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Published in:
Clinical Pharmacology & Therapeutics

DOI:
10.1002/cpt.1002

Publication date:
2018

Document Version
Publisher's PDF, also known as Version of record

Link to publication in Discovery Research Portal

Citation for published version (APA):
Response to “Influence of Diabetes on Antiplatelet Drug Efficacy”

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To The Editor: Gong et al. in their Letter to the Editor make valuable comments in relation to our study, which, as they correctly point out, included a substantial number of individuals with diabetes. First, they draw attention to the well-documented increased platelet reactivity in patients with diabetes, and a recent study showing increased platelet P2Y12 expression may be contributory. They also highlight a recent fairly small study indicating that in stroke patients the CYP2C19 loss-of-function allele had a reduced impact on clopidogrel response in individuals with higher glycated albumin levels independent of diabetes status. Such observations imply reduced efficacy of clopidogrel in this patient population. Indeed, a further study indicated impaired P2Y12 inhibition by clopidogrel following hospitalization for an arterial thrombo-occlusive event. As a consequence, we are not able to directly compare the overall efficacy of clopidogrel therapy with a comparator arm to investigate the impact of diabetes and related features. However, it is possible to exploit CYP2C19 genotype as a genetic instrument to probe the extent to which these features modify antiplatelet response. Our overall study comprised 72% individuals with diabetes and 77% of the outcome events occurred in this group. While we found that diabetes, obesity, smoking, and recent elevated glycated hemoglobin were, as expected, significantly associated with increased risk, we did not find any evidence of an interaction with the CYP2C19 genotype. While we probably do not have adequate power in our current study to formally avoid a type II error, a meta-analysis of clinical trials also found no substantial effect of either diabetes or obesity on clopidogrel efficacy. Our bioresource is currently expanding with the Scottish-wide GoSHARE study (www.goshare.org.uk). As a consequence, we anticipate the ability to explore these interesting and highly clinical relevant aspects in the future.

CONFLICT OF INTEREST

The authors declared no conflict of interest.

FUNDING

The GoDARTS Bioresource has been funded by a wide range of sources of many years. The present study was unfunded.

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Received 15 December 2017; accepted 17 December 2017; Advance online publication 30 December 1899. doi:10.1002/cpt.1002