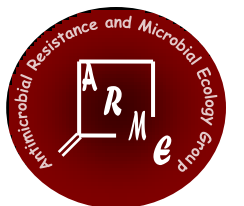


Antibiotic Resistance and the Environment...

...What we Know and What we Need to Know!

Dr. Dearbháile Morris

Antimicrobial Resistance and Microbial Ecology Group, School of Medicine,
Centre for Health from Environment, Ryan Institute,
National University of Ireland Galway.




Antimicrobial Resistance is a major public health problem




HIGH-LEVEL MEETING ON ANTIMICROBIAL RESISTANCE



21 SEPTEMBER 2016, UN HEADQUARTERS, NEW YORK



GLOBAL ACTION PLAN ON ANTIMICROBIAL RESISTANCE



World Health Organization

Review on Antimicrobial Resistance

Tackling drug-resistant infections globally

Home Background Industry Publications Infographics Media About us >

The Review on Antimicrobial Resistance has now completed its work. The content of this website is frozen as an archive of the Review's work, and will not be updated further. The legacy of the Review will be maintained by its sponsor organisations, the Wellcome Trust and the UK Department of Health.



For queries about the legacy of the Review, or for media enquiries for the Review's former chair, Jim O'Neill only, please contact, please contact info@amr-review.org.

"The Review on Antimicrobial Resistance (AMR), was commissioned in July 2014 by the UK Prime Minister, who asked economist Jim O'Neill to analyse the global problem of rising drug resistance and propose concrete actions to tackle it internationally. The Review on AMR was jointly supported by the UK Government and Wellcome Trust, although operated with full independence from both. Established as a two-year, time-limited process, the Review engaged widely with international stakeholders to understand and propose solutions to the problem of drug-resistant infections from an economic and social perspective, and produced its final report and recommendations in the summer of 2016.

"If we fail to act, we are looking at an almost unthinkable scenario where antibiotics no longer work and we are cast back into the dark ages of medicine" – David Cameron, former UK Prime Minister

The real implications of spreading drug resistance will be felt the world over, with developing countries and large emerging nations bearing the brunt of this problem. Routine surgeries and minor infections will become life-threatening once again and the hard won victories against infectious diseases of the last fifty years will be jeopardised. Hospital stays and expenses, for both public health care providers and for out-of-pocket payers will increase significantly. Drug resistant infections are already on the rise with numbers suggesting that up to 50,000 lives are lost each year to antibiotic-resistant infections in Europe and the US alone. Globally, at least 700,000 die each year of drug resistance in illnesses such as bacterial infections, malaria, HIV/AIDS or tuberculosis.

"We have reached a critical point and must act now on a global scale to slow down antimicrobial resistance" – Professor Dame Sally Davies, UK Chief Medical Officer



A European One Health Action Plan against Antimicrobial Resistance (AMR)

Antimicrobial Resistance is a major public health problem

iNAP
PROTECTING ANTIBIOTICS FOR THE FUTURE
ANTAIBHEATHAIGH A CHOSAINT DON TODHCHAÍ

IRELAND'S NATIONAL ACTION PLAN ON ANTIMICROBIAL RESISTANCE 2017 - 2020

An Roinn Sláinte
DEPARTMENT OF HEALTH

Talmhaíochta, Bia agus Mara
Department of Agriculture, Food and the Marine

Strategic Objective 1: Improve awareness and knowledge of antimicrobial resistance

Strategic Objective 2: Enhance surveillance of antibiotic resistance and antibiotic use

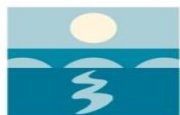
Strategic Objective 3: Reduce the spread of infection and disease

Strategic Objective 4: Optimise the use of antibiotics in human and animal health

Strategic Objective 5: Promote research and sustainable investment in new medicines, diagnostic tools, vaccines and other interventions



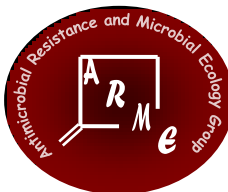
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Environment
Ionad Sláinte ó Chomhshaoil



Antimicrobial Resistance is a major public health problem



ANTIMICROBIALS IN AGRICULTURE AND THE ENVIRONMENT: REDUCING UNNECESSARY USE AND WASTE

THE REVIEW ON ANTIMICROBIAL RESISTANCE

CHAIRIED BY JIM O'NEILL

DECEMBER 2015

3.6 CLOSE KNOWLEDGE GAPS ON AMR IN THE ENVIRONMENT AND ON HOW TO PREVENT TRANSMISSION

AMR is a good example of a One Health matter in which human health is connected to that of animals and the environment. Only a multidisciplinary effort can provide an adequate response. There is a major lack of knowledge about the release and spread of resistant organisms in the environment and the threats and risks this poses to human and animal health. For example, the release of antimicrobials into the environment through human, animal and manufacturing waste streams should be assessed and new technologies developed to enable efficient and rapid degradation of antimicrobials in wastewater treatment plants, organic waste streams or the environment.

The feasibility and implementation of monitoring programmes need to be further studied, including the development of harmonised monitoring of antimicrobials and microorganisms resistant against antimicrobials in the environment. Using harmonised monitoring and research data, risk assessment methodologies should be developed to evaluate risks to human and animal health. In the agri-food sector, the links between farming practices, animal health and AMR development and spread need to be further investigated.

The Commission will:

- ▶ support research into knowledge gaps on the release of resistant microorganisms and antimicrobials into the environment and their spread;
- ▶ explore risk assessment methodologies, with the support of scientific agencies and bodies, and use them to evaluate the risks to human and animal health from the presence of antimicrobials in the environment;
- ▶ support research into and the development of new tools for monitoring antimicrobials and microorganisms resistant against antimicrobials in the environment;
- ▶ support the development of technologies that enable efficient and rapid degradation of antimicrobials in wastewater and the environment and reduce the spread of AMR.

2.4 BETTER ADDRESSING THE ROLE OF THE ENVIRONMENT

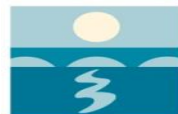
The environment is increasingly acknowledged as a contributor to the development and spread of AMR in humans and animals, in particular in high risk areas due to human, animal and manufacturing waste streams, but strong evidence is still required to better inform decision-making in this area. Specific actions to improve the knowledge base are considered in section 3. Once relevant monitoring and research data become available, risk assessment methodologies should be developed to evaluate the risks to human and animal health.

The Commission will:

- ▶ adopt an EU strategic approach to pharmaceuticals in the environment¹³;
- ▶ maximise the use of data from existing monitoring, e.g. Watch List monitoring under the Water Framework Directive¹⁴, to improve knowledge of the occurrence and spread of antimicrobials in the environment, including by using the Information Platform for Chemical Monitoring (IPChem) to access relevant monitoring data¹⁵;
- ▶ reinforce the role of the Scientific Committee on Health and Environmental Risks (SCHER) in providing the expertise on environment-related AMR issues.



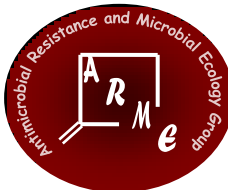
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Environment
Ionad Sláinte ó Chomhshaoil



Antimicrobial Resistance is a major public health problem

- Different types of antimicrobial resistant bacteria.
- Carbapenemase-producing *Enterobacteriaceae* (CPE) are resistant to almost all available antibiotics
- Treatment of infection with CPE very difficult.
- Different types of carbapenemase enzymes – most common in Ireland and the U.K. are KPC, OXA-48 and NDM.
- January 2017 - first report of death due to an untreatable infection caused by NDM-producing *Klebsiella pneumoniae*¹

1. Chen L, Todd R, Kiehlbauch J, et al. Pan-resistant New Delhi metallo-beta-lactamase-producing *Klebsiella pneumoniae* — Washoe County, Nevada, 2016. *MMWR Morb Mortal Wkly Rep* 2017; 66: 33

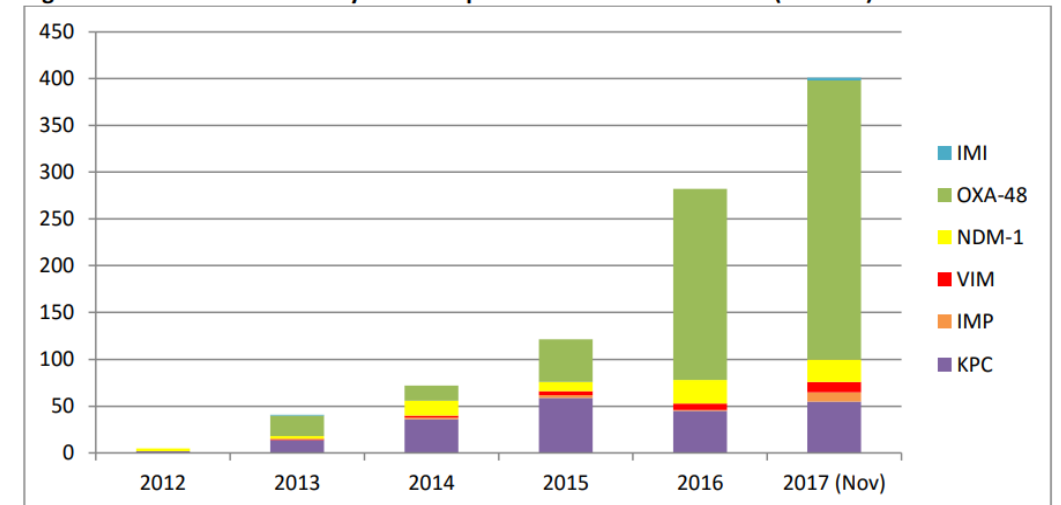
Public Health Emergency Plan to tackle CPE



CPE – Carbapenemase-producing *Enterobacteriaceae* (CPE) also described as (CRE) – bacteria which become resistant to all conventionally used antimicrobials.

- The Minister for Health has activated the Public Health Emergency Plan to address CPE in our health system.

Figure 1: Total number of newly detected patients with CPE 2012-2017 (Jan-Nov)



Source: CPE Reference Laboratory Data

CAUSES OF ANTIBIOTIC RESISTANCE



Antibiotic resistance happens when bacteria change and become resistant to the antibiotics used to treat the infections they cause.



Over-prescribing of antibiotics



Patients not finishing their treatment



Over-use of antibiotics in livestock and fish farming



Poor infection control in hospitals and clinics



Lack of hygiene and poor sanitation



Lack of new antibiotics being developed

www.who.int/drugresistance

#AntibioticResistance



World Health Organization



Ireland

One Health Report on Antimicrobial Use & Antimicrobial Resistance

Ireland's first One Health Report on Antimicrobial Use and Antimicrobial Resistance 2016

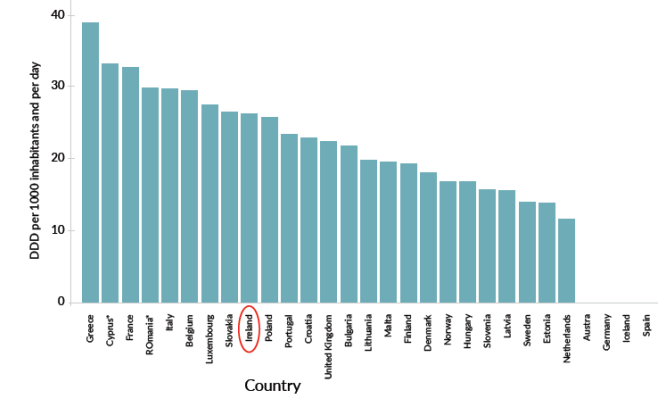
January 2019

AMU in Humans

Ireland ranked 9th highest of 25 EU/ EEA Member States for antibacterial consumption in humans in 2016.

26.1 defined daily doses (DDD) per 1,000 inhabitants per day (DID).

Consumption of Antibacterials for systemic use (ATC group J01) in the community and hospital sector in Europe, reporting year 2016



Source: TESSy, The European Surveillance System on 2018-03-06

AMU in Animals

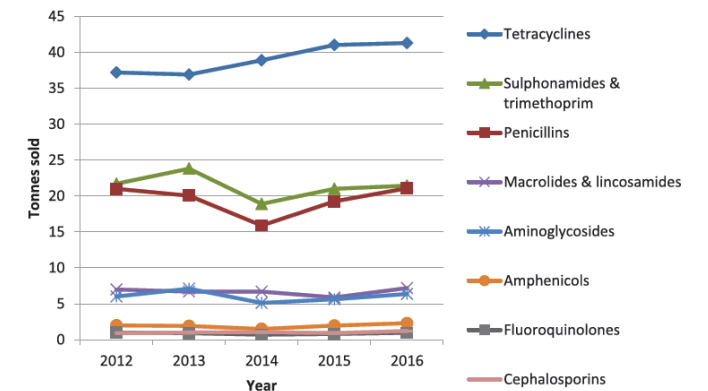
Ireland ranked 17th highest of 30 EU/ EEA member states for antimicrobial use in animals (mg/kg biomass) in 2016.

Based on sales data, most animal AMU in Ireland (66.6%) is formulated as premixes or oral remedies, presumed to be predominantly used as in-feed or in-water medication for the intensive pig and poultry sectors.

The antimicrobials most commonly sold for animal use in Ireland (by weight) are:

- Tetracyclines (39.9%)
- Sulphonamides & trimethoprim (20.7%)
- Penicillins (20.4%)

Sales (tonnes sold) of veterinary antibiotics 2012 - 2016



Source: HPRA



Ireland

One Health Report on Antimicrobial Use & Antimicrobial Resistance

Ireland's first One Health Report on Antimicrobial Use and Antimicrobial Resistance 2016

January 2019

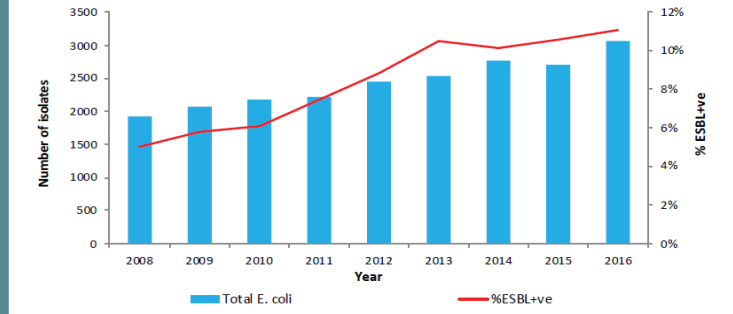
AMR in bacterial isolates from Humans

Extended spectrum beta lactamase (ESBL)-producing *E. coli*

E. coli is by far the most common causative pathogen of bloodstream infection (BSI) in Ireland, with around 3,000 cases per year. When *E. coli* acquire the capacity to produce ESBLs, this enables them to resist the activity of most beta lactam antimicrobials (e.g. penicillins, cephalosporins, monobactams). This increases dependence on carbapenems for effective treatment of infection.

In 2016, of those *E. coli* causing blood stream infection, 11% were ESBL-producing *E. coli*. This is the highest level since surveillance began. Concerns about increased risk of ESBL-producing *E. coli* may in turn drive increased carbapenem use for empiric treatment of invasive infection.

Annual total *E. coli* BSI: 2008-2016 (%ESBL producing = red line)



Source: HPSC

Indicator (non-pathogenic) Bacteria

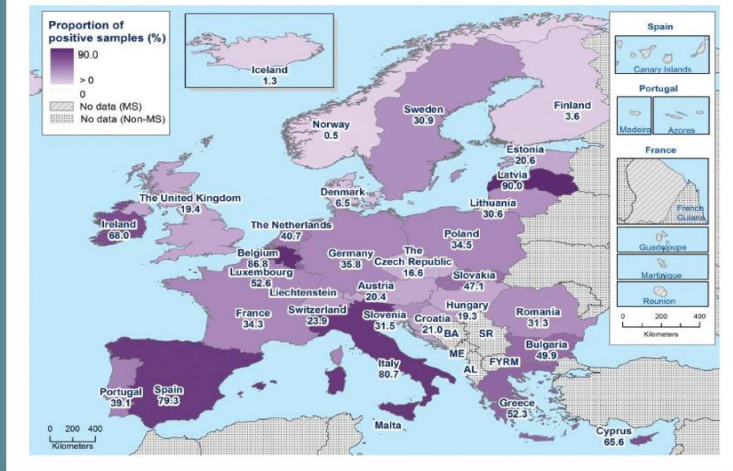
Indicator commensal *E. coli*

- More frequently resistant to antimicrobials commonly used in veterinary medicine.
- Proportion of MDR was higher in poultry compared to pig isolates.

ESBL-producing *E. coli*

- In 2016, the proportion of Irish poultry meat samples harbouring ESBL *E. coli* was above the EU average.

Presumptive ESBL *E. coli* from broilers, 2016 (specific monitoring)



Source: EFSA and ECDC, 2018. The European Union summary report on antimicrobial resistance in zoonotic and indicator bacteria from humans, animals and food in 2016. EFSA Journal 2018;16(2):5182, 270 pp. <https://doi.org/10.2903/j.efsa.2018.5182>

Antimicrobial Resistance - Humans

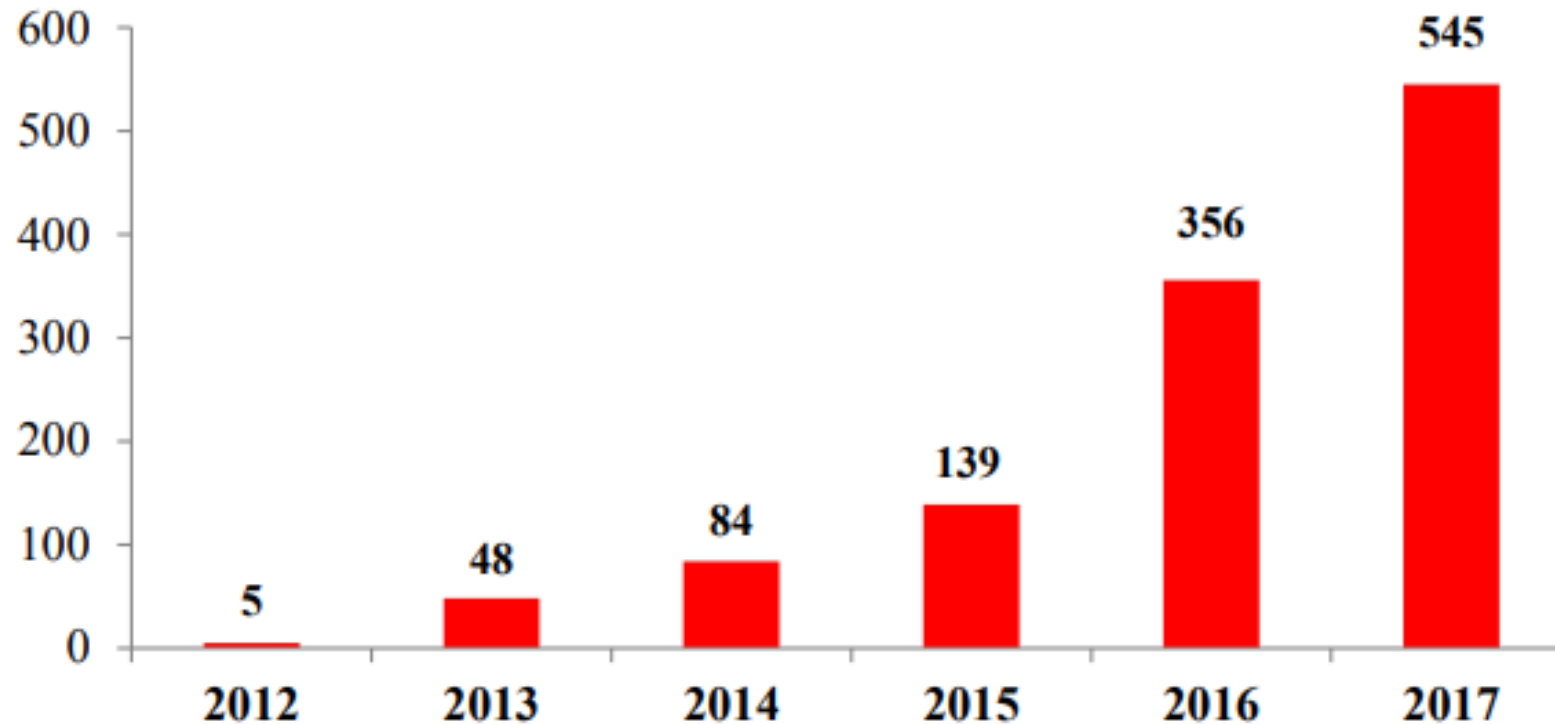


Figure 1: Carbapenemases detected in clinical isolates of Enterobacterales in Ireland Sept 2012 to Dec 2017.

Source: National Carbapenemase Producing Enterobacterales Reference Laboratory service Annual Report 2017.



Ireland

One Health Report on Antimicrobial Use & Antimicrobial Resistance

Ireland's first One Health Report on Antimicrobial Use and Antimicrobial Resistance 2016

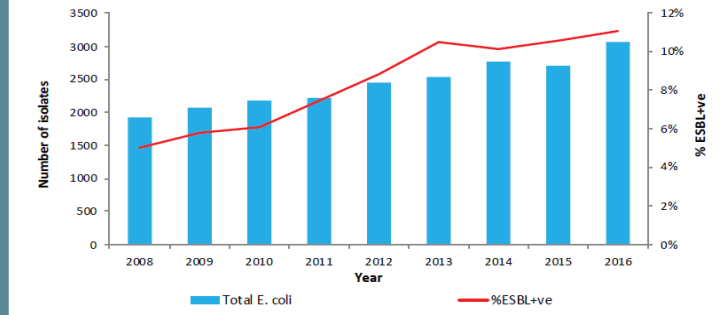
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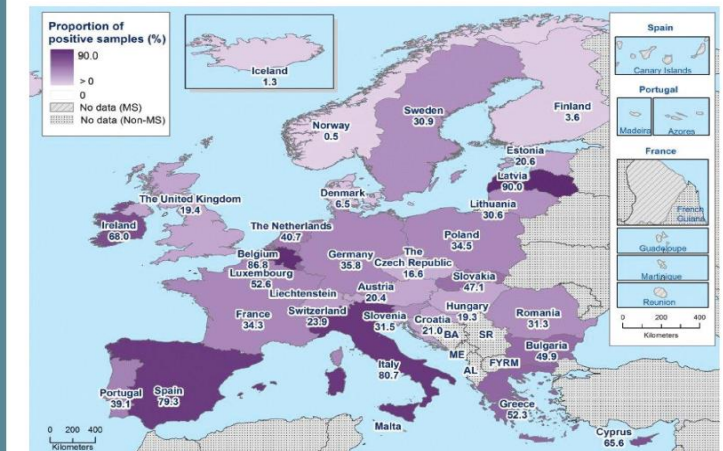
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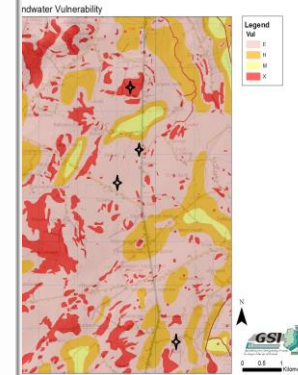
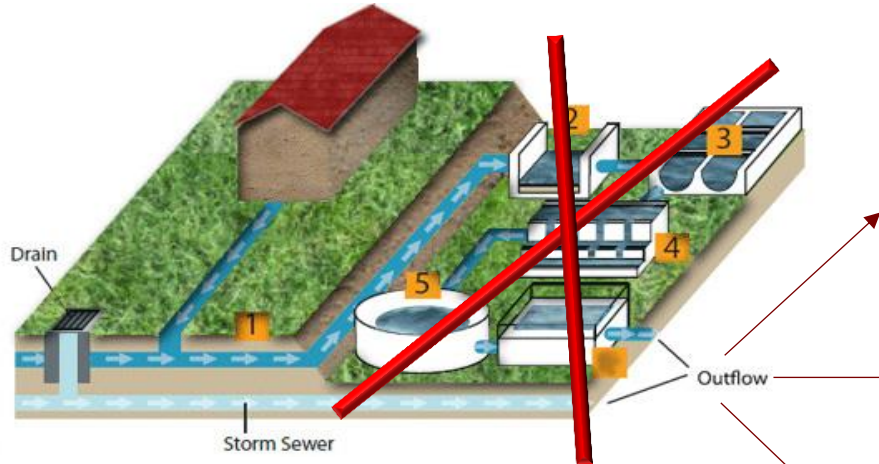
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Where Antimicrobials and Antimicrobial Resistant Bacteria Go



What we Know and What we Need to Know

- Dispensing raw sewage in the vicinity of recreational bathing areas is a major risk for transmission of antimicrobial resistant organisms.



The screenshot shows the Eurosurveillance journal interface. At the top, it features the Eurosurveillance logo with '20 years 1996-2016' and the tagline 'Europe's journal on infectious disease epidemiology, prevention and control'. Below the logo is a navigation menu with links for HOME, ARCHIVES, ABOUT US, EDITORIAL POLICY, FOR AUTHORS, FOR REVIEWERS, LINKS, and 20-YEAR. The main content area displays the article title: 'INDISTINGUISHABLE NDM-PRODUCING *ESCHERICHIA COLI* ISOLATED FROM RECREATIONAL WATERS, SEWAGE, AND A CLINICAL SPECIMEN IN IRELAND, 2016 TO 2017'. The authors listed are BM Mahon, C Brehony, E McGrath, J Killeen, M Cormican, P Hickey, S Keane, B Hanahoe, A Dolan, and D Morris. There are also navigation buttons for 'Back to Table of Contents', 'Tweet', and 'Next'.



Source: EPA, Urban Waste Water Treatment in 2017

What we Know and What we Need to Know

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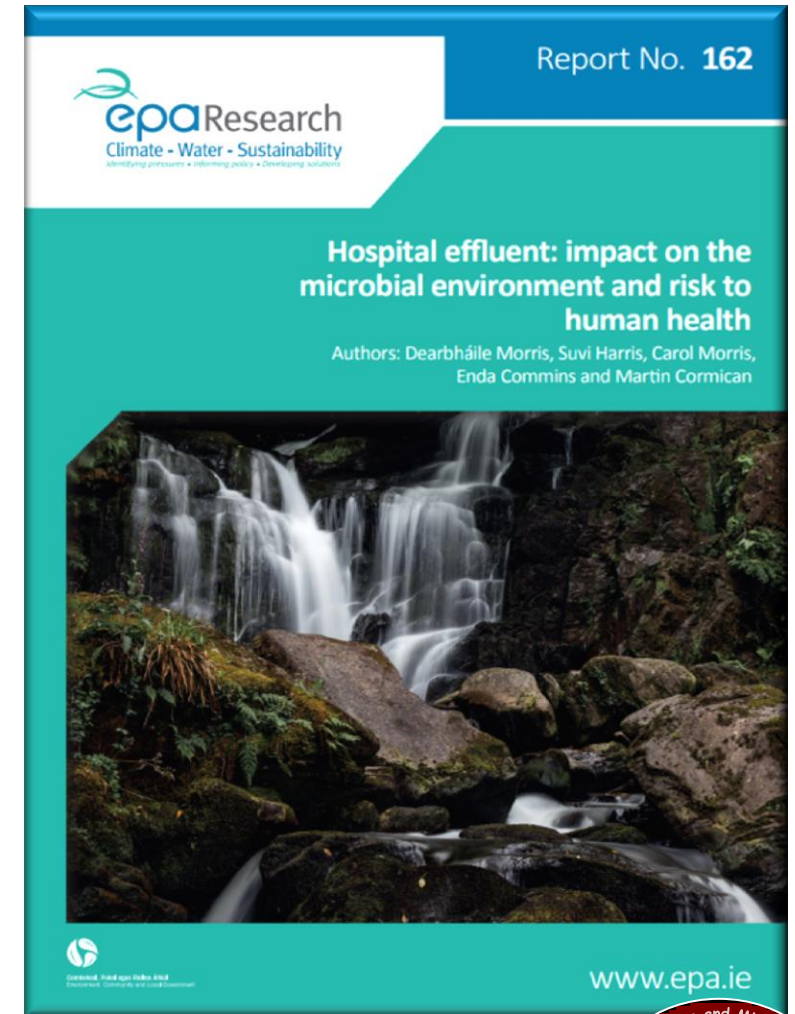
- Do current drinking and bathing water quality regulations protect public health?



Source: EPA, Urban Waste Water Treatment in 2017

What we Know and What we Need to Know

- Current waste water treatment processes do not remove all antimicrobial resistant organisms (AROs)
- What level and type of waste water treatment is effective for the removal of AROs?
- Integrated constructed wetlands are a natural alternative to conventional wastewater treatment processes but are they effective at removing AROs?



What we Know and What we Need to Know

- Land spreading of organic wastes (animal manure and sewage sludge) is an important part of the circular economy
- Studies have shown land spreading of manures increased the proliferation of ARO and abundance of antimicrobial resistance genes in soils^{1,2}.

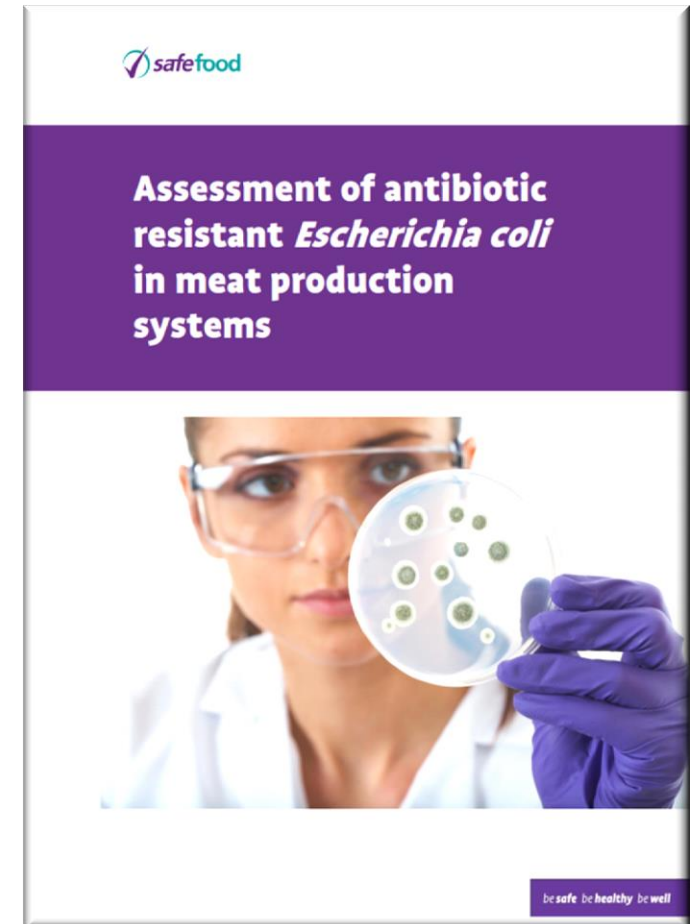


1. Udikovic-Kolic, N., Wichmann, F., Broderick, N. A. & Handelsman, J. Bloom of resident antibiotic-resistant bacteria in soil following manure fertilization. *Proc. Natl. Acad. Sci.* **111**, 15202–15207 (2014).

2. Muurinen, J. *et al.* Influence of Manure Application on the Environmental Resistome Under Finnish Agricultural Practice with Restricted Antibiotic Use. *Environ. Sci. Technol.* acs.est.7b00551 (2017). doi:10.1021/acs.est.7b00551

What we Know and What we Need to Know

- Retail meats: 99% of chicken meats, 33% of pork products and 14% of beef products contaminated with AROs¹
- What is the level of AROs in manures from different sources?
- What treatment is effective at removing them?
- What impact does application have on soil?



¹https://www.safefood.eu/SafeFood/media/SafeFoodLibrary/Documents/Publications/Research%20Reports/Assessment-of-antibiotic-resistant-Escherichia-coli-in-meat-production-systems_1.pdf



ONE HEALTH



AREST



AREST: 2018-2021

Antimicrobial Resistance and the Environment –
Sources, persistence, Transmission and risk management (2017-HW-LS-1)

Web: <http://www.nuigalway.ie/medicine-nursing-and-health-sciences/medicine/disciplines/bacteriology/research/arest/>



<https://www.researchgate.net/project/AREST-Antimicrobial-Resistance-and-the-Environment-Sources-Persistence-Transmission-and-Risk-Management>



O'Connell
NUI Galway



Technical University of Denmark



Cork
County Council
Comhairle Contae Chorcaí



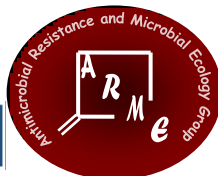
Comhairle Chontae na Gaillimhe
Galway County Council



Ryan
Institute



Centre for Health from
Environment
Ionad Sláinte ó Chomhshaoil





AREST: 2018-2021

Antimicrobial Resistance and the Environment – Sources, persistence, Transmission and risk management

Project Team



Dr. Dearbháile Morris
 Dr. John Cullinan
 Dr. Louise O'Connor
 Dr. Mark Healy
 Prof. Xinmin Zhan
 Dr. Liam Morrison
 Ms. Suzanne Nolan
 Prof. Martin Cormican*
 Dr. Carlos Chique
 Ms. Brigid hooban



Dr. Geraldine Duffy
 Dr. Kaye Burgess
 Dr. Fiona Brennan



Dr. Deirdre Prendergast



Feidhmeannacht na Seirbhíse Sláinte
Health Service Executive

Prof. Martin Cormican*



Dr. Fiona Walsh



Prof. Enda Cummins
 Prof. Fionola Leonard
 Prof. Seamus Fanning
 Prof. Barry McMahon



Comhairle Chontae na Gaillimhe
Galway County Council

Dr. Rita Gately



Cork County Council
Comhairle Contae Chorcaí

Dr. Dan Crowley



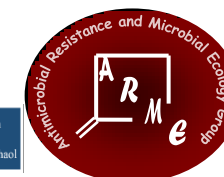
Technical University of Denmark



Cork County Council
Comhairle Contae Chorcaí



Comhairle Chontae na Gaillimhe
Galway County Council





AREST: 2018-2021

Antimicrobial Resistance and the Environment – Sources, persistence, Transmission and risk management

The AREST project will:

- Map hotspots of drivers of AMR in selected local authority areas:
Galway City Council, Galway County Council, Fingal County Council and Cork County Council.
- Assess the relative contributions of various sectors (healthcare, agriculture) to ARO in the environment.
- Assess efficiencies of treatment processes for removal of ARO from drinking water, wastewater and manure.
- Develop a risk ranking protocol to assess the relative contribution of various sectors on the sources and levels of AROs in the environment.



O'É Gaillimh
NUI Galway



Technical University of Denmark



Cork
County Council
Comhairle Contae Chorcaí



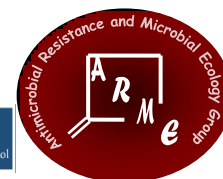
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Galway County Council



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Galway City & County Council



Fingal County Council



Cork County Council



Legend

ED Livestock Index

Index Value¹



Healthcare Facilities

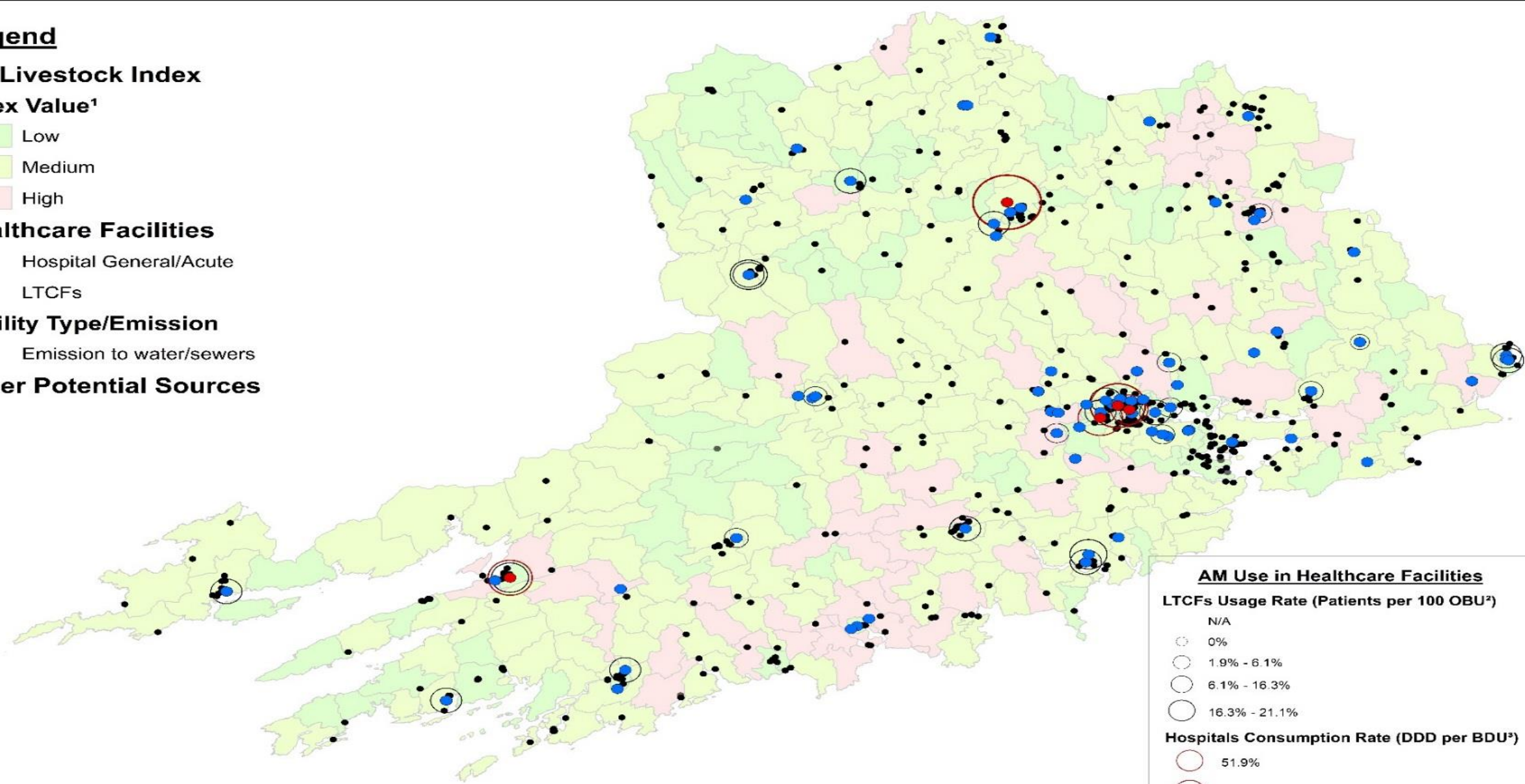
- Hospital General/Acute
- LTCFs

Facility Type/Emission

- Emission to water/sewers

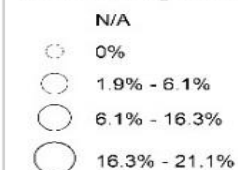
Other Potential Sources

-

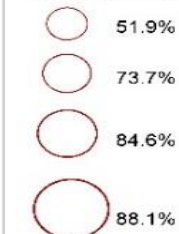


AM Use in Healthcare Facilities

LTCFs Usage Rate (Patients per 100 OBU²)



Hospitals Consumption Rate (DDD per BDU³)



¹ Based on CSO agricultural census estimates including (ED) cattle and sheep density and poultry/pig presence-absence data. The index weighting is based on estimated livestock category antibiotic applications.

² Usage per 100 occupied bed units

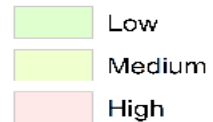
³ Consumption at daily defined dose per 100 bed days used



Legend

ED Livestock Index (2010)

Index Value¹



Healthcare Facilities

- Hospital General/Acute
- LTCFs

UWWTP

Treatment Type

- Raw Sewage Discharge
- Primary, Secondary or Tertiary

UWW Discharge Location

- UWW Discharge Location

Section 4 Discharges

Activity/Type

- Healthcare
- Animal/Food Production

Landfill Sites



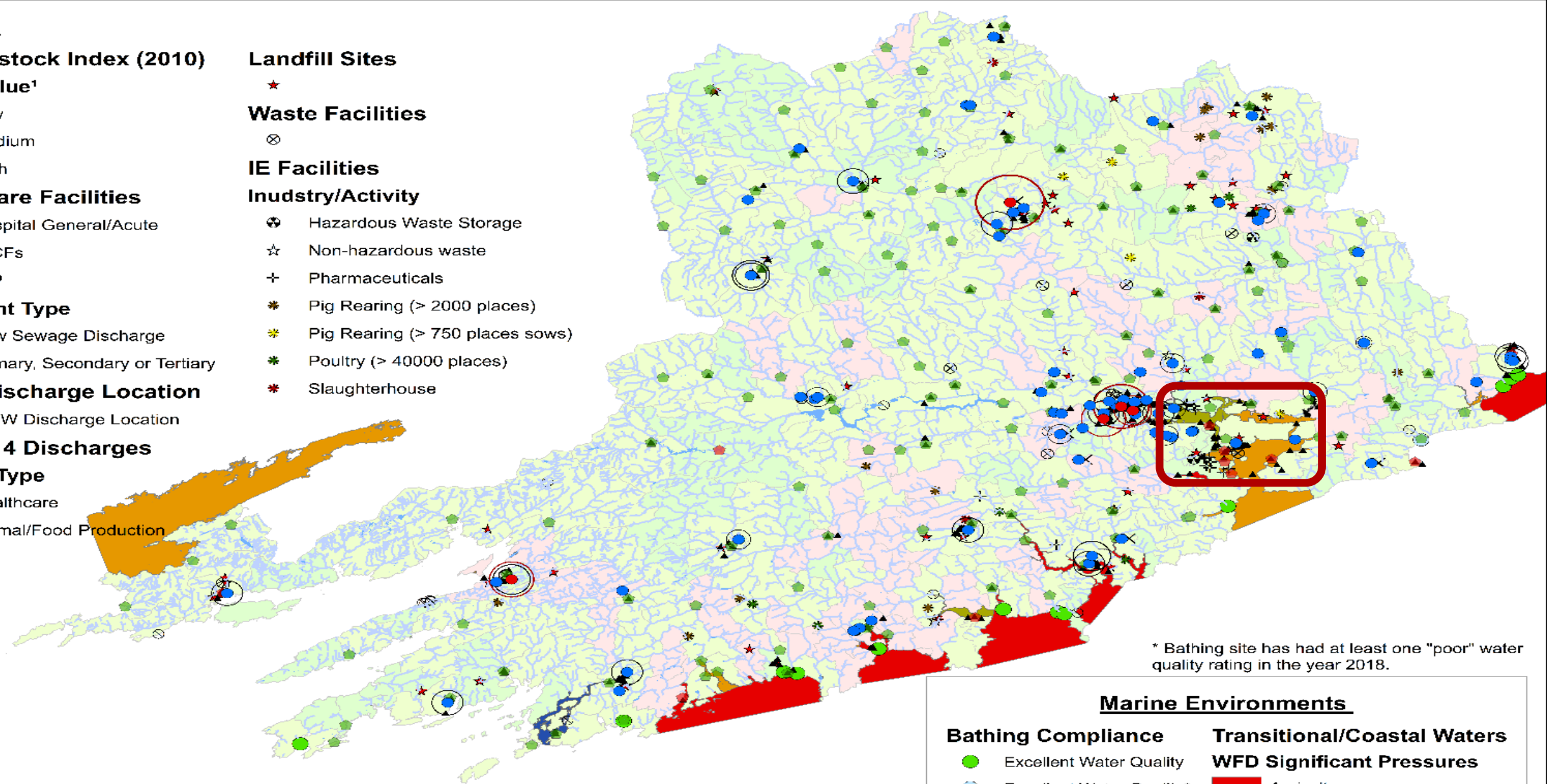
Waste Facilities



IE Facilities

Industry/Activity

- Hazardous Waste Storage
- Non-hazardous waste
- Pharmaceuticals
- Pig Rearing (> 2000 places)
- Pig Rearing (> 750 places sows)
- Poultry (> 40000 places)
- Slaughterhouse



* Bathing site has had at least one "poor" water quality rating in the year 2018.

Marine Environments

Bathing Compliance

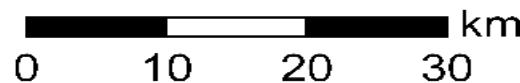
- Excellent Water Quality
- Excellent Water Quality*

Transitional/Coastal Waters

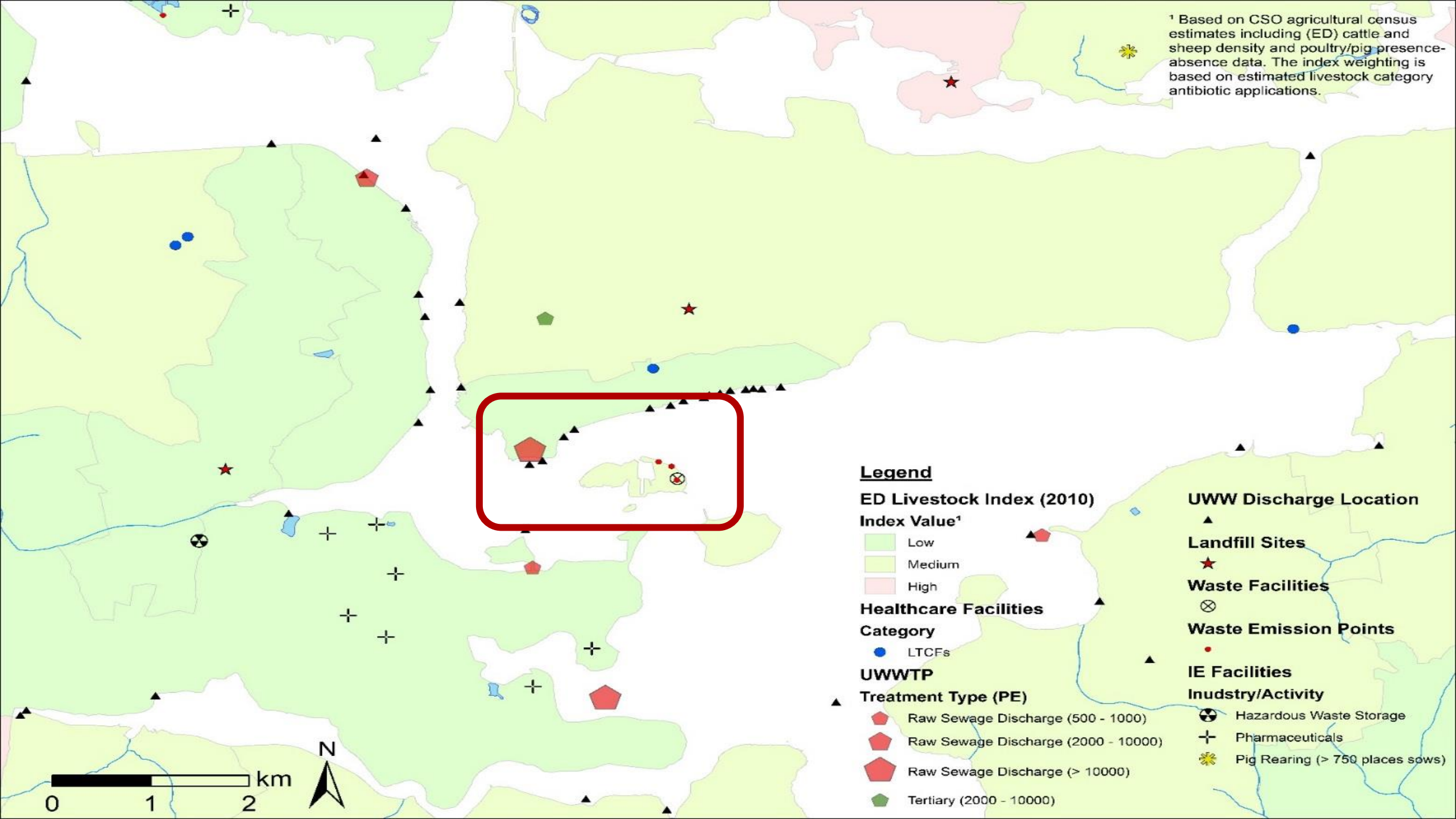
WFD Significant Pressures

- Agriculture
- Anthropogenic Pressures
- Urban Run-off
- Urban Waste Water
- Industry

¹ Based on CSO agricultural census estimates including (ED) cattle and sheep density and poultry/pig presence-absence data. The index weighting is based on estimated livestock category antibiotic applications.



¹ Based on CSO agricultural census estimates including (ED) cattle and sheep density and poultry/pig presence-absence data. The index weighting is based on estimated livestock category antibiotic applications.



Legend

ED Livestock Index (2010)

- Index Value¹
- Low
 - Medium
 - High

Healthcare Facilities

- Category
- LTCFs

UWWTP

Treatment Type (PE)

- Raw Sewage Discharge (500 - 1000)
- Raw Sewage Discharge (2000 - 10000)
- Raw Sewage Discharge (> 10000)
- Tertiary (2000 - 10000)

UWW Discharge Location

Landfill Sites

Waste Facilities

Waste Emission Points

IE Facilities

Industry/Activity

- Hazardous Waste Storage
- Pharmaceuticals
- Pig Rearing (> 750 places sows)





Legend

UWWTP

Treatment Type (PE)

Raw Sewage Discharge (> 10000)

UWW Discharge Location

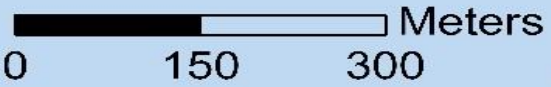
▲

Waste Emission Points

●

Waste Facilities

⊗



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Point Prevalence and Longitudinal Surveys:

- Hospital effluent
- Long term care facility effluent
- Agricultural waste – manures (pig, poultry, cattle)
- Wildlife
- Companion animals
- Recreational waters – freshwaters and seawaters
- Waste water treatment plants (influent & effluent)
- Integrated constructed wetlands (influent & effluent)
- Surface and ground waters
- Septic tanks
- Boreholes/private wells.

Legend

UWWTP

Treatment Type (PE)



Raw Sewage Discharge (> 10000)

UWW Discharge Location



Waste Emission Points



Waste Facilities



Service Layer Credits: Sources: Esri, HERE, Garmin, USGS, Intermap, INCREMENT P, NRCan, Esri Japan, METI, Esri China (Hong Kong), Esri Korea, Esri (Thailand), NGCC, © OpenStreetMap contributors, and the GIS User Community

0 150 300 Meters





AREST: 2018-2021

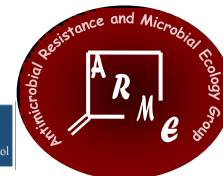
Antimicrobial Resistance and the Environment – Sources, persistence, Transmission and risk management

The AREST project will:

- Generate national level data on the key sources, hot spots and drivers of AMR in the environment from various sectors which will inform priority areas for action.
- Provide evidence of the extent of contamination of the environment with antimicrobial resistant organisms to support policy decisions and engage with health, agriculture and the local authority sectors on AMR.
- Produce engaging visual representations of data that will strongly support wider communication with the public and policy makers.
- Embed the “One Health” concept and build the capacity of Ireland’s research community to support Ireland’s National Action Plan on AMR.



Technical University of Denmark



PIER: 2019-2022

Public health Impact of Exposure to antimicrobial Resistance in coastal waters (2018-HW-LS-2)

Web: <http://www.nuigalway.ie/medicine-nursing-and-health-sciences/medicine/disciplines/bacteriology/research/pier/>

PIER: 2019-2022

Public health Impact of Exposure to antimicrobial Resistance in coastal waters (2018-HW-LS-2)

Project Team



Dr. Dearbháile Morris
Dr. Liam Burke
Dr. Louise O'Connor
Dr. Sinéad Duane
Dr. Easkey Britton
Prof. Martin Cormican



Feidhmeannacht na Seirbhíse Sláinte
Health Service Executive

Dr. Diarmuid O'Donovan
Dr. Áine McNamara
Dr. Regina Kiernan
Dr. Katharine Harkin

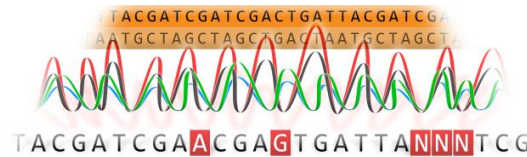
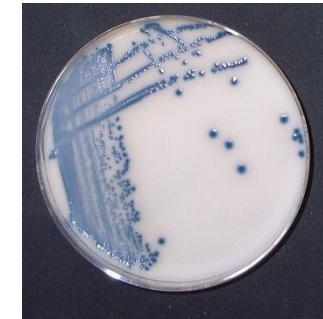


Prof. William Gaze
Dr. Anne Leonard

Public health Impact of Exposure to antimicrobial Resistance in coastal waters (2018-HW-LS-2)



Photo by [Guy Kawasaki](#) on [Unsplash](#)



DESIGN: 2019-2020

Detection of Environmental Sources of Infectious diseases in Groundwater Networks

DESIGN: 2019-2020

Detection of Environmental Sources of Infectious diseases in Groundwater Networks

Project Team



Dr. Jean O'Dwyer



Dr. Dearbháile Morris
Dr. Liam Burke

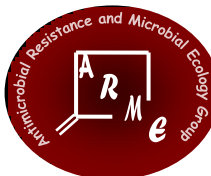


Dr. Paul Hynds



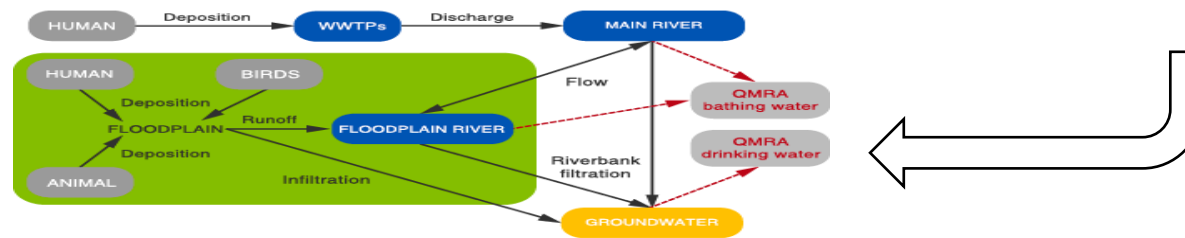
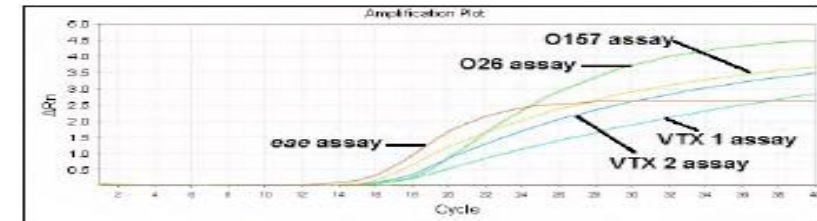
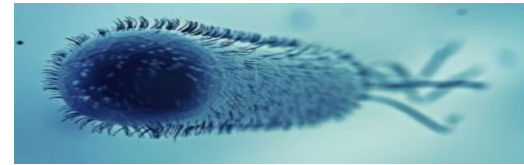
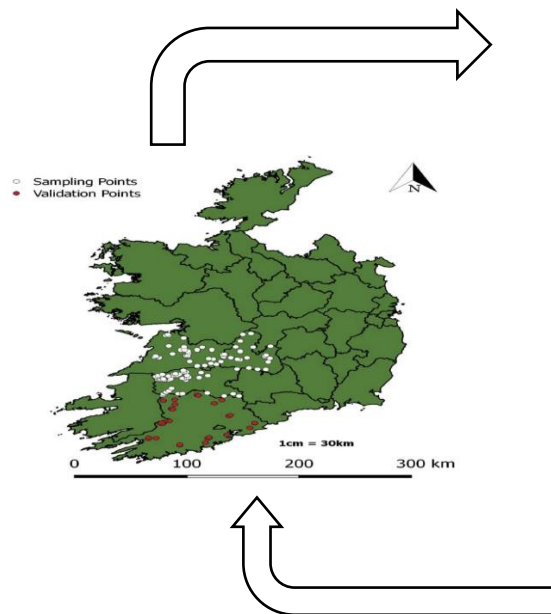
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LIMERICK
OILScoil LUIMNIGH

Dr. Michael Ryan



DESIGN: 2019-2020

Detection of Environmental Sources of Infectious diseases in Groundwater Networks



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Web: <http://www.nuigalway.ie/medicine-nursing-and-health-sciences/medicine/disciplines/bacteriology/research/>



ONE HEALTH
EJP ASM 2019



DUBLIN, MAY 22ND -24TH 2019
TEAGASC CONFERENCE CENTRE, ASHTOWN, DUBLIN
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