

Discovering crucial role of protein that could stop HPV virus infection

April 30 2018



Credit: University of Leeds

New drugs in the early stages of development by pharmaceutical companies could have an extra benefit – the ability to cancel out HPV virus, which can cause cervical cancer and skin conditions.

Scientists at the University of Leeds and University of Birmingham, whose research focuses on Human Papilloma Virus (HPV), say many people are unaware it is also responsible for numerous other cancers,



especially those affecting the head and neck, which are increasing and becoming more prevalent in men.

They have now discovered the role of one of the crucial proteins responsible for allowing HPV to infect <u>cells</u>. It is called STAT3 and is essential for HPV to replicate.

The team, led by Dr. Andrew Macdonald at the University's Astbury Centre for Structural Molecular Biology, said without STAT3 the <u>virus</u> could not work because the full set of properties it needed would not be available.

Completing the pathway

His team, funded by Wellcome and the Medical Research Council, has also found the individual enzymes which activates STAT3 in cells infected with HPV.

Uncovering the identity and role of these enzymes which 'switch on' STAT3 completes the chain and means scientists have a new pathway to work through to target HPV-based cancers.

Dr. Macdonald said the fact drugs were already being developed which targeted the same enzymes for other uses meant the possibility of a new treatment could be much closer to becoming a reality.

He said: "We need to find new ways of treating HPV cancers, so we have been working towards understanding how the virus works and interacts with cells.

"Now we have identified both the role of STAT3 and how it is activated, we have a potential target for future studies in combating HPV associated disease.



"We know there are various drugs being developed that have the potential to target the enzymes we are dealing with for other purposes. This is great news because it means that while there is still a long way to go, we are already over the first hurdle."

Controlling the virus

The team used samples of donated <u>human skin cells</u> to understand how the virus increased cell growth, and found that when STAT3 was not active, this was prevented.

Ph.D. researcher Ethan Morgan, who carried out much of the lab work to identify the roles of STAT3 and the enzymes that switch it on, added: "The process of uncovering these enzymes and the way they activate STAT3 has been really important because we have now found an approach which could prevent HPV infection. The next step for us will be more time in the lab refining our work and then confirming the role of STAT3 in HPV associated cancers."

Currently there is no screening programme for head and neck cancers and, while there is a HPV vaccine, it is only effective if administered before infection with the virus.

The research has been published in the journal PLoS Pathogens.

More information: Ethan L. Morgan et al. STAT3 activation by E6 is essential for the differentiation-dependent HPV18 life cycle, *PLOS Pathogens* (2018). DOI: 10.1371/journal.ppat.1006975

Provided by University of Leeds



Citation: Discovering crucial role of protein that could stop HPV virus infection (2018, April 30) retrieved 6 May 2024 from https://medicalxpress.com/news/2018-04-crucial-role-protein-hpv-virus.html

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