

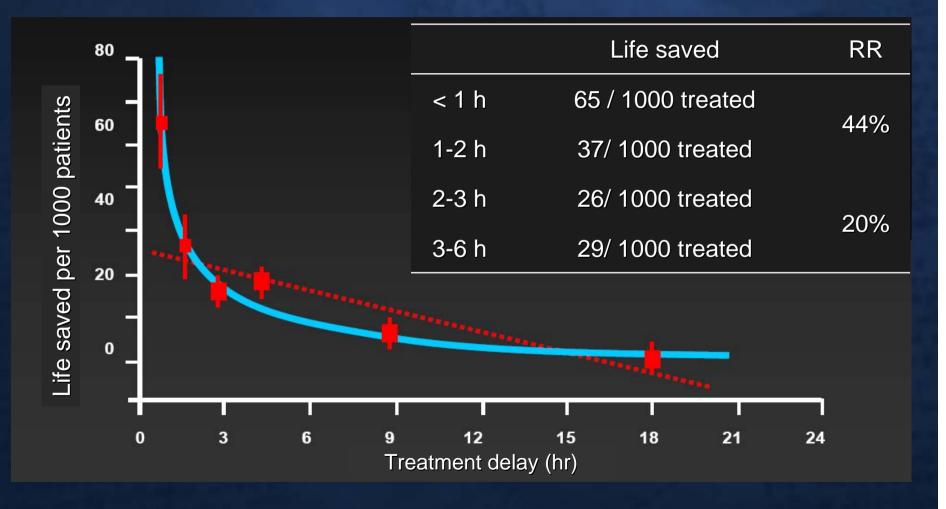


Time to Reperfusion in Acute Myocardial Infarction: Review on Controversies and the Result of Contrast MRI Study

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Life Saved by Fibrinolytic Therapy is a Function of Door-to-Needle Time

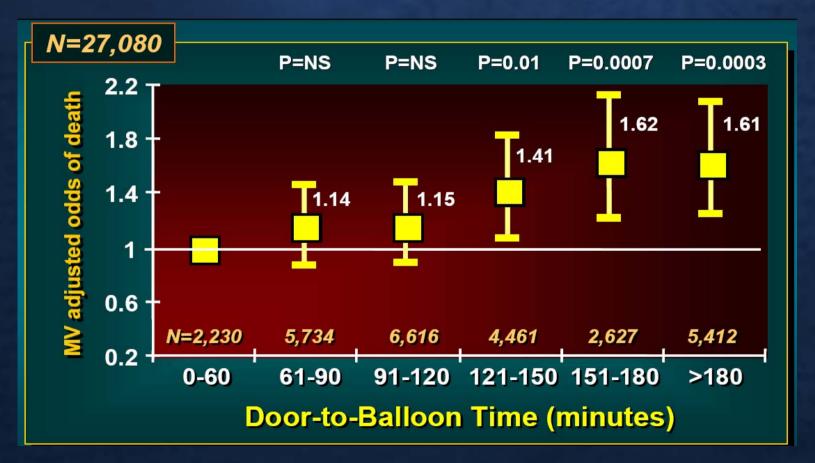




Door-to-Balloon Time and Mortality

AHA/ACC guideline 2004

Class I: Door-to-balloon time: \leq 90 minutes (Evidence B)



Cannon CP et al. NRMI-2 JAMA 2000;283:2941-7





Initial time Variable and Mortality in Primary PCI

Dependent on symptom-to-balloon time for all patients ✤ De Luca G, JACC 2003 Brodie BR, AHJ 2006 Time dependent only for high-risk patients ℜ Antoniucci D, AJC 2002 Relatively time independent after 2 h of symptom Time dependent with symptom-to-balloon time but NOT for door-toballoon time ℜ Cannon CP, JAMA 2000 McNamara RL, JACC 2006 ✓ No association between mortality and door or symptom-to-balloon time Zijlstra F, EHJ 2002





Symptom-to-Balloon Time and 30-Day Mortality Relatively time independent after 2 h of symptom

30 Day plus Late Cardiac Mortality 18 16 16.9 15.616.4 14 Mortality % 12 p = .0230 Day Mortality 10 • 9.5 О. 0.3 8 9.0 6 p = .04 4 4.3 $\mathbf{2}$ 0 <2 2-44-6 ≥ 6 Time to Reperfusion (hrs)

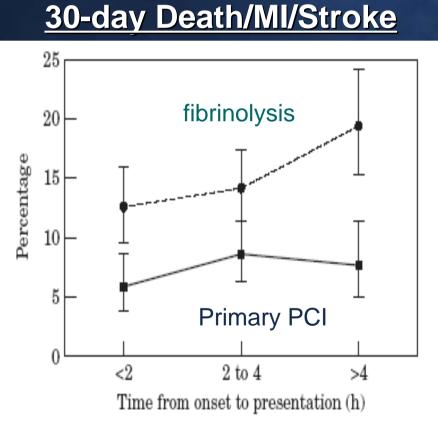
Brodie BR, JACC 1998



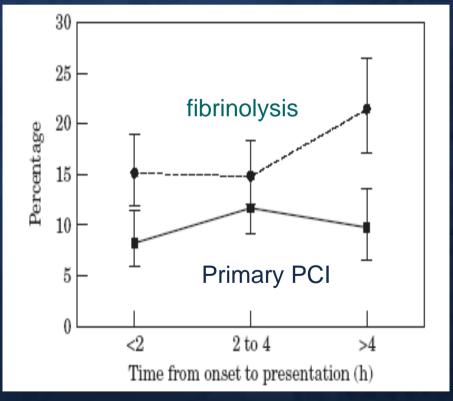


Symptom-to-Balloon Time

Time dependent for fibrinolysis but not for primary PCI



6-month Death/MI/Stroke



Zijlstra F, Eur Heart J 2002

Impact of Initial Delay of Primary PCI Korean AMI Registry (KAMIR)

- 1416 patients who met inclusion criteria, out of 5069 patients in KAMIR (2005. 11 – 2007.1)
- Inclusion criteria
 - Primary PCI within 12 hours of symptom onset in STEMI
 - Available 1-month follow-up data (92.5% FU rate)



Impact of Initial Delay of Primary PCI Korean AMI Registry (KAMIR)

- Symptom onset-to-door time
 - median time: 163 min [90 285]
 - ≤ 120 min : 36 %
- Door-to-balloon time
 - median time: **90 min** [65 136]
 - ≤ 90 min : 51 %
- Symptom onset-to-balloon time
 - median time : 274 min [185 442]
 - ≤ 240 min : 42 %
- 1-month mortality
 - 62 death / 1416 patients = 4.4%

106 min in NRMI-4

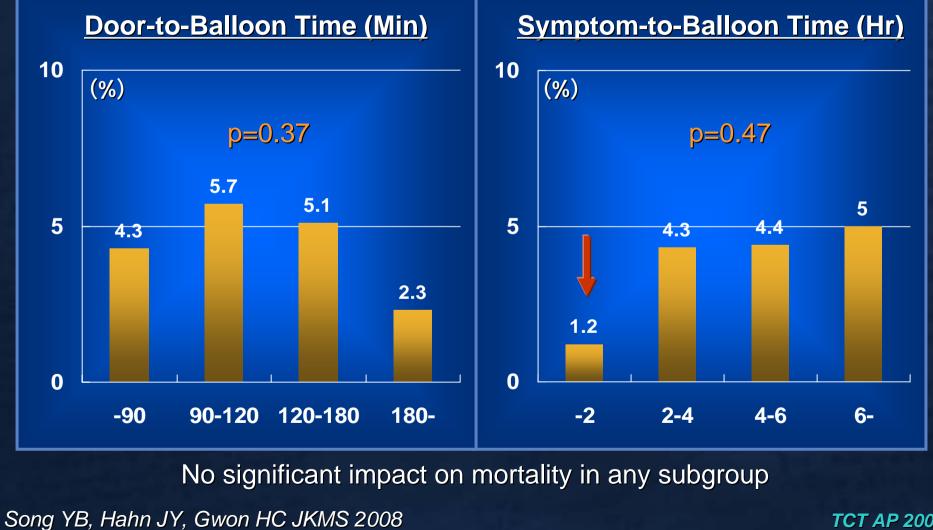
4.5% in NRMI-4

Song YB, Hahn JY, Gwon HC JKMS 2008



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Impact of Initial Delay of Primary PCI on 30-day Mortality in KAMIR



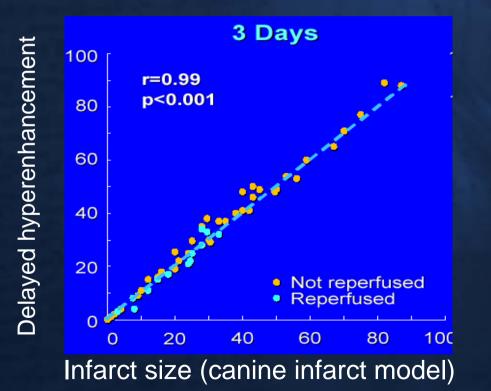


TTC

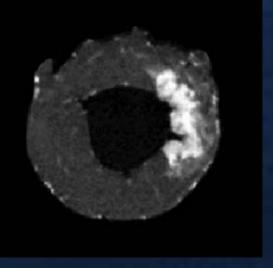
CE-MRI

Infarct Size Measurement by MRI

Delayed hyperenhancement (DHE) in contrast-enhanced magnetic resonance imaging (CE-MRI)



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Kim et al. Circulation 1999



Ischemic Duration Was Associated With Transmural Necrosis After Primary PCI

Table 3. Ischemic Time and Enzymatic Data According to the CE-MR Evidence of TN and/or SMO in Patients Without TIMI Flow Grade 3 of Infarct-Related Artery at Index Angiography

	TN-/SMO-	TN+/SMO-	TN+/SMO+
Patients, n (%)	19 (29.7)	22 (34.4)	23 (35.9)
Time to treatment, min*	90 ± 40	177 ± 101	255 ± 145
Peak of troponin I, µg/l†	53 ± 50	110 ± 107	137 ± 97

Limitation of the study

- No data on infarct size
- Semi-quantitative measurement of infarct transmurality
- Small sample size

Tarantini G et al. J Am Coll Cardiol 2005;46:1229 –35





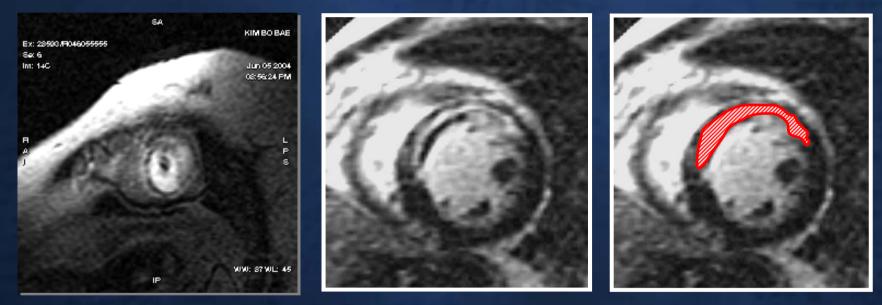
The Aim of the Study

 To evaluate the impact of time to reperfusion on myocardial infarct size & infarct transmurality in patients undergoing primary PCI for STEMI within 12 hours of symptom onset.

 CE-MRI: Quantitative measurements of infarct transmurality as well as infarct size



Infarct Size Measurement by MRI = Volume% of delayed Hyper-enhancement



n in Area measurement

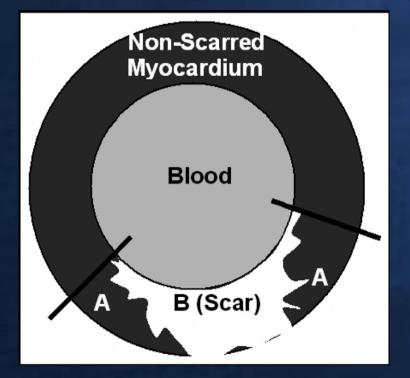
Delayed myocardial imaging 5-15 minutes after injection in short-axial planes with IR-prepared fast GRE seq. (FOV, 40 cm; slice thickness, 6 mm; TI, 200-300 ms)

> Sum of each area × Slice thickness = Infarct volume Infarct size = infarct volume / myocardial volume x 100 (%)

Hahn JY, Gwon HC. Am Heart J 2007;153:665.e1-8



Quantitative Measurement of Infarct Transmurality



Transmurality in each slice

= B / (A+B)

Infarct transmurality

= the average transmurality of all segments with infarction.





Baseline characteristics

- 77 patients enrolled
- Male: 66 (86%)
- Age: 56 years (49-70)
- Symptom-to-balloon time: 259 min (162-370)
- Door-to-balloon time: 100 min (70-125)
- MRI: within 1 month after primary PCI



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Baseline Characteristics Grouped by Symptom-to-Balloon Time

Symptom-to-balloon time, min	< 180 (n=23)	180 ~ 360 (n=34)	> 360 (n=20)	P value
Age (years)	54±9	57±11	57±12	0.45
Male	21 (91%)	29 (85%)	16 (80%)	0.57
Diabetes	5 (22%)	9 (27%)	3 (15%)	0.62
Hypertension	8 (35%)	10 (29%)	10 (50%)	0.31
Dyslipidemia	10 (44%)	11 (32%)	6 (30%)	0.59
Current smoker	19 (83%)	20 (59%)	8 (40%)	0.02
Anterior infarction	14 (61%)	17 (50%)	15 (75%)	0.19



Baseline Characteristics Grouped by Symptom-to-Balloon Time

Symptom-to-balloon time, min	< 180 (n=23)	180 ~ 360 (n=34)	> 360 (n=20)	P value
LAD / LCx / RCA	14 / 2 / 7	16 / 4 / 14	15/0/5	0.28
Multivessel	10 (44%)	15 (44%)	8 (40%)	0.96
Baseline TIMI 0/1	17 (74%)	25 (74%)	17 (85%)	0.59
Final TIMI 3 flow	22 (96%)	33 (97%)	18 (90%)	0.52
Collateral flow	1 (4%)	4 (12%)	4 (20%)	0.28
Myocardial blush 0/1	5 (22%)	15 (44%)	5 (25%)	0.15
No reflow	2 (9%)	1 (3%)	2 (10%)	0.52
GP IIb/IIIa inhibitor	4 (17%)	7 (21%)	11 (55%)	0.009
Distal protection	8 (35%)	7 (21%)	4 (20%)	0.41

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Baseline Characteristics Grouped by Door-to-Balloon Time

Door-to-balloon time, min	< 90 (n=30)	90 ~ 120 (n=27)	> 120 (n=20)	<i>P</i> value
Age (years)	59±10	54±10	55±11	0.27
Male	25 (83%)	26 (96%)	15 (75%)	0.11
Diabetes	5 (17%)	7 (26%)	5 (25%)	0.66
Hypertension	13 (43%)	10 (37%)	5 (25%)	0.42
Dyslipidemia	12 (40%)	8 (30%)	7 (35%)	0.72
Current smoker	15 (50%)	21 (78%)	11 (55%)	0.08
Anterior infarction	18 (60%)	16 (59%)	12 (60%)	0.99



Baseline Characteristics Grouped by Door-to-Balloon Time

Door-to-balloon time, min	< 90 (n=30)	90 ~ 120 (n=27)	> 120 (n=20)	P value
LAD / LCx / RCA	17 / 2 / 11	16 / 3 / 8	12 / 1 / 7	0.93
Multivessel	13 (43%)	12 (44%)	8 (40%)	0.95
Baseline TIMI 0/1	21 (70%)	22 (82%)	16 (80%)	0.54
Final TIMI 3 flow	30 (100%)	25 (93%)	18 (90%)	0.24
Collateral flow	3 (10%)	2 (7%)	4 (20%)	0.39
Myocardial blush 0/1	9 (30%)	9 (33%)	7 (65%)	0.93
No reflow	1 (3%)	3 (11%)	1 (5%)	0.47
GP IIb/IIIa inhibitor	5 (17%)	11 (41%)	6 (30%)	0.13
Distal protection	7 (23%)	5 (19%)	7 (35%)	0.42

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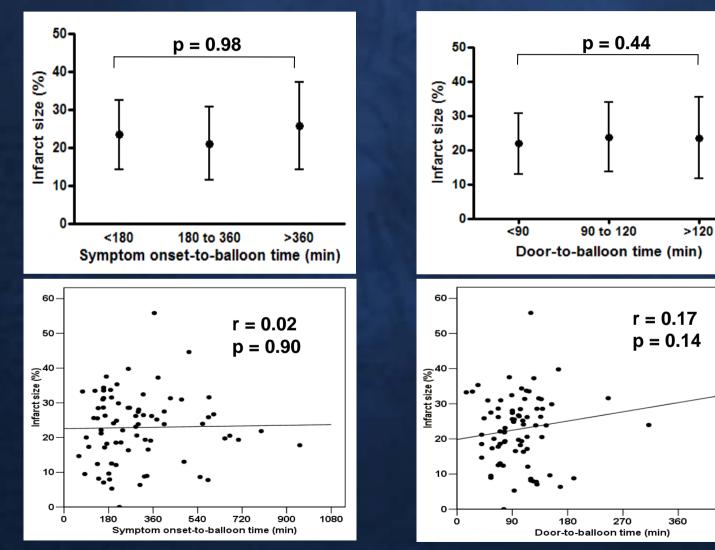
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Infarct Size

Symptom-to-Balloon Time

Door-to-Balloon Time



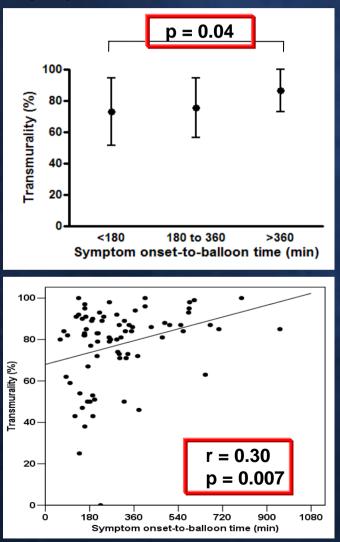
TCT AP 2008

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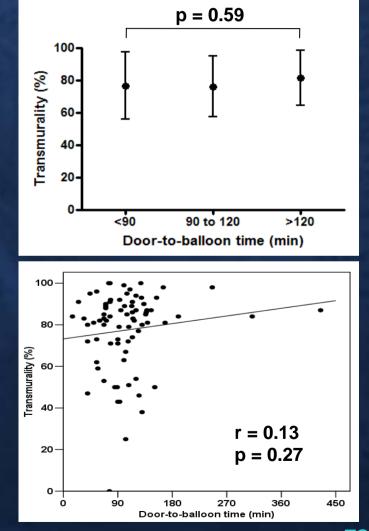


Infarct Transmurality

Symptom-to-Balloon Time



Door-to-Balloon Time





Independent Predictors of Infarct Size and Transmurality

	Odds ratio	95% CI	<i>P</i> value
Infarct size > 25%			
Anterior infarction	4.51	1.38 – 14.72	0.013
Transmurality > 75%			
Use of GP IIb/IIIa inhibitors	0.06	0.01 – 0.41	0.004
Symptom-to-balloon time (per 30 min)	1.34	1.08 – 1.66	0.008





Why the Outcome of Primary PCI is Less Time-dependent?

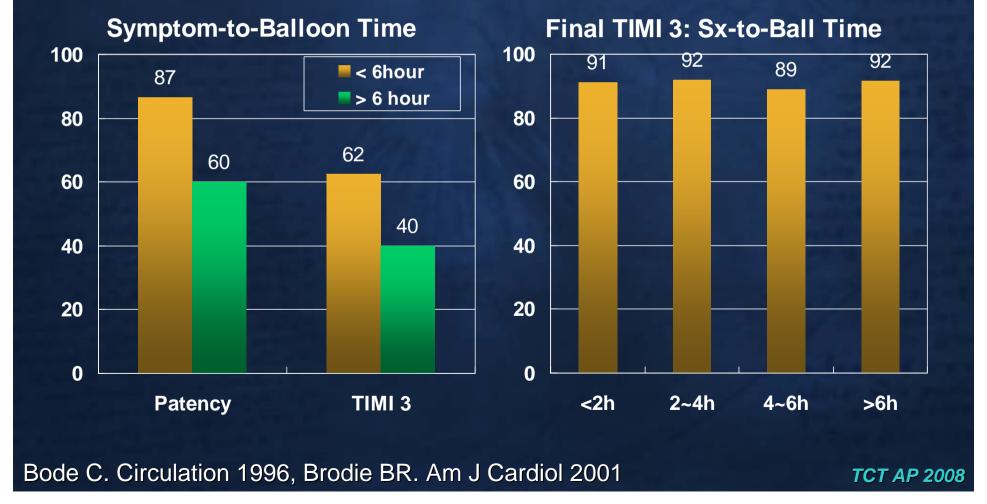
Successful reperfusion is more time-dependent after fibrinolysis than after primary PCI.
High TIMI 3 flow was achieved after primary PCI even in the patients with late presentation.



Successful Reperfusion is Time-dependent after Fibrinolysis, but not after Primary PCI

<u>Fibrinolysis</u>

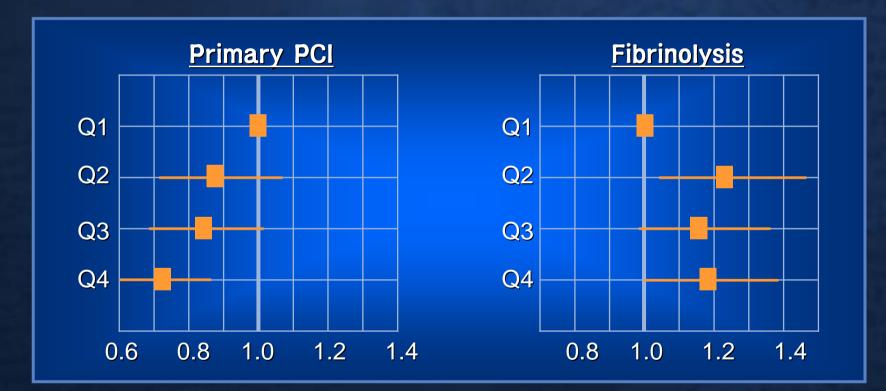
Primary PCI





PCI volume and Mortality

Relative risk of 30-day mortality



NRMI, NEJM 2000

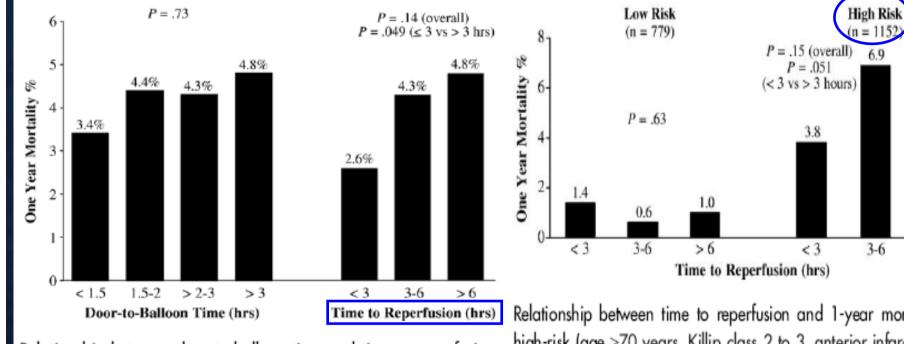


Summary and Conclusion

- Clinical outcome and infarct size is less dependent on the time-to-reperfusion in primary PCI, may reach a plateau in less than 3 hours of symptom onset.
- The effort to reduce the time delay is still valuable both in the hospital and outside of the hospital.



Symptom onset-to-Balloon Time & Mortality



Relationship between door-to-balloon time and time to reperfusion, and 1-year mortality.

Relationship between time to reperfusion and 1-year mortality in high-risk (age >70 years, Killip class 2 to 3, anterior infarction, or renal insufficiency) and low-risk patients.

Brodie BR et al. *Am Heart J* 2006;151:1231-8

TCT AP 2008

7.0

>6