

Supplementary Material 9: GRADE Evaluation of main outcomes

Outcome	Certainty assessment							Effect		Certainty
	Nº of studies	Study design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Relative (95% CI)	Absolute (95% CI)	
Total Myocardial Infarction	15	randomised trials	not serious	serious ^a	not serious	not serious	none	RR 0.86 (0.78 to 0.95)	1 fewer per 1,000 (from 1 fewer to 1 fewer)	⊕⊕⊕○ Moderate ^a
Fatal Myocardial Infarction	6	randomised trials	not serious	not serious	not serious	serious ^b	none	RR 0.87 (0.44 to 1.72)	1 fewer per 1,000 (from 2 fewer to 0 fewer)	⊕⊕⊕○ Moderate ^b
Non-Fatal Myocardial Infarction	13	randomised trials	not serious	not serious	not serious	not serious	none	RR 0.87 (0.79 to 0.97)	1 fewer per 1,000 (from 1 fewer to 1 fewer)	⊕⊕⊕⊕ High
Major Adverse Cardiovascular Event (MACE)	12	randomised trials	not serious	serious ^a	not serious	not serious	none	RR 0.87 (0.80 to 0.93)	1 fewer per 1,000 (from 1 fewer to 1 fewer)	⊕⊕⊕○ Moderate ^a
Stroke	11	randomised trials	not serious	not serious	not serious	not serious	none	RR 0.88 (0.82 to 0.94)	1 fewer per 1,000 (from 1 fewer to 1 fewer)	⊕⊕⊕⊕ High
Coronary Revascularization	6	randomised trials	not serious	serious ^a	not serious	not serious	none	RR 0.85 (0.72 to 0.99)	1 fewer per 1,000 (from 1 fewer to 1 fewer)	⊕⊕⊕○ Moderate ^a
Cardiovascular Mortality	13	randomised trials	not serious	not serious	not serious	not serious	none	RR 0.87 (0.82 to 0.93)	1 fewer per 1,000 (from 1 fewer to 1 fewer)	⊕⊕⊕⊕ High
Heart Failure	10	randomised trials	not serious	not serious	not serious	not serious	none	RR 0.95 (0.75 to 1.19)	1 fewer per 1,000 (from 1 fewer to 1 fewer)	⊕⊕⊕⊕ High
Unstable Angina	8	randomised trials	not serious	not serious	not serious	not serious	none	RR 0.97 (0.87 to 1.08)	1 fewer per 1,000 (from 1 fewer to 1 fewer)	⊕⊕⊕⊕ High

Legend: CI: confidence interval; RR: risk ratio

Explanation:

a. A serious rating for inconsistency was given in view of moderate to high heterogeneity as demonstrated by the I² and p-value for Cochran Q statistic

b. A serious rating for imprecision was given for the wide confidence interval demonstrated.