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# Deferred Stenting After MI

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## ACS encompasses a spectrum



Unstable angina

NSTEMI

STEMI

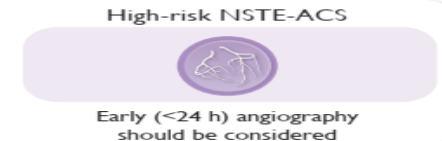
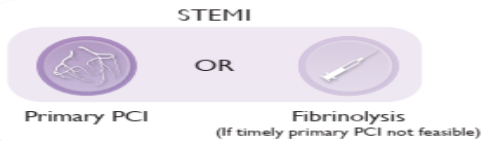
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### Think 'A.C.S.' at initial assessment



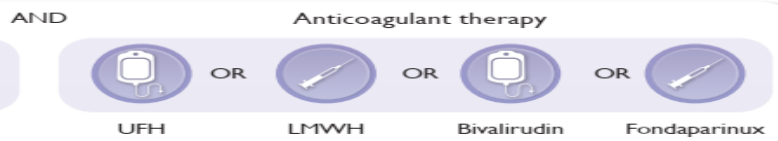
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### Think invasive management



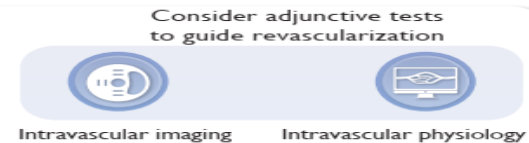
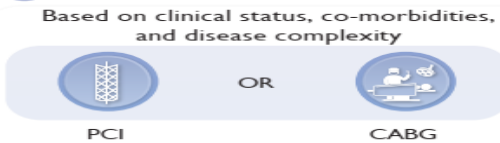
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### Think antithrombotic therapy



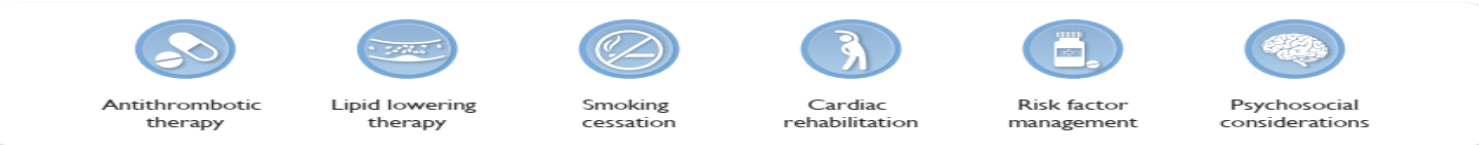
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### Think revascularization



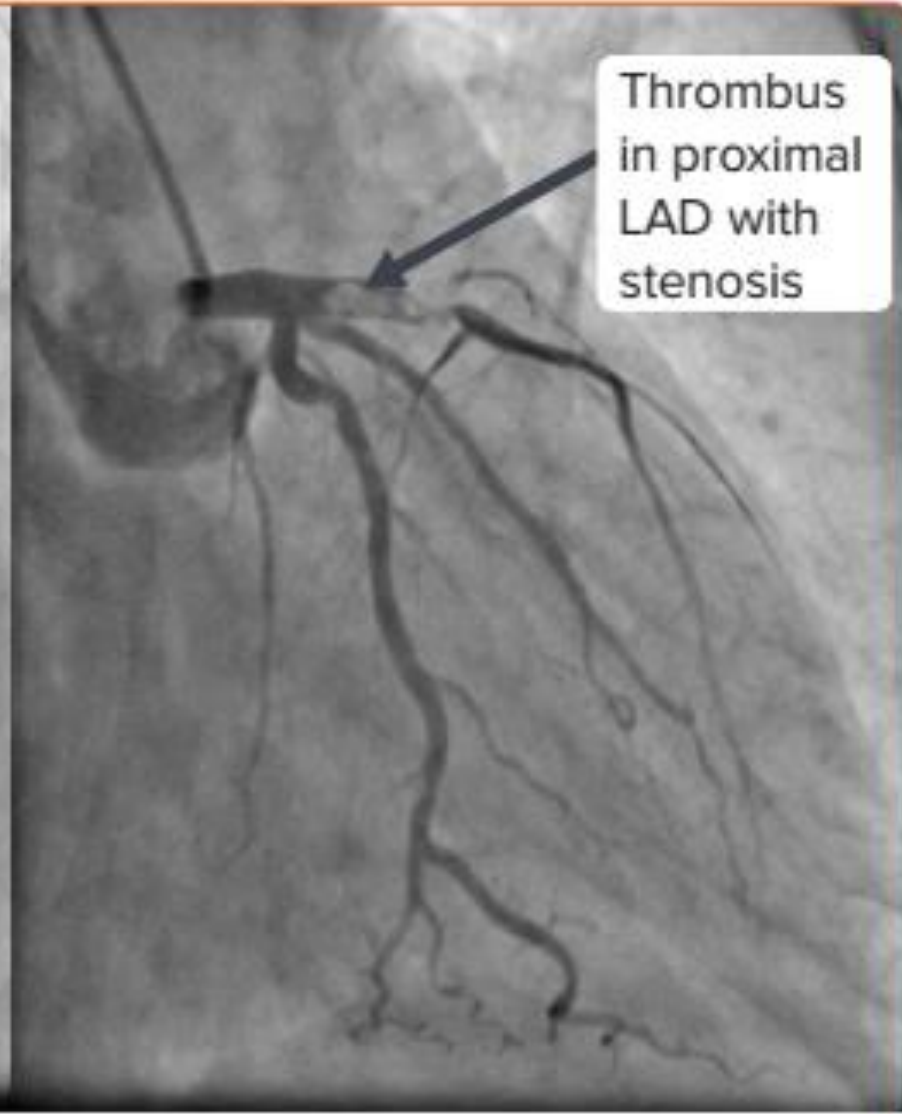
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### Think secondary prevention



# Case Vignette

- A 45-year-old man presented to ED
- Acute anterior-wall MI of 8 hours duration
- HTN & C/S
- CAG showed a significant stenosis with grade IV thrombus in proximal LAD with TIMI 2 flow





- Due to high thrombus burden, stenting was deferred and the patient was put on IV eptifibatide infusion for 18 hours followed by subcutaneous LMWH twice daily.
- Rescue PCI was planned in case the patient developed chest pain.
- After 5 days, his angiogram showed moderate stenosis and the thrombus in LAD was almost absent with improved TIMI 3 flow.





B



# Introduction and Definition

- Deferred stenting is a strategy that aims to postpone stent placement for a fixed time window after stable distal flow has been achieved.
- Other terminology used in the literature for a procrastinated stenting approach include **delayed** PCI and **secondary** PCI.

# Continue...

- In a fraction of patients reduced coronary blood flow (slow flow or no reflow) is seen despite epicardial vessel patency with PCI, and this is associated with a worse prognosis.
- Distal migration of thrombus and atherosclerotic debris are important contributors to interventional slow flow/no reflow.



# Continue...

- Presence of residual thrombus even after manual aspiration is one of the pitfalls and it predicts poor outcomes.
- Thrombus grading on angiography is done by the Gibson's angiographic score/TIMI criteria.
- A thrombus grade higher than 3 is usually considered as high thrombus burden.

## Table 1: Thrombolysis in MI Grading for Thrombus Burden

Grade	Characteristics
0	No angiographic evidence of thrombus
1	Possible thrombus: reduced contrast density or haziness, irregular lesion contour, a smooth convex meniscus at the site of a total occlusion suggestive but not diagnostic of thrombus
2	Definite thrombus, with greatest dimensions $\leq 1/2$ the vessel diameter
3	Definite thrombus, with greatest linear dimension $> 1/2$ but $< 2$ vessel diameters
4	Definite thrombus, with the largest dimension $\geq 2$ vessel diameters
5	Total thrombotic occlusion

Source: Gibson et al. 2001.<sup>6</sup>

# Continue...

- This time of deferment has multiple benefits:
  - Gradual clearing of the thrombus,
  - Improvement of microvascular flow,
  - Reduction of vasospasm
  - Prevention of distal embolisation,
  - Avoidance of slow flow/no reflow
  - Attenuated periprocedural MIs.
- Indeed, data suggests the coexistence of thrombus and spasm and hence a deferred strategy can lead to better stent selection (large and short stents).

# Approach

- There is a possible risk of reocclusion during the waiting period which can be mitigated by parenteral anticoagulants and GP IIb/IIIa inhibitors.
- A rescue PCI should be considered if necessary.
- A prolonged systemic anticoagulation can increase the risk of bleeding which can be detrimental.
- However, the use of the CRUSADE bleeding risk score can help to assess the baseline bleeding risk of the patient.

Predictor	Score
<b>Baseline haematocrit, %</b> <31 31–33.9 34–36.9 37–39.9 ≥40	9 7 3 2 0
<b>Creatinine clearance,<sup>a</sup> mL/min</b> ≤15 >15–30 >30–60 >60–90 >90–120 >120	39 35 28 17 7 0
<b>Heart rate (b.p.m.)</b> ≤70 71–80 81–90 91–100 101–110 111–120 ≥121	0 1 3 6 8 10 11
<b>Sex</b> Male Female	0 8
<b>Signs of CHF at presentation</b> No Yes	0 7
<b>Prior vascular disease<sup>b</sup></b> No Yes	0 6
<b>Diabetes mellitus</b> No Yes	0 6
<b>Systolic blood pressure, mmHg</b> ≤90 91–100 101–120 121–180 181–200 ≥201	10 8 5 1 3 5

Used with permission of *Circulation* 2009.

CRUSADE = Can Rapid risk stratification of Unstable angina patients Suppress ADverse outcomes with Early implementation of the ACC/AHA guidelines

# Continue...

- Scores  $<20$  indicate a very low risk of in-hospital bleeding.
- Score  $>20$  the use of GP IIb/IIIa inhibitor infusion should be avoided.
- Score  $>50$  are at a very high risk of bleeding and the duration of LMWH duration should be reduced to 72 hours instead of the standard proposed duration of 5–7 days.



# Continue...

- Minimally invasive mechanical intervention (MIMI) is an adjuvant technique during primary PCI before deferring stent placement in arteries with TIMI 0–1 flow.
- The strategy entails the use of a guidewire, an undersized balloon catheter and thrombus aspiration to establish distal coronary flow.
- The aim is to restore the flow with minimal forward propagation of thrombus.

# Review of Literatures...

- Predictors of greater benefit from deferred stenting
  - Male sex,
  - younger age,
  - larger size of culprit artery
  - higher thrombus burden at baseline
- In a Danish pilot study, the need for subsequent stenting was reduced by 38% without any risk of reocclusion at 3 months with a deferred strategy.
- In Ke et al., subsequent stents were avoided in 23% of patients.

# Continue...

- In a French study, Souteyrand et al. used OCT to guide deferred stenting.
- The study tested the safety of three different strategies – acute (<2 days), early (up to 7 days) and late deferral (up to 1 month) in the setting of STEMI with large thrombus burden on angiogram.
- There were no MACE recorded between initial and final procedure.
- The thrombus presence as assessed by OCT continued to diminish from acute phase (94.1%) to early phase (78%) to late phase (32%).
- This study demonstrated that OCT-guided postponement of stent implantation led to good procedural outcomes with 100% success and alleviation of no-reflow events.

# Continue...

- The SUPER-MIMI study tested a longer deferral time of 7 days in 155 patients with STEMI.
  - There was an improvement in TIMI flow,
  - Decrease in thrombus burden
  - Stenosis severity diminished.
- More importantly, stenting was also avoided in 38% cases with a minimal chance of reocclusion (1.3%).

# Continue...

- In the DEFER-STEMI study, patients were randomized to either conventional stenting or deferred stenting.
- Patients with STEMI along with angiographic or clinical features for risk of slow flow/no reflow were enrolled for the study.
- There was significant reduction in incidence of the primary endpoints in the deferred group.
  - Slow flow/no reflow
  - There were also fewer thrombotic events
  - Final TIMI flow was higher.
  - higher salvaged myocardium on CMR

# Continue...

- On the contrary, the DANAMI 3-DEFER trial **failed** to show any benefit of deferred stenting on **clinical outcomes**.
- About 1,200 patients were randomised to a deferred stent strategy versus an immediate stenting technique.
- There was no significant difference in the primary outcome
- In addition, there was a slightly higher, although not significant, chance of reocclusion rates (2%) in the deferred stenting group.
- However, there was an insignificant improvement in LVEF in the deferred stenting group.
- An MRI sub-study also failed to find any benefit on myocardial infarct size, microvascular obstruction and myocardial salvage index.
- **However, in patients with lesion length/stent >24 mm, the deferred strategy significantly reduced infarct size.**



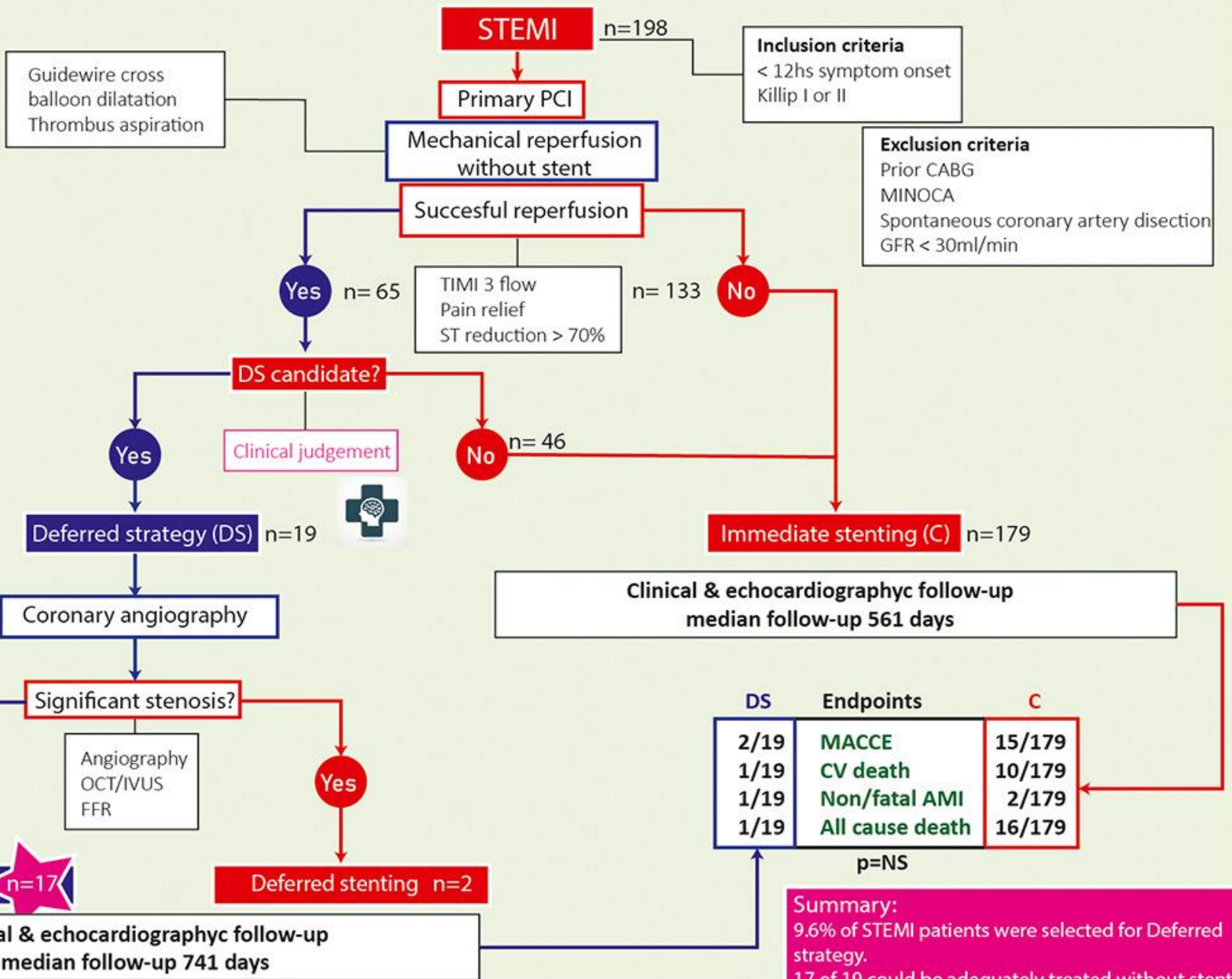
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- Why are there contrasting results from two large RCTs on deferred PCI?
  - First, the DEFER STEMI enrolled patients at high risk of slow flow based on clinical angiographic features, whereas DANAMI 3-DEFER was an all-comer primary PCI study. A deferral strategy should only be applied after careful angiographic selection.
  - Second, DEFER STEMI was an angiographic and MRI endpoint study whereas DANAMI 3-DEFER looked at clinical outcomes. We know that clinical outcomes are affected by many variables and imaging features are only one of the facets.
  - Third, DANAMI 3-DEFER was a larger, multicentre, randomised study, in contrast to DEFER, which was a small, single-centre, proof of concept study.
  - Fourth, the use of GPIIb/IIIa inhibitors in DANAMI 3-DEFER was significantly lower compared to DEFER STEMI.

# Continue...

- The INNOVATION study did not find any merit in a routine defer strategy during primary PCI at two centres in South Korea.
- In the subset of anterior infarction, the primary endpoint – infarct size and microvascular obstruction – was significantly attenuated.

**Graphical abstract**



DS	Endpoints	C
2/19	MACCE	15/179
1/19	CV death	10/179
1/19	Non/fatal AMI	2/179
1/19	All cause death	16/179

p=NS

**Summary:**  
 9.6% of STEMI patients were selected for Deferred strategy.  
 17 of 19 could be adequately treated without stent implantation.

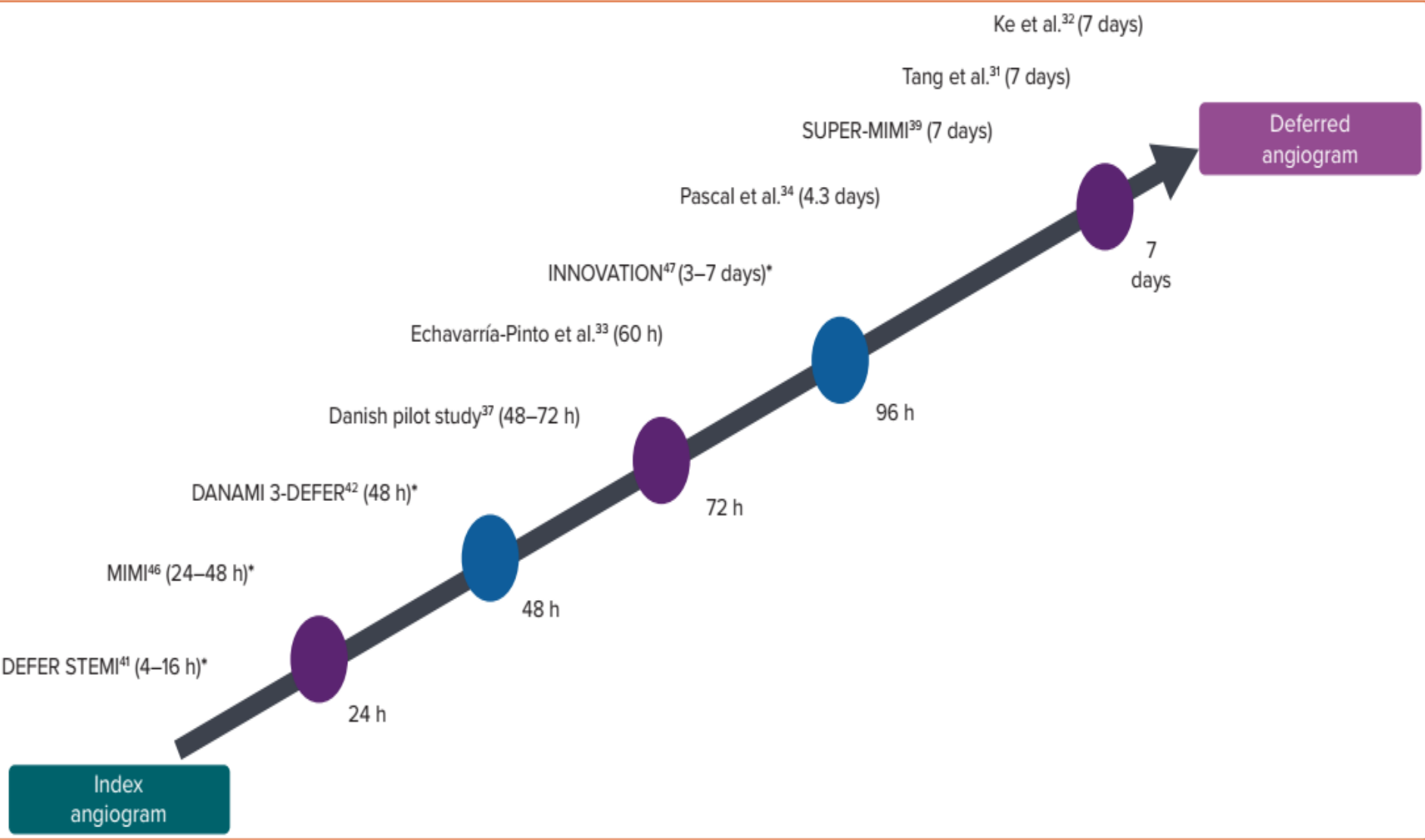
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- The DS group showed:
  - Significantly lower rate of stent implantation and a
  - Higher use of thrombus aspiration and
  - Higher use of GP IIb/IIIa inhibitors.
- No significant differences were observed between the groups in terms of all-cause mortality or MACCE.

# Continue...

- A meta-analysis by Freixa et al., which encompasses six studies, 283 patients:
  - Three coronary reocclusions occurring
  - Improved left ventricular function
  - Lower MACE rate.
- Subsequently, Qiao et al. in their meta-analyses of nine studies found:
  - No difference in incidence of slow flow/no reflow
  - Improvement in LVEF in the long term
  - No difference in MACE.



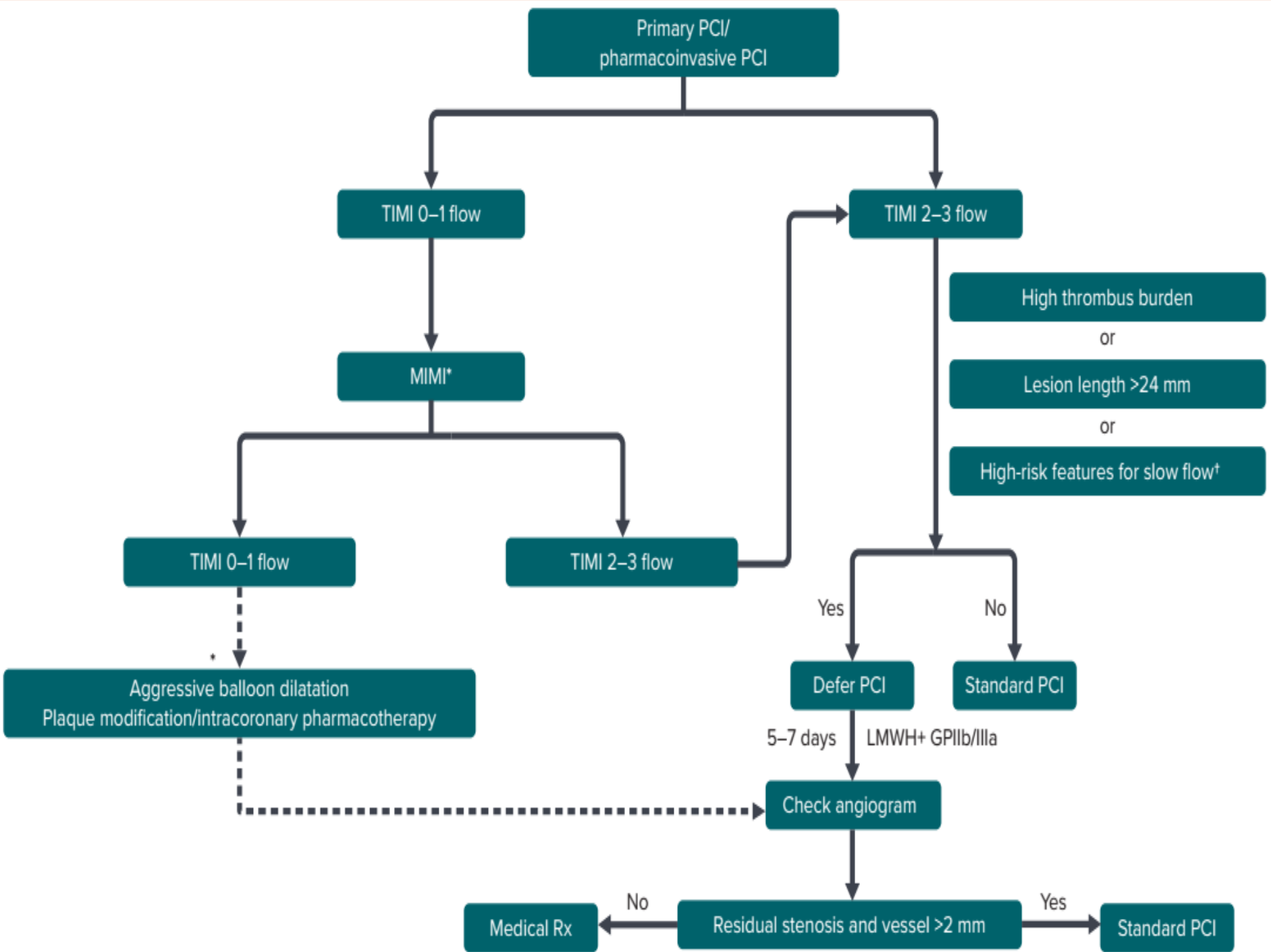


\*Randomised trials



## What Should Be the Ideal Deferral Time?

- A 7-day deferral is plausible with additional thrombus attenuation and improved periprocedural advantages though only a minority of RCTs enrolled such patients.
- Administer IV GPIIb/IIIa inhibitor for 12–16 hours after the procedure followed by low molecular weight heparin for 5–7 days or until the next angiogram depending upon the baseline CRUSADE score.
- In patients with a CRUSADE score  $>20$ , the GP IIb/IIIa infusion is avoided and only intracoronary bolus is provided.
- For those at score  $>50$ , the duration of heparin is also reduced.



Primary PCI/  
pharmacoinvasive PCI

TIMI 0-1 flow

TIMI 2-3 flow

High thrombus burden

or

Lesion length >24 mm

or

High-risk features for slow flow\*

MIMI\*

TIMI 0-1 flow

TIMI 2-3 flow

Aggressive balloon dilatation  
Plaque modification/intracoronary pharmacotherapy

Defer PCI

Standard PCI

Yes

No

5-7 days LMWH+ GPIIb/IIIa

Check angiogram

No

Yes

Medical Rx

Residual stenosis and vessel >2 mm

Standard PCI

# Deferred stenting

```
graph LR; A((Deferred stenting)) --- B((Disadvantages)); A --- C((Problems with immediate stenting)); A --- D((Advantages));
```

## Disadvantages

- Reocclusion
- Unplanned revascularisation
- Increase bleeding from extended parenteral anticoagulation
- Increased medical expenses

## Problems with immediate stenting

- Increased slow flow/no reflow
- Increased stent thrombosis
- Late stent malapposition
- Smaller stent sizes due to vasospasm

## Advantages

- Low thrombus burden
- Decreased periprocedural MI
- Improved thrombolysis in MI flow
- Slow flow/no reflow prevented
- Larger stent size
- Lesser number of stents implantation
- Smaller infarct size

# Future Directions

- INNOVATIONCORE (NCT03744000): 2024

To evaluate the impact of deferred versus immediate stenting in patients with acute ST-segment elevation anterior wall myocardial infarction (STEMI)

- OPTIMAL(NCT03282773): 2022

The OPTIMAL study is designed to compare the clinical performance of deferred stenting with that of immediate stenting for AMI caused by left main coronary artery occlusion.



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