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## **ACCEPTED MANUSCRIPT**

Influence of metformin on mitochondrial subproteome in the brain of apoE

knockout mice.

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#### **Abstract**

Neurodegenerative diseases are the set of progressive, age-related brain disorders, characterized by an excessive accumulation of mutant proteins in the certain regions of the brain. Such changes, collectively identified as causal factors of neurodegeneration, all impact mitochondria, imminently leading to their dysfunction. These observations predestine mitochondria as an attractive drug target for counteracting degenerative brain damage.

The aim of this study was to use a differential proteomic approach to comprehensively assess the changes in mitochondrial protein expression in the brain of apoE-knockout mice (apoE<sup>-/-</sup>) and to investigate the influence of prolonged treatment with metformin – an indirect activator of AMP-

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