

## Bacterial immune system has a better memory than expected

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A sample with phages growing on a bacterium is pipetted onto agar. The clear spots, called plaques, arise if a single phage particle multiplies and kills all the surrounding bacteria. At number 3, more than 100 plaques can be seen, which indicates that the same number of phages capable of infecting this type of bacterium is present in the sample.

Bacteria's memories of hostile viruses are stronger than thought. Even when the intruders change their DNA sequence, the immune system of bacteria can recognise these and subsequently destroy them. That is the conclusion of NWO Vidi researcher Stan Brouns in the journal *PNAS*.

Bacteria are continuously attacked and killed by intruding viruses. To survive this conflict <u>bacteria</u> have developed an immune system, called



CRISPR-Cas (clustered regularly interspaced short palindromic repeats—CRISPR associated). Although there are other less-developed immune mechanisms, the immune system CRISPR-Cas has a 'memory', as a result of which the bacteria are protected against viral infections.

## Prior knowledge

Bacteria add pieces of DNA from intruding phages (small viruses that only infect bacteria) or plasmids (DNA that exchanges information between unicellular organisms) to the memory system of CRISPR-Cas. The DNA of the phages is subsequently broken down and the intruder is rendered harmless. However, phages avoid being recognised by the CRISPR-Cas system by changing their DNA sequence. This makes it harder for the immune system of the bacteria to identify the viral DNA.

Microbiologist Dr Stan Brouns from Wageningen University has discovered that the immune system of bacteria is more robust than had previously been thought: it can recognise viruses by making use of vague memories. Somehow or other the <a href="immune system">immune system</a> takes up pieces of the now altered DNA and destroys the intruder. In a certain sense the bacterium repeats the same trick but now with prior knowledge. As a vague memory has already been stored in the memory the process now proceeds faster and more efficiently. The CRISPR-Cas system therefore seems able not only to resist recent intruders but also old intruding viruses and intruders related to new viruses.

## **Yoghurt**

The CRISPR-Cas system appears to have a better memory against intruders than had previously been thought; it makes bacteria resistant to external pathogens and therefore gives bacteria an advantage in the evolutionary struggle with intruding <u>viruses</u>. This knowledge can be used



to accelerate the resistance of bacteria to phages that threaten milk fermentations (like in yoghurt and cheese).

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