

4to SIMPOSIO DE MEDICINA CARDIOVASCULAR “CENTENARIO 2018”

¿Cómo se hace la mejor angioplastia primaria en 2018?

Datos para una reperfusión efectiva

Leandro Lasave

Instituto Cardiovascular de Rosario

Sanatorio Parque Rosario

Director Curso de Fellows ProEducar SOLACI

www.icronline.com

Datos para una reperfusión efectiva

- 1- Tiempos
- 2- DAPT
- 3- Vía de Acceso
- 4- Anticoagulación – Antitrombosis
- 5- Tromboaspiracion
- 6- STENT: DES vs BMS – Directo vs Diferido
- 7- Solo ARI o MV?

Datos para una reperfusión efectiva

1- Tiempos

2- DAPT

3- Vía de Acceso

4- Anticoagulación – Antitrombosis

5- Tromboaspiracion

6- STENT: DES vs BMS – Directo vs Diferido

7- Solo ARI o MV?

TIEMPOS

empos:

es de traslado y activación.

riage prehosp adecuado hacia centros

4/7

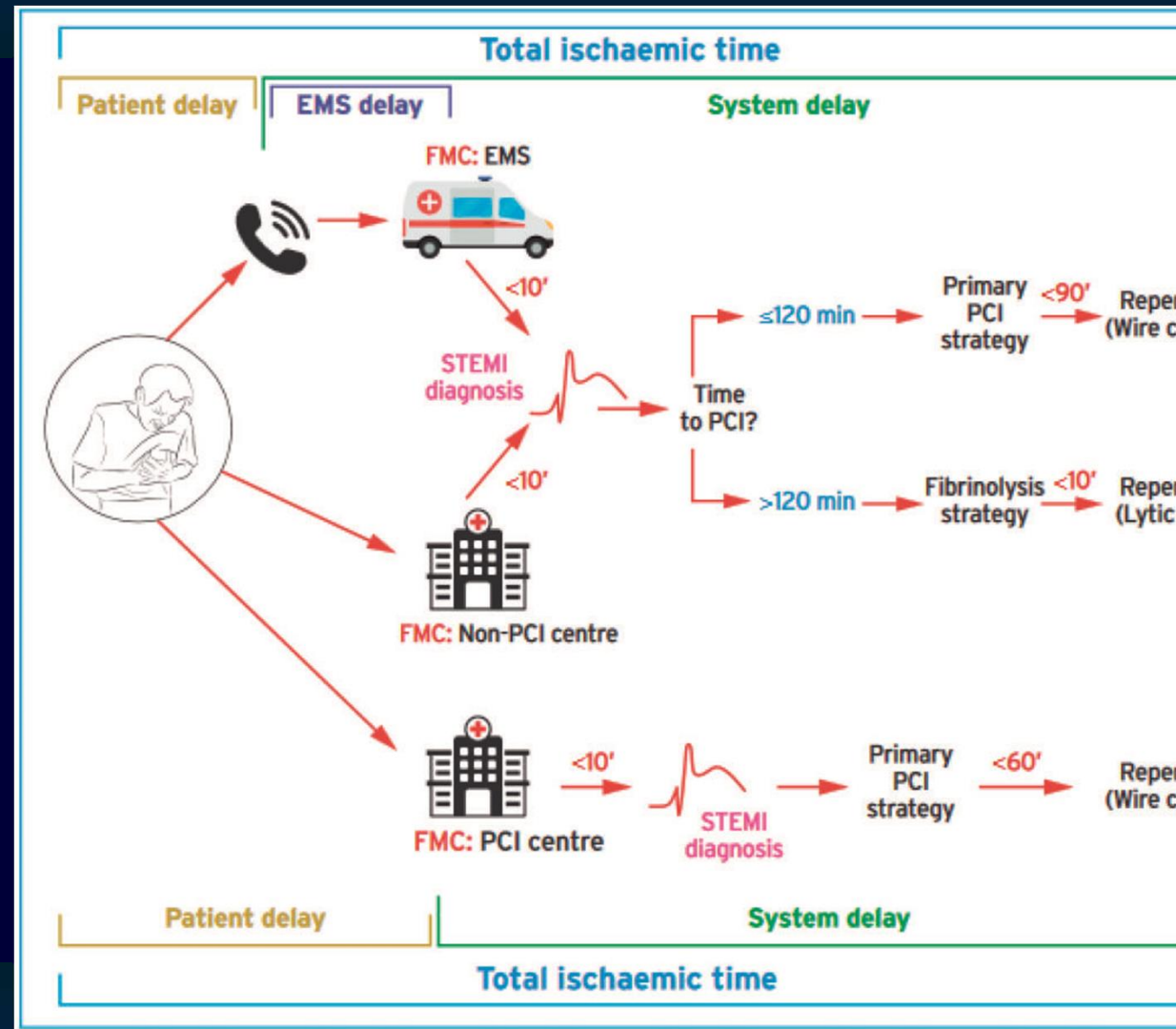
reactivar al Centro con envío de ECG

evitar el paso por la sala de guardia e ir

directo a sala de hemodinamia

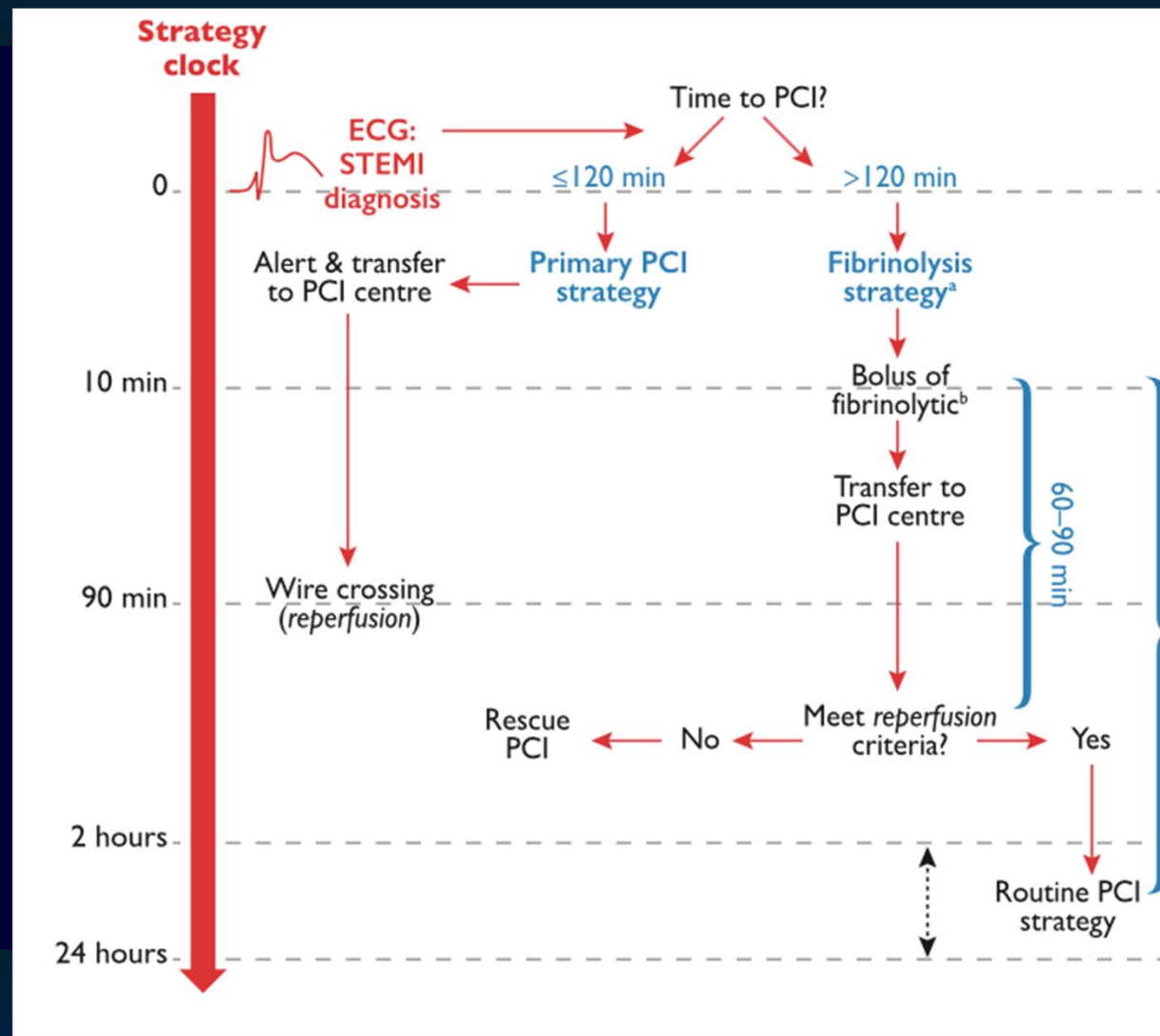
Maximo tiempo esperable para PPCI

120 min FMC



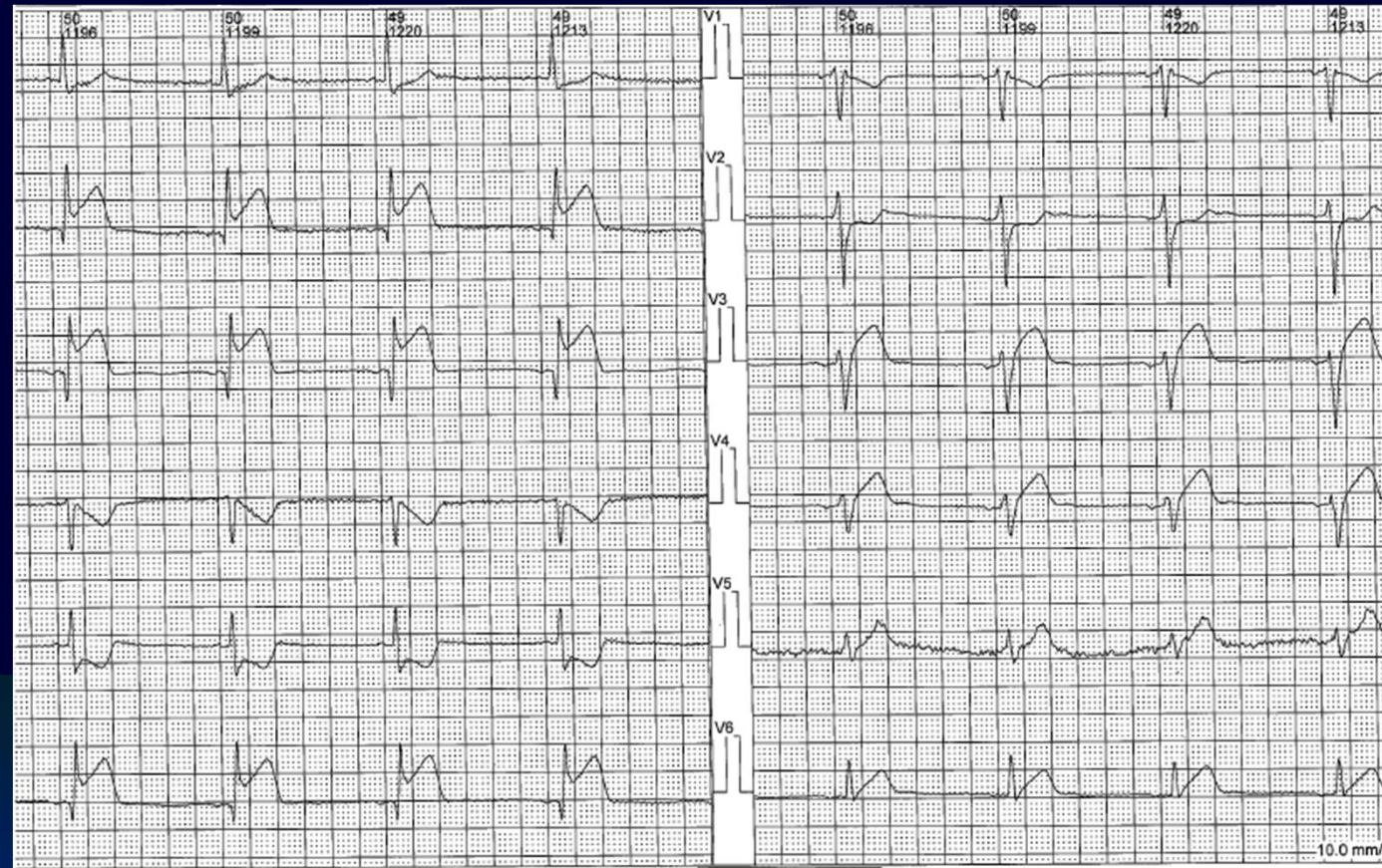
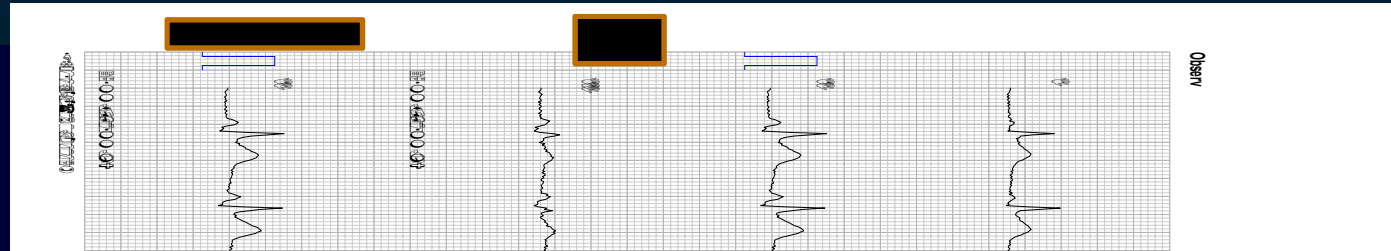
TIEMPOS

Intervals	Time targets
Maximum time from FMC to ECG and diagnosis ^a	≤10 min
Maximum expected delay from STEMI diagnosis to primary PCI (wire crossing) to choose primary strategy over fibrinolysis (if this target time can be met, consider fibrinolysis)	≤120 min
Maximum time from STEMI diagnosis to wire crossing in patients presenting at primary PCI centres	≤60 min
Maximum time from STEMI diagnosis to wire crossing in transferred patients	≤90 min
Maximum time from STEMI diagnosis to bolus or start of fibrinolysis in patients unable to meet primary PCI target times	≤10 min
Delay from start of fibrinolysis to evaluation of efficacy (success or failure)	60–90 min
Delay from start of fibrinolysis to reperfusion (if fibrinolysis is successful)	2–24 hours



CASO CLINICO

on, 58 años
antecedentes
gina típica, 1 h evolucion
mada al SEM 12:20 hs
ivo SEM 12:37 hs
G: 12:41 hs
ivación Centro 24/7: 12:51hs
reso sala hemodinamia 13: 24 hs



Datos para una reperfusión efectiva

1- Tiempos

2- DAPT

3- Vía de Acceso

4- Anticoagulación – Antitrombosis

5- Tromboaspiración

6- STENT: DES vs BMS – Directo vs Diferido

7- Solo ARI o MV?

ANTIAGREGACIÓN PLAQUETARIA

Todo paciente derivado a PPCI debe recibir DAPT

- AAS + inhibidor P2Y12

AAS:

- Oral: 150 – 300 mg
- IV: 250 – 500 mg

P2Y12:

Ticagrelor 180 mg ud + 90 mg cada 12 hs

Prasugrel 60 mg ud + 10 mg dia

Clopidogel 600 mg ud + 150 o 75 mg dia

Cangrelor IV (no recomendado)

Inhibidores GPIIbIIIa: no en forma rutinaria. Solo como rescate

ACUTE Trial. Thromb Haemost 2017;117(3):625–635.

No diferencia en administrar en ambulacion vs sala hemodi
Estudio ATLANTIC. N Engl J Med 2014;371(11):1016–10

CURRENT-OASIS 7. Lancet . 2010;376(9748):1233–124

ANTIAGREGACIÓN PLAQUETARIA

Recommendations	Class ^b	Level ^c
Antiplatelet therapy		
A potent P2Y ₁₂ inhibitor (prasugrel or ticagrelor), or clopidogrel if these are not available or are contraindicated, is recommended before (or at latest at the time of) PCI and maintained over 12 months, unless there are contraindications such as excessive risk of bleeding. ^{186,187}	I	A
Aspirin (oral or i.v. if unable to swallow) is recommended as soon as possible for all patients without contraindications. ^{213,214}	I	B
GP IIb/IIIa inhibitors should be considered for bailout if there is evidence of no-reflow or a thrombotic complication.	IIa	C
Cangrelor may be considered in patients who have not received P2Y ₁₂ receptor inhibitors. ^{192–194}	IIb	A

Antiplatelet therapies	
Aspirin	Loading dose of 150–300 mg orally or of 75–250 mg i.v. if oral ingestion is not possible, followed by a maintenance dose of 75–100 mg/day
Clopidogrel	Loading dose of 600 mg orally, followed by a maintenance dose of 75 mg/day
Prasugrel	Loading dose of 60 mg orally, followed by a maintenance dose of 10 mg/day In patients with body weight ≤60 kg, a maintenance dose of 5 mg/day is recommended Prasugrel is contra-indicated in patients with previous stroke. In patients ≥75 years, prasugrel is generally not recommended, but a dose of 5 mg/day should be used if treatment is deemed necessary
Ticagrelor	Loading dose of 180 mg orally, followed by a maintenance dose of 90 mg <i>b.i.d.</i>
Abciximab	Bolus of 0.25 mg/kg i.v. and 0.125 µg/kg/min infusion (maximum 10 µg/min) for 12 hours
Eptifibatide	Double bolus of 180 µg/kg i.v. (given at a 10-min interval) followed by an infusion of 2.0 µg/kg/min for up to 18 hours
Tirofiban	25 µg/kg over 3 min i.v., followed by a maintenance infusion of 0.15 µg/kg/min for up to 18 hours

www.icronline.com

2017 Guidelines STEMI. European Heart Journal (2018) 39, 119

Datos para una reperfusión efectiva

1- Tiempos

2- DAPT

3- Vía de Acceso

4- Anticoagulación – Antitrombosis

5- Tromboaspiración

6- STENT: DES vs BMS – Directo vs Diferido

7- Solo ARI o MV?

ACCESOS

- Radial
- Femoral



ACCESOS

- Radial
- Femoral

Desventajas del acceso Transradial en la PPCI

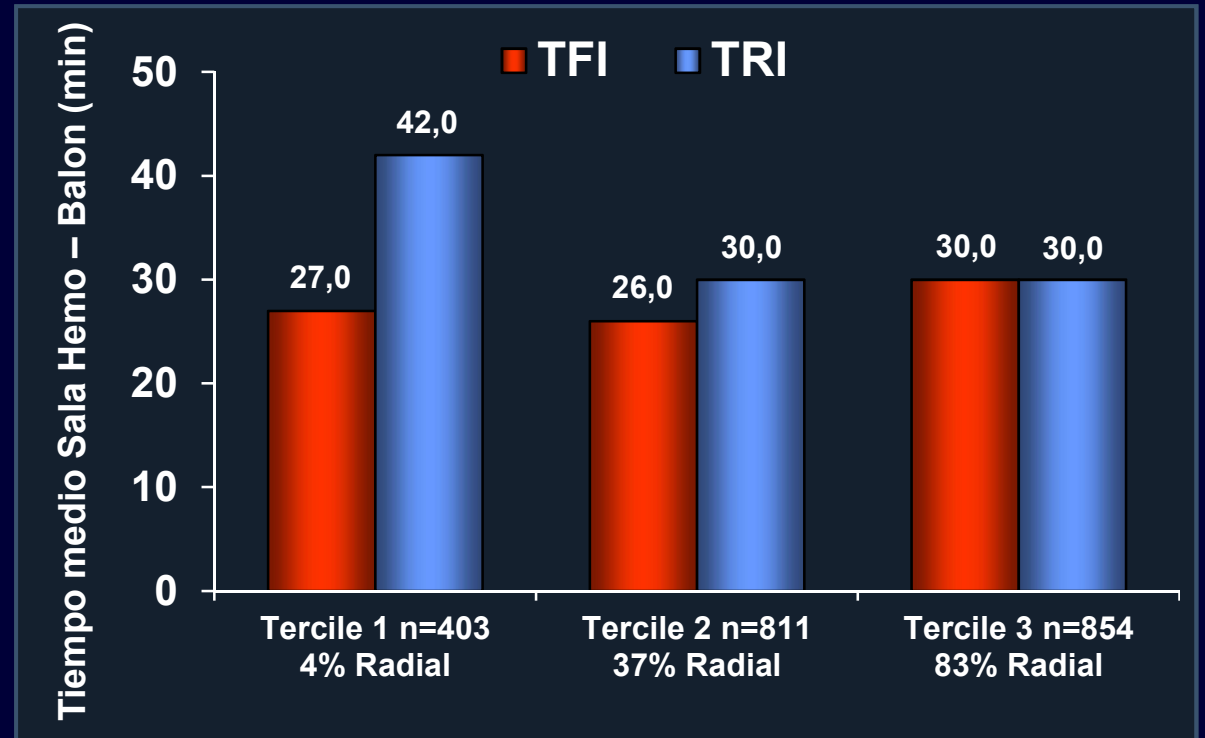
- Tasa de Crossover 5-7% en Radial STEMI Trials
 - espasmo, tortuosidad, soporte guía
- Aumento tiempo de fluoroscopia
- Aumento volumen de contraste
- Retraso en la reperfusión
- Todas las medidas y resultados dependen de la experiencia del centro y del volumen

Curva de aprendizaje

www.icronline.com

ACCESOS

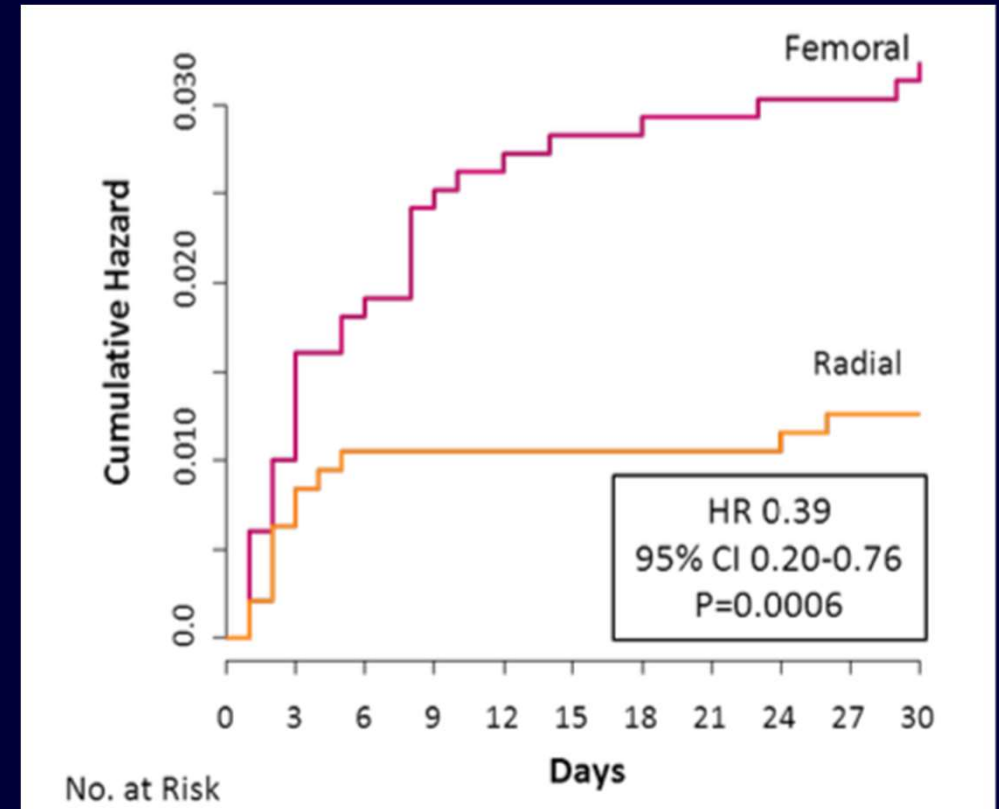
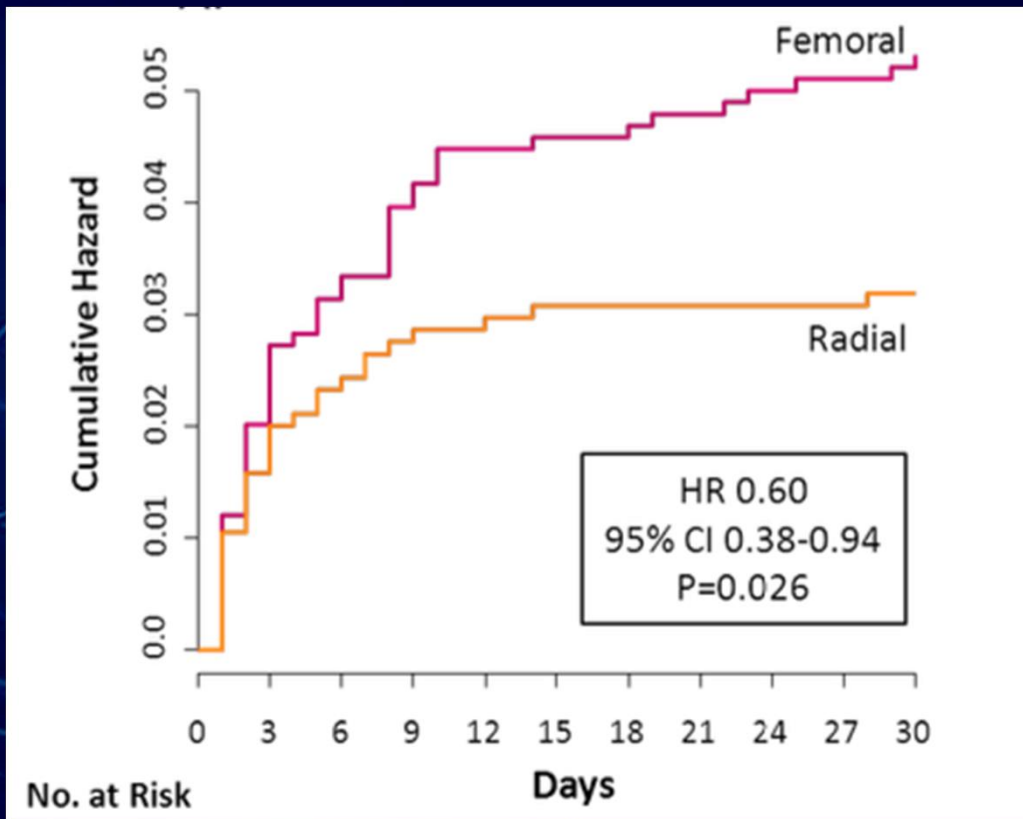
- Tiempo de reperfusión en PPCI según la experiencia del centro en TRI



Cantor W, et al. *Circ Cardiovasc Interv.* 2015;8:e002097

Evidencia TRI en PPCI

RIVAL Trial. Menor tasa de MACE y Mortalidad



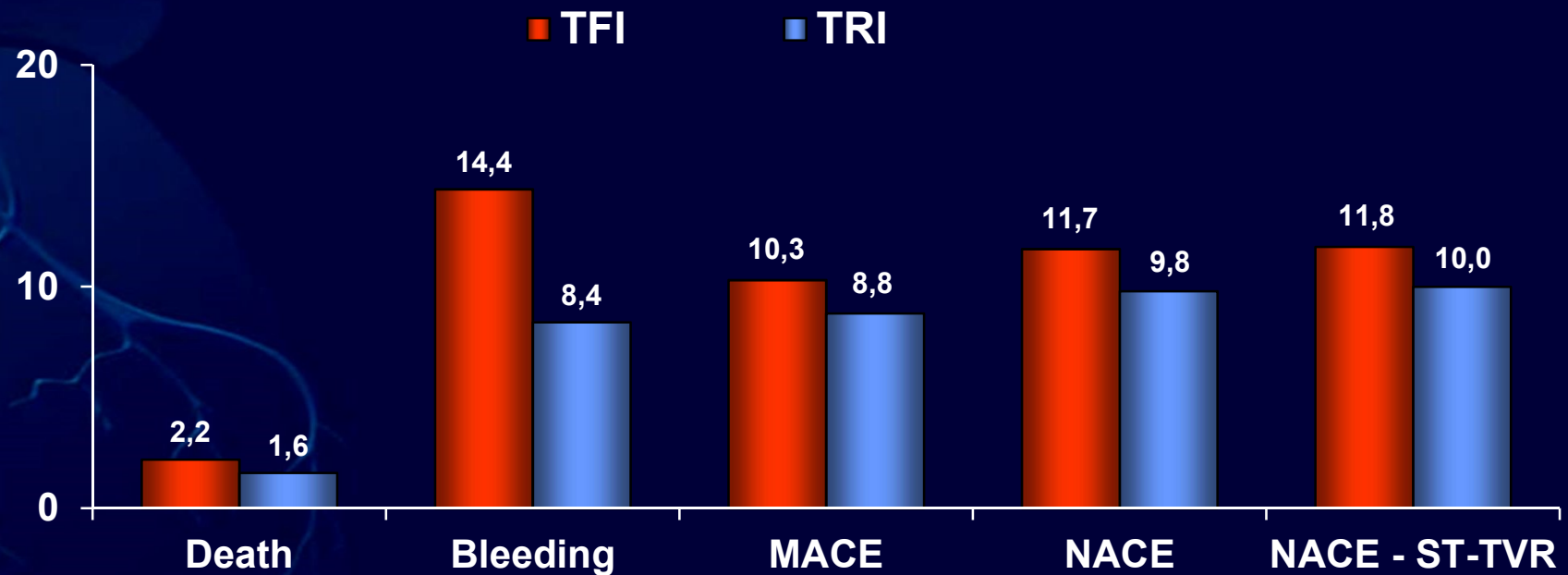
Muerte CV, IAM, ACV, Major Bleeding

Metha S, et al. RIVAL study. J Am Coll Cardiol 2012;60:2490-9

www.icronline.com

Evidencia TRI en PPCI

MATRIX Trial. 8,404 pts. Menor tasa de eventos a 30 días



MACE (Major Adverse Cardiovascular Events) : All cause mortality - Myocardial infarction - Stroke.

NACE (Net Adverser Clinical Events): MACE +BARC 3 or 5 bleeding.

ST: Stent Thrombosis. TVR: Urgent Target Vessel Revascularization

Valgimigli M, et al. MATRIX trial. Lancet 2015; 385: 2465–76

www.icronline.com

– ACCESOS

CHANGE IN RECOMMENDATIONS 2017

Radial access^a MATRIX ¹⁴³	
DES over BMS EXAMINATION ^{144,145} COMFORTABLE-AMI ¹⁴⁶ , NORSTENT ¹⁴⁷	
Complete Revascularization^b PRAMI ¹⁴⁸ , DANAMI-3-PRIMULTI ¹⁴⁹ , CVLPRUT ¹⁵⁰ , Compare-Acute ¹⁵¹	
Thrombus Aspiration^c TOTAL ¹⁵² , TASTE ¹⁵³	
Bivalirudin MATRIX ¹⁵⁴ , HEAT-PPCI ¹⁵⁵	
Enoxaparin ATOLL ^{156,157} , Metz-analysis ¹⁵⁸	
Early Hospital Discharge^d Small trials & observational data ^{159–162}	
when 95% AVOID ¹⁶³ , DETO2X ¹⁶⁴	Oxygen when SaO ₂ <90%
100% TNK-tPA all patients STREAM ¹⁶⁵	Dose I.V. TNK-tPA half in Pts ≥75 years

2017 NEW RECOMMENDATIONS

- Additional lipid lowering therapy if LDL >1.8 mmol/L (70 mg/dL) despite on maximum tolerated statins IMPROVE-IT¹⁶⁶, FOURIER¹⁶⁷
- Complete revascularization during index primary PCI in STEMI patients in shock
Expert opinion
- Cangrelor if P2Y₁₂ inhibitors have not been given CHAMPION¹⁶⁸
- Switch to potent P2Y₁₂ inhibitors 48 hours after fibrinolysis
Expert opinion
- Extend Ticagrelor up to 36 months in high-risk patients PEGASUS-TIMI 54¹⁶⁹
- Use of poly-pill to increase adherence FOCUS¹⁷⁰
- Routine use of deferred stenting DANAMI 3-DEFER¹⁷¹

I
IIa
IIb
III

IRA technique

Stenting is recommended (over balloon angioplasty) for primary PCI. ^{146,147}	I
Stenting with new-generation DES is recommended over BMS for primary PCI. ^{148–151,178,179}	I
Radial access is recommended over femoral access if performed by an experienced radial operator. ^{143–145,180}	I
Routine use of thrombus aspiration is not recommended. ^{157,159}	III
Routine use of deferred stenting is not recommended. ^{153–155}	III

2017 Guidelines STEMI. European Heart Journal (2018) 39, 119–177

Datos para una reperfusión efectiva

1- Tiempos

2- DAPT

3- Vía de Acceso

4- Anticoagulación – Antitrombosis

5- Tromboaspiración

6- STENT: DES vs BMS – Directo vs Diferido

7- Solo ARI o MV?

– ANTICOAGULACIÓN

- ✓ Heparina no fraccionada
- ✓ Heparina Bajo Peso Molecular – Enoxaparina. ATOLL Trial
- ✓ Bivalirudina

– ANTICOAGULACIÓN

- ✓ Heparina no fraccionada
- ✓ Heparina Bajo Peso Molecular – Enoxaparina. ATOLL Trial
- ✓ Bivalirudina

– ANTICOAGULACIÓN

Bivalirudina: beneficios potenciales

- 1- Disminuir eventos isquémicos y mortalidad
 - Trombosis stent
- 2- Disminuir Sangrado

– ANTICOAGULACIÓN

Bivalirudina: beneficios potenciales

- 1- Disminuir eventos isquémicos y mortalidad
 - Trombosis stent
- 2- Disminuir Sangrado

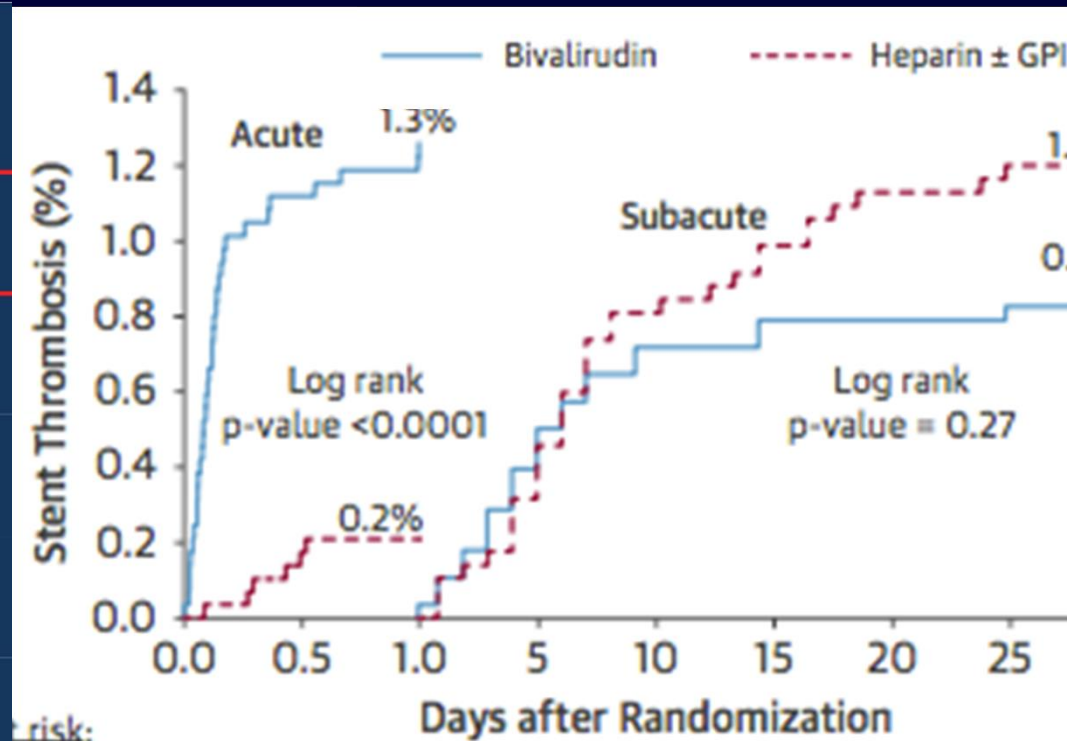
EAT-PPCI

0 Ptes STEMI - HNF 70 U/k vs Bivalirudina // GPI bailout (15y13%) // 80% Ra
MACE 30 ds - PFS: sangrado mayor

	Bivalirudin group (n=905)	Heparin group (n=907)	p value
Primary efficacy outcomes measures	79/905 (8.7%)	52/907 (5.7%)	0.01
Death	46/905 (5.1%)	39/907 (4.3%)	0.43
Stroke	15/905 (1.6%)	11/907 (1.2%)	0.43
Myocardial infarction or reinfarction	24/905 (2.7%)	8/907 (0.9%)	0.004
Additional unplanned target lesion revascularisation	24/905 (2.7%)	6/907 (0.7%)	0.001
Net thrombosis rate	24/697 (3.4%)	6/682 (0.9%)	0.001
Definite	23/697 (3.3%)	5/682 (0.7%)	0.001
Probable	1/697 (0.1%)	1/682 (0.1%)	>0.99
Acute (≤ 24 h)	20/697 (2.9%)	6/682 (0.9%)	0.007
Subacute (>24 h to 28 days)	4/697 (0.6%)	0/682 (0.0%)	0.12

TROMBOSIS STENT

	UFH + GP IIb/IIIa (N=1553)	Bivalirudin (N=1571)	P Value
definite or probable*	1.9%	2.5%	0.33
definite	1.4%	2.2%	0.11
probable	0.5%	0.3%	0.26
acute (≤24 hrs)	0.3%	1.3%	0.0009
subacute (>24 hrs – 30d)	1.7%	1.2%	0.30

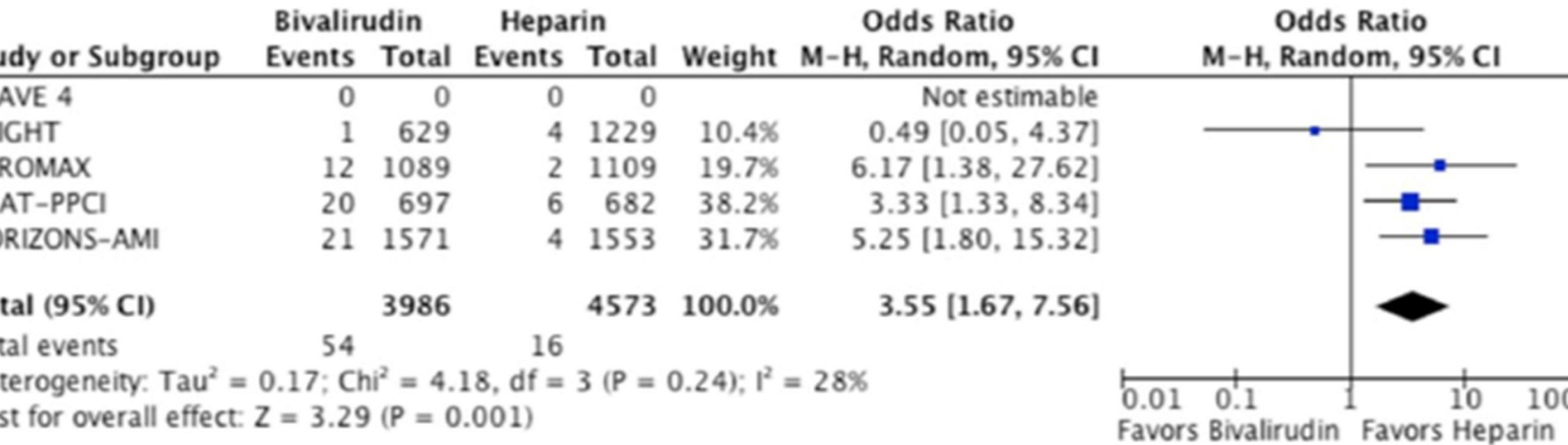


Stone, G.W. et al. J Am Coll Cardiol. 2015; 65(1):27–

Metanalysis.

TROMBOSIS STENT

(c) Definite or probable acute stent thrombosis (0-24 hours)



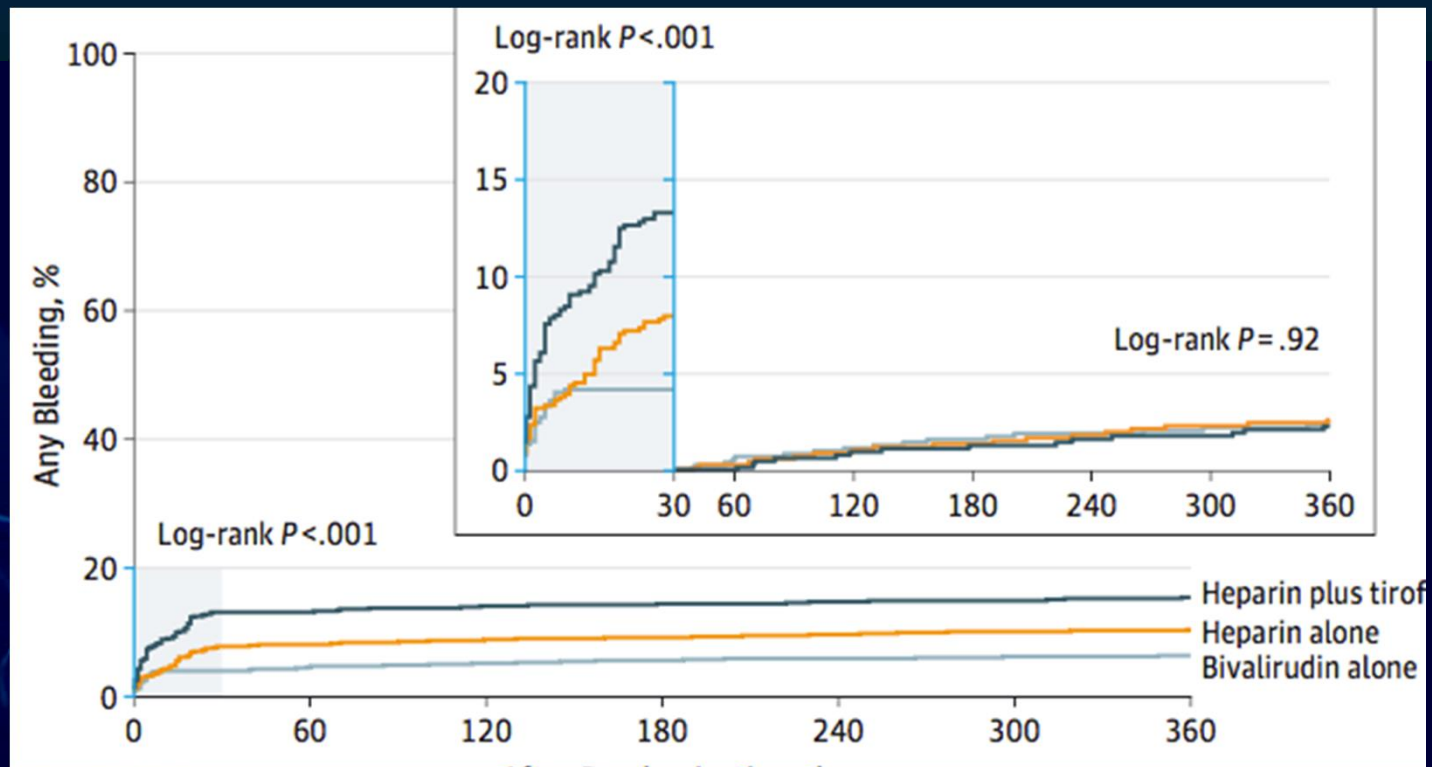
– ANTICOAGULACIÓN

Bivalirudina: beneficios potenciales

- 1- Disminuir eventos isquémicos y mortalidad
 - Trombosis stent
- 2- **Disminuir Sangrado**

BRIGHT

CUALQUIER SANGRADO (BARC 2-5)



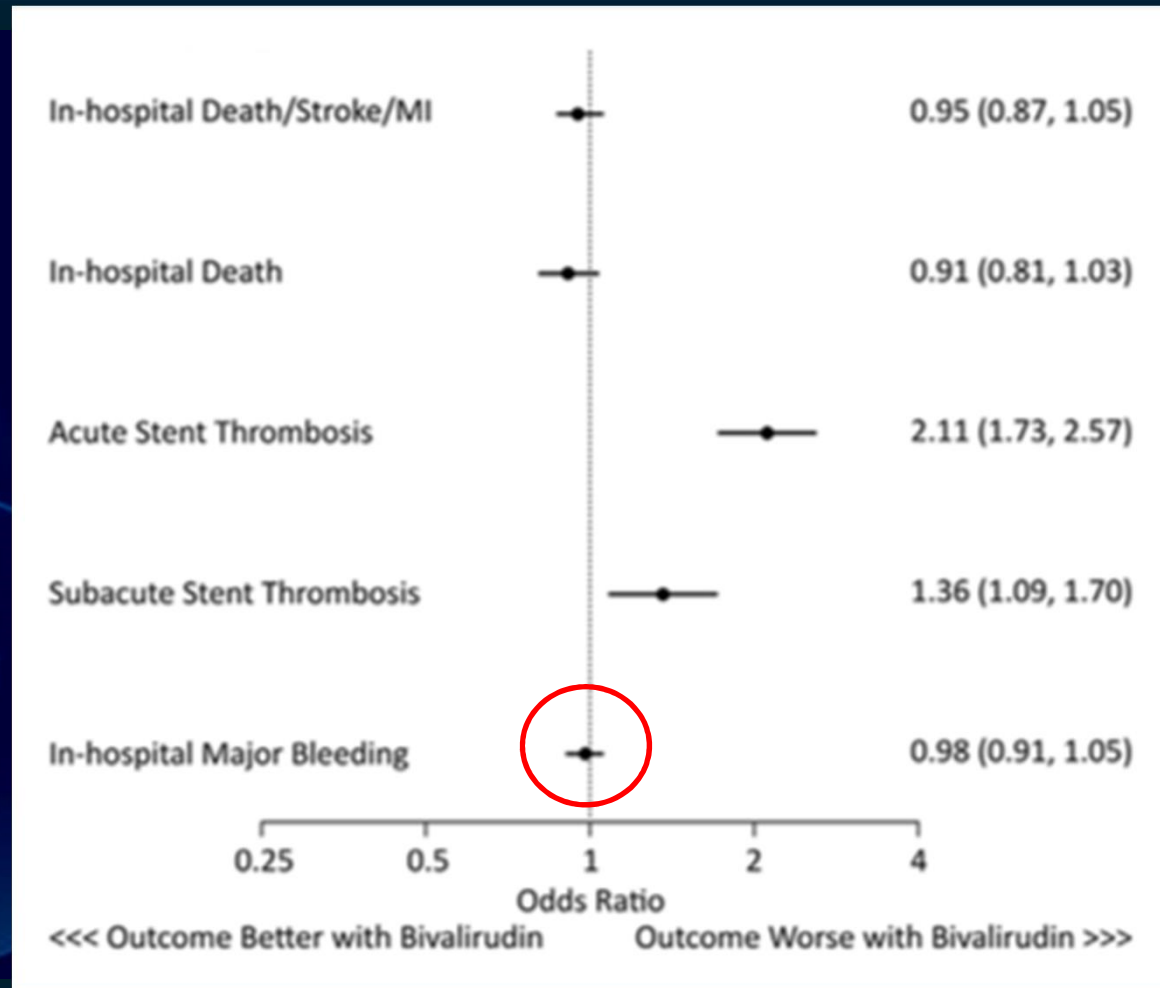
	BIVAL	HEPAR	HEPA+GPI			
bleeding	30 (4.1)	55 (7.5)	90 (12.3)	-3.5 (-5.9 to -1.1)	-8.2 (-11.0 to -5.5)	-4.8 (-7.8 to -1.8)
BARC 2-5	9 (1.2)	26 (3.6)	37 (5.1)	-2.3 (-3.9 to -0.8)	-3.8 (-5.6 to -2.1)	-1.5 (-3.6 to 0.6)
BARC 3-5	4 (0.5)	11 (1.5)	15 (2.1)	-1.0 (-2.0 to 0.1)	-1.5 (-2.7 to -0.4)	-0.5 (-0.2 to 0.8)

VALIDATE-SWEDEHEART

6006 Ptes STEMI - HNF 70 U/k vs Bivalirudina // 90% Radial
 PFP: Muerte, IAM, sangrado, ACV

Outcome	Bivalirudin Group (N=3004)	Heparin Group (N=3002)	Hazard Ratio (95% CI)	P Value
	<i>no. (%)</i>			
Primary end point: death from any cause, myocardial infarction, or major bleeding	216 (7.2)	241 (8.0)	0.89 (0.74–1.07)	0.21
Death from any cause	57 (1.9)	52 (1.7)	1.10 (0.75–1.60)	0.63
Death from cardiovascular causes	54 (1.8)	49 (1.6)	1.10 (0.75–1.62)	0.62
Myocardial infarction	23 (0.8)	33 (1.1)	0.70 (0.41–1.18)	0.18
Major bleeding	152 (5.1)	169 (5.6)	0.89 (0.72–1.11)	0.32
BARC type 2	100 (3.3)	114 (3.8)		
BARC type 3	53 (1.8)	53 (1.8)		
BARC type 5	4 (0.1)	3 (0.1)		
Stent thrombosis†	50 (1.7)	53 (1.8)	0.94 (0.64–1.39)	0.77
Definite	8 (0.3)	20 (0.7)	0.40 (0.18–0.91)	0.03
Probable	11 (0.4)	11 (0.4)		
Stroke	21 (0.7)	23 (0.8)	1.00 (0.52–1.92)	1.00
Primary hemorrhagic stroke	5 (0.2)	0		
Death, myocardial infarction, major bleeding, or stroke	226 (7.5)	253 (8.4)	0.89 (0.74–1.06)	0.19
Death, myocardial infarction, or BARC type 3 or 5 bleeding	124 (4.1)	131 (4.4)	0.94 (0.74–1.21)	0.64

Bivalirudina vs Heparina. STEMI via Radial



Jovin I, et al. JAmCollCardiolIntv2017;10:1102–11

www.icronline.com

Anticoagulant therapy		
Anticoagulation is recommended for all patients in addition to antiplatelet therapy during primary PCI.	I	C
Routine use of UFH is recommended.	I	C
In patients with heparin-induced thrombocytopenia, bivalirudin is recommended as the anticoagulant agent during primary PCI.	I	C
Routine use of enoxaparin i.v. should be considered. ^{200–202}	IIa	A
Routine use of bivalirudin should be considered. ^{209,215}	IIa	A
Fondaparinux is not recommended for primary PCI. ¹⁹⁹	III	B

Datos para una reperfusión efectiva

- 1- Tiempos
- 2- DAPT
- 3- Vía de Acceso
- 4- Anticoagulación – Antitrombosis
- 5- Tromboaspiración**
- 6- STENT: DES vs BMS – Directo vs Diferido
- 7- Solo ARI o MV?

– TROMBOASPIRACIÓN

- ✓ Estudios pequeños o de centros únicos demostraron cierto beneficio en la tromboaspiración manual
- ✓ 2 ECR con poder estadístico para superioridad no demostraron beneficios en la aspiración de rutina (TASTE y TOTAL)
 - ✓ TOTAL (10732 pts): Aumento de ACV en rama TBA
- ✓ En el subgrupo con alta carga de trombo, la TBA se asoció con:
 - ✓ Menor tasa de mortalidad (HR 0.80)
 - ✓ Mayor tasa de AIT y ACV (OR 1,56)

– TROMBOASPIRACIÓN

- ✓ TBA no es recomendada de forma rutinaria
- ✓ En caso de gran carga trombótica puede ser considerado

IRA technique		
Stenting is recommended (over balloon angioplasty) for primary PCI. ^{146,147}	I	A
Stenting with new-generation DES is recommended over BMS for primary PCI. ^{148–151,178,179}	I	A
Radial access is recommended over femoral access if performed by an experienced radial operator. ^{143–145,180}	I	A
Routine use of thrombus aspiration is not recommended. ^{157,159}	III	A
Routine use of deferred stenting is not recommended. ^{153–155}	III	B

2017 Guidelines STEMI. European Heart Journal (2018) 39, 119–177

Datos para una reperfusión efectiva

- 1- Tiempos
- 2- DAPT
- 3- Vía de Acceso
- 4- Anticoagulación – Antitrombosis
- 5- Tromboaspiracion
- 6- STENT: DES vs BMS – Directo vs Diferido**
- 7- Solo ARI o MV?

– STENT

- ✓ Implante de Stent es el procedimiento indicado en la PPCI
- ✓ BMS superior a Balon: menor reinfarto y TVR, sin alteraciones mortalidad
- ✓ DES 1º Generación superior a BMS: menor tasa de TVR
- ✓ DES 2º Generación superior a BMS: menor tasa de TVR, Trombosis stent mortalidad a 5 años
 - ✓ CONFORTABLE-AMI (Biolimus polimero bioabsorb vs BMS)
 - ✓ EXAMINATION (Everolimus vs BMS)
 - ✓ NORDIC (Biolimus vs BMS)
- ✓ DES segunda generación superior a DES 1ºG: menor tasa Trombosis y ReIAM

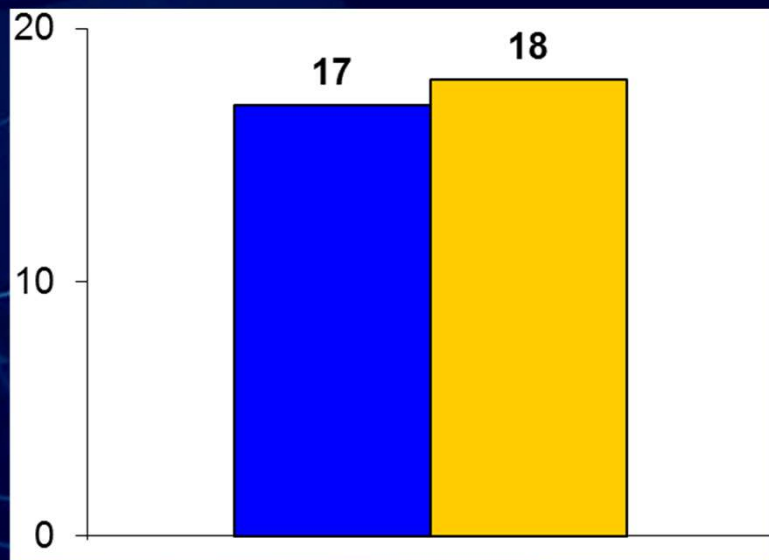
COMFORTABLE AMI randomized trial. JAMA 2012;308(8):777–787

EXAMINATION Trial. Lancet 2012;380(9852):1482–1490. www.icronline.com

NORSTENT . Investigators. Drug-eluting or bare-metal stents for coronary artery

– STENT DIFERIDO

✓ Estudio DANAMI 3 DEFER: Pts con IAM con flujo TIMI 2-3 espontáneo o conseguido por el operador, randomizado a implante inmediato o diferido a 48 hs. FUP 4 años



Primary endpoint

Deferred stenting
(n = 603)

Immediate PCI
(n = 612)

Resultados: Diferido vs Inmediato:

- MACE: 17% vs. 18%; $p = 0.92$
- Mortalidad: 7% vs. 9%, $p = 0.37$;
- MI: 7% vs. 7%, $p = 0.77$;
- TVR: 7% vs. 4%, $p = 0.03$

Kelbæk H, et al. Lancet 2016;387:2199-2206

– STENT



IRA technique		
Stenting is recommended (over balloon angioplasty) for primary PCI. ^{146,147}	I	A
Stenting with new-generation DES is recommended over BMS for primary PCI. ^{148–151,178,179}	I	A
Radial access is recommended over femoral access if performed by an experienced radial operator. ^{143–145,180}	I	A
Routine use of thrombus aspiration is not recommended. ^{157,159}	III	A
Routine use of deferred stenting is not recommended. ^{153–155}	III	B

2017 Guidelines STEMI. European Heart Journal (2018) 39, 119–177

Datos para una reperfusión efectiva

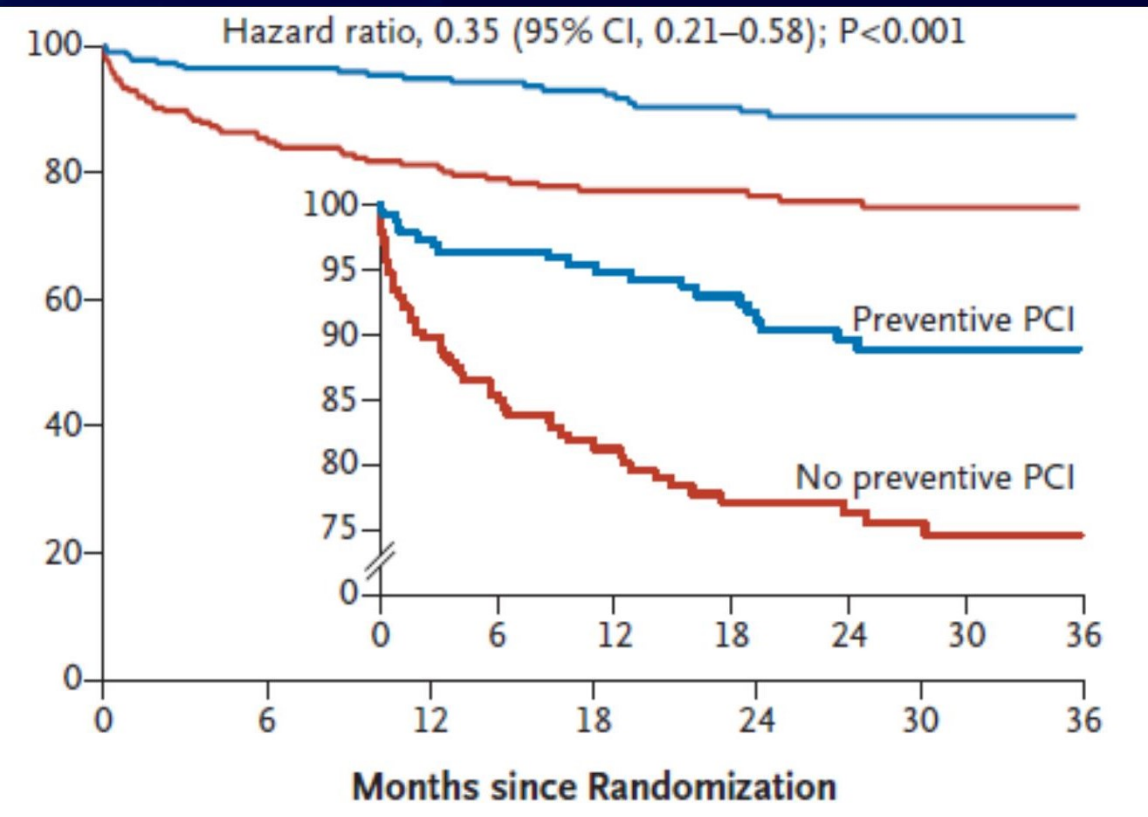
- 1- Tiempos
- 2- DAPT
- 3- Vía de Acceso
- 4- Anticoagulación – Antitrombosis
- 5- Tromboaspiracion
- 6- STENT: DES vs BMS – Directo vs Diferido
- 7- Solo ARI o MV?**

– ARI vs REVASC COMPLETA

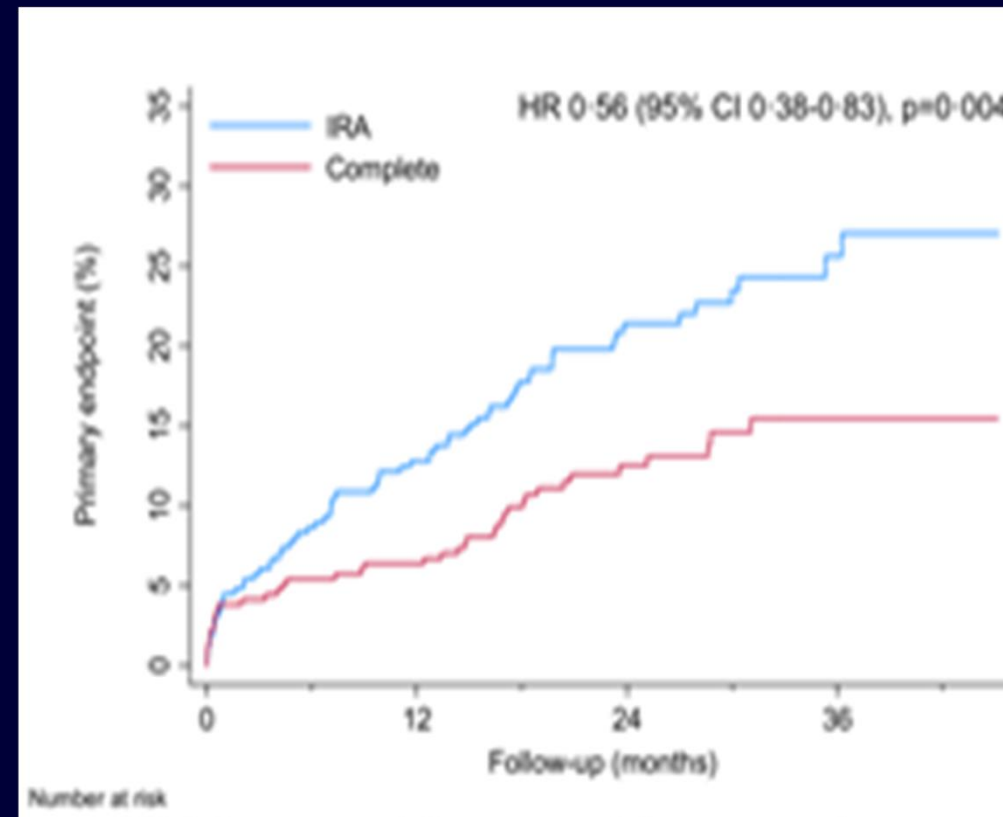
- ✓ Aprox el 50% de los pts con IAM tienen enfermedad de multiples vasos
- ✓ Siempre tratar la ARI
- ✓ La revascularización completa es recomendable
 - ✓ No durante el procedimiento índice
 - ✓ Antes del alta, inmediata o diferida
 - ✓ Guiada por Angio o isquemia
- ✓ PRAMI (465 pts)
- ✓ CvLPRIT (296 pts)
- ✓ DANAMI 3 PRIMULTI (627 pts) - FFR
- ✓ COMPARE Acute (885 pts) - FFR

- ARI vs REVASC COMPLETA

PRAMI



DANAMI 3 PRIMULTI



– ARI vs REVASC COMPLETA

Non-IRA strategy		
Routine revascularization of non-IRA lesions should be considered in STEMI patients with multivessel disease before hospital discharge. ^{167–173}	IIa	A
Non-IRA PCI during the index procedure should be considered in patients with cardiogenic shock.	IIa	C
CABG should be considered in patients with ongoing ischaemia and large areas of jeopardized myocardium if PCI of the IRA cannot be performed.	IIa	C

2017 Guidelines STEMI. European Heart Journal (2018) 39, 119–177

CONCLUSIONES

Factores para una reperfusión efectiva

- 1- Tiempos
- 2- DAPT
- 3- Vía de Acceso
- 4- Anticoagulación – Antitrombosis
- 5- Tromboaspiración
- 6- STENT: DES vs BMS – Directo vs Diferido
- 7- Solo ARI o MV?

CONCLUSIONES



R

www.icronline.com

Muchas gracias