

Eliminating resistant bacteria with nanoparticles

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The researchers have developed nanoparticles (red) that can kill resistant bacteria (yellow) incorporated by cells of the body (coloured electron microscopy image). Credit: Empa

Novel nanoparticles developed by researchers at ETH Zurich and Empa detect multi-resistant bacteria hiding in body cells and kill them. The scientists' goal is to develop an antibacterial agent that is effective where conventional antibiotics remain ineffective.



In the arms race "mankind against <u>bacteria</u>," bacteria are currently ahead of us. Our former miracle weapons, antibiotics, are failing more and more frequently when germs use tricky maneuvers to protect themselves from the effects of these drugs. Some species even retreat into the inside of <u>human cells</u>, where they remain "invisible" to the <u>immune system</u>. These particularly dreaded pathogens include multi-resistant staphylococci (MRSA), which can cause life-threatening diseases such as sepsis or pneumonia.

In order to track down the germs in their hideouts and eliminate them, a team of researchers from ETH Zurich and Empa is now developing nanoparticles that use a completely different mode of action from conventional antibiotics: While antibiotics have difficulty in penetrating human cells, these nanoparticles can penetrate the membrane of affected cells. Once there, they can fight the bacteria.

Bioglass and metal

The team led by Inge Herrmann, a professor of Nanoparticulate Systems at ETH Zurich and researcher at Empa in St. Gallen, used cerium oxide, a material with antibacterial and anti-inflammatory properties in its nanoparticle form. The researchers combined the cerium oxide with a bioactive ceramic material known as bioglass and synthesized nanoparticle hybrids from the two materials.

In cell culture and using electron microscopy, they investigated the interactions between the hybrid nanoparticles, human cells and bacteria. When the scientists treated cells infected with bacteria with the nanoparticles, the bacteria inside the cells began to dissolve. However, if the researchers specifically blocked the uptake of the hybrid particles into the cells, the antibacterial effect was gone.



Development of resistance less likely

The particles' exact mode of action is not yet fully understood. It has been shown that other metals also have antimicrobial effects. However, cerium is less toxic to human cells than, for instance, silver. Scientists currently assume that the nanoparticles affect the cell membrane of the bacteria, creating <u>reactive oxygen species</u> that lead to the destruction of the germs. Since the cell membrane of human cells is structured differently than that of bacteria, our cells are not affected by this process.

The researchers think that resistance is less likely to develop against a mechanism of this kind. Next, the researchers aim to analyze the interactions of the particles in the infection process in more detail in order to further optimize the structure and composition of the nanoparticles. Their goal is to develop a simple and robust antibacterial agent that is effective inside infected <u>cells</u>.

More information: Martin T. Matter et al. Inorganic nanohybrids combat antibiotic-resistant bacteria hiding within human macrophages, *Nanoscale* (2021). DOI: 10.1039/d0nr08285f

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