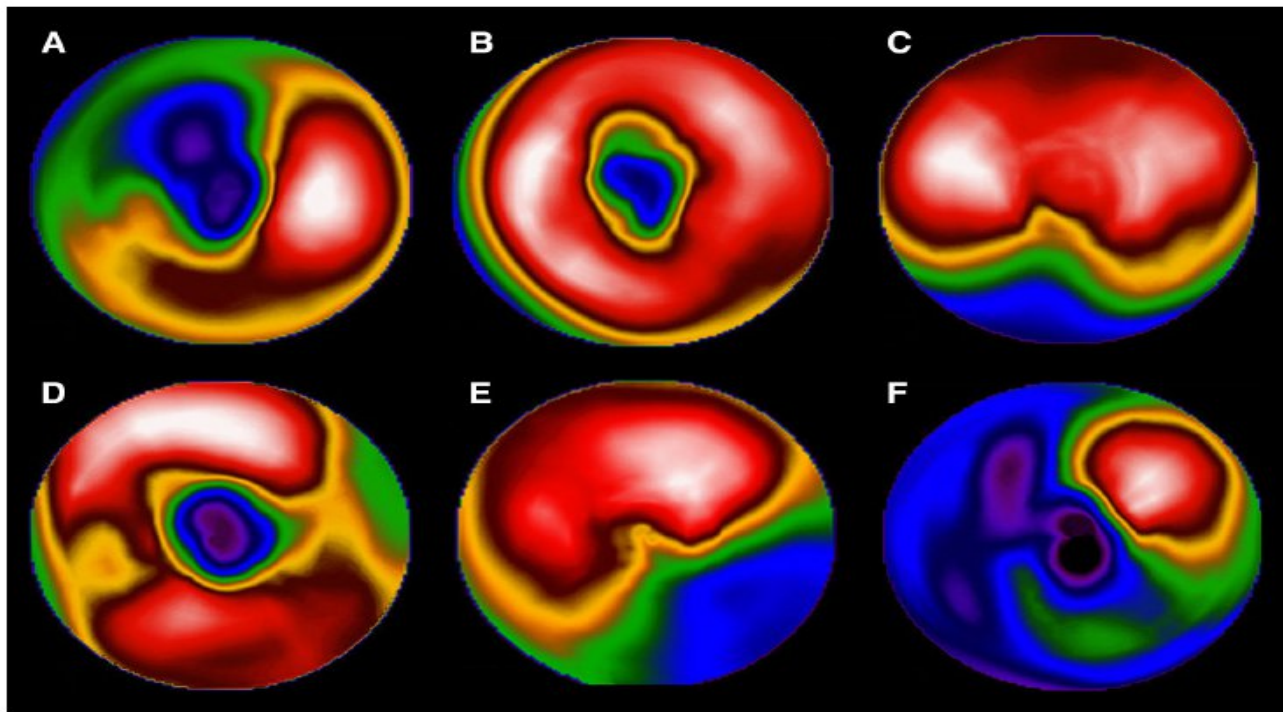
## SPECT imaging: a novel and clinically-relevant segmentation approach



Nudi F, et al (J Nucl Cardiol 2014)



## Vessel-related ischemia



Nudi et al, (J Nucl Cardiol 2016)



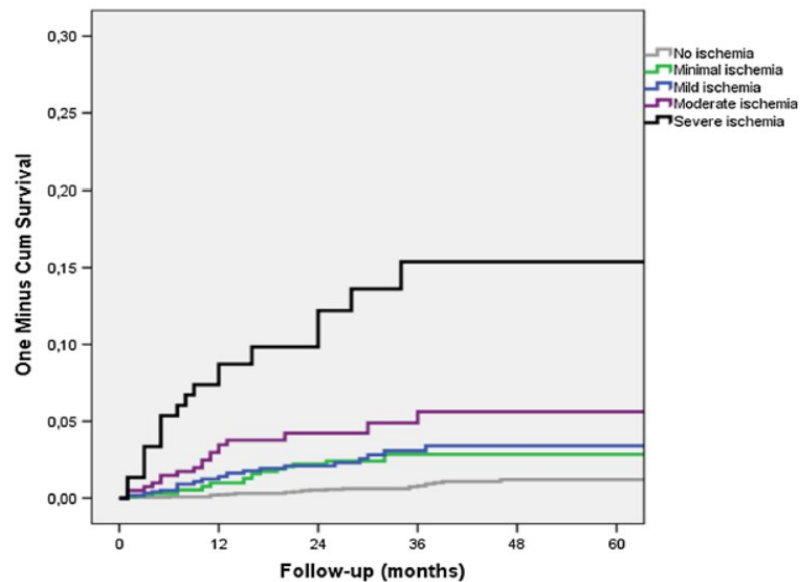
## Vessel-related ischemia (VRI): 1 vs 2 vs 3

MIS 1 No LAD	MIS 1 LAD	MIS 1 Both	MIS 2 No LAD	MIS 2 LAD	MIS 2 Both	MIS 3 No LAD	MIS 3 LAD	MIS 3 Both	MIS 4 No LAD	MIS 4 LAD	MIS 4 Both
13	14	15	16	17	18	19	20	21	22-28		
12	13	14	15	16	8	9	10	11	12		
11	12	4	5	6	7	8	9	10	11		
19	20	21	4	5	6	7	8	9	10		
9	10	11	12	13	14	15	16	17	18		
8	9	10	11	12	4	5	6	7	8		
5	6	7	8	9	3	4	5	6	7		
7	8	9	10	11	12	13	14	3	4		
3	4	5	6	7	8	3	4	5	6		
4	5	6	7	2	3	4	5	6	2		
0	1	2	3	1	2	3	4	2	3		

Nudi F, et al (J Nucl Cardiol 2015)



## Risk of death according to maximal ischemia score (MIS)



Patients at risk according to maximal ischemia score:

No	4161	4147	2663	1724	778	395
Minimal	1399	1380	628	457	137	85
Mild	1566	1507	757	506	197	109
Moderate	1020	915	558	381	184	78
Severe	568	510	348	256	153	66

Nudi F, et al (J Nucl Cardiol 2014)





# Non-invasive imaging in coronary syndromes: recommendations of the European Association of Cardiovascular Imaging and the American Society of Echocardiography, in collaboration with the American Society of Nuclear Cardiology, Society of Cardiovascular Computed Tomography, and Society for Cardiovascular Magnetic Resonance

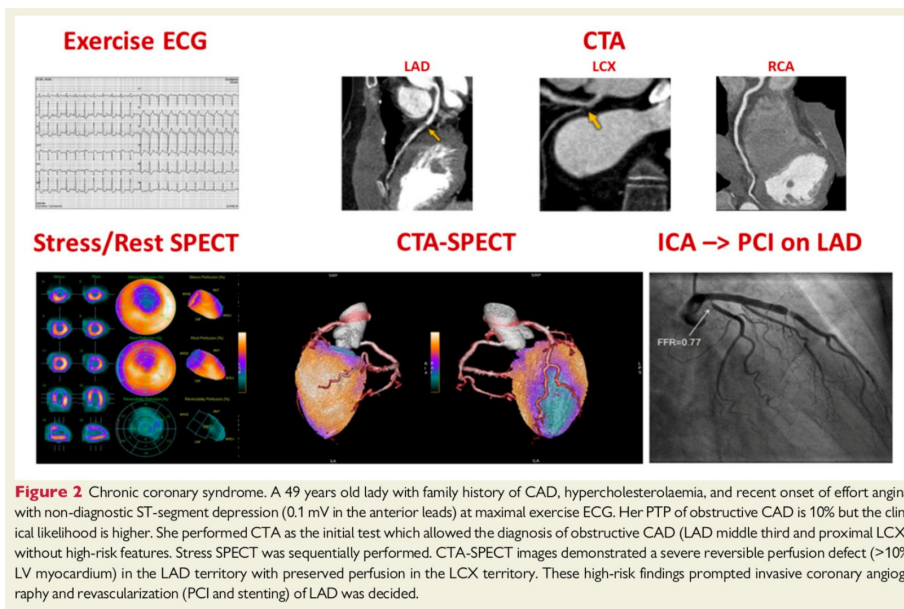
Thor Edvardsen<sup>1,\*</sup> (chair), Federico M. Asch<sup>2</sup> (co-chair), Brian Davidson<sup>3,4</sup>, Victoria Delgado<sup>5</sup>, Anthony DeMaria<sup>6</sup>, Vasken Dilsizian<sup>7</sup>, Oliver Gaemperli<sup>8</sup>, Mario J. Garcia<sup>9</sup>, Otto Kamp<sup>10</sup>, Daniel C. Lee<sup>11</sup>, Danilo Neglia<sup>12</sup>, Aleksandar N. Neskovic<sup>13</sup>, Patricia A. Pellikka<sup>14</sup>, Sven Plein<sup>15</sup>, Udo Sechtem<sup>16</sup>, Elaine Shea<sup>17</sup>, Rosa Sicari<sup>18</sup>, Todd C. Villines<sup>19</sup>, Jonathan R. Lindner<sup>20</sup>, and Bogdan A. Popescu<sup>21</sup>

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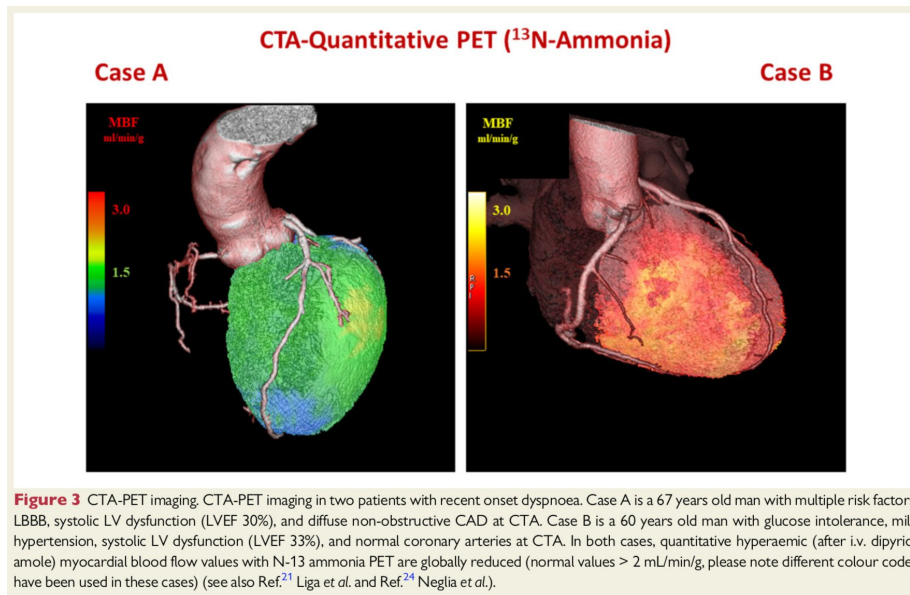


## Non-invasive imaging in coronary syndromes





## Non-invasive imaging in coronary syndromes





# Non-invasive imaging in coronary syndromes



## Conclusions and future directions

While multiple technologies and approaches to diagnose CAD in the acute and chronic stages have been developed, they address different aspects and stages of the disease. The imaging modality applied in any clinical situation should depend upon the information that is being sought. All modalities can provide information regarding LV structure and function, although echocardiography and CMR clearly have advantages. Similarly, with the general exception of CTA, all techniques can detect ischaemia and viability; although echocardiography and nuclear imaging have the largest imprint in clinical practice, CMR is increasingly being utilized. CTA is the non-invasive procedure of choice to visualize coronary anatomy. As none of them is perfect or can provide all the needed information, there is a need for clinicians to have a deep understanding of the disease within the coronary arteries and beyond. Such critical view of the disease and of our patients, together with the comprehensive knowledge of each diagnostic tool, will allow for development of the appropriate diagnostic strategies for each patient and situation, which is ultimately the goal of this document.

With the advent of newer technologies such as hybrid systems that combine nuclear imaging with CT, opportunities for obtaining complementary anatomic and functional information in a single imaging session may prove to have higher value than do current approaches. Furthermore, developments in high-definition imaging such as CT or CMR may allow for more detailed plaque evaluation and provide an opportunity to evaluate coronary plaques with a different goal, by assessing not only for stenosis and ischaemia, but to also evaluate plaque morphology and activity to improve the identification of patients at risk of acute coronary events.



## Hybrid imaging with CT and MPCl

**TABLE 4** Meta-Analysis for the Diagnostic Performance of Various Hybrid Cardiac Imaging Modalities

	N	Sensitivity (95% CI)	Specificity (95% CI)	Positive Likelihood Ratio (95% CI)	Negative Likelihood Ratio (95% CI)	Diagnostic Odds Ratio (95% CI)
Per-patient analysis						
SPECT/coronary CTA	5	0.92 (0.88–0.95)	0.90 (0.85–0.93)	10.38 (3.60–29.94)	0.08 (0.03–0.27)	158.16 (21.11–1185.00)
PET/coronary CTA	4	0.87 (0.80–0.92)	0.96 (0.92–0.99)	22.12 (5.20–94.00)	0.12 (0.05–0.29)	213.68 (25.94–1760.10)
CMR/coronary CTA	3	0.91 (0.83–0.96)	0.94 (0.88–0.98)	12.86 (5.90–28.02)	0.13 (0.07–0.26)	120.36 (35.42–408.98)
Per-vessel analysis						
SPECT/coronary CTA	3	0.91 (0.86–0.95)	0.95 (0.93–0.96)	18.51 (8.01–42.76)	0.11 (0.05–0.24)	174.33 (52.59–577.89)
PET/coronary CTA	3	0.81 (0.75–0.86)	0.97 (0.95–0.98)	28.42 (7.68–105.17)	0.15 (0.04–0.51)	202.03 (19.51–2091.8)
CMR/coronary CTA	2	0.80 (0.73–0.87)	0.87 (0.79–0.92)	6.37 (2.69–15.07)	0.13 (0.01–2.07)	53.95 (13.48–215.83)
CMR = cardiac magnetic resonance; PET = positron emission tomography; SPECT = single-photon emission computed tomography; other abbreviations as in Table 3.						

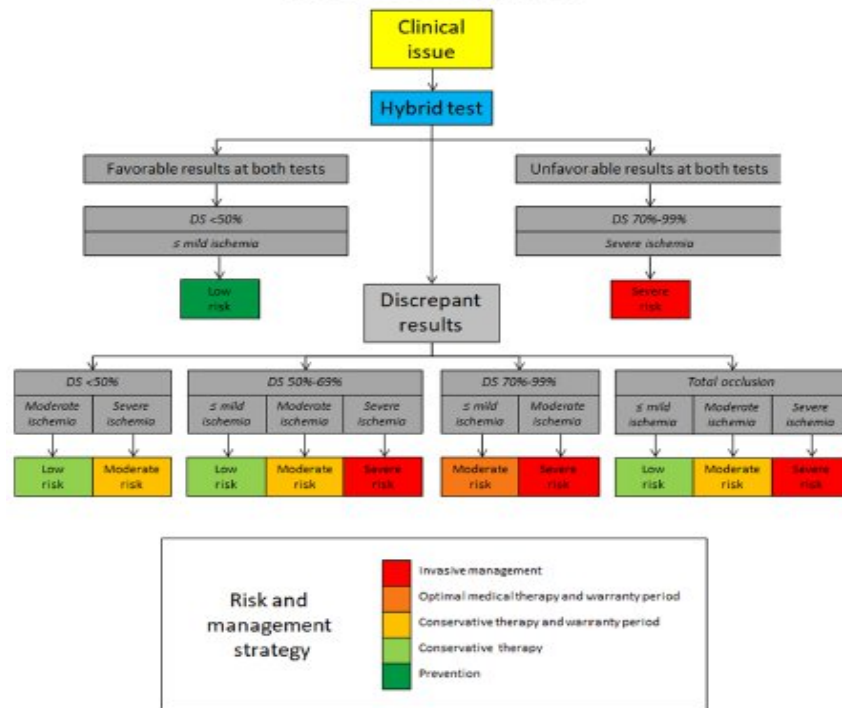
Rizvi et al, JACC Imaging 2017





# Hybrid cardiac imaging: always beneficial

## HYBRID TEST PARADIGM



Nudi et al, J Nucl Cardiol 2019





## Promise of hybrid SPECT/CT

*Revolution CT:*

CACS=0.7 mSv

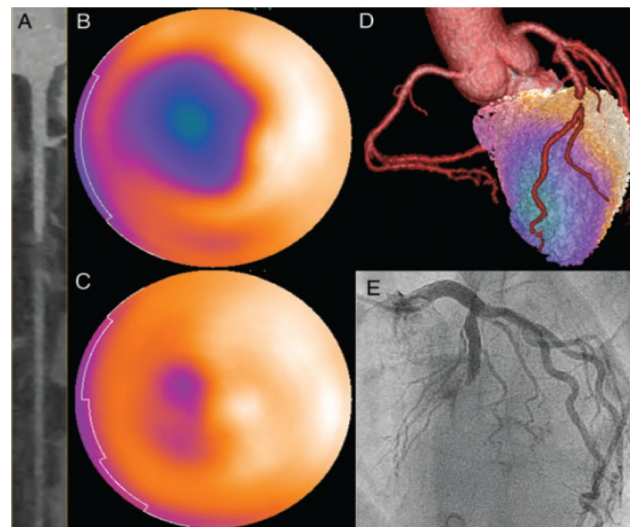
CCTA=0.5 mSv

*Discovery*

530NMc:

SPECT=0.9 mSv

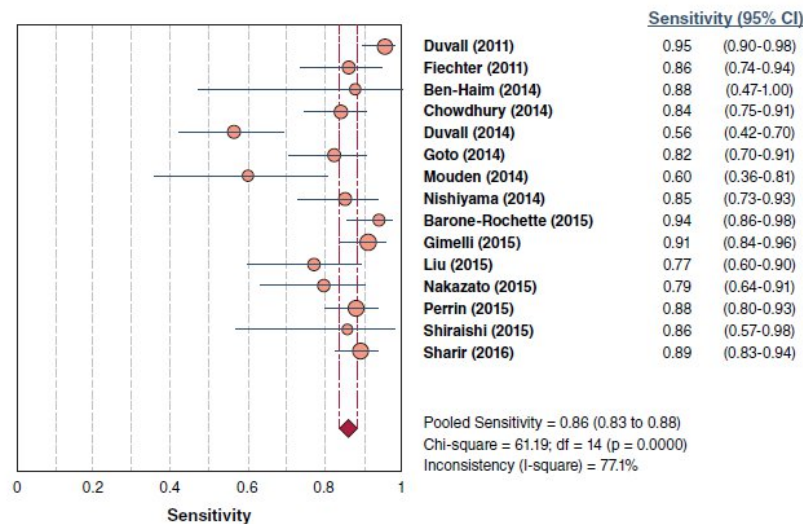
**TOTAL=2.1 mSv**





## CZT SPECT to diagnose CAD

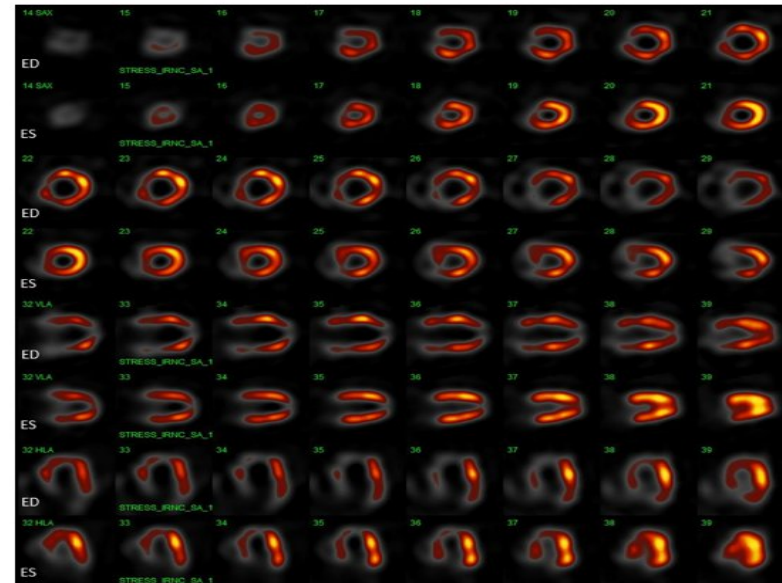
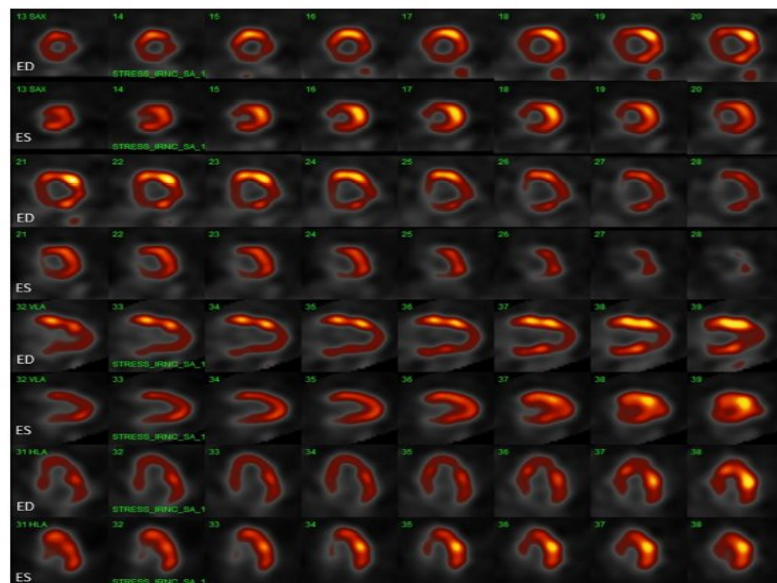
**FIGURE 2** Forest Plot of Univariate Analysis for Sensitivity



Heterogeneity was appraised using the chi-square test, with corresponding degrees of freedom (df) and p value. CI = confidence interval.



## Benefits of CZT SPECT for improved diagnosis



Nudi et al, J Nucl Cardiol 2017



## Myocardial Blood Flow (ml/min/g) with CZT SPECT

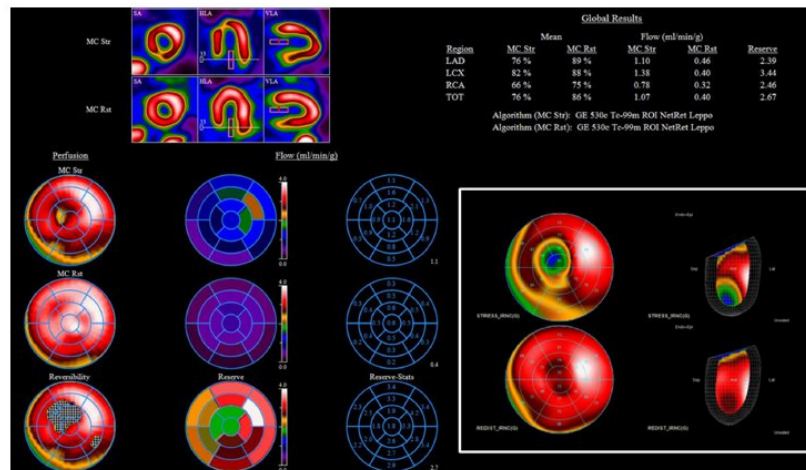
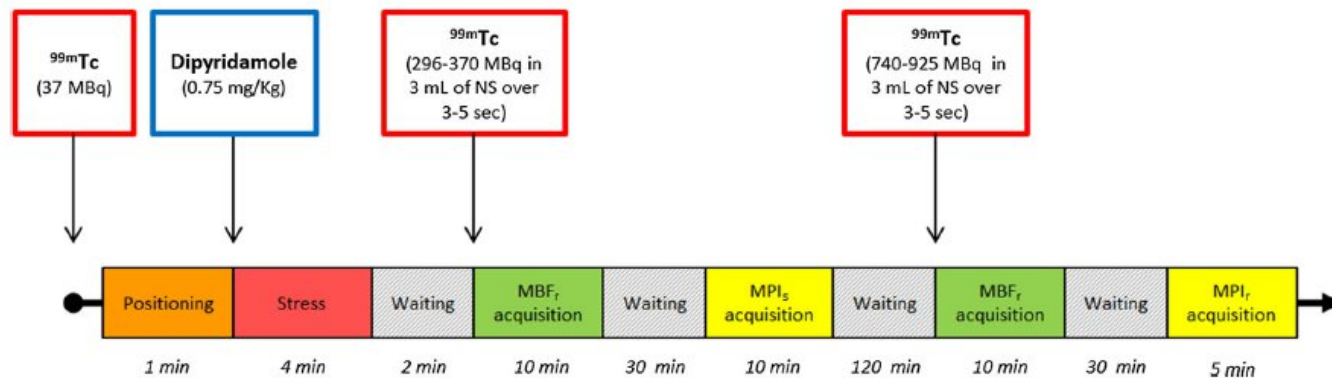


Figure 2. Myocardial blood flow and myocardial perfusion reserve (MPR), plus myocardial perfusion imaging (MPI) (right bottom inset) in a 60-year-old gentleman with effort angina, family history of coronary artery disease (CAD), active smoking, dyslipidemia and hypertension. MPI showed ischemia in a large apical (maximal ischemia score = 3) region and the inferior (maximal ischemia score = 1) region, whereas per-vessel and global MPR were apparently normal. Notably, MPR was conversely abnormal in the apical segments. This patient later underwent invasive coronary angiography which highlighted multivessel angiographically significant CAD, and was then treated with coronary artery bypass grafting.



## Myocardial Blood Flow (ml/min/g) with CZT SPECT



**Figure 1.** Study protocol. *MBF*, myocardial blood flow; *min*, minutes; *MPI*, myocardial perfusion imaging; *NS*, normal saline; *Tc*, technetium.



## Treadmill Exercise Produces Larger Perfusion Defects Than Dipyridamole Stress N-13 Ammonia PET

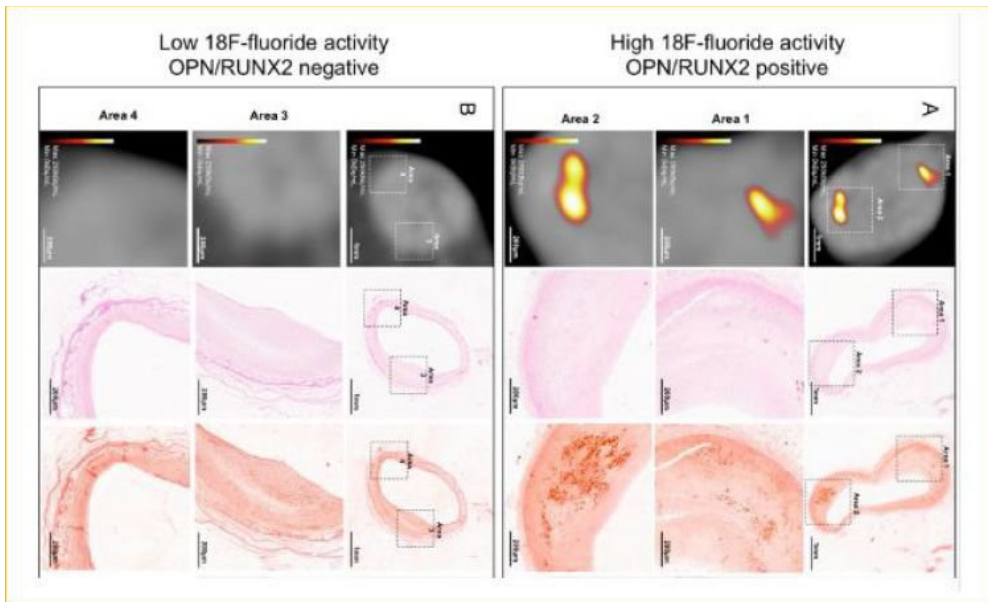
	Exercise Stress	Dipyridamole Stress	p Value	Correlation	Significance of Correlation
Summed stress score	9.1 ± 5.7	6.9 ± 5.9	<0.01	0.80	<0.001
Summed rest score	3.4 ± 3.3	3.2 ± 2.8	NS	0.79	<0.001
Summed difference score	5.8 ± 4.7	3.7 ± 4.6	<0.02	0.61	0.001
LV defect size (stress) (%)	19.3 ± 11.5	13.8 ± 13.6	<0.02	0.62	0.001
LV defect size (rest) (%)	8.1 ± 9.4	6.4 ± 6.5	NS	0.57	0.003





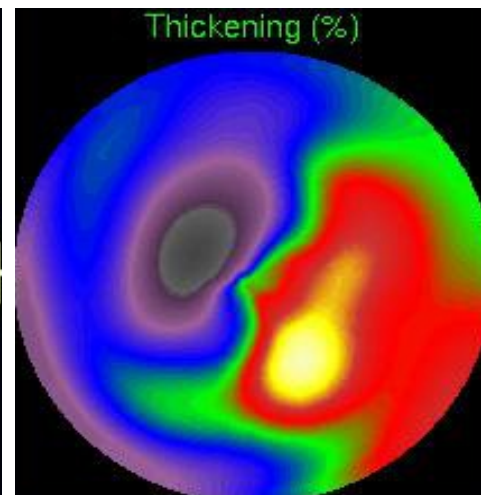
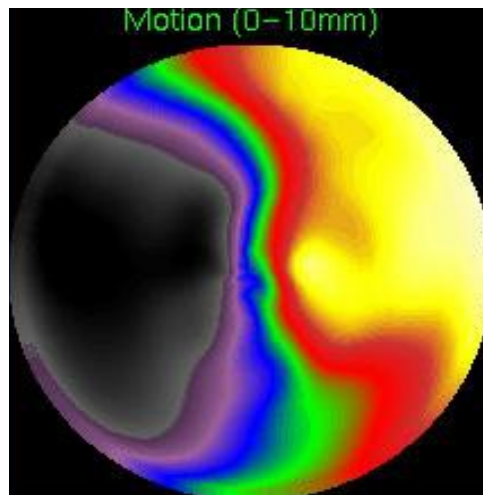
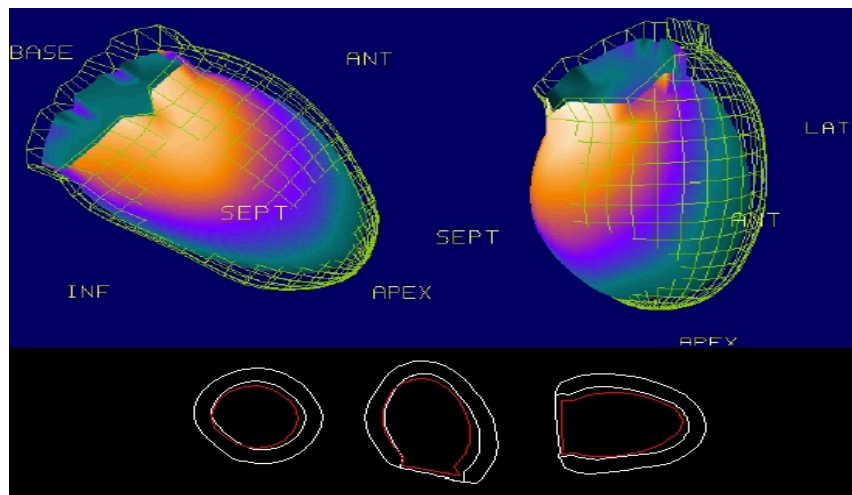
## Studio PRE18FFIR

Confronto tra imaging PET con 18F-fluoruro di sodio (lesioni “calde”), reperti istologici di placca aterosclerotica.



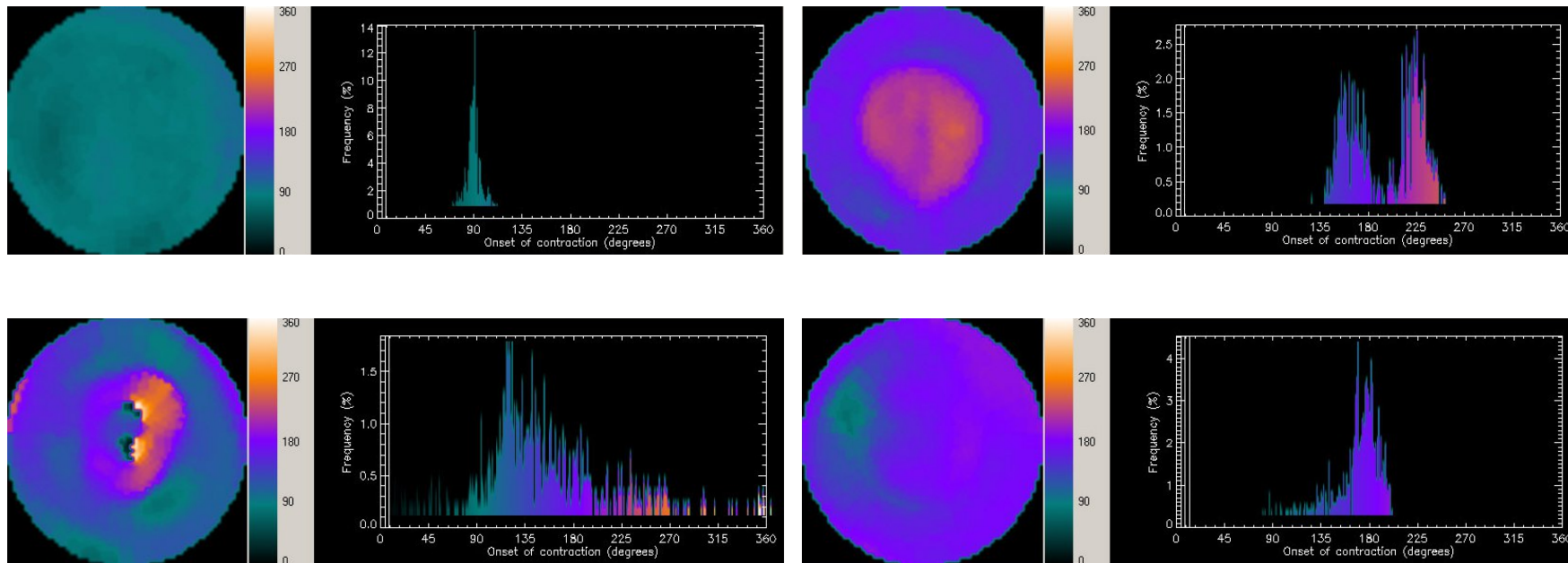


## SPECT Imaging



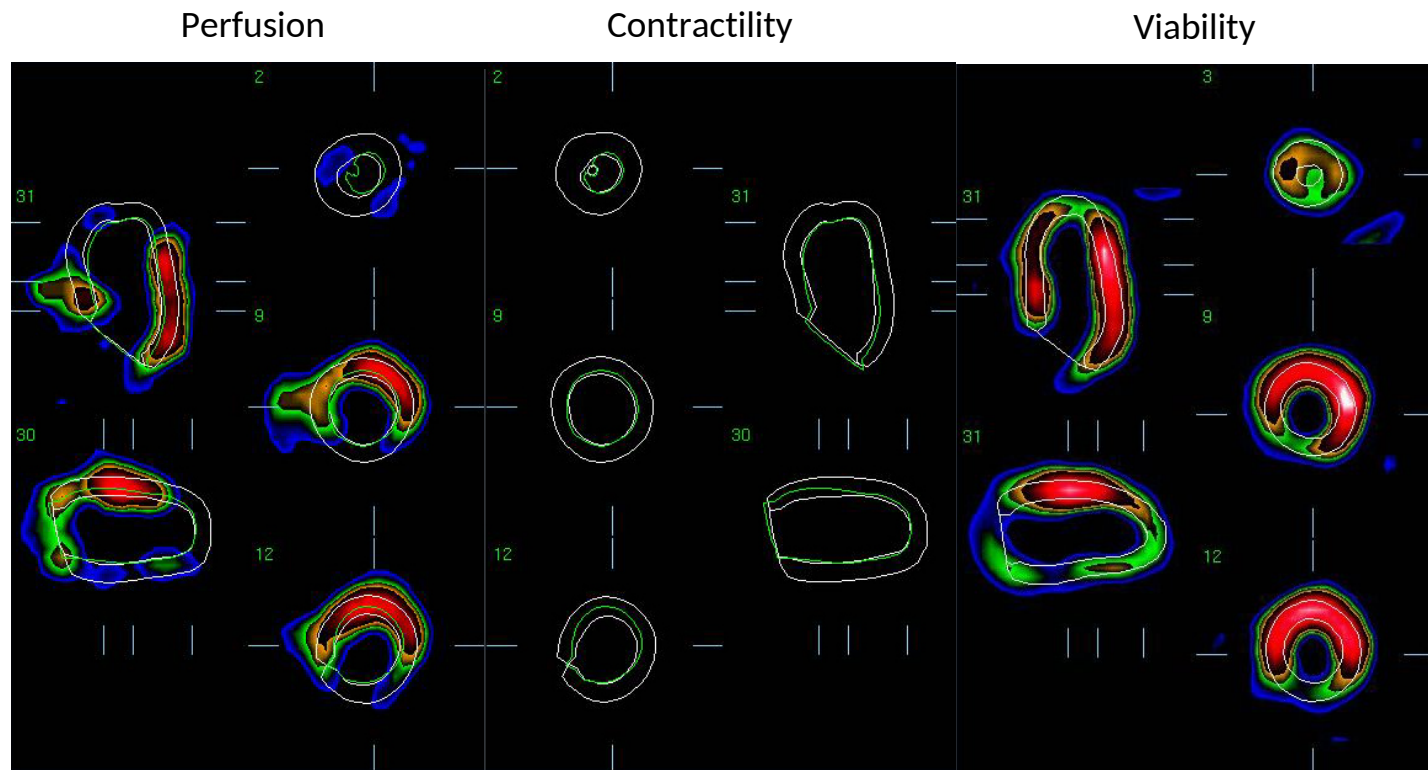


## Phase Analysis of Gated Myocardial Perfusion SPECT for the Assessment of Left Ventricular Dyssynchrony



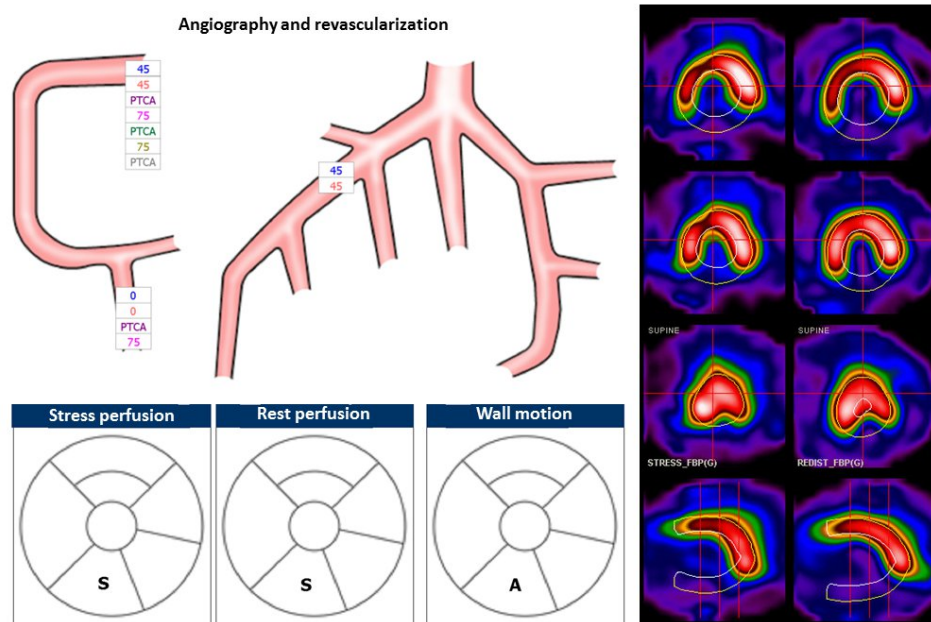


## Metabolism-Perfusion Mismatch





# Assessment of the Fate of Myocardial Necrosis by Serial Myocardial Perfusion Imaging





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**Nudi Francesco**







## Guidelines ESC 2019

