SUPPLEMENTARY APPENDIX

CONTENTS

Page 1

Supplementary Figure I
Effect of physiological pattern of disease on the impact of placebo-controlled PCI on symptom endpoints

Page 2

Supplementary Table I
Sensitivity analysis showing only the results for patients in whom the pre-randomization FFR value was <0.80.

Supplementary Table II
Sensitivity analysis showing only the results for patients in whom the pre-randomization iFR value was <0.89.
Supplementary Figure I

Effect of physiological pattern of disease on the impact of placebo-controlled PCI on symptom endpoints

The association between physiological pattern of CAD as assessed by iFR-pullback and the benefit of PCI over placebo for (A) exercise time, (B) SAQ angina frequency and (C) SAQ derived freedom from angina, after adjustment for baseline FFR values.

$P_{interaction} = 0.797$
Supplementary Table I

| Entry Criterion | PCI (n) | Placebo (n) | Endpoint                  | Focal vs Diffuse P_{interaction} | Focal vs Diffuse P_{interaction} |
|-----------------|--------|-------------|---------------------------|---------------------------------|---------------------------------|
|                 |        |             |                           | Unadjusted for baseline FFR     | Adjusted for baseline FFR       |
| FFR ≤0.80       | 52     | 43          | Stress echo score         | 0.067                           | 0.076                           |
|                 | 62     | 56          | Exercise time             | 0.619                           | 0.646                           |
|                 | 62     | 56          | SAQ Angina Frequency      | 0.691                           | 0.683                           |

Sensitivity analysis showing only the results for patients in whom the pre-randomization FFR value was <0.80.

Abbreviations: FFR = Fractional flow reserve, PCI = Percutaneous coronary intervention, SAQ = Seattle Angina Questionnaire

Supplementary Table II

| Entry Criterion | PCI (n) | Placebo (n) | Endpoint                  | Focal vs Diffuse P_{interaction} | Focal vs Diffuse P_{interaction} |
|-----------------|--------|-------------|---------------------------|---------------------------------|---------------------------------|
|                 |        |             |                           | Unadjusted for baseline iFR     | Adjusted for baseline iFR       |
| iFR ≤0.89       | 46     | 42          | Stress echo score         | 0.030                           | 0.052                           |
|                 | 56     | 56          | Exercise time             | 0.749                           | 0.703                           |
|                 | 56     | 56          | SAQ Angina Frequency      | 0.545                           | 0.568                           |

Sensitivity analysis showing only the results for patients in whom the pre-randomization iFR value was <0.89.

Abbreviations: iFR = instantaneous wave-free ratio, PCI = Percutaneous coronary intervention, SAQ = Seattle Angina Questionnaire