Please note: Dr. Tatonetti is a compensated advisor to Advera Health, Inc. Mr. Lorberbaum has reported that he has no relationships relevant to the contents of this paper to disclose.

REFERENCES

1. Lorberbaum T, Sampson KJ, Chang JB, et al. Coupling data mining and laboratory experiments to discover drug interactions causing QT prolongation. J Am Coll Cardiol 2016;68:1756-64.

2. Lorberbaum T, Sampson KJ, Woosley RL, et al. An integrative data science pipeline to identify novel drug interactions that prolong the QT interval. Drug Saf 2016;39:433-41.

3. Observational Health Data Sciences and Informatics. Available at: https:// www.ohdsi.org/. Accessed March 3, 2017.

4. Ryan PB, Madigan D, Stang PE, et al. Medication-wide association studies. CPT Pharmacometrics Syst Pharmacol 2013;2:e76.

5. Hripcsak G, Ryan PB, Duke JD, et al. Characterizing treatment pathways at scale using the OHDSI network. Proc Natl Acad Sci USA 2016;113: 7329-36.

Vitamin E May Protect Against Contrast-Induced Acute Kidney Injury

McCullough et al. (1) reviewed the pathophysiology and treatment options for contrast-induced acute kidney injury (CI-AKI). They stated that no effective adjunctive pharmaceutical had been demonstrated that either prevented or treated CI-AKI. However, they also suggested that of the agents being investigated, statins were the most promising. We would like to point out that strong evidence has also emerged regarding the effect of vitamin E against CI-AKI, which was not mentioned in their review.

Three randomized placebo-controlled trials found that vitamin E significantly prevented CI-AKI, with point estimates ranging from 52% to 75% for the decrease in the incidence of CI-AKI (2-4). All participants had chronic kidney disease and had been subjected to coronary catheterization or angiography. The latest trial reported CI-AKI cases in 14.1% of the placebo group, but in only 6.7% of the vitamin E group, which corresponded to 7.4% of participants benefitting from the vitamin, with a number needed to treat (NNT) of 13.5 (4). The 2 earlier studies found NNTs of 5.8 (2) and 10.6 (3). In each study, approximately one-half of the patients were on statin therapy; therefore, the effects of vitamin E might have also been beneficial in addition to statins.

Two of the vitamin E trials were carried out in Thailand (2,3), and 1 was carried out in Iran (4). Thus,

it is not known whether the findings can be directly generalized to Western countries. Even if the positive findings might only be applicable to less developed countries, the findings are important for the populations of such countries.

Vitamin E is an essential nutrient, and therefore, its potential benefit in preventing CI-AKI is interesting. Furthermore, vitamins E and C may interact. Vitamin E decreased total mortality in male smokers aged older than 65 years if their dietary vitamin C intake level was high, but not if their vitamin C intake was low (5). Thus, a large factorial trial seems warranted to examine the effect of statins and vitamins E and C, and their combinations to discover the optimal protocol to prevent CI-AKI.

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REFERENCES

1. McCullough PA, Choi JP, Feghali GA, et al. Contrast-induced acute kidney injury. J Am Coll Cardiol 2016;68:1465-73.

2. Tasanarong A, Piyayotai D, Thitiarchakul S. Protection of radiocontrast induced nephropathy by vitamin E (alpha tocopherol): a randomized controlled pilot study. J Med Assoc Thai 2009;92:1273-81.

3. Tasanarong A, Vohakiat A, Hutayanon P, et al. New strategy of alpha- and gamma-tocopherol to prevent contrast-induced acute kidney injury in chronic kidney disease patients undergoing elective coronary procedures. Nephrol Dial Transplant 2013;28:337-44.

4. Rezaei Y, Khademvatani K, Rahimi B, et al. Short-term high-dose vitamin E to prevent contrast medium-induced acute kidney injury in patients with chronic kidney disease undergoing elective coronary angiography: a randomized placebo-controlled trial. J Am Heart Assoc 2016;5:e002919.

5. Hemilä H, Kaprio J. Modification of the effect of vitamin E supplementation on the mortality of male smokers by age and dietary vitamin C. Am J Epidemiol 2009;169:946-53.

REPLY: Vitamin E May Protect Against Contrast-Induced Acute Kidney Injury



We appreciate the comments from Drs. Hemilä and Rezaei concerning small randomized trials of shortterm vitamin E for the prevention of contrastinduced acute kidney injury (CI-AKI). In the trial