

# Changing Practices to Reduce Antibiotic Resistance



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University of Arizona College of Agriculture and Life Sciences and  
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Addressing Antibiotic Resistance Hotspots Session, APHL Annual Meeting  
June 2, 2018

# Antibiotic Resistance: the Quintessential One Health Issue

## Multiple disciplines

Clinical, Environmental  
Policy  
Statisticians

**Working** locally, nationally,  
and globally

**To attain health** for people,  
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What is **known**? What is **not known**?

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# Antibiotic Resistant Bacteria

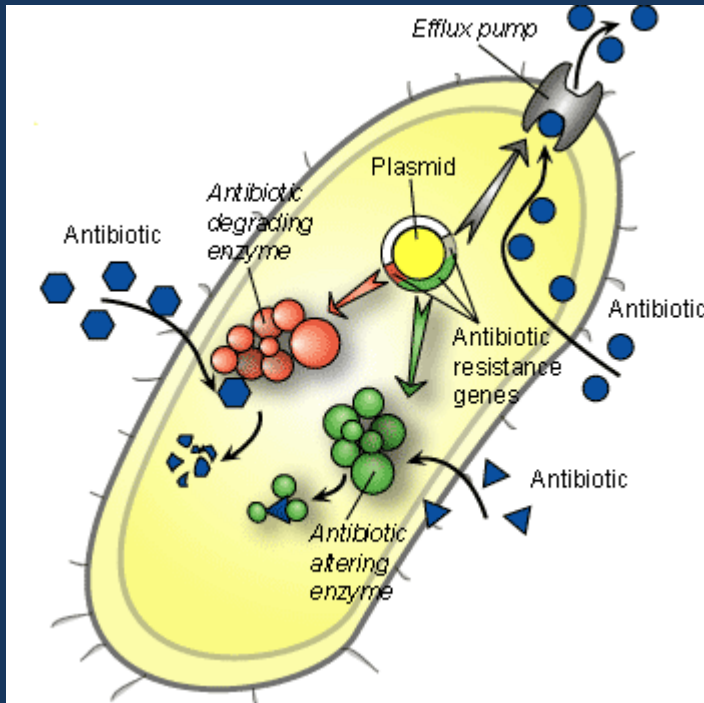
The ability of a bacterium to **prevent an antibiotic from adversely affecting** that isolate, strain, or group.

**Horizontal Gene Transfer** confers antibiotic resistance in response to selective pressure

Clinical settings

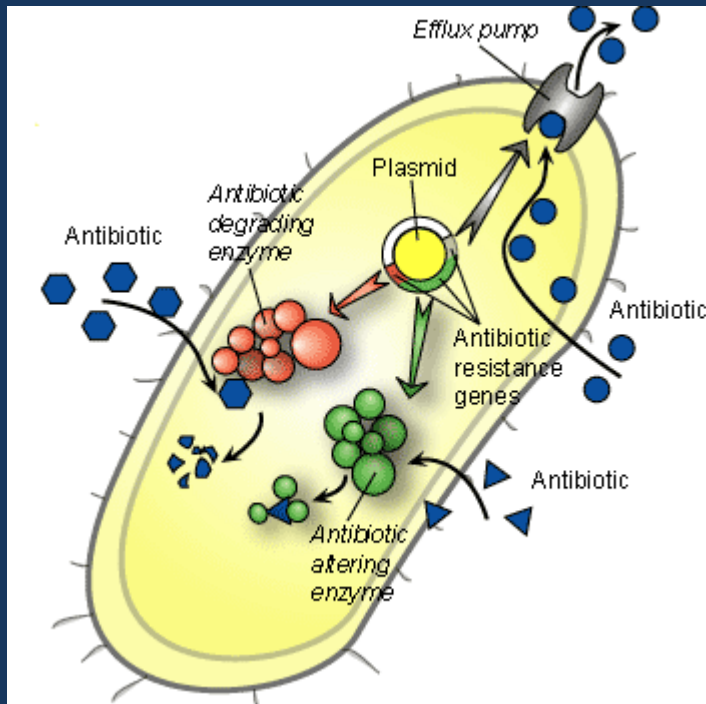
High antibiotic dosages

**Resistance – anthropogenic?**



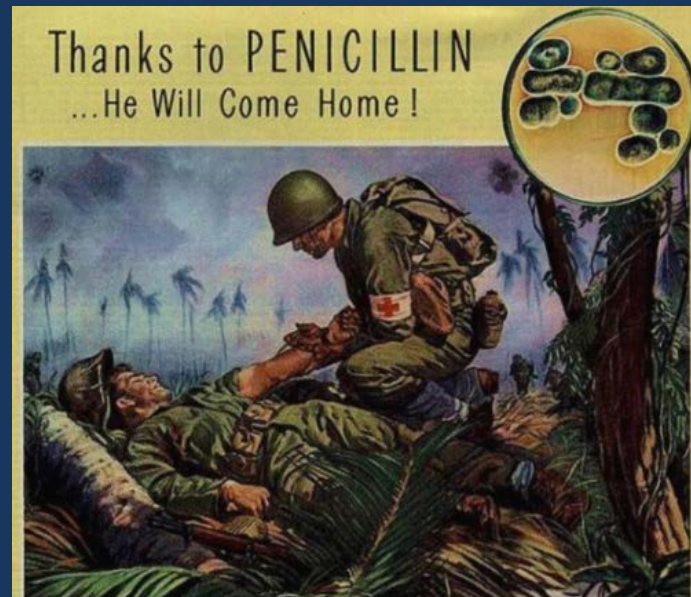
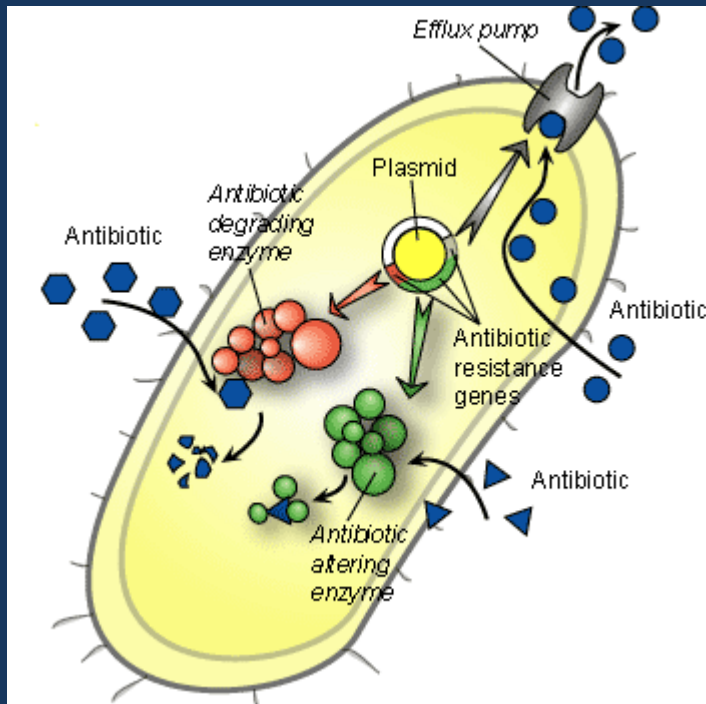
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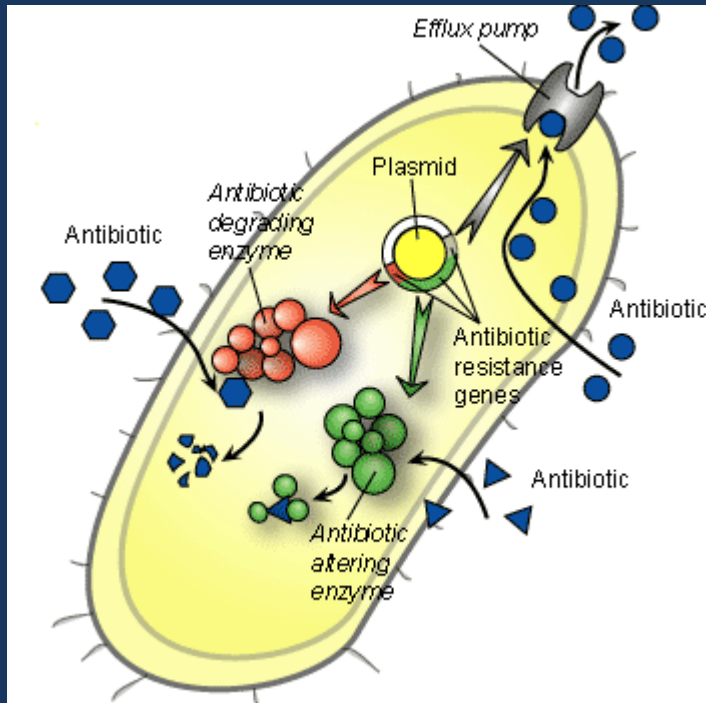
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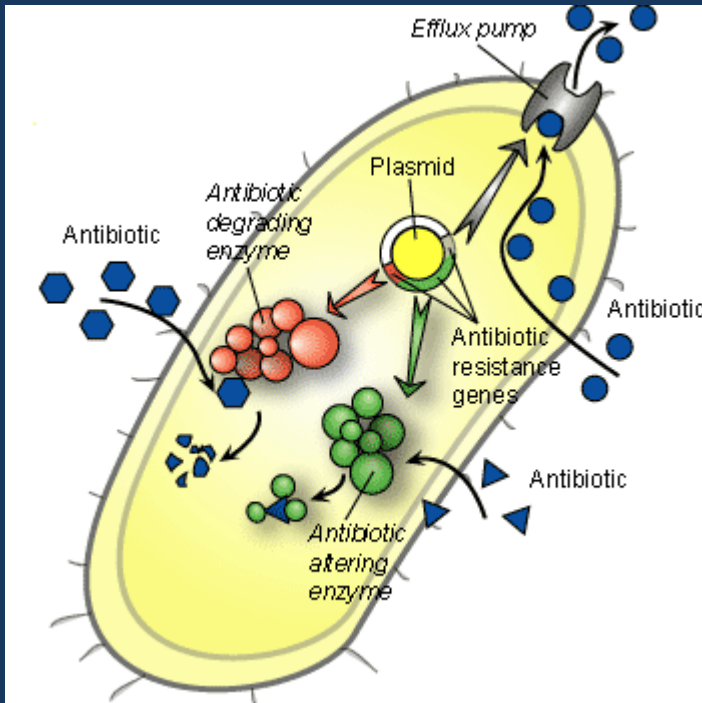


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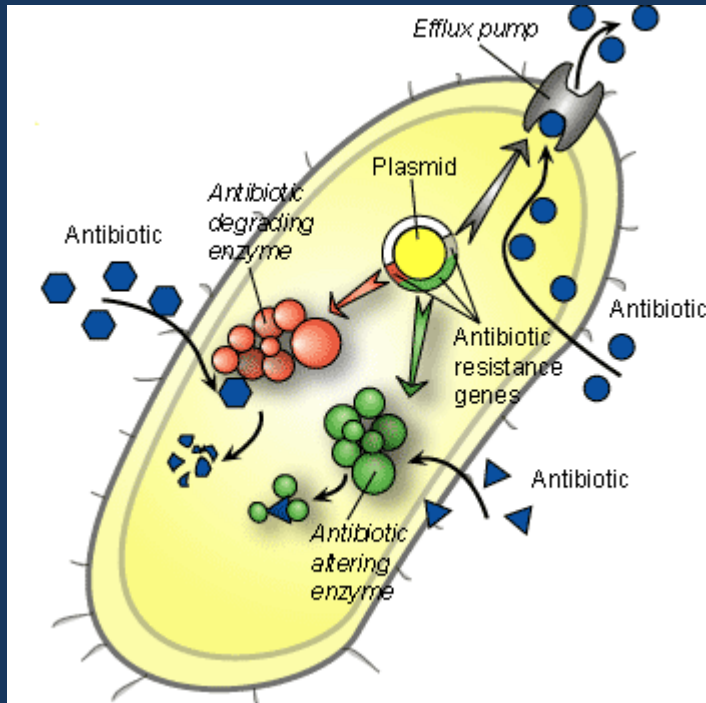
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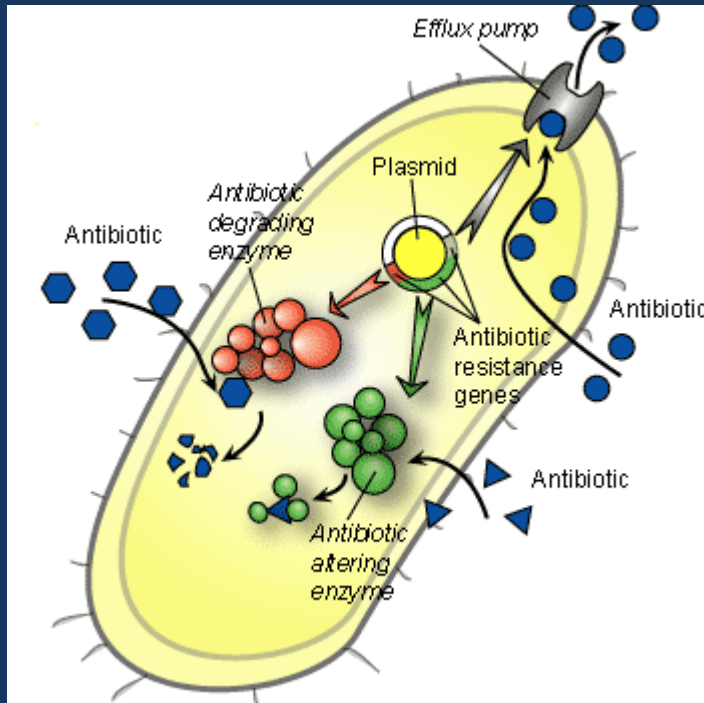
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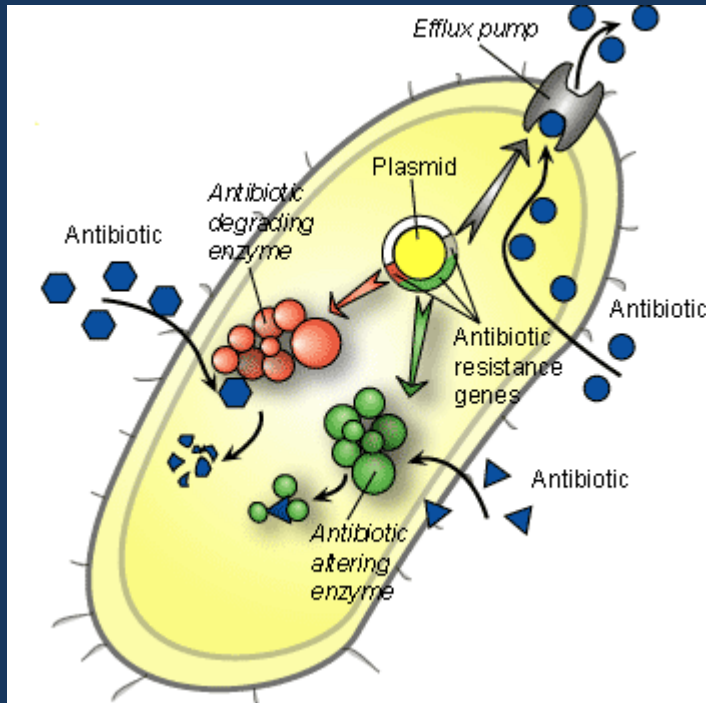
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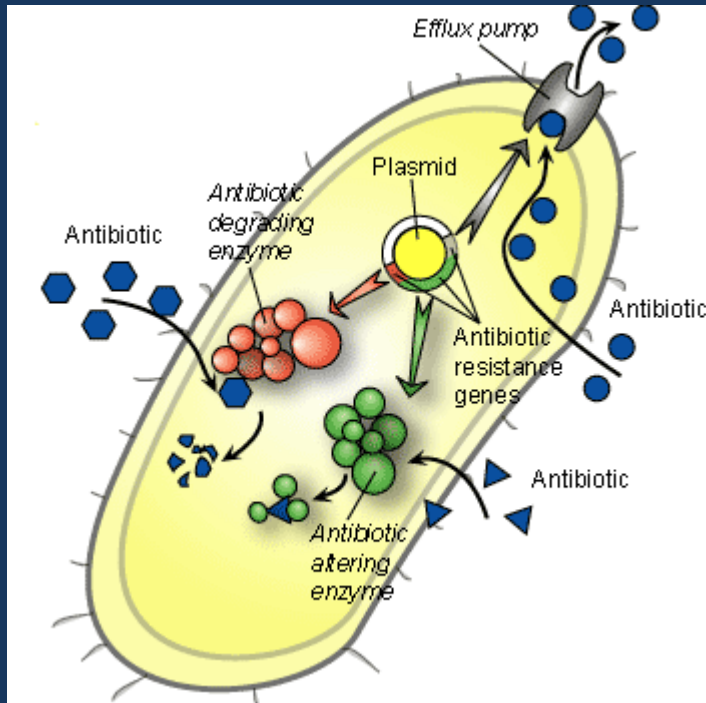
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**Environment has a role in  
emergence and spread**

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Proposed that this is a “key source of resistance to the environment”



# Does Recycled Municipal Wastewater Induce Antibiotic Resistance?





# Gilbert Riparian Preserve

Created in 1986, seven  
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Control site: agricultural  
irrigation retention pond

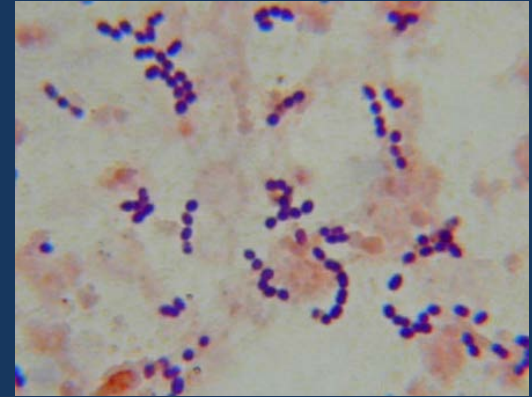


# Field Sampling over Two Years

*Enterococcus* spp.

GI tracts of humans and animals;  
environmental persistence

Great capacity for **gene transfer**



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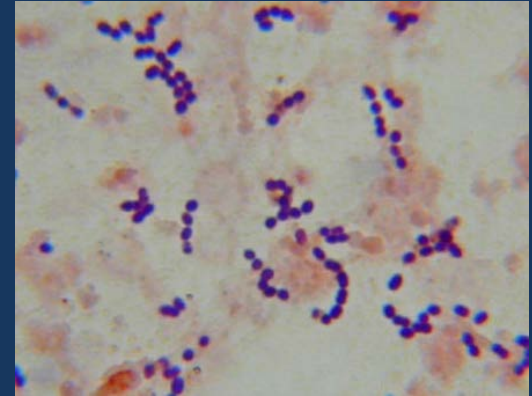
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Emergence of multiple drug-  
resistant strains in clinical settings

**Ideal bacterial group** for  
investigating the ecology of  
resistance development





# High Level Antibiotic Resistance

Antimicrobial	% Isolates with High-Level Resistance	
	Groundwater	Wastewater
Tigecycline	14.3	6.1
Erythromycin	42.8	21.2
Tetracycline	21.4	0.0
Ciprofloxacin	57.1	24.2
Chloramphenicol	7.1	0.0
Penicillin	14.3	6.1
Daptomycin	57.1	51.5
Vancomycin	7.1	3.0
Streptomycin	0.0	0.0
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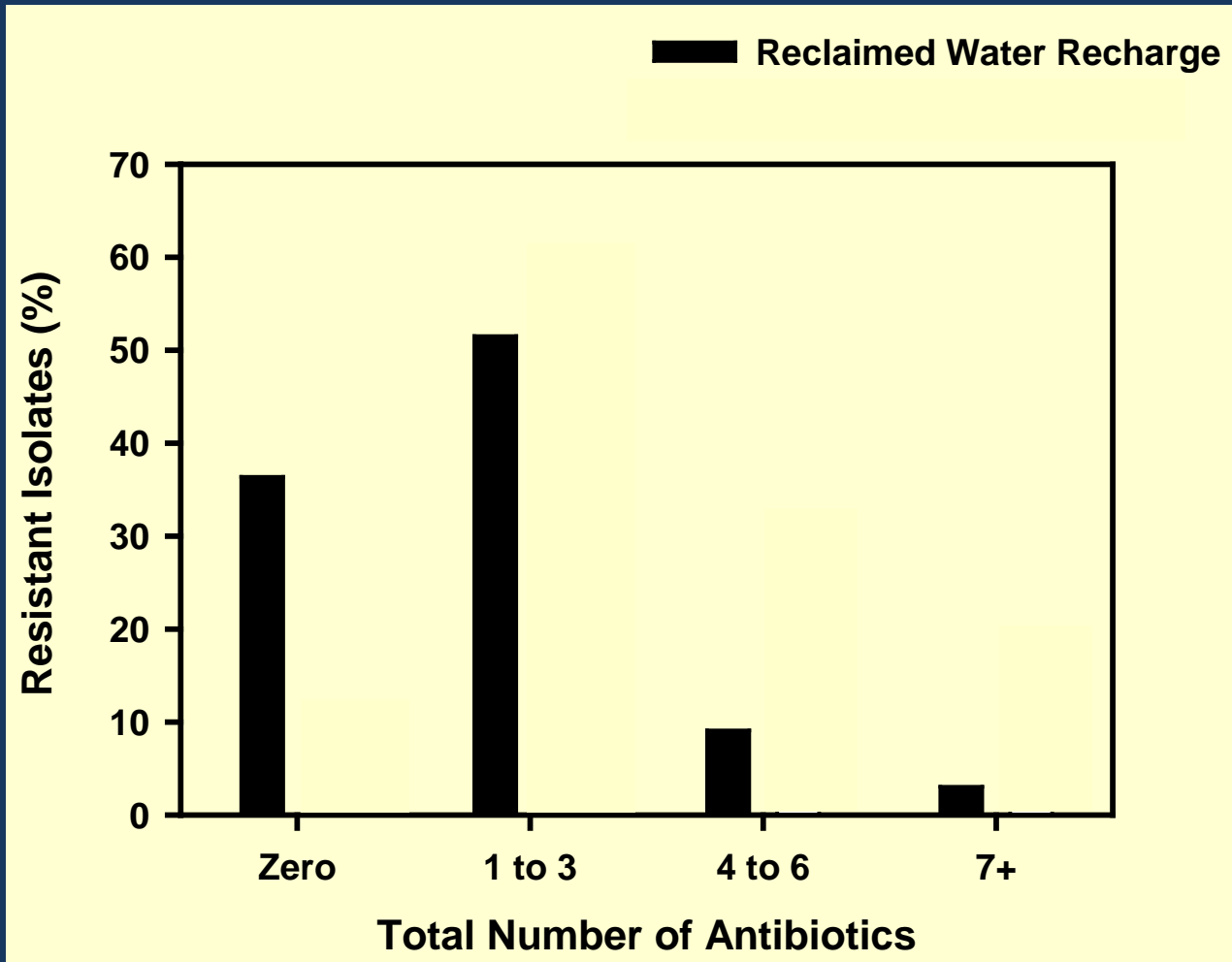
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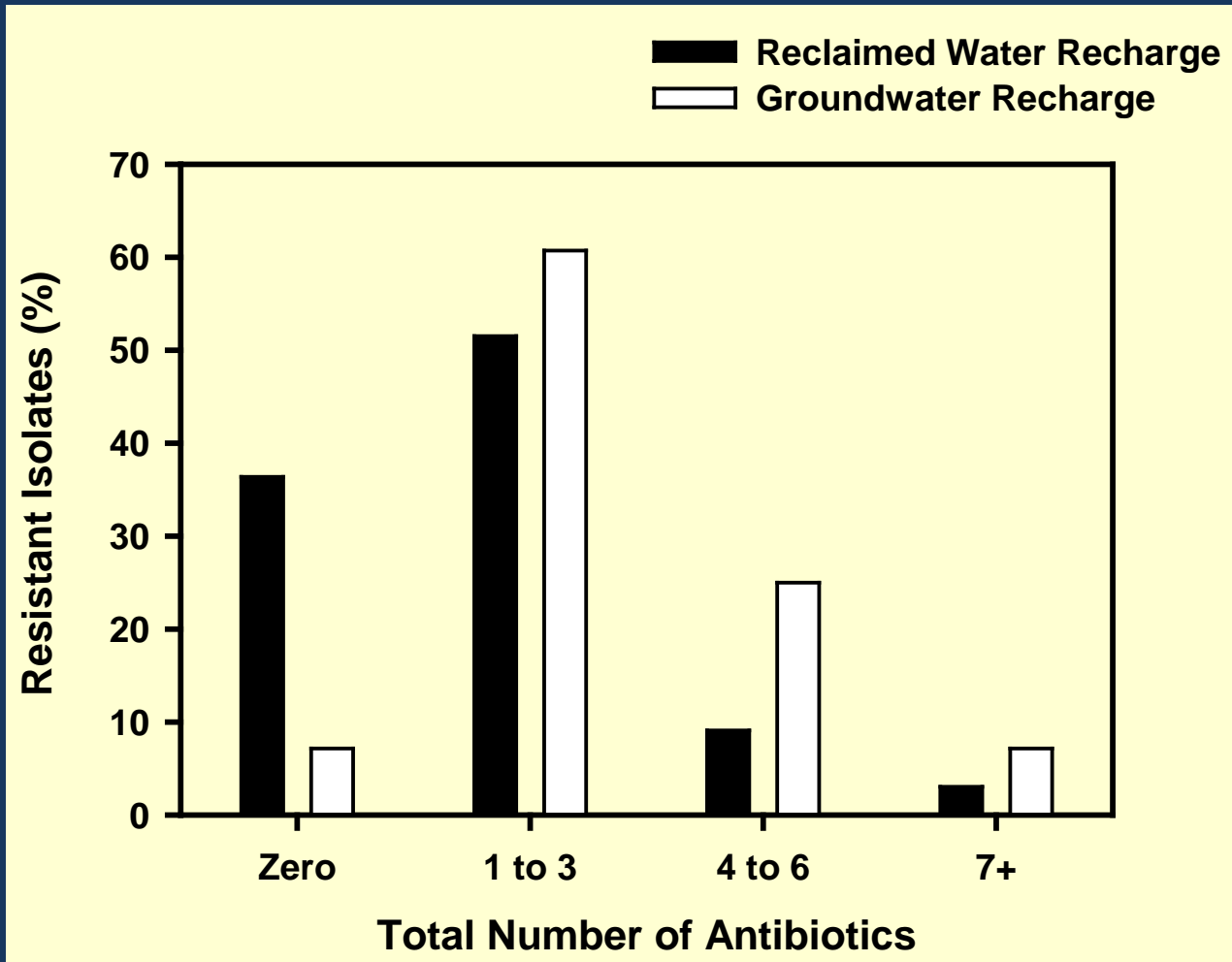
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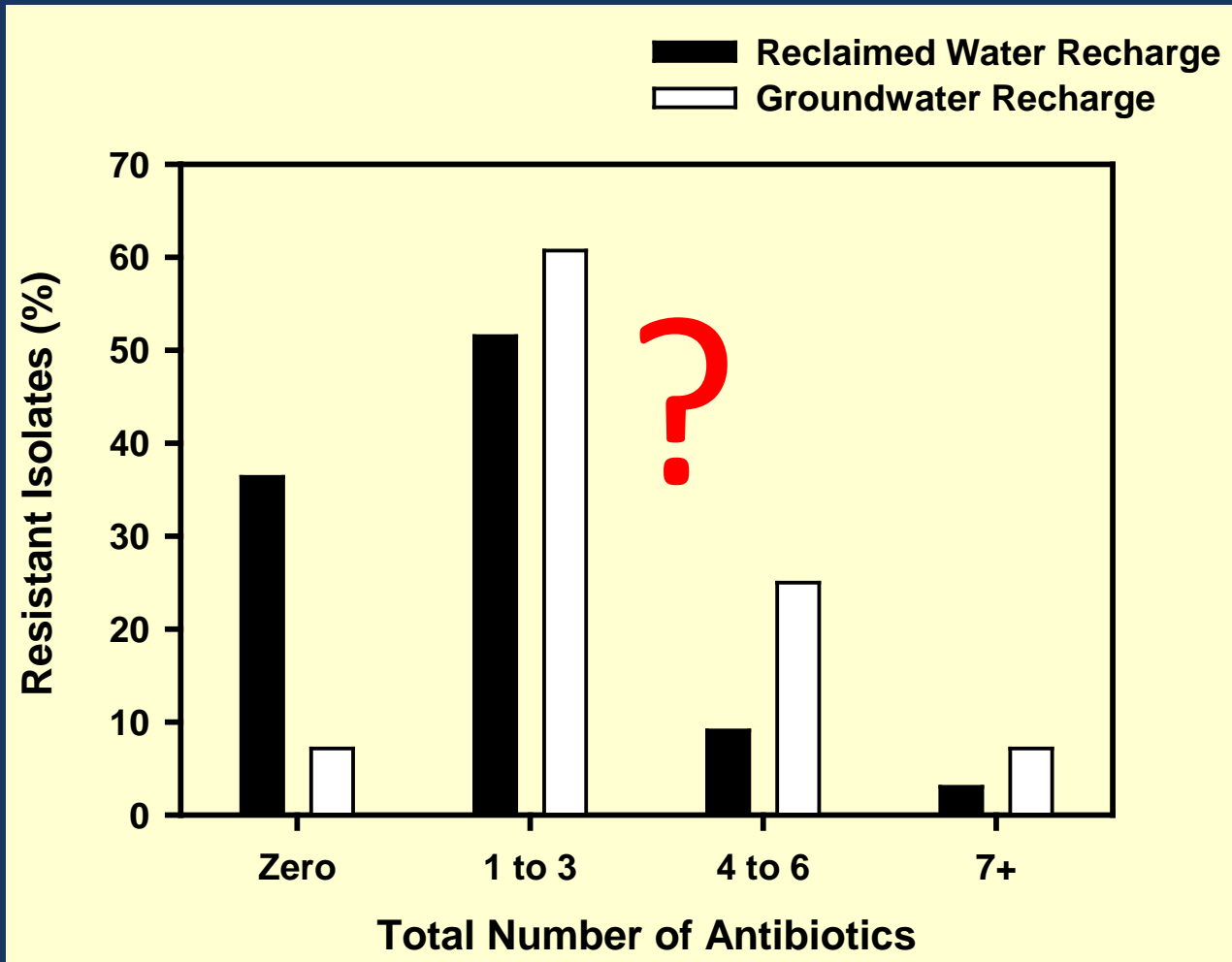
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Antibiotic resistance:

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Environmental complexity – not a  
direct cause/effect



# Pollution via Antibiotic Resistance Genes



Science of The Total Environment

Volume 447, 1 March 2013, Pages 345-360



Review

## Urban wastewater treatment plants as hotspots for antibiotic resistant bacteria and genes spread into the environment: A review

L. Rizzo <sup>a</sup>  , C. Manaia <sup>b</sup>, C. Merlin <sup>c</sup>, T. Schwartz <sup>d</sup>, C. Dagot <sup>e</sup>, M.C. Ploy <sup>f</sup>, I. Michael <sup>g</sup>, D. Fatta-Kassinos <sup>g</sup>

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Urban waste  
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**Review**

## Antibiotic resistance genes as an emerging environmental contaminant

Haley Sanderson,<sup>a</sup> Colin Fricker,<sup>a</sup> R. Stephen Brown,<sup>a</sup> Anna Majury,<sup>ab</sup>  
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ELSEVIER

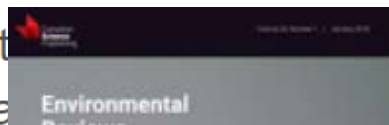
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**ENVIRONMENTAL**  
Science & Technology

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## Cell-free DNA: A Neglected Source for Antibiotic Resistance Genes Spreading from WWTPs

Yan Zhang<sup>†</sup>, Aolin Li<sup>‡</sup>, Tianjiao Dai<sup>§</sup>, Feifei Li<sup>||</sup>, Hui Xie<sup>‡</sup>, Lujun Chen<sup>\*†‡</sup>, and Donghui Wen<sup>\*§</sup> 

<sup>†</sup> Jiangsu Key Laboratory of Anaerobic Biotechnology, School of Environment and Civil Engineering, Jiangnan University, Wuxi 214122, China

<sup>‡</sup> School of Environment, Tsinghua University, Beijing 100084, China

<sup>§</sup> College of Environmental Sciences and Engineering, Peking University, Beijing 100871, China

<sup>||</sup> School of Water Resource and Environment, China University of Geosciences, Beijing, 100083, China

<sup>‡</sup> Zhejiang Provincial Key Laboratory of Water Science and Technology, Department of Environmental Technology and Ecology, Yangtze Delta Region Institute of Tsinghua University, Zhejiang Jiaxing 314050, China



# Antibiotic Resistance Genes and “Cell-Free DNA”

Water Environment  
Research Foundation Study  
(2014-2017) by Gerrity,  
Rock, and McLain



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Water Environment  
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- As much as 50% of total DNA was “free DNA”
- Quantified intact resistance genes by PCR, qPCR





# What is KNOWN and What is UNKNOWN about Antibiotic Resistance in Environmental Bacteria?

It is ancient – studies need control sites



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Culturing studies do not show cause-and-effect



## Case Study #2: Antibiotics in Animal Feed

**USA Growth promotion** – 34.3 million pounds in 2015 (7.7 million pounds for humans) – since 1940s



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EU – began weaning in 1997; ‘judicious use rules’ in 2006

Appropriate drug

Effective drug

At right dose

Correct length of time



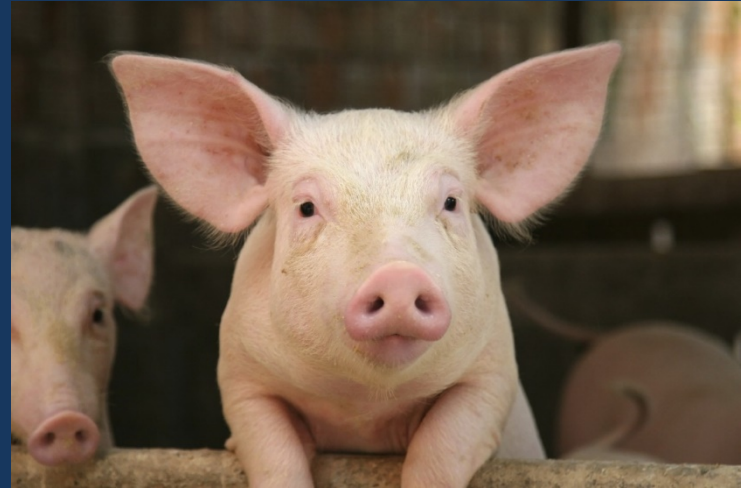


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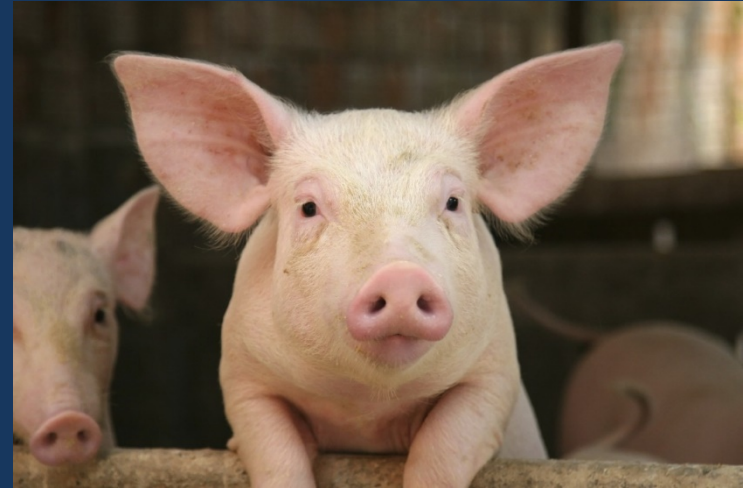
USA – FDA established rules aligning with EU in 2018

Aimed at controlling antibiotic resistant bacteria moving off farms



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\*DANMAP (2001) Danish Veterinary Laboratory, Copenhagen

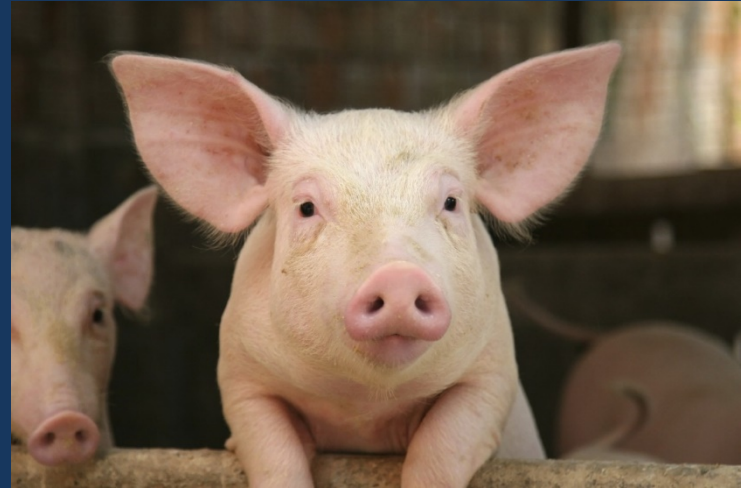


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**Increase in human *Salmonella* infection and resistance**



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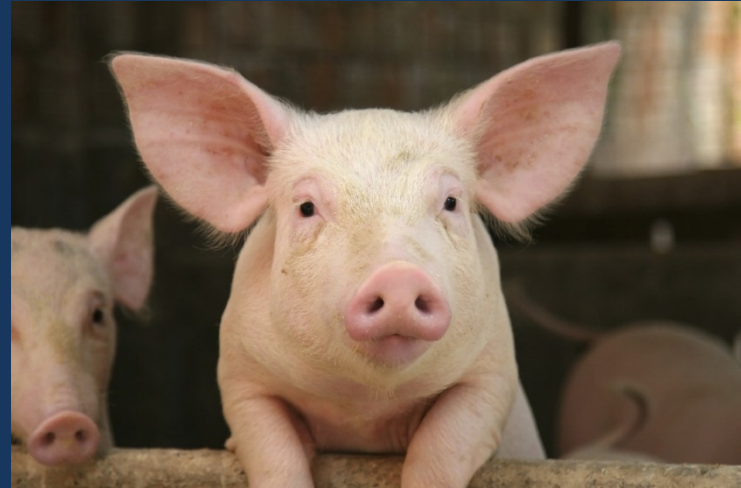
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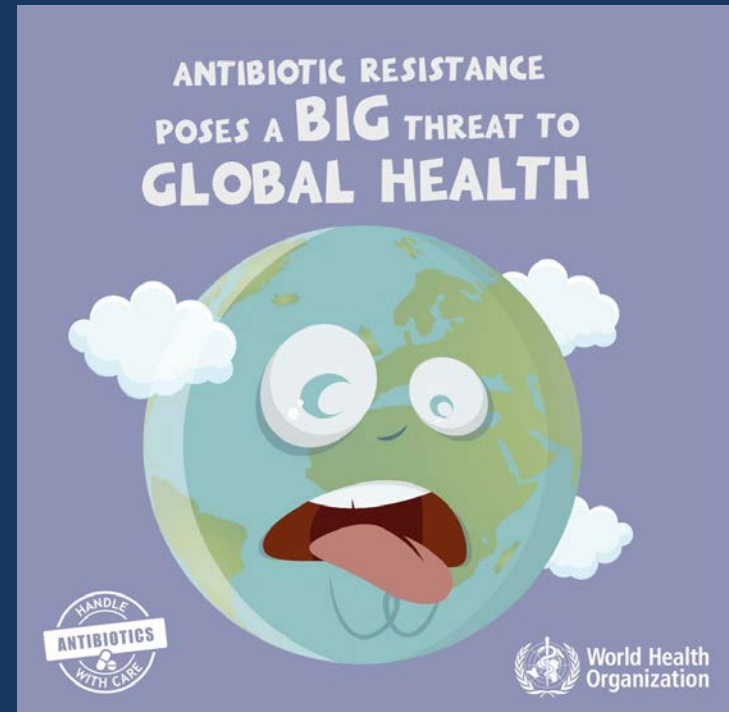
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Increased use of therapeutic antibiotics for food animals (48 tons in 1996; 94 tons in 2001)



# Changing Practices – is it effective?

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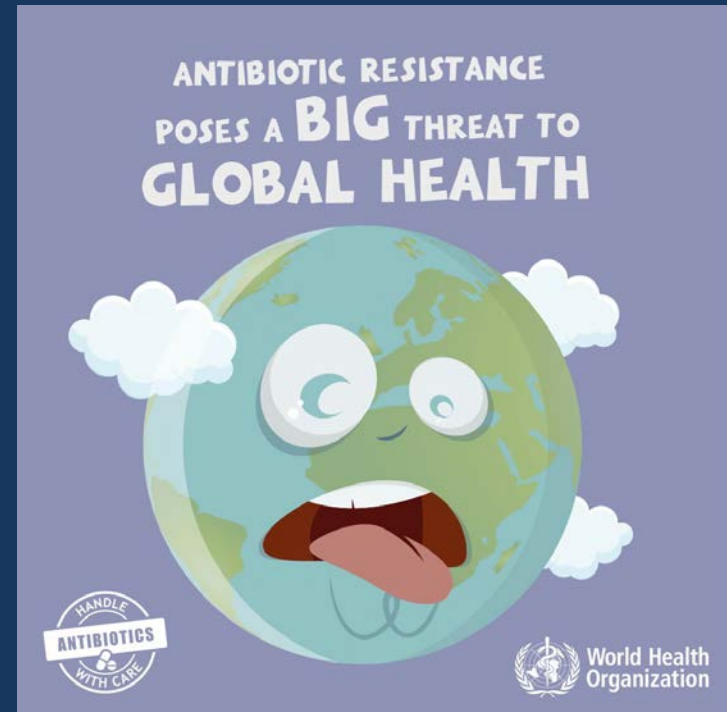




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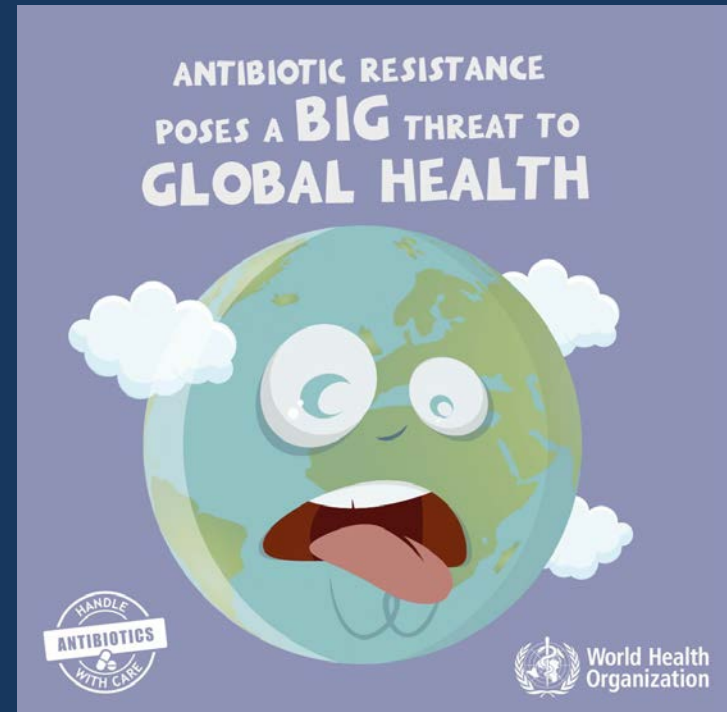


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Accurate assessments of **environmental quality impacts** – and accurate assessments of **human health risk** – increase in importance



# Questions?

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