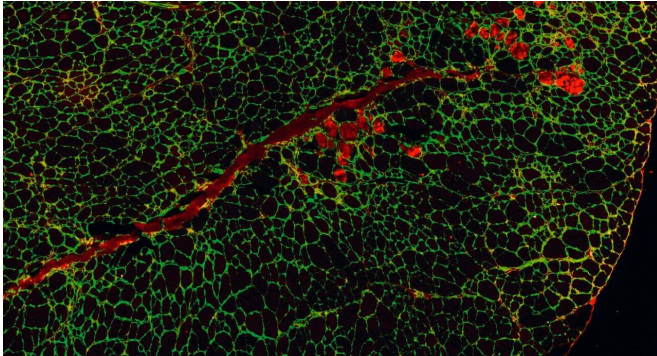


Urolithin A appears effective against muscular dystrophy

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Regenerating muscle cells thanks to Urolithin A. Credit: © Amazentis / EPFL Auwerx Lab

Progression of Duchenne Muscle Dystrophy (DMD) can be delayed in mice by supplementing their diets with Urolithin A, according to new results reported today. The findings, published in *Science Translational Medicine*, raise hopes that new treatment options could one-day be developed for DMD, an incurable genetic condition characterized by progressive muscle degeneration. About 1 in 3,500 boys are born with DMD, which usually develops in childhood and significantly reduces life expectancy.

The new research carried out at the laboratory of Professor Johan Auwerx, MD, Ph.D. at the Swiss Federal Institute of Technology EPFL and the University of Lausanne in collaboration with scientists at the Swiss life science company Amazentis, highlights the important role that defective mitochondria can play in DMD. The powerhouses of cells, mitochondria produce the energy necessary for normal [muscle](#) function. But [muscle cells](#) taken from both human DMD patients and from mice bred to mimic the condition show significant defects in mitochondrial activity, the study finds. Specifically, patterns of gene expression show the development of DMD is

associated with a marked decrease in mitophagy—the process cells rely on to remove and recycle defective mitochondria and maintain energy levels high.

"Duchenne Muscle Dystrophy is the most common fatal genetic disease diagnosed in childhood with still no cure available," says Johan Auwerx, MD, Ph.D., lead-author and Professor at the EPFL. "Our work represents a significant breakthrough in the search for new therapeutic approaches for muscular dystrophies."

The natural compound Urolithin A is known to activate mitophagy and improve mitochondrial health in both mice and humans. When the study scientists and lead authors, Peiling Luan and Davide D'Amico, fed the compound to DMD mice for just ten weeks, they saw mitophagy levels rise effectively restoring them to normal. This led to a significant reduction of muscle damage and improvement in muscle health and performance. The DMD mice administered Urolithin A saw grip strength increase by 31% and running performance increase by 45% compared with control untreated animals. And they lived longer—survival increased by 40%.

Importantly for the human disease, Urolithin A reduced a damaging condition called fibrosis in muscles of the DMD mouse heart and diaphragm by 36% and 39%, respectively. Similar damage seen in DMD patients typically leads to fatal cardiac or respiratory failure. Urolithin A was also able to enhance the regeneration of mouse muscle stem cells. This is particularly relevant to the disease in humans as the onset of DMD is linked with the exhaustion of functional stem cells.

Davide D'Amico, Ph.D., Project Leader at Amazentis and a first author of the paper, said: "Prior to this study, it was understood that the dramatic loss of muscle function in DMD patients was associated with mitochondrial dysfunctions.

Here we discovered that defective mitophagy, the removal and recycling of dysfunctional mitochondria, plays a key role in the progression of DMD."

Chris Rinsch, Ph.D., Co-founder and CEO of Amazentis, said, "The rigorous science being published in *Science Translational Medicine* strengthens the scientific evidence of Urolithin A as a potent enhancer of muscle function. It's exciting to see this natural metabolite can support not only healthy muscle, but also shows promise for progressive muscle diseases in pre-clinical research."

More information: "Urolithin A improves muscle function by inducing mitophagy in muscular dystrophy" *Science Translational Medicine* (2021). [stm.sciencemag.org/lookup/doi/...
scitranslmed.abb0319](https://stm.sciencemag.org/lookup/doi/10.1126/scitranslmed.abb0319)

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