



La Technologie au service du coronarien

Les approximations de la FFR

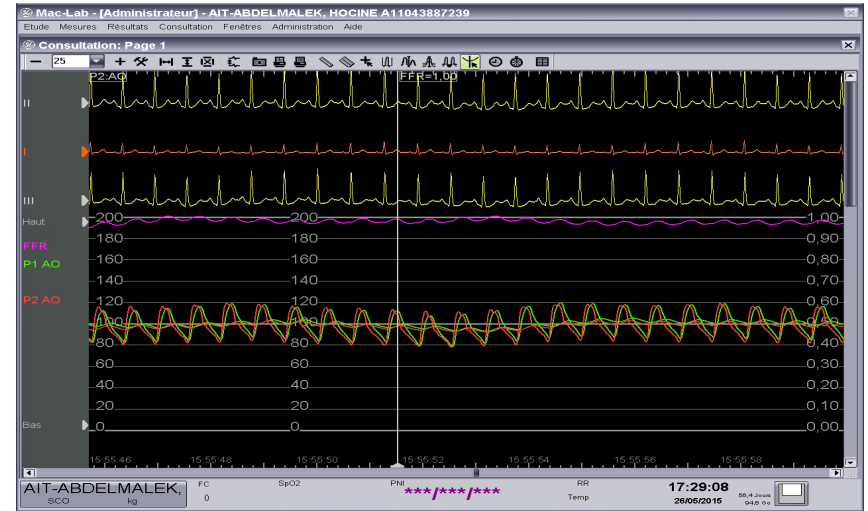
Patrick Dupouy

PCVI Antony-Melun

Les approximations techniques de la mesure de FFR

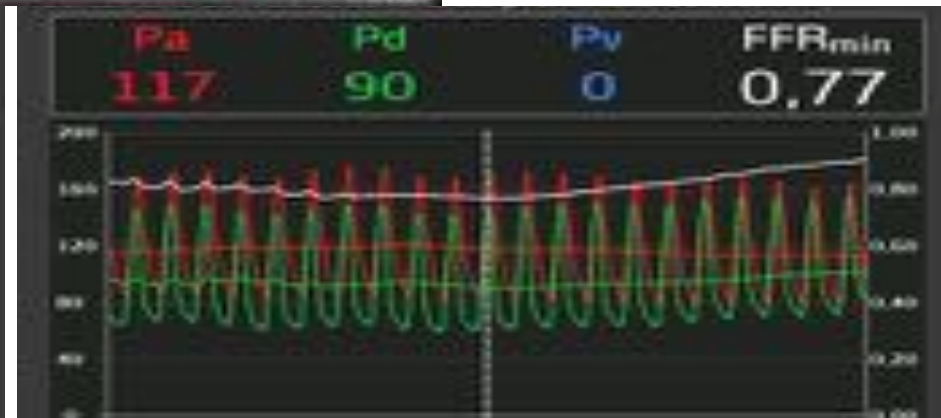
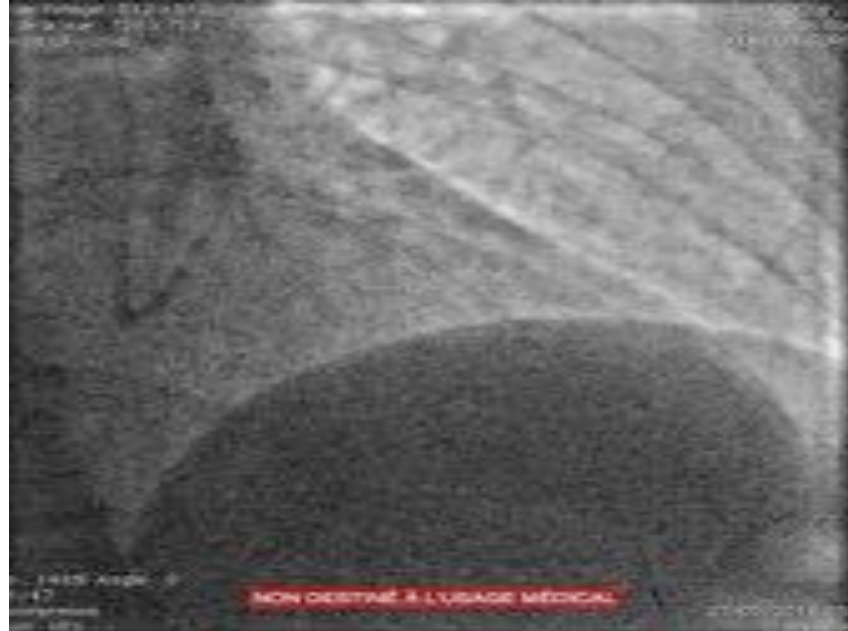
- Dose d'adénosine
- Stabilité de la ligne iso et du signal
- Sonde coro ou sonde ATL (4F)
- Sonde dans l'ostium ou dégagée
- Hémodynamique de base

Stabilité du signal



Variations hémodynamiques



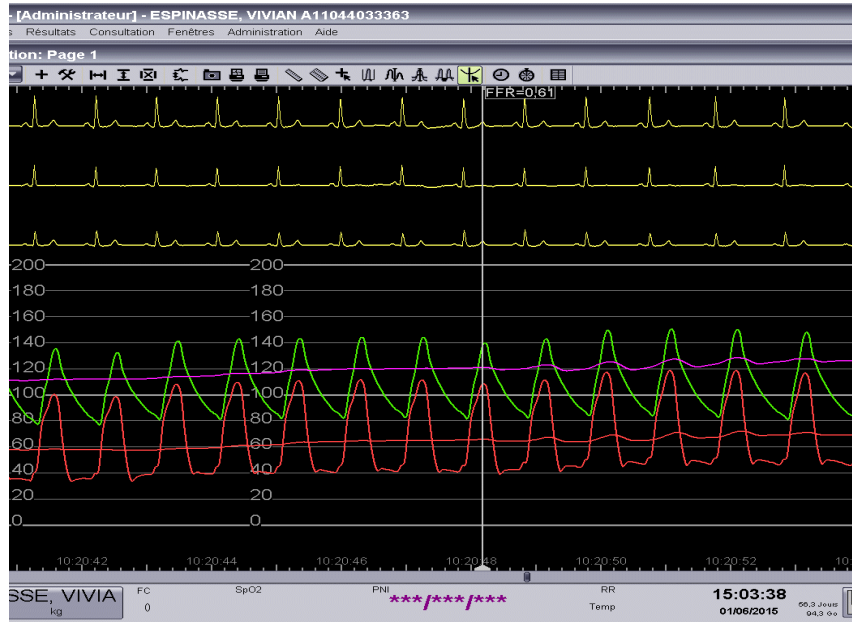




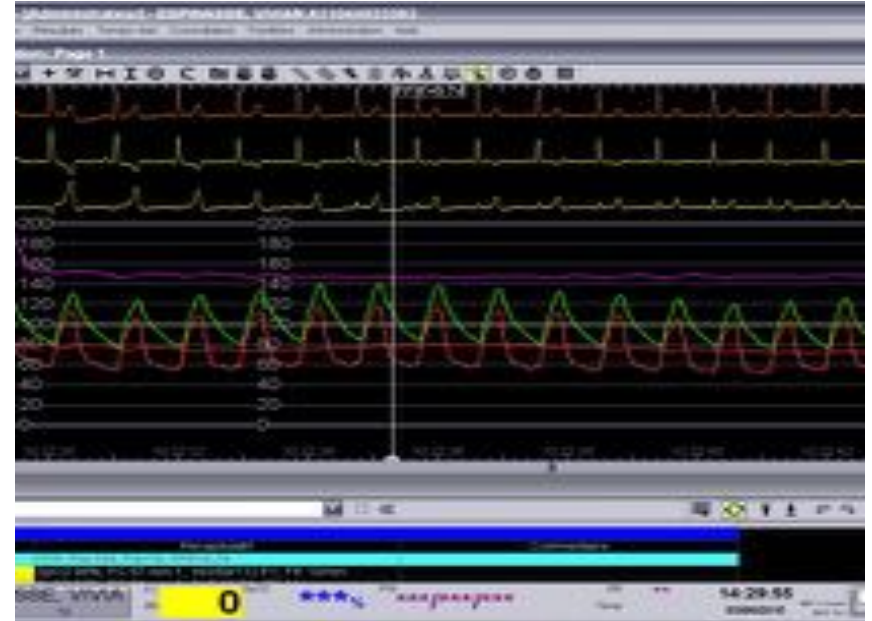
Vasodilatation épicardique



PRE RISORDAN FFR 0,60

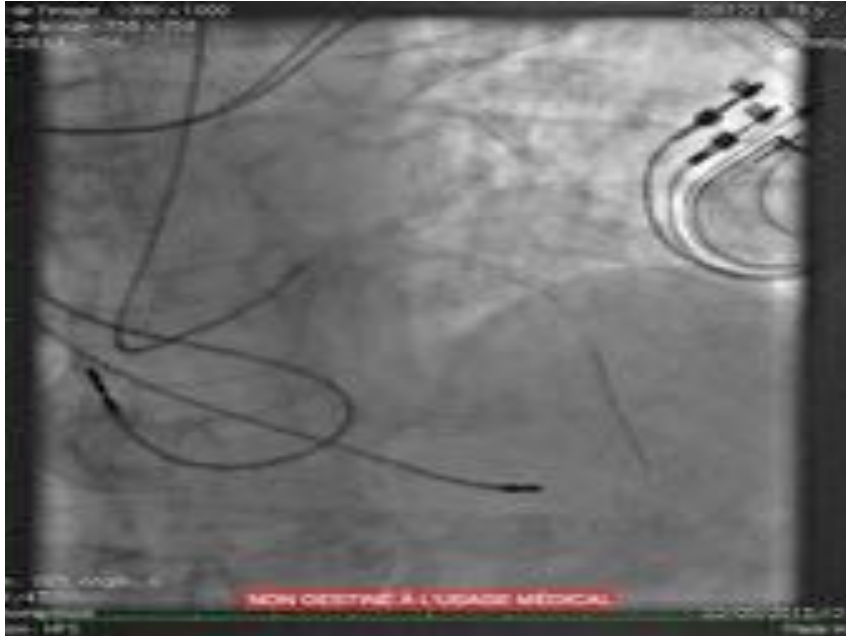


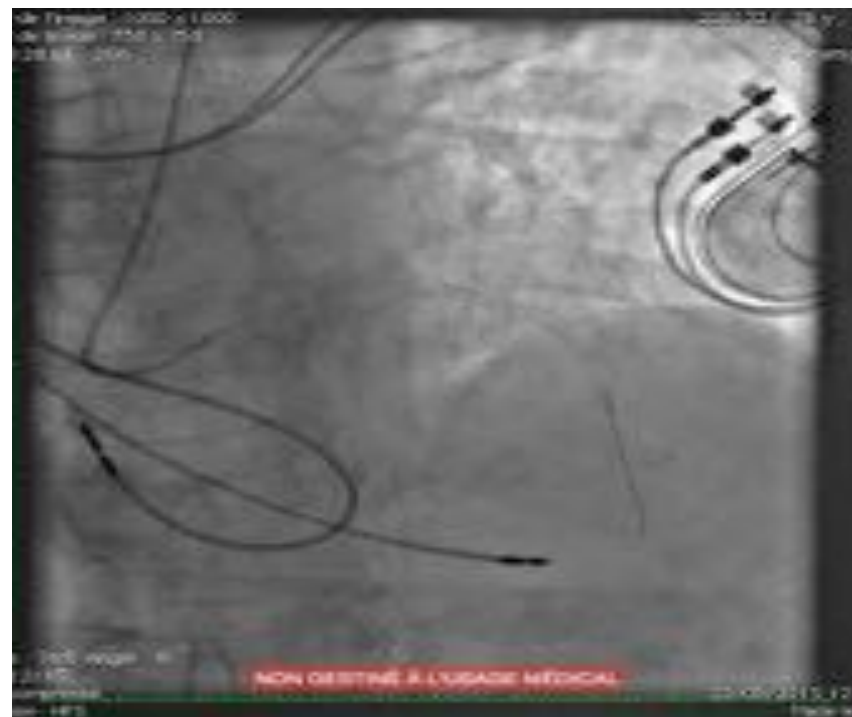
POST RISORDAN FFR 0,74

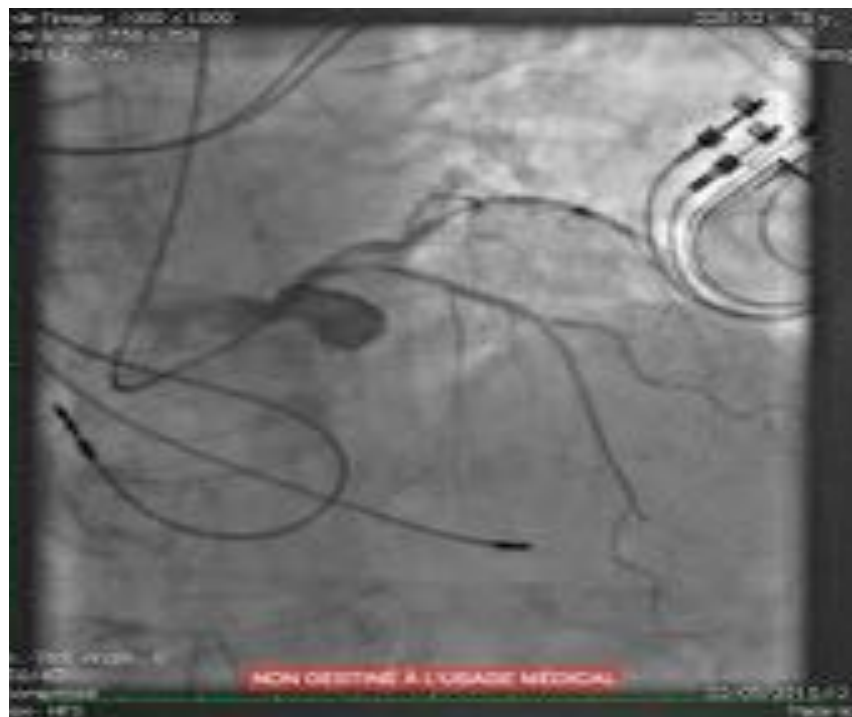


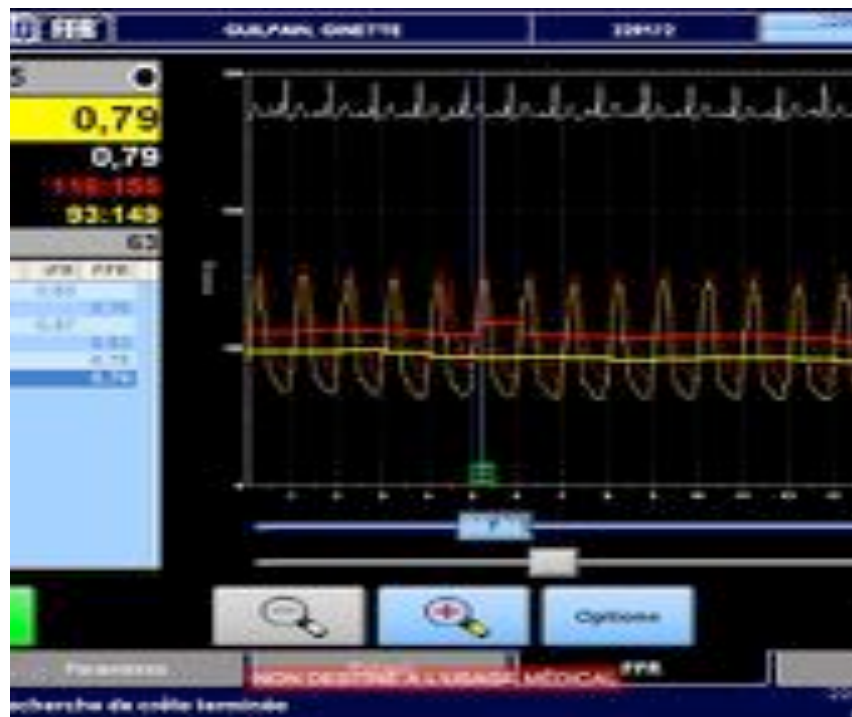
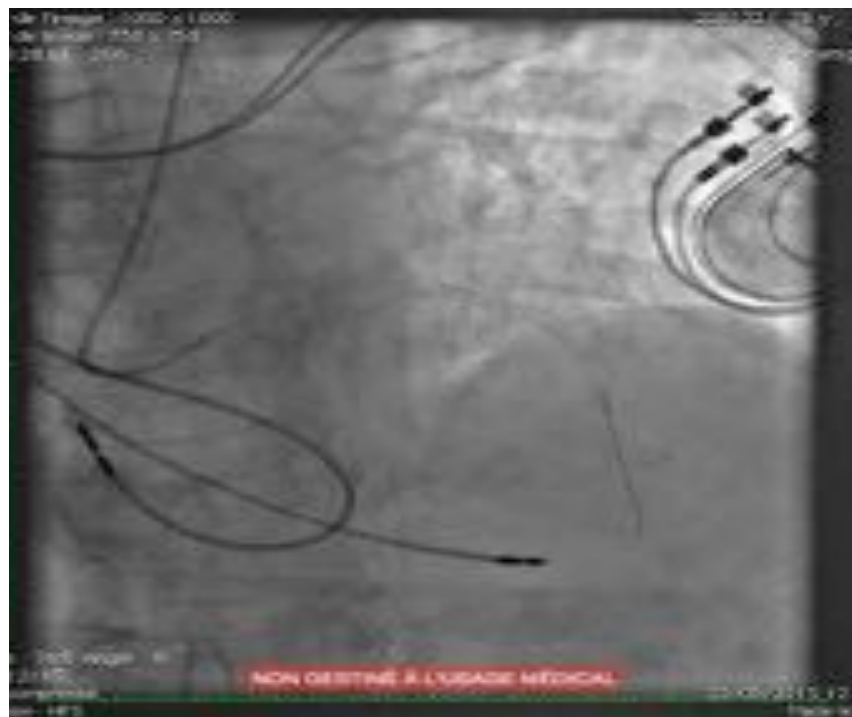


Se dégager de l'ostium

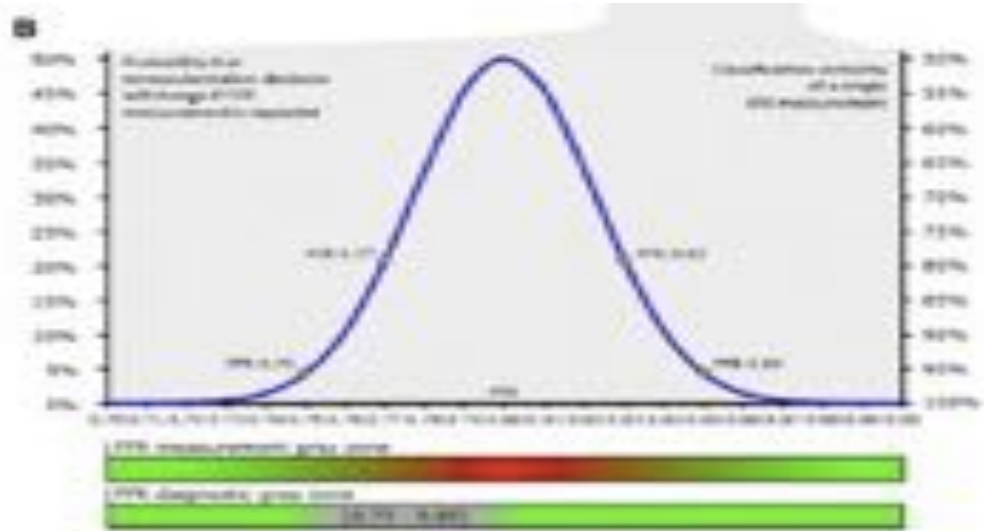
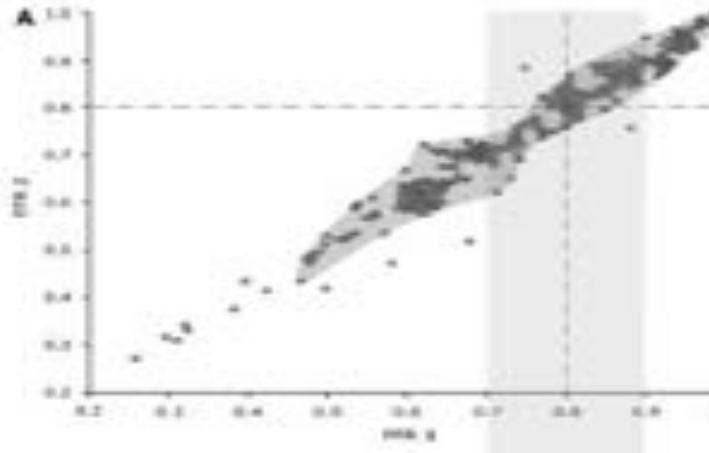








Une FFR peu en cacher une autre ..



Des Alternatives à l'adénosine ?

- Gradient de repos
- FFR contraste
- iFR
- FFR Adénosine
- FFR Adénosine + contraste
- Quel intérêt ?
- Gain de temps
- Moins d'injection
- Moins de toxicité ?

TABLE 2. Mean P_2/P_1 Ratio, Time to Peak Action, Duration of the Plateau Phase, and Electrocardiographic Changes Induced by the Different Vasodilatory Stimuli in Group 1 Patients

	P_2/P_1	Time to Peak Action, s	Duration of the Plateau Phase, s	Pain Score, 0-10
Place IC 20 mg	0.87 ± 0.20	33 ± 5	33 ± 7	0
Ado IC 20 mg	0.82 ± 0.20	13 ± 2	7 ± 2	0
Ado IC 40 mg	0.82 ± 0.19	15 ± 2	6 ± 1	0
ATP IC 20 mg	0.82 ± 0.20	14 ± 3	4 ± 1	0
ATP IC 40 mg	0.80 ± 0.16	14 ± 3	3 ± 1	0
CM IC 0 mL	0.88 ± 0.21	10 ± 3	2 ± 1	0
Ado fem $140 \mu\text{g} \cdot \text{kg}^{-1} \cdot \text{min}^{-1}$	0.81 ± 0.18	89 ± 5	NA	5.3 ± 1.4
Ado fem $180 \mu\text{g} \cdot \text{kg}^{-1} \cdot \text{min}^{-1}$	0.81 ± 0.18	NA	NA	6.1 ± 2.7
ATP fem $140 \mu\text{g} \cdot \text{kg}^{-1} \cdot \text{min}^{-1}$	0.81 ± 0.19	79 ± 29	NA	4.8 ± 3.5
ATP fem $180 \mu\text{g} \cdot \text{kg}^{-1} \cdot \text{min}^{-1}$	0.81 ± 0.17	NA	NA	4.8 ± 3.5
Ado per1 $140 \mu\text{g} \cdot \text{kg}^{-1} \cdot \text{min}^{-1}$	0.81 ± 0.16	113 ± 48	NA	4.3 ± 2.7
Ado per1 $180 \mu\text{g} \cdot \text{kg}^{-1} \cdot \text{min}^{-1}$	0.80 ± 0.17	NA	NA	6.0 ± 3.46
ATP per1 $140 \mu\text{g} \cdot \text{kg}^{-1} \cdot \text{min}^{-1}$	0.81 ± 0.17	104 ± 36	NA	4.3 ± 1.4
ATP per1 $180 \mu\text{g} \cdot \text{kg}^{-1} \cdot \text{min}^{-1}$	0.82 ± 0.20	NA	NA	3.8 ± 2.7
Place IC 20 mg	0.87 ± 0.18	21 ± 3	21 ± 6	0

Values are mean $\pm$ SD. Place indicates placebo; Ado, adenosine; CM, contrast media; IC, intracoronary; fem, femoral artery; and per1, peripheral vein.

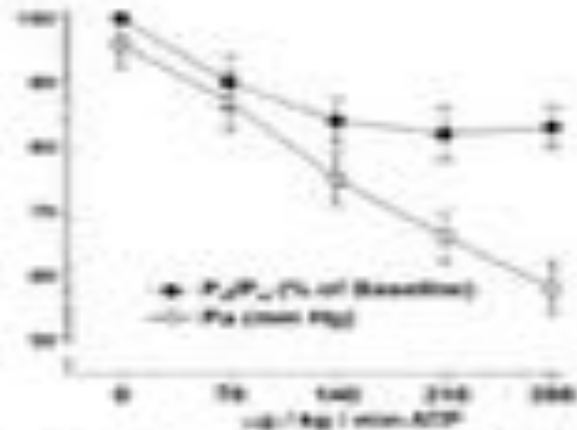
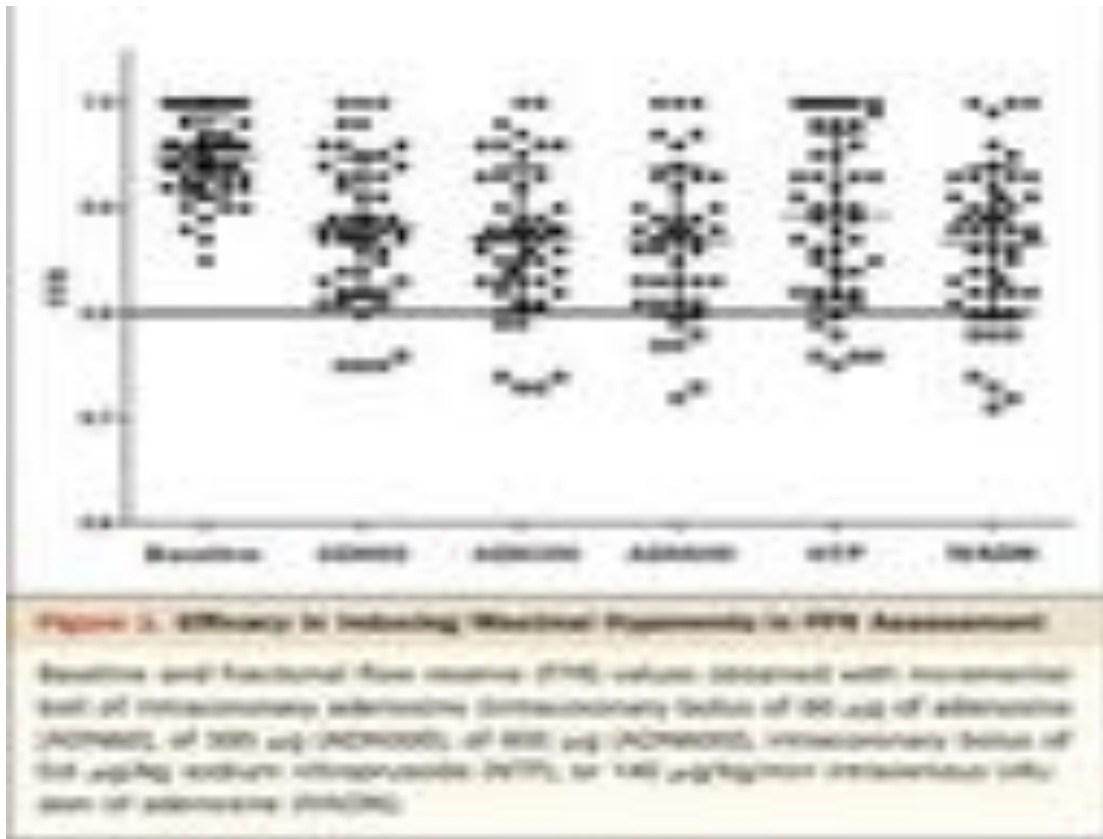


Fig 4. Plots of the mean values ($\pm$ SEM) of P_2 and of P_2/P_1 in function of the dosage of intravenous infusion of ATP (2 patients [stenotic artery]). Above a dosage of 140 min^{-1} , no further decline in P_2/P_1 was observed despite decline in mean P_2 .

Effets de l'adénosine

Table 2. Results: Effect of Different Doses of IC or IV Adenosine and IC Sodium Nitroprusside on Profile, Symptoms, Atrioventricular Block, and FFR

	n	HR (beats/min)	SBP (mm Hg)	DBP (mm Hg)	MBP (mm Hg)	Symptoms	AVB	FFR
Baseline	30	70 ± 8 71 (64–75)	144 ± 20 146 (130–160)	79 ± 9 80 (74–82)	101 ± 11 100 (95–107)	0 (0%)	0 (0)	0.96 ± 0.04 0.95 (0.92–0.98)
ADN60	30	68 ± 12 68 (63–75)	142 ± 22* 140 (130–150)	75 ± 10† 75 (70–80)	97 ± 12† 96 (90–103)	1 (3%)†	8 (18%)	0.88 ± 0.07† 0.88 (0.83–0.93)
ADN300	48	64 ± 12 66 (57–74)	144 ± 23* 142 (135–160)	74 ± 9† 75 (70–80)	97 ± 10† 97 (91–103)	4 (8.3%)†	13 (27%)	0.87 ± 0.07† 0.87 (0.83–0.93)
ADN600	43	62 ± 12 64 (50–73)	143 ± 25* 144 (130–156)	76 ± 9† 75 (70–80)	96 ± 10† 97 (91–102)	3 (6.9%)†	10 (23%)	0.87 ± 0.07† 0.87 (0.81–0.93)
ICN	48	74 ± 11 75 (68–82)	123 ± 19† 120 (110–130)	67 ± 8† 66 (60–72)	84 ± 10† 87 (80–93)	4 (8.4%)†	0 (0%)	0.88 ± 0.07† 0.90 (0.83–0.96)
INADN	50	75 ± 12 76 (66–80)	137 ± 23† 130 (120–150)	75 ± 12† 75 (70–80)	96 ± 14† 91 (87–102)	20 (40%)†	1 (2%)	0.87 ± 0.07† 0.88 (0.82–0.94)

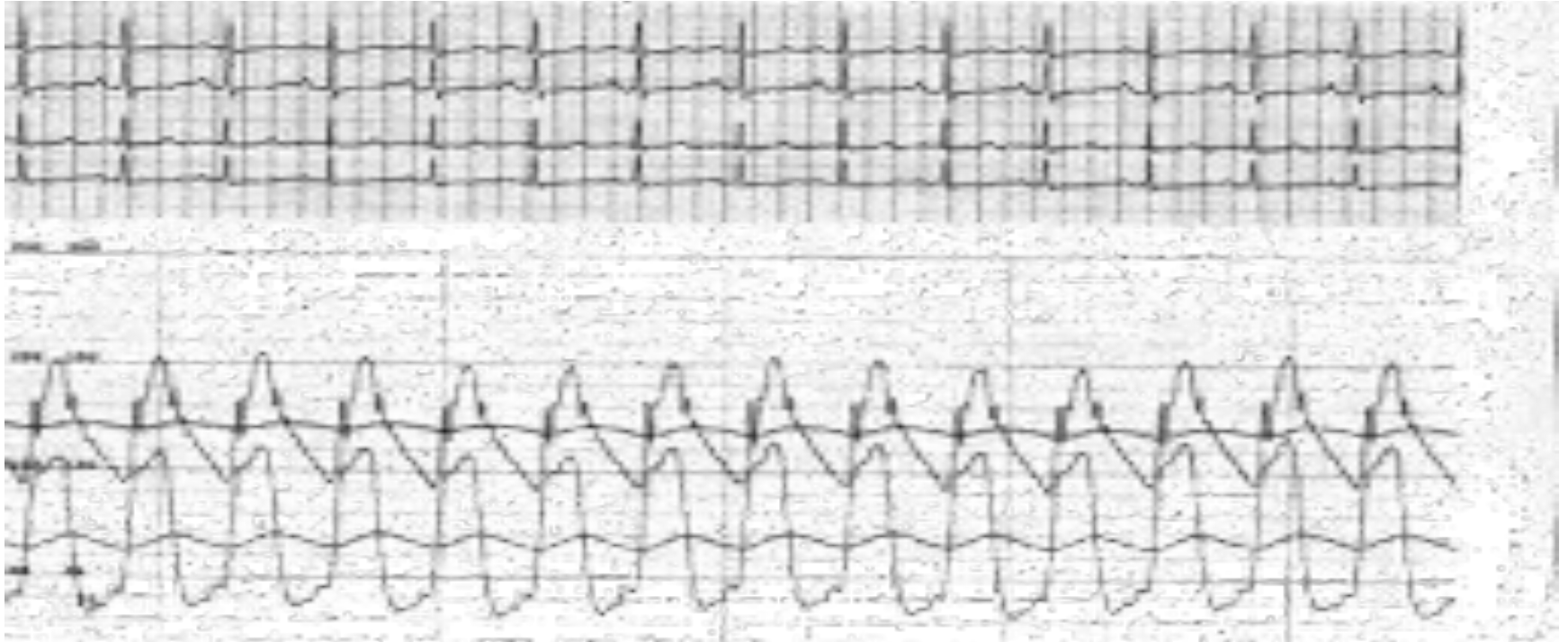




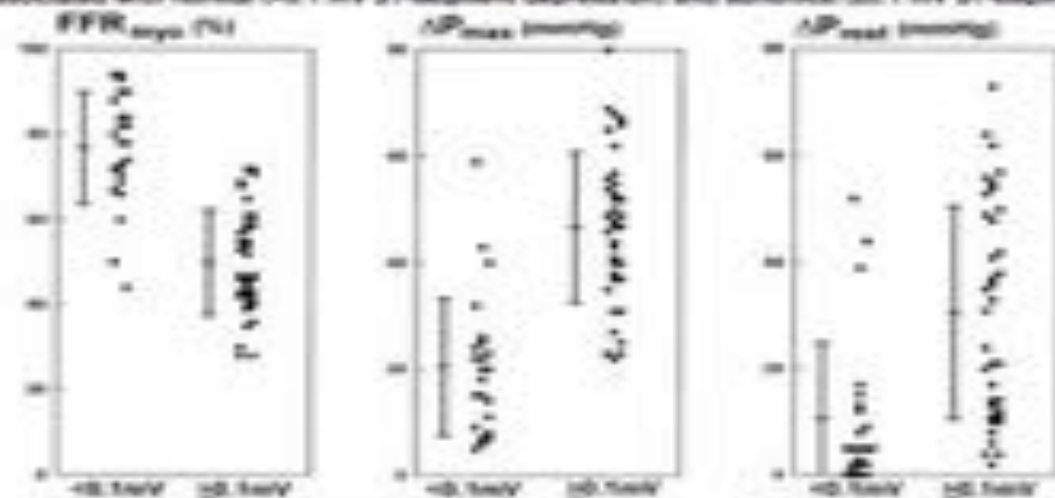
$$\text{FFR} = \text{Pm Co} / \text{Pm Ao}$$

EN HYPERHEMIE

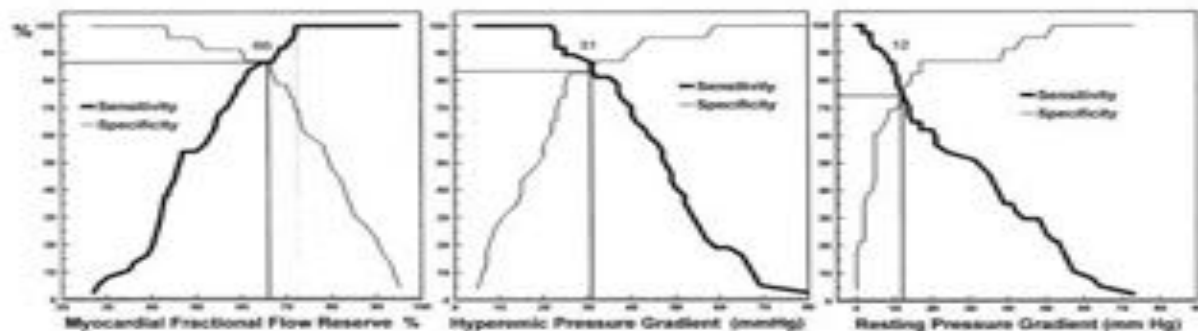
Gradient de Repos



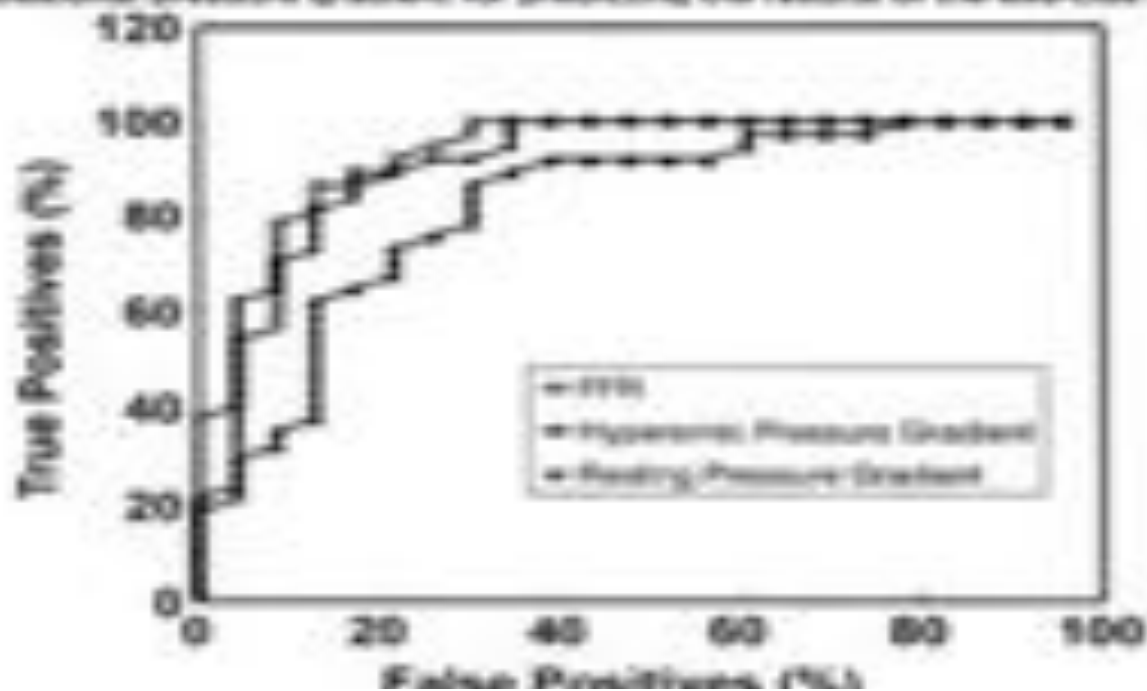
Scatterplots showing values of myocardial fractional flow reserve (FFR_{myo}), hyperemic transitional pressure gradient (ΔP_{max}), and resting transitional pressure gradient (ΔP_{rest}) associated with normal (<0.1 mV ST-segment depression) and abnormal (≥ 0.1 mV ST-segment

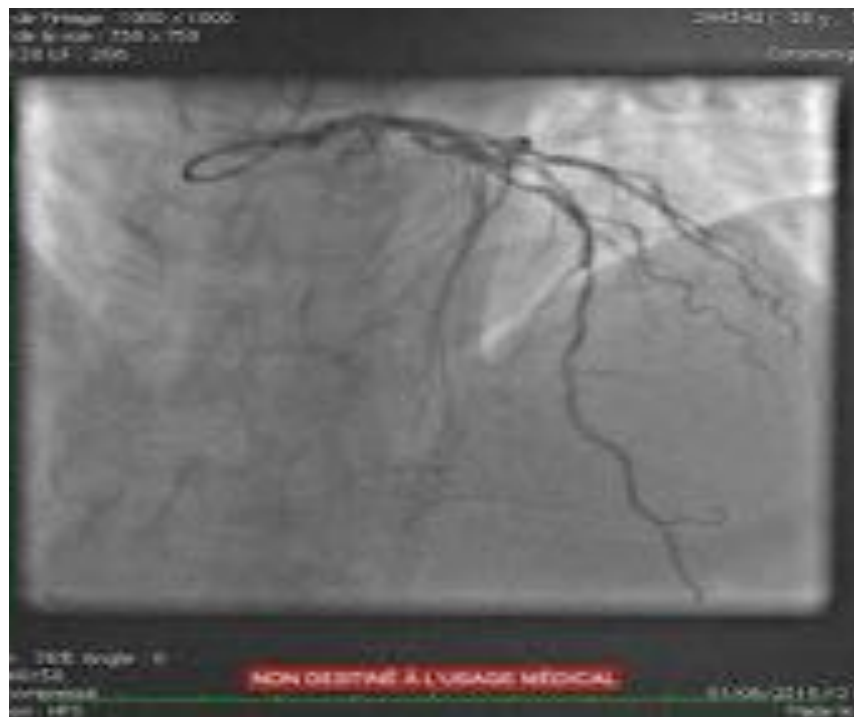


Source: Kastrup et al. Circulation. 1993;88:1000-1008

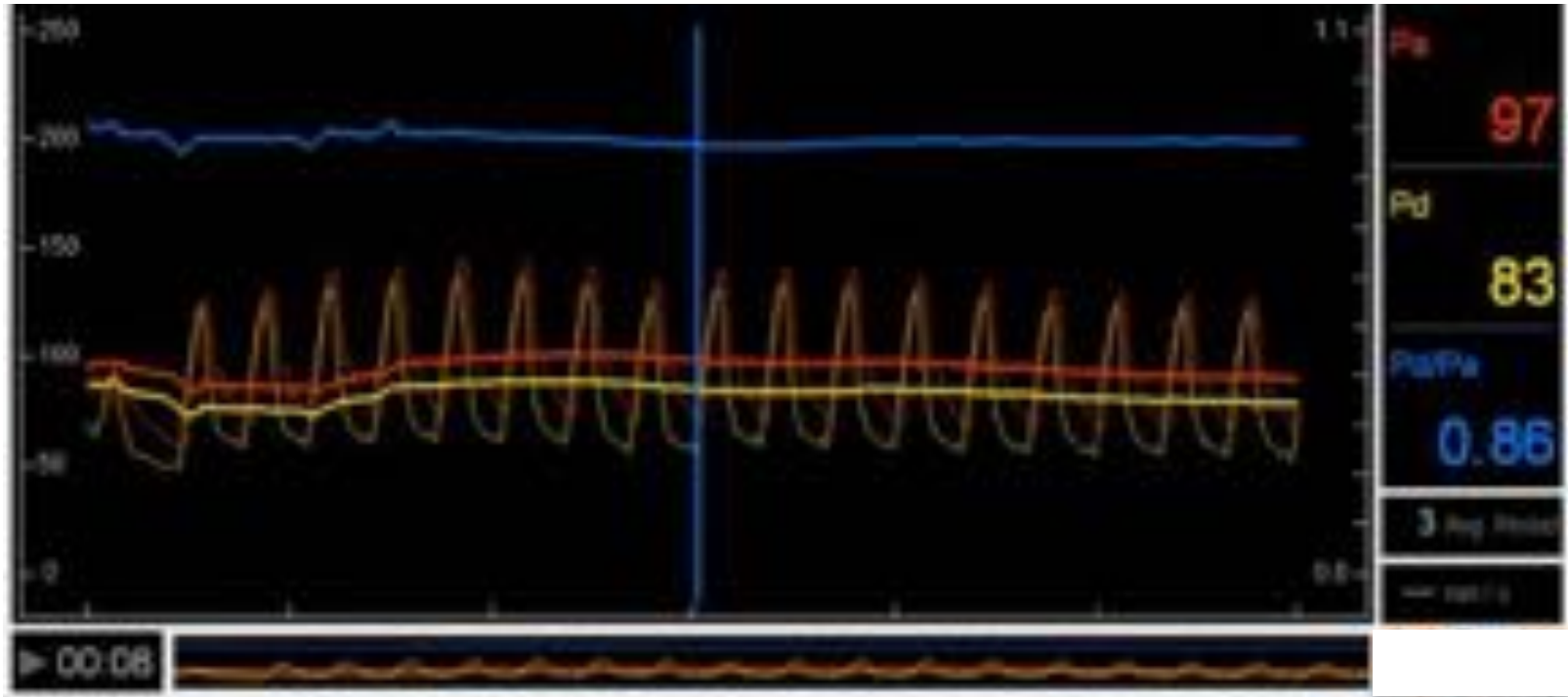


transcatheter pressure gradient for predicting the results of the exercise ECG.

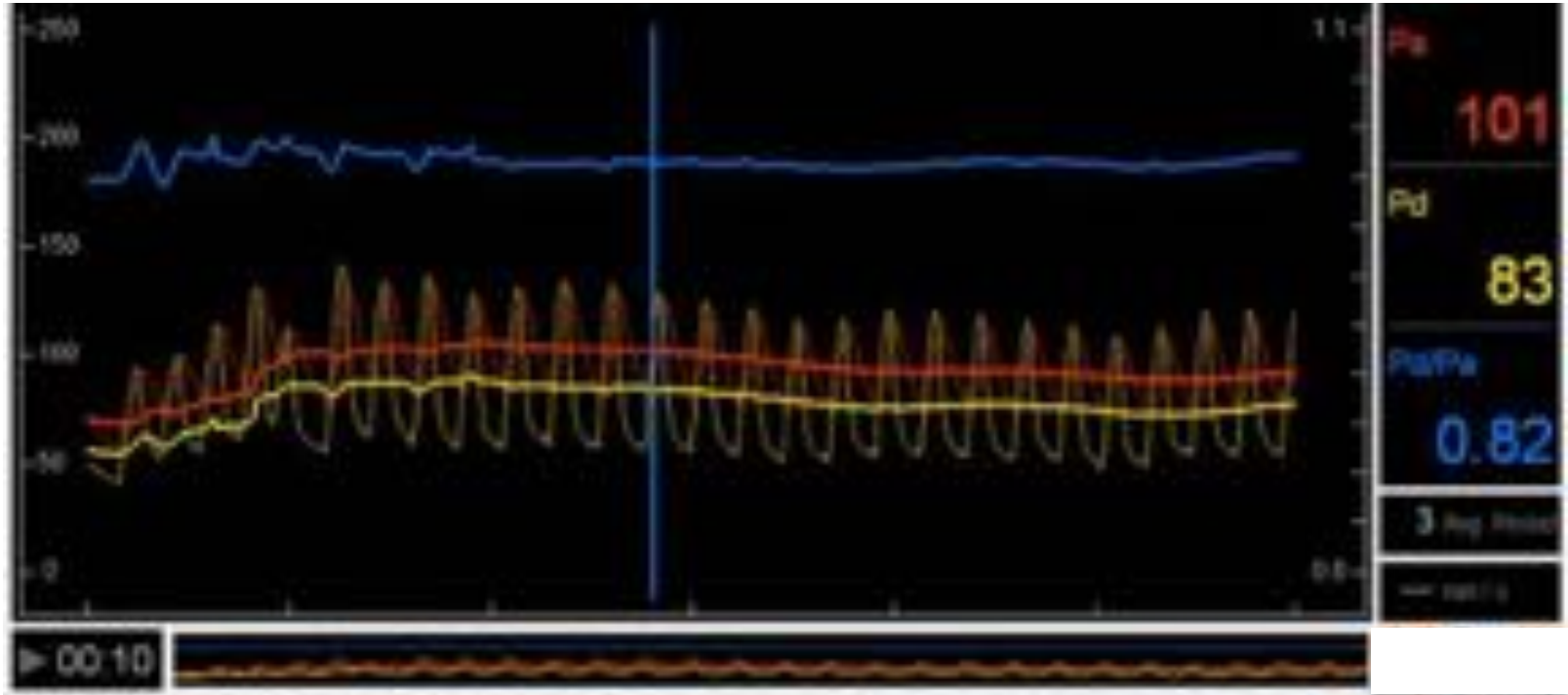




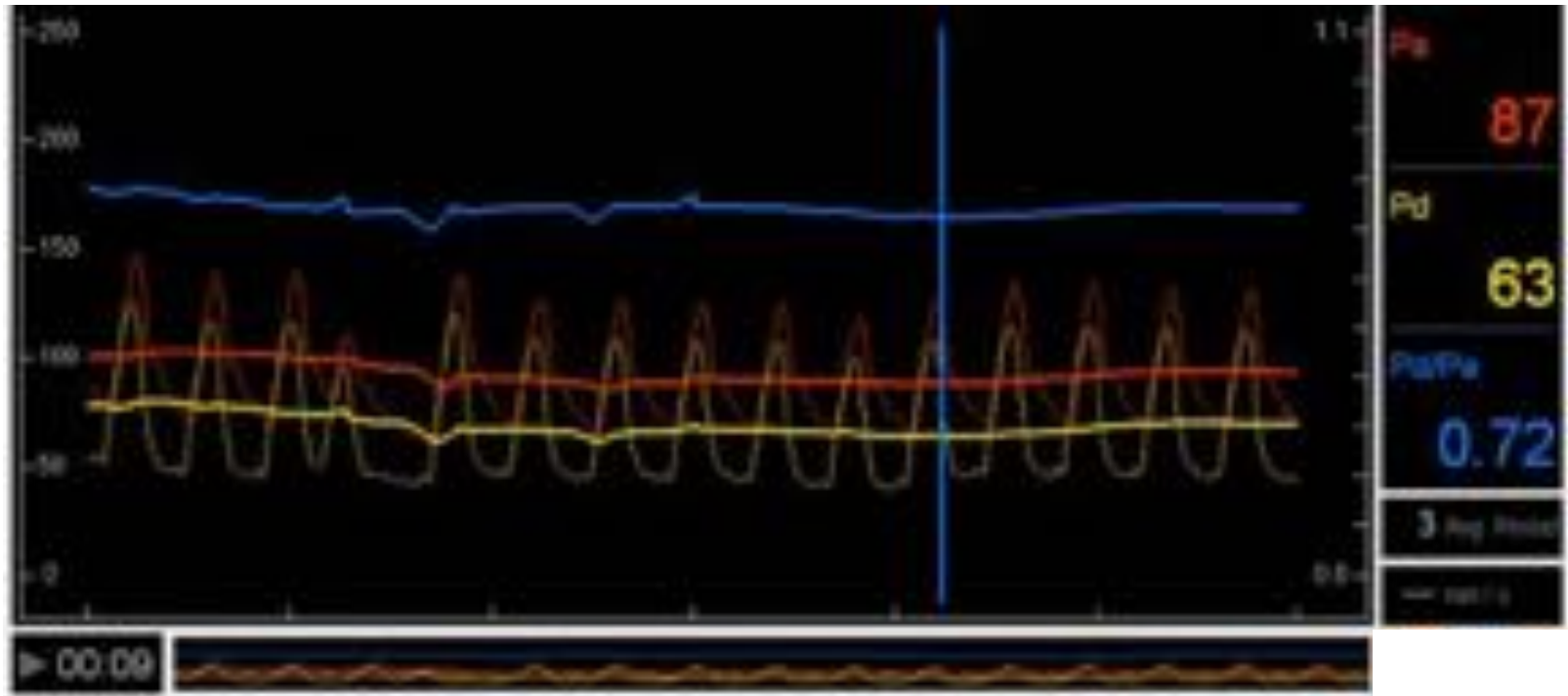
Gradient de base



Gradient contrast

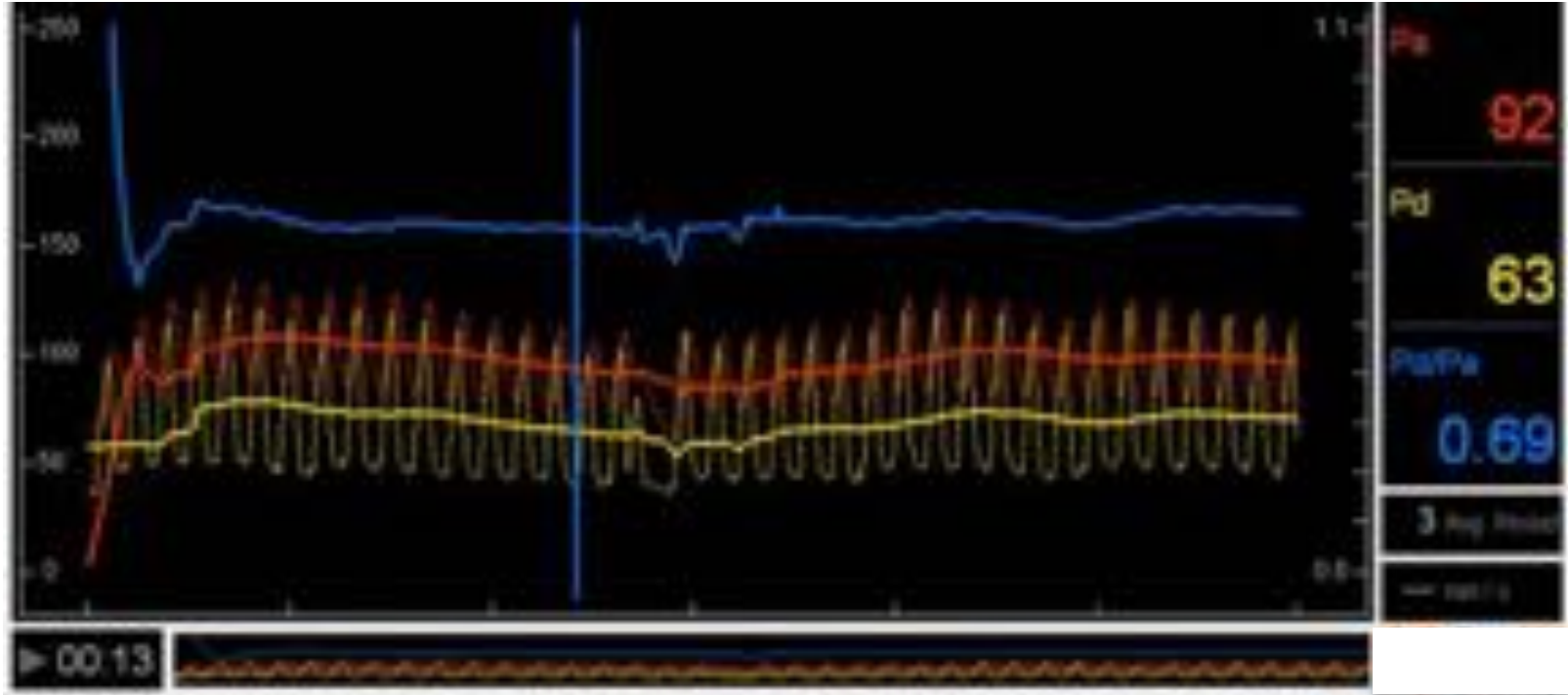


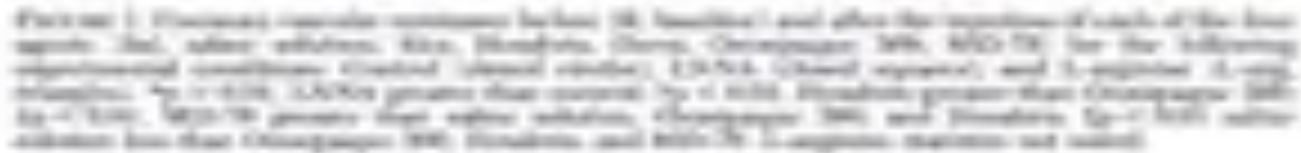
Gradient Adenosine 140ug bolus



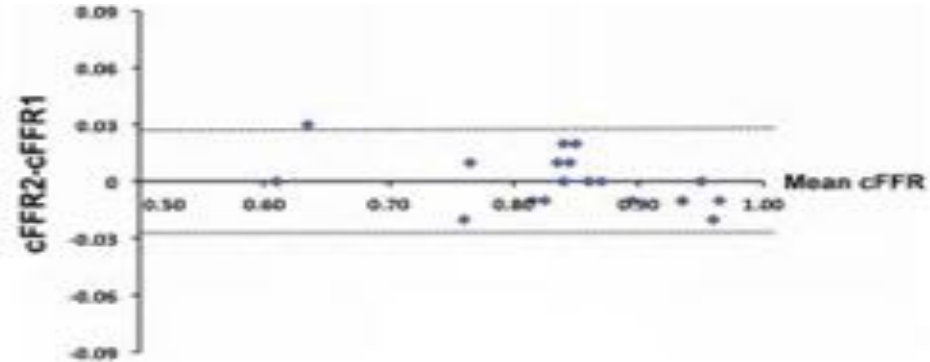
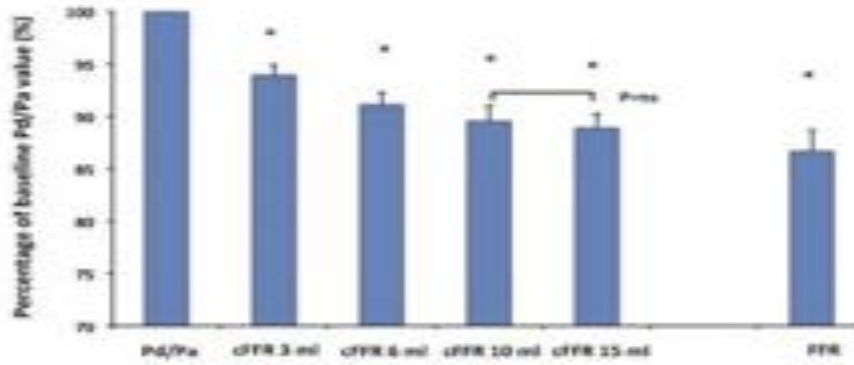


Adenosine + contraste





cFFR $n=138$



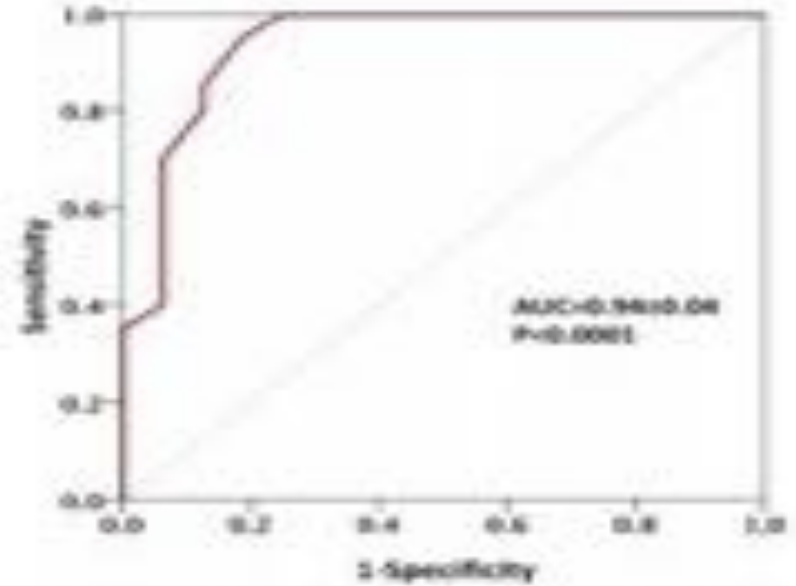
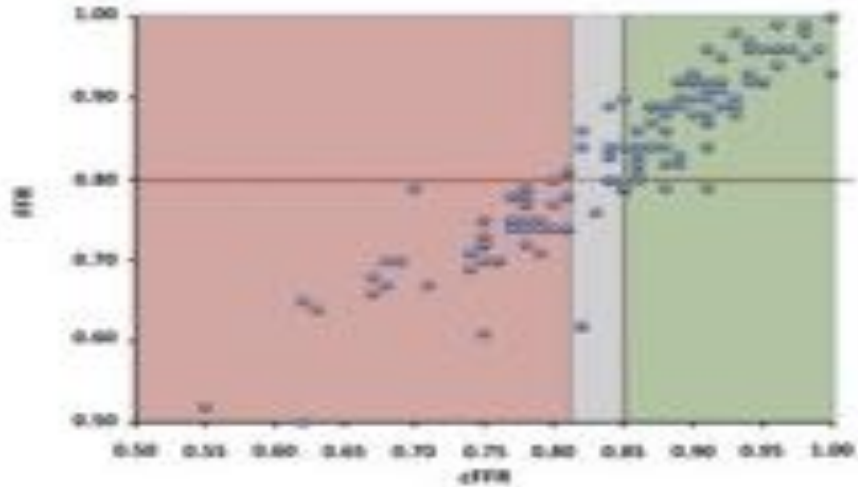


Figure 2. A receiver-operating characteristic curve was calculated using the threshold cutoff for contrast-enhanced fractional flow reserve (cFFR) of 0.83 in the exploration cohort. The receiver-operating characteristic was found to have an area under the curve (AUC) of 0.94 ± 0.04, which suggests the high accuracy rate of contrast-enhanced FFR in predicting FFR.

RINASCI Study $n = 104$

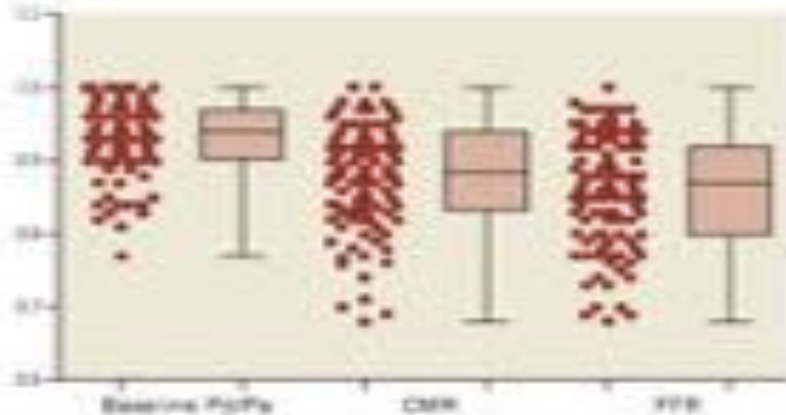


Figure 1. Values of resting Pd/Pa, contrast medium induced Pd/Pa ratio (CMR) and fractional flow reserve (FFR).

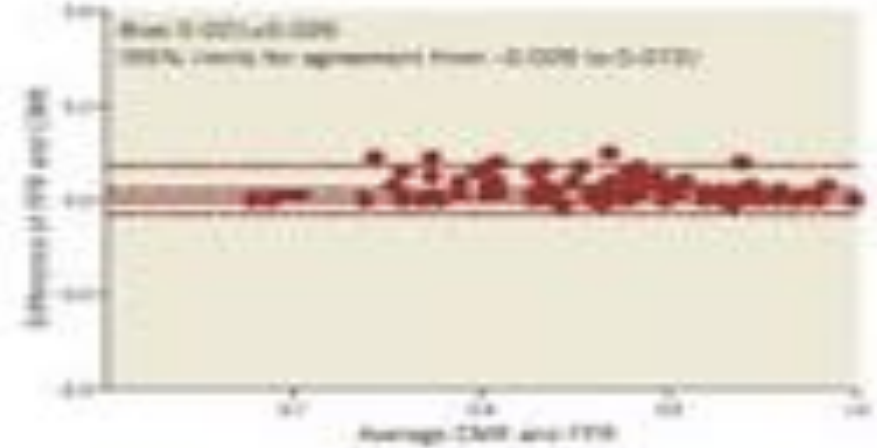


Figure 2. The Bland-Altman plot demonstrated a good agreement between contrast medium induced Pd/Pa ratio (CMR) and fractional flow reserve (FFR) across the entire range of coronary severity (0.52 to 0.92, 95% CI of disagreement: -0.03 to 0.07).

RINASCI Study $n = 104$

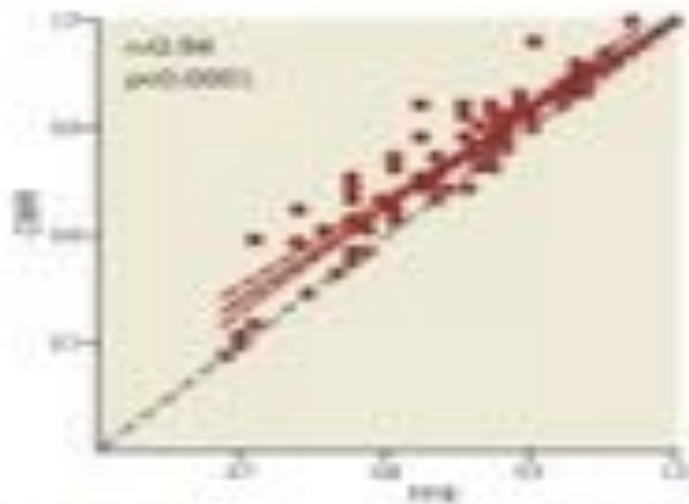
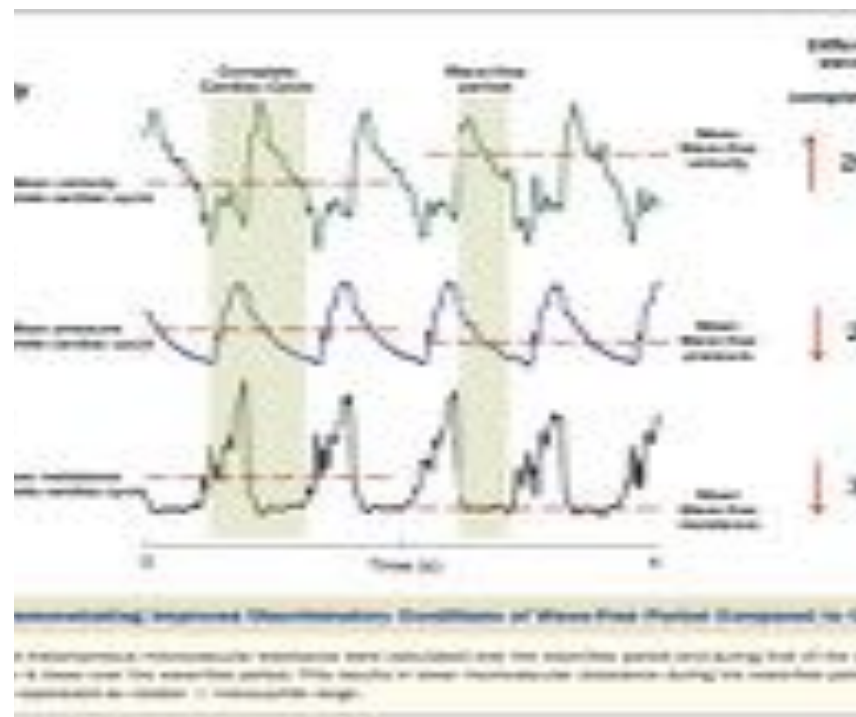
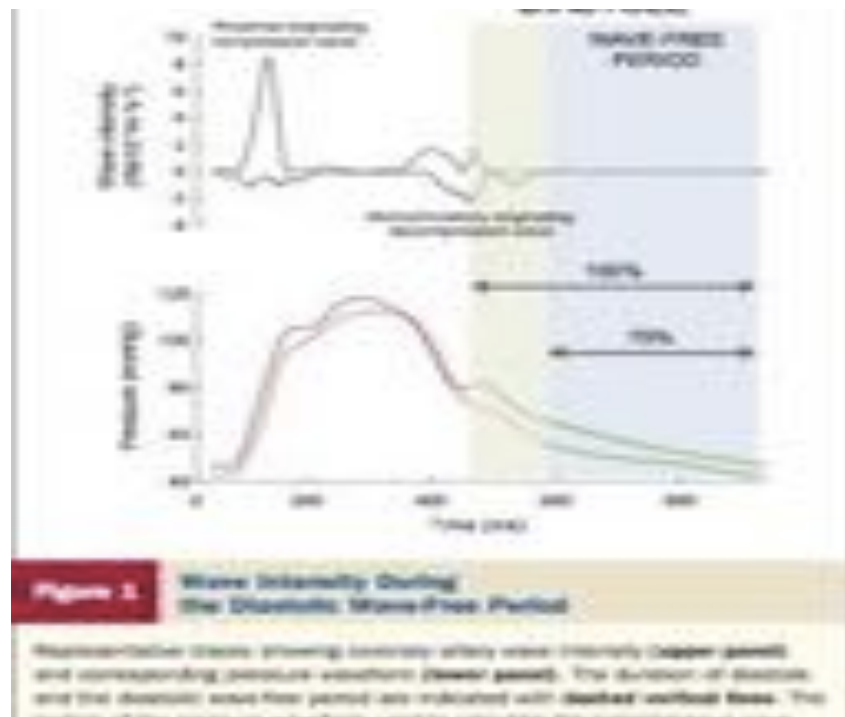


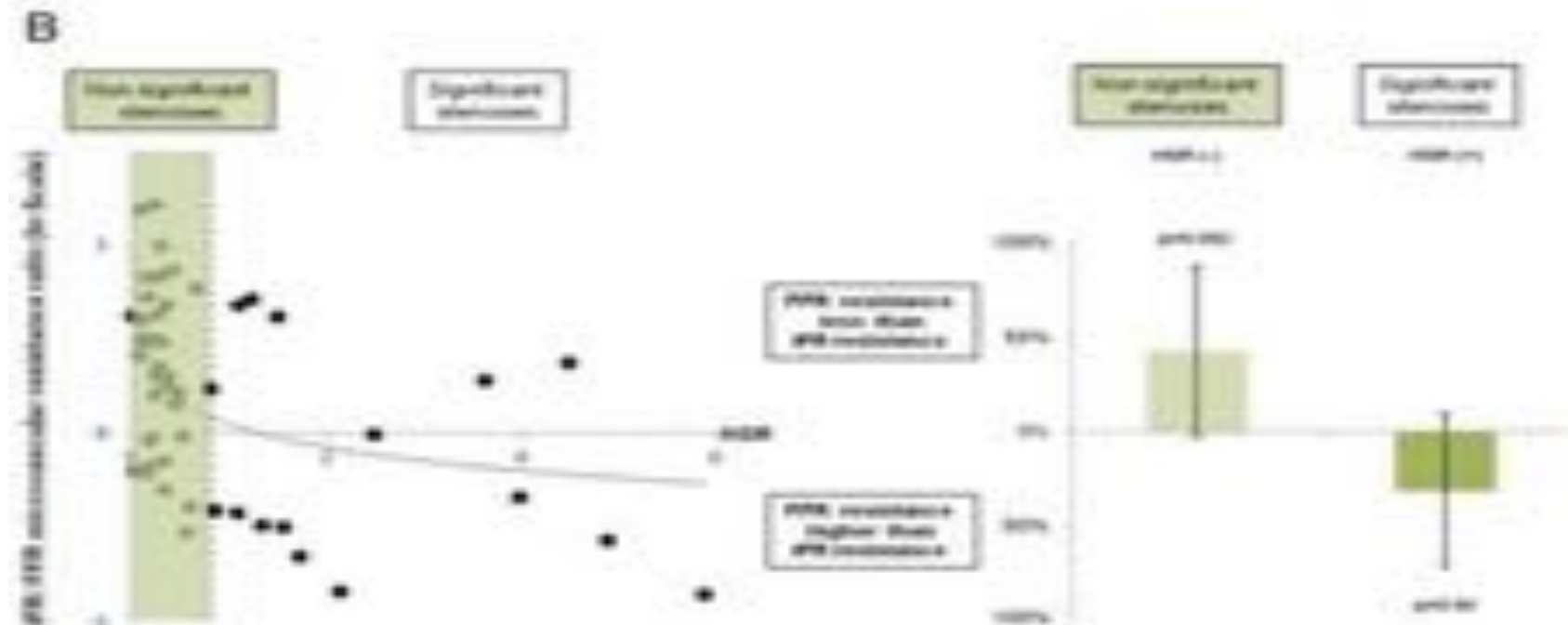
Fig 2. Correlation between contrast medium induced PDPs (mL) and fractional flow reserve (FFR) values.



Figure 3. We propose a hybrid CME/FFR approach it summarized in a simple algorithm that could allow limiting adenosine administration only in doubtful cases. We consider a CME value <0.82 to be significant and consequently we suggest performing PCI. A CME value >0.88 is not significant and consequently we suggest deferring PCI, and inducing maximal hyperemia using adenosine for FFR assessment when CME is between 0.84 and 0.87. As consequence, PCI would be performed when FFR is <0.80 and deferred when FFR is >0.88 .



iFR

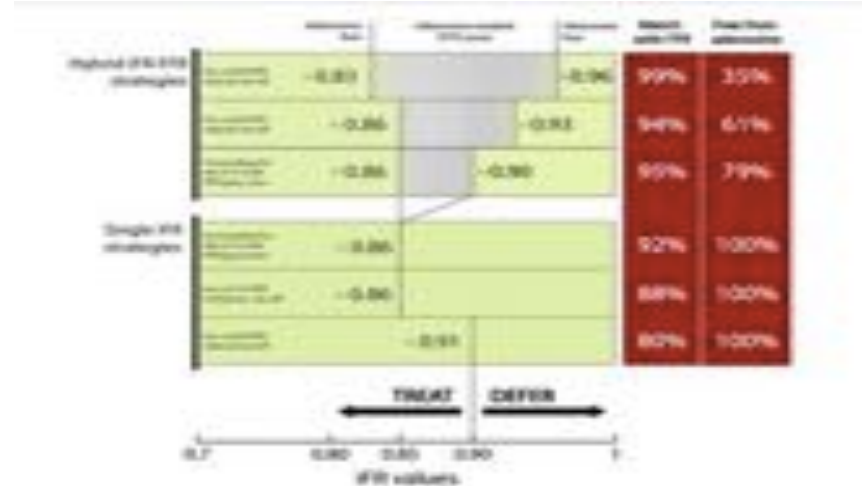


Advise Study

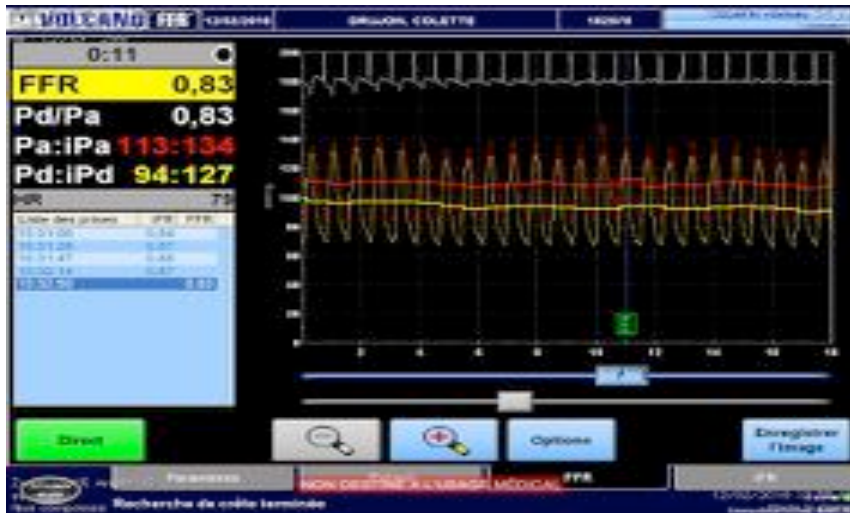
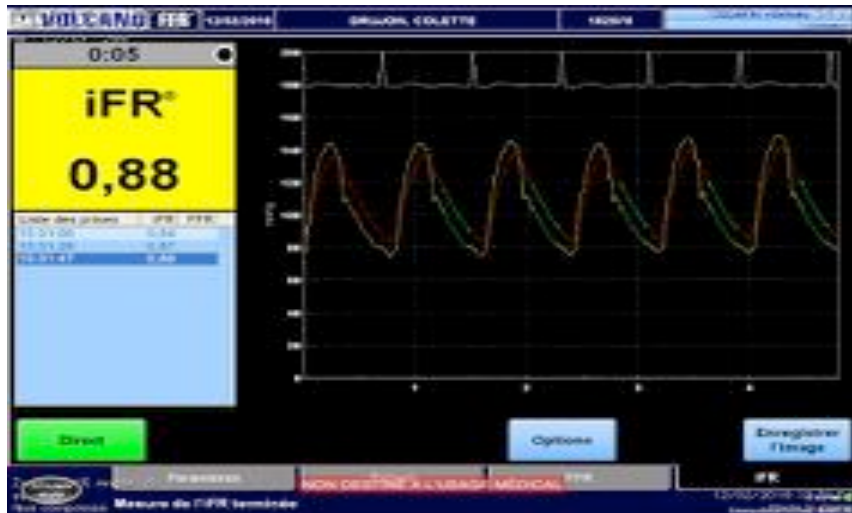
Classification agreement between iFR and FFR

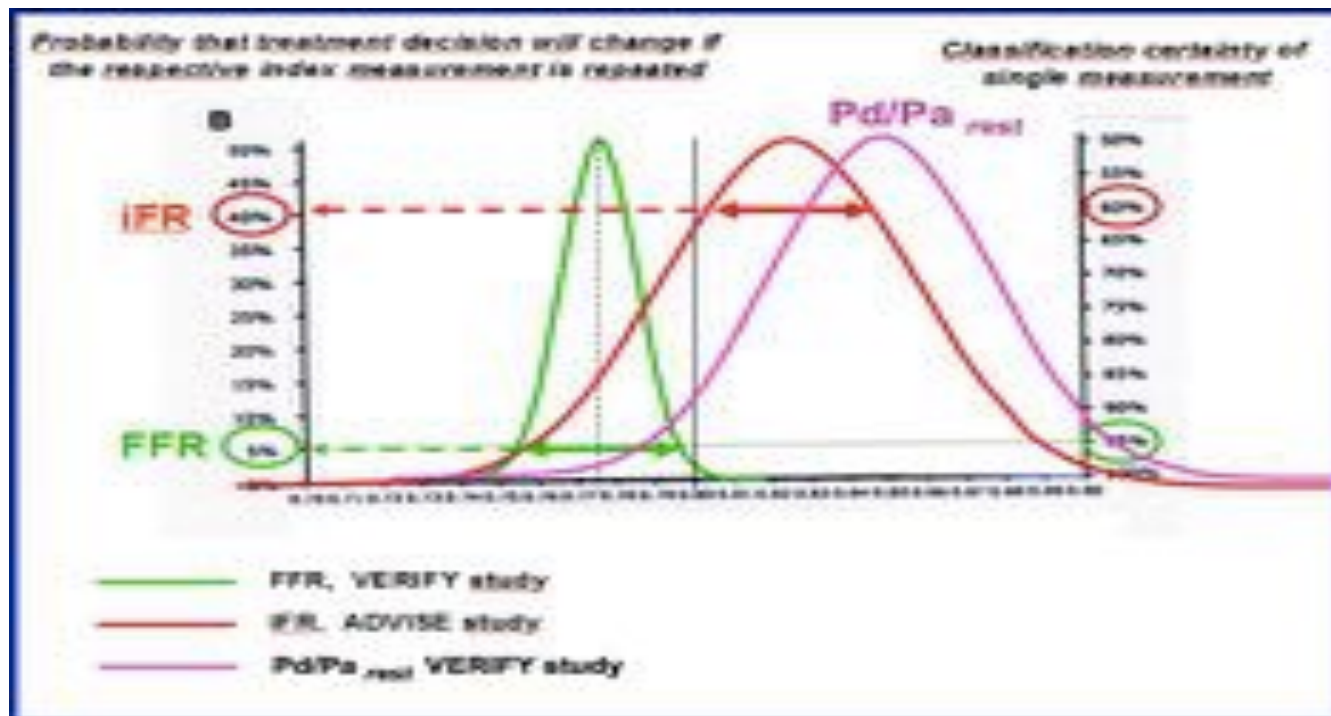
	FFR 0.8	FFR 0.75	FFR grey zone [†]
iFR cut-off	0.9	0.85	0.85
ROC AUC	0.87	0.90	0.93
Classification match	80%	88%	92%
Sensitivity	81%	75%	82%
Specificity	79%	92%	94%
PPV	71%	72%	88%
NPV	87%	92%	95%

[†] Accounting for the 0.75 - 0.8 FFR grey zone

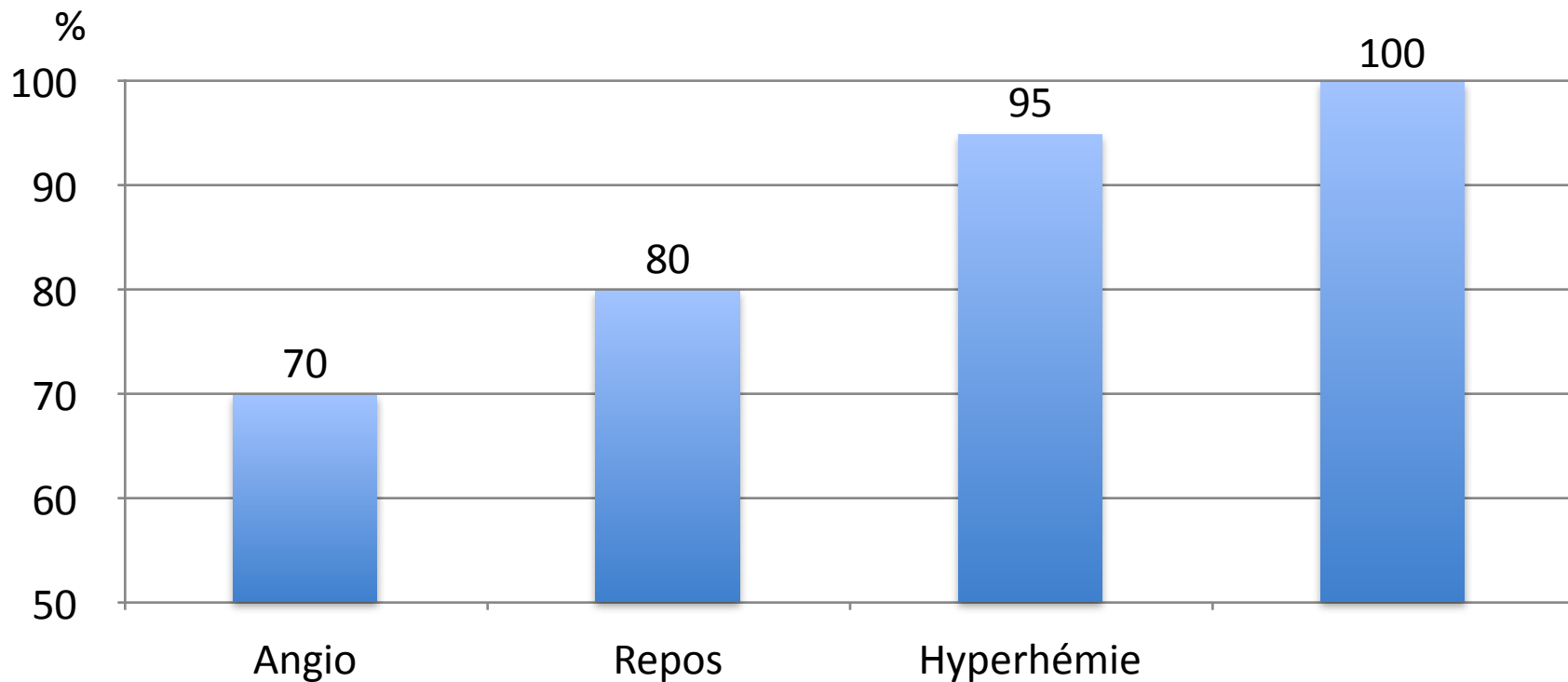








Précision diagnostique lésion ischémique

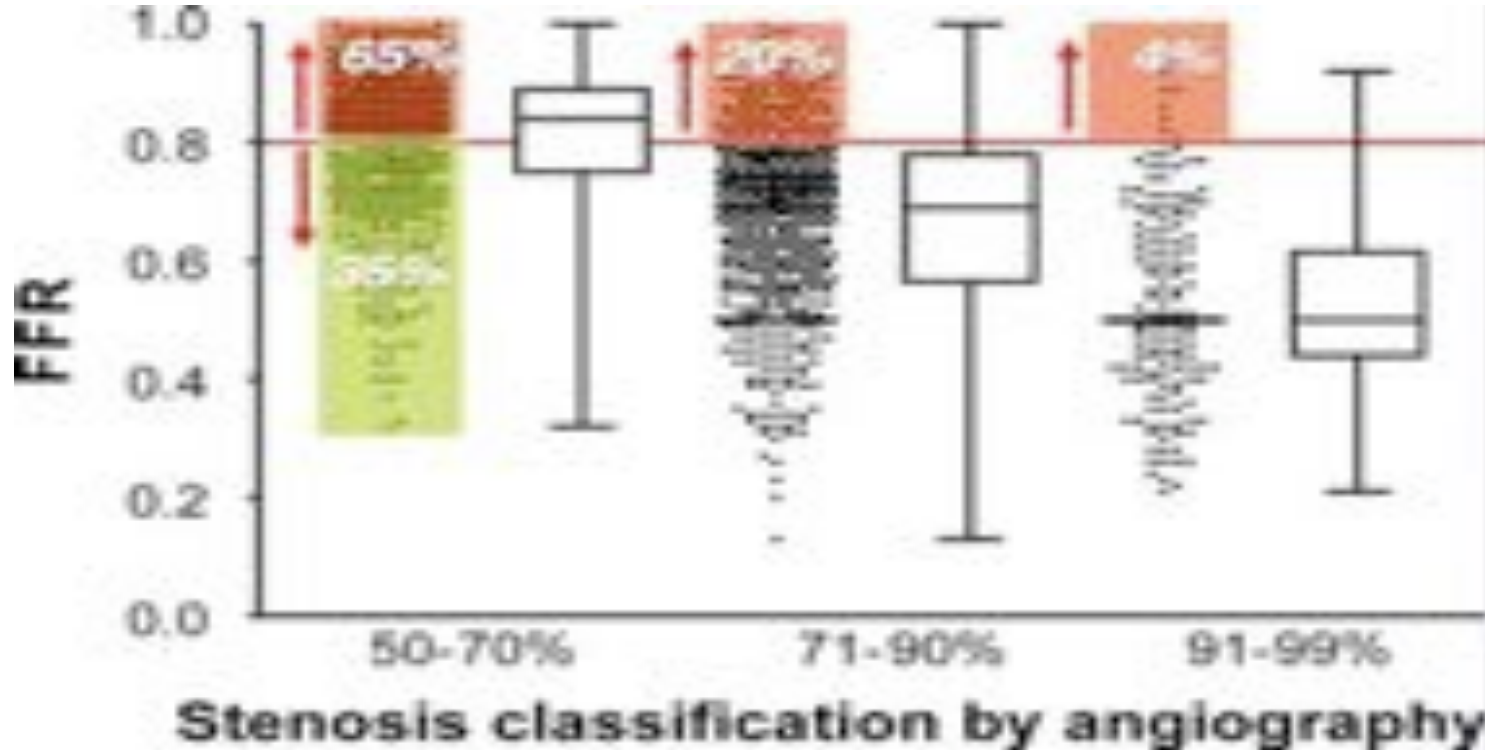


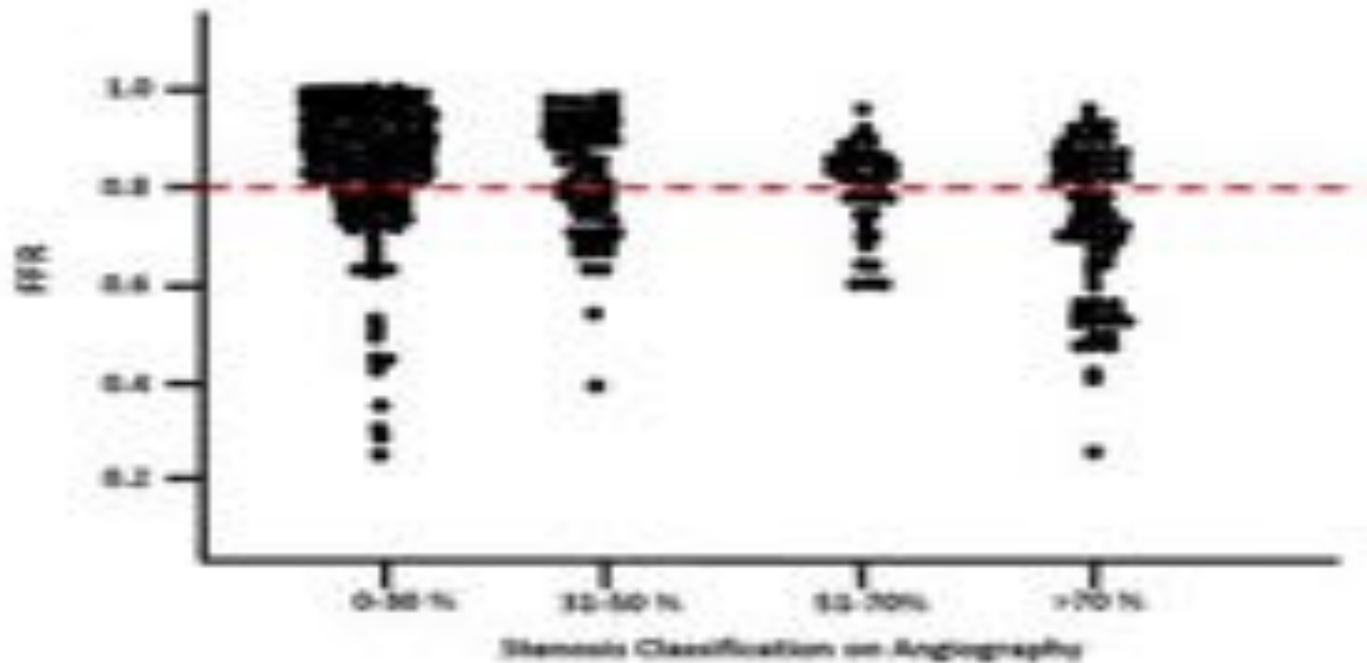
Conclusions

- L'utilisation du guide pression sans stimulus adjuvant est possible
- Quelque soit la technique utilisée, un valeur $< 0,80$ signe la fonctionnalité de la sténose
- Quelle valeur seuil retenir pour
 - Rapport Pd/Pa de repos – aucune étude
 - Contraste 0,83 zone grise : 0,84-0,88 ?
 - iFr 0,9 zone grise : 0,86-0,93
- Manque études de validation
- Define-Flair IFR SwedHeart Studies

INFLUENCE DE LA FFR SUR L'ACTIVITE ANGIOPLASTIE

FFR vs Angiographie





R3F registry

The FFR real life in France

- French prospective and multicentric Registry
- Inclusions 2008-2010
- **1101 Patients**
- Hop, 6 month and 1 year follow-up
- Electronic CRF

Circulation



Outcome Impact of Coronary Revascularization Strategy Revascularization With Fractional Flow Reserve at Time of Diagnostic Angiography: Insights From a Large French Multicenter Fractional Flow Reserve Registry

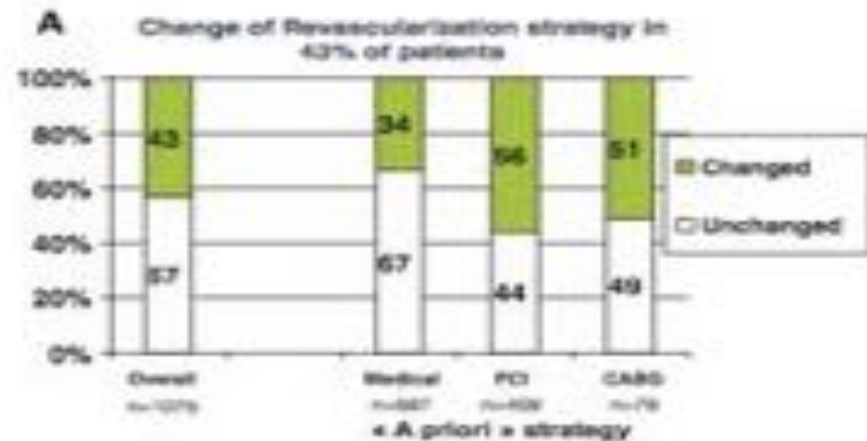
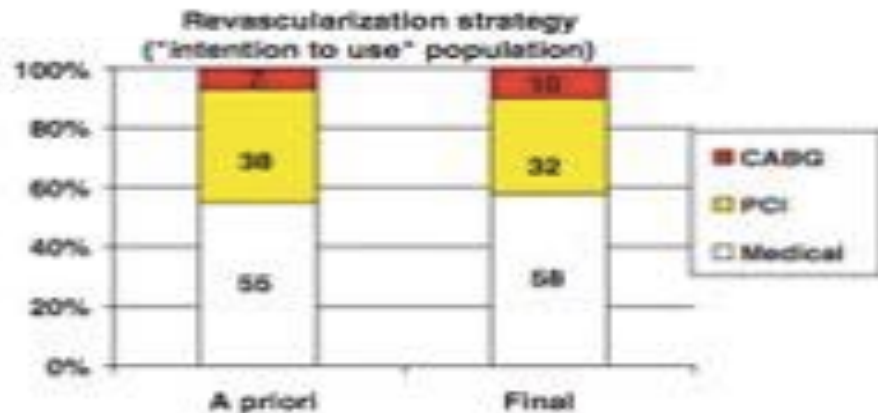
Eric Van Belle, Gilbert Rioufol, Christophe Pouillon, Thomas Cosset, Karim Bougrini, Emmanuel Teiger, Stéphane Champagne, Louis Belle, Didier Barreau, Michel Hansson, Cyril Renaud, Raphael Dauphin, Jean Dillenger, Youssef El Hachimi, Georgios Sidiropoulos, Christophe Berteille, Nicolas Lhoest, Pierre Barrey, Laurent Laborgne and Patrick Dupouy

Circulation, published online November 19, 2013;

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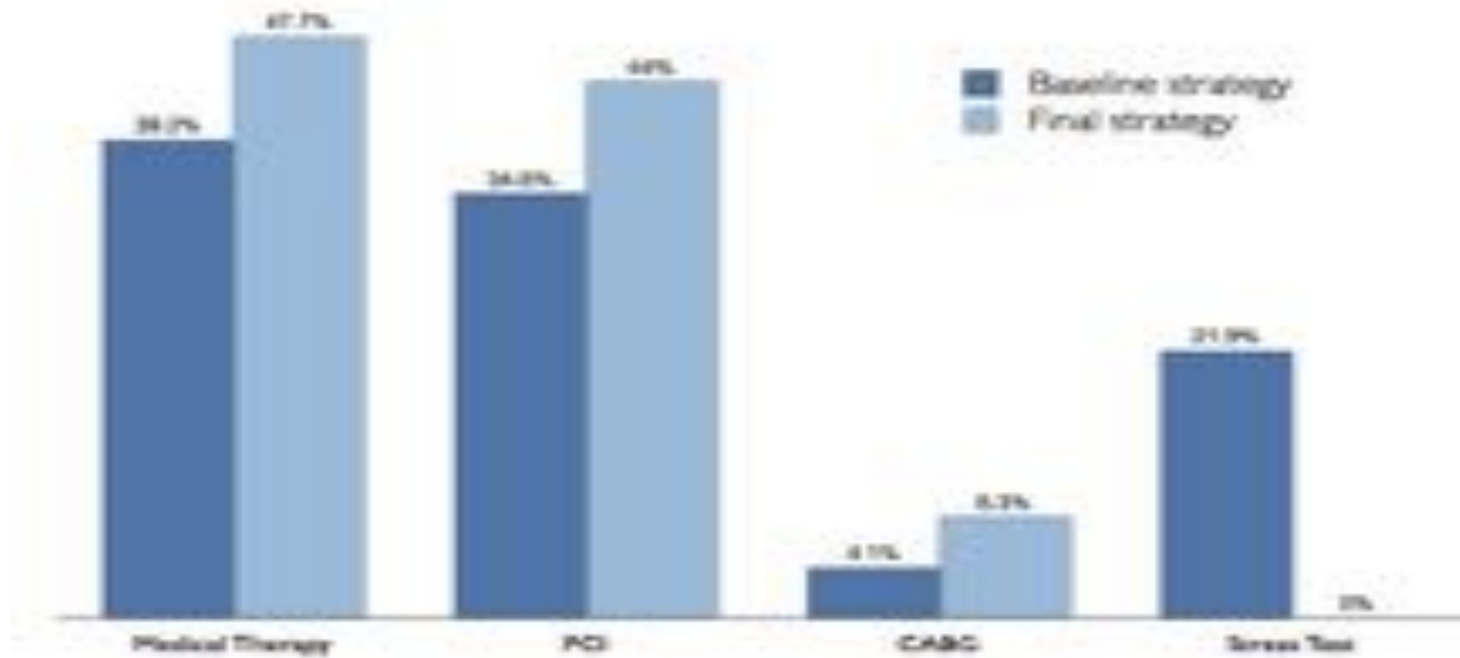
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-6% angioplastie

Post-It study



Routine FFR

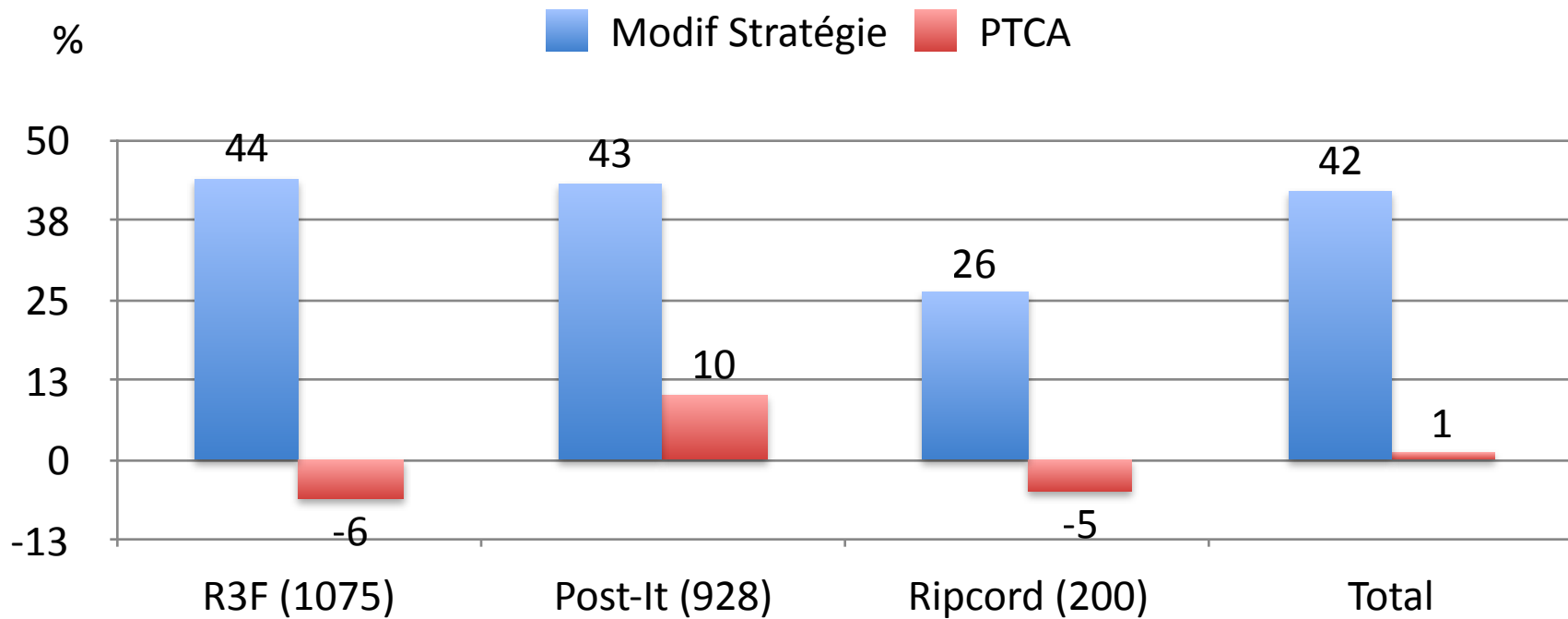
		FFR				Total
		Medical	PCI	CABG	Further info	
PLAN 1 ANGIO	Medical	63	6	3	0	72
	PCI	24	64	2	0	90
	CABG	1	3	19	0	23
	Further info	1	7	6	1	15
	Total	89	80	30	1	200

Fishers exact test $p < 0.0001$

Summary

- Agreement about category of management in 147 out of 200 (74%)
 - ie after FFR management change in 26% of cases

-10% ANGIOPLASTIES

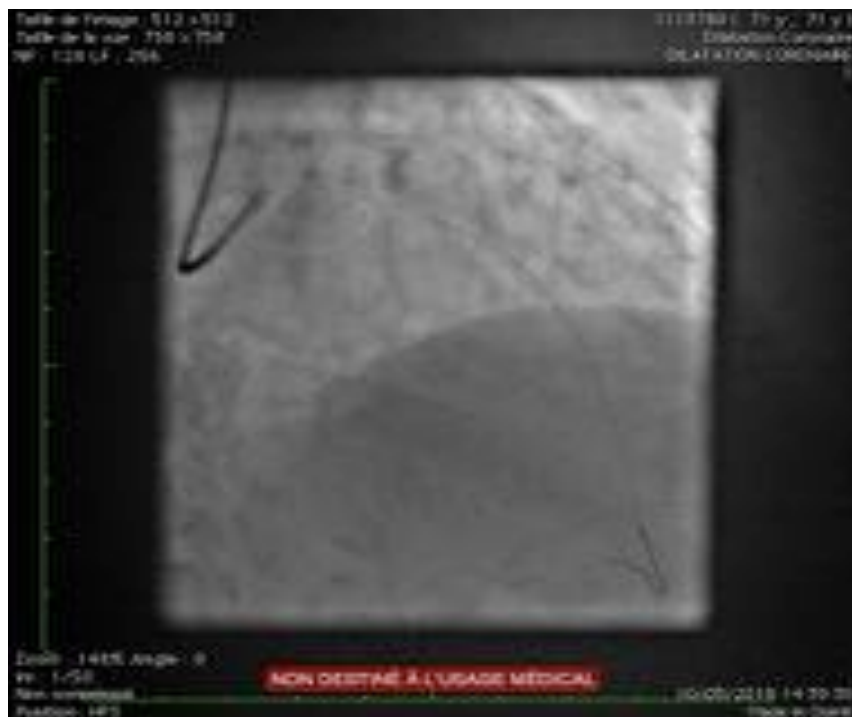


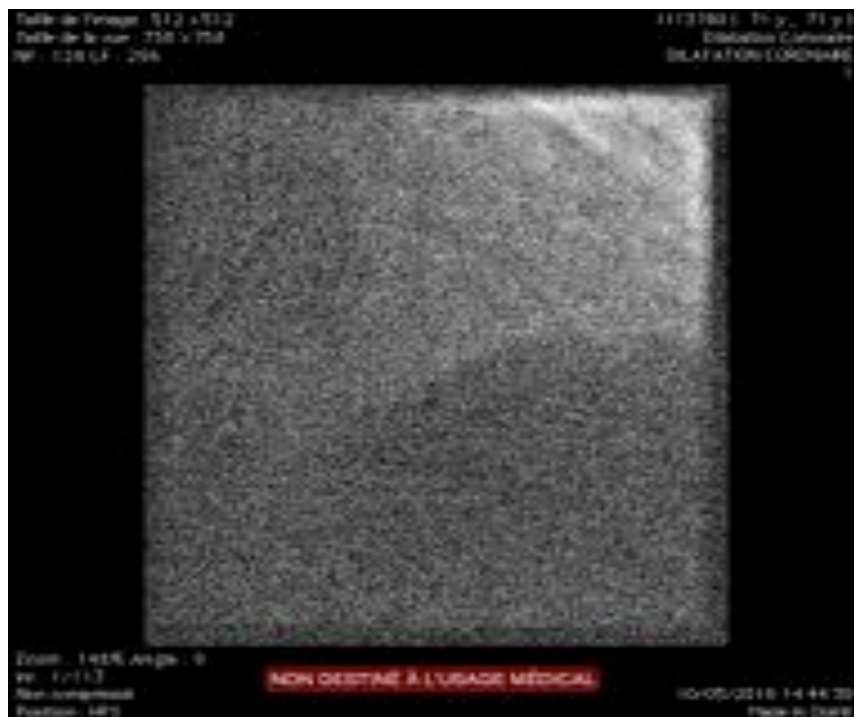
Conclusions

- L'utilisation de la FFR a peu d'impact sur le taux de revascularisation par angioplastie

La FFR dans les lésions de bifurcation

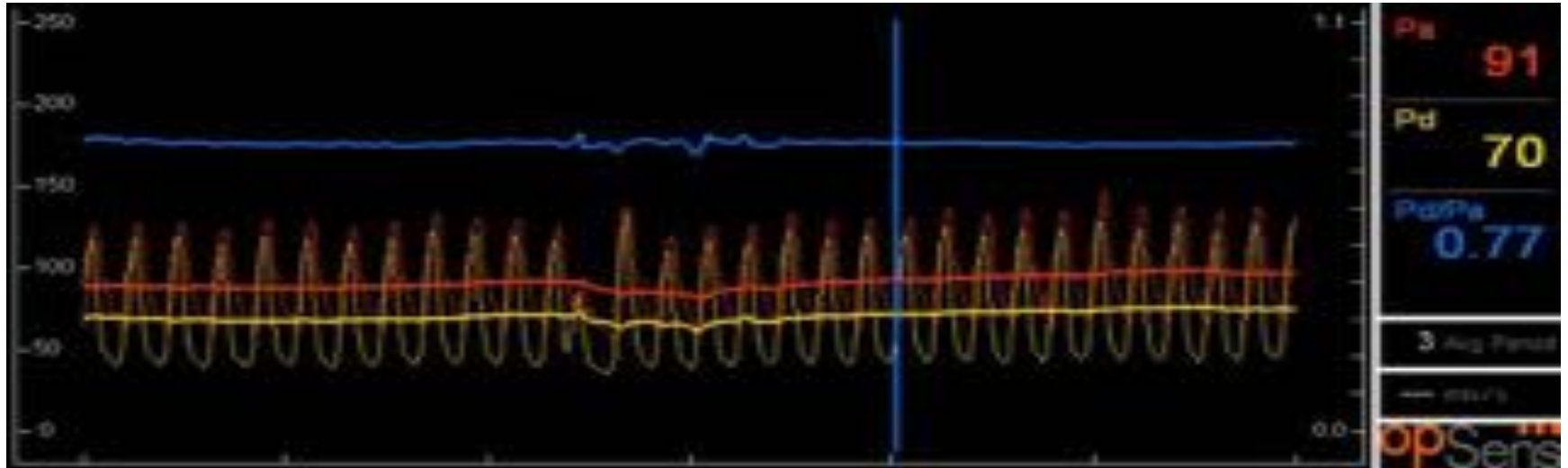






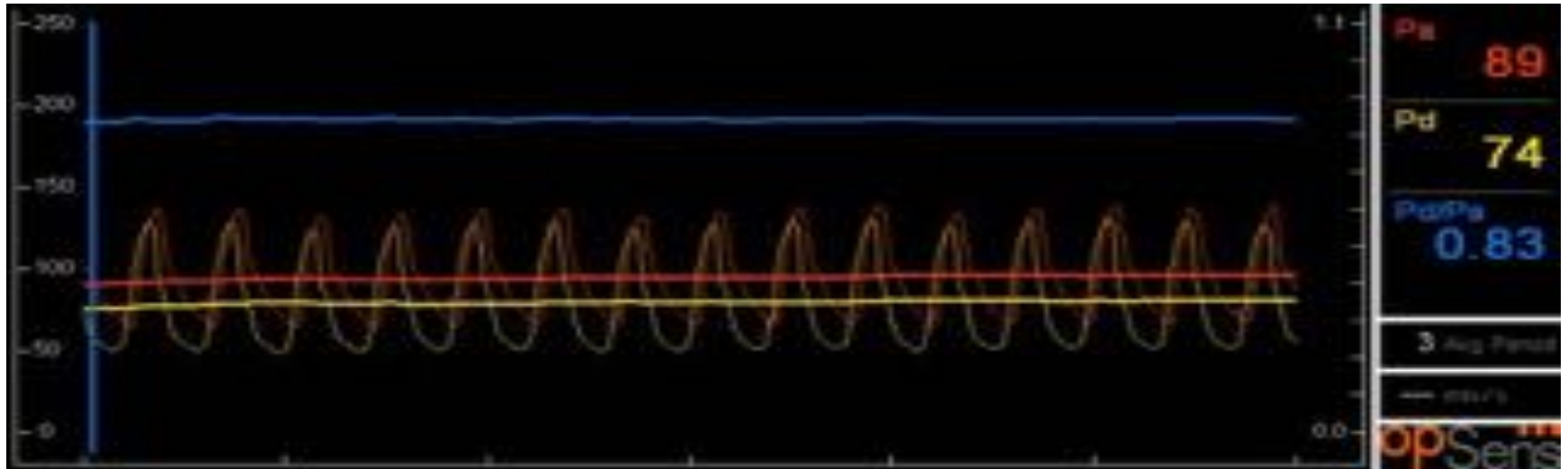


FFR Diag après stent IVA





FFR après KB Diag



FFR dans les bifurcations

Table 1. FFR during bifurcation intervention.

	FFR is useful	FFR is generally not recommended
Pre-intervention	To assess the functional significance of MB To assess the functional significance of pure SB stenosis	Small SB To determine functional significance of SB when there is a significant MB stenosis SB FFR to predict the functional significance of jailed SB
Post MB stenting	To assess the functional significance of jailed SB and to predict the outcomes	Small SB Long diffuse, highly angulated or calcified SB SB slow flow
Post SB angioplasty	To assess SB procedural success and to predict the outcomes after KBI	SB slow flow SB severe dissection
Post SB stenting	To evaluate residual ischaemia	To predict procedural outcomes of complex two stenting
FFR: fractional flow reserve; KBI: kissing balloon inflation; MB: main branch; SB: side branch		

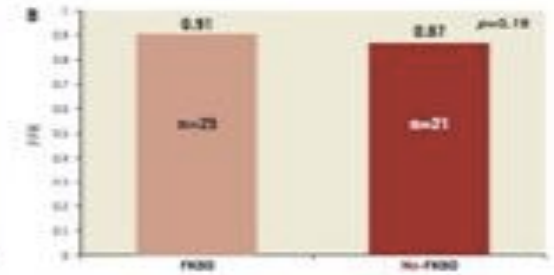
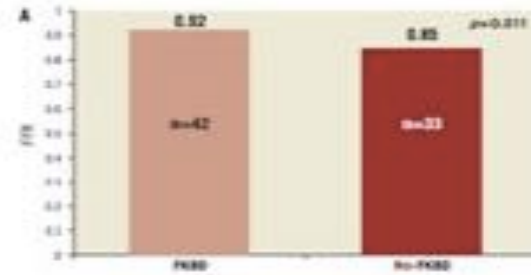
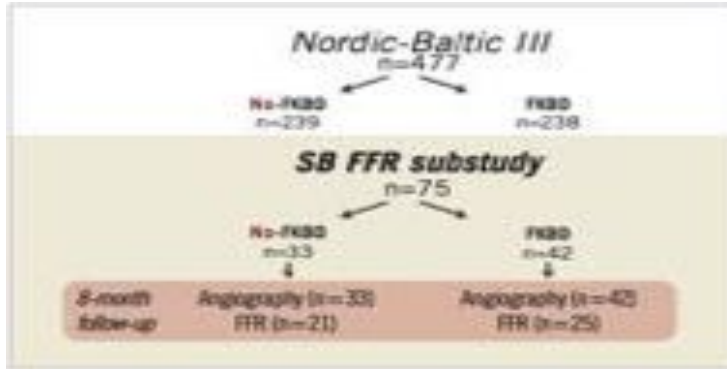
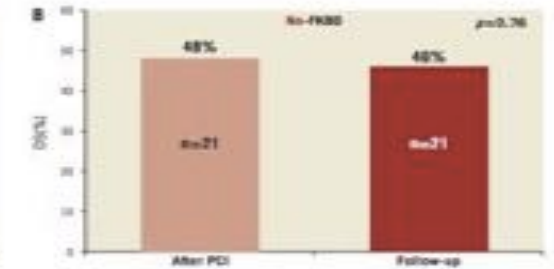
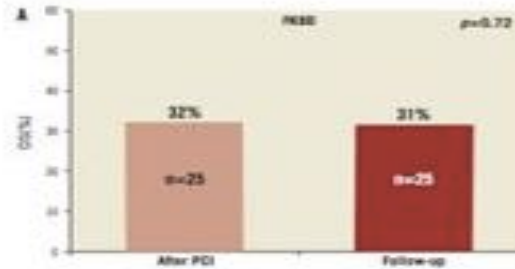
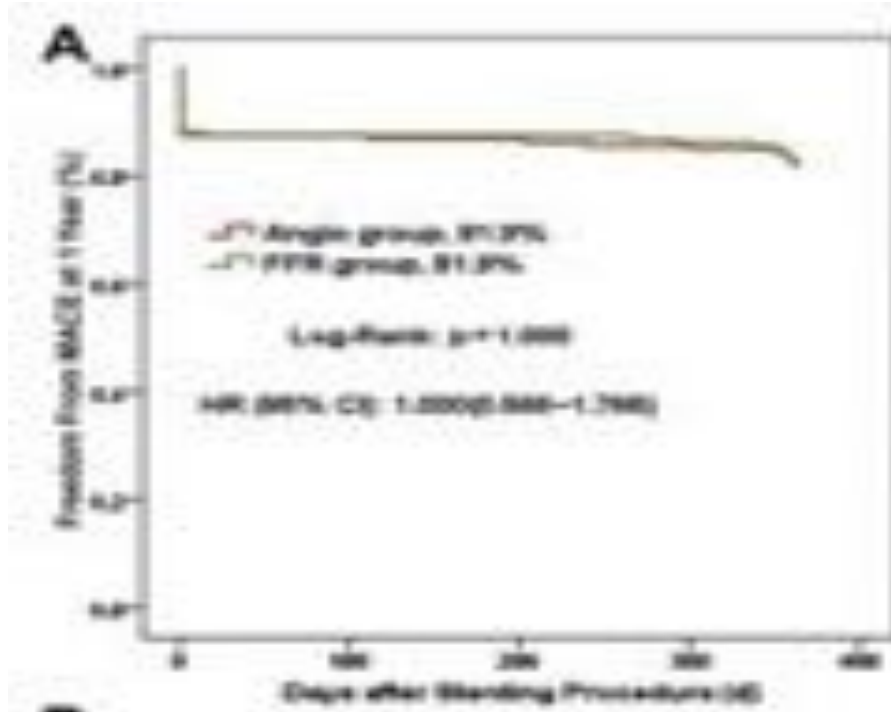
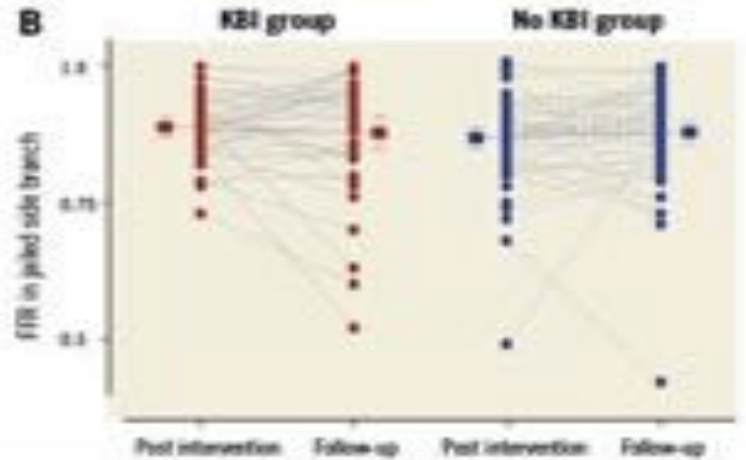
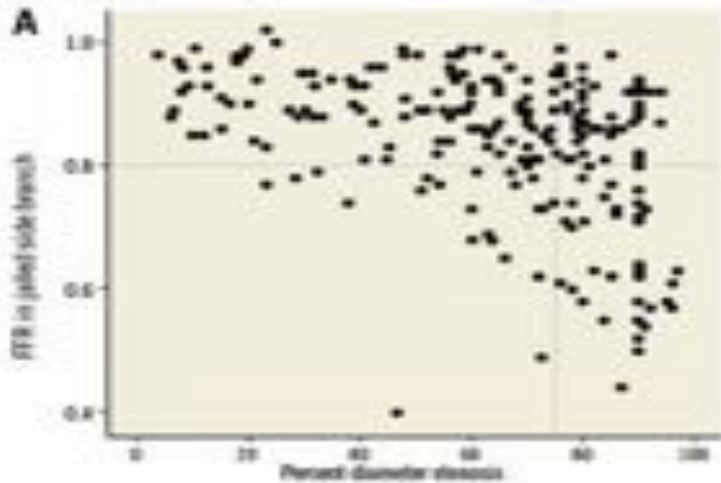


Figure 4. A) Mean FFR in SB after PCI. B) Mean FFR in SB at 8-month follow-up.



DKCRUSCH-IV Study





Serial FFR measurement in jailed side branch

Roc BK et al 2008	100 patients with provisional strategy Repeated SB FFR at 6-month follow-up (n=65).	- At 6-month follow-up, there were no changes in FFR in lesions with $(0.85 \pm 0.06$ to 0.84 ± 0.01, $p=0.4$) and without SB before angioplasty $(0.87 \pm 0.06$ to 0.89 ± 0.07, $p=0.12$). - Binary restenosis rate was 48%; however, functional restenosis (FFR <0.75) rate was 8% (5/65). There were no changes in SB FFR during the 6-month follow-up period $(0.92$ to 0.91, $p=0.80$ in KBI group and 0.87 to 0.87, $p=0.90$ in no KBI group).
Nordic-Baltic Bifurcation II 2012	75 patients with provisional strategy Repeated SB FFR at 6-month follow-up (n=40)	There were no changes in SB FFR during the 6-month follow-up period $(0.92$ to 0.91 , $p=0.80$ in final KBI group and 0.87 to 0.87 , $p=0.90$ in no final KBI group).

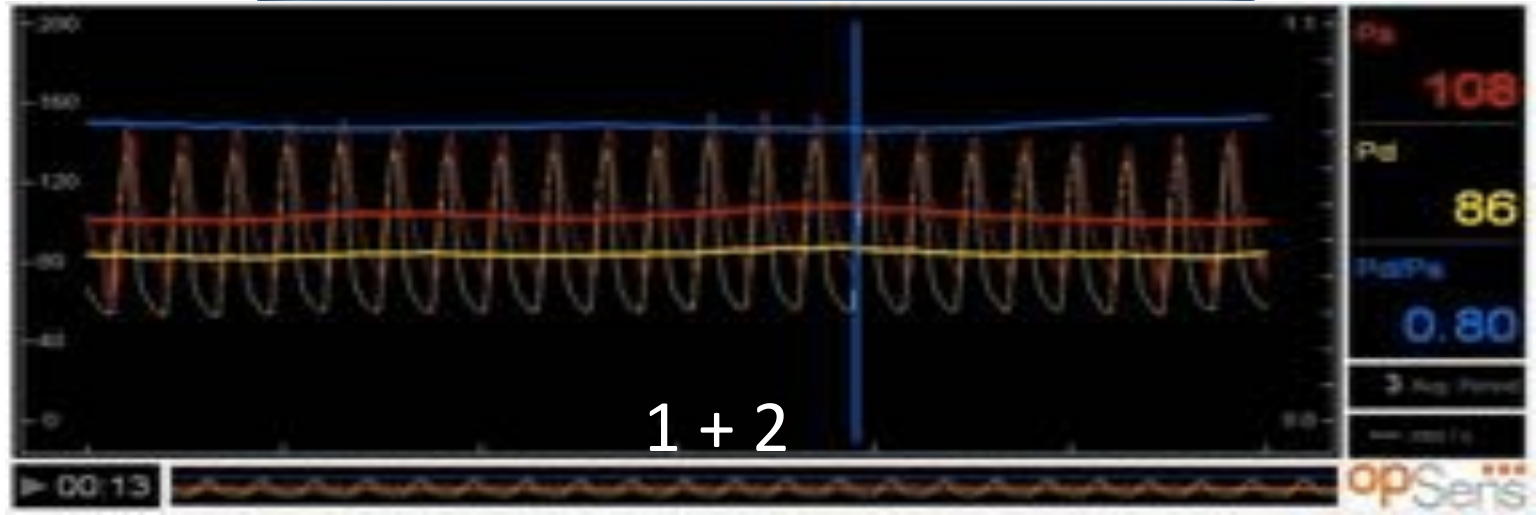
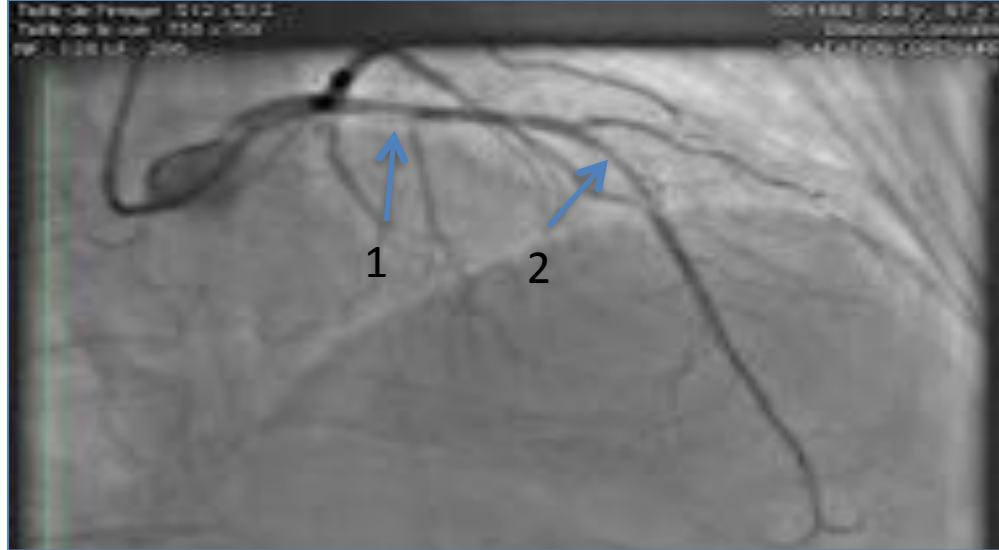
FFR-guided PCI vs. Angio-guided PCI for jailed side branch

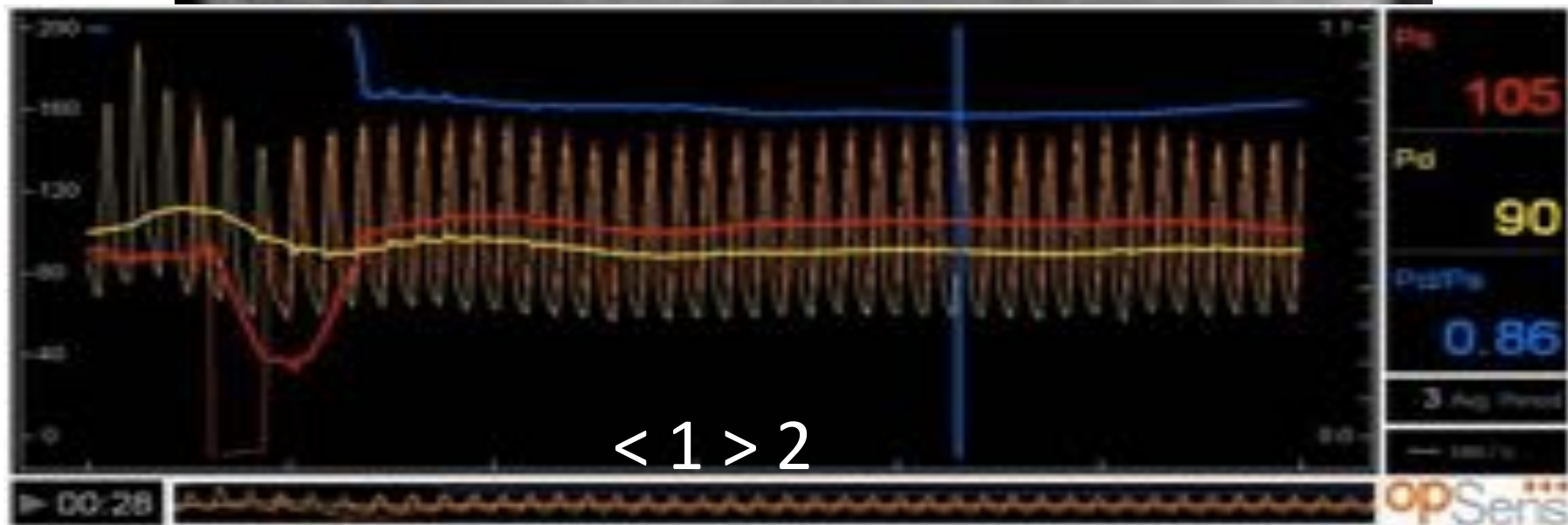
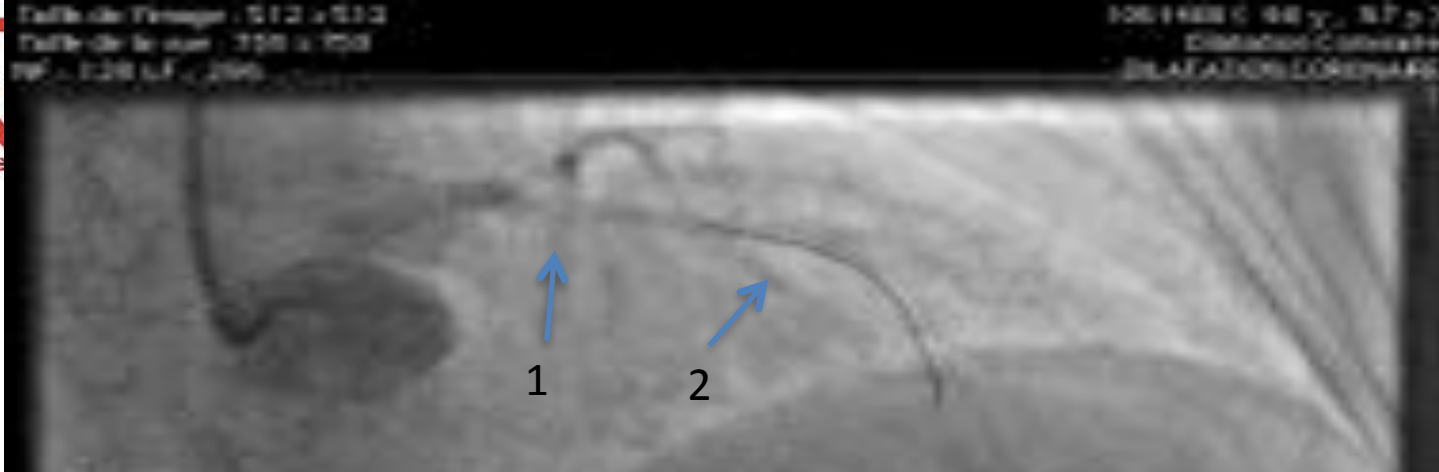
Roc BK et al 2008	110 patients with provisional strategy, SB intervention when FFR <0.75 . Control group 110 patients without FFR measurements.	- The FFR-guided group showed significantly less frequent SB intervention (30% in FFR-guided vs. 42% in angiography-guided group, $p=0.03$). - There was no difference in 9-month TMR (4.6% vs. 3.7%, $p=0.7$).
DIORIS-B-VI 2014	320 patients with Medina 1,1,1 or 0,1,1 bifurcation lesions. Randomly assigned to FFR-guided FFR <0.80 or angiography-guided SB treatment.	- Treatment of SB was less in FFR-guided group than in angiography-guided group (SB stenting: 25.9% vs. 38.1%, $p=0.01$). - NACE (cardiac death, MI, TMR) rate at 1 year was comparable (18.1% vs. 18.1%, $p=1.00$). Restenosis at distal SB was more frequent in angiography-guided group than in FFR-guided group (9.2% vs. 1.2%, $p=0.01$).

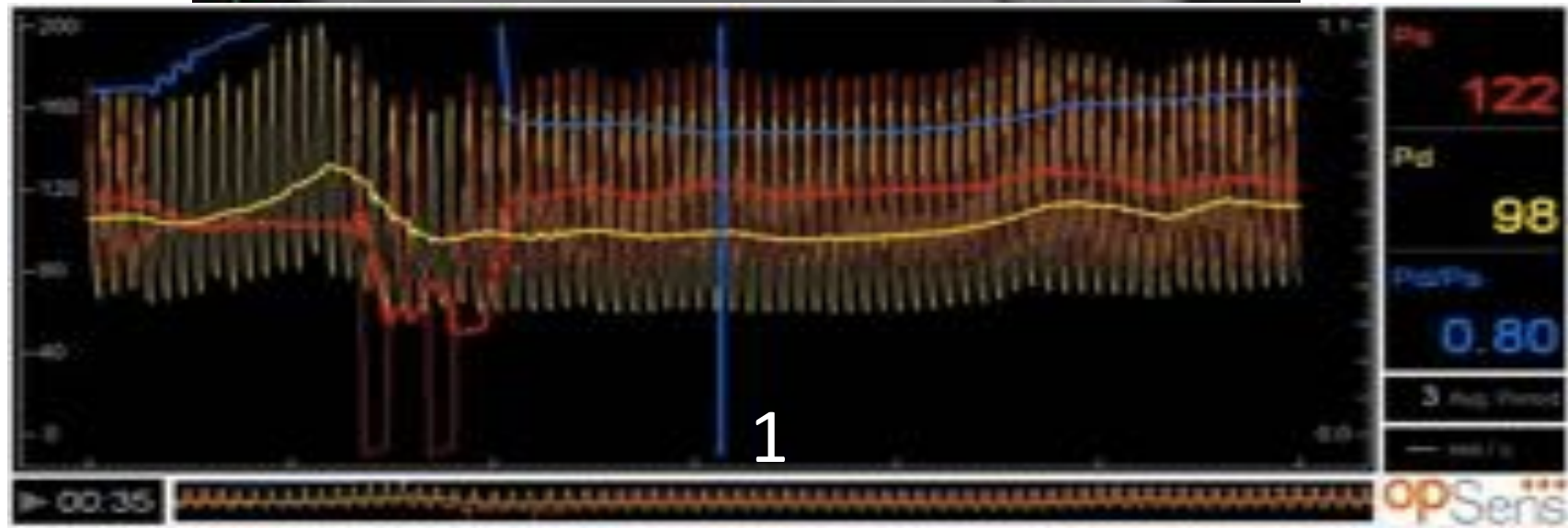
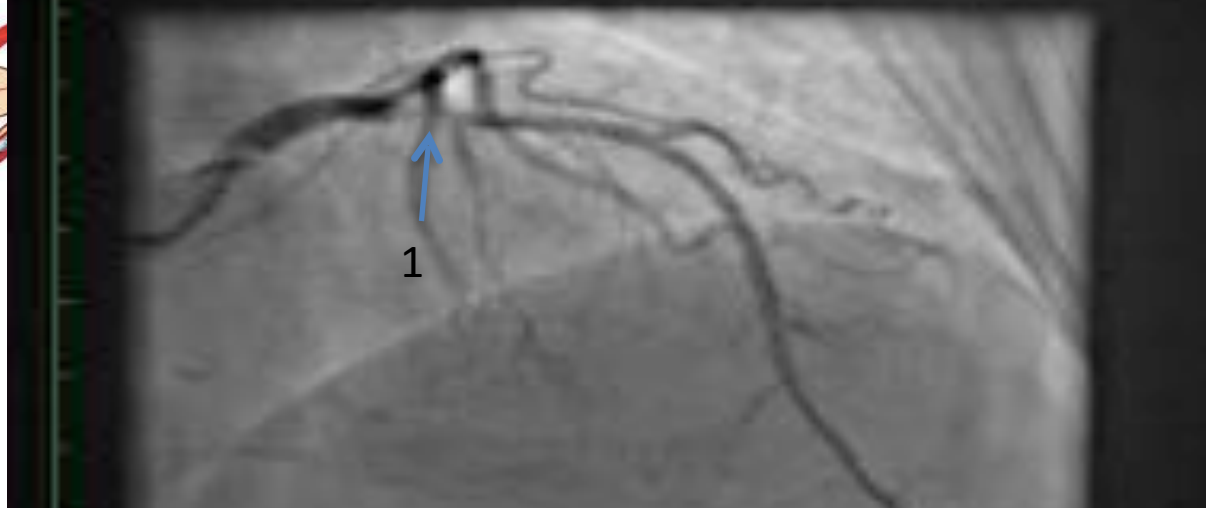
Conclusion

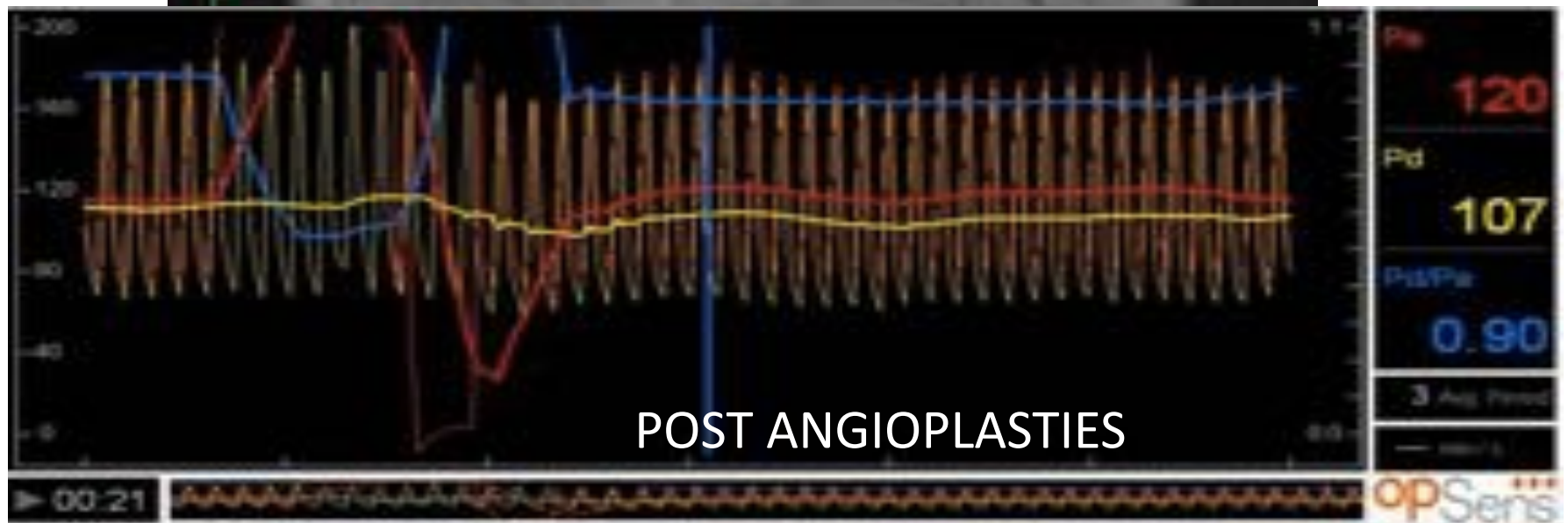
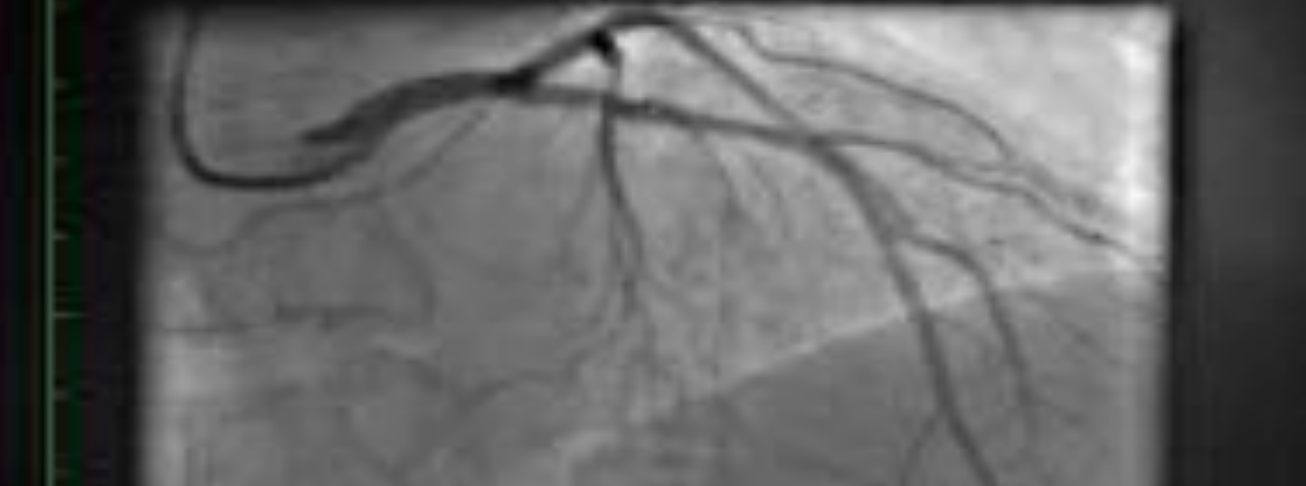
- Quelques avantages .. À la marge :
- Il est difficile de montrer un bénéfice clinique à l'utilisation de la FFR dans les bifs
- Souvent, petit territoire (sauf TC)
- Avantage au niveau de la simplification de la procédure en limitant le nombre de kissing

LÉSIONS MULTIPLES







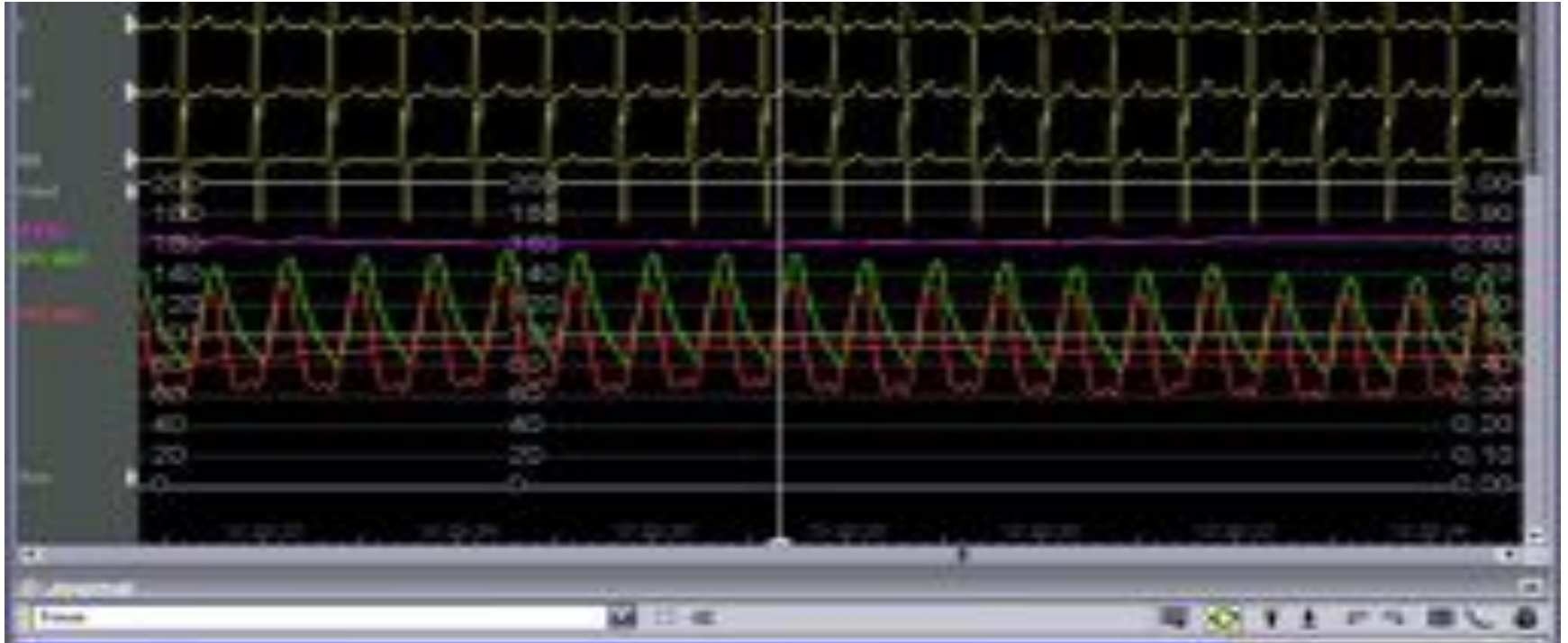


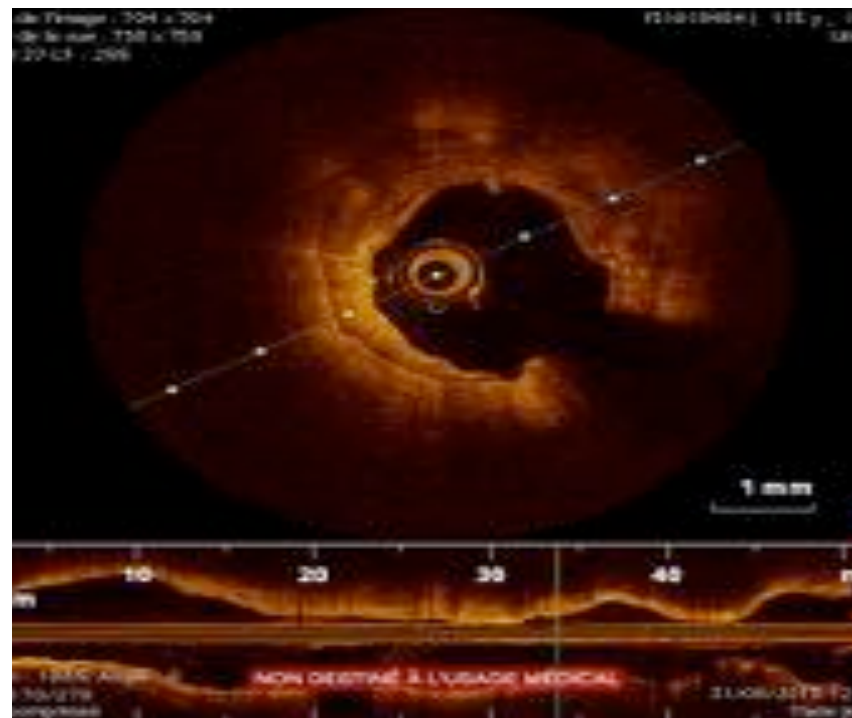
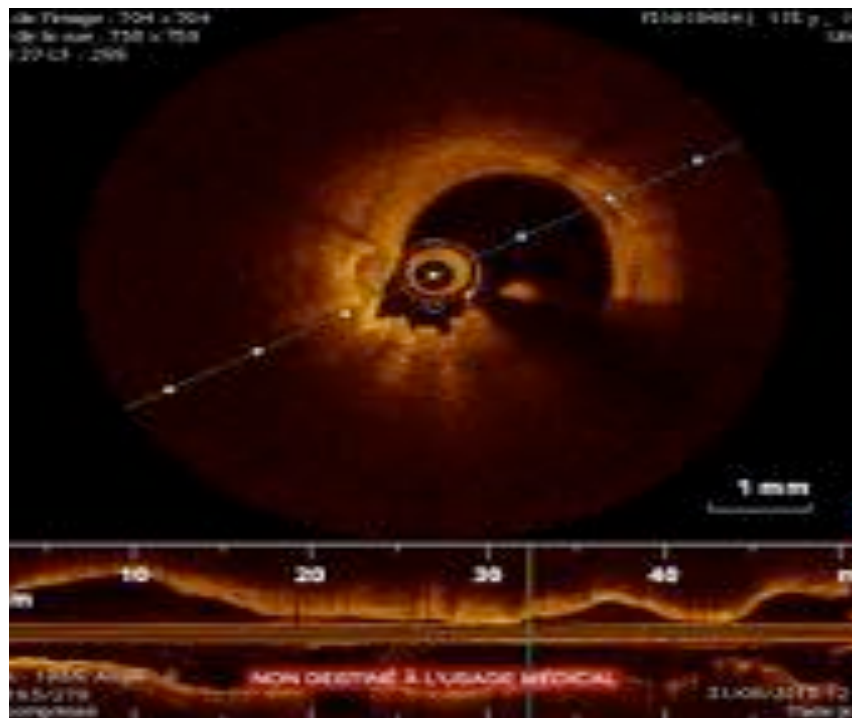
IMPORTANCE DU TERRITOIRE DEPENDANT DE LA LESION





$FFR = 0,80 \dots$



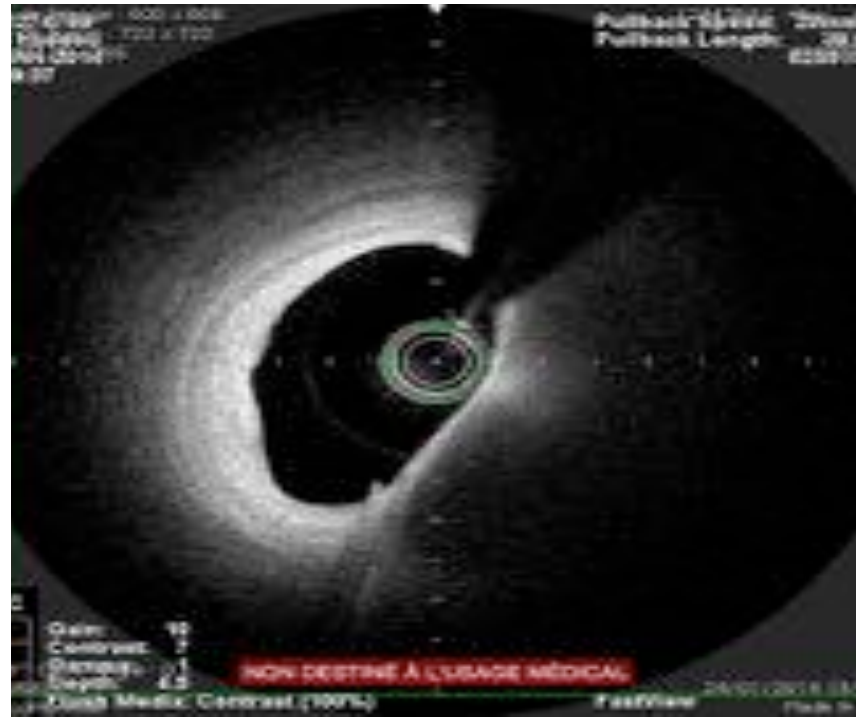




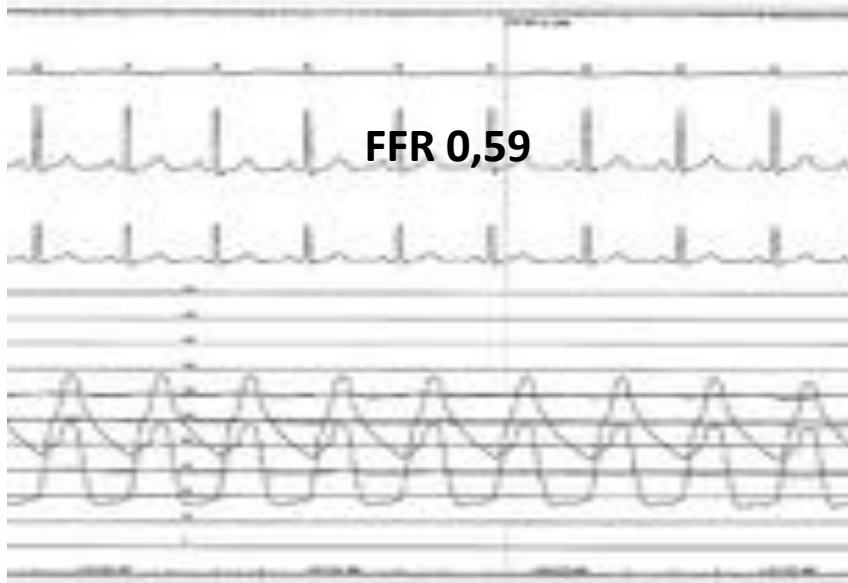
40 y Woman. Chest pain at stress



LM ostium 5 mm²



FFR LAD



FFR LCX

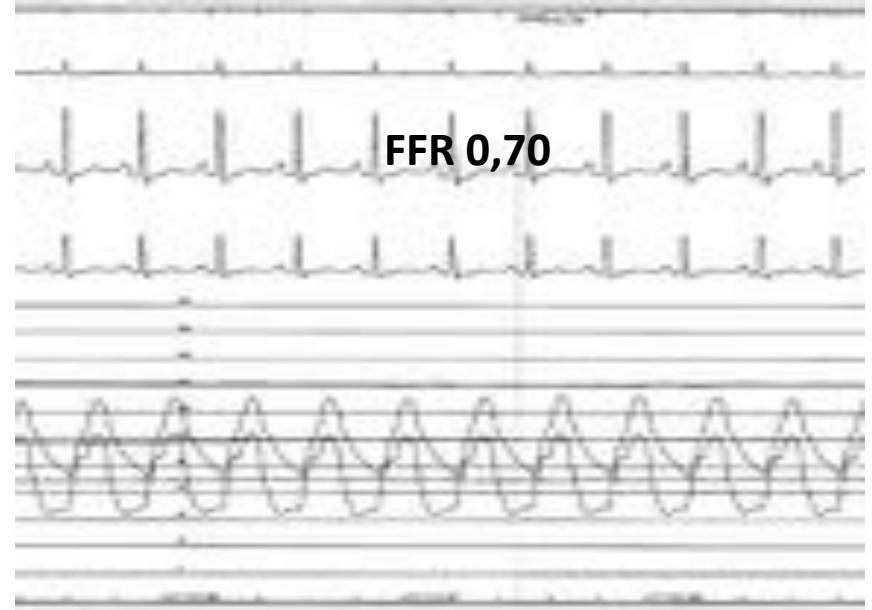


Table 5. Correlates of FFR

Lesion severity (1% decrease of % stenosis)

Number of diseased vessels by angiography

1

2

3

Lesion complexity ACC/AHA: B2/C vs A/B1 by angiography†

Diabetes mellitus

Age (1-y increase)

Smoking

Nonischemic testing (reference group: not done)

Not treated

Positive

Oral intensity

Lesion length (1-mm increase)

Pharmacologic (including nitro) (reference group: <50%)

30%–50%

>50%

LAD location

Male sex

Previous revascularization

1

2

3

Age (1-y increase)

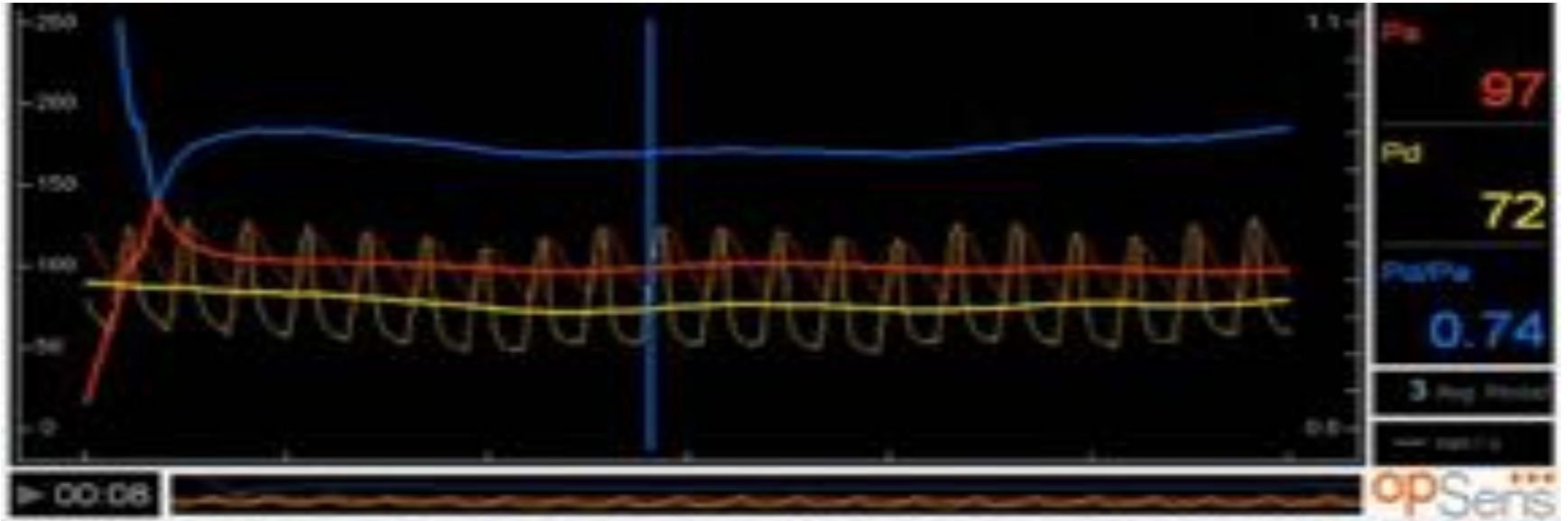
Lesion complexity ACC/AHA: B2/C vs A/B1 by angiography*

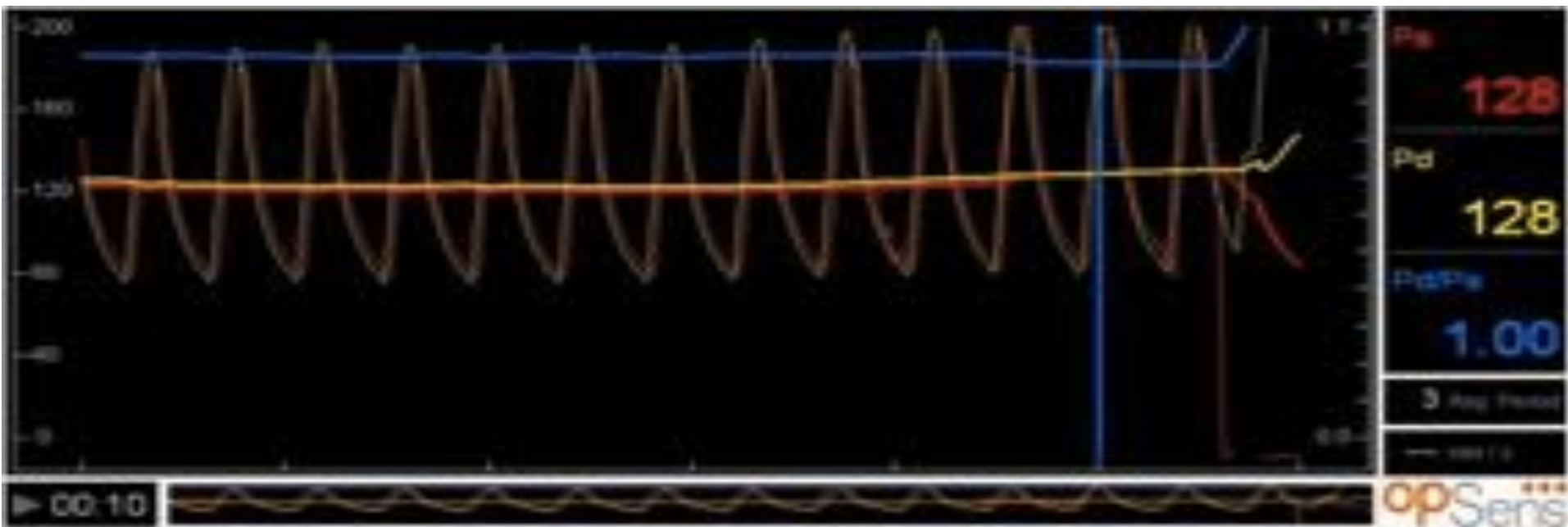
	Odds Ratio*	Lower CI	Upper CI	PValue
Lesion severity (1% decrease of % stenosis) by angiography*	1.03	1.02	1.04	<0.0001
Number of diseased vessels by angiography				<0.0001
1	2.41	1.52	3.84	
2	2.64	1.64	4.24	
3	3.75	2.31	6.08	
Lesion complexity ACC/AHA: B2/C vs A/B1 by angiography†	1.52	1.20	1.93	0.0005
Diabetes mellitus	1.42	1.13	1.78	0.003
Age (1-y increase)	0.98	0.97	0.99	0.005

APRES FFR

	Estimates	Lower CI	Upper CI	PValue
LAD location*	−0.05	−0.06	−0.03	<0.0001
Lesion length (1-mm increase)*	−0.01	−0.01	−0.01	<0.0001
Lesion severity (1% decrease of % stenosis)*	−0.01	−0.01	−0.01	<0.0001
Number of diseased vessels (reference group=0)				<0.0001
1	−0.02	−0.03	−0.01	
2	−0.04	−0.06	−0.03	
3	−0.03	−0.05	−0.02	
Age (1-y increase)	0.01	0.01	0.01	<0.0001
Lesion complexity ACC/AHA: B2/C vs A/B1 by angiography*	−0.01	−0.02	−0.01	0.001

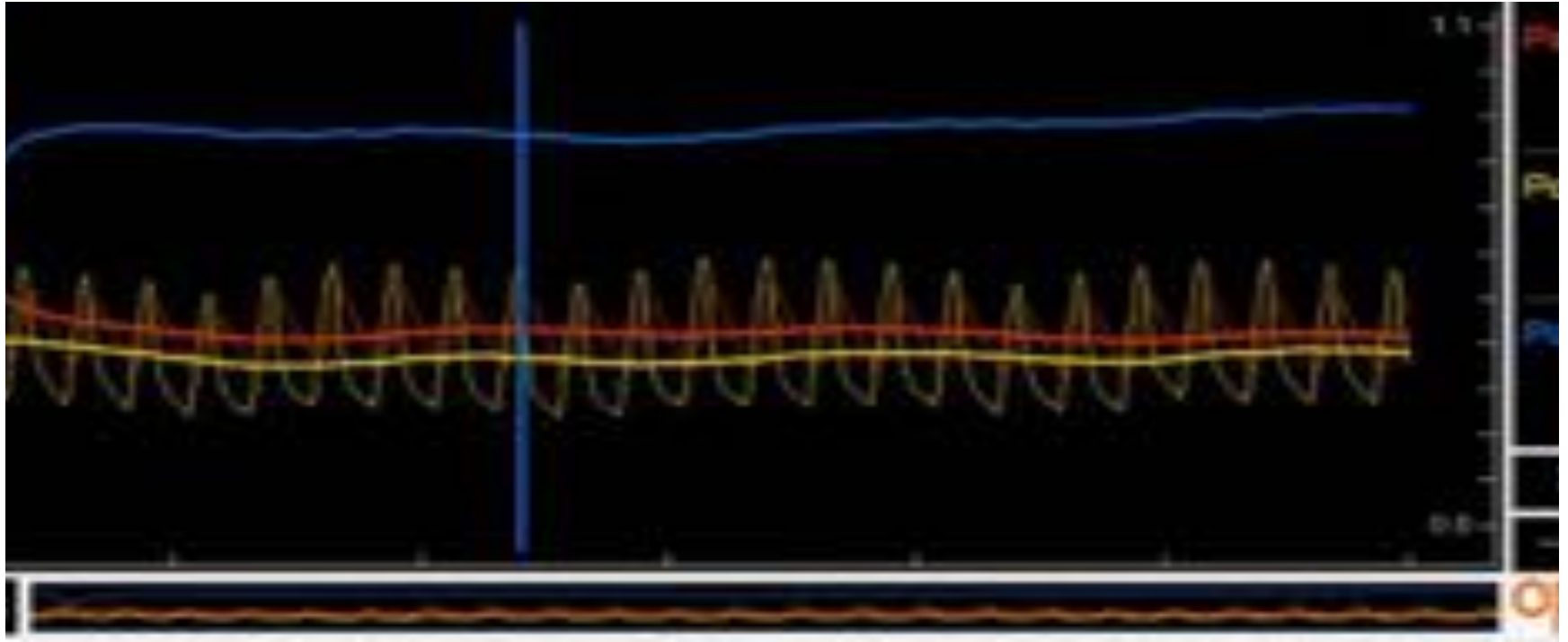
Gradient adenosine

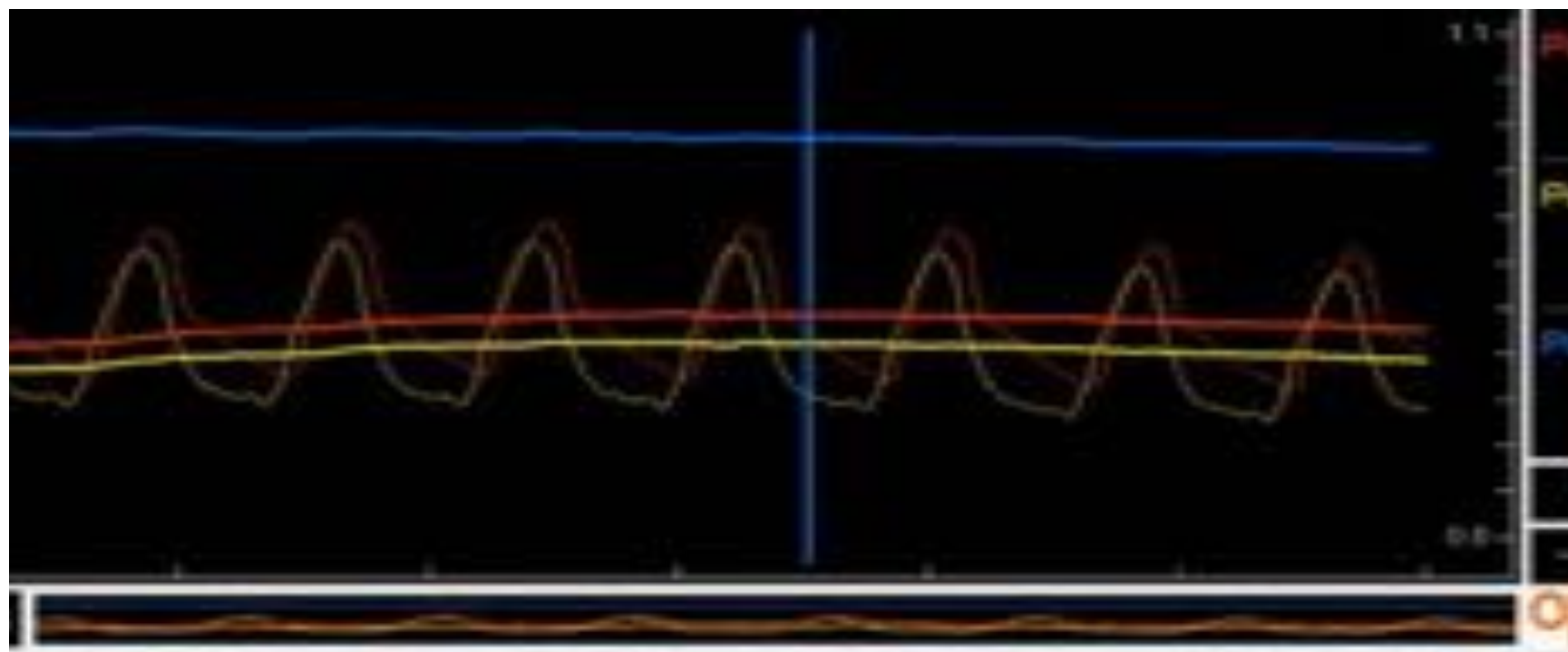


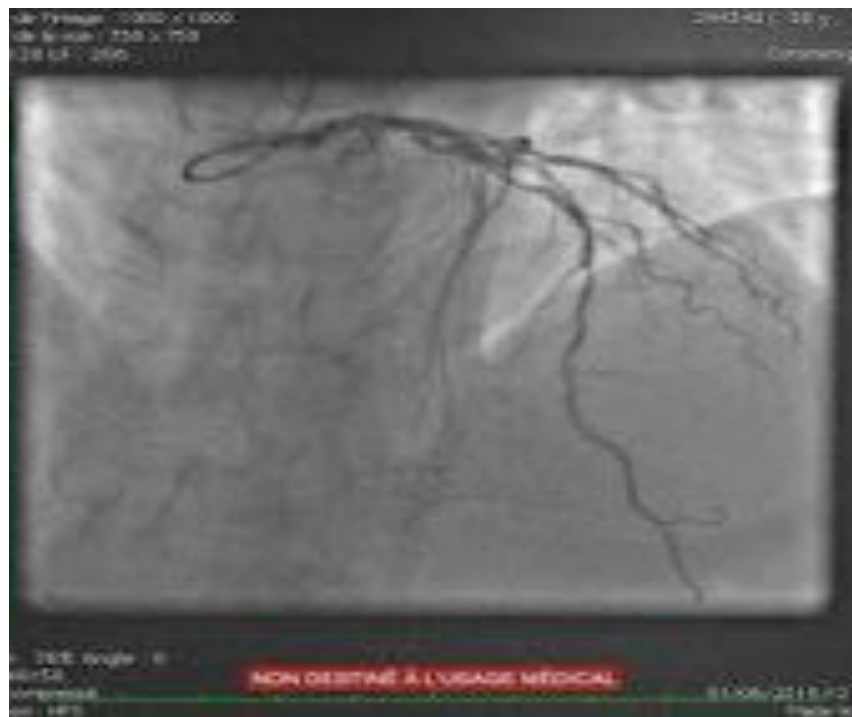




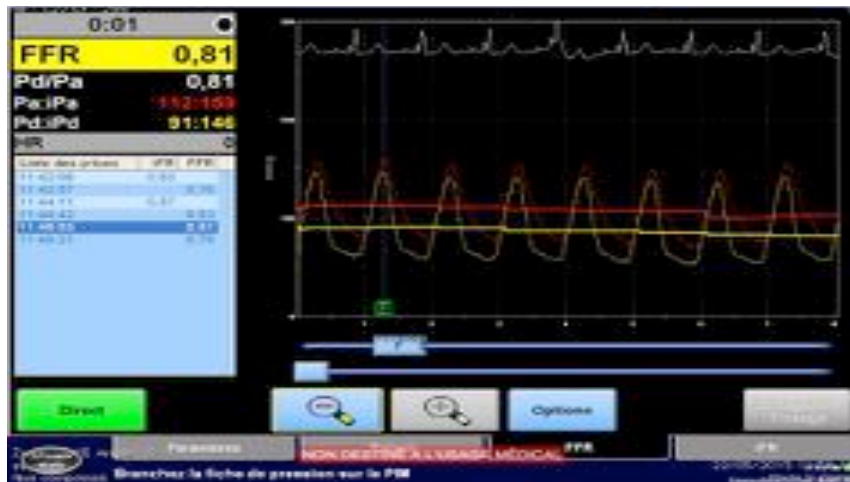
Adénosine



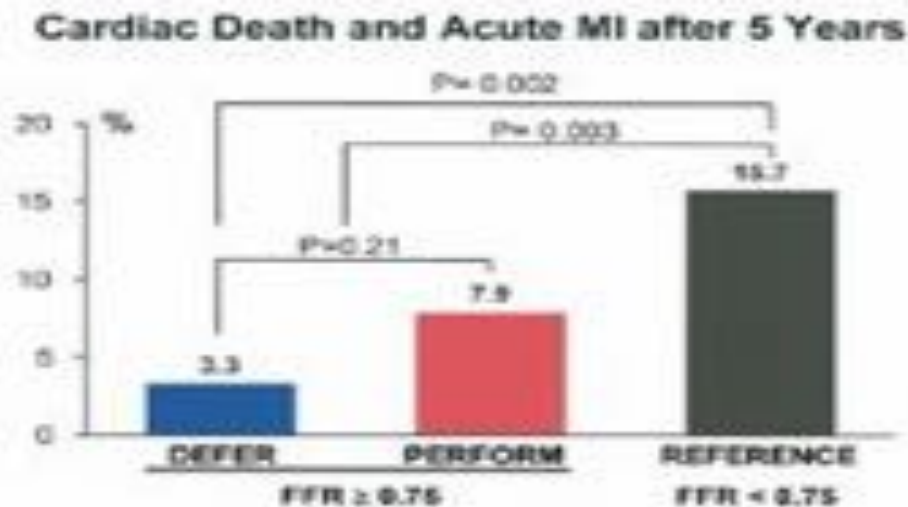




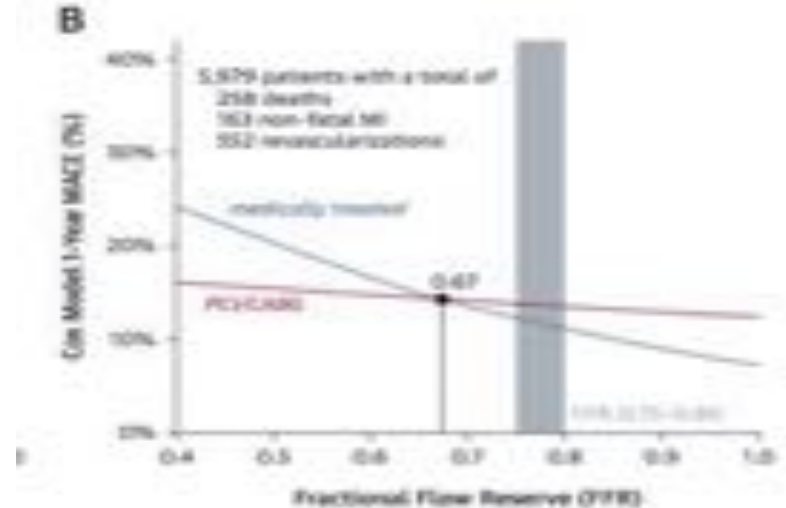
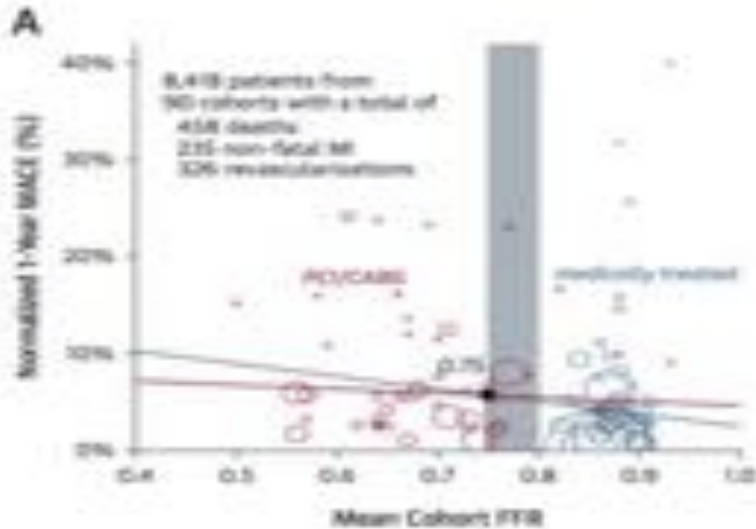
- Traitement médical ?
- Revascularisation .. Quand même ?
- Defer like mais une autre époque



J Am Coll Cardiol 2007;49:2105–11



FFR marqueur d'événements CV



Euro
PCR
2015

Overall FFR/patients

Study diagram



Prevalence of
gray zone

2781 (16%)
FFR 0.76 - 0.80

6531 excluded

Multiple measurements
Multiple stenoses
Bypass grafts
In-stent restenosis
Myocardial bridge
Heart transplants

17380 FFR
measurements

7995 (46%)
FFR 0.70 - 0.85

1459 (8%)
FFR 0.70 - 0.85

FFR/patients in the
study range

Med Rx 1010 (70%)

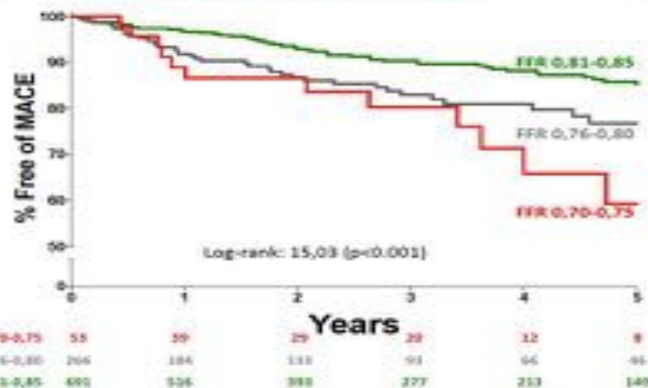
5 years Follow up

Revasc 449 (30%)

Euro
PCR
2015

MACE-Rate per FFR strata

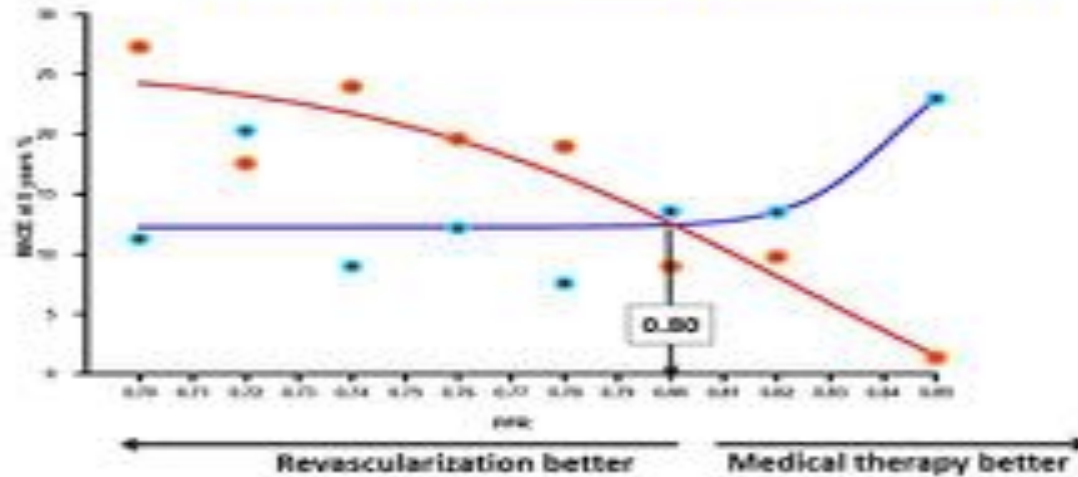
Pts with MT (n=1010)



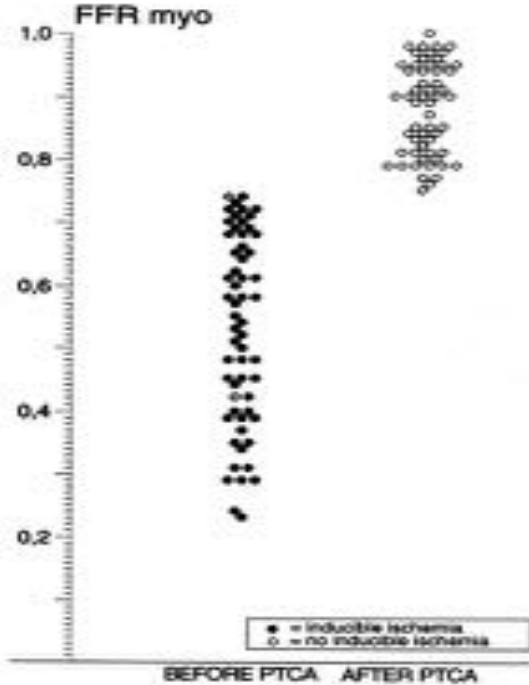
MACE-Rate per FFR strata

Pts with MT (n=1010)

Pts with Revascularisation (n=449)



L'évolution du concept de FFR



- Concept physiologique
- Concept Fonctionnel
- Concept Pronostic clinique
- Marqueur d'événements

Pijls N et al. Circulation

1995;92:2422-2434