

Mesoporous silica capsules

When you next eat a curry you could be helping to protect yourself from cancer.



Curry contains the spice turmeric and a major component of turmeric is curcumin. Curcumin is known to have anti-tumour and anti-cancer capabilities. Indian people, whose diet includes turmeric, have significantly lower rates of bowel and intestinal cancer than people who eat a western diet. However, curcumin is poorly absorbed from the intestine so to benefit from these properties a lot has to be eaten.

Scientists at the Centre for Strategic Nano-Fabrication at The University of Western

Australia have been working on a way to encase curcumin in tiny silica capsules to ensure better uptake by the body through a 'slow-release' delivery system.

Mesoporous silica (SiO_2) capsules can be used to carry curcumin because they are non-toxic and chemically inert.



How capsules are made

A multi-step, self-assembly approach is used to make these capsules. The first step uses a chemical called cetyltrimethylammonium bromide (CTAB). CTAB is a surfactant, that is, its molecule has a hydrophobic (water-hating) end and a hydrophilic (water-loving) end. This means it can form bonds with both water and oil molecules.

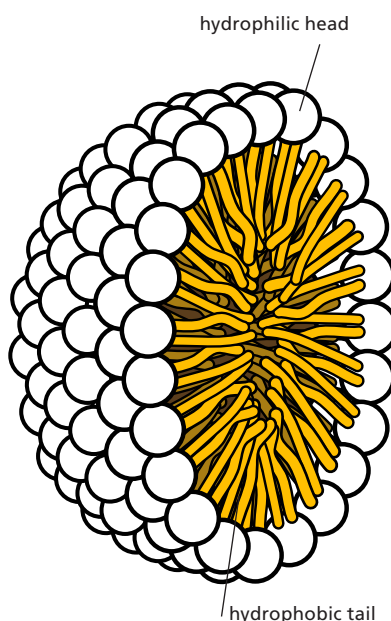


Figure 1: Micelle in aqueous solution.

CTAB molecules clump together in aqueous solution to form surfactant micelles. Hydrophilic heads of the molecules form the outside layer, and hydrophobic tails all point inwards in this spherical structure.

Curcumin, like oil, is hydrophobic and therefore not soluble in water. However, when mixed with the CTAB solution, it is taken up into the centre of micelles where it is surrounded by the hydrophobic tails. Under the right conditions, micelles can be made to aggregate in rod-shaped structures (micellar rods) instead of spheres. In this case curcumin molecules are trapped in the middle of the rods.

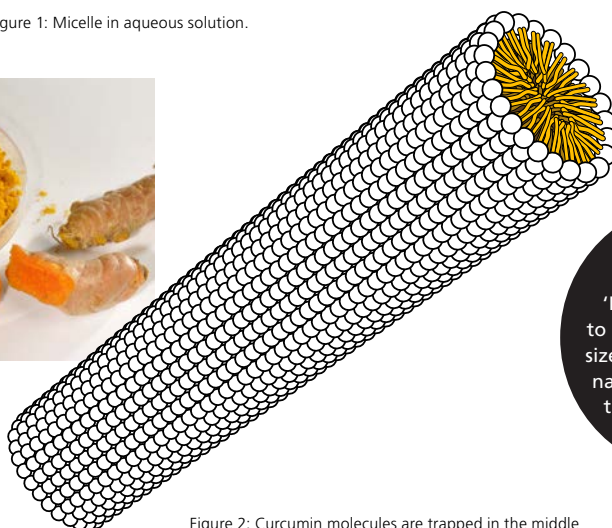
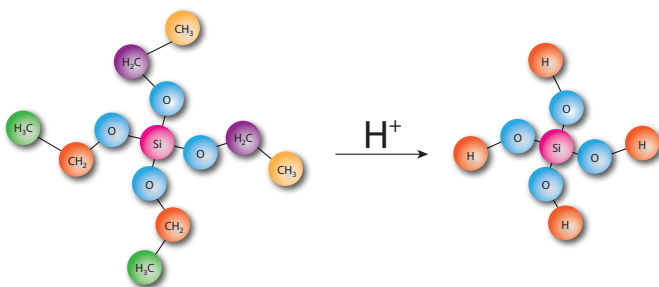
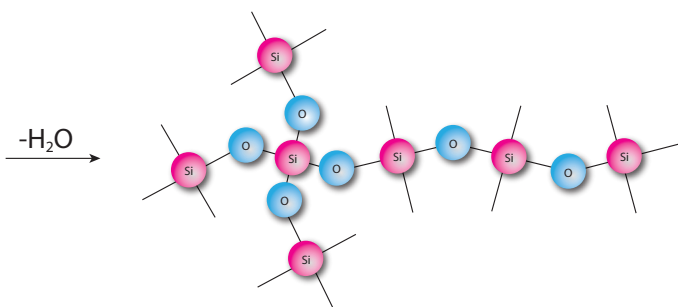


Figure 2: Curcumin molecules are trapped in the middle of micellar rods.

Did you know?

'Mesoporous' refers to materials with nano-sized pores in them — a nanometer is a million times smaller than a millimetre!

These rods are used as templates for the capsules by coating them with silica. A silica compound, tetraethyl orthosilicate (TEOS), is added to the CTAB/curcumin mixture. This covers the outside of the micellar rods. When acid is added, ethyl groups are removed from TEOS to form a coating of $\text{Si}(\text{OH})_4$. Water is then removed from $\text{Si}(\text{OH})_4$ to form covalent network silica (SiO_2), which 'wraps up the parcel' ready for delivery.

Figure 3: Formation of Si(OH)_4 by addition of acid to TEOS.Figure 4: Removal of water creates a SiO_2 network.

Pores are created as silica wraps around the rods. Pore size, which is important in controlling the rate of release of curcumin, can be determined by varying the surfactant used. To be classified as mesoporous, pores need to be 2 to 50 nm in diameter.

Micellar rods line up in regular structures as they are coated. The silica coating then bends and folds into various 'origami' shapes, ranging in size from 5 to 10 μm (about the same size as a red blood cell). About 150 silica capsules would fit across a pinhead!

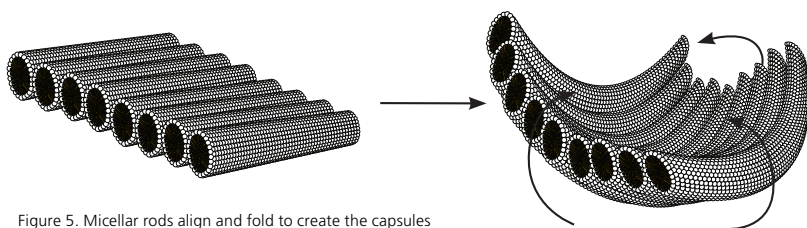


Figure 5: Micellar rods align and fold to create the capsules



Nigel Clifford

Nigel Clifford is a Green Chemistry honours graduate from The University of Western Australia.

The Green Chemistry program offered by The University of Western Australia is focussed upon chemistry that is non-hazardous for both people and the environment.

As part of his honours degree Nigel developed a way of using mesoporous silica capsules as controlled release carriers of curcumin in the human body. Controlled-release medicines mean far fewer side effects and more effective medicinal uptake.

Nigel's research offers a new approach to the delivery of medicines to the human body, and may well be the drug delivery system of the future.

References

- 1) Clifford, N. W., Iyer, K. S. and Raston, C. L. (2008). Encapsulation and controlled release of nutraceuticals using mesoporous silica capsules. *Journal of Materials Chemistry*, 18, 162-165.
- 2) Lee, Y. S., Surjadi, D. and Rathman, J. F. (1996). Effects of Aluminate and Silicate on the Structure of Quaternary Ammonium Surfactant Aggregates. *Langmuir*, 12, 6202-6210.

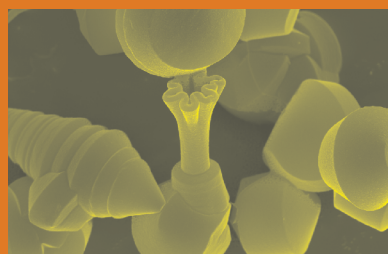
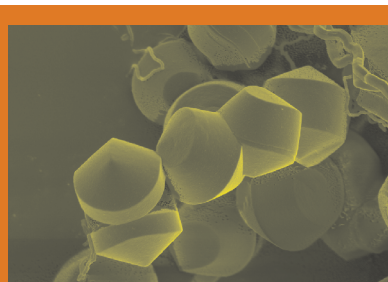


Figure 6: Capsules form a wide range of shapes, depending on the reactants used.

Slow release of curcumin from capsules has been measured in test tubes with the same acid concentration (pH) that is found in blood. Controlled release should lead to much better uptake of curcumin, but at this time not enough is known about the effects of silica capsules on the body for them to be trialled in animals or humans.

In the future this method of drug delivery may be ideal for expensive or poorly absorbed drugs. Curcumin was used in this research as a model for hydrophobic drugs in general. The challenge ahead for the research group is to adapt these methods to a range of drugs and to turn it into a commercial product.