

Cardiac Imaging—Physiologic Assessment of Coronary Artery Lesion

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Abstract

Keywords

- ► fractional flow reserve
- coronary flow reserve
- instantaneous wave free ratio
- ► resting full cycle ratio
- coronary physiology

Thirty percent of angiographically significant coronary lesions may be functionally significant. Physiological assessment is essential to treat intermediate coronary lesions before taking the decision for percutaneous intervention. Even when coronary artery bypass is planned as treatment modality in triple-vessel coronary artery disease, the SYNTEX II trial has shown that functionally significant lesion treatment by bypass grafts improves the outcome of these patients. In this article, possible ways of estimating the physiologic assessment of coronary lesion, and the advantages and disadvantages of methods of estimating the physiologic assessment of coronary lesions is discussed.

Introduction

Even though the coronary angiogram is the gold standard test for the detection of the presence and severity of coronary artery disease, the physiological significance is better to treat a particular coronary lesion. The severity of coronary lesion only on angiogram may underestimate or overestimate. In addition, other than luminal narrowing, multiple factors also contribute to ischemia, like the length and serial lesions and the amount of viable myocardium. That is the reason why a similar degree of the lesion in two different patients differs in producing the severity of myocardial ischemia.^{1,2}

There are noninvasive methods also for the assessment of the physiologic significance of coronary lesions, like stress echocardiogram, magnetic resonance imaging, and nuclear myocardial perfusion scan. However, none give lesion-specific data within the coronary artery.² The fractional flow reserve (FFR) measurement was introduced in the 1980s. At present, the recommendation is to show the ischemia of lesion before doing percutaneous coronary intervention (PCI).³

Coronary Circulation

Epicardial coronaries serve a conduit artery function, whereas resistance arteries (100–400 μ m) and arterioles (>100 μ m) contribute to coronary resistance. Normal coronary blood flow at rest is 250 mL/min and can increase up to five times

during stress. The increase in oxygen demand during stress is met by increasing blood flow and dilatation of microcirculation (**-Fig. 1**). Myocardial ischemia results from epicardial coronary artery stenosis of 50% or microcirculatory dysfunction. To some extent, only coronary angiography (CAG) gives the anatomical severity of coronary artery disease (CAD); for physiological significance, we require different varieties of tests.

Methods to Determine the Physiologic Assessment of Coronary Artery Lesion

There are two methods to determine the physiologic assessment of coronary artery lesion: (1) invasive methods and (2) noninvasive methods.

Invasive Methods

The invasive method of estimating coronary lesions can be done along with coronary angiogram itself or at the time of planned PCI. There are different indices for the physiologic assessment of coronary artery lesion, invasively.

Fractional Flow Reserve

Fractional flow is derived by the ratio of the pressure distal to the lesion (Pd) to the pressure in the aorta (Pa). FFR is the ratio of the maximal achievable blood flow through the stenotic artery to the maximal normal flow (**-Fig. 2**). So, maximal

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Fig. 1 Metabolic control of coronary circulation. Abbreviations: PLA2, phospholipase A2; AA, arachidonic acid; PG, prostaglandins.



Fig. 2 Diagrammatic representation of FFR calculation.

vasodilation is mandatory for the calculation of the FFR. The original equation contains the venous pressure also in the calculation, but it is so low in normal conditions that is omitted. Heart failure and conditions associated with elevated venous pressures require consideration of the venous pressure also.⁴

Physics of FFR

On oxygen demand, autoregulation of the coronary circulation causes adequate coronary flow. When microvascular resistance

is low there is linear relation between the pressure and flow (**>Fig. 3ab**).

Types of FFR Wires

- 1. RADI FFR wire—Traditional FFR has hollow wire, sensor housing, and a tip. Sensor housing has a drift sensitive piezoelectric sensor, 10 mm away from the tip and there is abrupt transition proximal and distal to the sensor. Due to hollow wire body, torque control of the wire is poor.
- 2. COMET FFR wire—Due to Synergy's laser-cut high-torque sleeve there is improved deliverability and performance of the wire. It has optical pressure sensor which records accurate measurements and is prone to less pressure drift. Ashahi tip and coating facilitates better tip transition. In addition, it has free spin handle and wireless connection.
- 3. Dual-coil technology wire—It has Ashahi tip with actone with Synergy's laser-cut technology. This actone is a silicone coating on Ashahi ultrathin film with stainless steel core. The wire body is hydrophilic coated. This wire is highly torquable, flexible, and retains the tip shape for longer time.
- 4. Navvus microcatheter—RXi rapid exchange FFR system, featuring an ultrathin monorail pressure microcatheter, Navvus catheter, showed lesser FFR values than the traditional pressure wire, but this was not statistically significant.
- 5. iLab Polaris multimodality system—This is integrated with intravascular ultrasound (IVUS). This simplifies the procedure and decreases the procedure time.



Fig. 3 (a) Pressure and flow response in normal artery, during resting and hyperemic conditions. (b) Comparison of pressure and flow during resting and hyperemic conditions of stenosed artery with normal artery.

Preparation of the Patient

If intravenous (IV) adenosine is planned for hyperemia, then large-caliber access in the femoral or cubital vein is required. Theophylline should be stopped 12 hours before FFR, but β -blockers need not be stopped. Caffeine intake within 1 hour of FFR should be avoided.

Procedure

Even though the FFR wire passes through the 6F diagnostic catheter, a guide catheter (of at least 5F) without side holes is preferred for FFR due to its larger size and if FFR is positive, then PCI can be performed directly. Guide catheter with side holes may give false-positive FFR, as it creates a false gradient at the side hole and tip of the catheter. Also, maximal vasodilation by the adenosine may not be achieved as the drug partially goes to the aorta, not entirely into the coronary artery. It is better to avoid larger catheters and deep engagement. Coaxial engagement of the guide catheter is required. Anticoagulation should be given as per the PCI protocol. Nitroglycerin 100 to 200 µg should be given for epicardial vasodilatation.

For FFR measurement, proper zeroing of aortic pressure is required.

FFR is calculated by passing a specialized wire with a sensor distal to the coronary lesion. The catheter has to flush to remove the residual contrast, which can cause signal drift. Then equalization of the pressure sensor 1 to 2 mm distal to be tip of the guide catheter. Nonleaking thin introducer needle can be a leftover during FFR measurement, it is better to remove it otherwise. While wiring, minimal or no contrast should be used to prevent signal drift. The pressure transducer should be at least 20 to 30 mm distal to the lesion. The pressure distal to the lesion is represented as Pd, and at the guide that is aorta as Pa. Pd/Pa represents the basal gradient across the lesion. For a wire in a narrow vessel or severe tortuosity, the sensor may interact with the vessel wall and produce the artifacts. Pd/Pa ratio reflects FFR after maximum vasodilation. This means the important step in FFR calculation is the effective vasodilatation with pharmacological agents.

When there is a resting gradient <0.80, then measurement under hyperemia is not necessarily required to make a clinical decision. To achieve maximum hyperemia, it is better to give nitroglycerin to dilate epicardial coronaries first. IV adenosine (140 µg/kg/min) is preferred over intracoronary injection, especially where there are tandom lesions are there and the wire is required to be pulled back. If FFR is in the borderline zone (0.75-0.8), then IV infusion may be increased to 180 µg/kg/min. Usually left main coronary artery (LMCA) requires >200 µg and >100 µg for right coronary artery (RCA) if adenosine is given intracoronary. With adenosine transient, atrioventricular blocks can happen. Another drug that can be used for hyperemia is regadenoson. It is a selective A 2A receptor antagonist, and a single IV bolus injection of 400 µg induces maximal hyperemia within the 30 seconds. It can produce a burning nonischemic chest pain. Papaverine 8 mg for the right coronary artery and 12 mg for the left coronary artery can also be used for the induction of hyperemia. This drug may cause transient QT prolongation, and very rarely ventricular tachycardia or torsade pointes. Nitroprusside 0.6 µg/kg in bolus is another alternative drug for hyperemia.

Interpretation of FFR

FFR value is the lowest Pd/Pa ratio registered during the steady state. In the case of IV adenosine administrations, pressure values can fall to a minimum before reaching a steady state. These values should not be interpreted as FFR values. Ectopic beats or values measured under atrioven-tricular block/bradycardia can cause false-low FFR values, which must be excluded.

The normal value of FFR is 1. FFR value 0.7 means that 70% of the blood flow without stenosis is only flowing through the stenosis. The pioneer work of Pijls et al⁵ suggested that FFR < 0.75 is significant and requires revascularization.

This is arrived at by comparing with sequential tests of exercise-stressed tests, thallium scintigraphy, and dobutamine-stress echocardiography. This cutoff value was tested in the DEFER trial⁶ 140 µg/kg/min continuous IV infusion of adenosine is the gold standard for achieving maximal hyperemia.⁷ Even though IV adenosine is known to produce side effects like chest pain, atrioventricular blocks, and bronchospasm, all are of shorter duration and it is safe to use adenosine. A much simpler method of producing effective hyperemia is intracoronary 50 to 200 µg of adenosine bolus.8 However, this route of administration is not useful in the FFR measurement in sequential lesions where pullback is required. Other than adenosine, drugs used for hyperemia are nicorandil, regadenoson, nitroprusside, and papaverine. The advantage of nicorandil is that it produces vasodilation of both macro- and microvascular systems.9,10

Advantages of FFR

FFR is reliable at all heart rates, blood pressure, and contractility. It a high-reproducibility and low intra-individual variability. This is independent of gender and CAD risk factors. It varies less with common doses of adenosine than does CFR.

Limitations of FFR

FFR has limited value in a patient with microvascular dysfunction, previous myocardial infarction, left ventricular hypertrophy, and diabetic microangiopathy as nonviable myocardium due to previous myocardial infarction does not vasodilate during hyperemia. So with similar coronary stenosis, a patient with nonviable myocardium will show high FFR.

Controversies in FFR

The controversies in FFR are as follows:

1. Acute coronary syndrome (ACS) - In ACS patients, significant vasoactive substances from the ruptured plaque and thrombus are generated, which preclude the proper hyperemic response for FFR estimation. During acute myocardial infarction (AMI), false-negative FFR may occur as there is microcirculatory defect which causes decrease in flow across the lesion leading to high FFR. However, an FFR of <0.8 is truly functionally significant. But, DANAMI-3-PRIMULTI and Compare-acute trails in ACS situation showed 30% approximate relative risk reduction of the composite of cardiac death or myocardial infarction using FFR-guided treatment.

2. Assessment of serial stenoses—In sequential stenotic lesions, the calculation of FFR is not simple. There will be an effect of distal lesion on the proximal lesion and vice versa. By keeping the transducer in between two lesions is also not reliable. Keeping the wire distal to both lesions shows the combined significance of both lesions. We can calculate accurately by measuring the coronary occlusive wedge pressure during hyperemia, which is not a practical method.¹¹ Wire pull back during hyperemia has to be done first to see the step up at different lesions and determine the maximum pressure gradient producing lesion. To treat that lesion first and then reassess the FFR once again, to decide whether other lesion requires treatment or not.

Artifacts and Pitfalls in FFR

Pitfalls may be related to preparation, in measurement, or in the tracing interpretation. During preparation care should be taken to calibrate and equalize. During measurement attention should be paid to prevent drifting, wedging the catheter, and avoiding whipping of the pressure trace (**- Fig. 4**), and ensuring that proper hyperemia has occurred. The drift is suspected when both pressure curves are identical in shape with proper dicrotic notch but shifted parallelly. On the contrary, when there is true gradient between the two pressure tracings then both curves do not look identical



Fig. 4 Typical artifacts caused by mechanical interaction of the pressure sensor with vessel wall (whipping artifacts).

and have different morphologies; they may show diastolic pressure difference without a dicrotic notch. If guide catheter is wedged, then calculated FFR is going to be high, as wedging produces the diastolic fall of the aortic pressure. So, the true diastolic gradient between the aortic and distal pressures is less. While interpreting the tracing the cursor should be adjusted to record the actual FFR, but not the lowest recorded FFR which may falsely be very low immediately after changing the pressure transducer from closed position to open position after adenosine injection for recording.

Algorithm on How to Use FFR

Fig. 5 gives a flowchart to show the indications to the usage of FFR.

Indications and Guidelines for FFR

Concusses of multiple associations' recommendation of FFR is for the intermediate coronary lesions in chronic stable angina with Class IIa, level of evidence A.

Conditions Where FFR Is Useful

FFR is useful in the following conditions:

- 1. Pre-PCI
- 2. Post-PCI—Ideally, after PCI, 0.95 FFR is the best. Studies demonstrated that for up to 0.9 of FFR, there was no increase in cardiac events, but between 0.8 and 0.9, the event rate increased.
- 3. Bifurcation lesion—In the case of bifurcation lesion PCI, revascularization of the side branch is not necessary if FFR is >0.80 in the side branch.

FFR in Different Situations

LMCA—Some precautions have to be taken while doing the FFR to LMCA. FFR measurement has to be done in downstream vessels without stenosis, optimally in both the left circumflex coronary artery and the left anterior descending coronary artery.

Continuous intravenous administration of adenosine is preferred than bolus intracoronary administration of adenosine & withdrawal of the guide catheter from the ostium of LMCA, for FFR measurement.

Serial stenosis—It can only be evaluated jointly. The distally measured value represents the combined effect of both lesions. Values measured between two stenoses are



Fig. 5 Algorithm for FFR usage.

unreliable. If FFR of the combined effect of two lesions is <0.8, then stent the tighter lesion and reassess the FFR once again.

Why FFR was Not Popularized

Even though it is a guideline recommended for revascularization, doing FFR was confined to a few cases as it requires additional time for hyperemia. However, now these other modes of physiological evaluation hyperemia induction are not required.

Instantaneous Wave-Free Ratio

The concept of instantaneous wave-free ratio (iFR) is to see the flow and pressure in the wave-free period of diastole (**-Fig. 6**). This corresponds to the beginning from 25% into diastole to 5 milliseconds before the end of diastole. The major advantage is that hyperemia is not required, and individual lesion severity can be assessed in serial lesions. With the co-registration software, iFR values at different segments of coronaries can be displayed by moving the cursor on the angiogram. In turn, iFR gives lesion-specific data and allows even the estimation of the improvement expected by



Fig. 6 Physics of iFR.

treating the lesion and length of the stent required to get that improvement in the physiology of the lesion.

In the ADVISE study, the optimal cutoff value of iFR to predict FFR < 0.80 was 0.83 with sensitivity, specificity, positive predictive value, and negative predictive values of 85%, 91%, 91%, and 85%, respectively¹² (\succ Fig. 7).

According to the ADVISE 2 and RESOLVE studies, there is 80 to 85% concordance between iFR and FFR when iFR = 0.89.^{13,14}

Algorithm to Use iFR

Fig. 8 is a flowchart demonstrating the indications of iFR and the scenarios where iFR has to be done in addition to FFR.

Resting Full-Cycle Ratio

This measures the mean resting Pd/Pa over the entire cardiac cycle. It is simple and it does not require the detection of a specific point in the cardiac cycle.

Coronary Flow Reserve

The major limitation of both FFR and iFR is considering flow and pressure to be equivalent. Coronary flow reserve (CFR) directly measures the coronary flow (**- Fig. 9**). It is derived by the ratio of blood flow during maximal vasodilation divided by coronary blood flow during resting conditions. CFR is calculated by the thermodilution method or by Doppler flow velocity.

Normal CFR is 3.5 to 5. Studies have demonstrated that CFR of <2.7 was a risk factor for CAD. The ischemic threshold of CFR is <2.

Another drawback of FFR and iFR is that these indices detect only epicardial arterial disease, which constitutes only 10% of coronary circulation, whereas CFR can detect even microvascular disease, which accounts for 90% of the coronary circulation.

FFR wire from the Radi company can be used for calculation of CFR as the transducer is even thermosensitive. In addition, FFR wire can be used as a workhouse wire.

rCFR is the maximal flow in a coronary artery with stenosis to maximal flow in a normal coronary artery without stenosis. The normal range of rCFR is 0.8 to 1.0. The major limitations of this index are not applicable in multivessel



Fig. 7 Comparison of cutoff values of FFR and iFR.

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Fig. 8 Algorithm for iFR. Abbreviations: CSA, chronic stable angina; ACS, acute coronary syndrome.



Fig. 9 Concept of coronary flow reserve (CFR). FFR = Pd/Pa, BMR = Pd/Vd (basal), HMR = Pd/Vd (hyperemia). CFR = 1 + FFR[(BMR/HMR) - 1].

disease, in a patient with myocardial infarction, left regional ventricular dysfunction, or asymmetric hypertrophy. This is because the microvascular circulatory response is variable in the different regions of the myocardial bed, but rCFR assumes that microvascular circulatory response is uniform throughout the myocardial beds.

Hyperemic Stenosis Resistance

After primary PCI, there are no measures to detect microcirculatory dysfunction, which is one of the major determinants of short- and long-term prognosis in AMI. Resistive reserve ratio (RRR) expresses the microcirculation vasodilatory capacity. This is calculated by the ratio between basal and hyperemic microcirculatory resistance (**Fig. 10**). This detects the suboptimal reperfusion. In one study, it was shown that RRR correlated better than CFR or index of myocardial resistance (IMR) in predicting the events rates at six months.

Microvascular Assessment

Among the deferred PCI patients of FAME 2, 14.6% had persistence of angina and 9% had clinical events at 2-year



Fig. 10 Diagrams to show the cause of increased coronary resistance. (a) R2—at the arteriolar level. (b) R3—at the intramural prearteriole. P1 and P2 = pressures proximal and distal to the abnormal resistance, respectively; R1 = the normally responsive subepicardial arteriole.

follow-up.¹¹ This observation gives the information that the ischemia is not entirely contributed by the epicardial lesion.¹⁵ The coronary circulation contains three important components—conductive epicardial coronary arteries, arterioles, and capillaries. Ischemia in any one of these three components of coronary circulation produces myocardial ischemia.

The studies showed that CFR could be used to risk-stratify in high FFR patients.¹⁶⁻¹⁸ An index of microcirculatory resistance (IMR) can quantify the degree of microvascular dysfunction by using distal coronary pressure. IMR reflects more precisely the microvascular dysfunction than CFR, which is a combined assessment of the macro- and microcirculation.

In **Fig. 11** the FFR_{myo} is explained.^{19,20} The ratio that represents the fraction of normal maximum flow that is preserved despite the presence of the stenosis is called FFR_{myo}.

The myocardial blood flow depends on the basal pressure gradient across the lesion (FFR_{myo}), but not on the hyperemic pressure gradient. As shown in **– Fig. 12**, FFR_{myo} does not change even though the pressure change occurs, so FFR_{myo} is considered as lesion-specific index of stenosis severity.²¹

Studies on Invasive Physiologic Assessment of Coronary Artery Lesion

- FFR Studies-FFR is useful in many intermediate coronary lesions including LMCA,²² except in tortuous heavily calcific lesions, where negotiation of relatively till stiff FFR wire is difficult. In ► Table 1 important landmark trails of FFR are mentioned.
- 2. iFFR STUDIES–In ► Table 2 iFR studies are mentioned.

Noninvasive Methods

FFR assessment can be done reliably by complete noninvasive method (e.g. CT scan) or by partial noninvasive method (e.g. FFR angio).

1. CT-derived FFR—Noninvasively FFR can be calculated using computational fluid dynamics (CFD) technology with CT (FFRCT). With cardiac computed tomography angiography (cCTA), the first three-dimensional (3D)

Fig. 11 Concept of FFR_{myo}

Fig. 12 Independence of FFR_{mvo} from the loading conditions.

Table 1 Landmark trails with FFR

anatomical coronary arterial model is constructed. Using allometric scaling law, total coronary flow is estimated from myocardial mass, from which the basal coronary outlet resistances are determined. With the morphometric law, the resistance of the downstream vessel to the vessel size at each outlet is determined. A mathematical model of hyperemic condition simulating adenosine-induced vasodilatation is derived. Then CFD analysis is performed based on the discretized model of patient-specific geometry and boundary conditions to numerically solve the governing equations of fluid dynamics, that is, Navier Stokes equations, as a Newtonian fluid. To detect the spatial distribution of FFRCT, the numerical solutions of coronary flow and pressure fields are computed.

2. FFRangio—The CathWorks FFRangio software device gives information of FFR noninvasively by deriving the flow dynamics from two conventional angiographic views of 25° apart. The FFR values at each segment of angiogram are displayed when the cursor is moved on the 3D reconstructed angio. The major advantage of

Study Name	Clinical presentation	FFR cutoff	Outcomes
DEFER ²⁷ 2001	intermediate stenosis in CSA	0.75	No benefit stenting a nonischemic stenosis
FAME ²⁸ 2009	Multivessel CAD	0.80	FFR to all lesions in multivessel PCI with DES, improves 1-y MACE
FAME 2 ²⁹ 2012	Stable CAD and hemodynamically significant stenoses	0.80	Even though angiographically severe lesion, but FFR negative, then OMT showed excellent outcomes
Muller et al ³⁰ 2011	Angiographically intermediate isolated proximal LAD stenosis	0.80	Angiographically equivalent lesion, but FFR negative, then OMT showed excellent outcomes
Mayo Registry ³¹ 2013	Patients undergoing PCI, excluding STEMI or cardiogenic shock	<0.75→PCI 0.75–0.80→Operator discretion>0.8→OMT	FFR-guided treatment strategy is associated with a favorable long-term outcome with decreased MACE
Van Belle et al ³² 2014	Angiographically ambiguous lesion	0.80	FFR during diagnostic angiography is safe and reclassifies 50% of the patient into other category of revascularization
RIPCORD ³³ 2014	Stable angina	0.8	FFR guides for which coronary artery lesions to be treated
DANAMI 3 PRIMULTI ³⁴ 2015	Patients with STEMI with multivessel disease and undergone only PCI of an infarct-related coronary artery	0.80	FFR-guided complete staged revascularization, reduces the risk of future events
Compare-Acute ³⁵ 2017	Patients with STEMI with multivessel disease and undergone only PCI of an infarct-related coronary artery	0.80	FFR-guided complete revascularization in acute condition, reduces the MACE
IRIS-FFR ³⁶ 2017	Patients with at least one coronary lesion	0.75	The risk of MACE was not significantly different between deferred and revascularized lesions depending on FFR cutoff value of 0.75
FORZA ³⁷	In patients with angiographically intermediate coronary lesions (AICL), FFR vs. OCT	FFR ≤ 0.80	The two tests offer different benefits, with OCT holding a slight edge

Note: SYNTEX II, ongoing; FAME 3, ongoing, J, confirmed, ³⁸ PRIME FFR.³⁹

Study name	Clinical presentation	iFR cutoff	Outcomes
DEFINE PCI ⁴⁰	Pts with stable or unstable angina with either multivessel or multilesion coronary artery disease with an abnormal baseline iFR	iFR ≤0.89	Residual ischemia after angiographically successful PCI was seen in 24.0% pts by iFR estimation (0.84 \pm 0.06)
ADVISE II ¹³	To assess the clinical value of iFR to characterize, coronary stenosis severity as determined with FFR	iFR 0.89	Minimum iFR of 0.89 has FFR agreement is ≥80%
RESOLVE ¹⁴	FFR and iFR in CSA patients		Endothelial dysfunction (by EndoPAT), plaque superficial irregularities (by OCT), and gene polymorphisms (in ENOS and HO-1) can be different in patients with discrepancy between FFR and iFR
DEFINE—FLAIR ⁴¹	iFR (n = 1,242) vs. FFR (n = 1,250)	iFR value was ≤0.89	The primary outcome, incidence of all-cause death, MI, or unplanned revascularization at 12 months, occurred in 6.8% of the iFR group vs. 7.0% of the FFR group ($p < 0.001$ for noninferiority)
SWEDEHEART ⁴²	Patients with CSA or ACS randomized to treatment guided by iFR or FFR	iFR value was ≤0.89	An iFR-guided revascularization strategy was noninferior to an FFR-guided revascularization strategy for MACE at one year

Table 2 Important iFR studies

Abbreviations: ADVISE II, ADenosine Vasodilator Independent Stenosis Evaluation II; DEFINE—FLAIR, Functional Lesion Assessment of Intermediate Stenosis to Guide Revascularisation; DEFINE PCI, Physiologic Assessment of Coronary Stenosis Following PCI; RESOLVE, multicenter core laboratory comparison of the instantaneous wave-free ratio and resting Pd/Pa with fractional flow reserve; SWEDEHEART, coronary angiographic findings and outcomes in patients with sudden cardiac arrest without ST-elevation myocardial infarction.

FFRangio is that it does not require a wire, contrast, or vasodilator, and can be done on proper images of angiogram only.

Studies on CT-Derived FFR

- 1. DISCOVER-FLOW (Diagnosis of Ischemia-Causing Stenoses Obtained via Noninvasive Fractional Flow Reserve) virtual PCI technology—Diagnostic results from this noninvasive version of FFR correlated well with conventional FFR (R = 0.72; p < 0.0001), with a slight underestimation (0.022 ± 0.116; p = 0.016) at the per-vessel level but no difference on a per-patient basis (p = 0.131). Use of CT-derived FFR yielded a false-positive rate of 11.3% and a false-negative rate of 4.4%.²³
- 2. DeFACTO—In CSA patients, FFRCT improved the diagnostic accuracy in detecting the severity of coronary lesion. But a major drawback of this study is that it did not achieve the prespecified primary goals.²⁴
- 3. NXT trail—In this study FFRCT was compared with invasive FFR and showed high diagnostic accuracy for CAD detection. Also, FFRCT increased the specificity for CAD detection when compared with only anatomical assessment of coronary anatomy by CTA.²⁵
- 4. PLATFORM—In CSA patients, both groups (invasive coronary angiography vs. CTA with selective FFRCT) showed same clinical outcomes and quality of life at 1-year follow-up.²⁶

Conflict of Interest

None declared.

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