

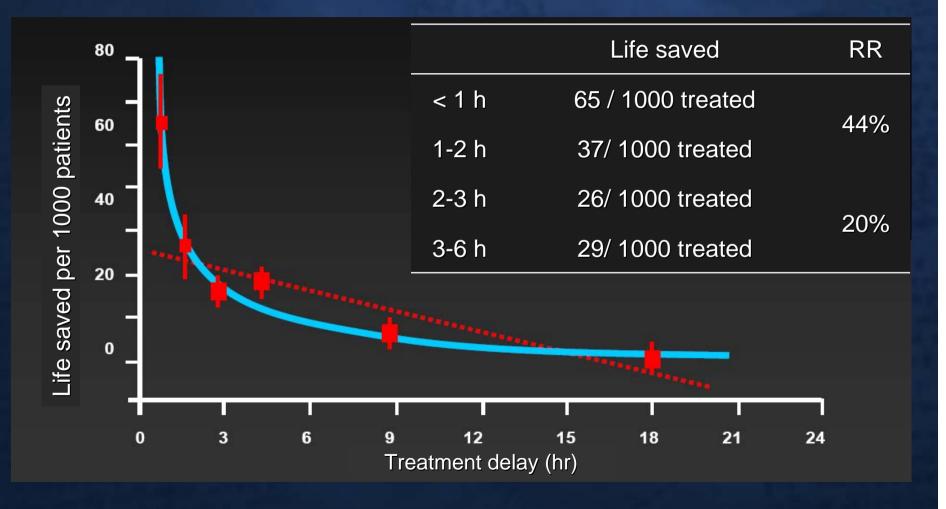


Time to Reperfusion in Acute Myocardial Infarction: Review on Controversies and the Result of Contrast MRI Study

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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)

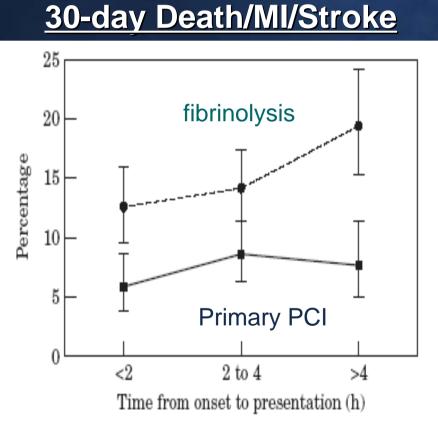
Brodie BR, JACC 1998



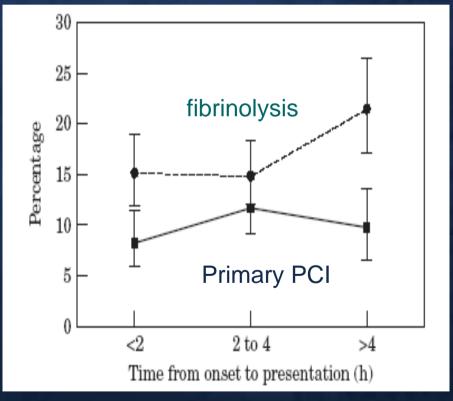


Symptom-to-Balloon Time

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

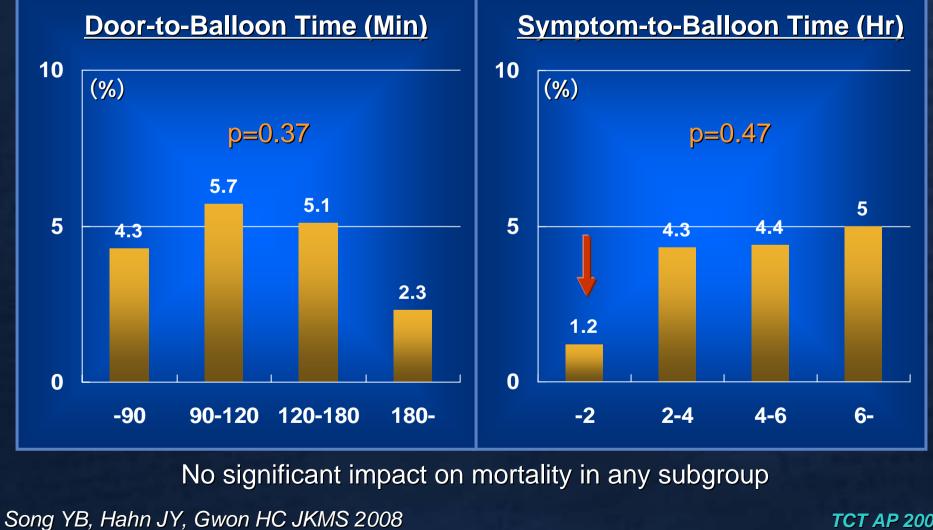
4.5% in NRMI-4

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



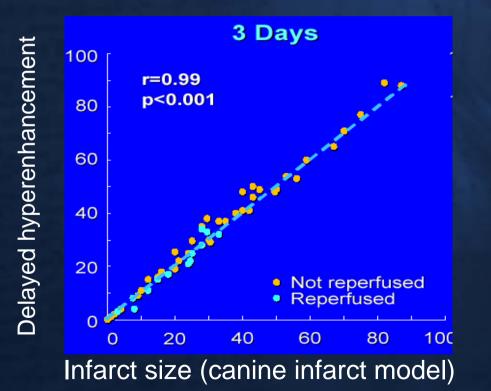


TTC

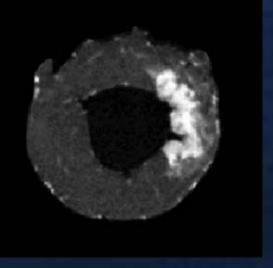
CE-MRI

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





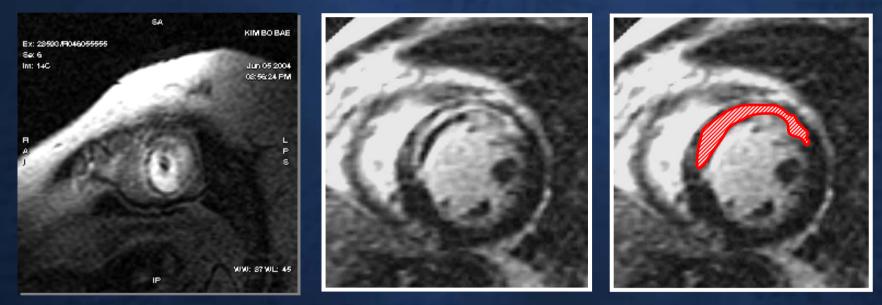
The Aim of the Study

 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

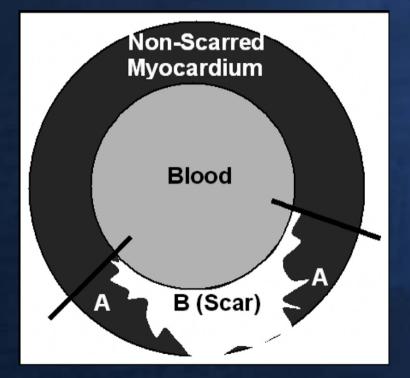
Delayed myocardial imaging 5-15 minutes after injection in short-axial planes with IR-prepared fast GRE seq. (FOV, 40 cm; slice thickness, 6 mm; TI, 200-300 ms)

> 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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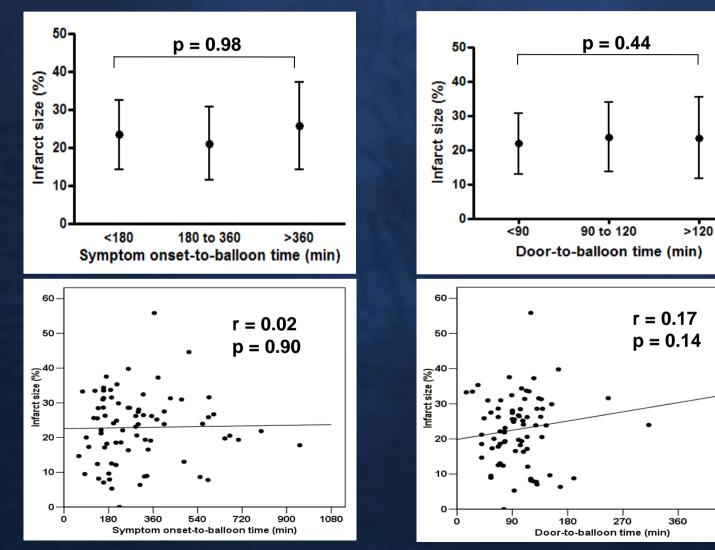
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Infarct Size

Symptom-to-Balloon Time

Door-to-Balloon Time



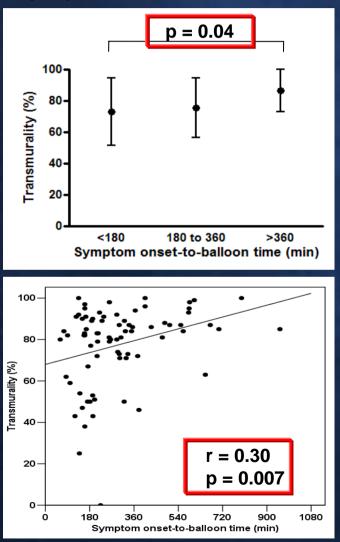
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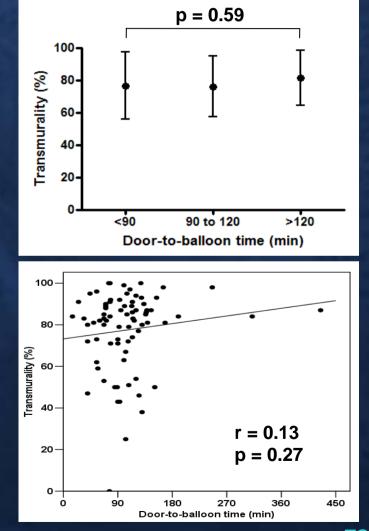


Infarct Transmurality

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?

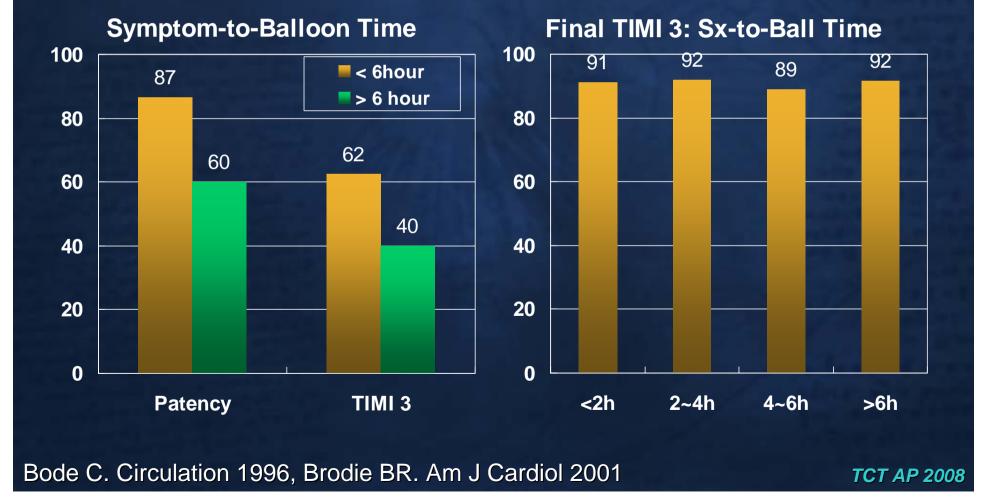
Successful reperfusion is more time-dependent after fibrinolysis than after primary PCI.
High TIMI 3 flow was achieved after primary PCI even in the patients with late presentation.



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

<u>Fibrinolysis</u>

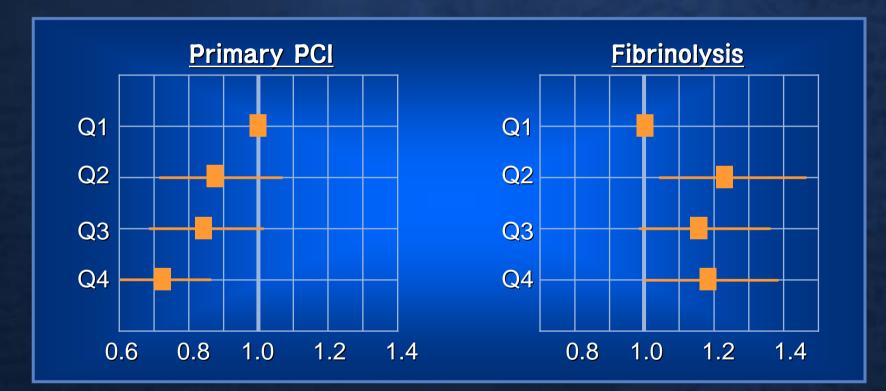
Primary PCI





PCI volume and Mortality

Relative risk of 30-day mortality



NRMI, NEJM 2000

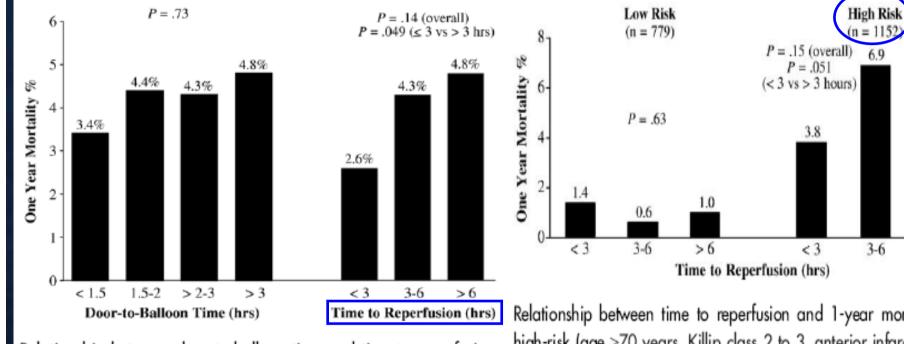


Summary and Conclusion

- 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

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