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Genetics of Coronary Artery Disease and Myocardial Infarction

Xuming Dai, Szymon Wiernek, James P. Evans, Marschall S. Runge

Abstract

Atherosclerotic coronary artery disease (CAD) comprises a broad spectrum of clinical entities that include asymptomatic subclinical atherosclerosis and its clinical complications, such as **25** angina pectoris, myocardial infarction (MI) and sudden cardiac death. **47** CAD continues to be the leading cause of death in industrialized society. The long-recognized familial clustering of CAD suggests that genetics **3** plays a central role in its development, with the heritability of CAD and MI estimated at approximately 50 to 60%. Understanding the genetic architecture of CAD and MI has proven to be difficult and costly due to the heterogeneity of clinical CAD and the underlying multi-decade complex pathophysiological processes that involve both genetic and environmental interactions. This review describes the clinical heterogeneity of CAD and MI to clarify the disease spectrum in genetic studies, **50** provides a brief overview of the historical understanding and estimation of the heritability of CAD and MI, recounts major gene discoveries of potential causal mutations in familial CAD and MI, summarizes CAD and MI-associated genetic variants identified using candidate gene approaches and genome-wide association studies, and summarizes the current status of the construction and validations of genetic risk scores for lifetime risk prediction and guidance for preventive

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