

Scientists pave way to reducing antibiotic resistance build-up in waterways

October 21 2020



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Researchers have developed a swift new method that could help reduce the problem of antibiotic resistance increasing via wastewater systems.

Around 70 per cent of the antibiotics we take as medicine ends up in the



natural environment, through trace residues excreted by patients and inappropriate disposal of medicines, among other sources. Bacteria are also present in these wastewaters, and when they are exposed to antibiotics they can evolve resistance within these environments. This could mean an increased threat to <u>human health</u>, if <u>resistant bacteria</u> enter and colonise the gut, for example through swallowing water while swimming. The bacteria could then transfer resistance to human-associated bacteria, meaning antibiotics are less likely to work when they are needed.

Previous research by the team at the University of Exeter and AstraZeneca had suggested the need to introduce safe thresholds on the concentration of antibiotics to release into sewers and other wastewater systems.

Now, in research published in *Environmental Health Perspectives* and funded by the Natural Environment Research Council, the Biotechnology and Biological Sciences Research Council and AstraZeneca, the team has developed a way to solve the problem. They have developed a rapid, cost-effective method to predict the lowest concentration of an antibiotic that increases resistance. The method opens the door to calculating what level of antibiotics are safe to release into wastewaters to minimise the impact on the rise of antibiotic resistance. The method can also be used to assess the impact of combinations of chemicals and antibiotics.

Antibiotic resistance is recognised by the World Health Organisation as one of the greatest health threats of our time. By 2050, up to 10 million deaths each year could be caused by antibiotics and other antimicrobial drugs no longer working to treat common diseases, including respiratory tract, sexually transmitted and urinary tract infections. The threat of resistance could also increase the risk of contracting infection after basic surgical procedures.



Dr. Aimee Murray, of the University of Exeter, who led the research said: "As part of the global battle to stop millions of deaths from the failure of antibiotics, we urgently need to reduce resistance building up in our natural environment. At the moment, environmental risk assessment guidelines don't require any tests that measure increases in antimicrobial resistance when antibiotics are released into the environment. Our research provides data that can underpin safe limits, a way to generate that data quickly, and could provide evidence to inform decisions about safe levels of antibiotics in the environment."

The method extracts bacteria from wastewater and exposes them to many different antibiotic concentrations. When the bacteria are actively growing and dividing under these conditions, the method finds the lowest antibiotic concentration that reduces growth, compared to when no antibiotic is present. The team show this is very similar to the lowest antibiotic concentration that increases numbers of key antibiotic resistance genes, meaning the new method can generate the same result in significantly less time, using fewer resources, specialised equipment and techniques.

The research team validated the method using these more complex and previously published experimental techniques and generated the largest available experimental dataset on the lowest antibiotic concentrations that promote resistance using a single method.

The research team also used the new method, called SELECT, on a number of commonly-prescribed <u>antibiotics</u>, and revealed which pose the highest risk of developing resistance in the natural environment.

Professor Will Gaze, of the University of Exeter, oversees the research programme. He said: "Our results indicate that the antibiotic ciprofloxacin poses the most significant risk of promoting antimicrobial resistance in the environment, supporting its inclusion on the current



Water Framework Directive's 'Watch List'. Our data also indicate trimethoprim, cefotaxime and azithromycin pose risks for increasing levels of antimicrobial resistance in the most impacted environments."

Professor Jason Snape, Global Head of Environment in AstraZeneca said "There is a clear environmental dimension to antimicrobial resistance and it is imperative that we have the tools to identify safe concentrations in the <u>environment</u>, from antibiotic manufacture and patient use Our existing tools protect wildlife, specifically primary productivity (fixation of carbon dioxide by algae and cyanobacteria), but they don't address the potential to select or enrich for resistance. The SELECT assay offers us this potential."

More information: The 'SELection Endpoints in Communities of bacTeria' (SELECT) method: A novel experimental assay to facilitate risk assessment of selection for antimicrobial resistance in the environment' *Environmental Health Perspectives* (2020). DOI: 10.1289/EHP6635

Provided by University of Exeter

Citation: Scientists pave way to reducing antibiotic resistance build-up in waterways (2020, October 21) retrieved 26 April 2024 from <u>https://phys.org/news/2020-10-scientists-pave-antibiotic-resistance-build-up.html</u>

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