

■ The Authors Respond

The letter by Dr. Ole Hauch regarding our article “Contemporary Trends in Oral Antiplatelet Agent Use in Patients Treated with Percutaneous Coronary Intervention for Acute Coronary Syndrome” provides thoughtful insight into the contemporary use of oral antiplatelet agents in patients who received percutaneous coronary intervention (PCI) for acute coronary syndrome (ACS). As supported by other studies conducted in the United States and other countries, an increase in the use of ticagrelor among ACS-PCI patients has been universally observed, although the rates of replacement across countries has varied. In interpreting these utilization differences, characteristics of the underlying datasets, as well as the health care systems examined in each study, have to be carefully considered. For example, centrally regulated insurance systems such as those in Greece and Sweden use direct price controls, international price comparisons, and value-based administration, which may encourage the use of efficacious but costly medications.^{1,2} Conversely, lump sum medication cost reimbursement would be considered as a potential control for prescription drug use in a commercially insured U.S. population, likely furthering acceptance of ticagrelor in comparison with Europe.

Our trend analysis comes with limitations in terms of characterizing factors in the selection of oral antiplatelet agents. Nonetheless, the results illustrate that recent use of antiplatelet medication generally corresponds to guideline and package insert recommendations. For example, prasugrel appears to be relatively discouraged in older patients and in patients with a history of stroke, transient ischemic stroke, or intracranial hemorrhage. Further, the 3 studies cited in Dr. Hauch’s letter found that the prevalence of ST-segment elevation was higher in patients prescribed either prasugrel or ticagrelor rather than clopidogrel, which is generally consistent with our results. That said, we also observed contraindicated uses. Moreover, as the letter pointed out, prescriber preferences and factors other than clinical characteristics have to be considered. Because of the inherent limitations of retrospective analyses using an administrative claims database, the influence of prescriber preferences and plan design on the use of cardiovascular medication remain relevant areas for future research.

Dr. Hauch’s letter affirms that our results suggest that a real-world effectiveness study would be valuable in understanding the use of oral antiplatelet agents in ACS patients treated with PCI. With respect to real-world outcomes associated with the 3 oral antiplatelet agents, while there have been several retrospective cohort analyses, the outcomes associated with the 3 agents have not been comprehensively covered, and there is no conclusive recommendation.^{3,4} To help fill this void, our research team is following up with studies based on comparative

TABLE

Preliminary Data Analyses: Comparative Resource Utilization of 3 Oral P2Y12 Antagonists over a 6-Month Period After ACS-PCI Discharge

	Third-Generation Agents vs. Clopidogrel	Ticagrelor vs. Prasugrel
Risk ratio, all-cause admissions	0.89 (0.78; 0.92)	1.02 (0.83; 1.19)
Risk ratio, all-cause office visits	0.99 (0.87; 1.03)	1.03 (0.87; 1.22)

effectiveness and health care resource utilization. Preliminary analyses, similar to the results from the pivotal randomized trials, have concluded that the use of third-generation antiplatelet agents, relative to clopidogrel, was associated with a decrease in the hazard rate of all-cause hospital admissions. In addition, patients who had initiated clopidogrel were associated with more hospital admissions, compared with those patients who were prescribed the newer agents, suggesting that there may be a meaningful cost offset to the higher price of ticagrelor or prasugrel (see Table). Whether these differences in prescription use in a commercially insured U.S. population, as compared with other countries, indicate underutilization of the newer agents will be further assessed in future studies.

Kibum Kim, PhD

Pharmacotherapy Outcomes Research Center
Department of Pathology
University of Utah

Robert J. DiDomenico, PharmD, BCPS, FCCP

Center for Pharmacoepidemiology and
Pharmacoeconomic Research
Department of Pharmacy Practice, College of Pharmacy
University of Illinois at Chicago

Daniel R. Touchette, PharmD, MA, FCCP

Center for Pharmacoepidemiology and
Pharmacoeconomic Research
Department of Pharmacy Systems, Outcomes and Policy,
College of Pharmacy, University of Illinois at Chicago

Surrey M. Walton, PhD

Center for Pharmacoepidemiology and
Pharmacoeconomic Research
Department of Pharmacy Systems, Outcomes and Policy,
College of Pharmacy, University of Illinois at Chicago
walton@uic.edu

DISCLOSURES

None of the authors of this study are involved in financial or personal relationships with agencies, institutions, or organizations that inappropriately influenced the statistical analysis plan or interpretation of the study results. DiDomenico received an honorarium from Amgen for the preparation of a heart failure drug monograph for *Pharmacy Practice News* and was a co-investigator on funded research for the Patient-Centered Outcomes Research Institute. DiDomenico also serves as an advisory board member for a heart failure program at Otsuka America Pharmaceuticals and as an advisory board member at Novartis Pharmaceuticals. Touchette has received unrestricted grant funding from Cardinal Health and Sunovion Pharmaceuticals and has also served as a consultant to, and director of, the American College of Clinical Pharmacy Practice-Based Research Network on a study funded by Pfizer.

REFERENCES

1. Karampli E, Souliotis K, Polyzos N, Kyriopoulos J, Chatzaki E. Pharmaceutical innovation: impact on expenditure and outcomes and subsequent challenges for pharmaceutical policy, with a special reference to Greece. *Hippokratia*. 2014;18(2):100-06.
2. Lundkvist J. Pricing and reimbursement of drugs in Sweden. *Eur J Health Econ*. 2002;3(1):66-70.
3. Bae JP, Faries DE, Ernst FR, et al. Real-world observations with prasugrel compared to clopidogrel in acute coronary syndrome patients treated with percutaneous coronary intervention in the United States. *Curr Med Res Opin*. 2014;30(11):2207-16.
4. Molife C, Frech-Tamas F, DeKoven M, et al. Comparison of healthcare resource utilization and costs in patients hospitalized for acute coronary syndrome managed with percutaneous coronary intervention and receiving prasugrel or ticagrelor. *J Med Econ*. 2015;18(11):898-908.