

For cells, some shapes are easier to swallow than others

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Credit: David Richards

Scientists have probed the process that allows cells to swallow up particles, finding that some shapes are easier to swallow than others. Cells take in small particles and other objects such as bacteria in a process called engulfment. Single-celled organisms use engulfment to take in food, while in humans it forms the first line of immune system defence, as white blood cells engulf harmful bacteria and other foreign bodies.

In a new study, researchers at the University of Exeter and Imperial College London have used a simple model to investigate how the basic shapes of particles and bacteria can affect how easily they are engulfed by cells.



This could help when designing the optimal <u>shape</u> for drugs, helping them to be more easily taken up by target cells. It could also help explain why some bacteria are especially good at avoiding being engulfed. The results are published today in *Proceedings of the National Academy of Sciences*.

During engulfment, the membrane is pushed out and reaches around the foreign particle, enveloping it and pulling it into the cell to be broken down. Until now, models of this process imagined the particle as a simple sphere, despite the wide range of different particle and bacteria shapes found in nature.

The new model reveals that the easiest shapes to engulf are rod shapes and ovals – but only if they are approached by the cell end-on. On their side, they become more cumbersome. The most difficult shapes to engulf are hourglass-types that can be created, for example, by bacteria dividing while being engulfed. These shapes appear to get stuck and are not fully engulfed.

Co-author Dr Robert Endres from Imperial's Department of Life Sciences said: "Engulfment is difficult to observe experimentally as <u>immune cells</u> don't like the laser light we usually use for imaging, which is why only simple spheres have been used in theoretical models so far.

"However, our technique paves the way for models closer to reality, with the dual advantage of understanding bacteria's defence mechanisms and aiding drug design."

The team's model combines two crucial elements. The first is a mathematical model of an expanding membrane cup shape that closely resembles models describing the freezing of water into ice, mathematically known as the supercooled Stefan problem.



The other is information about the active process that cells employ when engulfing particles. When a harmful object enters the body, the immune system emits signalling molecules. The model takes account of how these molecules help guide the bending of the membrane around the shape.

Co-author David Richards, from the Centre for Biomedical Modelling and Analysis at the University of Exeter, said: "This study tells us how the shape of bacteria affects how easily our immune cells can destroy them. The shape of some bacteria might make them extremely difficult for the body to destroy, which could be deadly.

"This work could be useful in the pharmaceutical industry, as it might pave the way to novel designs for very small drug-carrying particles that the body cannot accidentally destroy."

The team hopes to expand their simple model to actual <u>bacteria</u> and engineered drug particles with wide-ranging material properties.

More information: David M. Richards et al. Target shape dependence in a simple model of receptor-mediated endocytosis and phagocytosis, *Proceedings of the National Academy of Sciences* (2016). DOI: <u>10.1073/pnas.1521974113</u>

Provided by University of Exeter

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