

# Chest Pain of Recent Onset Insights from the ISCHAEMIA Trial

Dr Rosica Panayotova Consultant Cardiologist Stockport NHS Foundation Trust June 2023





### Coronary artery disease – clinical presentation

#### **Chronic Coronary Syndromes**

Stable plaque

#### **Acute Coronary Syndromes**

• Unstable plaque

NSTEMI

that results in injury and infarct to the

subendocardial myocardium.

During an NSTEMI, the plaque rupture and throm-

bus formation causes partial occlusion to the vessel

3

 Plaque rupture → acute vessel occlusion



ECG

TROPONINS

#### 1 STABLE ANGINA

Angina pain develops when there is increased demand in the setting of a stable atherosclerotic plaque. The vessel is unable to dilate enough to allow adequate blood flow to meet the myocardial demand.



The plaque ruptures and a thrombus forms around the ruptured plaque, causing partial occlusion of the vessel. Angina pain occurs at rest or progresses rapidly over a short period of time.

Supply ischemia, no infarct



Subendocardial infarct



Normal, Inverted T waves, or ST depression

Elevated

STEMI
A STEMI is characterized by compl

A STEMI is characterized by complete occlusion of the blood vessel lumen, resulting in transmural injury and infarct to the myocardium, which is reflected by ECG changes and a rise in troponins.



Hyperacute T waves or ST elevation

Elevated

Normal

Normal

Normal

Normal, Inverted T waves, or ST depression

This infographic was created by Paula Sneath and Leah Zhao for the Sirens to Scrubs series of CanadiEM.org.

Demand ischemia, no infarct



## Patient Mr X, 68 year old male

- Previous medical history of:
  - Hypertension on Lisinopril 10mg od
  - Smoker 5-10/day
- <u>Presents with:</u>
  - Fatigue, SOBOE and occasional chest discomfort following strenuous exertion
  - Can now only manage 9 holes of golf
  - Concerned as younger brother had a MI recently



# What would you do?

- 1. Reassure and advise smoking cessation?
- 2. Organise tests?
- 3. Prescribe medication and re-review?
- 4. Refer to General Cardiology clinic?
- 5. Refer to RACPC? (\*\*Does he meet referral criteria?)
- 6. Prescribe medication and refer?



# The ISCHEMIA study

320 sites from 37 countries: > 5000 patients



International Study Of Comparative Health Effectiveness With Medical And Invasive Approaches (ISCHEMIA)

**Primary Report of Clinical Outcomes** 

Funded by the National Heart, Lung, and Blood Institute

NYU School of Medicine

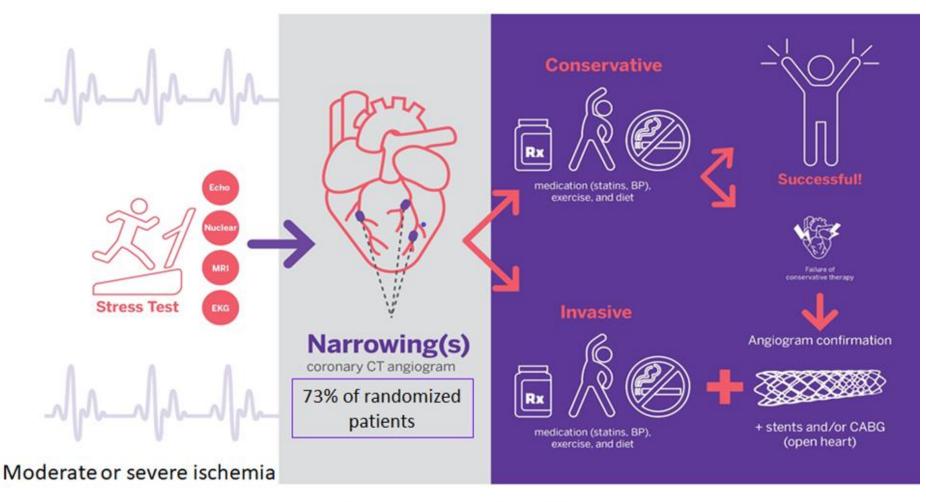
NYULangone #AHA19 Health

#### **Question:**

In patients with with <u>at least moderate ischemia on a stress test</u>, is there a benefit to adding cardiac catheterization and revascularization to optimal medical therapy?



## ISCHAEMIA trial design overview









## Endpoints

### Primary Endpoint:

 Time to CV death, MI, hospitalization for unstable angina, heart failure or resuscitated cardiac arrest

### **Major Secondary Endpoints:**

- Time to CV death or MI
- Quality of Life

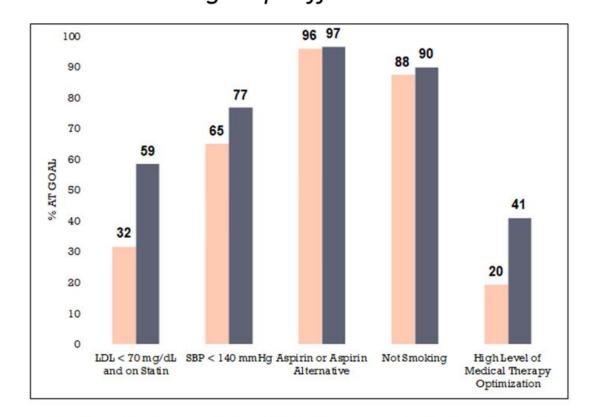
### **Other Endpoints include:**

- All-Cause Death
- Net clinical benefit (stroke added to primary endpoint)





### Risk Factor Management Baseline vs last visit No between group differences INV vs CON

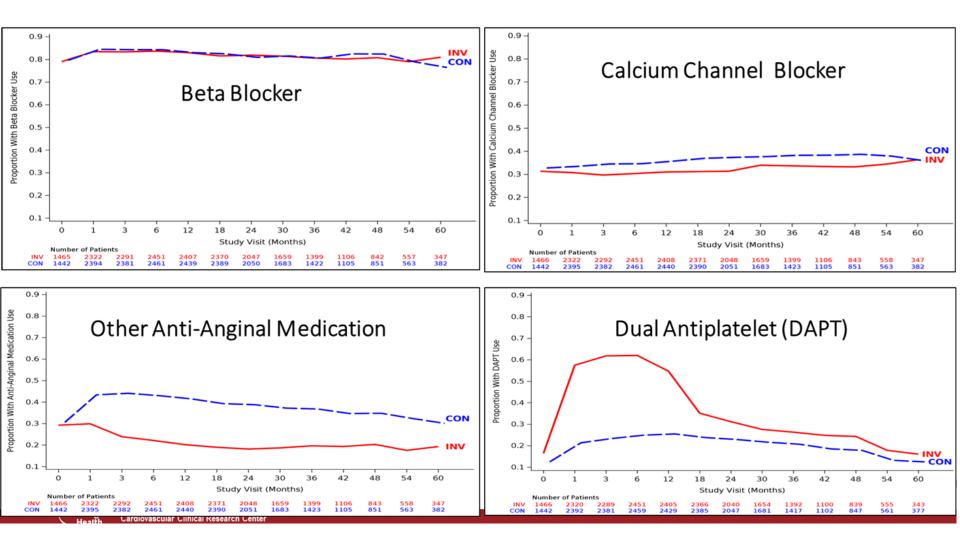


High Level of Medical Therapy Optimization is defined as a participant meeting all of the following goals: LDL < 70 mg/dL and on any statin, systolic blood pressure < 140 mm/Hg, on aspirin or other antiplatelet or anticoagulant, and not smoking. High level of medical therapy optimization is missing if any of the individual goals are missing.



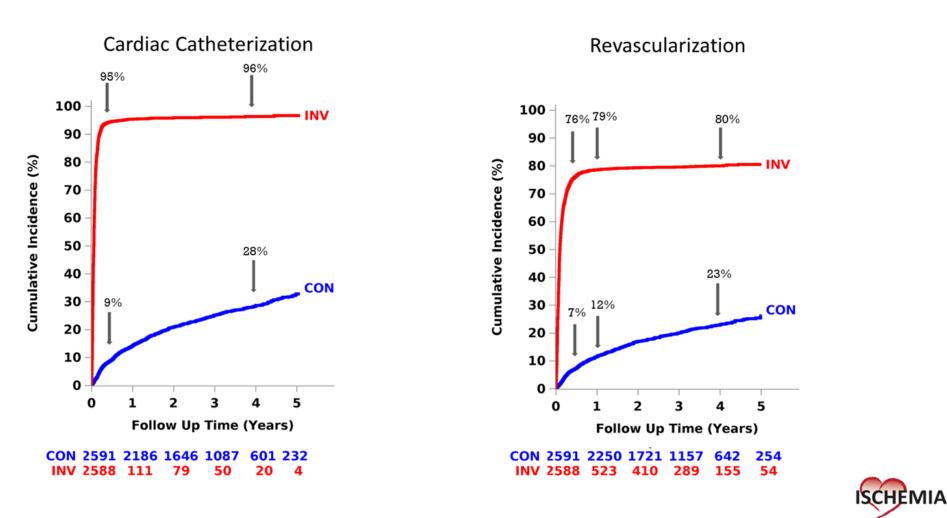


## Medication use over time

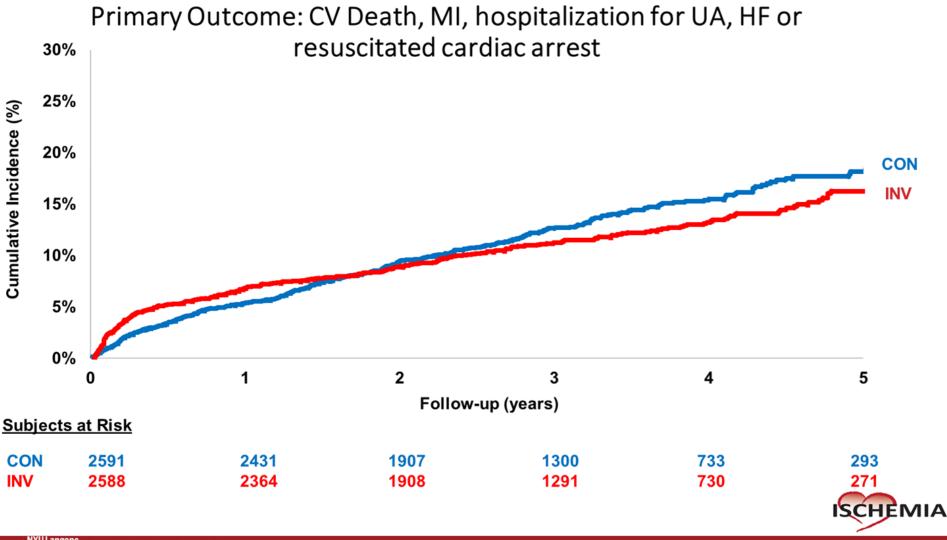




## Angiography and revascularisation



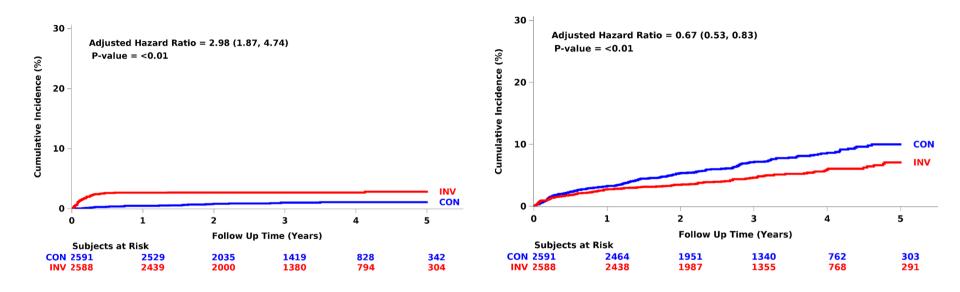






#### Procedural MI

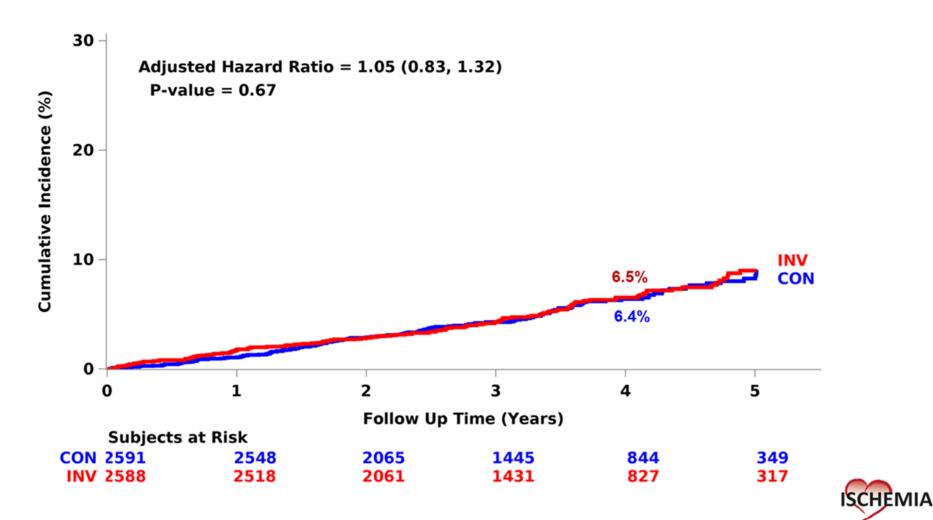
#### Spontaneous MI







## All cause mortality

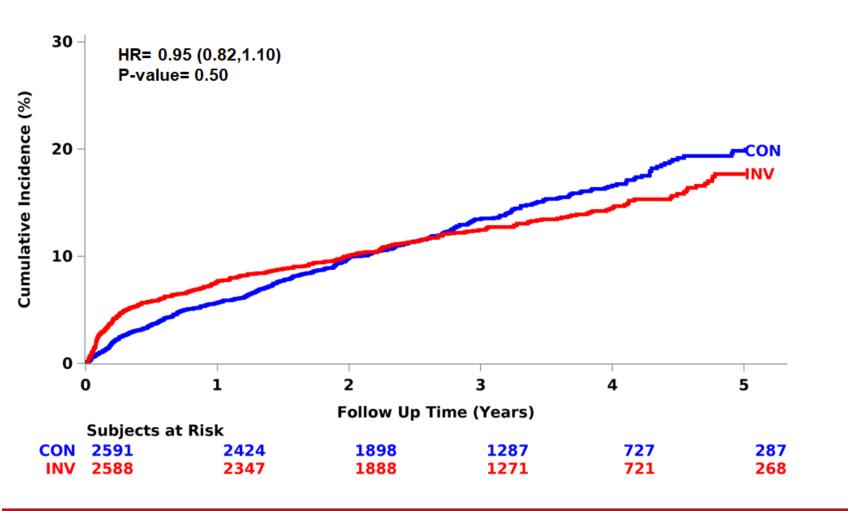






ISCHEMIA

## Net clinical benefit: CV death, MI, UA, HF, Stroke







# Conclusions

- ISCHEMIA is the largest trial of an invasive vs conservative strategy for patients with stable CAD
- Overall, an initial INV strategy vs an initial CON strategy did <u>not</u> demonstrate a reduced risk over the 3-4 year FU:
  - Primary endpoint CV death, MI, hospitalisation for UA, HF, RCA
  - Major Secondary endpoint CV death or MI



### Patient Mr X – symptoms suggestive of angina What would you do?

- 1. Reassure and advise smoking cessation?
- 2. Organise tests?
- 3. Prescribe medication and re-review?
- 4. Refer to General Cardiology clinic?
- 5. Refer to RACPC? (\*\*Does he meet referral criteria?)
- 6. Prescribe medication and refer?

## The RACPC



**Stepping Hill Hospital** 

Cardiology Department – Telephone number 0161 419 5077

**Rapid Access Chest Pain Clinic Referral Form** 

**Exclusion Criteria** – the following patients are not appropriate for this clinic but please consider either A&E or OP consultant cardiology referral instead

- Suspected or known significant aortic stenosis
- Exercise induced syncope or palpitations
- Known anaemia (Hb <100g/l)</li>
- Uncontrolled BP (persistently >200/100mmg Hg)
- NYHA Grade 3 or 4 heart failure
- Patients assessed in our RACPC within the last 24 months and had a negative test
- Patient has dizziness, breathlessness or palpitations
- If the patient is currently under follow-up with a cardiologist for another heart problem and develops chest pain



### The RACPC

Please ensure the following have been completed – incomplete referrals may cause delay

Reason for referral:

Medication:

Recent ECG: (within last 2 weeks)

Recent BP:

Interpreter required:

Recent bloods: (within 2 weeks, only attach if not analysed at SHH, to include U&E's, LFT, lipids, FBC, TFT's)

## The RACPC



Offer people optimal drug treatment for the initial management of stable angina. Optimal drug treatment consists of one or two anti-anginal drugs as necessary plus drugs for secondary prevention of cardiovascular disease.

Recommend aspirin 75 mg OD and atorvastatin 40mg OD if not already on these.

Choice	drug	dose
1 <sup>st</sup>	Bisoprolol	1.25 to 5 mg OD
2 <sup>nd</sup>	Amlodipine (or other calcium channel antagonist)	5 mg OD
3 <sup>rd</sup>	Isosorbide mononitrate Ranolazine	20 mg bd (am and lunchtime) 375mg BD (to 500mg bd after 2 weeks if well tolerated)

Nice Guidance: Stable angina: management

Clinical guideline [CG126] Published: 23 July 2011 Last updated: 25 August 2016

Guidance | Stable angina: management | Guidance | NICE

## Conclusion – *emphasis on 'OMT'*



In patients with suspected angina:

- The most important intervention, in the first instance, is '<u>medical</u> <u>optimisation</u>' – <u>OMT</u>:
  - prompt initiation on anti-ischaemic medical therapy (symptomatic & prognostic benefit)
  - risk factor and comorbidity management
- Initial assessment, incl tests to exclude other conditions:
  - AF
  - Valve disease (severe aortic stenosis)
  - Anaemia
  - Hyperthyroidism
- Onward referral (RACPC) for ischaemic testing, risk stratification & further treatment