

Rebreathing-Induced Hypoxia Improves Insulin Sensitivity in Adults with Type 2 Diabetes

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ABSTRACT

Hypoxia stimulates glucose uptake through an insulin-independent pathway. **PURPOSE:** To examine the acute effect of rebreathing-induced hypoxia on plasma glucose and insulin levels in adults with type 2 diabetes. We hypothesized that rebreathing-induced hypoxia would attenuate the increase in glucose and insulin levels during an oral glucose tolerance test. **METHODS:** Nine individuals with type 2 diabetes (4 women, age: 53 ± 10 years, body mass index: 35 ± 7 kg/m², HbA1c: $7.1 \pm 0.5\%$) visited the laboratory on two occasions. On both visits, a 2-hour, 75 g oral glucose tolerance test was conducted while simultaneously performing a rebreathing-induced hypoxia protocol (RIH) or breathing room air (Norm). Venous blood samples were collected 0, 30, 60, 90, and 120 min following ingestion of the high-glucose drink to measure plasma glucose and insulin levels. The rebreathing-induced hypoxia protocol consisted of two series of five 2-min rebreathing bouts in a low-volume, closed circuit system interspersed with two minutes of breathing room air. The first and second series of rebreathing bouts were performed within the first 30 min and 30-60 min after ingestion of the high-glucose drink, respectively. **RESULTS:** Rebreathing-induced hypoxia resulted in a nadir oxygen saturation of $88 \pm 4\%$ and a nadir fraction of inspired oxygen of $12 \pm 5\%$. Plasma glucose responses to the oral glucose tolerance test were not different between conditions, however, insulin levels were lower during rebreathing-induced hypoxia than normoxia (RIH vs. Norm: 0: 17 ± 13 vs. 21 ± 14 ; 30: 37 ± 28 vs. 42 ± 26 ; 60: 57 ± 32 vs. 77 ± 52 ; 90: 88 ± 67 vs. 116 ± 81 ; and 120: 114 ± 96 vs. 136 ± 111 uIU/ml, $p=0.03$), suggesting an acutely improved insulin sensitivity. Accordingly, the insulin area under the curve was lower during rebreathing-induced hypoxia than normoxia (124 ± 84 vs. 157 ± 102 , $p=0.02$). **CONCLUSION:** Exposure to short and intermittent rebreathing-induced hypoxia following ingestion of a high-glucose drink acutely improves insulin sensitivity in adults with type 2 diabetes. Rebreathing-induced hypoxia could therefore represent a novel and simple strategy to improve glycemic control in individuals living with type 2 diabetes.