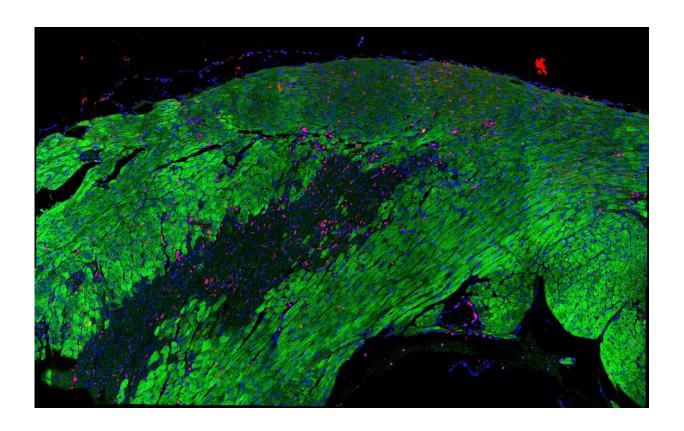


Researchers map heart recovery after heart attack with great detail

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Shown is an infarcted heart in which green represents cardiomyocytes (heart muscle cells), red shows B2M expression and blue represents cell nuclei. The area deprived of green is the infarcted area. Credit: Louk Timmer © Hubrecht Institute

Researchers from the Hubrecht Institute have mapped the recovery of the heart after a heart attack with great detail. They found that heart

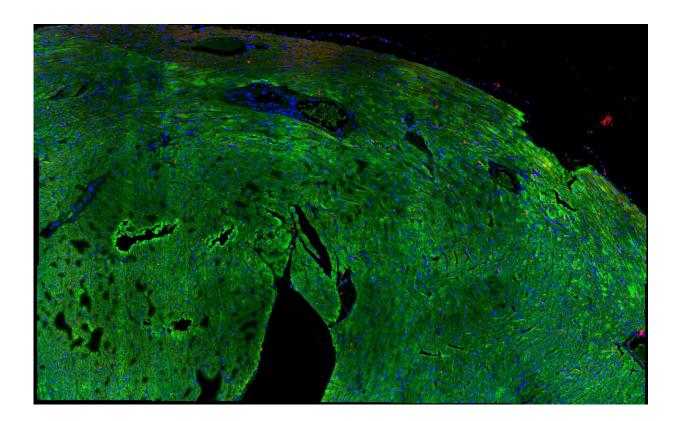


muscle cells called cardiomyocytes play an important role in intracellular communication after a heart attack. The researchers documented their findings in a database that is accessible for scientists around the world. This brings the research field a step closer to the development of therapies for improved recovery after heart injury. The results were published in *Communications Biology* on the 29th of January.

In the Netherlands, an average of 95 people end up in the hospital each day because of a heart attack. During a heart attack, the blood supply to a part of the heart is blocked, for example, due to a blood clot in a coronary artery. Attempts to restore the blood supply known as reperfusion, are made as soon as possible. However, a part of the heart has already been without oxygen for some time. Depending on the size and duration of the infarction, this causes cardiomyocytes to die. This can result in the formation of scar tissue, which is stiffer than normal heart tissue and therefore makes it more difficult for the heart to properly contract. This can cause the pumping function of the heart to deteriorate, which can eventually lead to heart failure.

In other words, insight into the <u>recovery</u> of the heart after a heart attack and how this leads to the formation of scar tissue is extremely important. However, much is still unknown. Researchers from the lab of Eva van Rooij have therefore examined this further. They studied how the hearts of mice recover at three different time points following a heart attack. To this end, they used single-cell sequencing, a technique that enables the examination of the RNA of individual <u>cells</u>. The researchers generated an enormous dataset with information about the role of different types of cells during the recovery process after a heart attack.





Shown is a control heart in which green represents cardiomyocytes (heart muscle cells), red shows B2M expression and blue represents cell nuclei. Credit: Louk Timmer © Hubrecht Institute

Communication network

Consequently, they used the data to map a <u>communication network</u>. Louk Timmer, researcher on the project, says, "Cells communicate with each other by secreting molecules. These molecules then trigger the recipient cell to take a specific action, which may be important for the recovery process. We have now mapped with great detail how different cells communicate with each other at different time points after a heart attack. That had never been done so thoroughly before." This communication network is now documented in a database and accessible to scientists around the world.



Formation of scar tissue

The role of cardiomyocytes in the recovery following a heart attack has been largely unknown, partly because of technical difficulties. However, another recent paper from Van Rooij's lab overcame these obstacles, allowing the researchers to specifically study the function of cardiomyocytes in the recovery process. "We noticed that at the earliest time point measured after the heart attack, cardiomyocytes were secreting increased amounts of a molecule called B2M. Subsequent experiments showed that the secretion of B2M can result in the activation of so-called fibroblasts—cells responsible for the formation of scar tissue," says Timmer. Cardiomyocytes thus seem to indirectly stimulate the production of scar tissue early in the recovery process. "Intuitively, we already thought that cardiomyocytes play an important role in intracellular communication during heart recovery, and it is great that we have now been able to confirm that."

When asked about the next research steps, Timmer emphasized the importance of additional studies. "Various scientists and experts can use this data, which enables us to gain a better understanding of the cells and molecules that are involved in the recovery of the heart and the way they communicate with each other. Hopefully, we can eventually improve the recovery process so that people end up with less damage after a heart attack."

More information: "Single-cell transcriptomics following ischemic injury identifies a role for B2M in cardiac repair". Bas Molenaar, Louk T. Timmer, Marjolein Droog, Ilaria Perini, Danielle Versteeg, Lieneke Kooijman, Jantine Monshouwer-Kloots, Hesther de Ruiter, Monika M. Gladka & Eva van Rooij. *Communications Biology*, 2020. DOI: 10.1038/s42003-020-01636-3



Provided by Hubrecht Institute

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