Supplementary Online Content

Maddox TM, Stanislawski MA, Grunwald GK, et al. Nonobstructive coronary artery disease and risk of myocardial infarction. *JAMA*. doi:10.1001/jama.2012.9630. 10.1001/jama.2014.14681

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This supplementary material has been provided by the authors to give readers additional information about their work.

eTable. Kaplan-Meier Estimates of 1-Year MI, All-Cause Mortality, and Combined Outcome Rates by Mild Non-Obstructive, Moderate Non-Obstructive, and **Obstructive CAD**

	No Apparent CAD ^a	Mild Non- Obstructive CAD	Moderate Non- Obstructive CAD	1-Vessel Obstructive CAD	2-Vessel Obstructive CAD	3- Vessel/Left Main Obstructive CAD
1-Year MI ^b ,	0.11	0.39 (0.20,	0.50 (0.30,	1.19 (1.00,	2.25 (1.90,	2.50 (2.10,
%, (95%	(0.10,	0.60)	0.80)	1.50)	2.70)	3.00)
CI°)	0.20)					
1-Year	1.38	1.89 (1.50,	2.17 (1.70,	2.24 (1.90,	3.40 (2.90,	4.33 (3.80,
Mortality,	(1.10,	2.30)	2.70)	2.60)	4.00)	4.90)
%, (95%	1.70)					
CI)						
1-Year	1.48	2.21 (1.80,	2.59 (2.10,	3.11 (2.70,	5.05 (4.50,	6.24 (5.60,
Combined	(1.20,	2.70)	3.20)	3.50)	5.70)	6.90)
Outcomes,	1.80)					
%, (95%	-					
CI)						

^aCAD – coronary artery disease; ^bMI – myocardial infarction; ^cCI – confidence interval

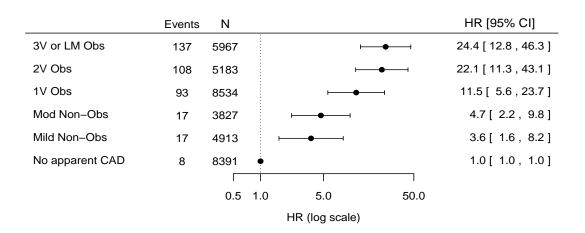
No apparent CAD - coronary stenoses <20% and/or luminal irregularities Mild Non-Obstructive CAD - maximal coronary stenosis of 20-49%

Moderate non-obstructive CAD - maximal coronary stenosis of 50-69%

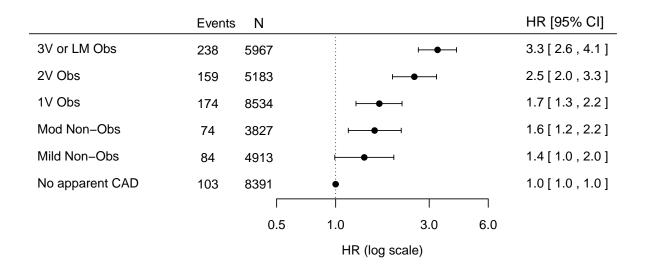
Obstructive CAD -any stenosis ≥50% in the left main coronary artery and/or ≥70% in any other coronary artery. Patients were categorized by CAD severity in a single, double, or triple-vessel distribution. Patients with ≥50% left main artery stenosis included in 3-vessel obstructive CAD.

eFigure 1. Adjusted Cox Model Results for 1-Year MI, Mortality, and Combined MI or Mortality by Mild Non-Obstructive, Moderate Non-Obstructive, and Obstructive CAD, Relative to No Apparent CAD

MI



Mortality



Combined MI or Mortality

	Events	s N		HR [95% CI]		
3V or LM Obs	357	5967	⊢•	4.7 [3.8 , 5.8]		
2V Obs	255	5183	⊢ •	3.8 [3.0 , 4.8]		
1V Obs	258	8534	⊢	2.3 [1.8 , 3.0]		
Mod Non-Obs	90	3827	⊢	1.8 [1.3 , 2.4]		
Mild Non-Obs	100	4913	⊢	1.6 [1.1 , 2.1]		
No apparent CAD	110	8391	•	1.0 [1.0 , 1.0]		
			İ	٦		
		0.5	1.0 3.0	6.0		
		HR (log scale)				

No apparent CAD - coronary stenoses <20% and/or luminal irregularities

Mild Non-Obstructive CAD - maximal coronary stenosis of 20-49%

Moderate non-obstructive CAD - maximal coronary stenosis of 50-69%

Obstructive CAD -any stenosis ≥50% in the left main coronary artery and/or ≥70% in any other coronary artery. Patients were categorized by CAD severity in a single, double, or triple-vessel distribution. Patients with ≥50% left main artery stenosis included in 3-vessel obstructive CAD.

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eFigure 2. Adjusted Cox Model Results for 1-Year MI, Mortality, and Combined MI or Mortality by CAD Extent, Stratified by Diabetes, Relative to No Apparent CAD

			М	I			
	Diabetes	Events	s N		HR [95% CI]		
No apparent CAD	Non-Diabetic	4	5670	•	1.0 [1.0 , 1.0]		
No apparent CAD	Diabetic	4	2721	•	1.0 [1.0 , 1.0]		
1V Non-Obs	Non-Diabetic	5	2792	⊢	2.3 [0.6 , 9.6]		
1V Non-Obs	Diabetic	5	1854	⊢	1.7 [0.5 , 6.4]		
2V Non-Obs	Non-Diabetic	5	1456	──	4.4 [1.1 , 17.9]		
2V Non-Obs	Diabetic	8	1149	⊢	4.4 [1.5 , 12.9]		
3V Non-Obs	Non-Diabetic	1	591 ⊢	• · · · · · · · · · · · · · · · · · · ·	2.0 [0.2 , 17.9]		
3V Non-Obs	Diabetic	5	542	⊢	5.5 [1.7 , 17.8]		
1V Obs	Non-Diabetic	59	5385	⊢	12.8 [4.7 , 34.6]		
1V Obs	Diabetic	42	4026	⊢	5.9 [2.0 , 17.5]		
2V Obs	Non-Diabetic	50	2899	⊢	19.7 [7.1 , 54.7]		
2V Obs	Diabetic	60	2553	⊢	13.4 [4.6 , 39.0]		
3V or LM Obs	Non-Diabetic	60	3182	⊢	22.6 [8.7 , 58.5]		
3V or LM Obs	Diabetic	77	2854	⊢	16.1 [6.2 , 42.2]		
			Г				
0.2 1.0 5.0 60.0							
	HR (log scale)						

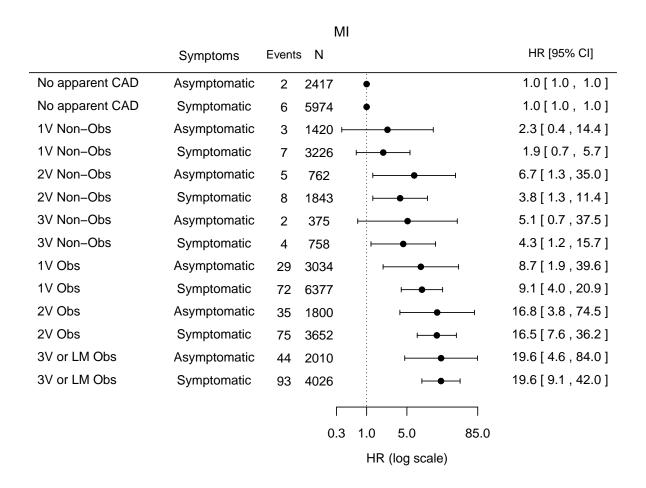
Mortality

	Diabetes	Events	N		HR [95% CI]	
No apparent CAD	Non-Diabetic	66	5670	•	1.0 [1.0 , 1.0]	
No apparent CAD	Diabetic	37	2721	÷	1.0 [1.0 , 1.0]	
1V Non-Obs	Non-Diabetic	51	2792	-	1.4 [1.0 , 2.2]	
1V Non-Obs	Diabetic	34	1854	⊢	1.3 [0.8 , 2.1]	
2V Non-Obs	Non-Diabetic	23	1456	—	1.3 [0.8 , 2.0]	
2V Non-Obs	Diabetic	12	1149	-	0.7 [0.3 , 1.4]	
3V Non-Obs	Non-Diabetic	16	591	├	2.0 [1.2 , 3.5]	
3V Non-Obs	Diabetic	11	542 H	•	1.2 [0.6 , 2.4]	
1V Obs	Non-Diabetic	97	5385	⊢● →	1.9 [1.4 , 2.6]	
1V Obs	Diabetic	95	4026	⊢	1.8 [1.1 , 2.9]	
2V Obs	Non-Diabetic	77	2899	⊢	2.8 [2.0 , 4.0]	
2V Obs	Diabetic	87	2553	⊢	2.6 [1.7 , 4.0]	
3V or LM Obs	Non-Diabetic	115	3182	⊢	3.6 [2.6 , 4.9]	
3V or LM Obs	Diabetic	124	2854	⊢	3.1 [2.0 , 4.7]	
				<u> </u>		
			0.3	1.0 2.0 6.0		
	HR (log scale)					

Combined MI or Mortality

	Diabetes	Events	N		HR [95% CI]	
No apparent CAD	Non-Diabetic	70	5670	•	1.0 [1.0 , 1.0]	
No apparent CAD	Diabetic	40	2721	•	1.0 [1.0 , 1.0]	
1V Non-Obs	Non-Diabetic	55	2792	——	1.5 [1.0 , 2.2]	
1V Non-Obs	Diabetic	38	1854 H	•	1.3 [0.8 , 2.1]	
2V Non-Obs	Non-Diabetic	27	1456	—	1.4 [0.9 , 2.2]	
2V Non-Obs	Diabetic	20	1149 ⊢	-	1.1 [0.6 , 1.8]	
3V Non-Obs	Non-Diabetic	17	591	├	2.0 [1.2 , 3.4]	
3V Non-Obs	Diabetic	15	542	<u> </u>	1.6 [0.9 , 2.8]	
1V Obs	Non-Diabetic	151	5385	⊢	2.5 [1.8 , 3.4]	
1V Obs	Diabetic	133	4026	⊢	2.2 [1.4 , 3.3]	
2V Obs	Non-Diabetic	122	2899	⊢•	3.7 [2.7 , 5.1]	
2V Obs	Diabetic	140	2553	⊢	3.7 [2.5 , 5.7]	
3V or LM Obs	Non-Diabetic	170	3182	⊢	4.6 [3.4 , 6.1]	
3V or LM Obs	Diabetic	188	2854	⊢•	4.2 [2.9 , 6.1]	
				<u>:</u>		
0.5 1.0 2.0 7.0						
	HR (log scale)					

eFigure 3. Adjusted Cox Model Results for 1-Year MI, Mortality, and Combined MI or Mortality by CAD Extent, Stratified by Anginal Symptoms, Relative to No Apparent CAD



Mortality HR [95% CI] **Events** Ν **Symptoms** 1.0 [1.0 , 1.0] No apparent CAD Asymptomatic 2417 36 Symptomatic 1.0 [1.0 , 1.0] No apparent CAD 67 5974 1V Non-Obs 1.4 [0.8 , 2.4] Asymptomatic 31 1420 1V Non-Obs Symptomatic 1.4 [0.9 , 2.2] 54 3226 Asymptomatic 2V Non-Obs 17 762 1.3 [0.7 , 2.3] 2V Non-Obs Symptomatic 0.8 [0.5 , 1.4] 18 1843 3V Non-Obs 1.4 [0.7 , 2.7] Asymptomatic 9 375 3V Non-Obs Symptomatic 1.8 [1.2 , 2.9] 18 758 1V Obs Asymptomatic 2.0 [1.2 , 3.4] 88 3034 1V Obs Symptomatic 104 6377 1.8 [1.2 , 2.7] 2V Obs Asymptomatic 2.8 [1.8 , 4.4] 74 1800 2V Obs 2.7 [1.8 , 4.0] Symptomatic 90 3652 3V or LM Obs Asymptomatic 2.8 [1.7 , 4.4] 85 2010 3V or LM Obs Symptomatic 4026 3.8 [2.8 , 5.3] 154 2.0 0.5 1.0 6.0 HR (log scale)

Combine MI or Mortality

