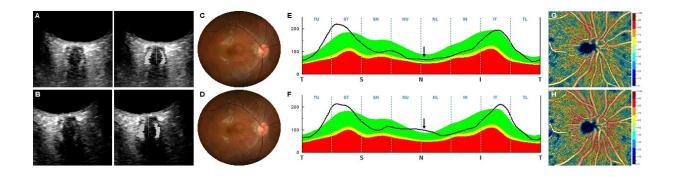


Retinal microvasculature is a potential biomarker for acute mountain sickness

April 7 2023



Data from the top and bottom rows are obtained before and after exposure to a simulated high altitude, respectively. (A/B) Ultrasonography image showing the optic nerve subarachnoid space area expansion (white shaded area) after exposure to a simulated high altitude (a) 6.855 vs. (b) 7.545 mm². (C/D) Fundus photograph showing the lack of disc edema during exposure to a simulated high altitude. (E/F) However, RNFL thickness increased after exposure (blue arrows). (G/H) Increased radial peripapillary capillary flow density is seen by optic coherence tomography angiography after exposure. Credit: Science China Press

A study led by Dr.Ningli Wang and Dr.Yuan Xie (Beijing Tongren Hospital, Capital Medical University) provides evidence that retinal microvasculature is a potential biomarker for cerebral microvasculature changes and acute mountain sickness(AMS) development during risk assessment of individuals at high altitudes.



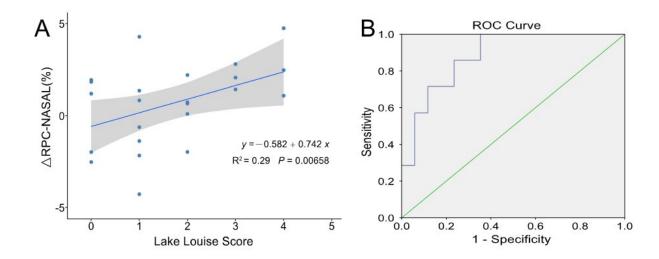
Since the retinal and cerebral vasculature share many morphological and physiological characteristics, direct evaluation of the more accessible retinal vasculature should theoretically provide insights into the cerebral circulation. Therefore, the role of retinal imaging to predict central nervous system diseases has received much attention from scholars.

China has the most extensive plateau in the world, millions of people now travel to <u>high altitudes</u> each year to ski, trek, climb, and work. Increased <u>cerebral blood flow</u> resulting from altered capillary level autoregulation at high altitudes leads to capillary overperfusion and then vasogenic cerebral edema, which is the leading hypothesis of <u>acute mountain sickness</u> (AMS).

The best intervention point is before the occurrence of irreversible brain lesions. However, due to technical limitations, cerebral vascular detection is currently limited to large vessels, and capillaries are hard to visualize in vivo.

This study aimed to investigate ocular microcirculation alterations, the only visualized capillaries in the central neural system (CNS), during early-stage AMS using a hypobaric chamber. They applied an well-documented AMS diagnostic criteria- Lake Louise Questionnaire Score (LL-S).





A indicated scatter plot showing the relationship between the changes in the radial peripapillary capillary flow density in the nasal sector (Δ RPC-NASAL) to the Lake Louise Questionnaire Score after exposure to a simulated high altitude using a hypobaric hypoxic chamber. B indicated an receiver operating characteristics curve showing sensitivities (true-positive rate) and specificities (false-positive rate) of the change in the radial peripapillary capillary flow density in the nasal sector (Δ RPC-NASAL) compared between all participants with biomarker-positive and biomarker-negative findings. Credit: Science China Press

Meanwhile, they comprehensively analyzed anatomical changes in the eye in early-stage AMS (3 ≤Lake Louise Questionnaire Score

Because retinal nerve fiber layers (RNFL) and white matter are both composed of axons, they applied optical coherence tomography (OCT) to assess RNFL variation. Furthermore, the area of the optic nerve subarachnoid space (ONSASA) was found to be significantly correlated with intracranial pressure in the previous study, thus they used ultrasound to measure ONSASA to indirectly assess cerebrospinal fluid pressure changes.



The results suggested that after high altitude simulation, the RNFL thickened in some locations and ONSASA enlarged. OCTA showed increased retinal radial peripapillary capillary (RPC) flow density, particularly on the nasal side of the nerve. The AMS-positive group had the largest increases in RPC flow density in the nasal sector.

Among multiple ocular changes, OCTA increase in RPC flow density was associated with simulated early-stage AMS symptoms. The changes in RPC flow density to predict early-stage AMS outcomes showed an powerful diagnostic power.

The findings further confirm that increased capillary perfusion pressure is an important mechanism for early AMS pathogenesis, and that OCTA measurement of retinal capillary blood flow density, especially peripapillary blood flow density, may be a potential biological indicator for predicting the development of AMS. This result lays the foundation for the role of fundus imaging in the diagnosis and management of cerebrovascular diseases.

The study is published in the journal Science China Life Sciences.

More information: Yuan Xie et al, Retinal microvasculature is a potential biomarker for acute mountain sickness, *Science China Life Sciences* (2023). DOI: 10.1007/s11427-022-2271-x

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