

Injection-free vaccination technique could address global vaccine challenge for HIV, malaria

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Scientists at King's College London have demonstrated the ability to deliver a dried live vaccine to the skin without a traditional needle, and shown for the first time that this technique is powerful enough to enable specialised immune cells in the skin to kick-start the immunising properties of the vaccine.

Funded by the Bill & Melinda Gates Foundation and published today in *Proceedings of the National Academy of Sciences*, researchers say although it is an early study this important technical advance offers a potential solution to the challenges of delivering live vaccines in resource-limited countries globally, without the need for refrigeration. A cheaper alternative to hypodermic needles, it would also remove safety risks from needle contamination and the pain-free administration could lead to more people taking up a vaccination. The researchers add that it could have an impact beyond infectious disease vaccination programmes, for example managing autoimmune and inflammatory conditions such as diabetes.

HIV, malaria and TB represent major global health challenges. Although promising research is underway to develop vaccines for these diseases, considerable stumbling blocks remain for countries where transporting and storing live vaccines in a continuously cold environment (around 2°C to 8°C or below) would not be possible. If a cold chain cannot be maintained for a live vaccine there is a high risk it could become unsafe



and lose effectiveness.

The team at King's used a silicone mould developed by US company TheraJect to create a microneedle array – a tiny disc with several microneedles made of sugar which dissolve when inserted into the skin. The team formulated a dried version of a live modified adenovirus-based candidate HIV vaccine in sugar (sucrose) and used the mould to create the microneedle array. They found that the dried live vaccine remained stable and effective at room temperature.

To test the effectiveness of the microneedle array, they applied it to mice. Using imaging (in collaboration with Professor Frederic Geissmann, King's College London) they observed how the vaccine dissolved in the skin and were able to identify for the first time exactly which specialised <u>immune cells</u> in the skin 'pick up' this type of vaccine and activate the immune system. The researchers found the first evidence that a sub-set of specialised dendritic cells in the skin were responsible for triggering this immune response.

When compared with a traditional needle vaccine method, the immune response generated by the dried microneedle vaccine (kept at room temperature) was equivalent to that induced by the same dose of injected liquid vaccine that had been preserved at -80°C.

Dr Linda Klavinskis from the Peter Gorer Department of Immunobiology at King's College London, said: 'We have shown that it is possible to maintain the effectiveness of a live vaccine by drying it in sugar and applying it to the skin using microneedles – a potentially painless alternative to hypodermic needles. We have also uncovered the role of specific cells in the skin which act as a surveillance system, picking up the <u>vaccine</u> by this delivery system and kick-starting the body's immune processes.



'This work opens up the exciting possibility of being able to deliver live vaccines in a global context, without the need for refrigeration. It could potentially reduce the cost of manufacturing and transportation, improve safety (as there would be no loss in potency), and avoids the need of hypodermic needle injection, reducing the risk of transmitting bloodborne disease from contaminated needles and syringes.

'This new technique represents a huge leap forward in overcoming the challenges of delivering a vaccination programme for diseases such as HIV and malaria. But these findings may also have wider implications for other infectious disease vaccination programmes, for example infant vaccinations, or even other inflammatory and autoimmune conditions such as diabetes.'

Provided by King's College London

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