

Targeting the brain's energy metabolism may hold the key for treating Parkinson's disease

March 2 2022

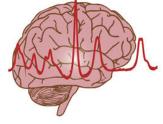
NADPARK

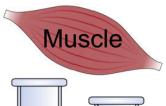
Parkinson's disease

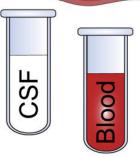












- ♠ Brain NAD levels
- Altered cerebral metabolism
- ↑ NAD-metabolome
- ↑ Transcription of mitochondrial genes
- ↑ Transcription of proteostasis genes
- ↓ Inflammatory cytokines



Graphical abstract. Credit: DOI: 10.1016/j.cmet.2022.02.001

Nicotinamide riboside (NR) is a member of the vitamin B3 family, which can be found in foods like fruits, vegetables, meat, and milk. It is also used as a nutritional supplement to boost cellular levels of NAD (nicotinamide adenine dinucleotide).

NAD is an essential metabolic factor, which is vital to a multitude of cellular processes, including <u>energy metabolism</u>, DNA damage repair, and control of gene expression. Increased cellular NAD levels have been linked to a longer and healthier life in animals and have been shown to protect <u>neuronal cells</u> against various forms of stress. The study shows for the first time that the NAD levels of the human brain can be increased by ingesting NR.

A potential new treatment

The goal of this randomized, double-blinded <u>phase</u> I trial was to establish whether NAD metabolism can be augmented in the brain of individuals with Parkinson's disease, by ingesting NR, an NAD-precursor. A total of 30 individuals with early Parkinson's disease received either 1,000 mg NR or placebo for a total of 30 days. The study showed that NR supplementation significantly increased NAD levels in the patient brain, and resulted in altered cerebral metabolism and decreased markers of inflammation in the fluid surrounding the brain and spinal cord. Moreover, a mild but significant improvement of Parkinson's disease symptoms was seen in participants who showed substantially increased brain NAD levels, and this correlated with the change in the brain's energy metabolism pattern.

The study was led by Prof. Charalampos Tzoulis, at Haukeland



University Hospital and the University of Bergen.

"We are very excited about these results. This trial represents a novel approach in experimental PD-therapy," Tzoulis says.

"We believe that augmenting the brain's NAD metabolism will not only target and rectify disease-related processes specific to PD, but may also optimize neuronal metabolism and fortify neurons, rendering them more resilient against age-related stress and neurodegenerative diseases. The results of the trial are highly encouraging and nominate NR as a potential neuroprotective therapy for PD, warranting further investigation in larger trials. A phase II study is already ongoing at our Center," the professor says.

More information: Brage Brakedal et al, The NADPARK study: A randomized phase I trial of nicotinamide riboside supplementation in Parkinson's disease, *Cell Metabolism* (2022). <u>DOI:</u> 10.1016/j.cmet.2022.02.001. www.cell.com/cell-metabolism/f ... 1550-4131(22)00045-6

Provided by University of Bergen

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