

Supplementary Online Content

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eTable 1. Site Personnel

eTable 2. Committees, CCC, Study Centers Personnel

eTable 3. Ischemia Definitions

eTable 4. Randomized Participant Baseline Vital Signs and Laboratory Values Intent-to-Treat Population

eTable 5. Randomized Participant Baseline Medications Intent-to-Treat Population

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

**eTable 1
Site Personnel**

Country (No. Randomizations)	Investigator(s)	Study Coordinator(s)	City & State (if applicable)	Institution (No. Randomizations)
*United States (853)				
Country Leader				
David J. Maron, MD				
Regional Leader for VA Sites				
William E. Boden, MD				
	Kreton Mavromatis, MD	John Doan, MD	Decatur, GA	Atlanta VA Medical Center (139)
	Jason Linefsky, MD	Raven Lee, CCRP		
		Risha Patel		
	Todd Miller, MD	So Yang Cho	Rochester, MN	Mayo Clinic (50)
		Susan Milbrandt		
		Dawn Shelstad		
	Subhash Banerjee, MD	Preeti Kamath, BDS, MHA, CCRP	Dallas, TX	V.A. North Texas Health Care System (35)
		Ishita Tejani, BDS, MS, MSPH		
	Harmony R. Reynolds, MD	Stanley E. Cobos, BA	New York, NY	NYU Langone Medical Center- Bellevue Hospital (26)
	Jonathan D. Newman, MD, MPH	Kirsten J. Quiles, MS		
	Sripal Bangalore, MD	Raven R. Dwyer, MPH		
	Robert M. Donnino, MD	Dalisa Espinosa, MBS		
	Lawrence M. Phillips, MD			
	Muhamed Saric, MD, PhD			
	Khaled Abdul-Nour, MD	Allison Schley, BS	Detroit, MI	Henry Ford Health System (21)
		Heather Golden		
	Peter H. Stone, MD	Hermine Osseni, MS	Boston, MA	Brigham & Women's Hospital, Harvard Medical School (21)
		Charlene Wiyarand		
		Peter Douglass, BA		
		Hayley Pomeroy, BA		
		Alexandra Craft, BA		
		Bethany Harvey, BA		
	James J. Jang, MD	Olivia Anaya	San Jose, CA	Kaiser Permanente San Jose (18)
	Gennie Yee, MD	Phoebe Goold, RN		
	Steven Weitz, MD	Steven Giovannone	Schenectady, NY	Cardiology Associates of Schenectady P.C. (17)
		Lori Pritchard, RN		

	Suzanne Arnold, MD	Rosann Gans, RN	Kansas City, MO	Saint Luke's Hospital (17)
	James Henry O'Keefe, Jr, MD (PI from 2012-2016)	Paul Kennedy, RN		
	Michael D. Shapiro, DO	Shobana Ganesan, PhD	Portland, OR	Oregon Health & Science University (17)
		David Schlichting, LPN		
		Aynun Naher		
	Mohammad El-Hajjar, MD		Albany, NY	Albany Medical Center Hospital (16)
	Mandeep S. Sidhu, MD, MBA			
	Steven A. Fein, MD	Wendy L. Stewart, MS		
	Mikhail T. Torosoff, MD, PhD	Kristin M. Salmi, BS		
	Radmila Lyubarova, MD			
	Sulagna Mookherjee, MD			
	Krzysztof Drzymalski, MD			
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	Rizwan A. Siddiqui, MD	Rebekah R. Herrmann, RN		
	Areef Ishani, MD			
	Ronnell A. Hansen, MD			
	Michel Georges Khouri, MD	Kristine Arges	Durham, NC	Duke University Medical Center (15)
		Melissa LeFevre		
		Jennifer Tomfohr		
	Jonathan L. Goldberg, MS, MD	Kimberly Ann Byrne	Cleveland, OH	Louis Stokes Cleveland Veterans Affairs Medical Center (14)
		Taissa Zappernick		
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		Meghana Kakade		
		Patricia Mieses		
		Stanley E. Cobos, BA	Brooklyn, NY	NYU-HHC Woodhull Hospital (12)
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	Philip Rogal, MD	Sherron C. Crook		
	Christopher McFarren, MD			
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		Saadat Khan		
		Mahmoud Mohamed		
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		Kathy E. Siegel, RN		
		Susan Derbyshire		
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		Gerald Leonard		
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		Nora Marchelletta		
		Kristina Wippler		
	David Booth, MD	Yvonne Taul, RN	Lexington, KY	University of Kentucky (8)
	Steve Leung, MD	Jennifer Isaacs, MS		
	Ahmed Abdel-Latif, MD, PhD	Viktoria Bulkley, RN		
	Hassan Reda, MD	Caroline Rodgers		
	Khaled Ziada, MD			
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		Kamalakar Surineni		
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	Jaekyeong Heo, MD			
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	Brian P. Shapiro, MD			
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		Pamela Julian			
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		Reyna Bhandari		
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	Claudia P Hochberg, MD	Paula Beardsley	Boston, MA	Boston Medical Center (4)
		Denise Fine		
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	Ray Wyman, MD	Joy Burkhardt, CCRP	Torrance, CA	Torrance Memorial Medical Center (4)
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	Amer Zarka, MD	Maria Aguirre		
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		Diana Parra		
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	Thomas Haldis, DO	Catherine Weick, BSRT(R)(VI)	Fargo, ND	Sanford Health (4)
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		Casey Riedberger		
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		Kirsten J. Quiles, MS		
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		Kimberly Miller-Cox, RN		
		Amy Ollinger, RN		
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	Jeffrey C. Milliken, MD	Andrea M. Lundeen, MA		
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	Kevin T. Harley, MD	Eduardo Hernandez-Rangel, MD		
	Michael A. Gibson, MD	Pam Singh		
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	Rita Coram, MD	Anne Marie Webb, BSN	Louisville, KY	University of Louisville (3)
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	Sabu Thomas, MD, MSc	Angela Kim, BS	Rochester, NY	University of Rochester (3)
	Ronald G Schwartz, MD, MS	Patrick Wilmot, BS		
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	Mahfouz El Shahawy, MD	Ramona Stevens	Sarasota, FL	Cardiovascular Center of Sarasota (3)
	James Stafford, MD	Loriane Black	Baltimore, MD	University of Maryland Medical Center (3)
	William B. Abernethy, MD	Amber B. Hull, RN	Asheville, NC	Asheville Cardiology Associates (3)
		Olivia J. Lim, RN		
		Helen C. Tucker		
		Natasha C. Putnam, RN		
		Linda L. Hall		
		Tia Cauthren		
		Trish Tucker		
	Andrew Zurick, MD	Hollie Horton	Nashville, TN	

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	Thomas M. Meyer, MD	Joyce R. White, MSN NP-C	Lynchburg, VA	Stroobants Cardiovascular Center (3)
	Ronald G. Morford, MD	Cynthia Baumann, RN		
	Bruce Rutkin, MD	Vidya Seeratan	Manhasset, NY	Northwell Health - Manhasset (3)
	Sabahat Bokhari, MD	Magnolia Jimenez	New York, NY	Columbia University Medical Center (3)
	Seth I. Sokol, MD	Cidney Schultz, RN	Bronx, NY	Jacobi Medical Center (3)
	Jay Meisner, MD	Jeanne Russo, RN		
	Ihab Hamzeh, MD		Houston, TX	Baylor College of Medicine (3)
	Arunima Misra, MD	Zohra Huda, RN, BSN, CCRP		
	Matthew Wall Jr., MD	Araceli Boan		
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	Thomas J. Mulhearn, MD			
	Arnold P. Good, MD	Beth A. Archer, BSN, RN	Columbus, OH	Ohio Health Grant Medical Center (3)
		Julia S. Dionne, BA		
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		Lindsey N. Sikora, BSN, RN		
		Jennifer H. Czerniak, RN		
		Jennifer A. Mull, MSN, RN		
		Elizabeth Ferguson		
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		Lori Christensen		
		Holly Park		
	Robert Chilton, MD	Joan Hecht	San Antonio, TX	Audie Murphy V.A. (2)
	Patricia K. Nguyen, MD	Davis Vo, BS	Palo Alto, CA	VA Palo Alto Healthcare System (2)
		James Hirsch		
	Matthew Jezior, MD	Jody Bindeman	Bethesda, MD	Walter Reed National Military Medical Center (2)
		Sara Salkind		
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		Lori-Ann Desimone, BSN		
	Paul C. Gordon, MD	Lina Felix-Stern		
	Thomas Crain, MD	Jassira Gomes		

		Catherine Gordon, BSN		
	Robert Stenberg, MD	Aimee Mann	Johnstown, PA	Conemaugh Valley Memorial Hospital (2)
		Theresa McCreary		
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		Raven R. Dwyer, MPH		
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		Kirsten J. Quiles, MS		
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		Dalisa Espinosa, MBS		
		Kirsten J. Quiles, MS		
	George J. Juang, MD	Candace Gopaul, BS	Brooklyn, NY	Coney Island Hospital (2)
		Karen Hultberg		
		Tauqir Huk		
		Afshan Hussain		
	Mohammed Al-Amoodi, MD	Yesenia Zambrano, BS	Yuma, AZ	Yuma Regional Medical Center (2)
		Sarah Medina Rodriguez		
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	David Wohns, MD	Abbey Mulder, RN	Grand Rapids, MI	Spectrum Health (2)
		Stacie Van Oosterhout, MEd		
	Ellis W. Lader, MD	Martha Meyer, RN, MSN	Kingston, NY	Mid Valley Cardiology (1)
	Michael Mumma, MD	Nancy L. Clapp, RN, BA, CCRC	Sarasota, FL	Sarasota Memorial Hospital (1)
		Heather Barrentine		
	Lekshmi Dharmarajan, MD	Jenne M. Jose, PA	Bronx, NY	NYU-HHC Lincoln Medical and Mental Health Center (1)
		Stanley E. Cobos, BA		
		Raven R. Dwyer, MPH		
		Dalisa Espinosa, MBS		
		Kirsten J. Quiles, MS		
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		Linda Schwarz, RN		
	Thomas R. Downes, MD (till Dec. 2016)	Scott M. Kaczowski	Loveland, CO	Medical Center of the Rockies (1)
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		Joel Klitch		
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	Kozhaya Sokhon, MD	Jessica Waldron	Sugar land, TX	Medicus Alliance Clinical Research Org., Inc. (1)
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		Lucilla Garcia		
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		Anjali Anand, MSc	Calicut	Government Medical College (208)
	Sajeev Chakanalil Govindan, MD, DNB, DM, PhD	Janitha Raj, B.Tech		
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	Sharad Chandra, DM	Shivali Patel		
		Suman Singh, PG		
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	S.Ramakrishnan, DM			
	Sandeep Seth, DM			
	Rakesh Yadav, DM			
	Sandeep Singh, DM			
	Ambuj Roy, DM			
	Neeraj Parakh, DM			
	Sunil Kumar Verma, DM			
	Rajiv Narang, DM			
	Sundeep Mishra, DM			
	Nitish Naik, DM			
	Gautam Sharma, DM			
	Shiv Kumar Choudhary, M.Ch			
	Chetan Patel, DNB			
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	Sanjeev Sharma, MD			
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		Ashok Seth, MD		
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	Atul Mathur, MD	Atul Verma, MD		
	Upendra Kaul, MD	Mona Bhatia, MD		
		Ankush Sachdeva, MD		
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		Nungshi Jungla, RN		
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		Patricia Blaise		
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		Camila Thais de Souza Ormundo		
		Carla Vicente		
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Poland (333)				
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	Edyta Kaczmarska, MD PhD			
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		Anna Fojt, MD		

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		Alexandra Vazquez		
China (246)				
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		Junqing Yang			
		Qi Zhong			
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		Zhulin Zhang			

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		Nier Zhong			
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		Xin Su			
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	Ana S. Mladenovic, MD, PhD			
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Jamie Rankin, MBBS (<i>past</i>)					
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	Elise van Dongen, MD	Elisabeth, M. Janzen, RN		
	Sander R. Niehe, MD			

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	Ana Santana, MD			
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	Lidia Sousa, MD			
	Hugo Marques, MD			
	Lino Patricio, MD	Mafalda Selas		
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	Pedro Rio, MD	Cláudia Freixo		
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	Rui Ferreira, MD			
	Tiago Silva, MD			
	Ines Rodrigues, MD			
	Pedro Modas, MD			
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	Jose Fragata, MD			
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	Angelo Nobre, MD	Maria Inês Caetano		
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	Claes Held, MD, PhD		Uppsala	Uppsala University (18)	

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	Shaiful Azmi Yahaya, MD	Irni Yusnida, RN			
		Humayrah Hashim			
* Countries participated in Economics Quality of Life (EQoL) Questionnaires					
**This site received one participant in transfer that was randomized at another site					

eTable 2
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**eTable 3
Ischemia Definitions**

Stress Test Modality	Severe Ischemia	Moderate Ischemia	Mild Ischemia	No Ischemia
Nuclear	≥15% Ischemic Myocardium	10-14.9% Ischemic Myocardium	5-9.9% Ischemic Myocardium	<5% Ischemic Myocardium
Echo	≥4/16 segments with stress-induced severe hypokinesia, akinesia or dyskinesia	3/16 segments with stress-induced severe hypokinesia, akinesia or dyskinesia	1-2/16 segments with stress-induced wall motion abnormality	No stress-induced wall motion abnormalities
Cardiac Magnetic Resonance (CMR) Perfusion	≥25% (8 of 32 subsegments) Ischemic Myocardium	12.5-24.9% (4-7 of 32 subsegments) Ischemic Myocardium	1-12.4% (1-3 of 32 subsegments) Ischemic Myocardium	No Ischemic Subsegments
Exercise Tolerance Test (ETT)	<ol style="list-style-type: none"> 1. Clinical history of typical angina or typical angina during the exercise test 2. Absence of resting ST-segment depression ≥1.0 mm or confounders that render exercise ECG non-interpretable (LBBB, LVH with repolarization, pacemaker, etc.) 3. As compared to the baseline tracing, additional exercise-induced horizontal or downsloping ST-segment depression ≥1.5 mm in 2 leads or ≥2.0 mm in any lead; ST-segment elevation ≥1mm in a non-infarct territory. 4. Either of the following: <ol style="list-style-type: none"> a. Workload at which ST-segment criteria are met is not to exceed completion of stage 2 of a standard Bruce protocol or 7 METs if a non-Bruce protocol is used or b. ST segment criteria are met at <75% of the maximum predicted HR <i>Note: Anatomic eligibility must be confirmed</i> 	<p>Moderate ischemia: either A or B.</p> <p>A) Meets ST segment criteria as stated in the protocol (items 2 + 3 below) during exercise and/or recovery AND either does not meet symptom criteria (item 1 below) OR does not meet workload criterion (item 4 below). An example would be a case meeting the ST depression criteria at >7 METs</p> <p>B) Ischemic ST depression ≥1mm during exercise and/or recovery and peak workload <7 METS but does not meet ISCHEMIA trial criteria (for severe ischemia). An example would be a case with 1.2 mm ST depression in ≥ 2 leads (check this) and peak workload 5 METs.</p>	<p>ST segment depression of at least 1.0 mm in the worst lead during exercise and/or recovery, but not meeting the moderate or severe ischemia definition and either meeting or not meeting the symptom criteria. An example would be 1.2 mm ST depression at 10 METs.</p>	No ischemia; Uninterpretable if there were confounders on the resting ECG precluding interpretation
Trial criteria for eligibility (indicated by unshaded boxes): moderate or severe ischemia by nuclear, echo or CMR, or severe ischemia by ETT				

eTable 4
Randomized Participant Baseline Vital Signs and Laboratory Values
Intent-to-Treat Population

Characteristic	All Randomized Participants (N=5,179)
Vital Signs	
Heart Rate (bpm)	
N	5,146
Median (25th, 75th)	67 (60, 75)
Systolic Blood Pressure (mmHg)	
N	5,150
Median (25th, 75th)	130 (120, 142)
Diastolic Blood Pressure (mmHg)	
N	5,150
Median (25th, 75th)	77 (70, 81)
Body Mass Index (BMI) (kg/m ²)	
N	5,124
Median (25th, 75th)	27.7 (25.0, 31.2)
Lab Values	
<i>Lipid Panel</i>	
Total Cholesterol (mg/dL)	
N	5,074
Median (25th, 75th)	154.3 (129.2, 185.6)
Triglycerides (mg/dL)	
N	4,984
Median (25th, 75th)	124.0 (91.2, 177.0)
HDL Cholesterol (mg/dL)	
N	5,023
Median (25th, 75th)	42.5 (36.0, 51.0)
LDL Cholesterol (mg/dL)	
N	4,941
Median (25th, 75th)	83.0 (63.0, 110.6)
HbA1c for Diabetics (%)	
N	1,960
Median (25th, 75th)	7.2 (6.5, 8.3)
<i>Complete Blood Counts</i>	
Hemoglobin (g/dL)	
N	4,886
Median (25th, 75th)	14.0 (13.0, 15.0)
Platelets (10 ⁹ /L)	
N	4,821
Median (25th, 75th)	220 (182, 261)
WBC (10 ⁹ /L)	
N	4,830
Median (25th, 75th)	7.2 (6.1, 8.6)
<i>Chemistry Panel</i>	
Enrollment Creatinine (mg/dL)	
N	5,179
Median (25th, 75th)	0.9 (0.8, 1.1)
Estimated GFR from Enrollment (mL/min)	
N	5,179

Median (25th, 75th)	81 (67, 97)
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eTable 5
Randomized Participant Baseline Medications
Intent-to-Treat Population

Characteristic	All Randomized Participants (N=5,179)
Anti-platelet Medications	4,871/5,175 (94.1%)
Aspirin	4,647/5,175 (89.8%)
Clopidogrel	1,344/5,175 (26.0%)
Other ¹	116/5,175 (2.2%)
Anticoagulant Medications	202/5,131 (3.9%)
Warfarin	105/5,131 (2.0%)
Dabigatran, Apixaban, Rivaroxaban	90/5,131 (1.8%)
Other ²	8/5,131 (0.2%)
Anti-platelet or Anticoagulant Medications	4,976/5,174 (96.2%)
Statins	4,904/5,173 (94.8%)
<i>Statin Type</i>	
Rosuvastatin	1,169/5,173 (22.6%)
Atorvastatin	2,948/5,173 (57.0%)
Simvastatin	652/5,173 (12.6%)
Other	404/5,173 (7.8%)
High-Intensity Statin ³	
Yes	1,900/5,173 (36.7%)
No/Unknown Dose	3,273/5,173 (63.3%)
Ezetimibe	211/5,175 (4.1%)
Fibrate	146/5,175 (2.8%)
Fish Oil	191/5,175 (3.7%)
Other Lipid Lowering Medication ⁴	38/5,175 (0.7%)
Anti-Hypertensive and Anti-Ischemic/Anginal Medications	5,017/5,175 (96.9%)
Beta Blocker	4,161/5,175 (80.4%)
Calcium Channel Blocker	1,576/5,175 (30.5%)
ACE Inhibitor	2,140/5,175 (41.4%)
Angiotensin Receptor Blocker (ARB)	1,308/5,175 (25.3%)
ACE Inhibitor or ARB	3,413/5,175 (66.0%)
Aldosterone Antagonist	115/5,175 (2.2%)
Diuretic	1,081/5,175 (20.9%)
Alpha 1 Blocker	184/5,175 (3.6%)
Long-acting Nitrate	1,672/5,175 (32.3%)
Ranolazine	260/5,175 (5.0%)
Trimetazadine	245/5,175 (4.7%)
Ivabradine	60/5,175 (1.2%)
Nicorandil	169/5,175 (3.3%)
Other ⁵	66/5,175 (1.3%)
¹ Includes Ticlopidine, Prasugrel, Ticagrelor, Cilostazol, and other anti-platelet medications, including types classified as "Unknown". ² Includes anticoagulant medications such as Enoxaparin and Dalteparin as well as types classified as "Unknown". ³ High-intensity dose of Rosuvastatin or Atorvastatin at the randomization visit. A "No/Unknown Dose" includes Not Applicable, Unknown, and NULL responses. ⁴ Includes medications such as Saroglitazar and Ranozex as well as medications classified as "Unknown". In addition, products such as red yeast rice and flax seed oil are also included. ⁵ Includes medications such as Renin Inhibitors, Hydralazine, Moxonidine as well as anti-hypertensive and anti-ischemic/anginal medications classified as "Unknown".	

