

Box 1 Definition of angina (NICE guidelines)¹⁰

Typical angina: (requires all three)

1. Constricting discomfort in the front of the chest or in the neck, shoulders, jaw or arms.
2. Precipitated by physical exertion.
3. Relieved by rest or sublingual glyceryl trinitrate within about 5 min
 - Presence of two of the features is defined as atypical angina.
 - Presence of one or none of the features is defined as non-anginal chest pain.
 - Stable angina may be excluded if pain is non-anginal provided clinical suspicion is not raised based on other aspects of the history and risk factors.
 - Do not define typical, atypical and non-anginal chest pain differently in men and women or different ethnic groups.

in support of numerical equivalency between these indices suggesting all can be used to guide treatment strategy.¹¹ Angina with underlying obstructive CAD allows symptom guided myocardial revascularisation (often with percutaneous coronary intervention (PCI)) and is effective in reducing ischaemic burden and symptoms (in many patients). Recent studies have served evidence that functional coronary disorders overlap and may contribute to angina even in patients with obstructive epicardial CAD. Dynamic changes in lesion or vessel ‘tone’ and propensity to vasoconstriction is important and may cause rest angina that is frequently overlooked in patients with obstructive CAD.¹² During invasive physiological assessment of ischaemia during exercise, *Assres et al* showed that ischaemia developed at FFR averaging=0.76 which is not often observed with adenosine induced hyperaemia.¹³ This finding implies there are other important drivers of sub-epicardial ischaemia (myocardial supply-demand factors). Furthermore, it reinforces that angina is not synonymous with ischaemia or flow-limiting coronary disease (eg, abnormal FFR or NIFPR). Coronary anatomy and physiology should not be considered in isolation but in the context of the patient.

Angina-myocardial ischaemia discordance

Although obstructive CAD or microvascular dysfunction may be present, the link between ischaemia and angina is not clearcut. The ‘ischaemic threshold’ (the heart rate-blood pressure product at the onset of angina) has intraindividual and inter-individual variability.¹⁴ Innate differences in vascular tone and endocrine changes (eg, menopause) may influence propensity to vasospasm while environmental factors including cold environmental temperature, exertion and mental stress are relevant. The large international CLARIFY registry highlighted the importance of symptoms, showing that angina with or without concomitant ischaemia, was more predictive of adverse cardiac events compared with silent ischaemia alone.¹⁵ Other potential drivers of discordance between angina and ischaemia include variations in pain thresholds and cardiac innervation (eg, diabetic neuropathy).

Symptoms and/or signs of ischaemia but no obstructive coronary artery disease (INOCA)

Cardiologists are inclined to adopt a ‘stenosis centric’ approach to patient management; however, as clinicians we must appreciate that all factors are relevant, including coronary anatomy and function but systemic health and the psychosocial background (Figure 2). First, systemic factors including heart rate, blood pressure (and their product) and myocardial supply-demand ratio (Bockberg index) are relevant.¹⁶ Reduced myocardial oxygen supply from problems such as anaemia or hypoxaemia should always be considered.

Second, coronary factors are well recognised but certain nuances are overlooked. In 2018, the first international consensus guidelines clarify that a definite diagnosis of MVA may be made in patients with angina with no underlying obstructive CAD, evidence of reversible ischaemia on functional testing and objective evidence of coronary microvascular dysfunction (table 1).¹⁷ ‘Probable MVA’ is defined by three of the above criteria. Coronary microvascular dysfunction may be structural



Figure 1 Reappraisal of ischaemic heart disease pathophysiology. Distinct functional and structural mechanisms can affect coronary vascular function and frequently coexist leading to myocardial ischaemia. CAD, coronary artery disease.

Contemporary Diagnosis And Management Of Bronchitisa

L Cohen



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