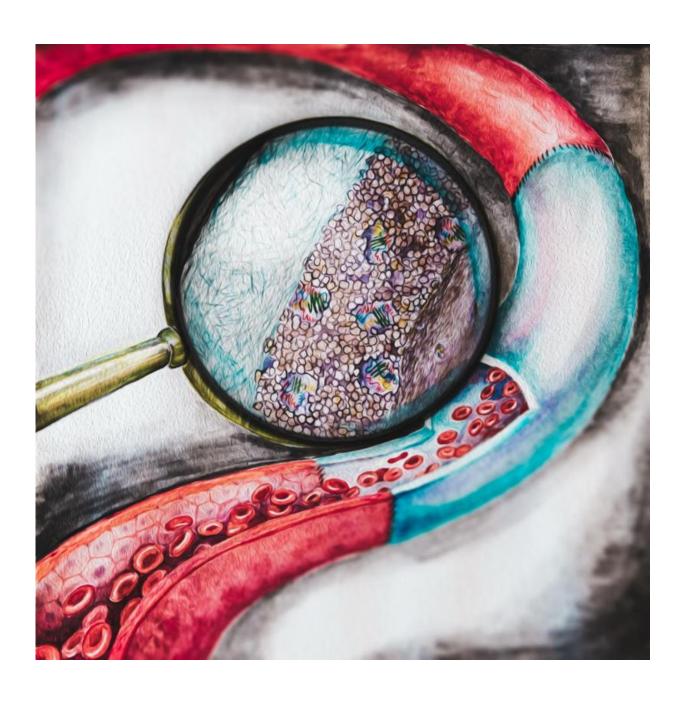


Artificial blood vessels become resistant to thrombosis

August 4 2015





This is an artist's representation of the improved vascular graft. The enlarged section shows the drug-entrapping coating attached to the inner surface of the graft. Credit: ITMO University/Yulia Chapurina

Scientists from ITMO University have developed artificial blood vessels that are not susceptible to blood clot formation. The achievement was enabled by a next-generation, drug-containing coating applied to the inner surface of the vessel. The results of the study were published in the *Journal of Medicinal Chemistry*.

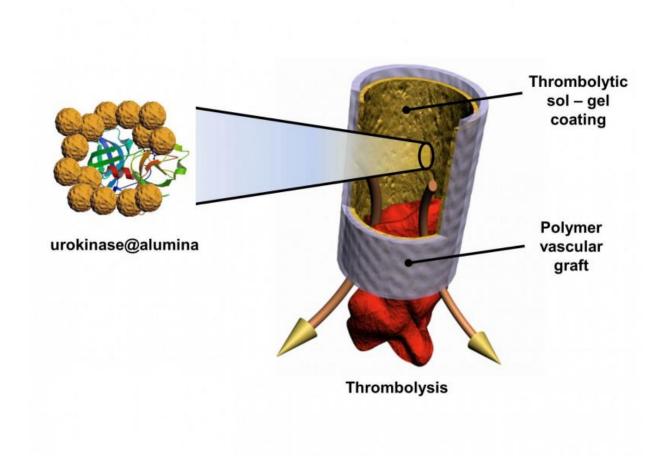
Surgery associated with cardiovascular diseases such as ischemia often requires the implantation of <u>vascular grafts</u>— <u>artificial blood vessels</u> that restore <u>blood</u> flow in a problematic part of the circulatory system. A serious disadvantage of vascular grafts is their tendency to become blocked due to clot formation, which results in compulsory and lifelong intake of anticoagulants among patients and sometimes requires additional surgical intervention.

In the study, a research team led by Vladimir Vinogradov, head of the International Laboratory of Solution Chemistry of Advanced Materials and Technologies at ITMO University, proposed a solution to the problem. The team synthesized a thin film made of densely packed aluminum oxide nanorods blended with molecules of a thrombolytic enzyme (urokinase-type plasminogen activator). Adhered to the innersurface of a vascular graft, the film causes the parietal area of the graft to fill with a stable concentration of a substance called plasmin that is capable of dissolving the clots.

The film's structure represents a porous matrix, accommodating the plasminogen activator. The matrix protects the plasminogen activator from the aggressive environment of the organism, at the same time



preserving the ability of the enzyme to interact with certain external agents through a system of pores. In particular, the matrix lets in plasminogen, a proenzyme naturally occurring in blood plasma. When plasminogen meets plasminogen activator inside the matrix, clot-dissolving plasmin forms.



A blood clot leaking through the graft is depicted. Credit: ITMO University

Yulia Chapurina, laboratory researcher and first author of the paper, set



up several in vitro experiments that demonstrated just how effective the film is:

"In order to test how our improved vascular graft worked, we grew an artificial clot made of blood plasma mixed with thrombin and placed it inside the graft. The results of the experiment amazed us. Very soon, the clot started to dissolve and leak through the graft. In vivo, our coating would destroy clots during formation, constantly ensuring an unobstructed blood flow in the graft."

Current-generation vascular grafts rely on drug-eluting technology, actively releasing medicine into the blood. The lifetime of such grafts is often determined by the amount of drug stored within the graft, which, in essence, means that they merely postpone the formation of clots. The new system is based on the entrapment of the drug inside a porous protective shell, which makes the lifetime of such a graft practically unlimited.

"Our approach is concept-based and can be applied not only to artificial blood vessels, but to any kind of implant. You just need to take the right kind of drug. For example, after the implantation of an artificial ureter, urease crystals often start to grow inside and doctors do not know how to deal with this problem. It is possible to apply a similar drug-containing coating that dissolves urease. The same approach may be used for kidney or liver surgery, but these are plans for the future," concludes Vladimir Vinogradov.

Provided by ITMO University

Citation: Artificial blood vessels become resistant to thrombosis (2015, August 4) retrieved 25 April 2024 from

https://medicalxpress.com/news/2015-08-artificial-blood-vessels-resistant-thrombosis.html



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