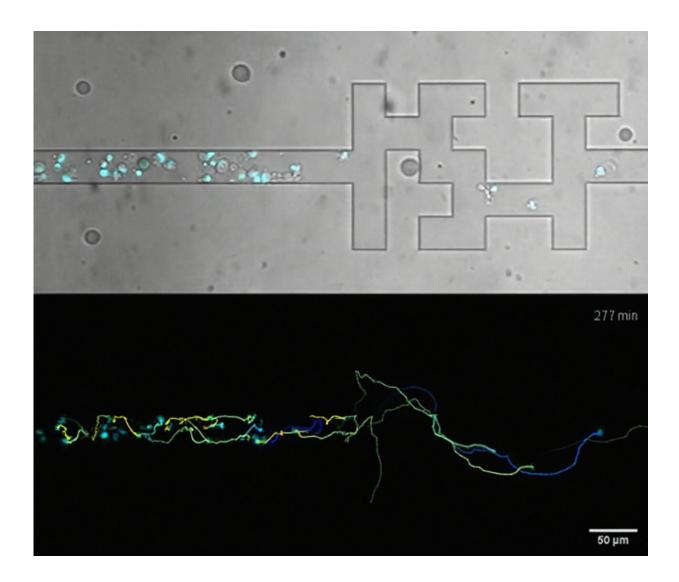


## Immune cells move more independently than previously thought

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Dendritic cells navigating in a microscopic labyrinth with the help of a chemokine gradient they have created. The nuclei of the cells are pictured in blue in the upper image, and the lines in the bottom image represent cell



movement. Credit: Jonna Alanko, University of Turku

Human immune cells are capable of coordinating their own movement more independently than previously thought. InFLAMES researcher Jonna Alanko has discovered that immune cells do not just passively follow the chemical cues in their environment. Quite the contrary, they can also shape these cues and navigate in complex environments in a selforganized manner.

Directional cell movement is an essential and fundamental phenomenon of life. It is an important prerequisite for individual development, reformation of blood vessels, and immune response, among others.

A study conducted by Postdoctoral Researcher Jonna Alanko focused on the movement and navigation of <u>immune cells</u> within the body. Chemokines, a class of signaling proteins, play a crucial role in guiding immune cells to specific locations.

Chemokines are formed, for instance, in the lymph nodes and create <u>chemical cues</u> called chemokine gradients for cells to follow within the body. According to Alanko, these chemokine gradients are like a trail of scent left in the air, it gets lighter the further you are from its source. The research article was published in the *Science Immunology* journal.

The traditional idea has been that immune cells recognize their target by following existing chemokine gradients. In other words, the cells following these cues have been seen as passive actors, which is not the case in reality.

"We were able to prove for the first time that contrary to the previous conception, immune cells do not need an existing chemokine <u>gradient</u> to



find their way. They can create gradients themselves and thereby migrate collectively and efficiently even in complex environments," explains Alanko.

## **Cells consume chemokines**

Immune cells have receptors with which they can sense a chemokine signal. One of these receptors is called CCR7 and can be found in <u>dendritic cells</u>.

Dendritic cells are professional antigen-presenting cells with an important role in activating the entire immune response. They need to locate an infection, recognize it, and then migrate to the lymph nodes with the information. In the <u>lymph nodes</u>, the dendritic cells interact with other cells of the immune system to initiate an <u>immune response</u> against pathogens.

The study conducted by Alanko revealed that dendritic cells do not only register a chemokine signal with their CCR7 receptor, but they also actively shape their chemical environment by consuming chemokines. By doing this, the cells create local gradients that guide their own movement and that of other immune cells. The researchers also discovered that T-cells, another type of an immune cell, can benefit from these self-generated gradients to enhance their own directional movement.

"When immune cells are capable of creating <u>chemokine</u> gradients, they can avoid upcoming obstacles in complex environments and guide their own directional movement and that of other immune cells," explains Jonna Alanko.

This discovery increases our understanding of how immune responses are coordinated within the body. However, it can also reveal how <u>cancer</u>



<u>cells</u> guide their movement to create metastases.

"The CCR7 receptor has also been discovered in many cancer types and in these cases, the receptor has been seen to boost cancer metastasis. Cancer cells may even use the same mechanism as immune cells to guide their movement. Therefore, our findings may help design new strategies to modify immune responses as well as to target certain cancers," notes Jonna Alanko.

Jonna Alanko is a postdoctoral researcher in the InFLAMES Flagship, at the MediCity research laboratory of the Faculty of Medicine at the University of Turku in Finland. She conducted a majority of her recently published study at the Institute of Science and Technology Austria (ISTA), in Austria, in a research group led by Professor Michael Sixt.

**More information:** Jonna Alanko et al, CCR7 acts as both a sensor and a sink for CCL19 to coordinate collective leukocyte migration, *Science Immunology* (2023). DOI: 10.1126/sciimmunol.adc9584

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