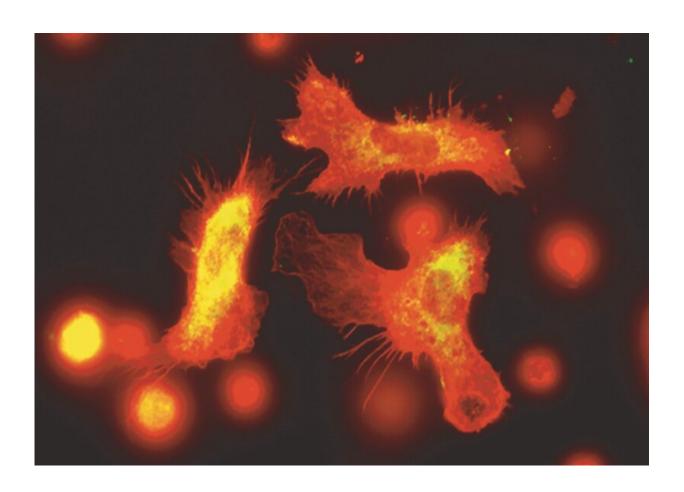


## Blood stem cells boost immunity by keeping a record of previous infections

March 13 2020



Immune cells by fluorescence microscopy: Blood stem cells remember a previous attack and produce more immune cells like these macrophages to fight a new infection. Credit: Sieweke lab / CIML

A Franco-German research team led by Prof. Michael Sieweke, from



the Center for Regenerative Therapies TU Dresden (CRTD) and the Center of Immunology of Marseille Luminy (CNRS, INSERM, Aix-Marseille University), today uncovered a surprising property of blood stem cells: not only do they ensure the continuous renewal of blood cells and contribute to the immune response triggered by an infection, but they can also remember previous infectious encounters to drive a more rapid and more efficient immune response in the future. These findings should have a significant impact on future vaccination strategies and pave the way for new treatments of an underperforming or overreacting immune system. The results of this research are published in *Cell Stem Cell* on March 12, 2020.

Stem <u>cells</u> in our bodies act as reservoirs of cells that divide to produce new stem cells, as well as a myriad of different types of specialized cells, required to secure tissue renewal and function. Commonly called <u>blood</u> <u>stem cells</u>, the <u>hematopoietic stem cells</u> (HSC) are nestled in the bone marrow, the soft tissue that is in the center of large bones such as the hips or thighs. Their role is to renew the repertoire of blood cells, including cells of the <u>immune system</u> which are crucial to fight infections and other diseases.

Until a decade ago, the dogma was that HSCs were unspecialized cells, blind to external signals such as infections. Only their specialized daughter cells would sense these signals and activate an immune response. But work from Prof. Michael Sieweke's laboratory and others over the past years has proven this dogma wrong and shown that HSCs can actually sense external factors to specifically produce subtypes of immune cells "on demand" to fight an infection. Beyond their role in an emergency immune response, the question remained as to the function of HSCs in responding to repeated infectious episodes. The immune system is known to have a memory that allows it to better respond to returning infectious agents. The present study now establishes a central role for blood stem cells in this memory.



"We discovered that HSCs could drive a more rapid and efficient immune response if they had previously been exposed to LPS, a bacterial molecule that mimics infection," said Dr. Sandrine Sarrazin, Inserm researcher and senior-author of the publication. Prof. Michael Sieweke, Humboldt Professor at TU Dresden, CNRS Research Director and last author of the publication, explained how they found the memory was stored within the cells: "The first exposure to LPS causes marks to be deposited on the DNA of the stem cells, right around genes that are important for an immune response. Much like bookmarks, the marks on the DNA ensure that these genes are easily found, accessible and activated for a rapid response if a second infection by a similar agent was to come."

The authors further explored how the memory was inscribed on the DNA, and found C/EBPb to be the major actor, describing a new function for this factor, which is also important for emergency immune responses. Together, these findings should lead to improvements in tuning the immune system or better vaccination strategies.

"The ability of the immune system to keep track of previous infections and respond more efficiently the second time they are encountered is the founding principle of vaccines. Now that we understand how blood stem cells book mark immune response circuits, we should be able to optimize immunization strategies to broaden the protection to infectious agents. It could also more generally lead to new ways to boost the <a href="immune response">immune response</a> when it underperforms or turn it off when it overreacts," concluded Prof. Michael Sieweke.

The research group of Prof. Michael Sieweke works at the interface of immunology and stem cell research. The scientists focus on the study of hematopoietic stem cells and macrophages, long-lived mature cells of the immune system that fulfil an important role in tissue regeneration. In 2018, Prof. Michael Sieweke received the most valuable research award



in Germany: the Alexander von Humboldt Professorship, which brings top international researchers to German universities. In addition to his position as Research Director at the Centre for Immunology at the University of Marseille Luminy, he now acts as Deputy Director at the Center for Regenerative Therapies at TU Dresden (CRTD). CRTD is academic home for scientists from more than 30 nations. Their mission is to discover the principles of cell and tissue regeneration and leveraging this for recognition, treatment and reversal of diseases. The CRTD links the bench to the clinic, scientists to clinicians to pool expertise in stem cells, developmental biology, gene-editing and regeneration towards innovative therapies for neurodegenerative diseases such as Alzheimer's and Parkinson's disease, hematological diseases such as leukaemia, metabolic diseases such as diabetes, retina and bone diseases.

**More information:** Bérengère de Laval et al. C/EBPβ-Dependent Epigenetic Memory Induces Trained Immunity in Hematopoietic Stem Cells, *Cell Stem Cell* (2020). DOI: 10.1016/j.stem.2020.01.017

## Provided by Dresden University of Technology

Citation: Blood stem cells boost immunity by keeping a record of previous infections (2020, March 13) retrieved 9 April 2024 from <a href="https://medicalxpress.com/news/2020-03-blood-stem-cells-boost-immunity.html">https://medicalxpress.com/news/2020-03-blood-stem-cells-boost-immunity.html</a>

This document is subject to copyright. Apart from any fair dealing for the purpose of private study or research, no part may be reproduced without the written permission. The content is provided for information purposes only.