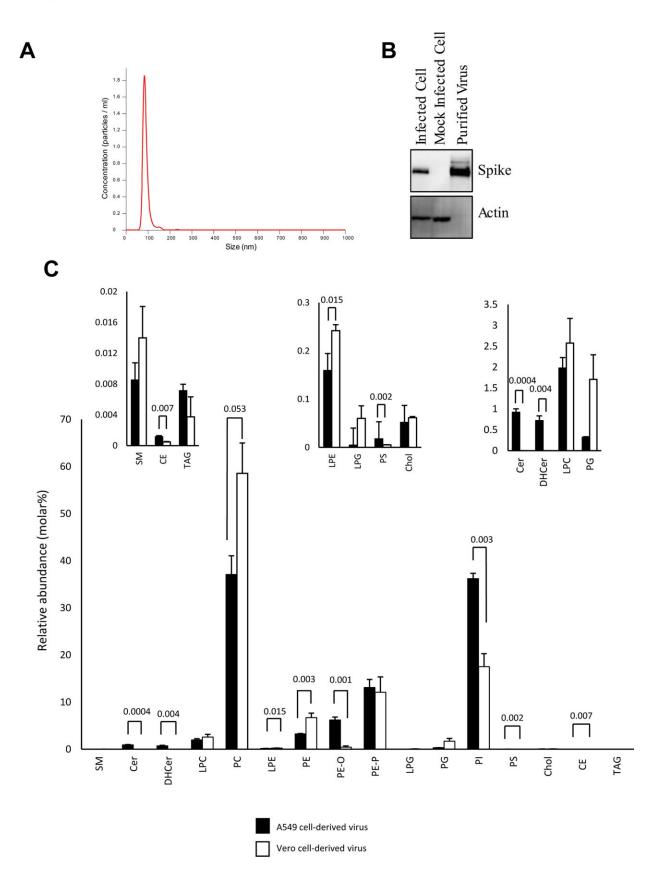


Make-up of SARS-CoV-2 outer envelope revealed for the first time

April 25 2022

Medical





Nanoparticle tracking analysis and western blotting confirm purity of SARS-CoV2 preparations, while targeted lipidomics demonstrates the membrane as a PL rich membrane mainly comprised of PE, PC and PI. Panel A-B purity analysis of gradient purified viral preparations. (A) Gradient purified virus was analyzed by nanoparticle tracking analysis and particle size plotted. (B) Proteins was solubilised in NuPAGE LDS buffer then separated by size on Bis-Tris gels, before being transferred to nitrocellulose and blotted for the indicated proteins, as outlined in Methods (see Supplementary Figure 1 for uncropped gels) Panel C. Lipidomics analysis of the total amounts of lipids in each category in Vero and A549 cells. Lipids were extracted from 3 preparations of virus from either Vero or A549 cells, and analyzed using LC/MS/MS, as indicated in Methods. The relative % of all detected lipid categories for all 3 preparations, with molecular species within each category combined to provide total values are shown. Amounts (ng) of all individual molecular species were added together for each preparation then converted to molar amounts using an average mass value per category. Molar% was then calculated following totalling of all lipid categories (n = 3, mean +/- SEM). Unpaired Students T test. Credit: Journal of Lipid Research (2022). DOI: 10.1016/j.jlr.2022.100208

Scientists from Cardiff University have detailed the molecular make-up of the fatty outer envelope of SARS-CoV-2 for the first time—and say it could prove an important new target for antivirals.

The team's work on the viral <u>lipid</u> envelope showed it had significant differences to healthy host cell membranes, suggesting the potential for it to be selectively targeted.

The research is published today in the Journal of Lipid Research.

"While vaccines and antivirals have targeted proteins or the replication cycle of SARS-CoV-2, there has been little research into the lipid envelope to date," said Professor Valerie O'Donnell, a lipid biochemist and co-author on the study from Cardiff University's School of



Medicine.

"Our study has mapped this essential viral component—indeed, there has been very little research into any viral lipid envelopes so far. Addressing this knowledge gap could enable selective therapeutic targeting that avoids damaging host membranes. We also suspect the membrane won't be impacted by mutation of the virus, unlike spike proteins, so it would be a consistent target."

The Cardiff team used <u>mass spectrometry</u> to create a <u>detailed map</u> of the little-understood lipid envelope and found it was comprised mainly of phospholipids but very little cholesterol, unlike host cell membranes. The findings suggest there is potential to target the virus in the <u>oral</u> <u>cavity</u> with therapeutics without collateral damage to the body's own cells.

As part of the same study, they carried out a small-scale clinical trial in hospitalized COVID-19 patients to test whether oral rinses containing lipid disrupting chemicals called surfactants could reduce infectivity in the mouth.

In the trial, 27 <u>hospital patients</u> with PCR-confirmed COVID-19 were given one of four different types of oral rinse, including two containing cetylpyridinium chloride (CPC), one containing povidone-iodine and another containing saline.

The most effective contained CPC and isopropyl myristate—a single 30-second rinse with this formulation in seven patients eliminated the virus in the saliva by 99.99% at one minute, and by 99.8% at 60 minutes. In three of the seven patients (around 40%) no live virus was detected in the saliva at any time after the initial rinse.

Other formulations containing compounds such as povidone-iodine or



saline had little to no effect, while the rinse containing CPC and benzoate did not show a persistent effect, suggesting formulation is "critical."

Professor Richard Stanton, co-author of the study from Cardiff University's School of Medicine, said: "The findings of our clinical trial suggest the lipid envelope could be targeted by specific formulations of oral rinses.

"Our results suggest certain mouthwashes could play a role in limiting COVID-19 exposure in healthcare settings, for example, destroying the virus for sufficient time to enable a dental or oral examination. It could also be a useful component of infection prevention and control for other respiratory enveloped viruses, such as influenza."

Professor David Thomas, co-author of the study from Cardiff University's School of Dentistry, said: "Larger <u>clinical trials</u> and population-based studies are warranted to determine if oral rinses have any impact on COVID-19 transmission in practice.

"We would encourage people to always use mouthwash safely and as per the manufacturer's guidelines."

An important caveat, say the researchers, is that mouthwash cannot target any virus in the <u>lower respiratory tract</u>, and it remains unclear whether <u>virus</u> that transmits to others arises from the upper or lower respiratory tract.

More information: Zack Saud et al, The SARS-CoV2 envelope differs from host cells, exposes pro-coagulant lipids, and is disrupted in vivo by oral rinses, *Journal of Lipid Research* (2022). DOI: 10.1016/j.jlr.2022.100208



Provided by Cardiff University

Citation: Make-up of SARS-CoV-2 outer envelope revealed for the first time (2022, April 25) retrieved 3 May 2024 from https://medicalxpress.com/news/2022-04-make-up-sars-cov-outer-envelope-revealed.html

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