

James & Esther King Biomedical Research Program

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*Medicine/Cardiology
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*2010 Program
New Investigator Research
(3-year project)*

Project Title: Effects of Cigarette Smoking on Clopidogrel-Induced Antiplatelet Effects in Patients with Coronary Artery Disease

Project Summary: Cardiovascular disease affects over 80 million people in the U.S. and is the most important cause of mortality. Smoking is a strong risk factor for cardiovascular disease as it has a number of adverse effects, including increasing platelet activation, which in turn increases blood clots. The P2Y₁₂ receptor is a key platelet receptor that influences blood clots as shown by studies in high-risk patients with coronary artery disease (CAD). Thus, there is a clinical benefit associated with antiplatelet agents that block this receptor as it plays a pivotal role in patients with CAD. Importantly, smoking affects the response to inhibitors of P2Y₁₂ receptor. This project will use comprehensive and innovative functional assessments to better elucidate how smoking affects P2Y₁₂ response. The project will test the central hypothesis that cigarette smoking enhances chemical alterations of clopidogrel (medication to prevent blood clots) and that the inhibition of platelet P2Y₁₂ effects are greater in smokers compared to non-smokers. The studies are clinically significant since they advance our knowledge of how smoking influences P2Y₁₂, a key therapeutic target for the treatment of CAD patients. These investigations are part of our long-term goal of defining the best antiplatelet treatment strategy in high-risk patients with CAD.