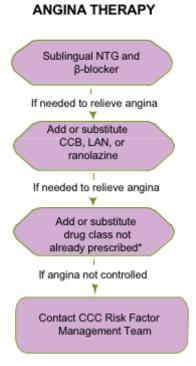
Supplementary Appendix

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

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eFigure 1. OMT Algorithm



*Consider (not in order of preference): ivabradine, nicorandil, perhexiline, trimetazidine, where approved. LAN=long-acting nitrate

LIPID LOWERING THERAPY Goal: LDLC <70mg/dL (1.8 mmol/L)



**Where available

HYPERTENSION THERAPY Goal: Systolic BP <130 mmHg Start β-blocker and ACE inhibitor *** Not at BP goal Increase ACE inhibitor or ARB Not at BP goal Add CCB and/or diuretic****

> Contact CCC Risk Factor Management Team

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***Or ARB if appropriate. Monitor serum potassium closely in participants not on dialysis

****Loop diuretic preferred

eTable 1. Strategies to minimize contrast induced acute kidney injury after cardiac catheterization/PCI

- Pre-, intra- and post-procedure hydration
 - Protocol used in the POSEIDON trial:
 - Initiate 3mL/kg/h of normal saline (NaCl 0.9%) IV, for at least 1 hour prior to angiography
 - Measure LVEDP prior to contrast administration
 - Adapt infusion rate based on LVEDP measurement as follows:
 - 5 mL/kg/hr for LVEDP < 13 mm Hg
 - 3 mL/kg/hr for LVEDP of 13 mm Hg to 18 mm Hg
 - 1.5 mL/kg/hr for LVEDP > 18 mm Hg
 - Continue fluid administration for 4 hours post procedure
 - Simplified protocol based on LVEF (expert opinion):
 - Participants with preserved EF
 - IV 0.9% NS at 1 cc/kg/hour for 12 hours pre- and postprocedure
 - Participants with EF<40%
 - IV 0.45% NS at cc/cc replacement (urine output should be match to maintain euvolemic state) for 12 hours pre- and postprocedure
- Pre-procedure high dose statins
- Avoid nephrotoxic agents for at least 48 hours prior
- Use iso- or low-osmolar contrast agents
- Limit contrast used: Ultra-low/Zero volume contrast techniques (IVUS guided PCI)
 - Use small diameter catheters (i.e., 5–6 French) without side-holes
 - All contrast injections require simultaneous cine angiogram, i.e., "no dye without the cine's eye."
 - Limit the volume of contrast injected from the catheter to 1–2 cm³ per injection using a 3-cm³ syringe.
 - During PCI, prior to exchange of devices such as balloon catheters, remove contrast from the guide catheter by back bleeding contrast out of the "Y" connector.
 - If available, display previous angiographic images (including angiography from past procedures) alongside active fluoroscopy screen as a reference to use as guidance during guide wire, balloon, stent and ultrasound passage.
 - Absolutely no contrast "puffing"/test injections during the procedure.
 - Use IVUS liberally for pre-PCI assessment of the lesion, selection of therapeutic modalities, and post-PCI result assessment.
 - Avoid ventriculography
 - Use of biplane if available
- Consider ischemia-guided revascularization
- Consider staged PCI for complex multivessel disease

N= intravenous; NUS= intravascular ultrasound; LVEDP= left ventricular end diastolic pressure; LVEF= left ventricular ejection fraction; PCI= percutaneous coronary intervention

eTable 2. Strategies to minimize acute kidney injury after CABG

Consider delay of surgery ≥7 days from time of cardiac catheterization

Use of off pump CABG may be reasonable

Renally dose all medications

In patients undergoing on pump CABG, maintain perioperative hematocrit > 19% and mean arterial pressure > 60 mmHg $\,$

CABG= coronary artery bypass graft surgery

eTable 3. Schedule of Follow-Up

Randomization visit (Baseline Visit) ۰ð Catheterization 8 PCI or CABG Screening visit Follow up 6m^B Frequency 1.5m^A 3m^A 12m^A 18m^E 24m 36m^c 30m beyond 36 Visit 1 Visit3 Visit 4 Visit6 Visit8 Visit 2 Visit5 Visit7 months Eligibility screen Х Informed consent (including Х biorepository consent if applicable) Creatinine and pregnancy test^D Х Medical History/Medical Status Х Х Х Х Х Х Х Х Х Х Q6m Cardiovascular medications Х Х Х Х Х Х Х Х Х Х Q6m NYHA* and CCS class** Х Х Х Х Х Х Х Х Х Х Q6m Release for medical records signed Х Х Х Х Q12m Safety assessment G Х Vital signs, weight, height^H Х Х Х Х Х Х Х Х Х Q12m Standard lab results¹ Х Х Х Х Х Х Х Q12m \mathbf{v}^{J} Biorepository blood draw Х Y Cardiac biomarkers^L Х Electrocardiogram (ECG) Х Х νN Х @ closeout Lifestyle Assessment (PACE)*** Х Х Х Х Х Q12m Lifestyle Counseling (PACE)*** Х Х Х Х Х Х Х Х Х Q6m Morisky Green Levine Medication Adherence Х Х Х Х Х Х Х Q6m Brief symptoms/QOL assessment^P Х Х Х Х Х Х Х Х Х Q6m Initiate Optimal Medical Therapy (OMT) Х Medical Therapy Evaluation and Optimization^Q Х Х Х Х Х Х Х Х Q6m Schedule catheterization for INV participants^R Х Hospitalization assessment Х Х Х Х Х Х Х Х Q6m Х Х Х Q6m Endpoint assessment Х Х Х Х Х Х

Schedule of Study Assessments and Procedures (Protocol Date: Jan. 06.2014)

Follow-up visits will be scheduled based on time since the date of randomization (baseline).

*NYHA- New York Heart Association **CCS- Canadian Cardiovascular Society ***PACE- Patient-centered Assessment and Counseling for Exercise and nutrition (PACE) assessment and counseling

^A 1.5, 3, and 12 month visits should be in clinic visits, depending on participant stability, risk factor control, and geography.

^B 6, 18, and 30 month visits may be via telephone, email, or in clinic depending on participant stability, risk factor control, and geography.

^C Follow ing the 36 month visit, follow -up visits should be in clinic visits at least every 12 months. Clinic visits can be replaced by email or phone depending on participant stability, risk factor control, and geography.

^D Creatinine if not done within 90 days and pregnancy test if premenopausal.

^F CCTA not performed if estimated glomerular filtration rate < 60ml/min (unless requested by the treating physician) and not performed in other selected participants (see ^G Safety Assessment (refer to section 13.4).

^H Height is only needed at randomization, assessments only required if visit is completed in clinic.

Required labs include: lipids (preferably fasting) at 3 month visit then semiannually only, and HbA1c (at visit 4, 6, 8 and annually thereafter for diabetic participants.

These lab

results will be requested from the participant's physician. If these results are not available they should be obtained by either the participant's treating physician or study staff. Creatinine values obtained clinically for participants with eGFR <60 at the three month follow-up visit and annually will also be recorded.

^J Additional lab required at randomization includes complete blood count Request from participant's physician, since it is expected that routine blood work will have been done within the last 6 months

^K May be requested.

^L For participants undergoing PCI: troponin and CK-MB pre-procedure and at 8-16 ± 2 hours post-PCI or at hospital discharge, whichever comes earlier. For participants undergoing CABG: troponin and CK-MB pre-procedure and at 18 ± 6 hours post-CABG. All biomarker measurements should be recorded on eCRF. A biomarker measurement should be obtained before and after all PCI and CABG procedures, whenever possible.

^M Send to ECG core lab; ECG required for all cardiac admissions and revascularizations; year 1 ECG optional (filed on site) and closeout.

 $^{\rm N}$ ECG done following procedure (60±30 mins post-PCl, 3 days post-CABG).

^O Seattle Angina Questionnaire/Duke Activity Status Index/Rand general health status item/Perceived Stress Scale/Patient Health Questionnaire/Life Orientation Test – Revised/EQ-5D/Demographic characteristics. Not required for the ISCHEMIA CKD ancillary trial.

^P Selected Seattle Angina Questionnaire/Rose dyspnea scale/EQ-5D.

^Q At every follow -up visit the research team, in collaboration with the treating physician(s), will evaluate effectiveness of medical therapy and optimize as needed according to guideline recommendations and study algorithms.

^R Planned cath and revascularization only in the INV group. See MOO for time windows for performing cath and revascularization after randomization. Catheterization and optimal revascularization treatment should be targeted within 30 days after randomization in the Invasive strategy group. In the Conservative group, catheterization and optimal revascularization is reserved for participants with refractory angina symptoms or acute ischemic events.

eTable 4. Definitions of Clinical Endpoints

<u>Death</u>

All deaths will be adjudicated and classified as cardiovascular, non-cardiovascular or undetermined. Cardiovascular deaths are defined as all deaths excluding those for which the principal and underlying cause is solely non-cardiovascular. Any death for which a cardiovascular contributing cause is suspected will also be considered a cardiovascular death.

Myocardial Infarction

Two versions of MI will be adjudicated in ISCHEMIA-CKD: a primary definition and secondary definition. Each definition includes a hierarchy of markers and threshold values as well as a set of rules for diagnosing MI when one or more key elements of the medical record are missing.

The <u>Primary Definition</u> is based upon the Universal Definition of MI, but relies upon site-reported MI decision limits for troponin (which may or may not be the same as the manufacturer 99%URL), and has selected unique marker criteria for MI after PCI or CABG (Type 4a, 5).

The <u>Secondary Definition</u> is also based upon the Universal Definition of Myocardial Infarction, but specifically uses the 99%URL from the assay manufacturer's package insert (which may or may not be the site's MI decision limit) and uses the same supporting criteria (eg. angiographic and ECG) as the UMI definition.

All MI events will be classified based on the Universal MI classification system as follows:

- Type 1: Spontaneous MI
- Type 2: Secondary MI
- Type 3: Sudden Death MI
- Type 4a: MI related to PCI
- Type 4b: MI related to stent thrombosis
- Type 4c: MI related to stent restenosis
- Type 5: MI related to CABG
- Silent MI

Spontaneous MI (Types 1, 2, 4b, 4c)

Diagnosis of spontaneous MI will be satisfied by a clinical setting consistent with acute myocardial ischemia and any one or more of the following criteria:

Marker elevation, as outlined below and <u>at least 1 of the following</u>:

- Symptoms of ischemia, usually lasting > 20 minutes in duration
- New ischemic ST and/or T wave and/or Q-wave ECG changes, or new LBBB, as described below
- Imaging evidence of new loss of viable myocardium in comparison to the baseline imaging test
- Angiographic evidence of intracoronary thrombus, stent thrombosis (4b) or high- grade instent restenosis (≥50%) (4c)

Marker data not available and <u>at least 2 of the following:</u>

- New ischemic ST and/or T wave and/or Q-wave ECG changes, or new LBBB, as described below
- Imaging evidence of new loss of viable myocardium in comparison to the baseline imaging test
- Angiographic evidence of intracoronary thrombus.

Autopsy evidence of a fresh myocardial infarction as stand-alone criterion

Spontaneous MI Marker Criteria

Troponin, including high-sensitivity troponin, is the preferred biomarker and takes precedence over CK-MB for both definitions.

<u>Primary Definition:</u> Preferentially uses a troponin threshold value reported as MI Decision Limit or the Upper Limit of Normal (ULN). Marker elevation is defined as troponin > ULN/MI decision limit. If troponin is not done or not available, then CK-MB > ULN will qualify. If both troponin and CK-MB are not done or not available, then CK > 2 x ULN will qualify.

<u>Secondary Definition:</u> Preferentially uses a troponin threshold reported by the manufacturer, namely, the manufacturer 99th percentile. Marker elevation is defined as troponin > 99th percentile. If the troponin 99th percentile is not reported, then troponin > ULN will qualify. If troponin is not done or not available, then CK-MB > ULN will qualify. If both troponin and CK-MB are not done or not available, then CK > 2 x ULN will qualify.

Spontaneous MI ECG Criteria

ECG criterion is considered to be met if any of the following:

ST elevation: New ST elevation at the J-point in two contiguous leads with the cutpoints: $\geq 0.2 \text{ mV}$ in men >age 40 and $\geq 0.25 \text{mV}$ in men <40 years or $\geq 0.15 \text{ mV}$ in women in leads V2–V3 and/or $\geq 0.1 \text{ mV}$ in other leads, or new LBBB.

Any new Q-wave in leads V2–V3 \ge 0.02 seconds or QS complex in leads V2 and V3 or Q-wave \ge 0.03 seconds and \ge 0.1 mV deep or QS complex in leads I, II, aVL, aVF, or V4–V6 in any two leads of a contiguous lead grouping (I, aVL, V6; V4–V6; II, III, and aVF) or R-wave \ge 0.04 seconds in V1–V2 and R/S \ge 1 with a concordant positive T- wave in the absence of a conduction defect.

ST depression and/or T-wave changes, new horizontal or down-sloping ST depression ≥ 0.05 mV in two contiguous leads; and/or T-wave inversion ≥ 0.1 mV in two contiguous leads. The ST-T wave criteria only apply in the absence of findings that would predude ECG analysis such as LBBB, LVH with repolarization abnormalities, pre-excitation and pacemakers.

Silent MI

This event includes evidence of new silent Q-wave MI detected during routine protocol or clinically obtained ECG follow-up. Silent MI events will be classified as a type 1 MI.

Sudden death MI (Type 3)

MI events in which a presentation consistent with infarction is present but the patient dies before the biomarkers are drawn or within the first few hours of the event before the biomarkers become positive. Sudden unexpected cardiac death, including cardiac arrest, often with symptoms suggestive of myocardial ischemia, accompanied by presumably new ST-segment elevation, or new LBBB, or evidence of fresh thrombus in a coronary artery by angiography and/or at autopsy, but death occurring before blood samples could be obtained, or at a time before the appearance of cardiac biomarkers in the blood.

PCI-Related MI (Type 4a)

Primary Definition

CK-MB is the preferred biomarker and takes precedence over troponin. For subjects with normal baseline biomarker level pre-PCI, peri-PCI MI requires a rise in CK-MB to

>5-fold the ULN (or a rise in troponin to >35 times the MI Decision Limit/ULN, when CK- MB is unavailable) within 48 hours post-PCI. If pre-PCI cardiac markers (CKMB or cTn) are elevated, they must be stable or falling as indicated by two samples at least 6 h apart. The post-PCI CKMB level should reflect a rise of >20% over pre-PCI levels. In addition to biomarker criteria, peri-PCI MI requires at least one of the following:

- Post- procedure angiographic TIMI 0/1 flow in a major coronary artery or a side branch with reference vessel diameter ≥2.0 mm which had TIMI 2-3 flow at baseline, or TIMI 2 flow in a major coronary artery or a side branch with reference vessel diameter ≥3.0 mm which had TIMI 3 flow at baseline or Type C dissection (NHLBI classification) or greater in the target vessel.
- New ECG changes (ST segment elevation or depression >0.1mV in 2 contiguous leads), new pathologic Q-waves in ≥2 contiguous leads, or new persistent LBBB present on a post-PCI ECG obtained at least 30 minutes and up to 48 hours post procedure in the absence of any intervening coronary event between the time of the PCI procedure and the ECG showing changes.

NOTE: A type 4a MI will be diagnosed with a rise in CK-MB to >10-fold the ULN (or when CK-MB is unavailable, a rise in troponin to >70 times the MI Decision Limit/ULN) as a stand-alone criterion. If biomarkers are missing, a type 4a MI will be diagnosed if BOTH ECG criteria (new ST elevation or depression, Q-wave criteria, or new and persistent LBBB) AND angiographic criteria above are present. If pre-PCI cardiac markers are missing, they will be assumed to be normal in those without a preceding event.

Secondary Definition

Elevation of troponin values >5 X 99th percentile URL within 48 hours post-PClin patients with normal baseline troponin values pre-PCl AND a rise of troponin values >20% if the baseline values are elevated pre-PCl and are stable or falling. If the troponin 99th percentile is not available, the MI Decision Limit / ULN may be used. If troponins are not available, CKMB elevation >5 X ULN will be used.

In addition to biomarker criteria, peri-PCI MI requires at least one of the following:

- Symptoms suggestive of myocardial ischemia (≥20 min)
- New ischemic ST changes or new pathological Q waves. (see "ECG Criteria" above) Note the UMI definition uses ≥0.05 mV of STD whereas the ISCHEMIA definition uses ≥0.1mV for PCI related ECG criteria
- Angiographic evidence of a flow limiting complication, such as loss of patency of a side branch, persistent slow-flow or no re-flow, embolization, or Type C dissection (NHLBI classification) or greater in the target vessel.
- Imaging evidence of new loss of viable myocardium or new regional wall motion abnormality.

NOTE: A type 4a MI will be diagnosed with a rise in troponin to >70 times the 99th percentile URL (or, when troponin is unavailable, a rise in CK-MB to >10 times the ULN) as a stand-alone criterion. If biomarkers are missing, a type 4a MI will be diagnosed if BOTH ECG criteria (new ST elevation or depression, Q-wave criteria, or new and persistent LBBB) AND angiographic criteria above are present. If pre- PCI cardiac markers are missing, they will be assumed to be normal in those without a preceding event.

CABG-Related MI (Type 5)

Primary Definition

CK-MB is the preferred serum biomarker and takes precedence over cardiactroponin. For subjects with normal baseline biomarker level pre-CABG, peri-CABG MI requires a rise in CK-MB to >10-fold the ULN (or a rise in troponin to >70 times MI Decision Limit/ULN when CK-MB is

unavailable) within 48 hrs post-CABG. In addition to biomarker criteria, peri-CABG MI requires at least one of the following:

- A new substantial wall motion abnormality by cardiac imaging (CEC assessed), except new septal and apical abnormalities. The CEC will have latitude in determining whether a new wall motion abnormality is "substantial" in the context of the clinical event.
- New pathologic Q-waves in ≥2 contiguous leads or new persistent LBBB is present on post CABG ECG obtained day 3 post CABG, or hospital discharge, whichever comes earlier in the absence of any intervening coronary event between the time of the CABG procedure and the ECG showing changes.

NOTE: A type 5 MI will be diagnosed with a rise in CK-MB to >15-fold the ULN (or when CK-MB is unavailable a rise in troponin to >100 times the MI Decision Limit/ULN) as a stand-alone criterion. If biomarkers are missing, an MI will be diagnosed if the ECG criteria (New pathologic Q waves or new persistent LBBB) AND new substantial wall motion abnormality are BOTH present. If pre-CABG cardiac markers are missing, they will be assumed to be normal in those without a preceding event.

Secondary Definition

Elevation of troponin values >10 X 99th percentile URL within 48 hrs post-CABG in patients with normal baseline troponin values (\leq 99th percentile URL). If the troponin 99th percentile is not available, the ULN may be used. If troponins are not available, CKMB elevation >10 X ULN will be used. In addition to biomarker criteria, peri-CABG MI requires at least one of the following:

- New pathologic Q waves or new LBBB
- Angiographic evidence of new graft or new native coronary artery occlusion.
- Imaging evidence of new loss of viable myocardium.

NOTE: A type 5 MI will be diagnosed with a rise in troponin to >100 times the 99th percentile URL (or when troponin is unavailable a rise in CK-MB to >15 times the ULN) as a stand-alone criterion. If biomarkers are missing, an MI will be diagnosed if the ECG criteria (New pathologic Q waves or new persistent LBBB) AND new substantial wall motion abnormality are BOTH present. If pre-CABG cardiac markers are missing, they will be assumed to be normal in those without a preceding event.

Complicated MI and Large MI

Complicated MI: Prognostically important MIs may also be identified as those with complications such as hemodynamic instability, cardiogenic shock, drop in EF >10% from baseline, electrical instability with life-threatening VT or VF, or heart failure complicating MI. Complicated myocardial infarctions may typically require ICU care, invasive support (eg. intubation, IABP, PA catheters) and intravenous medications (eg. inotropes or antiarrhythmics.) CEC adjudicators will identify complicated MIs based upon the information available to them in the eCRF and source documents.

- Hemodynamic instability: requiring fluids, inotropic or vasopressor support to maintain endorgan perfusion. May progress to shock if also accompanied by end-organ underperfusion.
- Shock: Compromise of end-organ perfusion due to hemodynamic instability and sustained hypotension. Often manifested by hypotension, increased creatinine, shock liver, and decreased mentation.
- Life-threatening VT or VF: Requiring antiarrhythmics or defibrillation to return sinus rhythm. Transient runs of VT (eg. during reperfusion) are not associated with hemodynamic instability are not usually considered life-threatening.
- Decreased EF ≥ 10%: EF assessment during the event which indicates a drop from prior assessments (eg. EF 30% from previous EF 55%)

- HF in the setting of an MI is defined on the basis of the physician's decision to treat HF with an intravenous (IV) diuretic, IV inotropic agent or IV vasodilator and at least 1 of the following:
 - Presence of pulmonary edema or pulmonary vascular congestion on chest radiograph believed to be of cardiac cause.
 - Rales greater than 1/3 up the lung fields believed to be due to HF.
 - Pulmonary Capillary Wedge Pressure (PCWP) or left ventricular end diastolic pressure (LVEDP) greater than 18 mmHg.
 - Dyspnea, with documented paO2 less than 80 mmHg on room air or O2 saturation less than 90% on room air, without significant lung disease

Large MI: The size of MI will be assessed by examining peak levels of cardiac biomarkers as a continuous function.

Hospitalization for Unstable Angina

Prolonged ischemic symptoms at rest (usually \geq 10 minutes in duration), or accelerating pattern of chest pain that occurs with a lower activity threshold (CCS class III or IV) considered to be myocardial ischemia upon final diagnosis resulting in an unscheduled visit to a healthcare facility resulting in an overnight stay <u>generally</u> within 24 hours of the most recent symptoms, cardiac biomarkers not meeting MI criteria, and at least one of the following:

- New or worsening ST or T wave changes on resting ECG* (core laboratory assessed)
- Angiographic evidence of a ruptured/ulcerated plaque, or thrombus in an epicardial coronary artery believed to be responsible for the ischemic symptoms/signs (core laboratory assessed).

*ECG Criteria:

<u>ST segment shifts and T-wave changes:</u> New horizontal or down-sloping ST depression ≥ 0.05 mV in two contiguous leads; and/or T inversion ≥ 0.1 mV in two contiguous leads, or new ST segment elevation ≥ 0.1 mV in 2 contiguous leads. The ST-T wave criteria only apply in the absence of findings that would preclude ECG analysis such as LBBB, LVH with repolarization abnormalities, pre-excitation and pacemakers.

Resuscitated Cardiac Arrest

Resuscitated cardiac arrest is defined as successful resuscitation for documented cardiac arrest out-of-hospital (or ER) in a patient subsequently admitted to hospital, and then discharged. A patient who is successfully resuscitated but dies before hospital discharge of complications related to the cardiac arrest (e.g., anoxic encephalopathy, septic shock), will be classified as a coronary heart disease death. An uncomplicated procedure-related cardiac arrest with prompt resuscitation and without adverse sequelae will not be counted as an event. Events that meet the MI criteria will be categorized as MI.

Hospitalization for Heart Failure

While patients may have multiple simultaneous disease processes, for the end point event of heart failure requiring hospitalization, the diagnosis of congestive heart failure would need to be the primary process. Heart failure (HF) requiring hospitalization is defined as an event that meets the following criteria:

a. Requires hospitalization defined as an admission to an inpatient unit or a visit to an emergency department that result in at least a 24 hour stay (or a date change if the time of admission/discharge is not available).

- b. Clinical symptoms of heart failure, including at least one of the following: New or worsening
 - Dyspnea
 - Orthopnea
 - Paroxysmal nocturnal dyspnea
 - increasing fatigue/worsening exercise tolerance

AND

- c. Physical signs of heart failure, including at least two of the following:
 - 1. Edema (> 2+ lower extremity)
 - 2. Pulmonary rales (pulmonary edema not occurring as the consequence of an arrhythmia in the absence of worsening heart failure. If pulmonary edema complicates acute MI event should be coded as MI)
 - 3. Jugular venous distension
 - 4. Tachypnea (respiratory rate > 20 breaths/minute)
 - 5. Rapid weight gain
 - 6. S3 gallop
 - 7. Increasing abdominal distension or ascites
 - 8. Hepatojugular reflux
 - 9. Radiological evidence of worsening heart failure
 - 10. A right heart catheterization within 24 hours of admission showing a pulmonary capillary wedge pressure (pulmonary artery occlusion pressure) ≥ 18 mm Hg and/or a cardiac output < 2.2 L/min/m2

NOTE: Biomarker results (e.g., brain natriuretic peptide (BNP)> 500 or Pro-NT BNP > 2500) consistent with congestive heart failure will be supportive of this diagnosis, but the elevation in BNP cannot be due to other conditions such as cor pulmonale, pulmonary embolus, primary pulmonary hypertension, or congenital heart disease. Increasing levels of BNP, although not exceeding the ULN, may also be supportive of the diagnosis of congestive heart failure in selected cases (e.g. morbid obesity).

AND

d. Need for additional/increased therapy

Initiation of, or an increase in, treatment directed at heart failure or occurring in a patient already receiving maximal therapy for heart failure and including at least one of the following:

- 1. Initiation of or a significant augmentation in oral therapy for the treatment of congestive heart failure
- 2. Initiation of intravenous diuretic, inotrope, or vasodilator therapy
- 3. Uptitration of intravenous therapy, if already on therapy
- 4. Initiation of mechanical or surgical intervention (mechanical circulatory support, heart transplantation or ventricular pacing to improve cardiac function), or the use of ultrafiltration, hemofiltration, or dialysis that is specifically directed at treatment of heart failure.

AND

e. No other non-cardiac etiology (such as chronic obstructive pulmonary disease, hepatic cirrhosis, acute renal failure, or venous insufficiency) and no other cardiac etiology (such as pulmonary embolus, cor pulmonale, primary pulmonary hypertension, or congenital heart disease) for signs or symptoms are identified.

<u>Stroke</u>

Stroke is defined as the rapid onset of a new neurologic deficit attributed to an obstruction in cerebral blood flow and/or cerebral hemorrhage with no apparent non-vascular cause (eg. trauma, tumor, or infection). Available neuroimaging studies will be considered to support the clinical impression and to determine if there is a demonstrable lesion compatible with an acute stroke.

Classification:

Transient Ischemic Attack

A Transient Ischemic Attack is defined as an acute episode of focal cerebral, spinal, or retinal dysfunction caused by an ischemia of central nervous system tissue which resolves within 24 hrs and without neuroimaging evidence of acute infarction.

Ischemic Stroke

Ischemic stroke is defined as an acute episode of focal cerebral, spinal, or retinal dysfunction caused by an infarction of central nervous system tissue.

Signs/ symptoms \geq 24 hrs regardless of neuroimaging findings: Ischemic stroke can be defined clinically- by persistence of signs and symptoms \geq 24 hrs, usually supported by evidence of infarction on neuroimaging (CT or MRI) although very early neuroimaging (usually with CT) may not demonstrate the infarction.

Signs/ symptoms < 24 hrs with neuroimaging evidence of infarction: Ischemic stroke can be defined by neuroimaging- where neuroimaging (usually MRI diffusion weighted or flair images) confirms the presence of acute infarction even if signs/ symptoms resolve within 24 hrs.

Patients admitted for an acute stroke treated with thrombolysis or interventions that have no residual neurologic symptoms after treatment will be classified as an ischemic stroke.

Ischemic Stroke with Symptomatic Hemorrhagic Conversion

Hemorrhagic conversion may be a consequence of ischemic stroke and may be symptomatic, resulting in neurologic deterioration, or asymptomatic. Symptomatic Hemorrhagic Conversion is defined neuroimaging evidence of hemorrhage within the area of infarction associated with clinical deterioration (eg. increase in NIHSS of \geq 4 points) or death, symptoms to hemorrhage related mass effect, or symptoms out of proportion to what would be expected from the ischemic stroke or cerebral edema alone. When an Ischemic Stroke with Symptomatic Hemorrhagic Conversion is identified, the date and time of stroke onset will refer to the first onset of the Ischemic Stroke and will not be counted as two events.

Hemorrhagic Stroke

Hemorrhagic stroke is defined as an acute episode of focal or global cerebral or spinal dysfunction caused by a non-traumatic intraparenchymal, intraventricular, or subarachnoid hemorrhage.

Undetermined- or Uncertain type- of Stroke

Undetermined stroke is defined as a stroke with insufficient information to allow categorization as Ischemic Stroke or Hemorrhagic Stroke. If possible, speculate on the stroke subtype and note in Comments. This is not to signify an indeterminate event where there is insufficient evidence to suspect a stroke had occurred.

eTable 5. Committee Members and Key Personnel

National Heart, Lung, and Blood Institute

Jerome L. Fleg, Project Officer Ruth Kirby

ISCHEMIA-CKD Clinical Coordinating Center (CCC)*

Study Leadership

Sripal Bangalore (Principal Investigator)

CCC Faculty

Judith S. Hochman (ISCHEMIA trial Chair) David J. Maron (ISCHEMIA trial Co-Chair) Jeffrey Berger (Director of the Biorepository, ISCHEMIA Regional Leader) Roy Mathew (Country Lead Nephrologist for US) Jonathan Newman (ISCHEMIA Regional Leader) Harmony R. Reynolds (ISCHEMIA Regional Leader) Mandeep Sidhu (US-VA Regional Co-Leader)

Program Director

Stephanie Mavromichalis

Project Managers

Gia Cobb Stephanie Ferket ** Andre Gabriel **

Clinical Research Associates

Diana Cukali Kevin McMahon **

Clinical Trial Assistants

Ahmed Ayoub Matthew Shinseki ** Paula Wilson Solomon Yakubov **

Data Analyst

Mark Xavier *see ISCHEMIA Trial Design Manuscript for complete listing **past members

ISCHEMIA Statistical and Data Coordinating Center, Duke Clinical Research Institute

Sean O' Brien, Principal Investigator See ISCHEMIA Trial Design Manuscript for complete listing

ISCHEMIA-CKD Committee Members Steering Committee

Sripal Bangalore, Principal Investigator Judith S. Hochman, ISCHEMIA trial Chair David J. Maron, ISCHEMIA trial Co-Chair Glenn M. Chertow, Nephrologist William Boden, ISCHEMIA trial Co-PI Bruce Ferguson, ISCHEMIA trial Co-PI Robert Harrington, ISCHEMIA trial Co-PI Gregg W. Stone, ISCHEMIA trial Co-PI David O. Williams, ISCHEMIA trial Co-PI

Renal Committee

Charles A. Herzog (Chair) Sripal Bangalore Carlo Briguori David M. Charytan Glenn M. Chertow Jerome Fleg Peter A. McCullough Roxana Mehran Ruth Kirby

Publications Committee

Sripal Bangalore (Chair) Karen Alexander Jerome Fleg Judith S. Hochman David J. Maron Roy Mathew Sean M. O'Brien Harmony R. Reynolds Mandeep Sidhu

Optimal Medical Therapy Committee

Same as the ISCHEMIA trial (See ISCHEMIA Trial Design Manuscript for complete listing)

Optimal Revascularization Therapy Planning Committee

Same as the ISCHEMIA trial (See ISCHEMIA Trial Design Manuscript for complete listing)

Clinical Event Review Committee

Same as the ISCHEMIA trial (See ISCHEMIA Trial Design Manuscript for complete listing)

BioRepository Committee

Same as the ISCHEMIA trial (See ISCHEMIA Trial Design Manuscript for complete listing)

EQOL Committee

Same as the ISCHEMIA trial (See ISCHEMIA Trial Design Manuscript for complete listing)

Recruitment for Women & Minorities

Same as the ISCHEMIA trial (See ISCHEMIA Trial Design Manuscript for complete listing)

DSMB Members

Same as the ISCHEMIA trial (See ISCHEMIA Trial Design Manuscript for complete listing)

Independent Statistical Analysis Center for DSMB Reporting

Same as the ISCHEMIA trial (See ISCHEMIA Trial Design Manuscript for complete listing)

ECG/ETT Core Lab

Same as the ISCHEMIA trial (See ISCHEMIA Trial Design Manuscript for complete listing)

Angiographic Core Lab

Same as the ISCHEMIA trial (See ISCHEMIA Trial Design Manuscript for complete listing)

Academic Research Organizations (AROs)

Same as the ISCHEMIA trial (See ISCHEMIA Trial Design Manuscript for complete listing)

Contract Research Organizations (CROs)

Same as the ISCHEMIA trial (See ISCHEMIA Trial Design Manuscript for complete listing)

eTable 6. Site Listing

country (No.Randomizations)	Investigator(s)	Study Coordinator(s)	City & State (if applicable)	Institution (No. Randomizations)
Inited States (159) ead Country Nephrologist				
Roy Mathew, MD	Mayil S. Krishnam, MD	Shirin Hey dari, MS	Orange, CA	University of California Irvine Medical Center
Roy Matnew, MD	Jef frey C. Milliken, MD Pranav M. Patel, MD Arnold H. Seto, MD Kev in T. Harley, MD (N) Michael A. Gibson, MD By ron J. Allen, MD Wei Ling Lau, MD (N)	Edgar Karanjah, MD Wanda C. Marfori, MD Eduardo Hernandez-Rangel, MD Pam Singh	orango, or c	(23)
			Rochester,	
	Patricia Pellikka, MD	Gay lin Petty, CVT	MN	May o Clinic (16)
	LaTony a J. Hickson, MD (N)	Susan K. Milbrandt Dawn D. Shelstad		
	Harmony R. Reynolds, MD Jonathan D. Newman, MD, MPH Sripal Bangalore, MD, MHA Lawrence M. Phillips, MD Muhamed Saric, MD	Stanley E. Cobos, BA Kirsten J. Quiles, MS Rav en R. Dwyer, MPH Dalisa Espinosa, MBS	New York, NY	NYU Langone Medical Center-Bellevue Hospital (15)
	Olga Zhdanov a, MD (N) Kreton Mav romatis, MD	John Doan, MD	Decatur, GA	Atlanta VA Medical Center (13)
	Jason Linefsky, MD Harold Franch, MD (N)	Raven Lee, CCRP Risha Patel		
	Anjali Acharya, MD (N) Seth Sokol, MD Jay Meisner, MD Amit Kakkar, MD Tarek Rashid, MD Hatem Elabd, MD	Jeanne Russo, RN Cidney Schultz, RN	Bronx, NY	Jacobi Medical Center (12)
	Charles Herzog, MD	Shari Mackedanz	Minneapolis,	Hennepin County Medical Center (9)
	Mengistu Simegn, MD	Barbara Wicklund	MN	
	Salvatore P. Costa, MD Terrance Welch, MD Michael Chobanian, MD (N)	Henry C. Stokes, RN Gay lin Petty, CVT	Lebanon, NH	Dartmouth Hitchcock Medical Center(7)
	Subhash Banerjee, MD	Preeti Kamath, BDS, MHA, CCRP Ishita Tejani, BDS, MS, MSPH	Dallas, TX	V.A. North Texas Health Care System (6)
	Adeday o Adeboye, MD	Amy Flowers	Columbia, SC	William Jennings Bryan Dom V.A. Medical Center (5)
	Roy Mathew, MD (N)	Kathry n Mason Anjana Rishmawi		
	Sudhanv a S. Hegde, MD	Stanley E. Cobos, BA Rav en R. Dwyer, MPH Dalisa Espinosa, MBS Kirsten J. Quiles, MS Caroly n J. Gruber, PA-C Noelle M. Durf ee, MS PA-C	Brookly n, NY	Kings County Hospital Center (9)
	Khaled Abdul-Nour, MD Lalathaksha Kumbar, MD (N)	Heather Golden Naima L. Ogletree, DNP, APRN- BC	Detroit, MI	Henry Ford Health System (4)
	Jerry Yee, MD (N)	Schawana Thaxton, DNP, NP-C		
	Alec Moorman, MD Bilal Malik, MD (N)	Fatima Ranjbaran, RN Bry n Smith, BS	Seattle, WA	University of Washington Medical Center (4

	Carly Ohmart		
Radmilar Ly ubarova, MD	Wendy L. Stewart, MS	Albany, NY	Albany Medical Center Hospital (3)
Mohammad El-Hajjar, MD	Kristin M. Salmi, BS		
Mandeep S. Sidhu, MD, MBA			
Steven A. Fein, MD			
Mikhail T. Torosoff, MD, PhD			
Radmila Ly ubarova, MD			
Sulagna Mookherjee, MD			
Krzy sztof Drzymalski, MD			
Rafia Chaudhry, MD (N)			
Krishnakumar Hongalgi, MD (N)			
Arif Asif, MD (N)(2012-2015) Loay Salman, MD (N)(2015- 2018)			
Patricia K. Nguyen, MD	Dav is Vo, BS	Palo Alto, CA	VA Palo Alto Healthcare System (3)
Yiming Lit, MD (N)	James Hirsch, BS		
Stev en P. Sedlis, MD	Leandro C. Maranan, CCRC	New York, NY	VA New York Harbor Health Care System (3)
Robert M. Donnino, MD			
Jeffrey Lorin, MD			
David Goldfarb, MD (N)			
Mohammad El-Hajjar, MD	Jennifer Thomson, MA	Albany, NY	Samuel Stratton VA Medical Center of Albany NY (2)
Paul Der Mesropian, MD (N)			
Joseph Sacco, MD			
Naveed Akhtar, MD			
Maris Orgera, MD			
Mandeep S. Sidhu, MD, MBA (2	012-2016)		
Roy Mathew, MD (N) (2012-			
2015) Elvira Gosmanova, MD (N) (201	5-2018)		
	Badhma Valaiy apathi, MD	Birmingham,	UAB Vascular Biology and Hypertension
Fadi Hage, MD	Baurina valaty aparti, ND	AL	Program (2)
Dana Rizk, MD (N)			
James E. Davies, MD			
Massoud Leesar, MD			
Jaeky eong Heo, MD Amy Iskandrian, MD			
Firas Al Solaiman, MD			
Satinder Singh, MD			
		Destes MA	Brigham & Women's Hospital, Harvard Medical
Peter H. Stone, MD	Hermine Osseni, MS	Boston, MA	School (2)
David Charytan, MD (N)	Charlene Wiy arand (BS)		
	Peter Douglass, BA		
	Hayley Pomeroy, BA		
	Alexandra Craft, BA		
	Bethany Harvey, BA		NVIII Wetherer (2)
Kev in Marzo, MD	Wendy Drewes, RN	Mineola, NY	NYU Winthrop (2)
Juan Gaztanaga, MD	Dipti Datal RN		
Shay an Shirazian, MD (N)	Dipti Patel, RN		NYU-HHC Lincoln Medical and Mental Health
Lekshmi Dharmarajan , MD	Jenne M. Jose, PA	Bronx, NY	Center (2)
	Stanley E. Cobos, BA		
	Raven R. Dwyer, MPH		
	Kirsten J. Quiles, MS		
Janani Rangaswami, MD (N)	Rachel Murphy, BS	Philadelphia, PA	Albert Einstein Medical Center (2)
Christian Witzke, MD	Kinnari Murphy, MPH		
Gregg Pressman, MD			
John B. Kostis, MD	Nora M. Cosgrove, RN	New	Cardiov ascular Institute, Rutgers RWJ Medical
JUNITE NUSUS, ME	NUIAINI. CUSYIUVE, KIN	Brunswick, NJ	School (1)
		Dianowick, No	
Abel E. Morey ra, MD Jonathan Lebowitz, MD (N)		Dranswick, rec	

Ellis W. Lader, MD Beth Stef anchik, MD (N)	Martha Mey er, RN, MSN	Kingston, NY	Mid Valley Cardiology (1)
Sampoornima Setty, MD	Kimberly E. Halverson, RHIT	La Crosse, WI	Gundersen Lutheran Medical Center (1)
Balaji Sriniv asan, MD (N)	Christine Roraff, RN		() ()
	Jonean Thorsen, RN		
Rita Coram, MD	Anne Marie Webb, BSN	Louisville, KY	University of Louisville (1)
	Ellie Fridell, BS		
	Heidi Wilson, BS		
David Booth, MD	Yvonne Taul, RN	Lexington, KY	Lexington VA Medical Center (1)
John Kotter, MD	Caroline Rodgers, RN		
Ahmed Abdel-Latif, MD, PhD	Jennifer Isaacs, MS		
Sadiq Ahmed, MD (N)	Viktoria Bulkley, RN		
	Laura True, RN		
	Alexandra Hunter, MPH		
Michelle Ratliff, MD	Roby n Elliott	Albuquerque, NM	New Mexico V.A. Healthcare System (1)
Karen Servilla, MD (N)	Jennif er Hogan		
James J. Jang, MD	Oliv ia Anaya	San Jose, CA	Kaiser Permanente San Jose (1)
Gennie Yee, MD			
Deepa Ramaswamy, MD (N)			
Michel Georges Khouri, MD	Kristine Arges	Durham, NC	Duke University Medical Center (1)
John Middleton, MD (N)	Melissa LeFevre		
	Jennif er Tomfohr		
Jason T. Call, MD	Stephanie, M. Lane, RN, BSN, CCRN	Winchester, VA	Winchester Cardiology and Vascular Medicine, PC (1)
David Sisson, MD (N)	Jennifer L. Stanford, RN, MSN		
Prakash Deedwania, MD	Antonia Vega	Fresno, CA	UCSF - Fresno Community Regional Medical Center (1)
Kiran Reddy , MD			
Mei Hwang, MD (N)			
Stev en Weitz, MD	Stev en Giovannone	Schenectady, NY	Cardiology Associates of Schenectady P.C. (1)
Page Salanger, MD (N)	Lori Pritchard, RN		
Ray Wyman, MD	Joy Burkhardt, CCRP	Torrance, CA	Torrance Memorial Medical Center (1)
	Suellen Hosino, RN, BSN, CCRP		
Khaled Dajani, MD	Carol M. Kartje, BSN	May wood, IL	Loy ola University Medical Center (1)
Holly Mattix-Kramer, MD (N)			
Verghese Mathew, MD			
Michael D. Shapiro, DO	Ay nun Naher, MBBS, MS	Portland, OR	Oregon Health & Science University (1)
Jose Rueda, MD (N)	David Schlichting, LPN		
Omar Almousalli, MD	Elizabeth Capasso-Gulve	Fairview Heights, IL	Advanced Heart Care Group / MEDICORICIUM, L.L.C. (1)
John Lehman, MD	Alaine Melanie Loehr	. 1019110, 12	
Norbert Urbanski, MD	Marlowe Mosley		

Russia (111)

Lead Country Cardiologist Olga Bockeria, MD, PhD Lead Country Nephrologist Ev geny Shutov, MD

Alexander M. Chernyavskiy, MD, PhD	Ivan A. Naryshkin, MD	Novosibirsk	E.Meshalkin National Medical Research Center of the Ministry of Health of the Russian Federation (73)
Evgeniy I. Kretov, MD			
Igor O. Grazhdankin, MD			
Alexander Sergeevich Borisov, N	/ID (N)		
Leo A. Bockeria, MD, PhD	Olga Bockeria, MD, PhD	Moscow	National Medical Research Center for Cardiov ascuar Surgery (34)
Karen Petrosyan, MD, PhD	Zalina Kudzoev a, MD		
Evgeny Shutov, MD (N)			
Leonid L. Bershtein, MD, PhD	Irina Subbotina	Saint Petersburg	North-Western State Medical University (4)
Sergey A. Sayganov, MD, PhD	Victoria Gumerova		

Anastasia M. Kuzmina-Krutetskaya, MD Elizav eta V. Zbyshevskaya, MD, PhD Nana O. Katamadze, MD, PhD Vladimir Ry asniansky, MD (N)

Poland (105)

Lead Country Cardiologists Radoslaw Pracon, MD, PhD Marcin Demkow, MD, PhD Lead Country Nephrologist Robert Malecki, MD

Tomasz Mazurek, MD, PhD Karolina Wojtera, MD	Jakub Maksym, MD	Warszawa	Medical University of Warsaw (57)
Anna Fojt, MD			
Ewa Szczerba, MD			
Piotr Pruszczy k, MD, PhD	Andrzej Łaby k, MD	Warszawa	Department of Internal Medicine and Cardiology, Infant Jesus Teaching Hospital, Medical University of Warsaw (22)
Marek Roik, MD, PhD	Agnieszka Szramowska, MD Olga Zdończy k, MD		
Marcin Demkow, MD, PhD	Olga Walesiak	Warsaw	Coronary and Structural Heart Diseases Department, Institute of Cardiology (19)
Radoslaw Pracon, MD, PhD	Katarzy na Malinowska		
Cezary Kepka, MD PhD			
Anna Teresinska, MD PhD			
Karolina Kry czka, MD PhD			
Jan Henzel, MD PhD			
Mateusz Solecki, MD PhD			
Edyta Kaczmarska, MD PhD			
Robert Malecki, MD (N)			
Jaroslaw Drozdz, PhD	Marta Swiderek, MA	Lodz	Cardiology Clinic, Medical University in Lodz (7)
Bartosz Czarniak, MD Malgorzata Frach (formerly Stasiak), MD	Ewelina Wojtala, MA		
Konrad Szy mczyk, MD			
Iwona Niedzwiecka, MD			
Sebastian Sobczak, MD			
Tomasz Ciurus, MD			
Piotr Jakubowski, MD			
Magdalena Misztal-Teodorczy k, MD			
Dawid Teodorczyk, MD			
Aleksandra Fratczak, MD			
Marcin Szkopiak, MD			
Patry cja Lebioda, MD			
Michal Wlodarczy k, MD			
Anna Plachcinska, MD			
Jacek Kusmierek, MD			
Magdalena Miller, MD			
Halina Marciniak, MD			
Karolina Wojtczak-Soska, MD			
Katarzy na Łuczak, MD			
Tomasz Tarchalski, MD			
Anna Cichocka-Radwan, MD			

India (92) Lead Country Cardiology Balram Bhargav a, DM Lead Country Nephrologist Sandeep Mahajan, MD

Sajeev Chakanalil Govindan, MD, DNB, DM, PhD

Anjali Anand, MSc

Rajesh Gopalan Nair, MD, DNB, DM	Janitha Raj, B.Tech		
Melemadathil Srilatha, MD, DM	Bachma Bayindran MSa		
(N)	Reshma Ravindran, MSc		
	Rajalekshmi VS, MSc, MScCRRA		
Atul Mathur, MD	Ajit Singh Narula, MD		Fortis Escort Heart Institute (13)
Upendra Kaul, MD	Vijay Kher, MD	New Delhi	
Sanjeev Gulati MD, DM (N)	Puneet Sodhi, MD		
Anoop Mathew, MD	Binoy Mannekkattukudy Kurian	Kolenchery	MOSC Medical College Hospital (12)
Eapen Punnoose, MD			
TA Kishore, MD (N) Satish Sankaranarayanan, MD (N)			
Ranjan Kachru, MD	Abhishek Dubey, PGDACR	New Delhi	Fortis Healthcare Fl.tLt. Rajan Dhall Hospital (11)
Sanjeev Gulati, MD (N)			
Balram Bhargav a, DM	Chandini Suv arna, BDS	New Delhi	All India Institute Of Medical Sciences (8)
Sandeep Mahajan, MD (N)			
G.Karthikeyan, DM			
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			
Gurpreet Gulati, MD			
Sanjeev Sharma, MD			
V K Bahl, DM			
Neeraj Pandit, MD, DM	Sheromani Bajaj	New Delhi	Dr Ram Manohar Lohia Hospital (5)
Ajay Sharma, MD, DM	Vandana Yadav, Msc.PGDACR		
Niruta Sharma MD	Girish Mishra, Msc, PGDACR		
Hemant Shakhar Mahapatra MD			
Cholenahally Nanjappa	Nandita Nataraj, BE(Biotech)	Bengaluru	Sri Jay adeva Institute of Cardiovascular
Manjunath, MD, DM	PGDICRCDM	Beligalulu	Sciences and Research (4)
Nagaraja Moorthy, MD, DM	Soundary a Nayak, BE(Biotech) PGDICRCDM		
Satvic Cholenahally Manjunath, MD,DM	Mahev amma Mylarappa, GNM (General Nursing)		
Sury aprakash Narayanappa, MBBS			
Umesh Lingaraj, MD (N)			
Veerabhadra Gupta, MD (N)			
Milind Av dhoot Gadkari, MD	Sheetal Rupesh Karwa, BHMS	Pune	KEM Hospital Pune (4)
Siddharth Gadage, MD DNB	Suv arna Kolhe, MSc		
Tapan Umesh Pillay, BHMS MSc			
NSC Valentine Lobo, MD (N)			
Johann Christopher, MD, DNB	K. Manjula Rani, MSc.	Hy derabad	Gurunanak CARE Hospital (3)
Nirmal Kumar, MD, DM	M. Sowjany a Reddy, BSc		
Suresh Kumar, MD, DM (N)	K. Preethi, BSc		
John Jose, MD	Anu Tharini	Vellore	Christian Medical College (3)
Vinoi George David, MD (N)	Anandaroop Lahiri		
· · ·	· ·	Ludbione	Hero DMC Heart Institute, Dayanand Medical
Gurpreet S. Wander, DM	Baljeet Kaur, MSc (Biotechnology)	Ludhiana	College and Hospital (2)
Rohit Tandon, MD	Sonika Gupta, MBA, B. Pharmacy		

Sarju Ralhan, M.Ch (CTVS) Naved Aslam, DM Abhishek Goyal, DM Vikas Makkar, DM (N)			
S.K. Dwiv edi, DM V.S. Narain, DM Sharad Chandra, DM	Roma Tewari, PG Meenakshi Mishra, PG Shiv ali Patel Suman Singh, PG	Lucknow	King George's Medical University, Departmer of Cardiology (2)
Johann Christopher, MD Praneeth Polamuri, MD Vikranth Reddy, MD, DNB (N)	Sowjany a Reddy Manjula Rani	Hy derabad	CARE Hospital (1)
Upendra Kaul, MD	Priy adarshani Arambam Bebek Singh	New Delhi	Batra Hospital and Medical Research Centre (BHMRC) (1)
Hong Cheng, MD Weijing Bian, MD Guoqin Wang , MD	Jing Dong, MD Xiaoy i Xu, MD	Beijing	Beijing Anzhen Hospital (24)
Jiyan Chen, MD Zhiming Ye, MD (N)	Haojian Dong Peiy u He Chunli Xia Junqing Yang Qi Zhong	Guangzhou	Guangdong General Hospital (15)
Xin Fu, MD Zhangsuo Liu, MD (N)	Dan Gao Dengke Jiang Ran Leng Xutong Wang Qianqian Y uan Lili Zhang	Zhengzhou	The First Affiliated Hospital of Zhengzhou University (13)
Shuy ang Zhang, MD, PhD Zheny u Liu, MD Xuemei Li, MD (N)	Ying Wang, MD Yechen Han, MM Lihong Xu, RN Zheny u Liu Gang Chen, MD Rongrong Hu	Beijing	Peking Union Medical College Hospital (11)
Yitong Ma, MD (N) Yining Yang, MD	Dongze Li Xiaomei Li Xiang Ma Zixiang Yu Qian Zhao	Urumqi	First Affiliated Hospital of Xinjiang Medical University (7)
Carlo Briguori, MD	Francesca De Micco	Naples	Clinica Mediterranea (52)
Gian Piero Perna, MD Marco Marini, MD Gabriele Gabrielli, MD Mario D'arezzo, MD (N)	Francesca Pietrucci, PhD	Ancona	Cardiology and CCU - Ospedali Riuniti Ancor (7)
Mario D'arezzo, MD (N) Marco Sicuro, MD Valentina Pellu, MD (N)	Gianpiero Leone, MD Francesco Pisano, MD Cristina Bare, BSc	Aosta	Ospedale Regionale Umberto Parini (1)

China (70)

Lead Country Cardiologist Lixin Jiang, MD, PhD Lead Country Nephrologists Xuemei Li, MD

Italy (62)

Lead Country Cardiologist Francesco Orso, MD

Paolo Calabro, MD	Fabio Fimiani	Napoli	AORN Dei Colli "V. Monaldi" UOC Cardiologia Università della Campania "L.Vanvitelli" (1)
Tiziana Formisano, MD Piero Tassinario, MD (N)			
Marcello Galv ani, MD	Chiara Attanasio	Forli	Ospedale "G.B. Morgagni – L. Pierantoni" For (AUSL della Romagna) (1)
Filippo Ottani, MD Marco De Fabritis, MD (N)			
Juan Manuel López Quijano, MD, MSc Alejandro Chev aile R amos, MD	Teresa Delgadillo	San Luis Potosi	Hospital Central Dr. Ignacio Morones Prieto (16)
(N) Jorge Carrillo Calvillo, MD			
Jorge Escobedo, MD	Ramon de Jesús-Pérez, RN	Benito Juarez	Instituto Mexicano del Seguro Social (10)
Rubén Baleón-Espinosa, MD Arturo S Campos-Santaolalla, MD			
Elihú Durán-Cortés, MD			
José M Flores-Palacios, MD			
Andrés García-Rincón, MD			
Moisés Jiménez-Santos, MD			
Joaquín V Peñafiel, MD			
José A Ortega-Ramírez, MD			
Aquiles Valdespino-Estrada, MD			
	María Pérez García	Mexico City	Instituto Nacional de Cardiología "Ignacio Chávez" (2)
Erick Alexánderson Rosas, MD			
Magdalena Madero Rovalo, DM			
Erick Alexánderson Rosas, MD Magdalena Madero Rov alo, DM (N) Guillermo Garcia-Garcia (N)	Lorena Lopez, BS	Guadalajara	Hospital Civil de Guadalajara Fray Antonio Alcalde (2)

Canada	(24)
--------	------

Lead Country Cardiologists Akshay Bagai, MD, MHS Kev in R. Bainey, MD, MSc Lead Country Nephrologist Ron Wald, MDCM, MPH

Mexico (30) Lead Country Cardiologist Jorge Escobedo, MD Lead Country Nephrologist Magdelena Madero, MD

Kev in R. Bainey, MD, MSc	Norma Hogg, RN	Edmonton, AB	University of Alberta (15)
Neesh Pannu, MD (N)	Suzanne Welsh, RN		
Asim N. Cheema, MD, PhD	Khry styna Kushniriuk, HBSc, MD	Toronto, ON	St. Michael's Hospital (3)
Akshay Bagai, MD, MHS	Mohammed Hussain		
Ron Wald, MDCM, MPH (N)	Olugbenga Bello		
Shaun Goodman, MD, MSc			
John Joseph Graham, MRCP,			
MB ChB, BSc Mark Peterson, MD, FRCSC,			
PhD			
Chi-Ming Chow, MD, CM, MSc			
Beth Abramson, MD, MSc			
Graham Wong, MD	Jackie Chow, BSN	Vancouver, BC	Vancouv er General Hospital (2)
Kenneth Gin, MD	Andrew Starov oytov, MD		
Christopher Fordyce, MD	Naomi Uchida, BSN		
	Ngaire Meadows		
Ariel Diaz, MD	Isabelle Roy, RN	Trois-Rivieres, QC	Centre Hospitalier de Regional Trois-Rivieres (1)
Philippe Rheault, MD	Patricia Alarie, RN		
Alejandro Gisbert, MD	Linda Arcand, RN		

Alain Raymond, MD	Estelle Montpetit		
Yanek Pépin-Dubois, MD			
Miguel Barrero, MD			
Carl-Éric Gagné, MD			
Mark Garand, MD			
Ricardo Costa, MD			
Catherine Lemay, MD			
Ying Tung Sia, MD			
Pierre Gervais, MD			
Alain Rheault, MD			
Pallav Garg, MBBS, MSc	Sandy Carr, RN	London, ON	London Health Sciences Centre (1)
Matthew Weir, MD (N)	Catherine Bone, RN		
Amar Uxa, MD	Nadia Asif	Toronto, ON	University Health Network (1)
Michael Farkouh, MD			
Christopher Chan, MD (N)	Suzana Tav ares		
			Centre Intégré Universitaire De Santé et de
Philippe Généreux, MD	Chantale Mercure, RN	Montréal, QC	Services Sociaux du Nord de l'île de Montréal /Hôpital du Scaré-Cœur de Montréal (1)
Jean Diodati, MD			
François Madore, MD (N)			

Singapore (14)

Lead Country Cardiologist Kian-Keong Poh, MD Lead Country Nephrologist Titus Lau, MD

Kian-Keong Poh, MD		Singapore	National University Heart Center Singapore (11)
Ping Chai, MD			
Titus Lau, MD (N)			
Joshua P. Loh, MD			
Edgar L. Tay , MD			
Kristine Teoh, MD	Sik-Yin V Tan, BSc		
Ly nette L. Teo, MD	Winnie C Sia, BSc		
Ching-Ching Ong, MD	Audrey W Leong, BSc		
RaymondC. Wong, MD			
Poay -Huan Loh, MD			
Theodoros Kofidis, MD			
Wan Xian Chan, MD			
Koo Hui Chan, MD			
David Foo, MBBS	Li Hai Yan, RN	Singapore	Tan Tock Seng Hospital (2)
Jason Loh Kwok Kong, MD			
Ching Min Er, MD			
Fahim Haider Jafary, MD			
Tracy Tan, MD (N)			
Terrance Chua, MD	Nasrul Ismail	Singapore	National Heart Centre Singapore (1)
	Min Tun Ky aw		
	Deborah Yip		
d)			
Whady Hueb, MD	My rthes Emy Takiuti, RN	Sao Paulo	Heart Institute (InCor) University of São Paulo (6)

Brazil (13) Lead Country Cardiologist

Renato D. Lopes, MD, PhD Lead Country Nephrologists Maria Eugenia Canziani, MD (Lead) Sergio Draibe, MD (Co-Lead)

> Eduardo Gomes Lima, MD Paulo Cury Rezende, MD Expedito Eustáquio Ribeiro Silv a, MD Alexandre Ciappina Hueb, MD

Marianna D. A. Dracoulakis, MD, PhD	Natalia S Oliveira, RN	Salv ador	Hospital da Bahia (5)
Rodolf o G. S. D Lima, MD			
Paulo Novis Rocha, MD (N)			
Alexandre Schaan de Quadros, MD		Porto Alegre	Instituto de Cardiologia de Porto Alegre (1)
Renato Abdala Karam Kalil, MD	Aline Peixoto Deiro		
José Luiz da Costa Vieira, MD	Alice Manica Muller		
Gabriel Grossmann , MD	Maria Antonieta Pereira de Moraes		
Pedro Píccaro de Oliveira, MD	Bruna Maria Ascoli		
Leonardo Bridi, MD	Sílvia Zottis Poletti		
Simone Savaris, MD			
Renato George Eick, MD (N)			
Paola Emanuela Poggio Smanio, MD, PhD	Leonardo Pizzol Caetano, PhD	São Paulo	Instituto Dante Pazzanese de Cardiologia (1)
Leda Lotaif, MD, PhD (N)			

Hungary(12)

Lead Country Cardiologist Andras Vertes, MD Lead Country Nephrologist Peter Voros, MD

Andras Vertes, MD	Judit Sebo, MD	Budapest	Eszszk-Szent Istvan Hospital (10)
Peter Voros, MD (N)	Zoltan Davidovits, MD		
	Laszlone Matics		
Bela Merkely, MD, PhD, DSc	Andrea Barty kowszki, MD	Budapest	Heart and Vascular Center, Semmelweis Univ ersity (1)
Mihaly Tapolyai, MD (N)	Pal Maurov ich-Horvat, MD, PhD, MPH		
Albert Varga, MD, PhD	Gergely Ágoston, MD	Szeged	University of Szeged (1)
Timea Boros, MD (N)			

Lithuania (12)

Lead Country Cardiologist Jelena Celutkiene, MD Lead Country Nephrologist Marius Miglinas, MD, PhD

Aleksandras Laucevicius, MD	Agne Juceviciene, MD	Vilnius	Vilnius University Hospital Santariskes Clinic (12)
Jelena Celutkiene, MD	Irma Kalibataite-Rutkauskiene, MD		
Marius Miglinas, MD (N)	Laura Keinaite		
	Monika Lauky te		
	Gelmina Mikolaitiene		
	Akvile Smigelskaite, MD		
	Ilona Tamasauskiene, MD		
	Agne Urboniene, MD		

Lisbon

Hospital de Santa Marta / Hospital Curry Cabral (8)

Portugal (10)

Lead Country Cardiologist Ruben Ramos, MD Lead Country Nephrologist Fernando Nolasco, PhD

Ruben Ramos, MD	Maf alda Selas	
Duarte Cacela, MD	Filipa Silv a	
Ana Santana, MD	Cláudia Freixo	
Antonio Fiarresga, MD		
Lidia Sousa, MD		
Hugo Marques, MD		
Lino Patricio, MD		
Luis Bernanrdes, MD		
Pedro Rio, MD		
Ramiro Carvalho, MD		
Rui Ferreira, MD		

	Tiago Silv a, MD Ines Rodrigues, MD Pedro Modas, MD Guilherme Portugal, MD Jose Fragata, MD Marina Vieira , MD Fernando Nolasco, PhD (N) Marina Vieira., MD Fernando Caeiro, MD			
	Pedro Farto e Abreu, MD	Maura Carina Nédio, BSc	Amadora	Hospital Professor Doutor Fernando Fonseca, EPE (1)
	Sérgio Brav o Baptista, MD, PhD Miguel Borges Santos, MD Patricia Carrilho, MD (N)			
	Fausto J. Pinto, PhD	Inês Zimbarra Cabrita, PhD	Lisbon	Santa Maria University Hospital, Cardiology Department, CHLN (1)
	Miguel Nobre Menezes, MD Guilhermina Cantinho Lopes, MD	Andreia Rocha, MSc Francisca Patuleia Figueiras, PhD		
	Ana Gomes Almeida, PhD	Andreia Coelho, BSc		
	Pedro Canas Silv a, MD	Marta Capinha		
	Angelo Nobre, MD	Maria Inês Caetano		
	Ana Rita Francisco, MD Jose Lopes, MD (N)	Susana Silva		
Spain (9) Lead Country Cardiologist Almudena Castro, MD Lead Country Nephrologist Raf ael Selgas, MD				
	Jose Lopez-Sendon, MD, PhD Almudena Castro, MD Elena Ref oyo Salicio, MD Gabriela Guzman, MD Gabriel Galeote, MD Silv ia Valbuena, MD Raf ael Selgas, MD (N)	Virginia Fernández-Figares, Pharm	Madrid	Hospital La Paz. IdiPaz (6)
	Jesús Peteiro, MD, PhD	Moisés Blanco-Calvo, PhD	A Coruna	Complexo Hospitalario Universitario A Coruña (CHUAC) Sergas, Department of Cardiology. INIBIC A Coruña. CIBER-CV. Universidad de A Coruña, Spain (2)
	María Dolores Martínez-Ruíz,	Encarnación Alonso-Álvarez, BSc		
	MD Ruth Pérez-Fernández, MD	Paula García-González, BSc		
	José J Cuenca-Castillo, MD Xacobe Flores-Ríos, MD Óscar Prada-Delgado, MD Gonzalo Barge-Caballero, MD Miguel Perez Fontan, MD (N)			
	Vicente Miro, MD Jose L Diez, MD Pilar Calv illo, MD	Begoña Igual, MD	Valencia	Hospital Universitario y Politecnico La Fe (1)
	Julio Hernandez Jaras, MD (N)			
Argentina (6) Lead Country Cardiologist Luis Guzman, MD Lead Country Nephrologist Raf ael Maldonado, MD				
	Mariano Rubio, MD Raf ael Maldonado, MD (N)	Graciela Scaro, MD	Cordoba	Clínica Privada Vélez Sarsfield (5)
	Julio César Figal, MD	Matías Nicolás Mungo	Ciudad Autonoma de Buenos Aires	Fundación Favaloro (1)

Oscar Méndiz, MD Claudia Cortés, MD Roberto René Favaloro, MD Pablo Raffaele, MD (N)

Peter Sizeland, MD (N)

France (6)

Lead Country Cardiologist Emmanuel Sorbets, MD, PhD Lead Country Nephrologist Eric Daugas, MD, PhD

Philippe Gabriel Steg, MD	Helene Abergel, MSc	Paris	Bichat Hospital (4)
Jean-Michel Juliard, MD	Axelle Fuentes, MSc		
Eric Daugas, MD, PhD (N)			
Emmanuel Sorbets, MD, PhD			
Christophe Thuaire, MD	Corine Thobois, RN	Chartres	C.H. Louis Pasteur (2)
Téodora Dutoiu, MD	Emilie Tachot, RN		
Catherine Albert, MD (N)	Christophe Laure, RN		
Bougrida Hammouche, MD (N)	Christel Vassaliere, RN		
Gerard Patrick Devlin, MD	Liz Low, RN	Hamilton	Waikato Hospital (6)
Raewyn Fisher, MD	Jay ne Scales, RN		

Kirsty Abercrombie, RN

United Kingdom (5)

New Zealand (6) Lead Country Cardiologist Gerard Patrick Devlin, MD Lead Country Nephrologist Peter Sizeland, MD

Lead Country Nephrologist David Wheeler, MD

Roxy Senior, MBBS, MD, DM	Grace M. Young , MSc, BSc (Hons)	Harrow	Northwick Park Hospital Harrow/ Royal Brompton Hospital London (2)
Ahmed Elghamaz, MB BCh	Christopher Kinsey		
Sothinathan Gurunathan, MBChB	Raisa Kavalakkat, MSc, BSc, RN		
Nikolaos Karogiannis, MBBS	Jo Evans, RN		
Benoy N Shah, MD, MBBS, BSc (Hons)	Ikraam Hassan, RN		
Richard HJ Trimlett, MBBS, CCST	Emma Howard, MSc, BSc		
Michael B Rubens, LRCP, MRCS, MBBS, DMRD	Ann Banfield, BSc, RN		
Edward D Nicol, MD, BMedSci, MBBS, DTM&H	Reinette Hampson, BSc (Hons), BA	(Hons)	
Tarun K Mittal, MD	Rory Collins, BSC		
Neill Duncan, MD (N)	Anastasia Vamvakidou, MBBS, MRCP		
Reto Andreas Gamma, MBBS	Sarah Williams, RN	Chelmsford	Broomfield Hospital (1)
Sumith Abey gunasekara, MD (N)	Kim Holland, RN		
	Karen Swan, RN		
Khaled Alfakih, MBBS, MD	Abigail Knighton, BSc., PG Dip.	London	King's College NHS Foundation Hospital (1)
Jonathan Byrne, PhD	Katherine Martin, RGN, Dip. N, MSc		
Ian Webb, PhD, MA (N)			
Dwayne S. G. Conway, MD	Judith Wright	Wakefield	Pinderfields Hospital (1)
	Donna Exley		

Serbia (5)

Lead Country Cardiologist Branko D. Beleslin, MD, PhD Lead Country Cardiologist

Sanja Simic Ogrizovic, MD

Branko D. Beleslin, MD, PhD	

Nikola N. Boskovic, MD Marija T. Petrovic, MD Milan R. Dobric, MD Zeljko Z. Markovic, MD, PhD Ana S. Mladenovic, MD, PhD Sanja Ogrizovic, MD (N)

Salamah Alfalahi, MD

Ana D. Djordjevic-Dikic, MD, PhD Vojislav L. Giga, MD, PhD Jelena J. Stepanovic, MD, PhD Belgrade

Faculty of Medicine, University of Belgrade; Cardiology Clinic, Clinical Center of Serbia (5)

Australia (4)				
Lead Country Cardiologist				
Joseph B. Selvanayagam, MBBS	S (Hons), Dphil			
Lead Country Cardiologist				
Magid Fahim, MBChB, FRACP	Jacob R. Colyong agom			Flinders Medical Centre and College of
	Joseph B. Selvanayagam, MBBS(Hons), DPhil	Sau Lee, PhD	Adelaide	Flinders Medical Centre and College of Medicine and Public Health (4)
	Majo X. Joseph, MBBS	Prince Thomas, RN		
	Jonathan M Gleadle, BM Dphil			
Austria (4)	(N)			
Lead Country Cardiologist				
Herwig Schuchlenz				
	Herwig Schuchlenz, MD	Gudrun Steinmaurer	Graz	LKH Graz West Austria (3)
	Stef an Weikl, MD		Oldz	
			<i></i>	Medical University of Vienna, Department of
	Irene Marthe Lang, MD	Max-Paul Winter, MD	Vienna	Cardiology (1)
Belgium (4)				
Lead Country Nephrologist				
Kathleen Claes, MD, PhD				
	Kaatje Goetschalckx, MD	Valerie Robesyn	Leuv en	University Hospital Leuven (3)
	Frans Van de Werf, PhD, MD	valono reobcogn	Louvon	
	Kathleen Claes, MD, PhD (N)			
	Christiaan Vrints, MD	Nathalie Brosens	Edegem	Universitair Ziekenhuis Antwerpen (1)
	Bharati Shivalkar, MD		Edogom	
	Amary Ilis Van Craenenbroeck, N	MD (N)		
Israel (4)				
	Yaron Arbel, MD	Daniela Puzhev sky	Tel Aviv	Tel Aviv Sourasky Medical Center (4)
	Doron Schwartz, MD (N)	Miri Revivo		, , , , , , , , , , , , , , , , , , , ,
	Orit Kliuk, MD			
Egypt (3)				
571-(1)	Magdy Abdelhamid, MD	Ahmed Talaat, MD	Cairo	Cairo University (3)
	Ahmed Kamal, MsC			
	Hossam Mahrous, MD			
	Mohamed Adel , MsC			
	Hussien El Fishawy, MD (N)			
United Arab Emirates (2)				
	Wael A. Almahmeed, MD	Virendra Misra, MD	Abu Dhabi	Sheikh Khalif a Medical City (2)
	Mohamed Hassan, MD (N)			
	Seema Nour, MD			
	Abdallah M. Abdallah, MD			

Germany (1) Lead Country Cardiologist Rolf Doerr, MD				
	Rolf Doerr, MD	Karin Ploetze, PhD	Dresden	Praxisklinik Herz und Gefaesse (1)
	Gregor Simonis, MD, PhD	Franziska Guenther		
	Juergen Stumpf, MD	Kerstin Bonin		
	Clemens T. Kadalie, MD	Kerstin Mikes, RN		
	Klaus Matschke, MD, PhD	Katharina Knaut		
	Doreen Reimann, MD (N)			
Macedonia (1)				
	Sasko Kedev , MD, PhD Irena Peov ska Mitevska, MD, PhD		Skopje	University Clinic of Cardiology (1)
	Elizabeta Srbinov ska Kostovska	a, MD, PhD		
	Hristo Pejkov, MD, PhD			
	Zvezdana Petronijevic, MD (N)			
	Liljana Tozija, MD (N)			
Netherlands (1)				
	Robert K. Riezebos, MD, PhD		Amsterdam	Cardio Research Hartcentrum OLVG (1)
	Pouneh Samadi, MD	Jeannette, J. M. Schoep, RN		
	Elise v an Dongen, MD	Elisabeth, M. Janzen, RN		
	Sander R. Niehe, MD			
	Yves Smets, MD (N)			
Romania (1)				
	Calin Pop, MD, PhD		Bucharest	Emergency County Hospital Baia Mare (1)
	Matei Claudia, MD, PhD			
Sweden (1)	Florina Chereches, MD (N)			
Lead Country Cardiologist Claes Held, MD, PhD				
	Claes Held, MD, PhD	Christina Björklund, RN	Uppsala	Uppsala University (1)
	Axel Åkerblom, MD, PhD Inga Soveri, MD, PhD (N)	Maria Andreasson, RN		

* (N) = Nephrologist