

New method maps chemicals in the skin

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3-D imaging of a skin section in which the nickel content is being measured. The



red area represents the presence of nickel. Credit: Per Malmberg

A new method of examining the skin can reduce the number of animal experiments while providing new opportunities to develop pharmaceuticals and cosmetics. Chemical imaging allows all layers of the skin to be seen and the presence of virtually any substance in any part of the skin to be measured with a very high degree of precision.

More and more chemicals are being released into our environment. For example, parabens and phthalates are under discussion as two types of chemicals that can affect us. But so far it has not been possible to find out how they are absorbed by the skin. A new study from Chalmers University of Technology and the University of Gothenburg shows how what is termed <u>chemical imaging</u> can provide comprehensive information about the human skin with a very high level of precision.

Investigations into how substances pass into and through the skin have so far taken place in two ways: by using tape strips to pull off the top "corneal" layer of skin for analysis, and through urine and blood testing to see what has penetrated through the skin. But we still know very little about what happens in the layers of skin in between. Chemical imaging now allows us to see all layers of the skin with very high precision and to measure the presence of virtually any substances in any part of the skin. This can lead to pharmaceutical products that are better suited to the skin, for example.

The new method was created when the chemists Per Malmberg, at Chalmers University of Technology, and Lina Hagvall, at the University of Gothenburg, brought their areas of research together.





Chemists Lina Hagvall and Per Malmberg prepare samples for analysis in the time-of flight secondary ion mass spectrometer (ToF-SIMS). During analysis the samples are inserted into the test chamber using the test arm shown at the bottom of the picture. Credit: Mats Tiborn

"With pharmaceuticals you often want as much as possible of the dose to be absorbed by the skin, but in some cases you may not want skin absorption, such as when you apply a sunscreen, which needs to remain on the surface of the skin and not penetrate it. Our method allows you to design pharmaceuticals according to the way you want the substance to be absorbed by the skin," says Hagvall.

Chemical imaging has until now mainly been used for earth sciences and cellular imaging, but with access to human skin from operations the



researchers have come up with this new area for the technology. The researchers now also see opportunities opening up for replacing pharmaceutical tests which currently involve <u>animal experiments</u>. Their methods provide more accurate results than tests on mice and pigs. Since it is not permissible to use animals to test cosmetics, this method may also create new opportunities for the cosmetics industry.

"Many animal experiments carried out by researchers and companies are no longer necessary as a result of this method. If you want to know something about passive absorption into the human skin, this method is at least as good. It's better to do your testing on <u>human skin</u> than on a pig," says Hagvall.

The new method can also provide a basis for determining the correct limits for harmful levels of substances that may come into contact with the skin. In order to establish those limits, you need to know how much of the dose on the skin's surface penetrates into and through the skin, which this method can show. It enhances our knowledge about what we are absorbing in our workplaces and in childcare facilities.





Test chamber in the time-of-flight secondary ion mass spectrometer (ToF-SIMS) in which the skin samples are exposed to ion beams. The samples are analysed in an ultra-high vacuum to avoid disruption. Credit: Mats Tiborn

"Our method can show everything with an image, whether you are looking for nickel, phthalates or parabens in the skin, or if you want to follow the drug's path through the skin. With just a skin sample we can essentially search for any molecules. We don't need to adapt the method in advance to what we are looking for," says Malmberg.

It will be possible to apply the results in the very near future. The technology itself is ready for use today. There is still a small amount of work left to do in optimising the tests to achieve the best results, but the researchers believe that the <u>method</u> will be ready for use within a year.



Facts: Chemical imaging

Chemical imaging involves the use of a laser or ion beam to analyse the sections of skin using a mass spectrometer. Every dot, or pixel, of the section which the beam strikes provides information, which is used to classify the chemicals present in the skin according to molecular weight. The <u>chemical</u> information from each dot can then be combined into a digital image which shows the distribution of a substance in the <u>skin</u>. A time-of-flight secondary ion <u>mass spectrometer</u> (ToF-SIMS), which provides a very <u>high spatial resolution</u> down to the nanometre range, was used in this particular study.

More information: Per Malmberg et al, Imaging mass spectrometry for novel insights into contact allergy - a proof-of-concept study on nickel, *Contact Dermatitis* (2017). <u>DOI: 10.1111/cod.12911</u>

Provided by Chalmers University of Technology

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