

Appendix Table 1: Cox-Model

Covariate	Survival HR (95% CI)	p-value	Myocardial Infarction HR (95% CI)	p-value	Repeat PCI/CABG HR (95% CI)	p-value
<b>Treatment</b>						
Medical therapy	referent		referent		referent	
Revascularization	0.76 (0.68-0.84)	<0.001	0.78 (0.72-0.85)	<0.001	0.59 (0.50-0.70)	<0.001
<b>Demographics</b>						
Age, yrs	1.06 (1.05-1.06)	<0.001	1.008 (1.003-1.014)	0.001	0.99(0.99-0.995)	0.001
Male gender	1.18 (1.07-1.30)	0.001	0.94 (0.88-0.998)	0.043	1.05 (0.99-1.12)	0.097
Rural	1.03 (0.90-1.18)	0.678	0.96 (0.86-1.07)	0.436	0.99 (0.90-1.08)	0.795
Income						
5	referent		referent		referent	
1	1.32 (1.18-1.48)	<0.001	1.24 (1.13-1.37)	<0.001	1.01(0.94-1.09)	0.702
2	1.15 (1.02-1.30)	0.023	1.10 (1.03-1.17)	0.004	1.00 (0.92-1.07)	0.916
3	1.04 (0.92-1.16)	0.561	1.05 (0.97-1.14)	0.234	1.03 (0.96-1.11)	0.431
4	1.15 (1.04-1.27)	0.005	0.99 (0.88-1.10)	0.812	0.96 (0.88-1.04)	0.277
<b>Medical comorbidities</b>						
PVD	1.54 (1.40-1.70)	<0.001	1.32 (1.22-1.43)	<0.001	1.02 (0.92-1.14)	0.657
Previous MI	1.03 (0.95-1.12)	0.429	1.88 (1.72-2.06)	<0.001	0.98 (0.92-1.04)	0.459
COPD	1.44 (1.29-1.61)	<0.001	1.09 (0.989-1.20)	0.085	0.99 (0.90-1.08)	0.761
Malignancy	1.26 (1.09-1.46)	0.002	0.78 (0.64-0.95)	0.015	0.99 (0.83-1.18)	0.918
Mean Charlson score	1.24 (1.20-1.29)	<0.001	1.22 (1.17-1.27)	<0.001	1.03 (0.99-1.08)	0.126
<b>Cardiac risk factors</b>						
Diabetes	1.09 (1.00-1.19)	0.053	0.95 (0.85-1.07)	0.395	1.08 (1.01-1.14)	0.021
Hypertension	1.08 (0.94-1.24)	0.289	1.29 (1.11-1.50)	<0.001	1.05 (0.97-1.11)	0.130
Hyperlipidemia	0.81 (0.72-0.91)	<0.001	0.93 (0.84-1.03)	0.153	0.99(0.94-1.05)	0.867
History smoking	1.10 (1.01-1.21)	0.033	1.01 (0.92-1.10)	0.909	1.03 (0.98-1.08)	0.313
<b>Cardiac status/testing</b>						
Native stenosis*						
LM	1.27 (1.12-1.44)	<0.001	1.09 (0.99-1.21)	0.088	0.89 (0.79-1.01)	0.063
Prox LAD	1.11 (1.03-1.19)	0.009	1.05 (0.97-1.12)	0.220	1.16 (1.09-1.23)	<0.001
Mid/distal LAD	0.99 (0.93-1.06)	0.745	1.11 (1.04-1.20)	0.004	1.20 (1.13-1.28)	<0.001
Circumflex	1.17 (1.09-1.27)	<0.001	1.16 (1.07-1.27)	<0.001	1.24 (1.17-1.32)	<0.001
RCA	1.29 (1.21-1.39)	<0.001	1.15 (1.06-1.24)	<0.001	1.33(1.27-1.39)	<0.001
Previous CABG	0.85 (0.76-0.95)	0.005	1.17 (1.08-1.27)	<0.001	0.71 (0.64-0.78)	<0.001
LV function						
≥50%	referent		referent		referent	
≤34%	1.93 (1.62-2.30)	<0.001	1.45 (1.30-1.61)	<0.001	0.83 (0.74-0.93)	0.002
35%-49%	1.36 (1.20-1.53)	<0.001	1.27 (1.17-1.38)	<0.001	0.90 (0.82-0.99)	0.038
NA	0.96 (0.86-1.07)	0.482	1.15 (1.04-1.28)	0.008	0.91 (0.83-1.00)	0.056
Exercise ECG risk						
Low risk	referent		referent		referent	
High risk	1.01 (0.87-1.16)	0.942	1.01 (0.84-1.23)	0.885	1.10(1.02-1.18)	0.012
Uninterpretable	1.33 (1.11-1.58)	0.002	0.94 (0.81-1.09)	0.384	0.90 (0.77-1.04)	0.158
NA	1.71 (1.51-1.93)	<0.001	1.42 (1.28-1.57)	<0.001	1.19(1.11-1.27)	<0.001
Functional imaging risk						
Low risk	referent		referent		referent	
High risk	1.11 (0.99-1.24)	0.075	0.97 (0.87-1.07)	0.501	1.00 (0.92-1.09)	0.998
Unknown/NA	1.14 (1.03-1.27)	0.014	1.03 (0.95-1.12)	0.506	1.12 (1.04-1.21)	0.002
<b>CCS class</b>						
0	referent		referent		referent	
1	0.86 (0.77-0.97)	0.011	0.97 (0.84-1.12)	0.704	1.13 (1.01-1.27)	0.036
2	0.82 (0.73-0.91)	<0.001	1.08 (0.94-1.23)	0.286	1.25(1.15-1.35)	<0.001
3	0.88 (0.79-0.99)	0.027	1.16 (1.04-1.30)	0.006	1.46 (1.34-1.58)	<0.001
4	0.96 (0.79-1.18)	0.724	1.28 (1.06-1.53)	0.009	1.39 (1.16-1.65)	<0.001
<b>Physician-level factors</b>						
Referral physician						
Specialty						
GP/FP	referent		referent		referent	
Cardiology	1.00 (0.91-1.10)	1.000	1.10 (1.01-1.19)	0.022	1.03 (0.96-1.11)	0.368
Internal medicine	0.98 (0.82-1.16)	0.784	1.07 (0.94-1.21)	0.323	1.02 (0.93-1.13)	0.685
Other	1.16 (0.98-1.36)	0.083	1.25 (1.11-1.42)	<0.001	0.97 (0.85-1.10)	0.612

Missing	1.06 (0.96-1.16)	0.278	1.10 (0.97-1.25)	0.150	1.10 (1.03-1.18)	0.007
<b>Hospital-level factors</b>						
Average annual volume per hospital						
Mean Cath volume	1.00 (1.00-1.00)	0.260	1.00 (1.00-1.00)	0.002	1.04 (0.87-1.24)	0.910
Hospital type						
CABG, PCI and Cath	referent		referent		referent	
Cath only	1.12 (0.97-1.29)	0.126	0.97 (0.83-1.12)	0.644	1.04 (0.87-1.24)	0.665
PCI and Cath only	0.92 (0.79-1.08)	0.313	1.14 (0.78-1.67)	0.499	1.13 (0.95-1.34)	0.167

HR=Hazard ratio, CI = confidence interval

CABG= coronary artery bypass grafting; COPD = chronic obstructive pulmonary disease;

MI = myocardial infarction; PCI = percutaneous coronary intervention;

GP/FP=practitioner/family practitioner

PVD = peripheral vascular disease, LM = left main, LAD = left anterior descending,

RCA = right coronary artery, LV = left ventricular, ECG = electrocardiogram,

CCS= Canadian Cardiovascular Society, Cath = catheterization, GP/FP = general

\*Income quintile: 1=lowest, 5 = highest;

† LM if  $\geq 50\%$  stenosis, Prox LAD if  $\geq 70\%$  stenosis, Mid/distal LAD if  $\geq 70\%$  stenosis,

Circumflex if  $\geq 70\%$  stenosis, RCA if  $\geq 70\%$  stenosis

NA implies not done or missing

**Appendix Table 2: Medication Use in Propensity Matched Cohort > 65 years**

Medication	Total (n=13,535)	Medical therapy patients (n=6,753)	Revascularized patients (n=6,782)	p-value
<b>Medications within 3 months pre-index, %</b>				
ACE or ARB	69.0	68.9	69.1	0.781
Anti-platelet agents (clopidogrel)	17.4	16.3	18.5	<0.001
β blockers	68.0	66.8	69.1	0.004
Calcium Channel blockers	38.6	38.3	38.9	0.471
Nitrates	45.5	41.5	49.6	<0.001
Statins	75.3	75.4	75.1	0.607
<b>Medications within 3 months post-index, %</b>				
ACE or ARB	71.7	71.6	71.8	0.757
Anti-platelet agents (clopidogrel)	47.2	19.7	74.5	<0.001
β blockers	68.0	66.8	69.1	<0.001
Calcium Channel blockers	39.9	41.5	38.2	<0.001
Nitrates	32.5	35.0	30.0	<0.001
Statins	83.6	81.7	85.4	<0.001
<b>Medications within 12 months post-index, %</b>				
ACE or ARB	81.4	80.9	81.9	0.826
Anti-platelet agents (clopidogrel)	52.3	27.1	77.3	<0.001
β blockers	76.6	76.3	76.9	0.443
Calcium Channel blockers	47.1	48.5	45.8	<0.001
Nitrates	45.0	47.1	42.8	<0.001
Statins	92.7	91.3	94.1	<0.001

ACE= angiotension converting enzyme inhibitors; ARB= angiotension receptor blockers

**Appendix Table 3: Baseline Characteristics of Propensity Matched COURAGE Cohort**

Covariate	Medical therapy patients (n=4,838)	Revascularized patients (n=4,838)	Total (n=9,676)	Excluded Patients (n=29,455)	p-value*
<b>Treatment</b>					
CABG	0.0	21.0	10.5	25.0	<0.001
PCI	0.0	79.0	39.5	40.0	
Medical therapy	100.0	0.0	50.0	35.0	
<b>Demographics</b>					
Mean age, yrs	65.77 ± 10.17	65.87 ± 10.15	65.82 ± 10.16	66.05 ± 10.35	0.054
Male gender	72.6	72.5	72.5	76.1	0.0
Rural	12.1	12.4	12.2	15.2	
Income					
1	18.8	19.3	19.0	18.4	
2	20.0	19.9	20.0	20.6	
3	20.3	20.2	20.3	20.2	
4	20.7	20.7	20.7	20.7	
5	20.1	19.9	20.0	20.1	
<b>Medical comorbidities</b>					
PVD	8.4	8.3	8.4	9.7	
Previous MI	29.1	28.1	28.6	28.0	
COPD	6.2	6.4	6.3	7.1	
Mean Charlson score	0.80 ± 1.22	0.81 ± 1.28	0.81 ± 1.25	0.88 ± 1.36	
<b>Cardiac risk factors</b>					
Diabetes	45.0	45.5	45.2	43.5	0.0
Hypertension	87.9	88.7	88.3	86.2	<0.001
Hyperlipidemia	82.9	83.3	83.1	80.1	<0.001
History smoking	31.3	31.3	31.3	31.7	0.5
<b>Cardiac status/testing</b>					
Native stenosis**					
LM	0.0	0.0	0.0	17.3	<0.001
Prox LAD	26.8	25.8	26.3	35.4	<0.001
Mid/distal LAD	49.6	49.1	49.3	49.7	0.515
Circumflex	48.2	48.7	48.5	52.5	<0.001
RCA	58.1	58.4	58.2	61.6	<0.001
Had previous CABG					<0.001
Yes	16.3	15.3	15.8	18.7	
No	83.6	84.6	84.1	81.2	
Unknown	0.1	0.1	0.1	0.1	
LV function					<0.001
NA	0.0	0.0	0.0	37.7	
<=34%	0.0	0.0	0.0	6.7	
35%-49%	16.4	15.4	15.9	11.7	
≥50%	83.6	84.6	84.1	43.9	
Exercise ECG risk					<0.001
High risk	33.6	33.1	33.4	25.8	
Low risk	37.2	38.2	37.7	17.4	
Uninterpretable	5.9	6.1	6.0	4.5	
NA	23.2	22.6	22.9	52.3	
Functional imaging risk					<0.001
High risk	41.7	41.9	41.8	28.8	
Low risk	35.6	35.4	35.5	19.0	
Unknown/NA	22.7	22.7	22.7	52.2	
<b>CCS class</b>					
0	14.7	15.6	15.2	16.9	
1	12.3	12.9	12.6	14.5	
2	27.4	26.7	27.1	41.9	
3	45.6	44.8	45.2	23.0	
<b>Physician-level factors</b>					

<b>Referral physician</b>					
Specialty	43.4	47.0	45.2	45.2	
Cardiology	43.4	47.0	45.2	45.2	
Internal medicine	9.5	9.1	9.3	9.3	
GP/FP	27.1	24.7	25.9	25.9	
<b>Hospital-level factors</b>					
Average annual volume per hospital					
Mean Cath volume	$4,143.37 \pm 1,679.79$	$4,139.48 \pm 1,701.16$	$4,141.43 \pm 1,690.42$	$4,076.20 \pm 1,698.09$	0.001
Hospital type					
Cath only	12.0	14.0	13.0	13.3	<0.001
PCI and Cath only	5.7	4.5	5.1	6.4	
CABG, PCI and Cath	82.2	81.5	81.9	80.4	

All covariates are presented as percentages, with the exception of mean values.

Mean values include standard deviation.

CABG= coronary artery bypass grafting; COPD = chronic obstructive pulmonary disease;

MI = myocardial infarction; PCI = percutaneous coronary intervention;

GP/FP=practitioner/family practitioner

PVD = peripheral vascular disease, LM = left main, LAD = left anterior descending,

RCA = right coronary artery, LV = left ventricular, ECG = electrocardiogram,

CCS= Canadian Cardiovascular Society, Cath = catheterization, GP/FP = general

\* p-value for comparison of COURAGE subgroup to excluded patients.

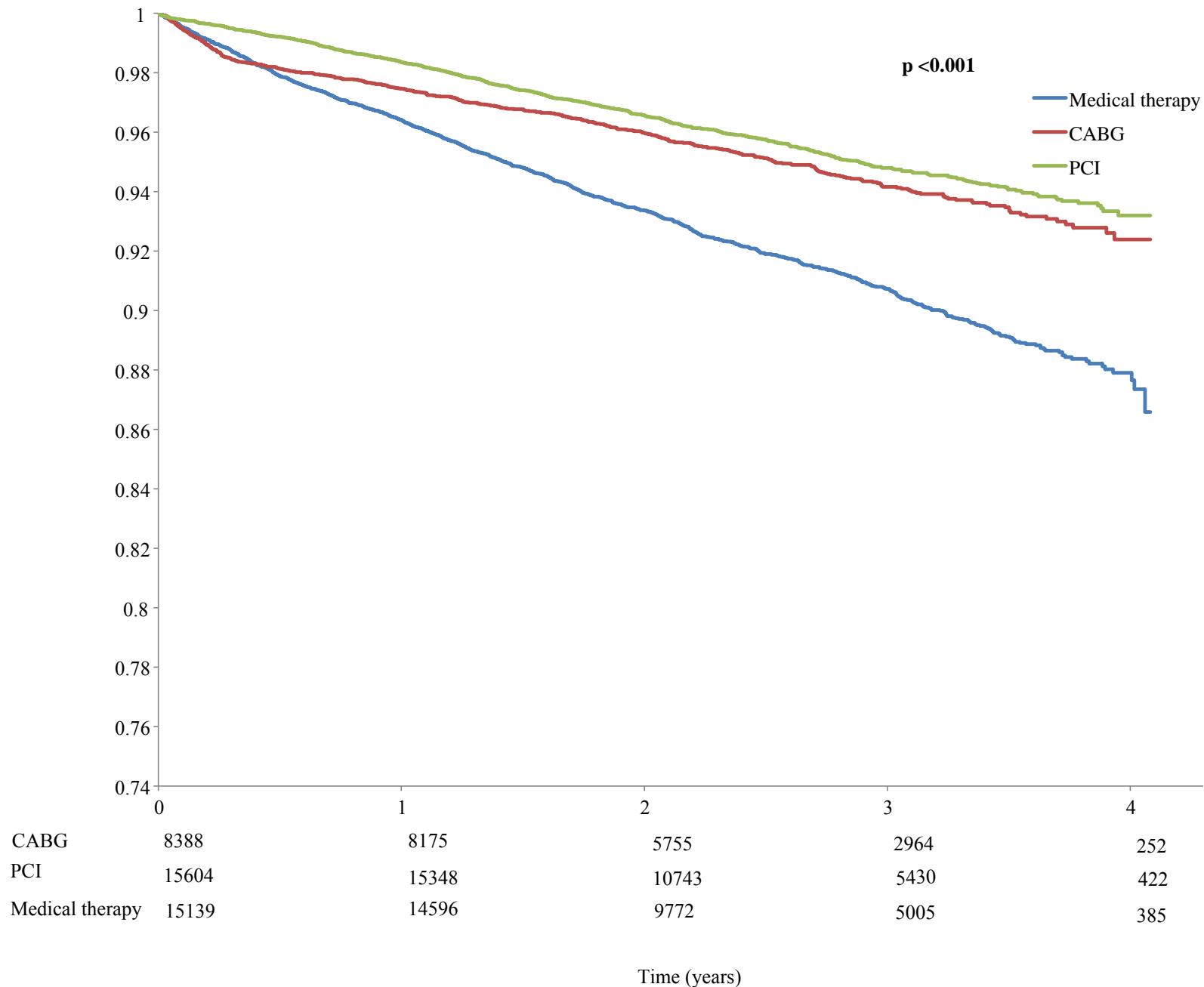
\*\*Income quintile: 1=lowest, 5 = highest;

† LM if  $\geq 50\%$  stenosis, Prox LAD if  $\geq 70\%$  stenosis, Mid/distal LAD if  $\geq 70\%$  stenosis,

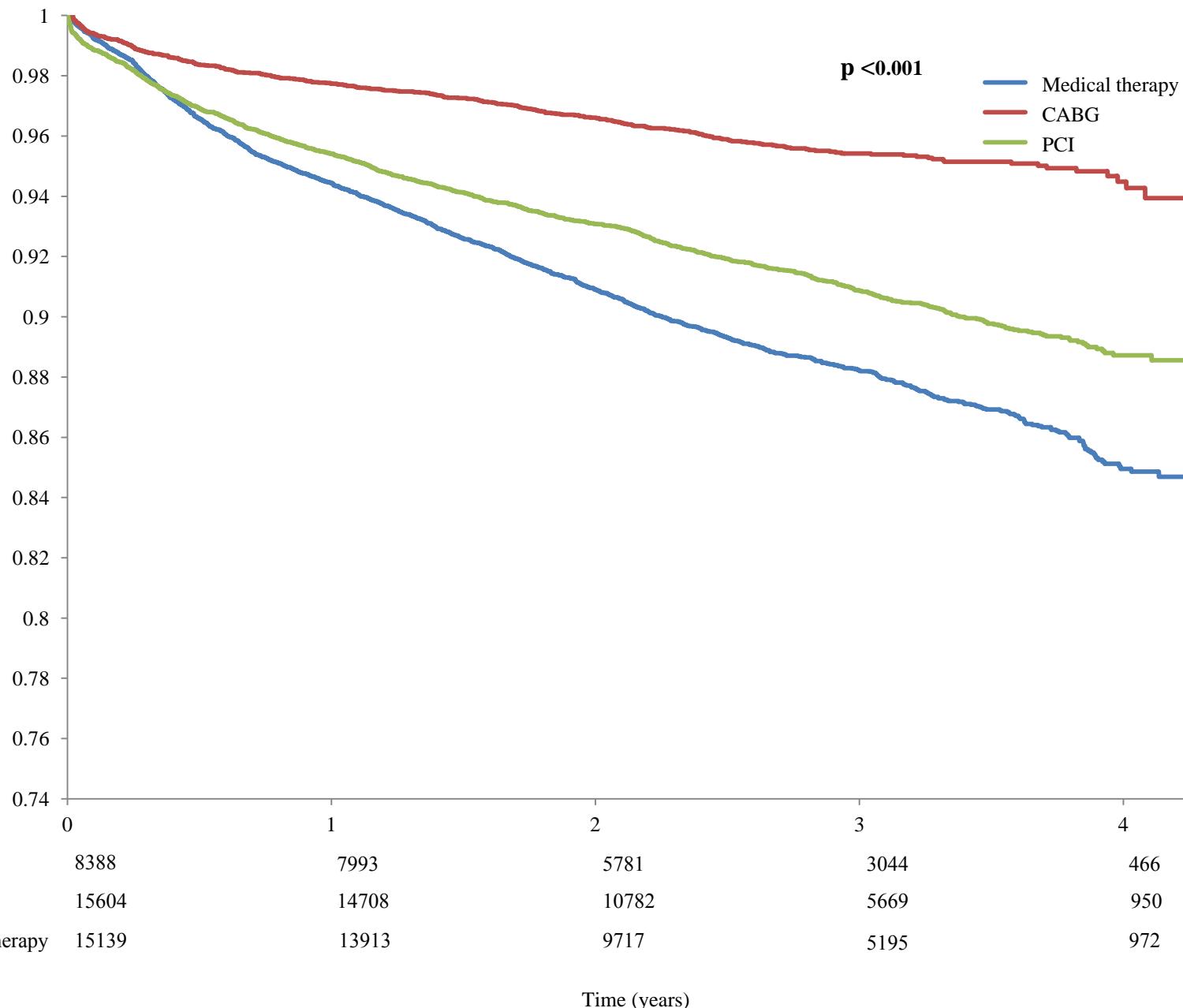
Circumflex if  $\geq 70\%$  stenosis, RCA if  $\geq 70\%$  stenosis

NA implies not done or missing

Appendix Figure 1a: Unadjusted KM Curves for Survival



Appendix Figure 1b: Unadjusted KM Curves for Myocardial Infarction



CABG 8388

PCI 15604

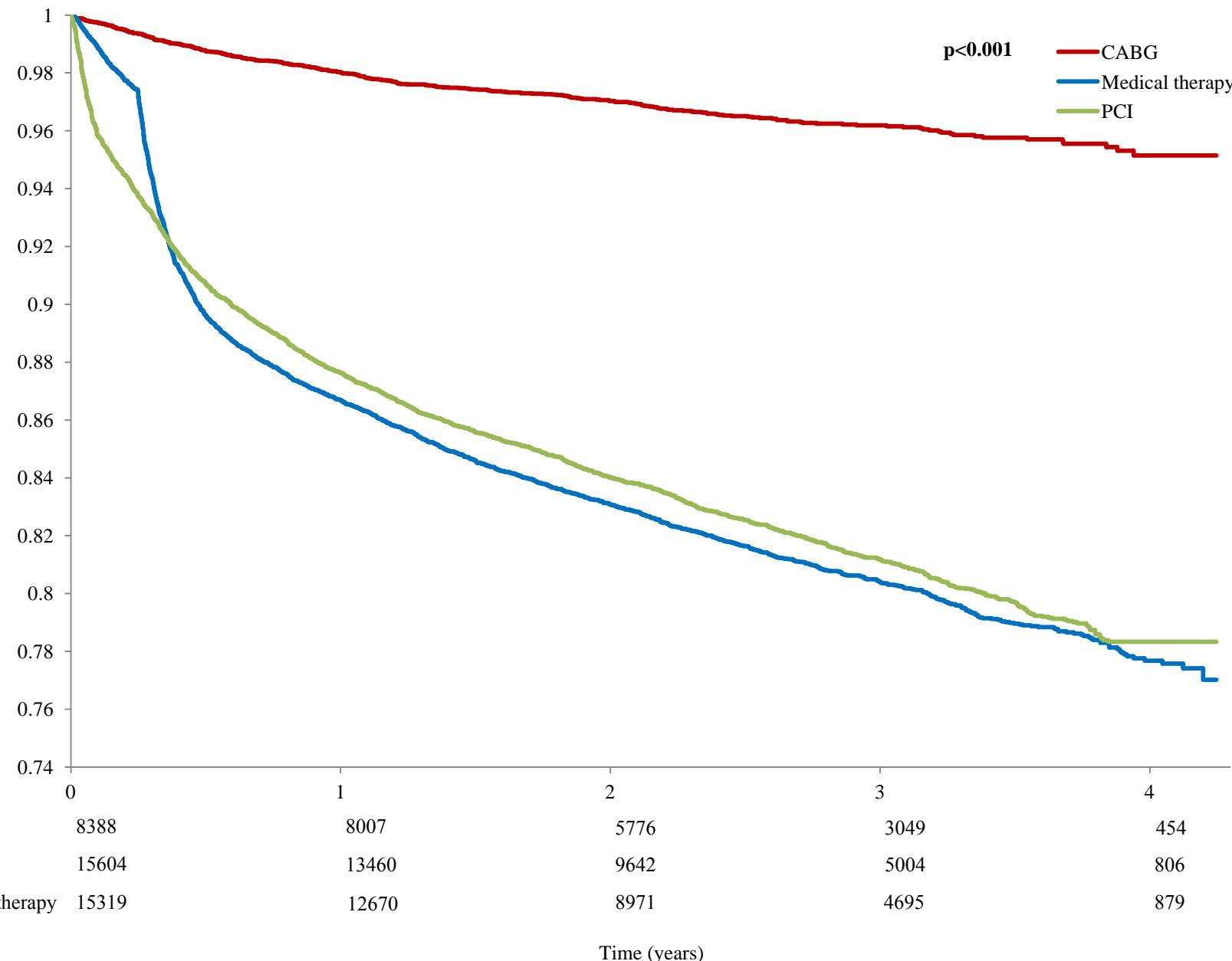
Medical therapy 15139

7993 5781 3044 466

14708 10782 5669 950

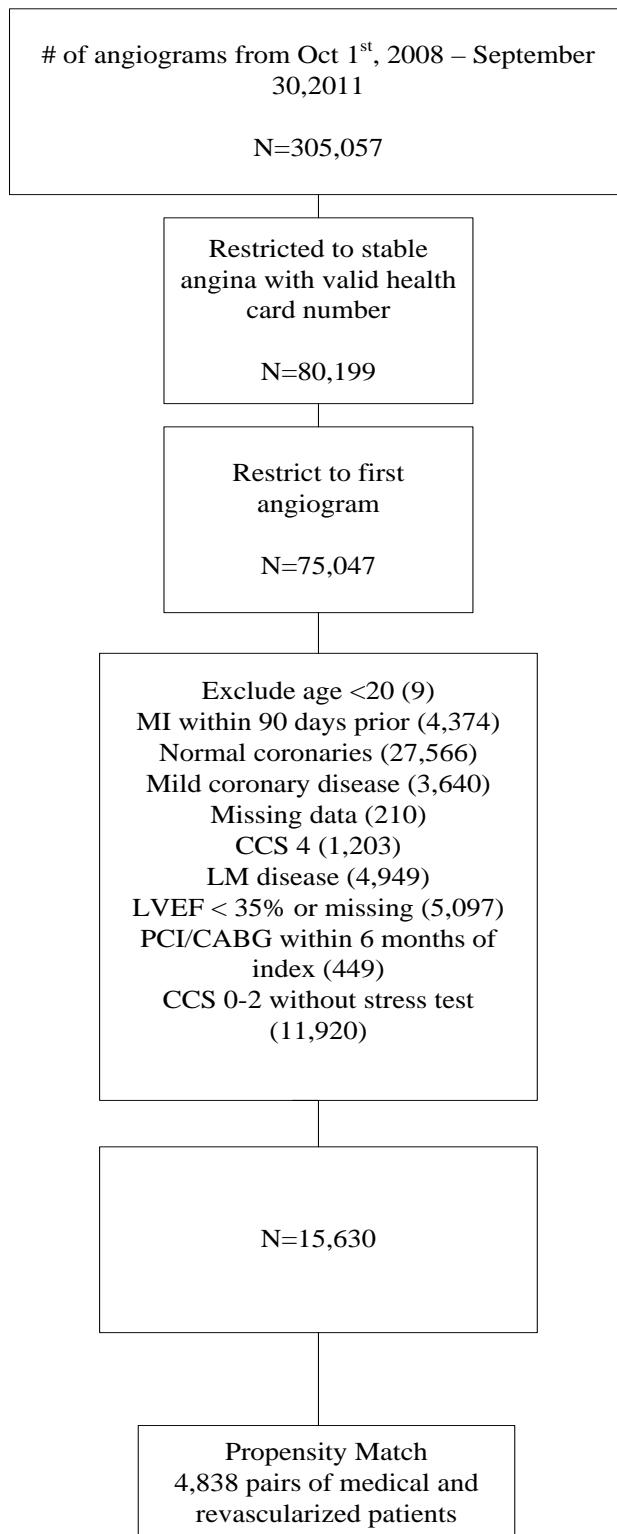
13913 9717 5195 972

Appendix Figure 1c: Unadjusted KM Curves for Repeat Revascularization

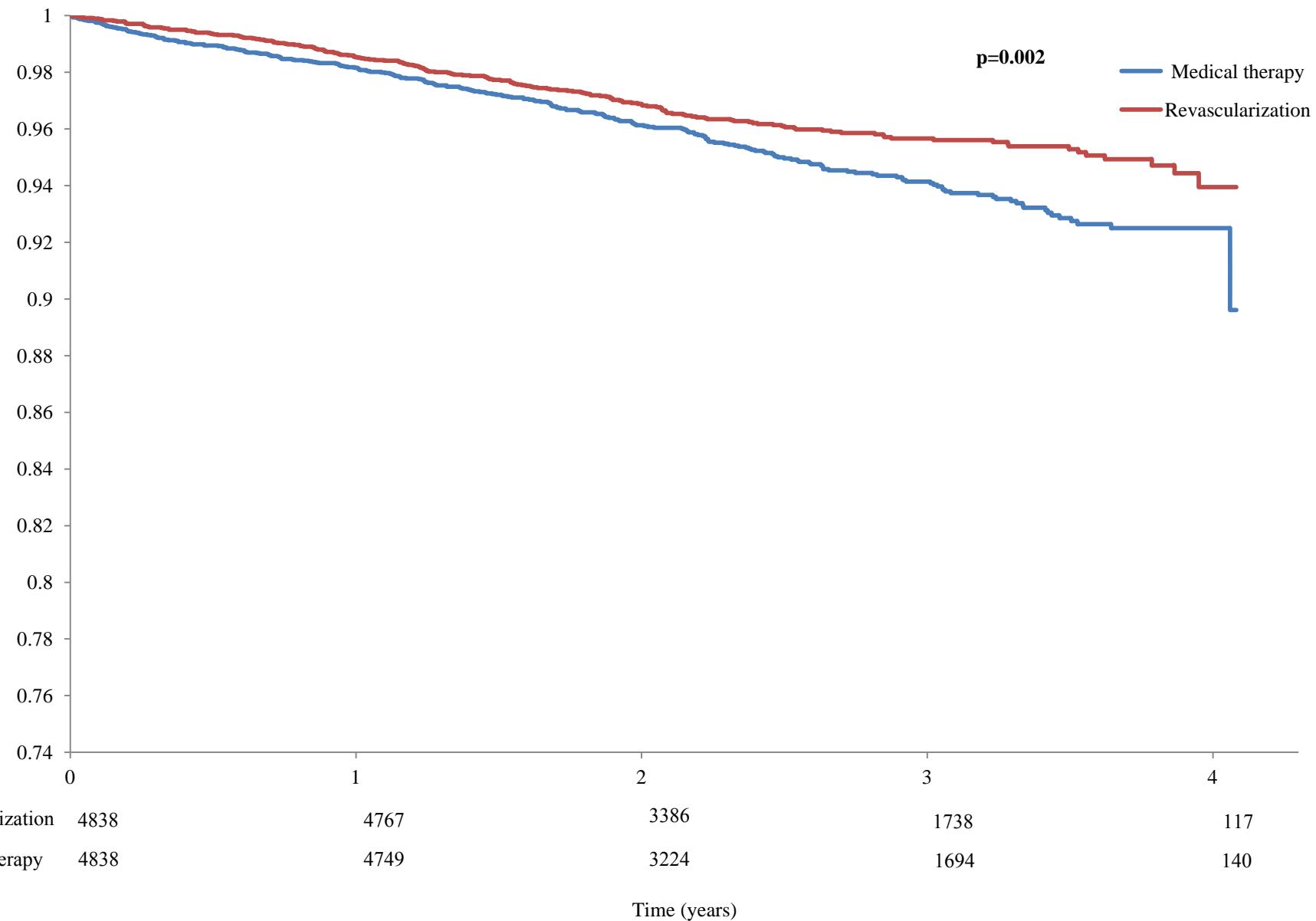


CABG	8388	8007	5776	3049	454
PCI	15604	13460	9642	5004	806
Medical therapy	15319	12670	8971	4695	879

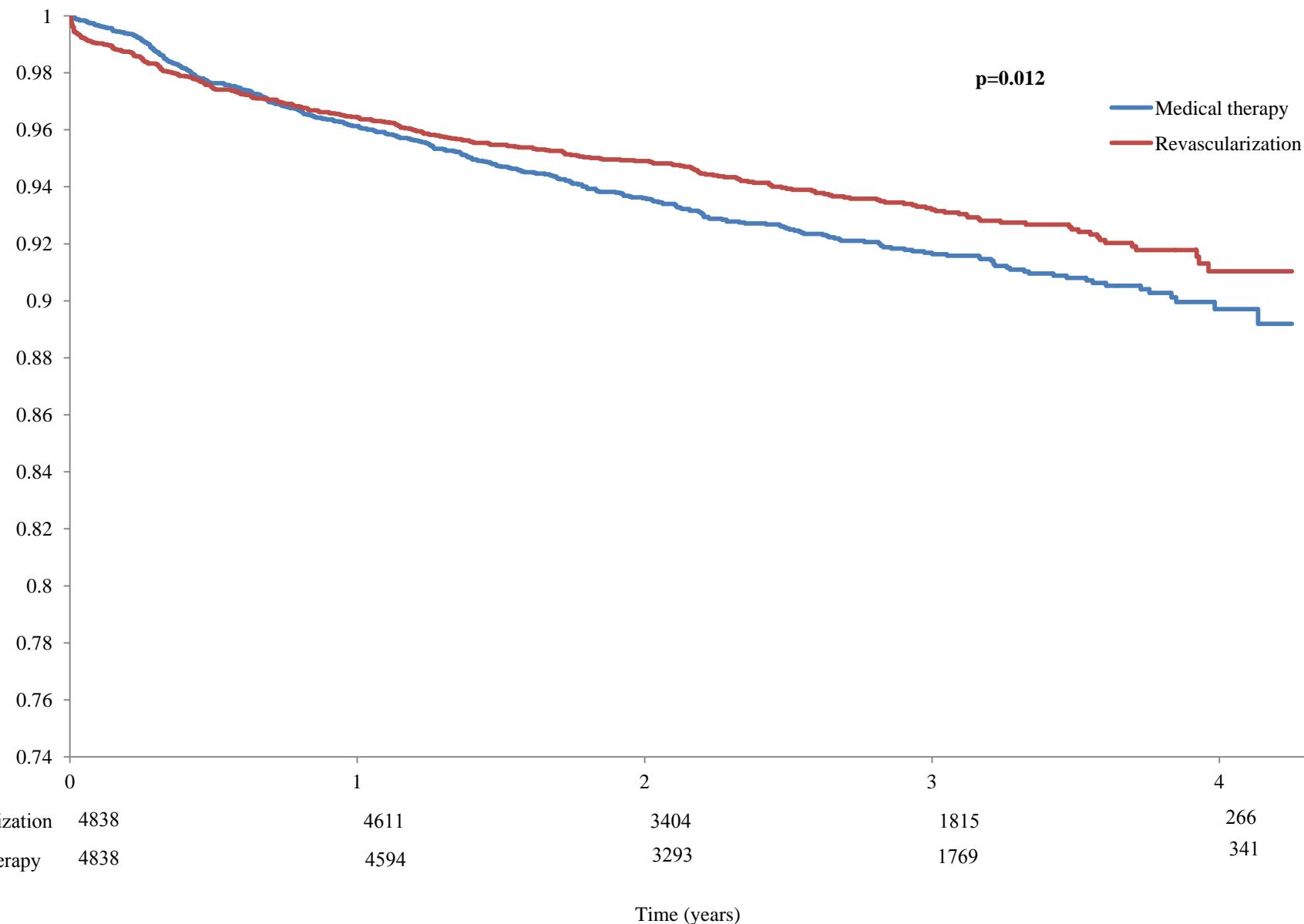
Appendix Figure 2: COURAGE Cohort



Appendix Figure 3a: COURAGE Propensity-Matched Cohort KM Curves for Death



Appendix Figure 3b: COURAGE Propensity-Matched Cohort KM Curves for Myocardial Infarction



Appendix Figure 3c: COURAGE Propensity-Matched Cohort KM Curves for Repeat Revascularization

