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# Retraction

Hoeldtke RD, Bryner KD, McNeill DR, Hobbs GR, Riggs JE, Warehime SS, Christie I, Ganser G, Van Dyke K: Nitrosative stress, uric acid, and peripheral nerve function in early type 1 diabetes. *Diabetes* 51:2817–2825, 2002

The editors received a request to publish a retraction of the above manuscript. What follows is the letter that provides the authors' explanation for this request.

FRANZ M. MATSCHINSKY  
EDITOR IN CHIEF

Dear Dr. Matschinsky,

We reported a few years ago that protein-bound nitrotyrosine was increased in early type 1 diabetes and showed a negative correlation with motor nerve conduction velocity (1). We measured nitrotyrosine and tyrosine by high-performance liquid chromatography (HPLC) followed by electrochemical detection and expressed our results as the ratio of nitrotyrosine to tyrosine, which we found to be  $0.34 \pm 0.12 \times 10^{-3}$  in the control subjects and  $0.778 \pm 0.04$  in diabetic patients. Unfortunately, at the time we published these data, we were unaware of a study by Frost et al. (2), who performed alkaline hydrolysis of the plasma protein followed by mass spectrometry and reported a much lower nitrotyrosine-to-tyrosine ratio ( $0.044 \times 10^{-3}$ ). A few years later, Shishehbor et al. (2) performed acid hydrolysis of serum protein and reported a ratio in nondiabetic patients of  $5.2 \times 10^{-6}$ , also much lower (100-fold) than we observed. Because of these discrepancies, we have reassessed our HPLC method and performed a more in-depth analysis of the electrical properties of the peaks we have previously identified as nitrotyrosine using a multichannel CoulArray liquid chromatograph (ESA, Chelmsford, MA). We observed that the electrical properties of the compound we "identified" as nitrotyrosine on the basis of HPLC retention time differed from the authentic compound in many samples and concluded that interfering substances were obscuring the true nitrotyrosine peaks. We have used a second HPLC column and eliminated the interferences from some of our samples, and this decreased the estimated ratio of nitrotyrosine to tyrosine by approximately a factor of 100. On

the basis of these considerations, we have concluded that our previously published nitrotyrosine data (1) are invalid. We therefore formally retract this publication. Increased nitrotyrosine has been documented in other laboratories in clinical diabetes using other methods (3,4).

## REFERENCES

1. Hoeldtke RD, Bryner KD, McNeill DR, Hobbs GR, Riggs JE, Warehime SS, Christie I, Ganser G, Van Dyke K: Nitrosative stress, uric acid, and peripheral nerve function in early type 1 diabetes. *Diabetes* 51:2817–2828, 2002
2. Frost MT, Halliwell B, Moore KP: Analysis of free and protein-bound nitrotyrosine in human plasma by a gas chromatography/mass spectrometry method that avoids nitration artifacts. *Biochem J* 345:453–458, 2000
3. Shishehbor MH, Aviles RJ, Brennan ML, Fu X, Goormastic M, Pearce GL, Gokce N, Keaney JF Jr, Penn MS, Sprecher DL, Vita JA, Hazen SL: Association of nitrotyrosine levels with cardiovascular disease and modulation by statin therapy. *JAMA* 289:1675–1680, 2003
4. Ceriello A, Mercuri F, Quagliaro L, Assaloni R, Motz E, Tonutti L, Taboga C: Detection of nitrotyrosine in the diabetic plasma: evidence of oxidative stress. *Diabetologia* 44:834–838, 2001

Sincerely,

ROBERT D. HOELDTKE  
KIMBERLY D. BRYNER  
DANIEL R. MCNEILL  
GERALD R. HOBBS  
JACK E. RIGGS  
SARAH S. WAREHIME  
IAN CHRISTIE  
GARY GANSER  
KNOX VAN DYKE  
LINDA CORUM