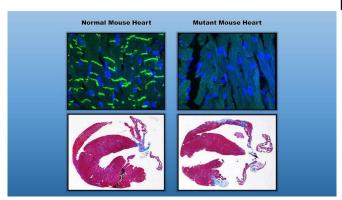


## Mental wellness may be important prevention factor for sudden death in genetic heart disease

18 February 2021



Proteins (green) normally found in the divisions between healthy heart cells (upper left) are found diffusely distributed throughout mouse heart cells that have a genetic mutation for arrhythmogenic cardiomyopathy (upper right). The mutant mouse hearts (bottom right) also have more scar tissue (blue) than healthy mouse hearts (bottom left). Credit: Stephen Chelko, Ph.D., Johns Hopkins Medicine

Studying mice modeling a human genetic disease that causes irregular heartbeats and can lead to sudden death, Johns Hopkins Medicine researchers have shown that emotional stress seems to promote symptoms of the disease and worsens its progression. The researchers say in their study, published Nov. 24, 2020, in the *Journal of Clinical Medicine*, that physicians and patients would be wise to focus together on good mental health—in addition to traditional exercise avoidance—to keep patients healthy and avoid triggering symptoms.

"There's much evidence that our body and mind are connected so that our psychological states can change the course of <u>disease</u>," says Jacopo Agrimi, Ph.D., a research fellow in the laboratory of Nazareno Paolocci, M.D., Ph.D., associate

professor of medicine at the Johns Hopkins
University School of Medicine. "Prolonged stress
may be like putting fuel on the fire in someone with
a chronic disease. So, we need to ensure that we
are taking patients' quality of life and psychological
wellness into account when managing their
disease."

The Johns Hopkins Medicine research study was conceived when the researchers heard anecdotal reports from patients at a seminar held as part of the Johns Hopkins ARVC/D Program.

Arrhythmogenic right ventricular dysplasia (ARVD), also known as arrhythmogenic right ventricular cardiomyopathy (ARVC), is a genetic heart condition and a leading cause of sudden death among young athletes, but it can affect people of all ages and all activity levels. ARVC/D was one of the diseases modeled by the mice in the study.

"Patients often reported that when they heard their defibrillator charging—which is needed to shock their heart beat back into normal rhythm when disrupted—they would panic due to past experiences resulting from the pain and discomfort of a defibrillator discharge," says Stephen Chelko, Ph.D., an adjunct assistant professor of medicine at the Johns Hopkins University School of Medicine, as well as an assistant professor of biomedical sciences at Florida State University. "As a result, many people also reported waking up after the shock, hearing the defibrillator recharge and become panicked again, thus creating a viscous cycle."

These patient stories led Chelko and Agrimi to survey patients with these genetic heart diseases about their stress levels during annual clinical follow-ups. They found that those with higher levels of perceived psychosocial stress showed a strong



correlation with clinical symptoms as measured by cardiac magnetic resonance imaging or electrocardiogram.

With this knowledge, the researchers then took mice with the second most common genetic mutation for arrhythmogenic cardiomyopathy and put half of the group in stressful situations by exposing them to large, aggressive and bullying mice for weeks.

Almost a third of the mice with the genetic heart condition and placed in the stressful situation died of <u>sudden death</u> compared with the mice with the heart problem in non-<u>stressful situations</u>. The researchers also observed that the stressed <u>mice</u> who remained alive showed more cardiac damage and more rapid disease progression.

"Genetic heart disease is stressful enough without adding psychological stress, with athletes having to give up exercise and participation in sports when they are diagnosed," says Chelko. "We need physicians to help patients navigate the disease and treat the whole condition, including the mental wellness aspect."

**More information:** Jacopo Agrimi et al. Psychosocial Stress Hastens Disease Progression and Sudden Death in Mice with Arrhythmogenic Cardiomyopathy, *Journal of Clinical Medicine* (2020). DOI: 10.3390/jcm9123804

Provided by Johns Hopkins University

APA citation: Mental wellness may be important prevention factor for sudden death in genetic heart disease (2021, February 18) retrieved 16 August 2022 from <a href="https://medicalxpress.com/news/2021-02-mental-wellness-important-factor-sudden.html">https://medicalxpress.com/news/2021-02-mental-wellness-important-factor-sudden.html</a>

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