

Which CTO should be intervented?

beyond guideline indications

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CASE Presentation

- 65y.o. Male
- Angina free
- HT, DM, Obese (145kg), Hlip, TIA, minor phys. activity
- Previous inf-post MI - ? years ago
- LVEF – 45%
- 2016 LAD PCI
- Admitted for carotis angioplasty (2017)

2017 Angiography



2017 LAD PCI



2023 Angiography

(presented with **NSTEMI**)

LVEF= 30%



2023 LAD PCI



What should be the next step?

- CX – RCA PCI?
- OMT?

CTOs are a common finding, present in **18-52%** of the patients with coronary artery disease who undergo coronary angiography

The presence of a CTO is independently associated with poorer prognosis, with an almost **2.9-fold increased risk of mortality** in patients presenting with a STEMI and concurrent CTO

What guidelines say?

ACC/AHA/SCAI CLINICAL PRACTICE GUIDELINE

2021 ACC/AHA/SCAI Guideline for Coronary Artery Revascularization: A Report of the American College of Cardiology/American Heart Association Joint Committee on Clinical Practice Guidelines

10.7. Treatment of CTO

Recommendation for Treatment of CTO

Referenced studies that support the recommendation are summarized in Online Data Supplement 29.

| COR | LOE | Recommendation |
|-----------|------------|--|
| 2b | B-R | 1. In patients with suitable anatomy who have refractory angina on medical therapy, after treatment of non-CTO lesions, the benefit of PCI of a CTO to improve symptoms is uncertain. ¹⁻⁴ |

What guidelines say?



ESC

European Society
of Cardiology

European Heart Journal (2019) 40, 87–165

doi:10.1093/eurheartj/ehy394

ESC/EACTS GUIDELINES

2018 ESC/EACTS Guidelines on myocardial revascularization

The Task Force on myocardial revascularization of the European Society of Cardiology (ESC) and European Association for Cardio-Thoracic Surgery (EACTS)

Percutaneous revascularization of CTOs should be considered in patients with angina resistant to medical therapy or with a large area of documented ischaemia in the territory of the occluded vessel.^{629,659–663}

IIa

B

Arguments against opening CTOs

- ✓ It is already occluded and can't get any worse
- ✓ It is worthless to open because most of the muscle must already be non-viable
- ✓ There is no need to intervene even if there is viable myocardium because the collateral circulation will suffice

Counter argument

- ✓ CTOs frequently serve viable myocardium and they are all flow limiting

Indications for opening CTOs

- Improving symptoms and quality of life
- Reducing mortality or myocardial infarction
- Improving tolerance to future myocardial infarctions
- Improving left ventricular function

• Improving symptoms and quality of life

- Reducing mortality or myocardial infarction
- Improving tolerance to future myocardial infarctions
- Improving left ventricular function

Open-CTO registry

EURO-CTO trial

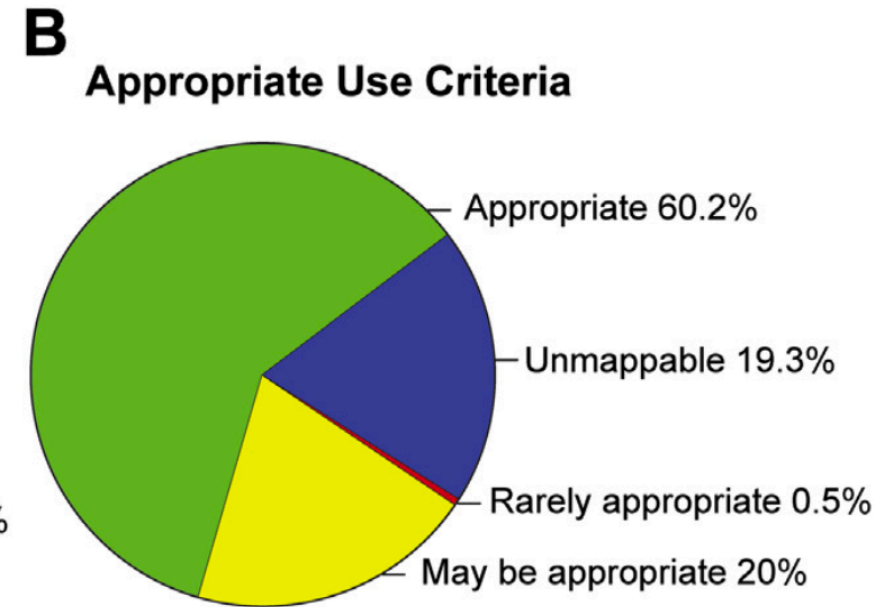
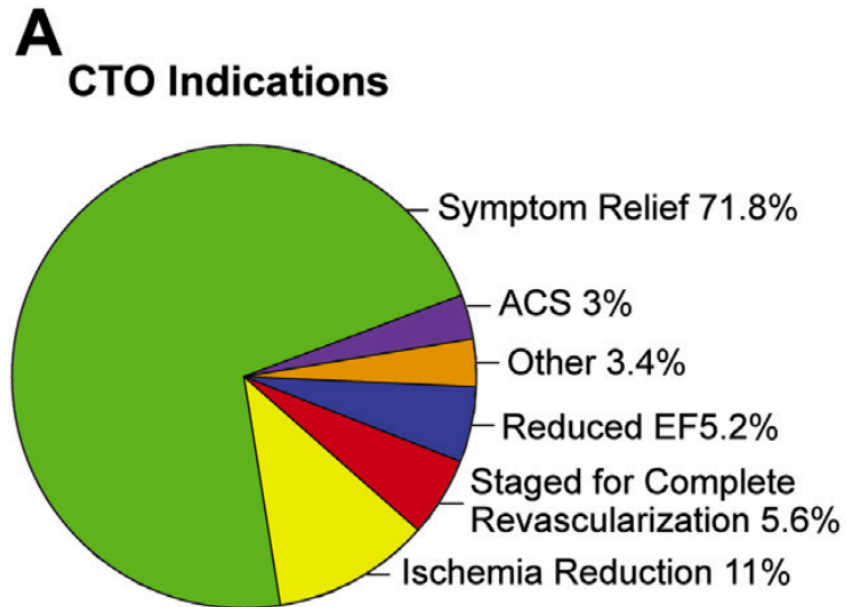
• Improving symptoms and quality of life

Open-CTO registry

- Reducing mortality or myocardial infarction
- Improving tolerance to future myocardial infarctions
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Early Procedural and Health Status Outcomes After Chronic Total Occlusion Angioplasty

A Report From the OPEN-CTO Registry (Outcomes, Patient Health Status, and Efficiency in Chronic Total Occlusion Hybrid Procedures)



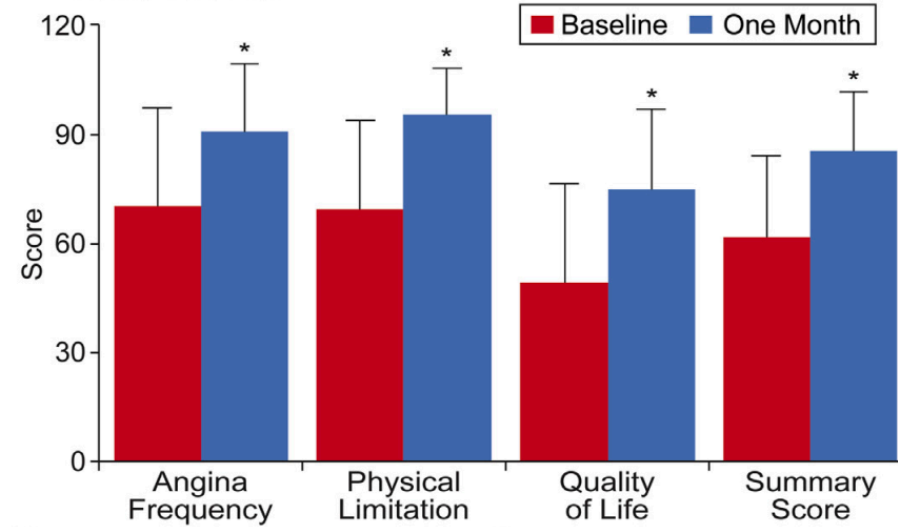
Distributions of the indications **(A)** for and appropriateness ratings **(B)** of the procedures. ACS = acute coronary syndrome; CTO = chronic total occlusion; EF = ejection fraction.

Improving symptoms and quality of life

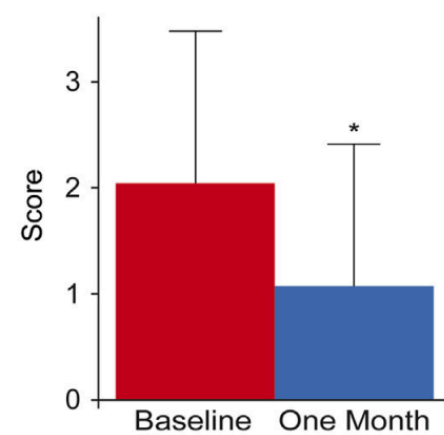
Open-CTO registry

- Reducing mortality or myocardial infarction
- Improving tolerance to future myocardial infarctions
- Improving left ventricular function

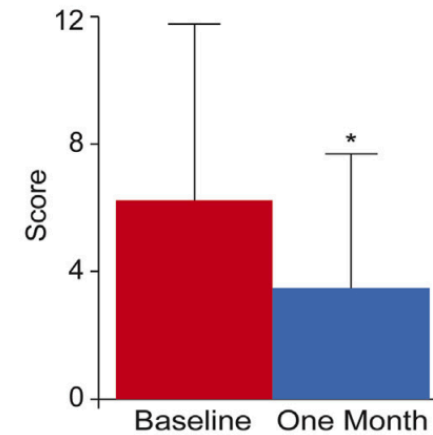
A SAQ Scores



B RDS Scores



C PHQ-8 Scores



*p < 0.01 vs. Baseline

Unadjusted mean \pm SEM health status scores at baseline and 1 month among those with paired data (n = 890). PHQ-8 = Physicians Health Questionnaire 8; RDS = Rose Dyspnea Scale; SAQ = Seattle Angina Questionnaire.

• Improving symptoms and quality of life

- Reducing mortality or myocardial infarction
- Improving tolerance to future myocardial infarctions
- Improving left ventricular function

A randomized multicentre trial to compare revascularization with optimal medical therapy for the treatment of chronic total coronary occlusions

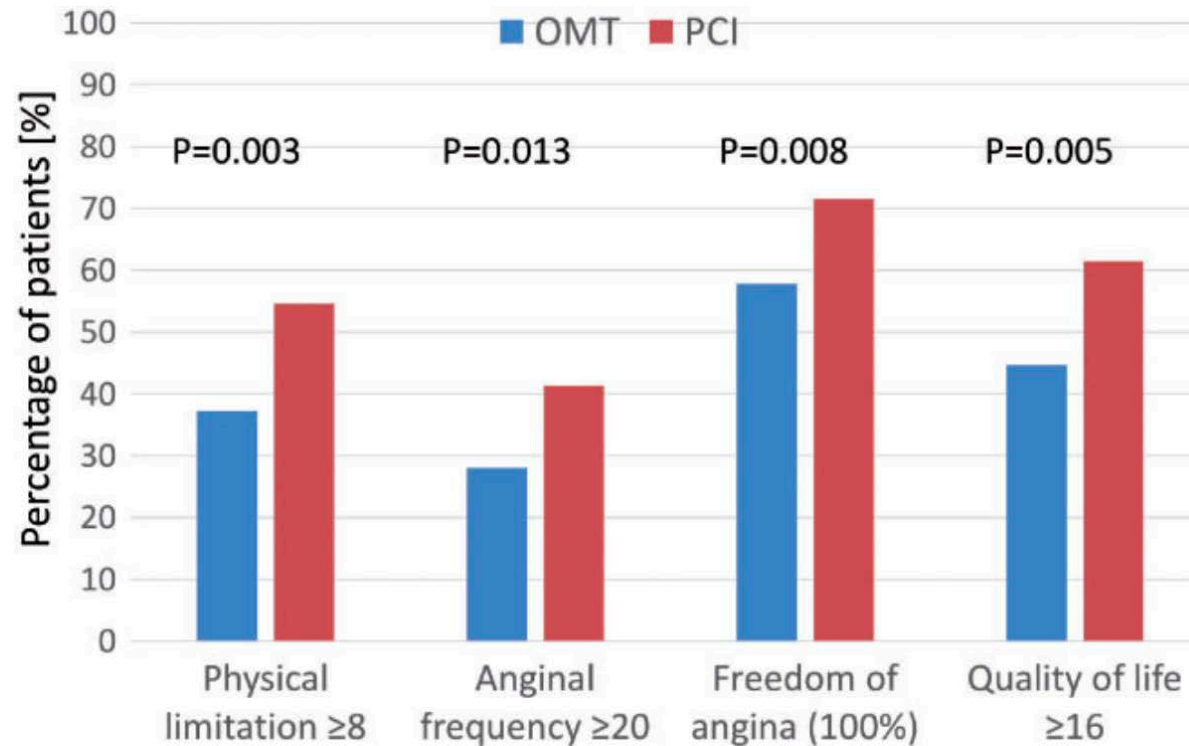


Figure 4 Comparison of significant changes in the Seattle angina questionnaire categories from baseline to follow-up between optimal medical therapy and percutaneous coronary intervention.

• Improving symptoms and quality of life

- Reducing mortality or myocardial infarction
- Improving tolerance to future myocardial infarctions
- Improving left ventricular function

| SAQ angina frequency | | | | |
|----------------------|-------------|-------------|-----------------------|------|
| 1 mo | 94.63±10.54 | 93.31±13.78 | 1.33 (−0.81 to 3.46) | 0.23 |
| 6 mo | 96.00±10.13 | 95.44±9.98 | 0.56 (−1.30 to 2.42) | 0.56 |
| 12 mo | 94.55±11.18 | 95.33±10.19 | −0.78 (−2.83 to 1.26) | 0.45 |
| 24 mo | 97.31±7.13 | 97.18±7.65 | 0.13 (−1.43 to 1.69) | 0.87 |
| 36 mo | 98.21±5.32 | 97.38±7.20 | 0.83 (−0.67 to 2.32) | 0.27 |

| SAQ quality of life | | | | |
|---------------------|-------------|-------------|----------------------|------|
| 1 mo | 66.16±19.87 | 64.26±19.65 | 1.90 (−1.55 to 5.35) | 0.28 |
| 6 mo | 72.08±17.54 | 69.74±17.48 | 2.34 (−0.90 to 5.58) | 0.16 |
| 12 mo | 72.19±19.06 | 71.89±16.6 | 0.30 (−3.12 to 3.71) | 0.86 |
| 24 mo | 77.37±17.43 | 75.91±17.77 | 1.45 (−2.25 to 5.16) | 0.44 |
| 36 mo | 78.26±17.39 | 77.53±16.69 | 0.73 (−3.26 to 4.72) | 0.72 |



Randomized Trial Evaluating Percutaneous Coronary Intervention for the Treatment of Chronic Total Occlusion

The DECISION-CTO Trial

Lee SW, Lee PH, Ahn JM, et al. Randomized Trial Evaluating Percutaneous Coronary Intervention for the Treatment of Chronic Total Occlusion. *Circulation*. 2019;139(14):1674-1683.

• Improving symptoms and quality of life

- Reducing mortality or myocardial infarction
- Improving tolerance to future myocardial infarctions
- Improving left ventricular function



Randomized Trial Evaluating Percutaneous Coronary Intervention for the Treatment of Chronic Total Occlusion

The DECISION-CTO Trial

SAQ treatment satisfaction

| | | | | |
|-------|-------------|-------------|----------------------|------|
| 1 mo | 83.07±12.75 | 80.42±15.03 | 2.66 (0.23 to 5.09) | 0.03 |
| 6 mo | 83.16±13.29 | 83.13±14.25 | 0.02 (-2.53 to 2.57) | 0.99 |
| 12 mo | 83.98±13.19 | 83.26±14.61 | 0.72 (-1.94 to 3.39) | 0.59 |
| 24 mo | 84.95±12.62 | 83.28±13.41 | 1.67 (-1.07 to 4.42) | 0.23 |
| 36 mo | 87.13±11.89 | 84.00±11.59 | 3.13 (0.38 to 5.89) | 0.03 |

• Improving symptoms and quality of life

- Reducing mortality or myocardial infarction
- Improving tolerance to future myocardial infarctions
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Initial Invasive or Conservative Strategy for Stable Coronary Disease

D.J. Maron, J.S. Hochman, H.R. Reynolds, S. Bangalore, S.M. O'Brien, W.E. Boden, B.R. Chaitman, R. Senior, J. López-Sendón, K.P. Alexander, R.D. Lopes, L.J. Shaw, J.S. Berger, J.D. Newman, M.S. Sidhu, S.G. Goodman, W. Ruzyllo, G. Gosselin, A.P. Maggioni, H.D. White, B. Bhargava, J.K. Min, G.B.J. Mancini, D.S. Berman, M.H. Picard, R.Y. Kwong, Z.A. Ali, D.B. Mark, J.A. Spertus, M.N. Krishnan, A. Elghamaz, N. Moorthy, W.A. Hueb, M. Demkow, K. Mavromatis, O. Bockeria, J. Peteiro, T.D. Miller, H. Szwed, R. Doerr, M. Keltai, J.B. Selvanayagam, P.G. Steg, C. Held, S. Kohsaka, S. Mavromichalis, R. Kirby, N.O. Jeffries, F.E. Harrell, Jr., F.W. Rockhold, S. Broderick, T.B. Ferguson, Jr., D.O. Williams, R.A. Harrington, G.W. Stone, and Y. Rosenberg, for the ISCHEMIA Research Group*

Total 5179 patients: 2588 undergo PCI, 2591 receive OMT

Follow up: median 3.2

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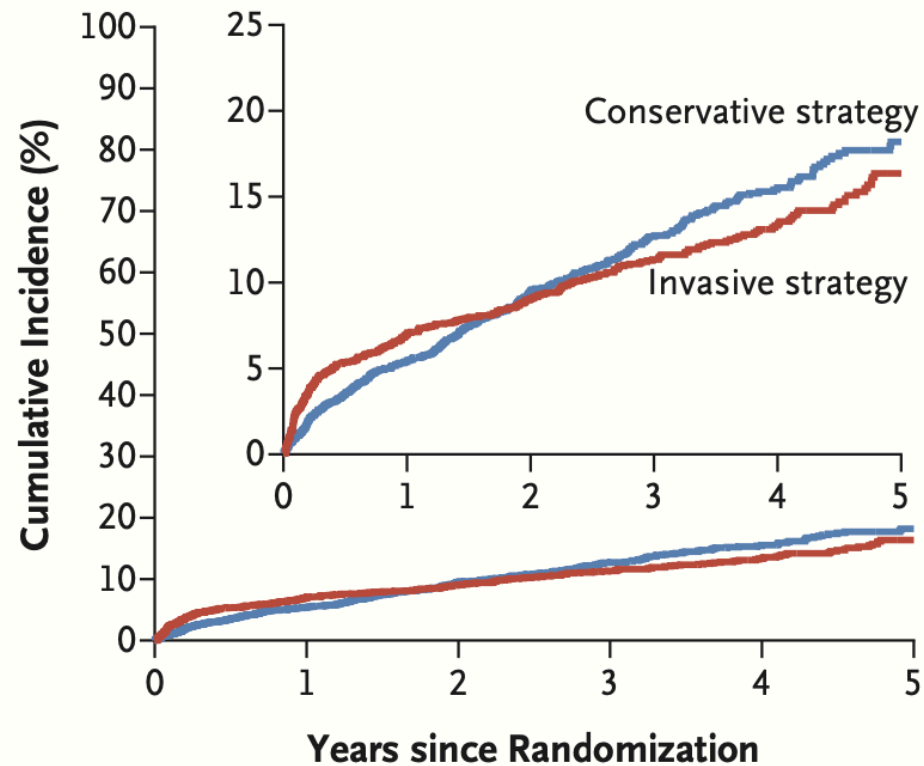
The NEW ENGLAND
JOURNAL of MEDICINE

ESTABLISHED IN 1812

APRIL 9, 2020

VOL. 382 NO. 15

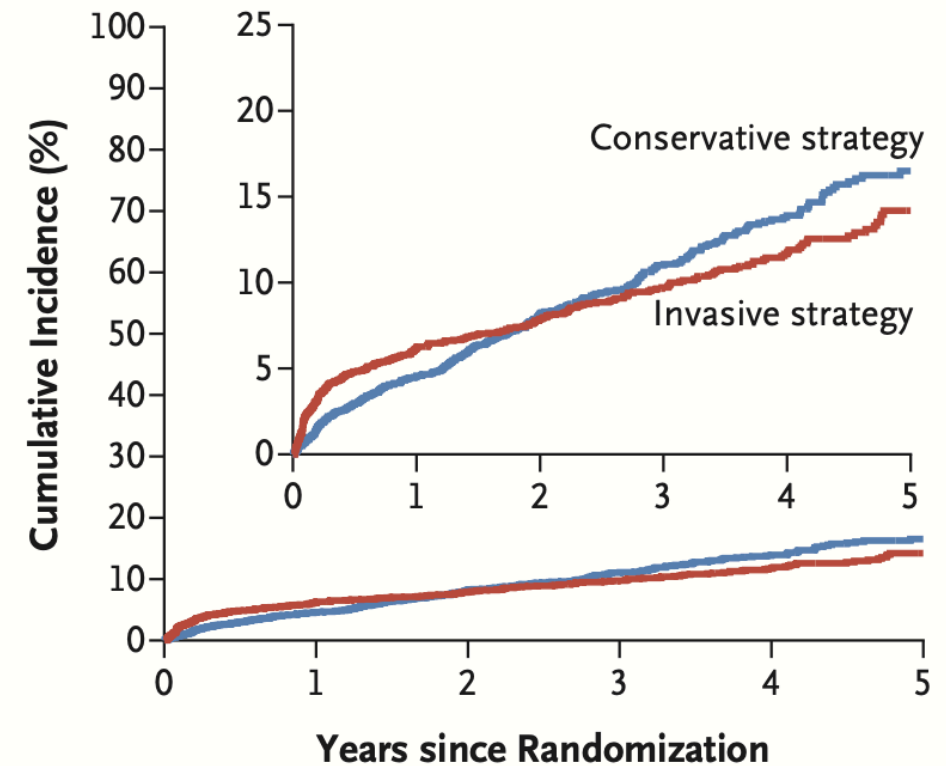
A Primary Composite Outcome



No. at Risk

| | | | | | | |
|-----------------------|------|------|------|------|-----|-----|
| Conservative strategy | 2591 | 2431 | 1907 | 1300 | 733 | 293 |
| Invasive strategy | 2588 | 2364 | 1908 | 1291 | 730 | 271 |

B Death from Cardiovascular Causes or Myocardial Infarction



No. at Risk

| | | | | | | |
|-----------------------|------|------|------|------|-----|-----|
| Conservative strategy | 2591 | 2453 | 1933 | 1325 | 746 | 298 |
| Invasive strategy | 2588 | 2383 | 1933 | 1314 | 742 | 282 |

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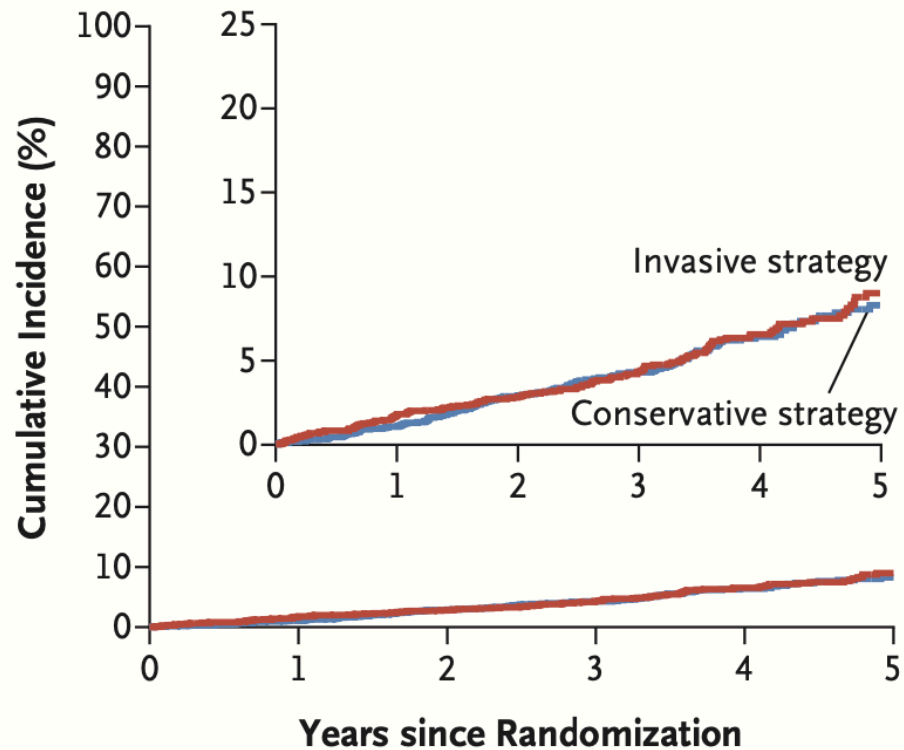
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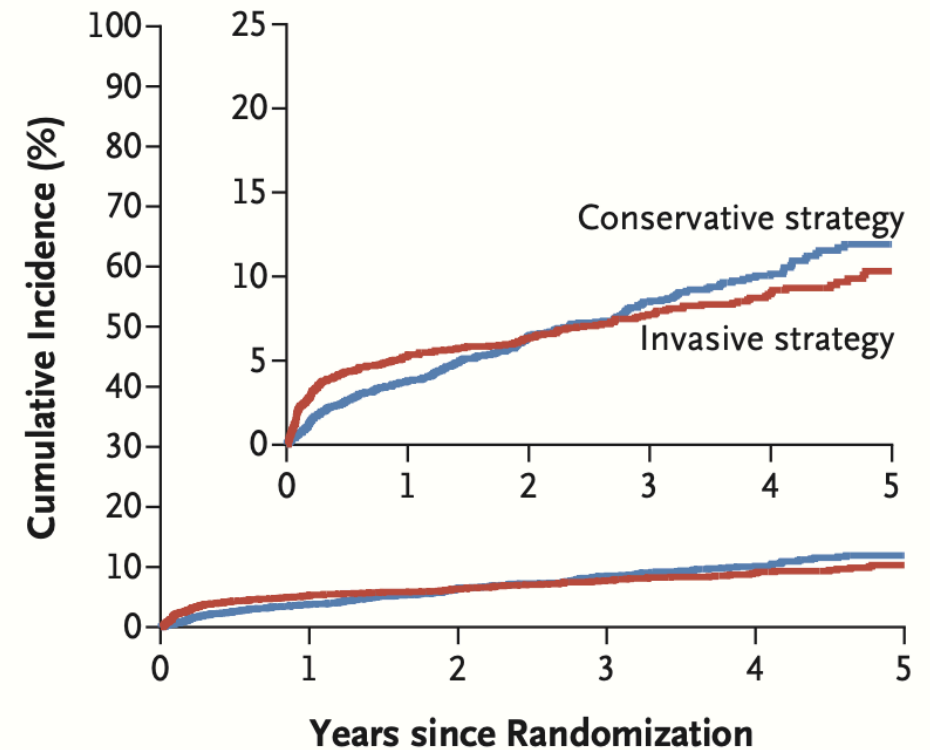
C Death from Any Cause



No. at Risk

| | | | | | | |
|-----------------------|------|------|------|------|-----|-----|
| Conservative strategy | 2591 | 2548 | 2065 | 1445 | 844 | 349 |
| Invasive strategy | 2588 | 2518 | 2061 | 1431 | 827 | 317 |

D Myocardial Infarction

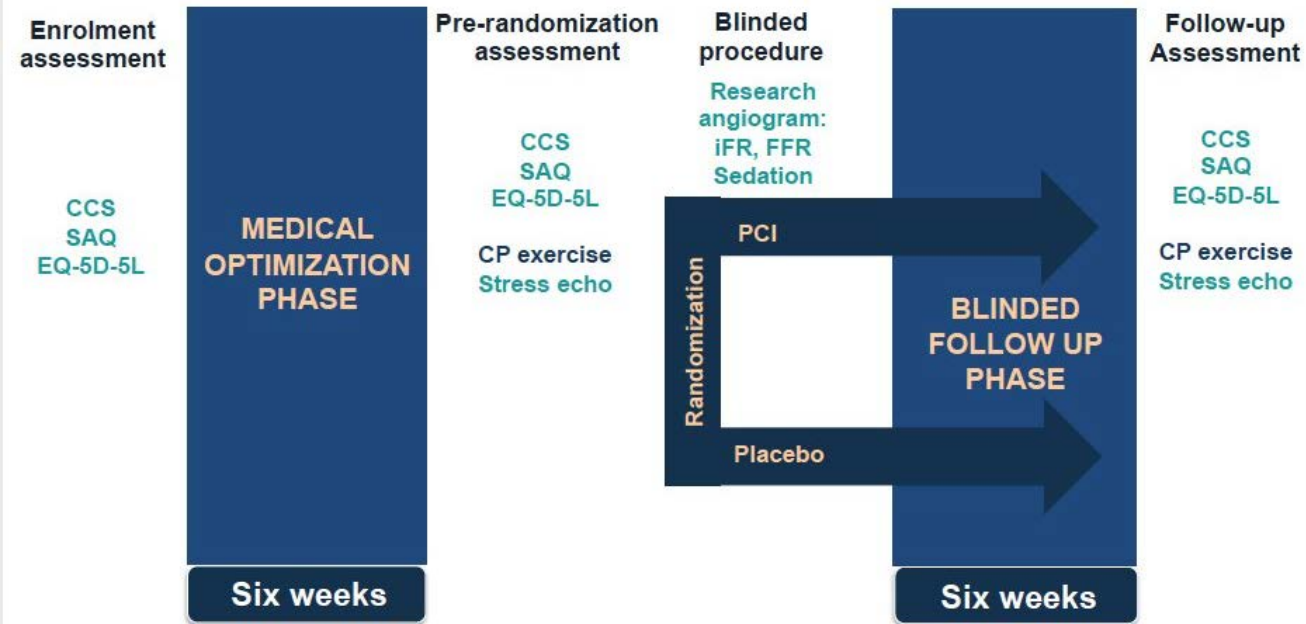


No. at Risk

| | | | | | | |
|-----------------------|------|------|------|------|-----|-----|
| Conservative strategy | 2591 | 2452 | 1931 | 1321 | 747 | 298 |
| Invasive strategy | 2588 | 2379 | 1931 | 1313 | 742 | 283 |

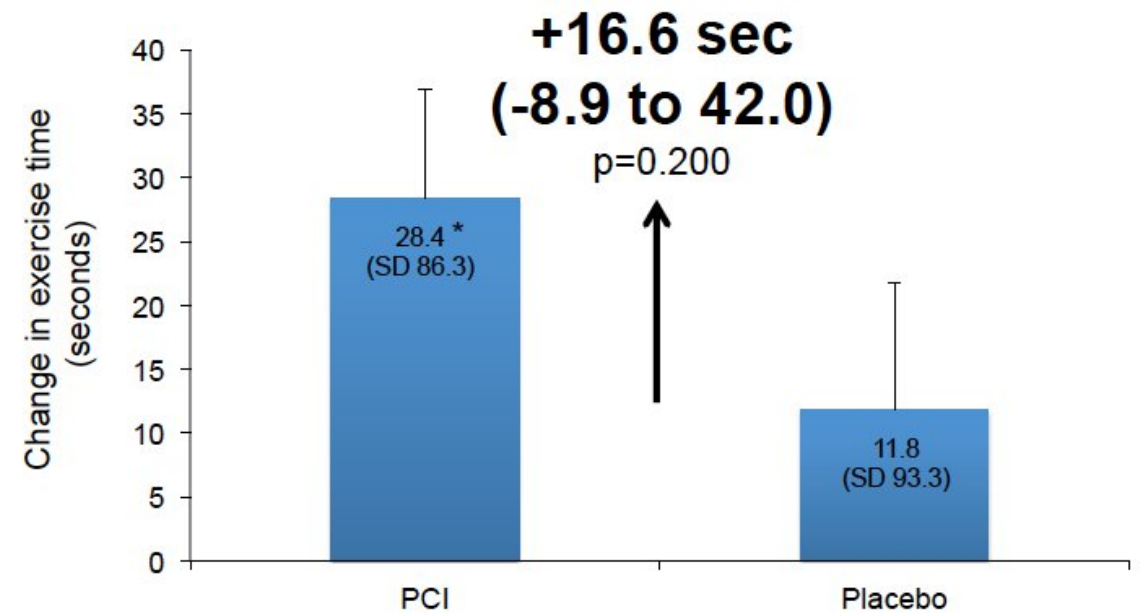
Orbita trial

Study design



Primary endpoint result

Change in total exercise time

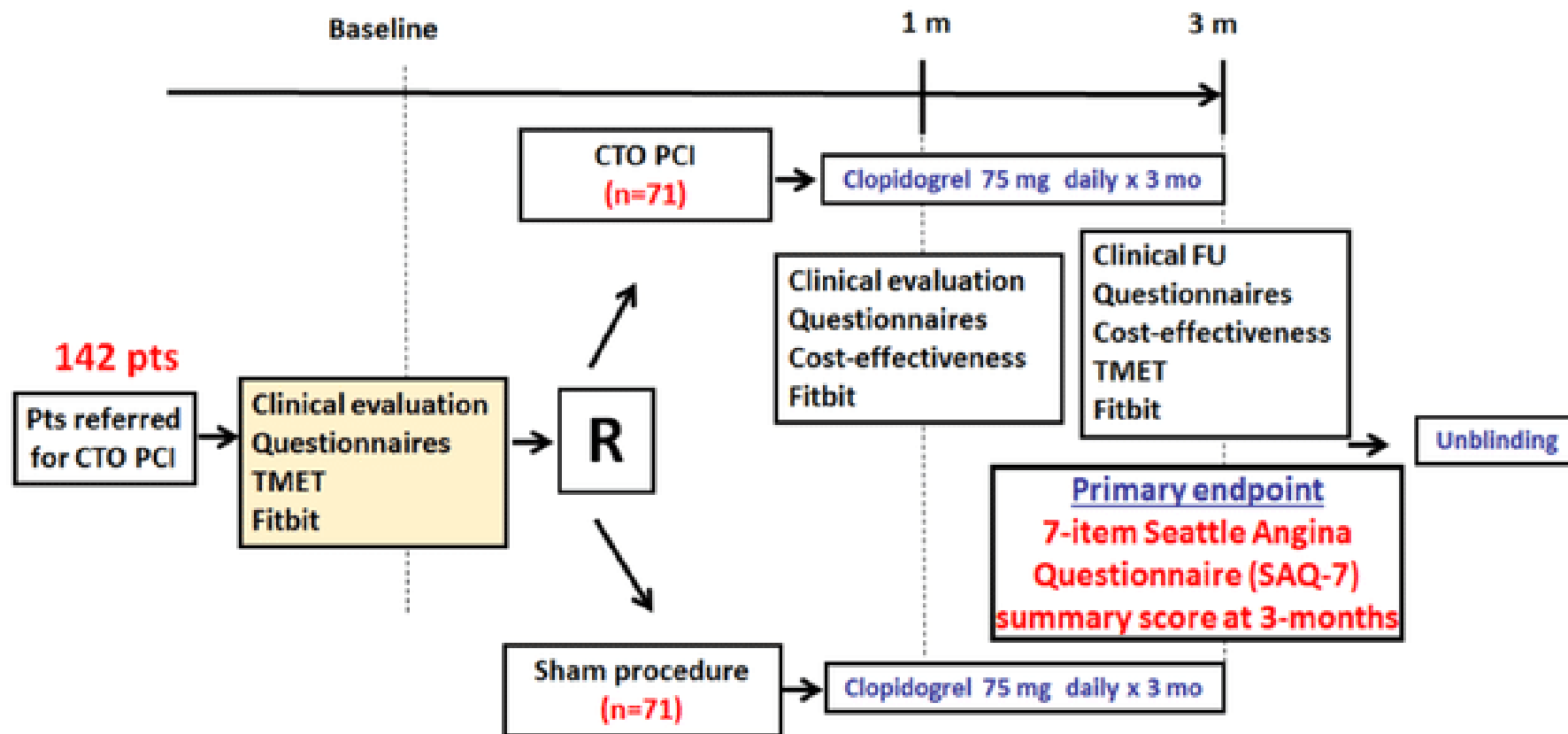


* = $p < 0.005$

SHINE-CTO

SHam-controlled INTervention
to improve QOL in CTOs

15 sites in the US, Europe, and Russia



Indications for opening CTOs

- Improving symptoms and quality of life
- Reducing mortality or myocardial infarction
- Improving tolerance to future myocardial infarctions
- Improving left ventricular function

- Improving symptoms and quality of life

• Reducing mortality or myocardial infarction

- Improving tolerance to future myocardial infarctions
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Randomized Trial Evaluating Percutaneous Coronary Intervention for the Treatment of Chronic Total Occlusion

The DECISION-CTO Trial

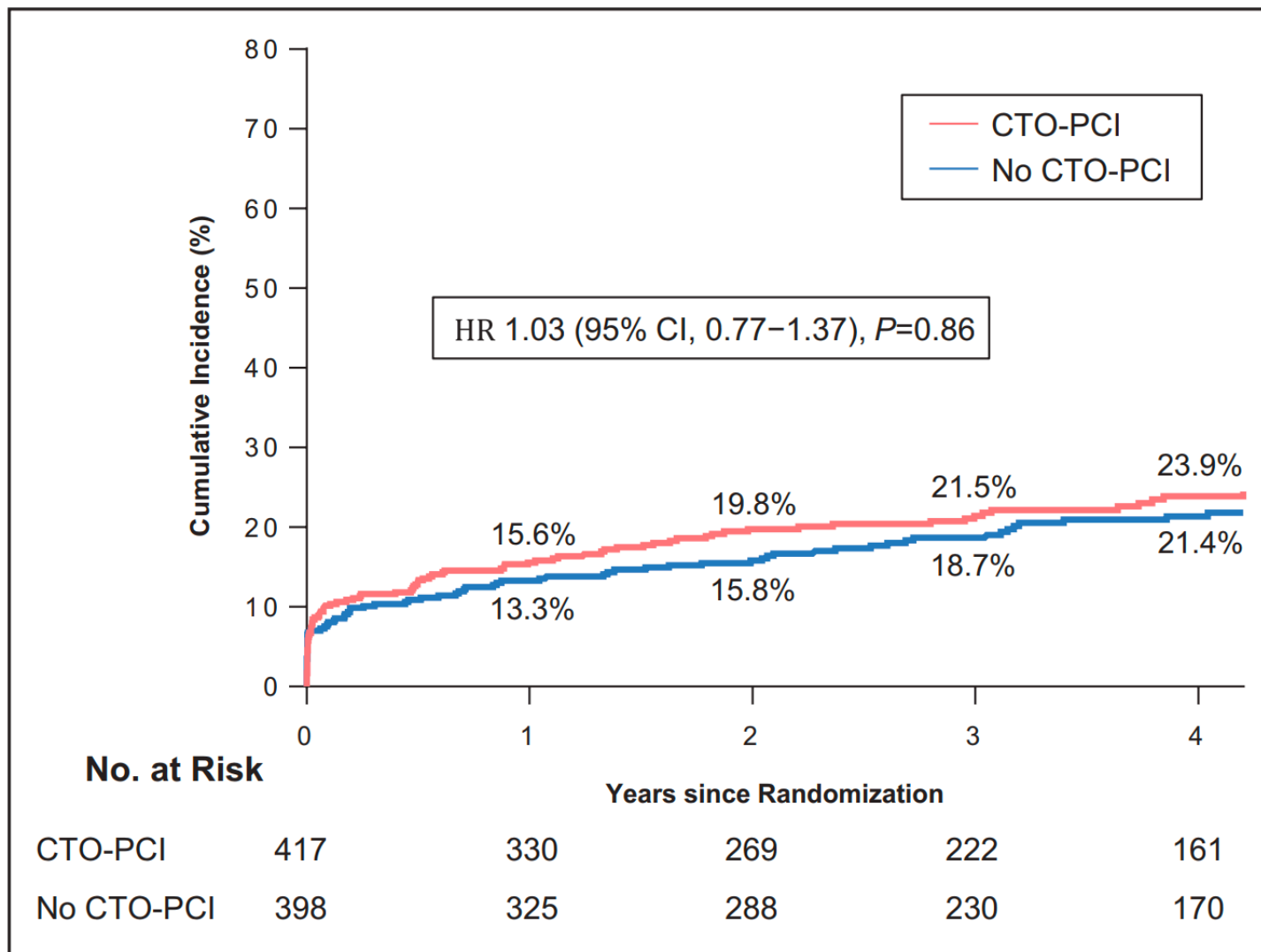


Figure 2. Cumulative incidence of the primary end point in the intention-to-treat population.

CTO indicates chronic total occlusion; HR, hazard ratio; and PCI, percutaneous coronary intervention.

- Improving symptoms and quality of life

• Reducing mortality or myocardial infarction

- Improving tolerance to future myocardial infarctions
- Improving left ventricular function

DECISION-CTO Trial

ORIGINAL RESEARCH ARTICLE



Randomized Trial Evaluating Percutaneous
Coronary Intervention for the Treatment
of Chronic Total Occlusion

The DECISION-CTO Trial

- The study was stopped prematurely after recruiting **only 417 (planned 1200)** patients to undergo PCI of all lesions **including a CTO and 398 patients to PCI of nonCTO lesions only**, which limits the power of the analyses.
- There were also **high crossover rates** from medical therapy to PCI
- The long-term mortality was 1.9% with complete revascularization vs. 3.6% with nonCTO PCI only, a non-significant 47% relative risk reduction, similar to the magnitude observed in the latest meta-analysis of observational studies.

- Improving symptoms and quality of life

• Reducing mortality or myocardial infarction

- Improving tolerance to future myocardial infarctions
- Improving left ventricular function

Meta-Analysis of Clinical Outcomes of Patients Who Underwent Percutaneous Coronary Interventions for Chronic Total Occlusions



Georgios E. Christakopoulos, MD^a, Georgios Christopoulos, MD^a, Mauro Carlino, MD^b, Omar M. Jeroudi, MD^a, Michele Roesle, RN, BSN^a, Bavana V. Rangan, BDS, MPH^a, Shuaib Abdullah, MD^a, Jerrold Grodin, MD^a, Dharam J. Kumbhani, MD, SM, MRCP^a, Minh Vo, MD^c, Michael Luna, MD^a, Khaldoon Alaswad, MD^d, Dimitri Karpaliotis, MD^e, Stephane Rinfret, MD^f, Santiago Garcia, MD^g, Subhash Banerjee, MD^a, and Emmanouil S. Brilakis, MD, PhD^{a,*}

Successful percutaneous coronary intervention (PCI) for chronic total occlusions (CTOs) has been associated with clinical benefit. There are **no randomized** controlled trials on long-term clinical outcomes after CTO PCI, limiting the available evidence to observational cohort studies. We sought to perform a weighted meta-analysis of the long-term outcomes of successful versus failed CTO PCI. **A total of 25 studies, published from 1990 to 2014, with 28,486 patients (29,315 CTO PCI procedures) were included.** We analyzed data on mortality, subsequent coronary artery bypass grafting (CABG), myocardial infarction, major adverse cardiac events, angina pectoris, stroke, and target vessel revascularization using random-effects models. Procedural success was 71% (range 51% to 87%). During a weighted mean follow-up of 3.11 years, compared with unsuccessful, successful CTO PCI was associated with lower mortality (odds ratio [OR] 0.52, 95% confidence interval [CI] 0.43 to 0.63), less residual angina (OR 0.38, 95% CI 0.24 to 0.60), lower risk for stroke (OR 0.72, 95% CI 0.60 to 0.88), less need for subsequent coronary artery bypass grafting (OR 0.18, 95% CI 0.14 to 0.22), and lower risk for major adverse cardiac events (0.59, 95% CI 0.44 to 0.79). There was no difference in the incidence of target vessel revascularization (OR 0.66, 95% CI 0.36 to 1.23) or myocardial infarction (OR 0.73, 95% CI 0.52 to 1.03). Outcomes were similar in patients who underwent balloon angioplasty only or stenting with bare metal or drug-eluting stents. Compared with failed procedures, successful CTO PCIs are associated with a lower risk of death, stroke, and coronary artery bypass grafting and less recurrent angina pectoris. Published by Elsevier Inc. (Am J Cardiol 2015;115:1367–1375)

Christakopoulos GE, Christopoulos G, Carlino M, et al. Meta-analysis of clinical outcomes of patients who underwent percutaneous coronary interventions for chronic total occlusions. Am J Cardiol. 2015;115(10):1367-75

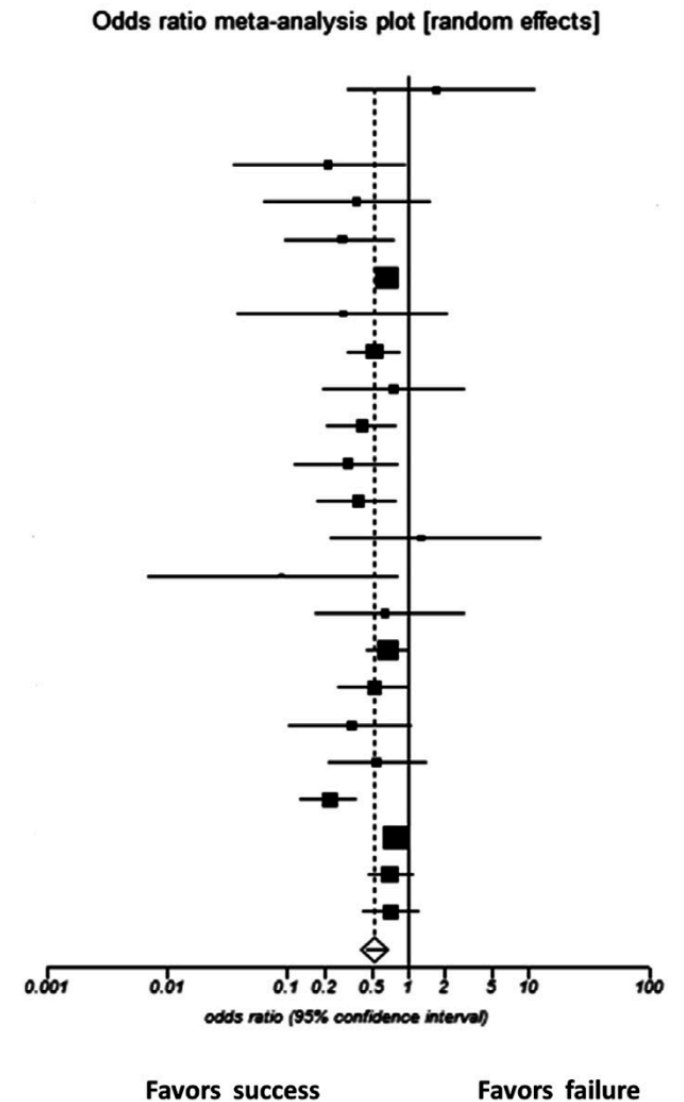
- Improving symptoms and quality of life

Reducing mortality or myocardial infarction

- Improving tolerance to future myocardial infarctions
- Improving left ventricular function

All-cause mortality

| Study | PCI success | | PCI failure | | Weight | Odds ratio [95%CI] |
|--------------|-------------|--------------|-------------|-------------|---------------|--------------------------|
| | Events | Total | Events | Total | | |
| Finci | 5 | 100 | 3 | 100 | 0.29 | 1.70 [0.32, 11.23] |
| Warren | 0 | 26 | 0 | 18 | 0 | * (excluded) |
| Ivanhoe | 3 | 317 | 7 | 163 | 0.94 | 0.21 [0.04, 0.95] |
| Angioi | 3 | 93 | 9 | 108 | 0.83 | 0.37 [0.06, 1.54] |
| Noguchi | 7 | 134 | 15 | 92 | 1.74 | 0.28 [0.09, 0.78] |
| Suero | 395 | 1491 | 180 | 514 | 20.22 | 0.67 [0.54, 0.84] |
| Olivari | 3 | 286 | 3 | 83 | 0.47 | 0.28 [0.04, 2.16] |
| Hoye | 37 | 567 | 36 | 304 | 4.50 | 0.52 [0.31, 0.87] |
| Drozd | 7 | 280 | 5 | 149 | 0.65 | 0.74 [0.20, 3.01] |
| Arslan | 19 | 117 | 37 | 115 | 3.21 | 0.41 [0.21, 0.80] |
| Aziz | 9 | 377 | 12 | 166 | 1.67 | 0.31 [0.12, 0.83] |
| Valenti | 17 | 344 | 17 | 142 | 2.35 | 0.38 [0.18, 0.83] |
| Labriole | 7 | 127 | 2 | 45 | 0.29 | 1.25 [0.23, 12.81] |
| Chen | 2 | 132 | 3 | 20 | 0.53 | 0.09 [0.01, 0.84] |
| Lee | 8 | 251 | 4 | 82 | 0.60 | 0.64 [0.17, 3.00] |
| Mehran | 74 | 1226 | 49 | 565 | 6.48 | 0.68 [0.46, 1.01] |
| Jolicoeur | 22 | 213 | 24 | 133 | 2.72 | 0.52 [0.27, 1.03] |
| Yang | 7 | 87 | 10 | 49 | 1.01 | 0.34 [0.10, 1.09] |
| Borgia | 19 | 237 | 9 | 65 | 1.34 | 0.54 [0.22, 1.44] |
| Jones | 26 | 582 | 44 | 254 | 6.01 | 0.22 [0.13, 0.38] |
| George S | 492 | 10199 | 259 | 4240 | 35.78 | 0.78 [0.67, 0.91] |
| Yamamoto | 92 | 1192 | 35 | 332 | 5.19 | 0.71 [0.47, 1.10] |
| Kim | 56 | 2045 | 20 | 523 | 3.18 | 0.71 [0.41, 1.26] |
| TOTAL | 1310 | 20423 | 783 | 8262 | 100.00 | 0.52 [0.43, 0.63] |



Christakopoulos GE, Christopoulos G, Carlino M, et al. Meta-analysis of clinical outcomes of patients who underwent percutaneous coronary interventions for chronic total occlusions. Am J Cardiol. 2015;115(10):1367-75

Figure 2. Forest plot for long-term all-cause mortality with successful versus failed CTO PCI.

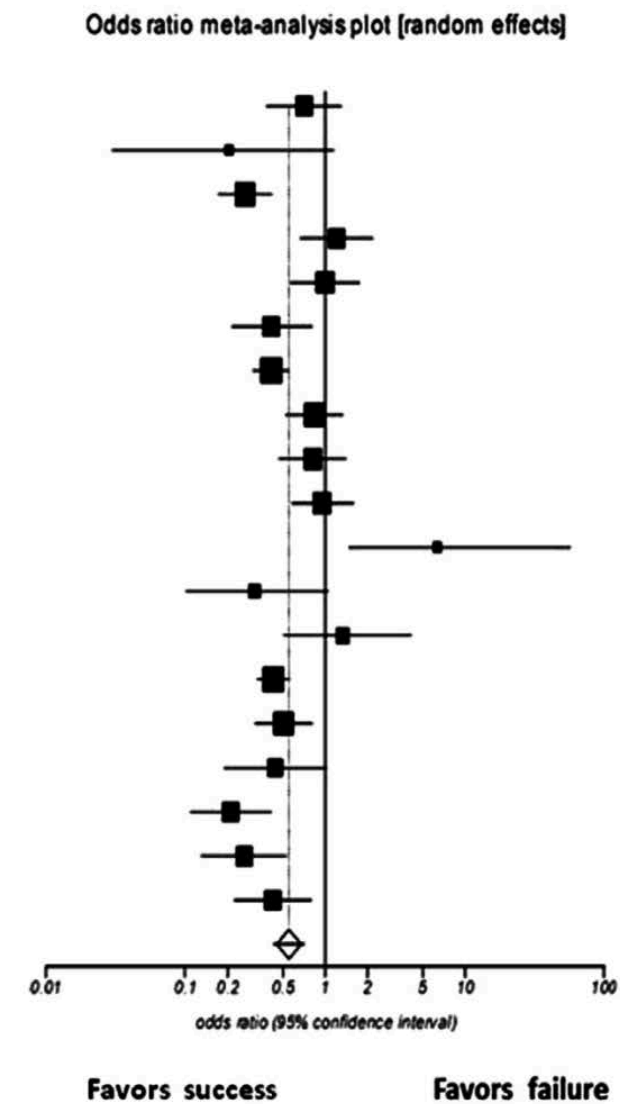
- Improving symptoms and quality of life

• Reducing mortality or myocardial infarction

- Improving tolerance to future myocardial infarctions
- Improving left ventricular function

MACE

| Study | PCI success | | PCI failure | | Weight% | Odds ratio [95%CI] |
|--------------|-------------|-------------|-------------|-------------|------------|--------------------------|
| | Events | Total | Events | Total | | |
| Finci | 33 | 100 | 41 | 100 | 3.36 | 0.71 [0.38, 1.31] |
| Warren | 3 | 26 | 7 | 18 | 0.90 | 0.20 [0.03, 1.15] |
| Ivanhoe | 63 | 317 | 76 | 158 | 9.94 | 0.27 [0.17, 0.41] |
| Angioi | 43 | 93 | 45 | 108 | 2.74 | 1.20 [0.66, 2.19] |
| Noguchi | 67 | 134 | 46 | 92 | 3.34 | 1.00 [0.57, 1.76] |
| Olivari | 35 | 286 | 21 | 83 | 3.50 | 0.41 [0.22, 0.80] |
| Hoye | 206 | 567 | 177 | 304 | 17.95 | 0.41 [0.30, 0.55] |
| Drozdz | 71 | 280 | 43 | 149 | 5.12 | 0.84 [0.53, 1.34] |
| Arslan | 55 | 117 | 60 | 115 | 3.92 | 0.81 [0.47, 1.41] |
| Valenti | 70 | 344 | 30 | 142 | 4.14 | 0.95 [0.58, 1.60] |
| Labriole | 29 | 127 | 2 | 45 | 0.28 | 6.36 [1.48, 57.01] |
| Chen | 19 | 132 | 7 | 20 | 1.27 | 0.31 [0.10, 1.06] |
| Lee | 24 | 251 | 6 | 82 | 1.00 | 1.34 [0.51, 4.16] |
| Mehran | 147 | 1226 | 137 | 565 | 20.19 | 0.43 [0.33, 0.56] |
| Jolicoeur | 75 | 213 | 69 | 133 | 6.73 | 0.50 [0.32, 0.80] |
| Yang | 19 | 87 | 19 | 49 | 2.32 | 0.44 [0.19, 1.02] |
| Borgia | 35 | 237 | 28 | 62 | 4.63 | 0.21 [0.11, 0.41] |
| Niccoli | 17 | 196 | 32 | 121 | 4.42 | 0.26 [0.13, 0.52] |
| Ciercwiez | 21 | 138 | 41 | 138 | 4.25 | 0.42 [0.22, 0.79] |
| TOTAL | 1032 | 4871 | 887 | 2484 | 100 | 0.55 [0.43, 0.71] |



Christakopoulos GE, Christopoulos G, Carlino M, et al. Meta-analysis of clinical outcomes of patients who underwent percutaneous coronary interventions for chronic total occlusions. Am J Cardiol. 2015;115(10):1367-75

Figure 4. Forest plot for long-term MACE with successful versus failed CTO PCI.

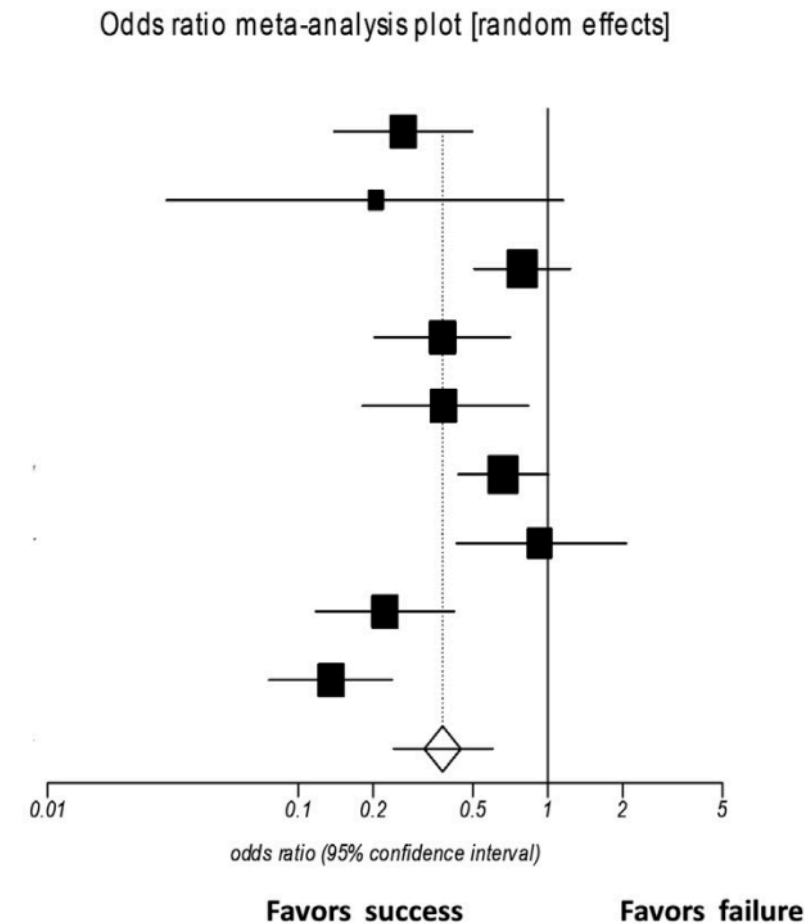
- Improving symptoms and quality of life

• Reducing mortality or myocardial infarction

- Improving tolerance to future myocardial infarctions
- Improving left ventricular function

Angina pectoris

| Study | PCI success | | PCI failure | | Weight% | Odds ratio [95%CI] |
|--------------|-------------|-------------|-------------|------------|------------|--------------------------|
| | Events | Total | Events | Total | | |
| Finci | 43 | 100 | 74 | 100 | c | 0.27 [0.14, 0.50] |
| Warren | 3 | 26 | 7 | 18 | 2.13 | 0.20 [0.03, 1.15] |
| Ivanhoe | 90 | 286 | 53 | 144 | 14.09 | 0.79 [0.51, 1.23] |
| Angioi | 27 | 93 | 56 | 108 | 10.72 | 0.34 [0.18, 0.64] |
| Olivari | 28 | 248 | 15 | 60 | 6.25 | 0.38 [0.18, 0.84] |
| Drozd | 120 | 280 | 79 | 149 | 17.19 | 0.66 [0.44, 1.01] |
| Jolicoeur | 21 | 213 | 14 | 133 | 4.53 | 0.93 [0.43, 2.06] |
| Borgia | 38 | 237 | 30 | 65 | 11.54 | 3.53 [1.87, 6.61] |
| Ciercwierz | 44 | 138 | 107 | 138 | 21.25 | 0.13 [0.07, 0.23] |
| TOTAL | 414 | 1621 | 435 | 915 | 100 | 0.38 [0.24, 0.60] |



Christakopoulos GE, Christopoulos G, Carlino M, et al. Meta-analysis of clinical outcomes of patients who underwent percutaneous coronary interventions for chronic total occlusions. Am J Cardiol. 2015;115(10):1367-75

Figure 5. Forest plot for angina with successful versus failed CTO PCI.

Indications for opening CTOs

- Improving symptoms and quality of life
- Reducing mortality or myocardial infarction
- Improving tolerance to future myocardial infarctions
- Improving left ventricular function

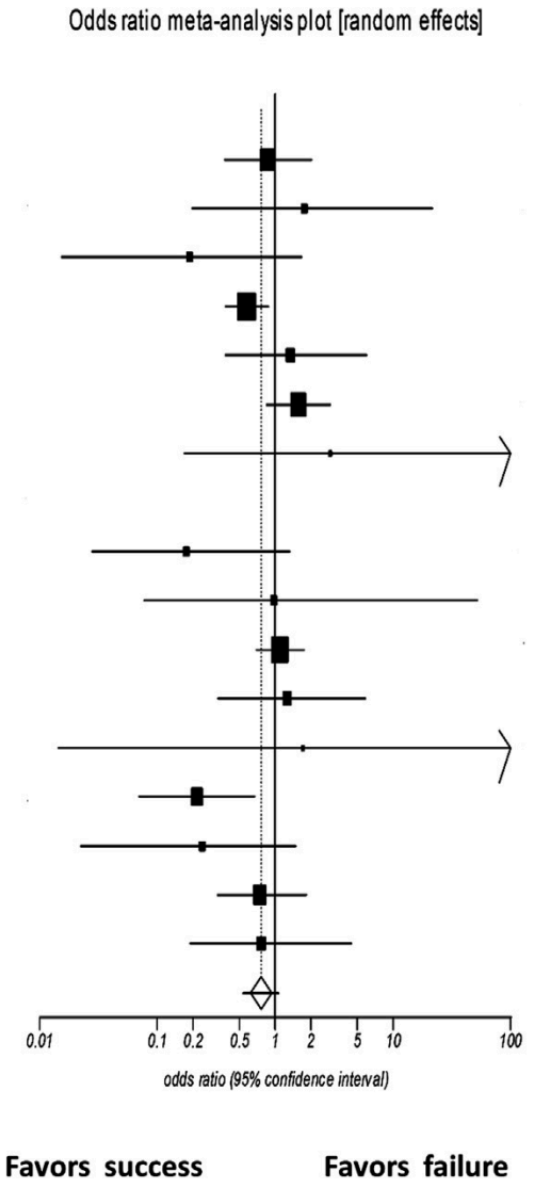
- Improving symptoms and quality of life
- Reducing mortality or myocardial infarction

• Improving tolerance to future myocardial infarctions

- Improving left ventricular function

Myocardial infarction

| Study | PCI success | | PCI failure | | Weight% | Odds ratio [95%CI] |
|--------------|-------------|-------------|-------------|-------------|------------|--------------------------|
| | Events | Total | Events | Total | | |
| Warren | 0 | 26 | 0 | 18 | 0 | * (excluded) |
| Ivanhoe | 19 | 317 | 11 | 158 | 7.16 | 0.85 [0.37, 2.04] |
| Angioi | 3 | 93 | 2 | 108 | 0.93 | 1.77 [0.20, 21.52] |
| Olivari | 2 | 286 | 3 | 83 | 2.40 | 0.19 [0.02, 1.68] |
| Hoye | 59 | 567 | 51 | 304 | 30.88 | 0.58 [0.38, 0.88] |
| Drozd | 10 | 280 | 4 | 149 | 2.61 | 1.34 [0.38, 5.96] |
| Arslan | 38 | 117 | 27 | 115 | 9.54 | 1.57 [0.85, 2.93] |
| Valenti | 3 | 344 | 0 | 142 | 0.36 | 2.92 [0.17, infinity] |
| Labriole | 0 | 127 | 0 | 40 | 0 | * (excluded) |
| Chen | 4 | 132 | 3 | 20 | 2.62 | 0.18 [0.03, 1.34] |
| Lee | 3 | 251 | 1 | 82 | 0.77 | 0.98 [0.08, 52.0] |
| Mehran | 71 | 1226 | 30 | 565 | 20.08 | 1.10 [0.70, 1.76] |
| Jolicoeur | 8 | 213 | 4 | 133 | 2.46 | 1.26 [0.33, 5.82] |
| Yang | 1 | 87 | 0 | 49 | 0.33 | 1.72 [0.01, infinity] |
| Borgia | 8 | 237 | 9 | 65 | 7.09 | 0.22 [0.07, 0.67] |
| Niccoli | 2 | 196 | 5 | 121 | 3.18 | 0.24 [0.02, 1.50] |
| Yamamoto | 30 | 1192 | 15 | 332 | 7.12 | 0.43 [0.23, 0.81] |
| Kim | 9 | 2045 | 3 | 523 | 2.47 | 0.77 [0.19, 4.42] |
| TOTAL | 270 | 7736 | 168 | 3007 | 100 | 0.73 [0.52, 1.03] |



Christakopoulos GE, Christopoulos G, Carlino M, et al. Meta-analysis of clinical outcomes of patients who underwent percutaneous coronary interventions for chronic total occlusions. Am J Cardiol. 2015;115(10):1367-75

Figure 3. Forest plot for MI with successful versus failed CTO PCI.

Indications for opening CTOs

- Improving symptoms and quality of life
- Reducing mortality or myocardial infarction
- Improving tolerance to future myocardial infarctions
- Improving left ventricular function

- Improving symptoms and quality of life
- Reducing mortality or myocardial infarction
- Improving tolerance to future myocardial infarctions

• Improving left ventricular function

Explore Trial

openheart Impact of collateralisation to a concomitant chronic total occlusion in patients with ST-elevation myocardial infarction: a subanalysis of the EXPLORE randomised controlled trial

| Table 5 Event rates at long-term follow-up | | | |
|---|-----------------------------|-----------------------------|-----------------|
| | GOODCOLL (n=162) | POORCOLL (n=140) | P values |
| All-cause death | 8 (5%) | 13 (9%) | 0.174 |
| Composite MACE | 16 (10%) | 20 (14%) | 0.286 |
| Cardiac death | 4 (3%) | 5 (4%) | 0.738 |
| All MI | 9 (6%) | 16 (11%) | 0.092 |
| CABG | 4 (3%) | 4 (3%) | 1.000 |

van Dongen IM, Elias J, van Houwelingen KG, et al. Impact of collateralisation to a concomitant chronic total occlusion in patients with ST-elevation myocardial infarction: a subanalysis of the EXPLORE randomised controlled trial. *Open Heart*. 2018;5(2):e000810. Published 2018 Jul 16.

- Improving symptoms and quality of life
- Reducing mortality or myocardial infarction
- Improving tolerance to future myocardial infarctions

• Improving left ventricular function

Explore Trial

openheart Impact of collateralisation to a concomitant chronic total occlusion in patients with ST-elevation myocardial infarction: a subanalysis of the EXPLORE randomised controlled trial

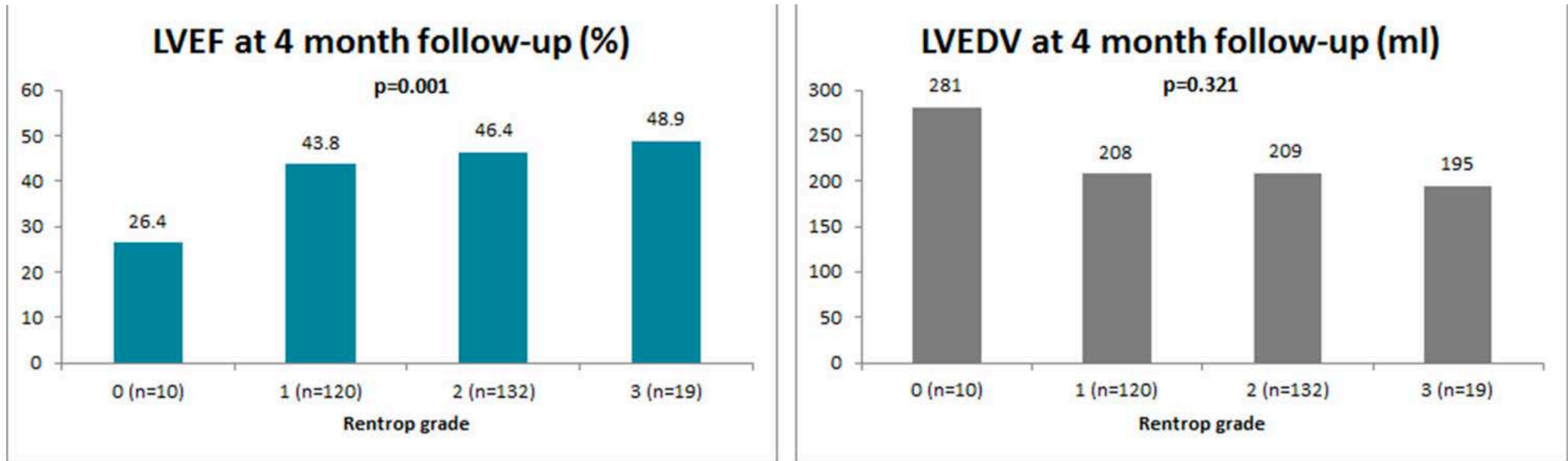


Figure 2 The impact of Rentrop grade collateralals to the coronary chronic total occlusion on left ventricular ejection fraction (LVEF) and left ventricular end-diastolic volume (LVEDV).

van Dongen IM, Elias J, van Houwelingen KG, et al. Impact of collateralisation to a concomitant chronic total occlusion in patients with ST-elevation myocardial infarction: a subanalysis of the EXPLORE randomised controlled trial. *Open Heart*. 2018;5(2):e000810. Published 2018 Jul 16.

- Improving symptoms and quality of life
- Reducing mortality or myocardial infarction
- Improving tolerance to future myocardial infarctions

[Explore Trial](#)

• Improving left ventricular function

- ✓ This study was assessing only ventricular remodelling in patients presenting with a STEMI with concurrent CTO, and hence does not reflect the majority of patients considered for CTO PC
- ✓ While no difference in the primary endpoint was observed, those with well-developed collaterals supplying the CTO territory (Rentropgrades 2-3) had significantly higher LVEF at 4 months ($46.2 \pm 11.4\%$ vs. $42.1 \pm 12.7\%$, $p=0.004$).
- ✓ Residual ischemia or viability in the CTO territory was not assessed prior to CTO PC

- Improving symptoms and quality of life
- Reducing mortality or myocardial infarction
- Improving tolerance to future myocardial infarctions

• Improving left ventricular function

[Explore Trial](#)

- ✓ Furthermore, the success rate of this study was relatively low, particularly compared to the other randomised trials published in the last few years, which may reflect either more complex disease, or recruitment in less experienced centres.

- Improving symptoms and quality of life
- Reducing mortality or myocardial infarction
- Improving tolerance to future myocardial infarctions

■ Improving left ventricular function

CTO specific studies, as well as general stable coronary disease studies, suggest that there is a survival benefit with revascularisation when the ischaemic territory **is greater than 12.5% of at risk myocardium**. It is thus unsurprising that the mortality benefit appears to be particularly related to PCI of the **LAD, rather than the RCA**

- Safley DM, Koshy S, Grantham JA, Bybee KA, House JA, Kennedy KF, et al. Changes in myocardial ischemic burden following percutaneous coronary intervention of chronic total occlusions. *Catheter Cardiovasc Interv* 2011;78(3):337–43.
- Claessen BE, Dangas GD, Godino C, Henriques JP, Leon MB, Park SJ, et al. Impact of target vessel on long-term survival after percutaneous coronary intervention for chronic total occlusions. *Catheter Cardiovasc Interv* 2013;82(1):76–82.

- Improving symptoms and quality of life
- Reducing mortality or myocardial infarction
- Improving tolerance to future myocardial infarctions

• Improving left ventricular function

Patients were stratified by their initial MPI summed difference score (SDS)

1. SDS <4—normal/minimal ischemia (<5% of ischemic myocardium);
2. SDS 4–8—mild ischemia (5–9.9%);
3. SDS 9–13—moderate ischemia (10–16%);
4. SDS >13—severe ischemia (>16%)

Changes in Myocardial Ischemic Burden Following Percutaneous Coronary Intervention of Chronic Total Occlusions

David M. Safley,^{1,2*} MD, Sindhu Koshy,¹ MD, J. Aaron Grantham,^{1,2} MD, Kevin A. Bybee,^{1,2} MD, John A. House,¹ MS, Kevin F. Kennedy,¹ MS, and Barry D. Rutherford,^{1,2} MD

Objectives: We assessed the potential for percutaneous coronary intervention (PCI) of a chronic total occlusion (CTO) to decrease myocardial ischemia and established objective criteria to predict post-procedure improvement. **Background:** Optimal treatment for CTO of coronary arteries is controversial, and selection criteria for PCI of CTO are subjective. **Methods:** All patients undergoing CTO PCI at a single center between 2002 and 2007 were included if myocardial perfusion imaging (MPI) was performed within 12 ± 3 months before and a follow-up study within 12 ± 3 months after PCI. Average summed difference scores were calculated and converted to percent ischemic myocardium to classify patients as having normal/minimal, mild, moderate, or severe ischemia. **A significant improvement in ischemia following PCI was classified as an absolute ≥5% decrease in ischemic myocardium.** Receiver operating characteristic (ROC) curves were used to identify ischemic thresholds predictive of decreased and increased ischemic burden on follow-up MPI. **Results:** In 301 patients, average baseline ischemic burden was 13.1% ± 11.9% and decreased to 6.9% ± 6.5% ($P < 0.001$) during follow-up. Overall, 53.5% of patients met criteria for improvement following PCI. These patients were more likely to be male, without diabetes, with CTO in the left anterior descending artery, and classified as having high ischemic burden at baseline. ROC analysis identified a baseline 12.5% ischemic burden as optimal in identifying those most likely to have a significantly decreased ischemic burden post-PCI. Those with a baseline ischemic burden less than 6.25% were more likely to have an increased ischemic burden post-PCI. **Conclusions:** Ischemic burden is reduced following CTO PCI, and the decrease is greater at high ischemic burden. **A threshold of 12.5% ischemic burden is suggested as a criterion for performing PCI in the setting of CTO.** © 2011 Wiley-Liss, Inc.

- Improving symptoms and quality of life
- Reducing mortality or myocardial infarction
- Improving tolerance to future myocardial infarctions

• Improving left ventricular function

NO ISHEMIA = HARM!

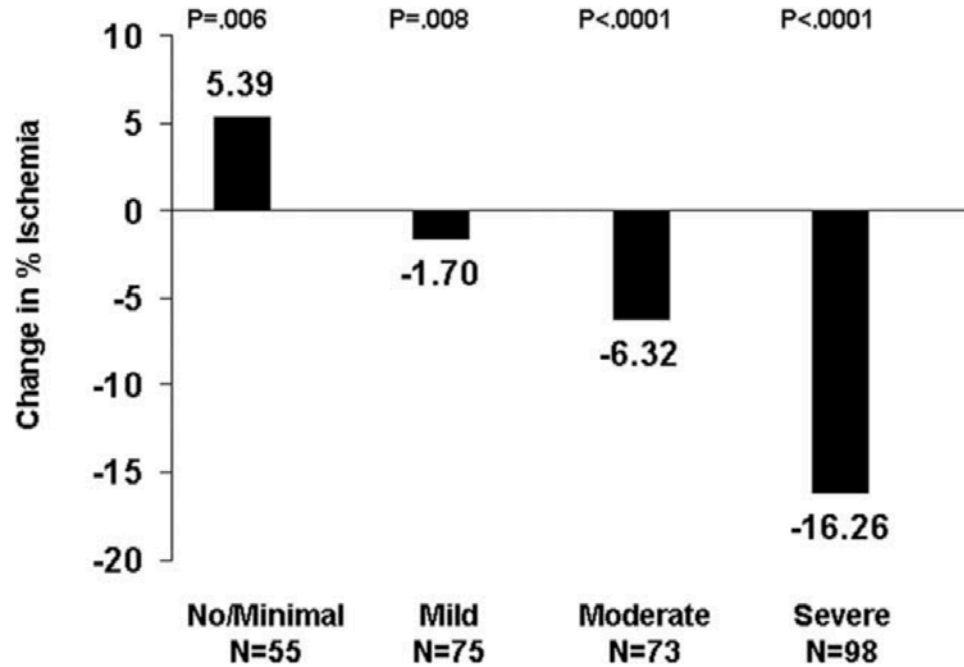
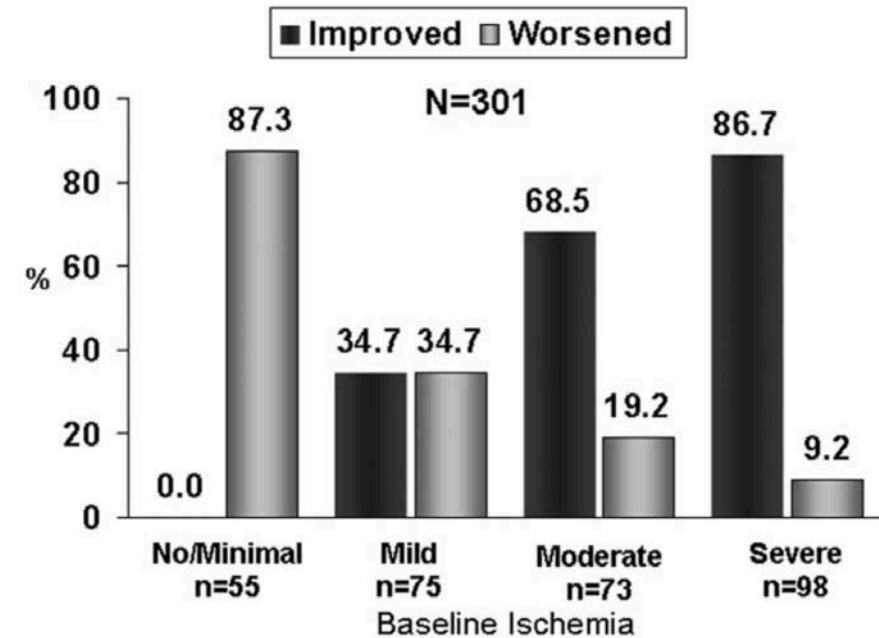


Fig. 1. Change in sum difference scores by severity of baseline ischemia assessed by myocardial perfusion imaging performed 12 months before and after percutaneous coronary intervention for chronic total occlusion.



P<.001 for the trend of more improvement with greater ischemia at baseline
P<.001 for the trend of more decrease with less ischemia at baseline

Fig. 2. Percentage of patients with changes of $\geq 5\%$ in ischemic myocardium after percutaneous coronary intervention for chronic total occlusion stratified by baseline ischemia.

- Improving symptoms and quality of life
- Reducing mortality or myocardial infarction
- Improving tolerance to future myocardial infarctions

• Improving left ventricular function

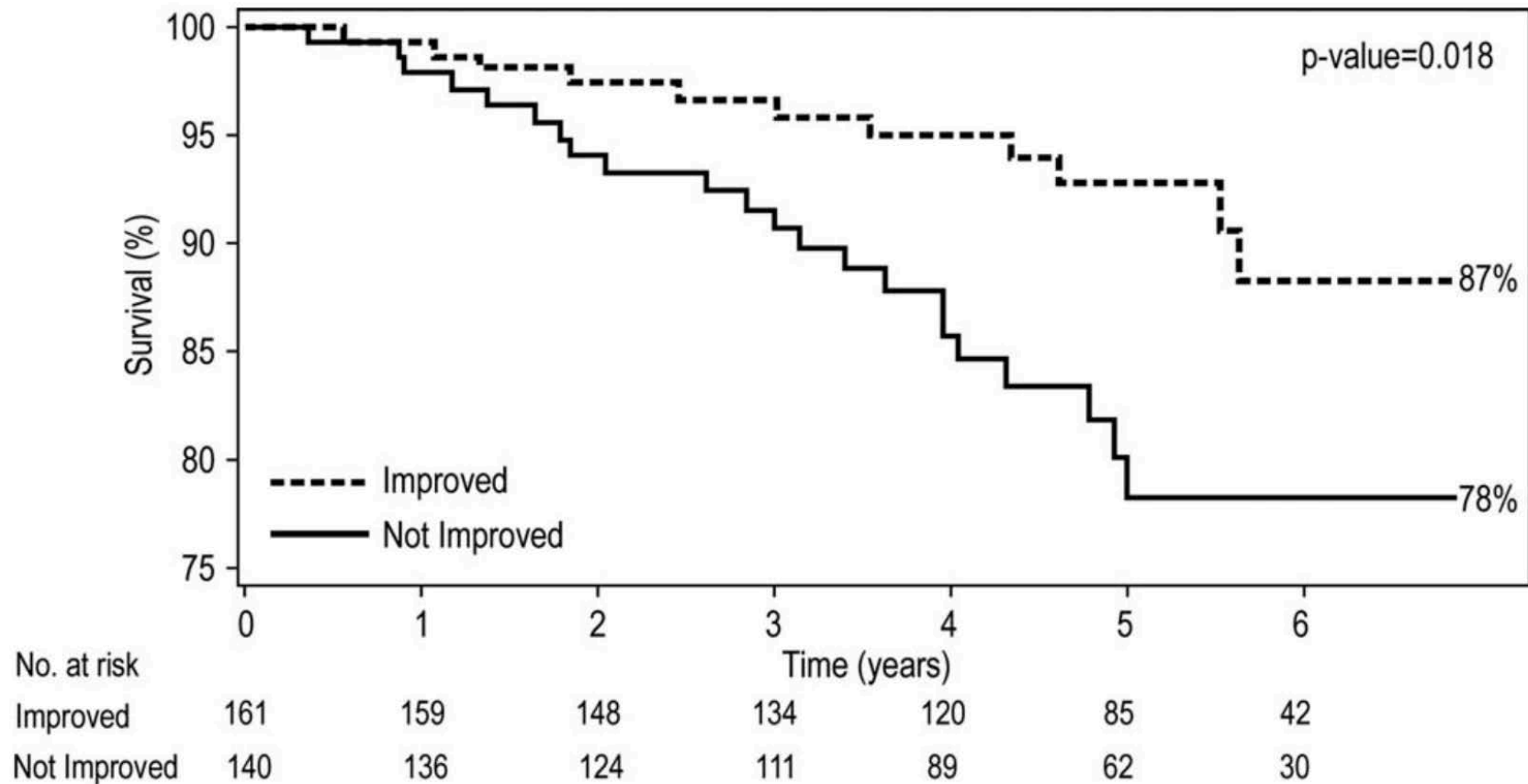


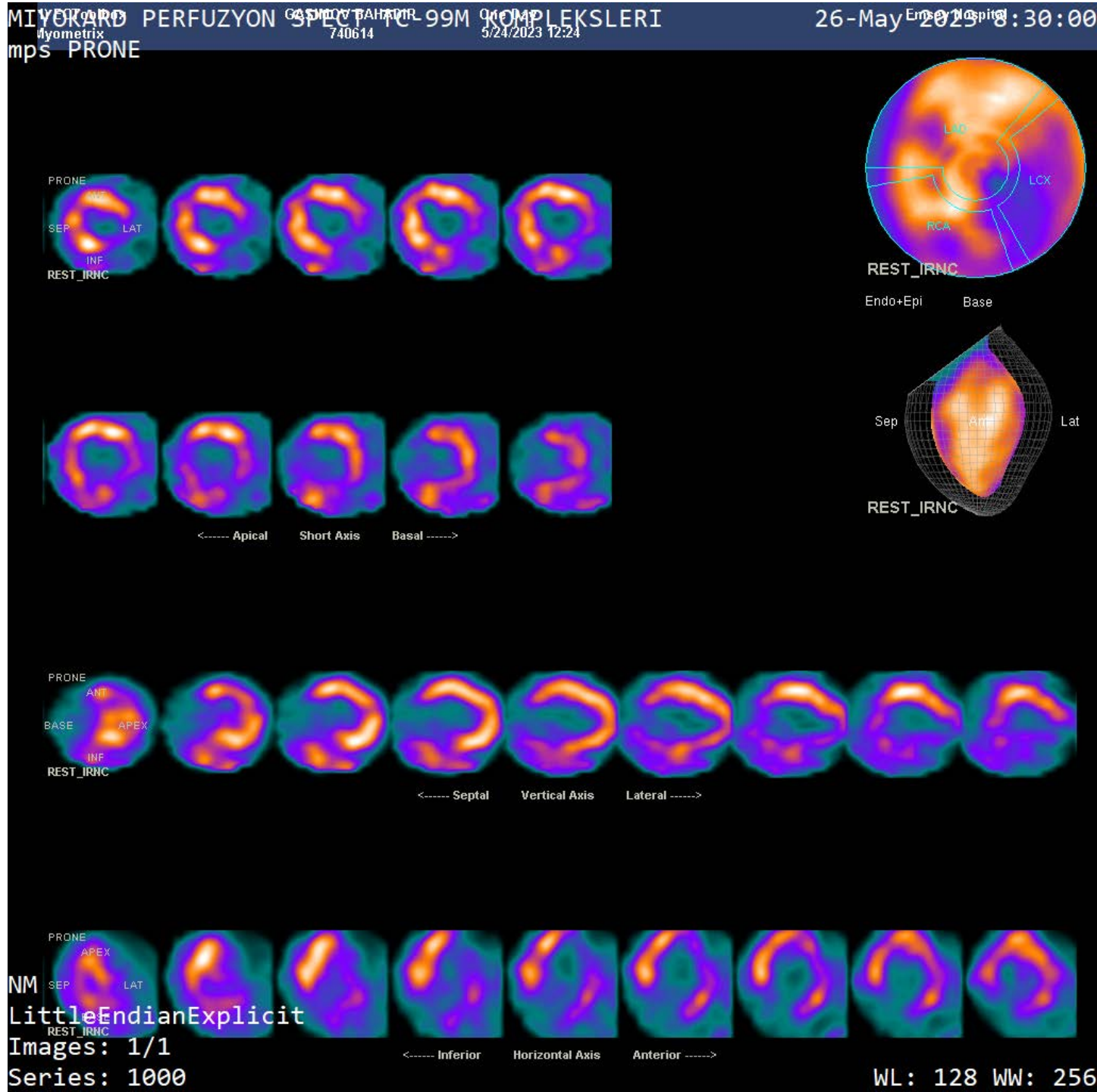
Fig. 3. Kaplan–Meier survival in patients with vs. without improvement in ischemia on myocardial perfusion imaging.

What should be the next step?

- CX – RCA PCI?
- OMT?

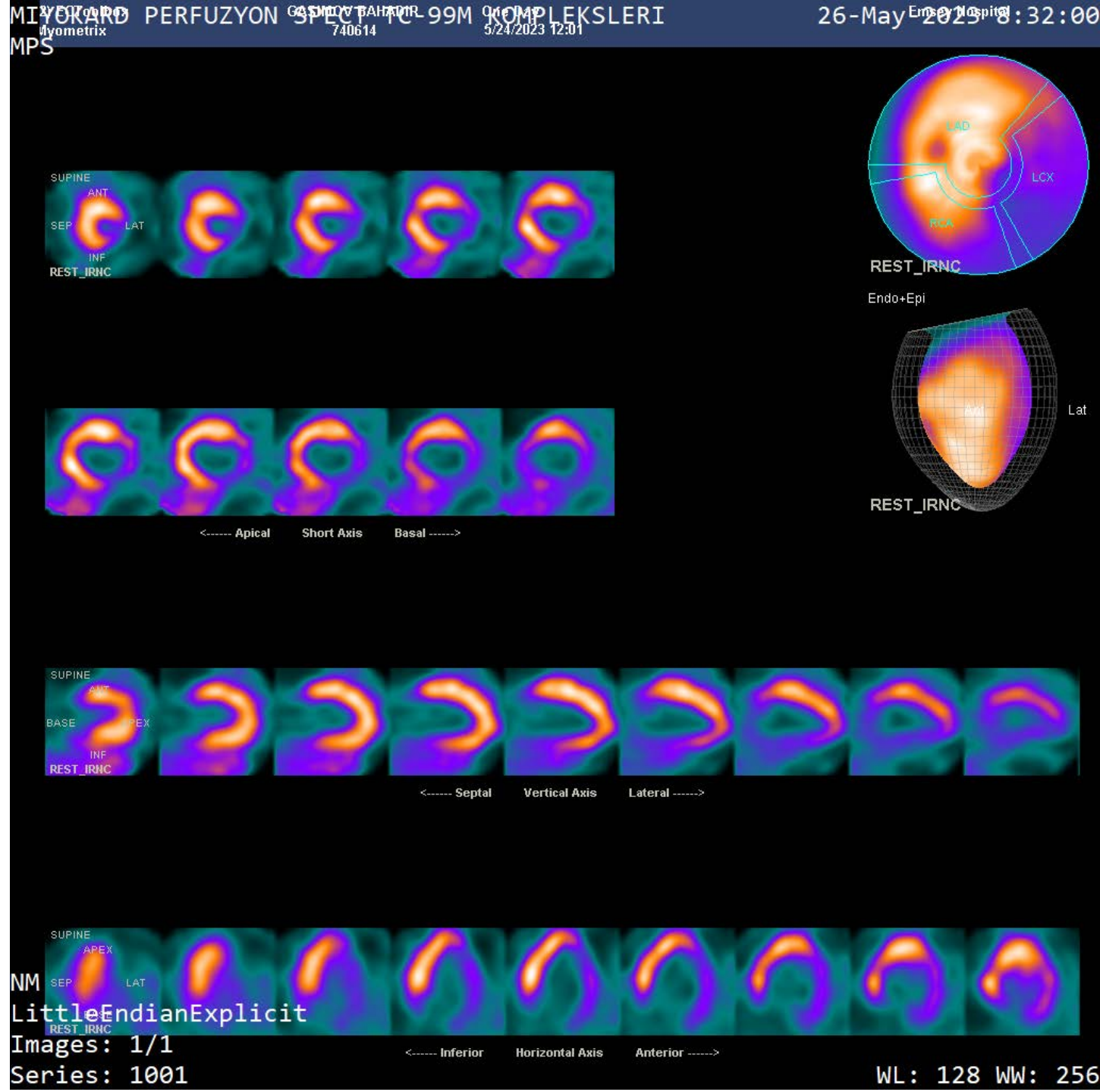
2023 SPECT

Apical infero-lateral wall **nonviable**
Other segments viable



2023 SPECT

Apical infero-lateral wall **nonviable**
Other segments viable



2023 CX PCI



Take Home Message

- Neither ISHEMIA nor other CTO-centric RCTs shows death and MI benefit in stable IHD.
- **LAD CTO PCI** shows more improvement in LVEF compared with OMT alone.
- There is a survival benefit with revascularisation when the ischaemic territory **is greater than 12.5% of at risk myocardium**
- With evident myocardial viability by MPI and CMR revascularization was associated with lower mortality
- Patients with good collaterals (Rentrop grade 2-3) had significantly higher LVEF and shows benefit in survey

