#### Ecosystem structure and functional responses to pharmaceutical and personal care products in aquatic ecosystems



### Emma J. Rosi-Marshall



#### **Collaborators**

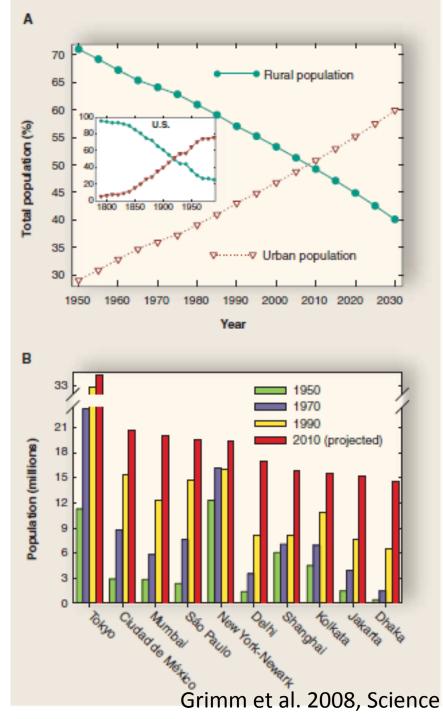
John Kelly, Heather Bechtold, Todd Royer, Dustin Kincaid, Brad Drury, Miguel Rojas, Paul Hoppe, Daniel Snow

#### **Funding Sources**

Illinois Water Resources Association Illinois Waste Management and Research NSF- BES LTER

### Currently there are more than 7 billion people on this planet

### Most of these people live in urban areas



### A typical urban stream

#### What you can't see

Antibiotics Antihistamines Antidepressants Painkillers Anticonvulsants Antimicrobials Hormones Fragrances Insect repellents Sunscreen

No wastewater treatment plant upstream

## A rapidly growing global industry

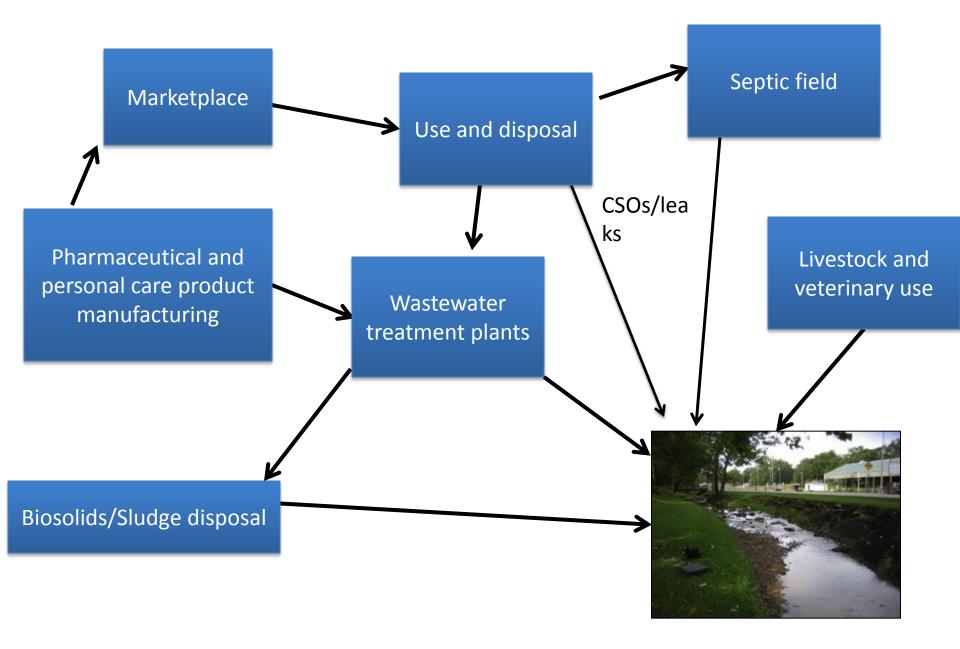
Global pharma market is expected to expand to more than US \$1 trillion by 2014, driven largely by Pharmerging markets



Source: IMS Health, Market Prognosis, Mar 2010. New Pharmerging definition

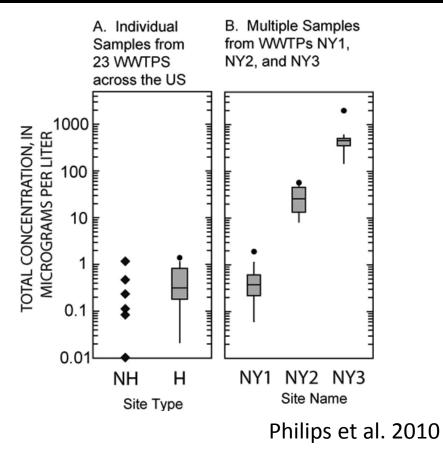


### How do drugs get into aquatic ecosystems?



### Your Pharmaceutical Footprint: Where do we get our drugs from?

Streams in New York that have pharmaceutical manufacturing facilities have much higher concentrations of PPCPs



### Your Pharmaceutical Footprint: Where do we get our drugs from?



Ciprofloxacin Concentration= 32 mg/L Typical dose= 250-750 mg From ~13% of a dose in 1 L of river water (Larsson et al. 2007)

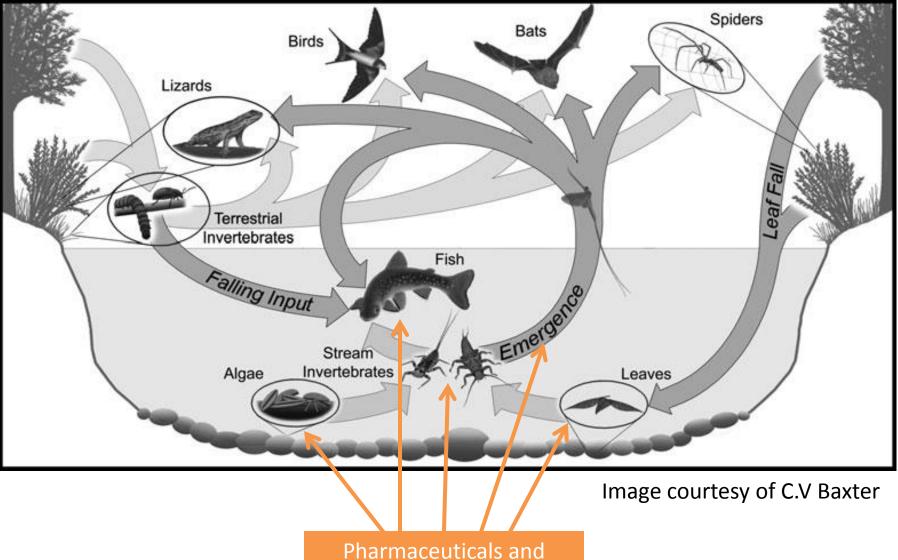
Pharmaceutical manufacturing plants release waste into one wastewater treatment plant

## Global treatment of wastewater

- UN estimates that only 10% of the world's wastewater is treated
- Untreated sewage enter US surface waters via CSOs and leaking infrastructure
- 40% of global population resides in coastal areas and 80-90% of sewage discharge in coastal zones is raw and untreated
- Estimated 200 million farmers grow crops in human wastes (Report from IWMI) and the WHO estimates that 10% of the world relies on this source of food

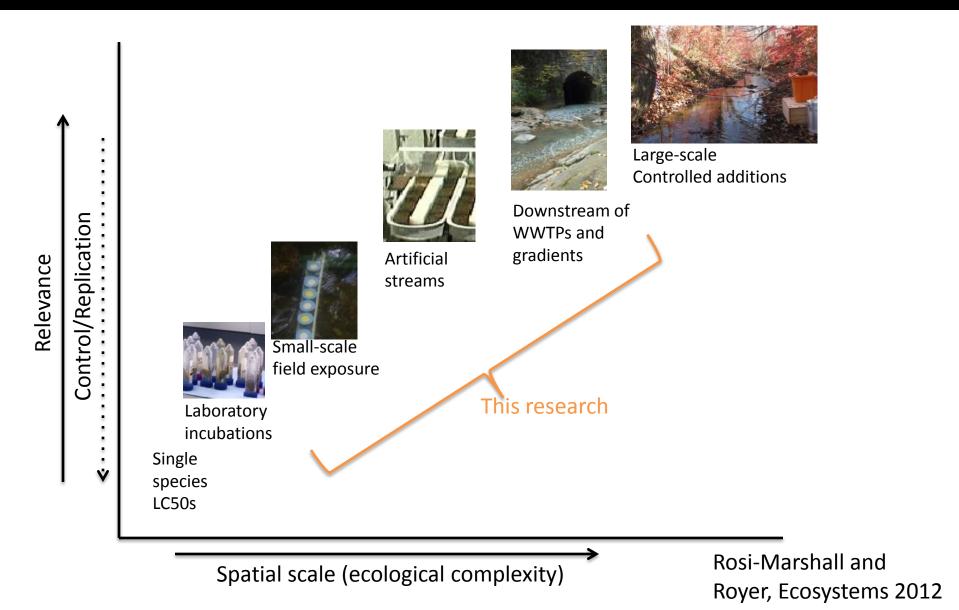


### Our rivers and lakes on drugs?



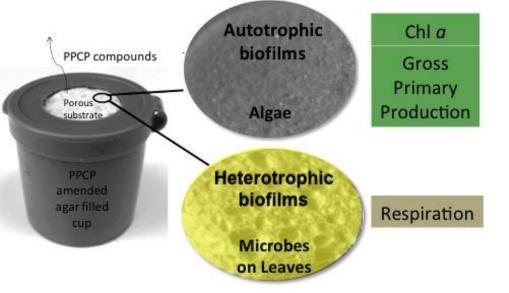
personal care products

## Measuring effects of drugs on streams using multiple scales of inquiry: from bottles to ecosystems



### Small-scale field exposure: New method

#### Pharmaceutical diffusing substrates





Compound	Medical Use
Caffeine	Stimulant
Cimetidine	Antihistamine (heartburn)
Ciprofloxacin	Antibiotic
Diphenhydramine	Antihistamine (allergens)
Metformin	Antidiabetic
Ranitidine	Antihistamine (heartburn)

# Pharmaceuticals can influence ecosystem function

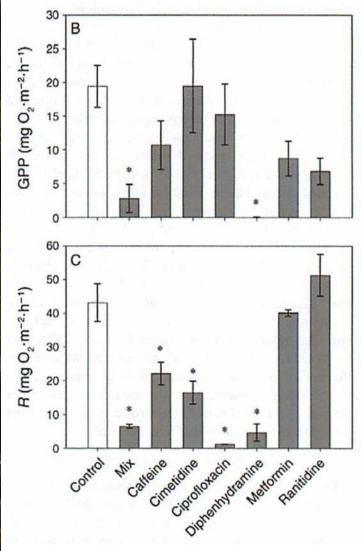
The responses vary by compound.

82% reduction in algal biomass by the Mix treatment.

Algal activity (GPP) was significantly suppressed by diphenhydramine (99% reduction) and Mix (86% reduction)

Respiration was significantly reduced by Cipro (97%), Diphenhydramine (89%), Caffeine (49%) Cimetidine (62%) and Mix (85%)





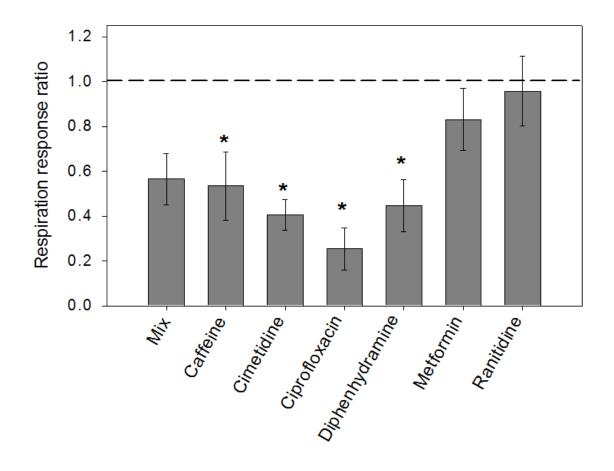
Rosi-Marshall et al. 2013, Eco Apps

# Consistent responses among seasons and streams

3 seasons in NY 1 season in IN 1 season in MD

Respiration of heterotrophic biofilms





Rosi-Marshall et al. 2013, Eco Apps

### Diphenhydramine can influence bacterial community structure

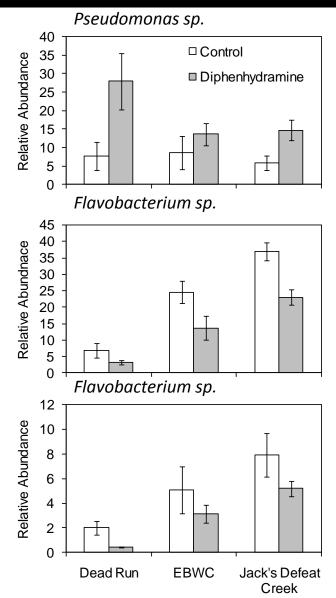


SIMPER Analysis of 16S rRNA tag pyroseqencing data sets for the control and diphenhydramine treatments from each site. OTU 4, OTU 1, and OTU 25 accounted for 14.33%, 12.25% and 3.43% of the variation between the control and diphenhydramine treatments.

Increase in a bacterial species known to degrade organic contaminants

Decrease in bacterial species known to rely of algal C

Rosi-Marshall et al. 2013, Eco Apps



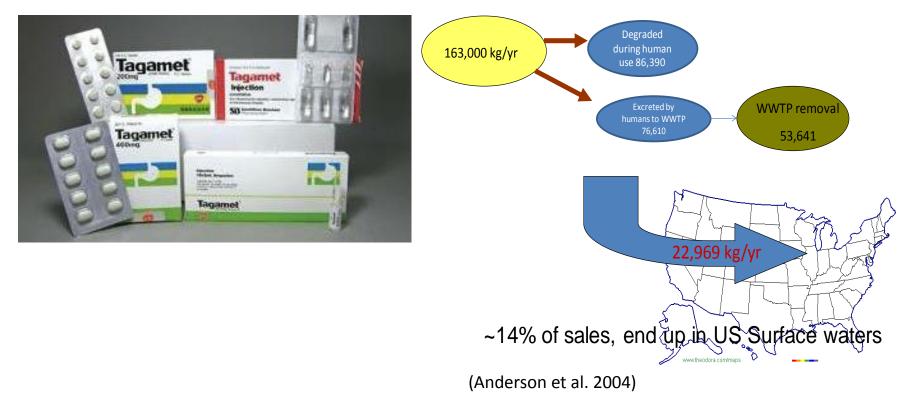
## Artificial streams



- Increases scale, duration of experiments, and ecological complexity
- Allows for examination of various endpoints and organisms

### Effects of a PPCP on aquatic invertebrates

- Tagamet<sup>®</sup> H<sub>2</sub> histamine antagonist that prevents secretion of stomach acid
- Cimetidine was approved by the FDA in 1977 and was the first drug ever to reach more than 1 billion dollars a year in sales.
- Histamine activates photoreceptors, olfactory receptors, and stomatogastric neurons (which control the motion of gut and foregut) in invertebrates
- These activities can be blocked by administration of antihistamines (Claiborne and Selverston 1984, Hardie 1988, Wachowiak 2002, Christie et al. 2004).



### Effects of a PPCP on an aquatic invertebrate

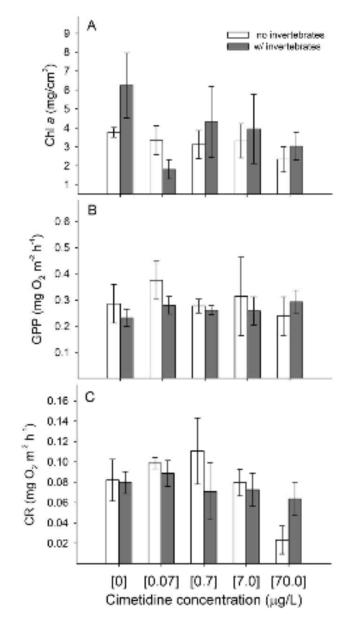
Long-term chronic exposure to 4 concentrations of cimetidine

Added cimetidine for 82 days to 32 streams.

No detectable effects on basal resources (algae and bacteria)



Hoppe, Rosi-Marshall and Bechtold, 2012, Freshwater Science



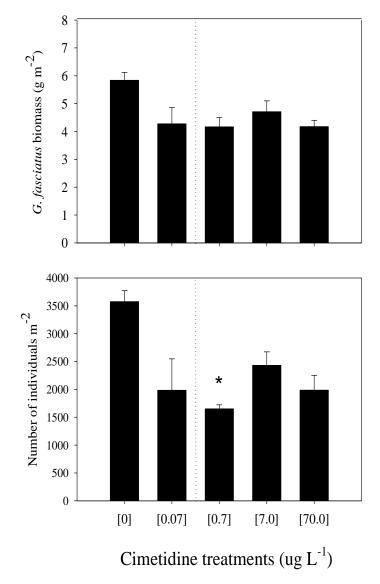
### Effects of a PPCP on aquatic invertebrates

- Long-term chronic exposure to 4 concentrations of cimetidine
- Had reproducing populations of invertebrates in streams
- Significant reduction in population growth and growth at concentrations of cimetidine observed in streams





Hoppe, Rosi-Marshall and Bechtold, 2012, Freshwater Science



# Effects of triclosan on riverine microbial communities

Triclosan: a generalized antimicrobial (bacteria, fungi and algae)

- Patented in 1966
- Nearly half of soaps in US contain triclosan
- Triclosan has been detected in surface waters, sediments, fishes, dolphins, human urine and breastmilk

What are the effects of triclosan on ecosystem structure and function?

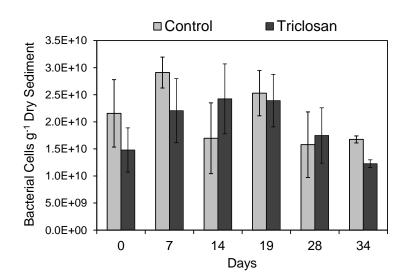


et al. 2013. ES&T

### Effects of triclosan on riverine microbial communities

Added triclosan in realistic concentrations to streams with active algal and bacterial communities

Initial decline in bacterial cells, with recovery after 7 days



Article

Artificial Streams at Loyola Univ. Chicago





#### Triclosan Exposure Increases Triclosan Resistance and Influences Taxonomic Composition of Benthic Bacterial Communities

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# Effects of triclosan on riverine microbial communities

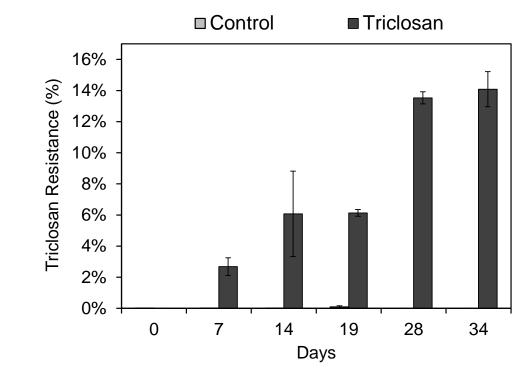
Increase bacterial resistance over time

This was evident after only 7 days and continued to increase until day 34

Examined the composition of bacterial communities with 16S rRNA tag pyrosequencing analysis

Significant change in the bacterial communities in response to triclosan additions





## "Natural" environmental releases: point sources

 Take advantage of release of PPCPs in the environment

• WWTPs, combined sewer overflows, sludge applications, artificial treatment wetlands, etc.

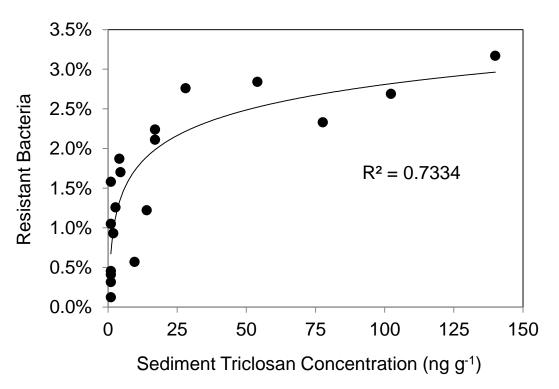
# Influence of wastewater effluent and triclosan on bacterial communities

- Conducted research in Chicago, IL
- Examined the microbial communities and function downstream of two wastewater treatment plants



## "Natural" environmental releases: point sources

• Measured triclosan, triclosan resistance, and bacterial composition

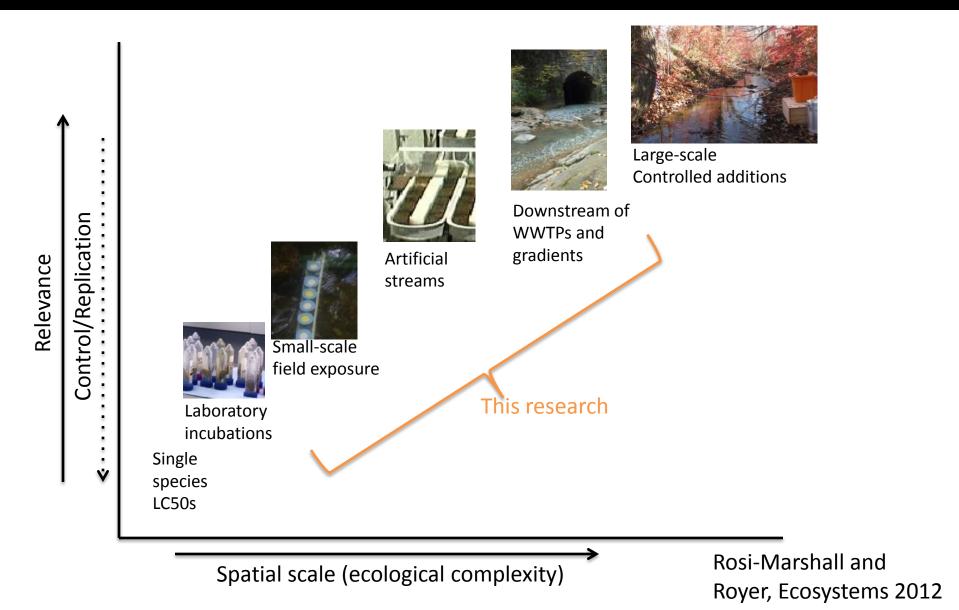


Triclosan concentrations in sediments correlated with resistant bacteria

Composition shifted in the field and some shifts were similar to shifts seen in artificial stream experiment

However, in another study we found that sewage discharges homegenize bacterial species composition (Drury et al. 2013 AEM)

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### Our rivers and lakes on drugs?

