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Title: Cerebral cortex blood flow, oxygen delivery and oxygenation during normoxic and hypoxic exercise in healthy humans

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Body: Background: During maximal hypoxic exercise a reduction in cerebral oxygen delivery may constitute a signal to terminate exercise. Aim: To investigate whether the rate of increase in cerebral cortex oxygen delivery is limited in hypoxic compared to normoxic exercise. Methods: We assessed frontal cerebral cortex blood flow using near infrared spectroscopy and the light-absorbing tracer indocyanine green dye, as well as frontal cortex oxygen saturation (StO₂%) in 11 cyclists during incremental exercise to the limit of tolerance (WR_{max}) in normoxia and acute hypoxia (F_IO₂:0.12). Results: In normoxia, cerebral cortex blood flow and oxygen delivery increased (p<0.05) from baseline to sub-maximal exercise reaching peak values at near-maximal exercise (80%WR_{max}: 287±9W; 81±23% and 75±22% increase relative to baseline, respectively), both leveling off thereafter up to WR_{max} (382±10W). Cerebral cortex StO₂% did not change from baseline (66±3%) throughout graded exercise. During hypoxic exercise, cerebral cortex blood flow increased from baseline to sub-maximal exercise peaking at 80%WR_{max} (213±6W; 60±15% relative increase) before declining towards baseline at WR_{max} (289±5W). Despite this, cerebral cortex oxygen delivery remained unchanged from baseline throughout graded exercise, being at WR_{max} lower than at comparable loads (287±9W) in normoxia (by 58±12%). Cerebral cortex StO₂% fell from baseline (58±2%) on moderate exercise in parallel with arterial oxygen saturation, but then remained unchanged to exhaustion (47±1%). Conclusion: Cerebral cortex oxygen delivery is limited in hypoxia compared to normoxia, thus potentially compromising maximal exercise capacity in hypoxia.