

Cholesterol-lowering drug improves function of heart's arteries

29 July 2020



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In a study published July 17 in the *Journal of the American Heart Association*, the Johns Hopkins researchers suggest there may be a way to limit cardiovascular disease risk in those living with HIV and other cardiovascular risk factors, such as high cholesterol, by improving the function of <u>blood</u> vessels.

"We hypothesized that PCSK9 mediates an inflammatory response that impairs vascular function in addition to its effects on <u>cholesterol</u> metabolism, and we tested this idea with the protein's inhibitor to learn whether it could help people who have impaired blood vessel function," says lead author of the study, Thorsten M. Leucker, M.D., Ph.D., assistant professor of medicine at the Johns Hopkins University School of Medicine. "We were surprised that this worked so well, but also heartened that there may be a way we can improve blood vessel function in those with increased inflammation."

For the study, 19 people with HIV and 11 people with high blood lipids but no HIV were given the drug evolocumab, a PCSK9 inhibitor, for six weeks. At the start of the study and following the course of treatment, the team used cardiac <u>magnetic</u> <u>resonance</u> imaging (MRI) to measure the area of, and blood flow in, the <u>right coronary artery</u> at rest and during a hand exercise, which normally results in relaxation of the blood vessels. When healthy people perform the test, the coronary artery



responds to the exercise and its area increases, enabling more blood to flow through. In people with impaired blood vessel function, such as those with HIV or <u>high cholesterol</u>, the artery doesn't get larger or may even constrict. Therefore, blood flow remains the same or decreases in those with impaired blood vessel function.

After the six-week treatment with the PCSK9 inhibitor, the participants living with HIV had an average 7.9% increase in coronary artery area and a 10.1% increase in blood flow during the hand grip exercise when compared to the resting value. These changes were significantly greater than the changes from rest to hand grip exercise during the baseline pre-treatment visit. Participants with high blood lipids also had improvements in coronary artery area and an increase in <u>blood flow</u> after six weeks of treatment.

The researchers say future studies will need to include more patients and study them over a longer time.

More information: Thorsten M. Leucker et al. Evolocumab, a PCSK9?Monoclonal Antibody, Rapidly Reverses Coronary Artery Endothelial Dysfunction in People Living With HIV and People With Dyslipidemia, *Journal of the American Heart Association* (2020). DOI: 10.1161/JAHA.120.016263

Provided by Johns Hopkins University School of Medicine APA citation: Cholesterol-lowering drug improves function of heart's arteries (2020, July 29) retrieved 11 October 2022 from <u>https://medicalxpress.com/news/2020-07-cholesterol-lowering-drug-function-heart-arteries.html</u>

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