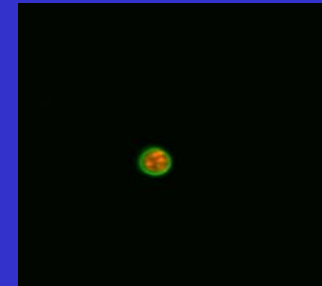
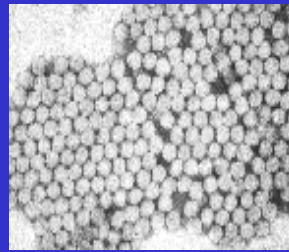
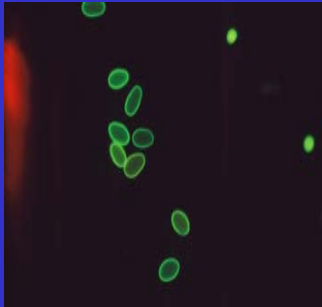


A Quantitative Microbial Risk Assessment Framework for the Great Lakes



Joan B. Rose
rosejo@msu.edu

Homer Nowlin Endowed Chair for Water Research

MICHIGAN STATE
UNIVERSITY











The Great Lakes

- Provides drinking water for 60 million U.S. and Canadian citizens
- 90% of the U.S. surface supply of freshwater
- Lakes cover 94,000 square miles
- 56 billion gallons used daily for municipal, agricultural and industrial use



The Great Lakes

- 10,000 miles of coastline in 8 different states
- 1/10th the population of U.S. is in basin
(Chicago, Detroit, Minneapolis, Milwaukee)
- \$4 billion commercial and sport fishing industry
- Over 500 recreational beaches

Challenges

- **Beach closures**
- **Nonpoint source pollution**
- **Small water systems**
 - **Groundwater protection**
 - **Septic systems**
- **Lakes & streams impairment (nutrients and Bacteria)**
- **Invasive Species**



Definitions used in risk analysis

| | |
|--------------------|--|
| Risk assessment | The qualitative or quantitative characterization and estimation of potential adverse health effects associated with exposure of individuals or populations to hazards (materials or situations, physical, chemical and or microbial agents.) |
| Risk management | The process for controlling risks, weighing alternatives, selecting appropriate action, taking into account risk assessment, values, engineering, economics, legal and political issues. |
| Risk communication | The communication of risks to managers, stakeholders, public officials, and the public, includes public perception and ability to exchange scientific information. |

PERCEIVED RISKS

Social, Economic, Legal and Political Context

RISK MANAGEMENT

Target must be defined
DALY, 10^{-4} , BAT reduction targets, will include motivational Factors.

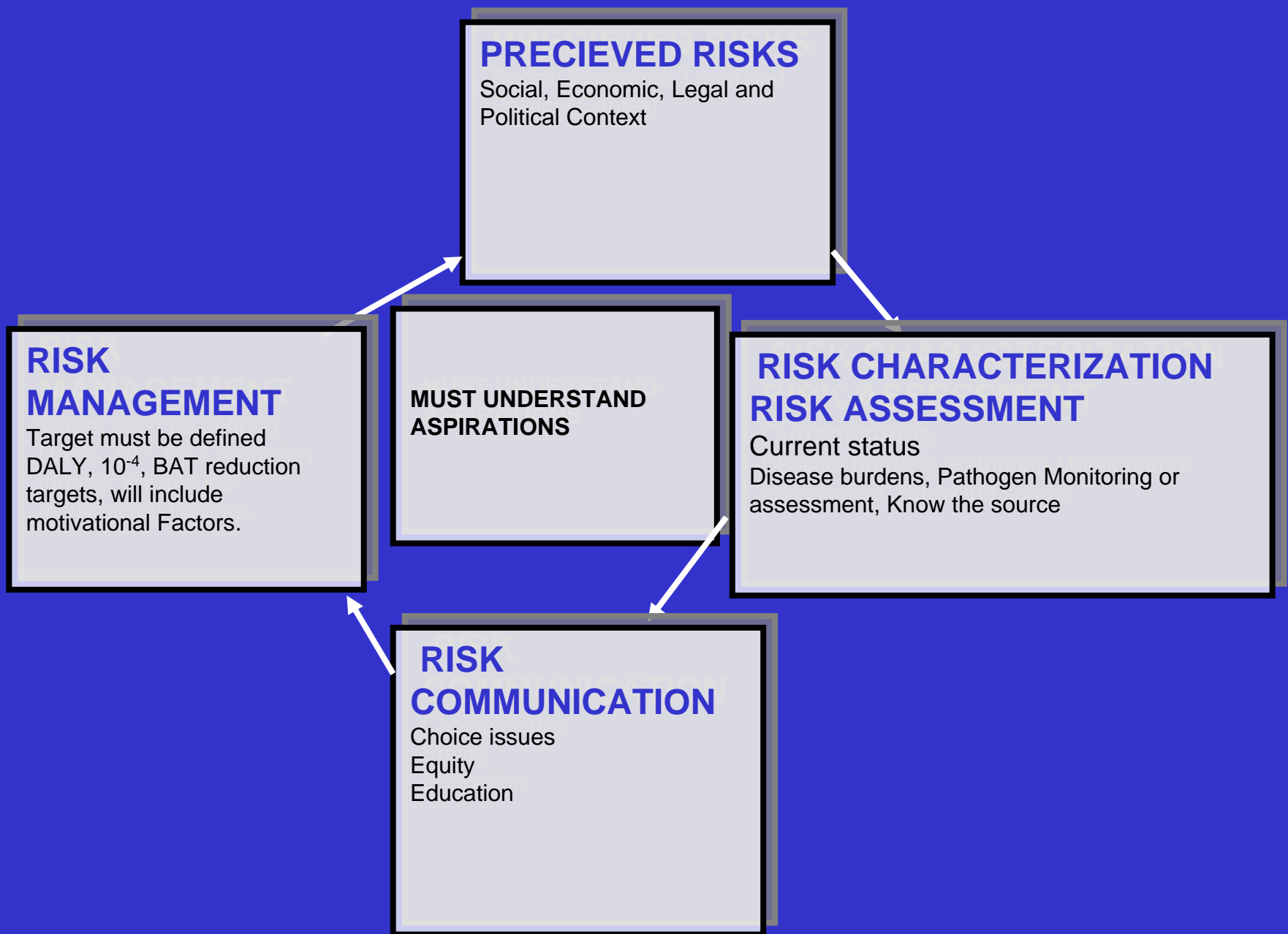
MUST UNDERSTAND ASPIRATIONS

RISK CHARACTERIZATION RISK ASSESSMENT

Current status
Disease burdens, Pathogen Monitoring or assessment, Know the source

RISK COMMUNICATION

Choice issues
Equity
Education



Risk assessment is a method to
examine qualitatively or
quantitatively the potential for
harm from exposure to
contaminants or specific hazards.

- Monitoring and data are some of the keys to establishing risks and therefore safety goals.

Quantitative Risk Assessment QRA

- Tool used to estimate adverse health effects associated with specific hazards.
- Elicits a statistical estimate or probability of harm.
- Used for risk management decisions.

NATIONAL ACADEMY OF SCIENCES RISK ASSESSMENT PARADIGM

HAZARD IDENTIFICATION

Types of microorganisms and disease end-points

DOSE-RESPONSE

Human feeding studies, clinical studies, less virulent microbes and health adults

EXPOSURE

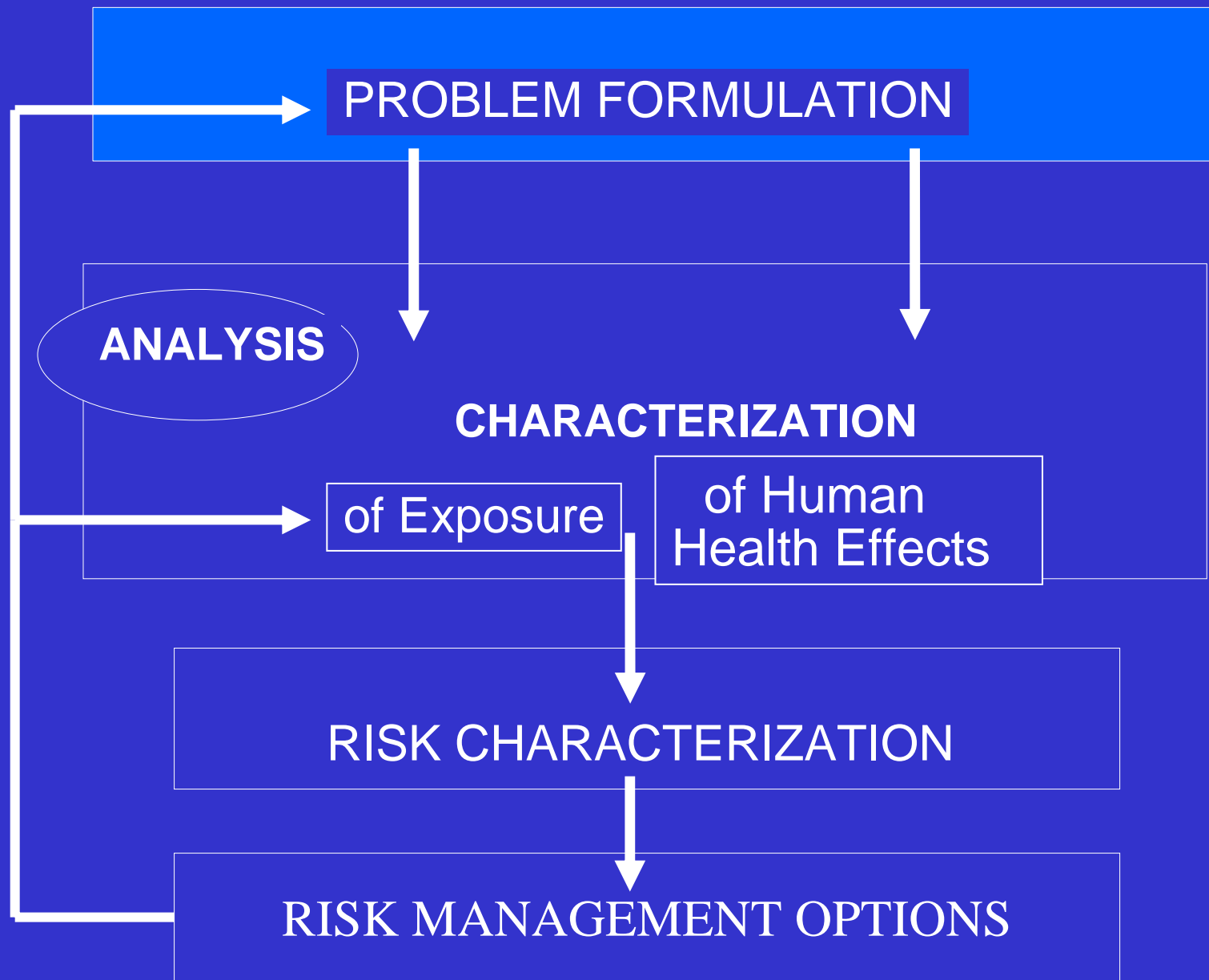
Monitoring data, indicators and modeling used to address exposure

RISK CHARACTERIZATION

Magnitude of the risk, uncertainty and variability

Evolution of QMRA





ANALYSIS PHASE

```
graph TD; subgraph Left; EA[Exposure Analysis]; PO["Pathogen Occurrence (detection/survival and spread)"]; EP[Exposure Profile]; end; subgraph Right; HE["Health Effects Disease Severity Secondary spread"]; DR[Dose-Response]; HPP[Host Pathogen Profile]; end; EP <--> HPP;
```

Exposure
Analysis

Pathogen
Occurrence
(detection/survival
and spread)

Exposure
Profile

Health
Effects
Disease
Severity
Secondary spread

Dose-Response

Host Pathogen
Profile

The Problem Formulation

Microbial Risks

PLAGUES

CONTAGION

EPIDEMICS

OUTBREAKS

1914

International Joint Commission Study

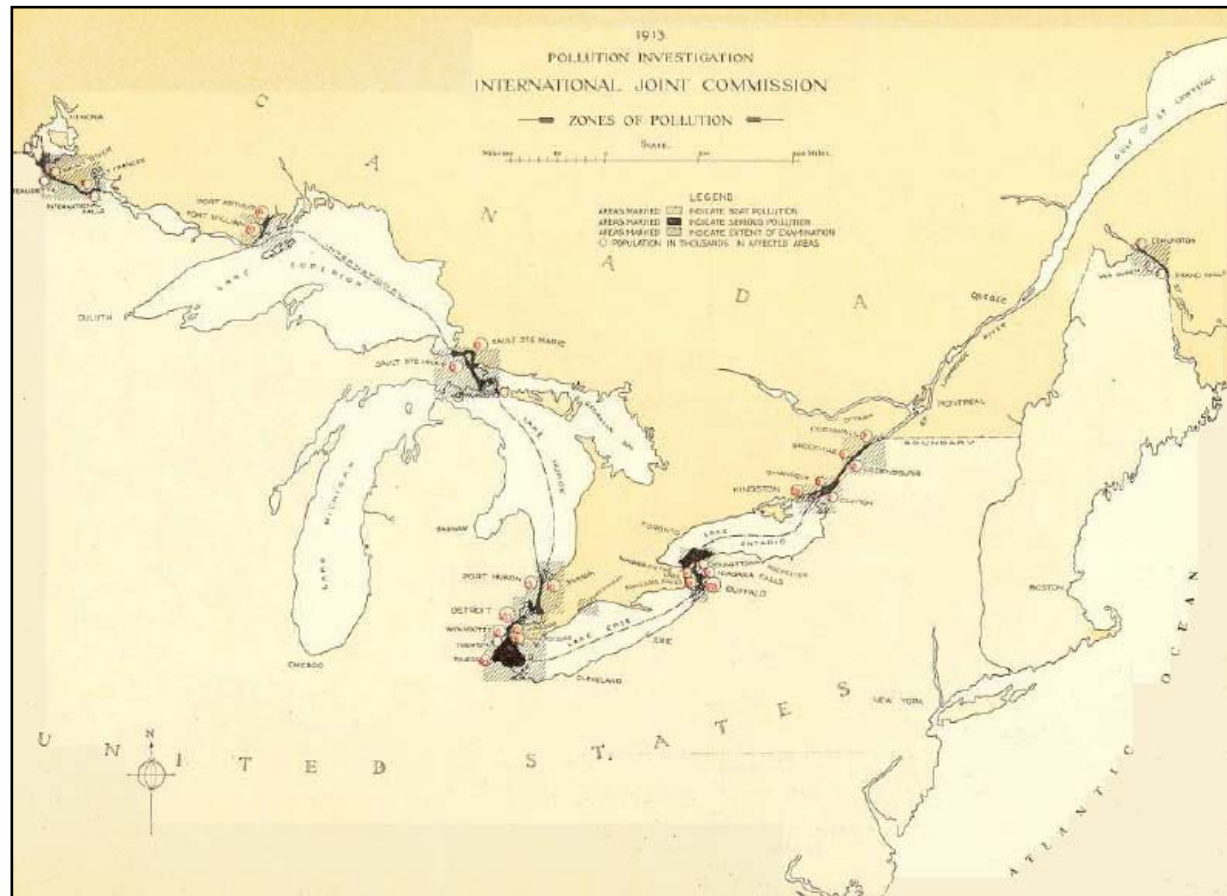
WATER QUALITY AGREEMENT

“Between 1912 and 1914 a massive bacteriological study of pollution in the boundary waters of the United States and Canada was conducted by the International Joint Commission: it was followed by another study of current and proposed sewage work. The findings of transboundary pollution from the bacteriological study led to a draft treaty on pollution in 1920. The bacteriological work was flawless.”

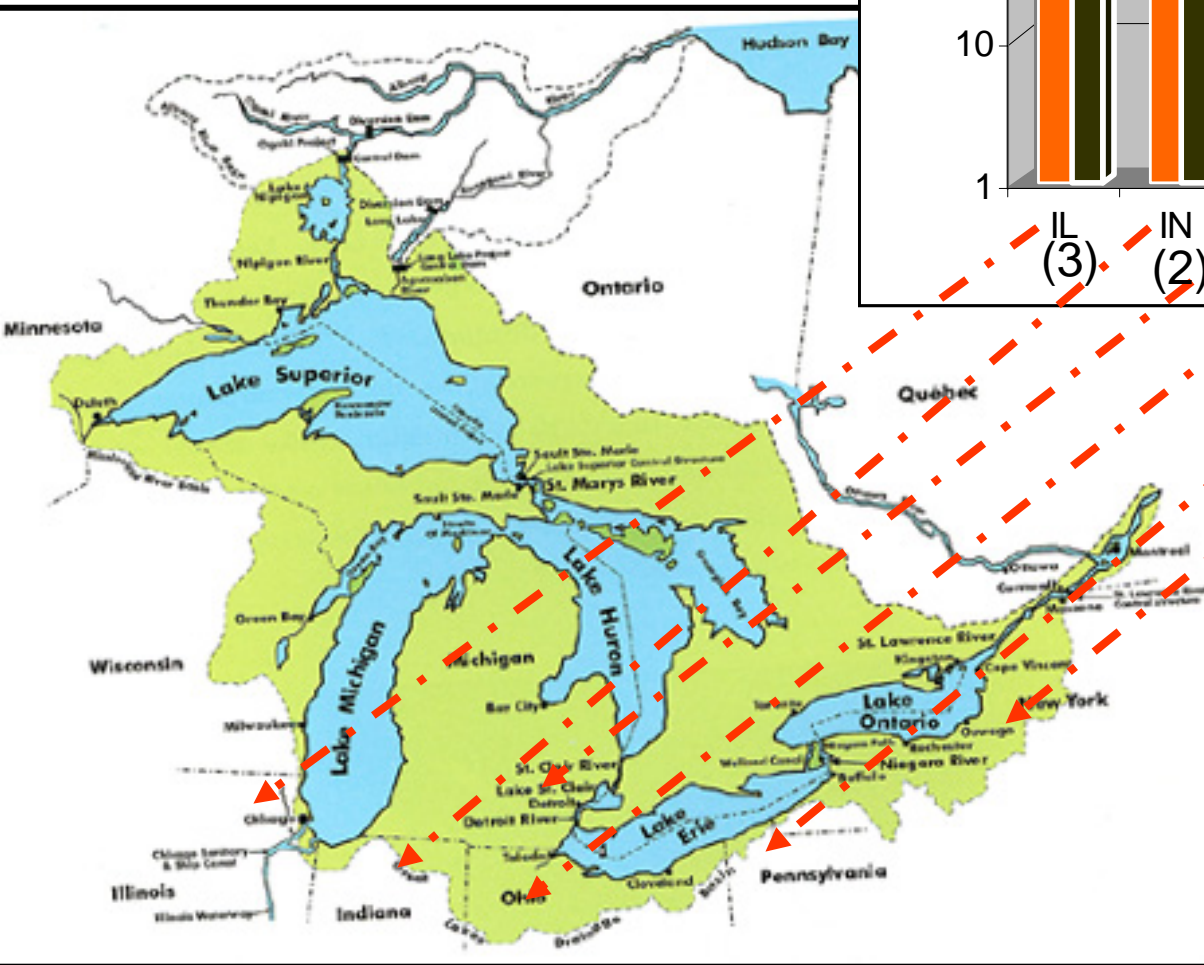
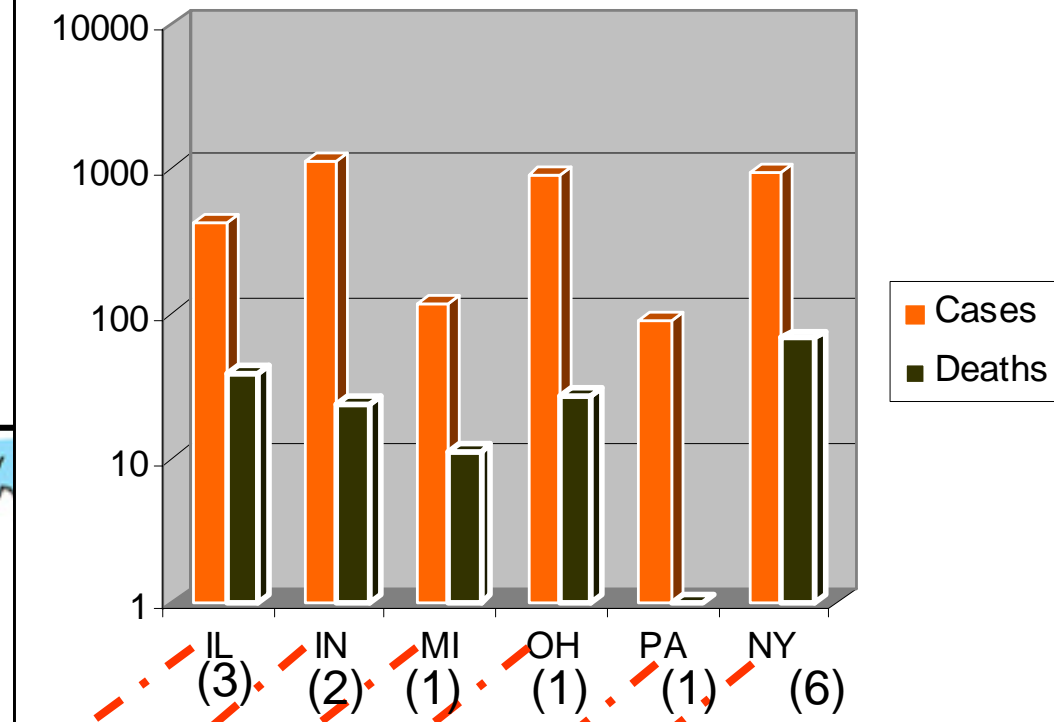
●

- **Municipal drinking water supplies were not safe – water intakes were located in highly polluted waters and water was not adequately treated**
- **Current sewage/drinking water treatment technologies were limited and expensive to implement**

- 1. Prohibit discharge of untreated sewage/ship ballast into boundary waters**
- 2. Discharge limit for *B. coli* (annual mean 500/100ml)**
- 3. Prohibit/restrict discharge of garbage, sawmill and industrial wastes**

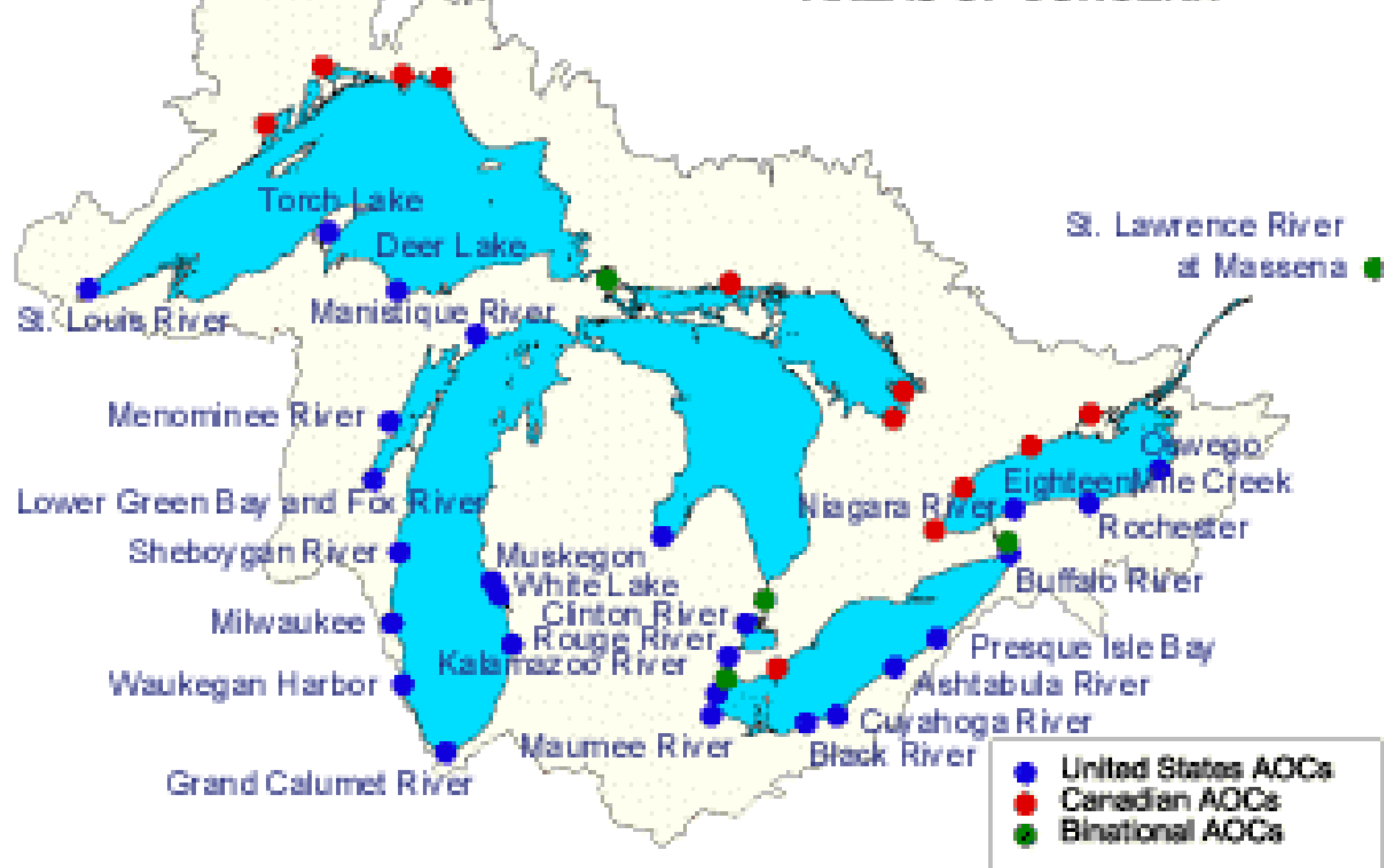


- **Largest Reported Typhoid Outbreaks in the US 1920-1929**



14 of top 25 outbreaks
(Wolman and Gorman, 1931)

GREAT LAKES AREAS OF CONCERN



**Emerging Waterborne Disease
Of the 21st Century: Largest
outbreak In the US 1993.
400,000 people ill (50% of
the population) 100 died
Cattle & Sewage blamed
Water met all requirements
Under the Safe Drinking
Water Act**



Investigation Continues Into Outbreak

Lake Michigan

**Linnwood Ave
Plant**
Capacity:
275 million
gallons a day

Waste from farms
upriver may have
contained the
Cryptosporidia
that are suspected
of entering the
water system.

Milwaukee

Texas Ave Station

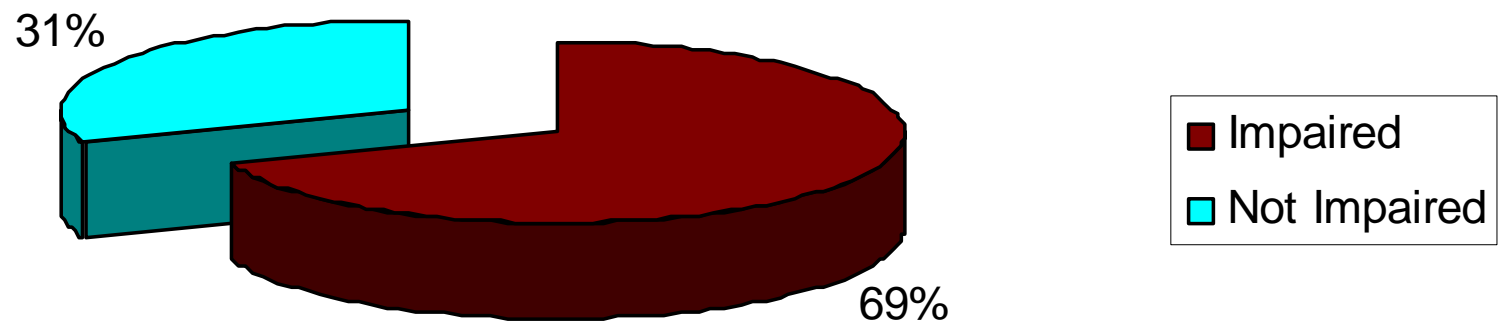
Intake
(7,800 ft offshore)

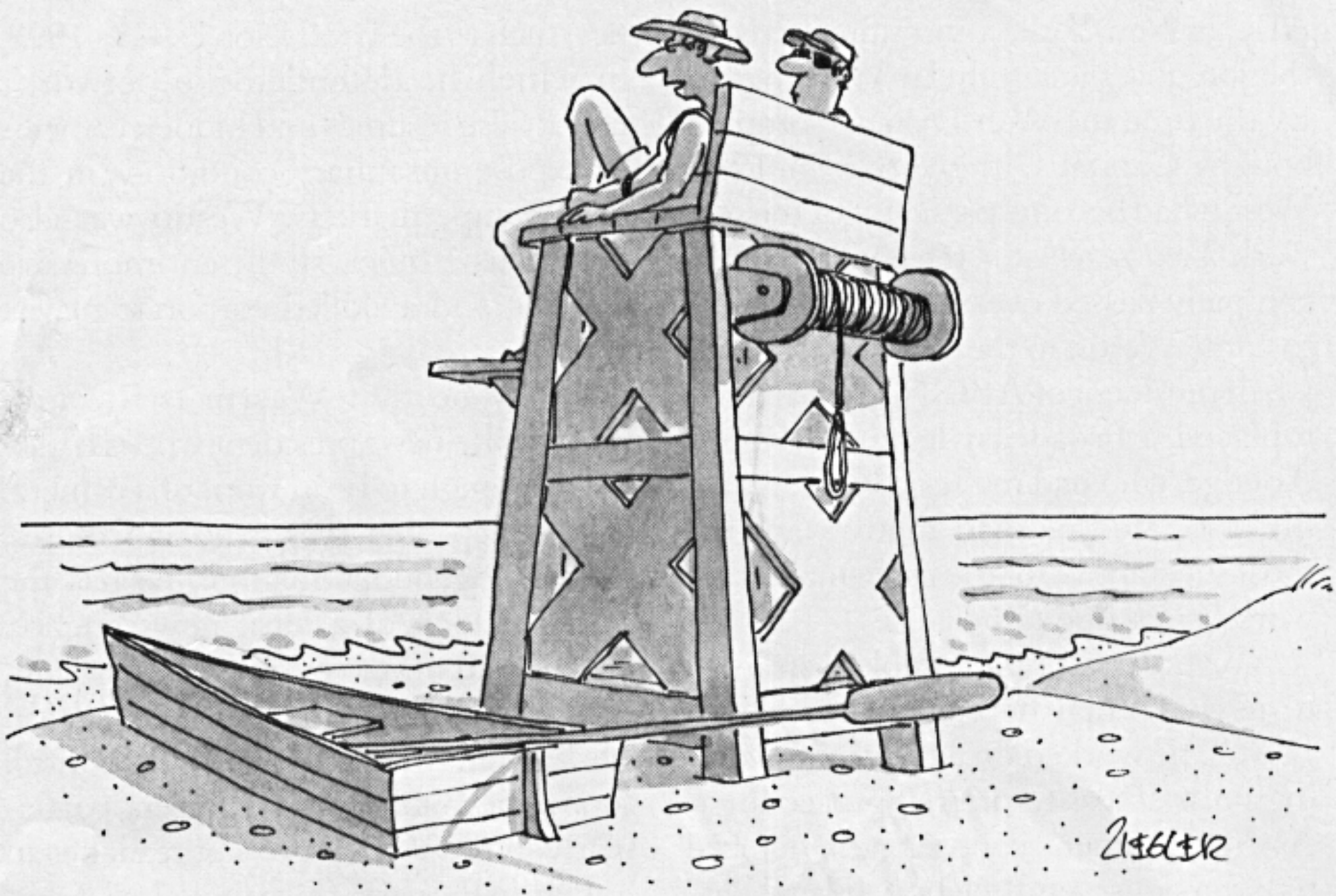
Howard Ave Plant

Capacity:
100 million
gallons a day

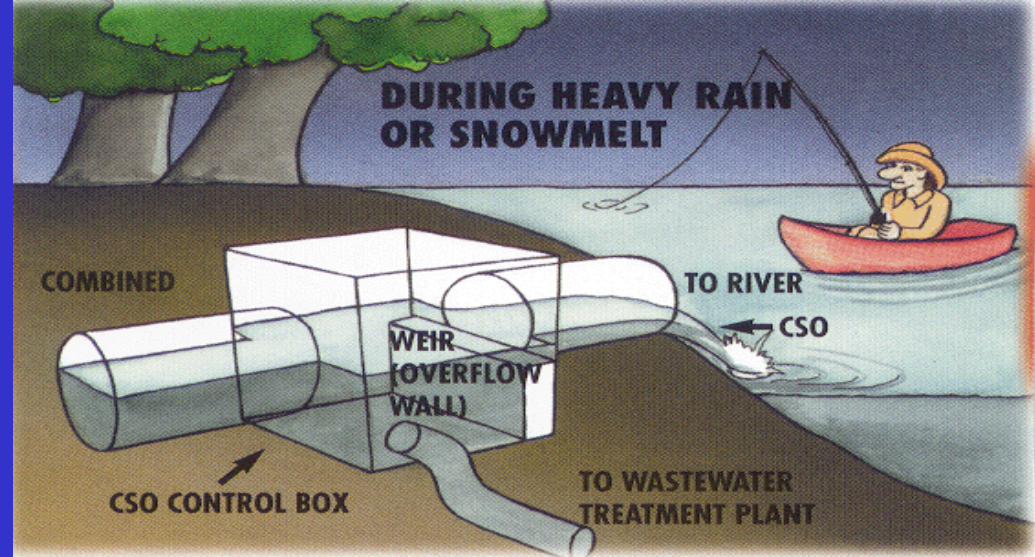
● **Pumping stations**

Recreational Water Impairment in the AOCs

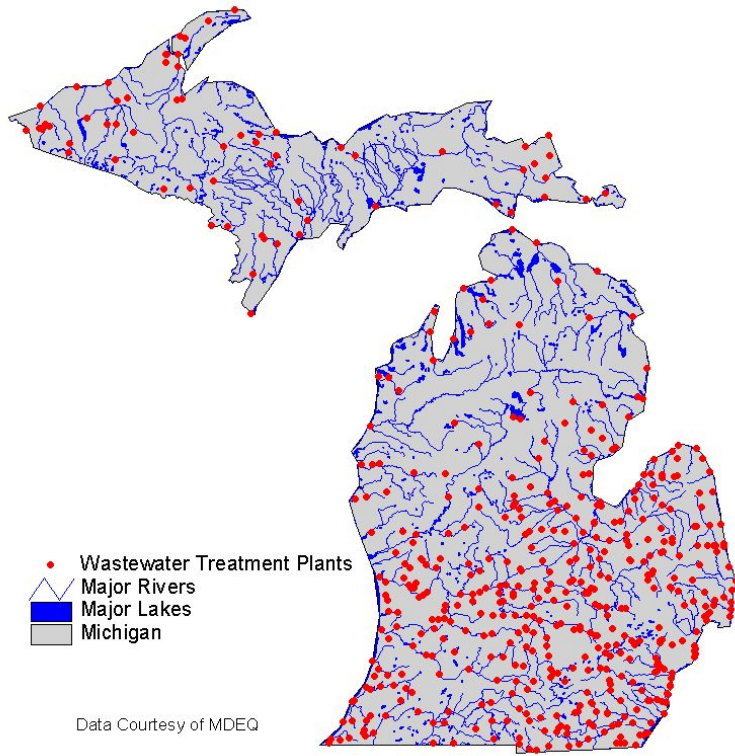




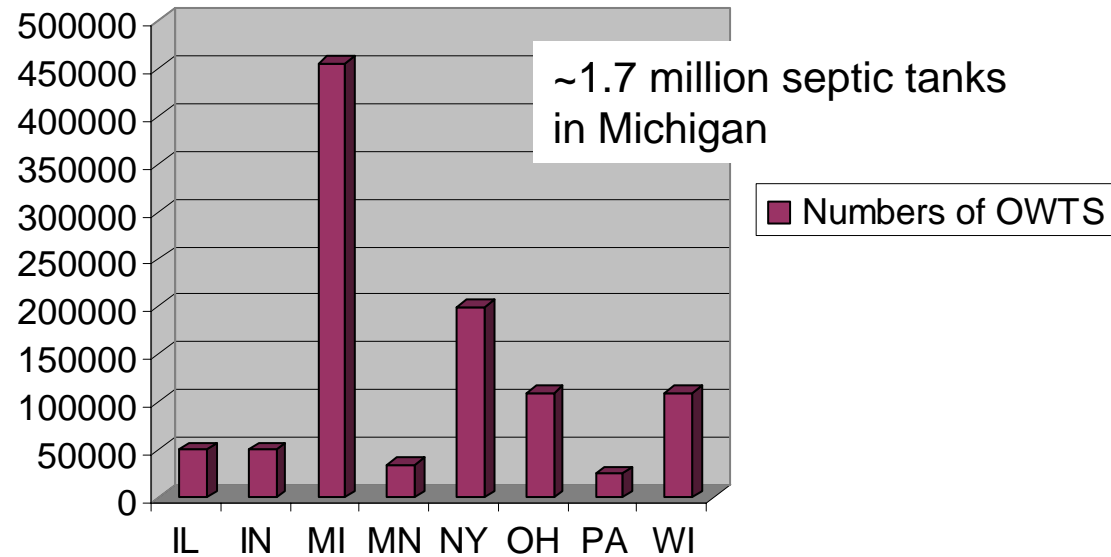
"I adore the beauty and tranquillity of these raw-sewage days."



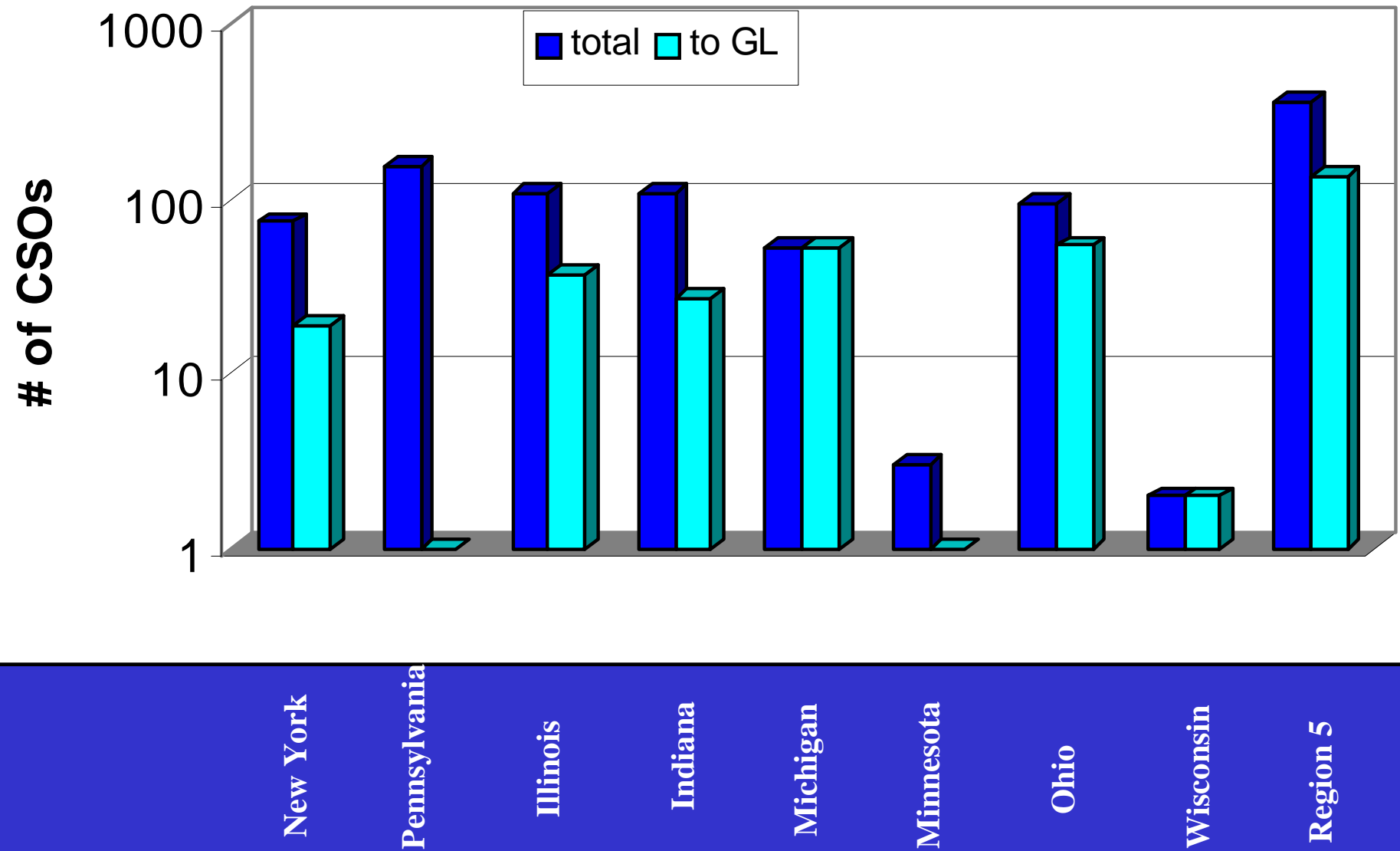
On site Wastewater Systems in the Great Lakes



Data Courtesy of MDEQ



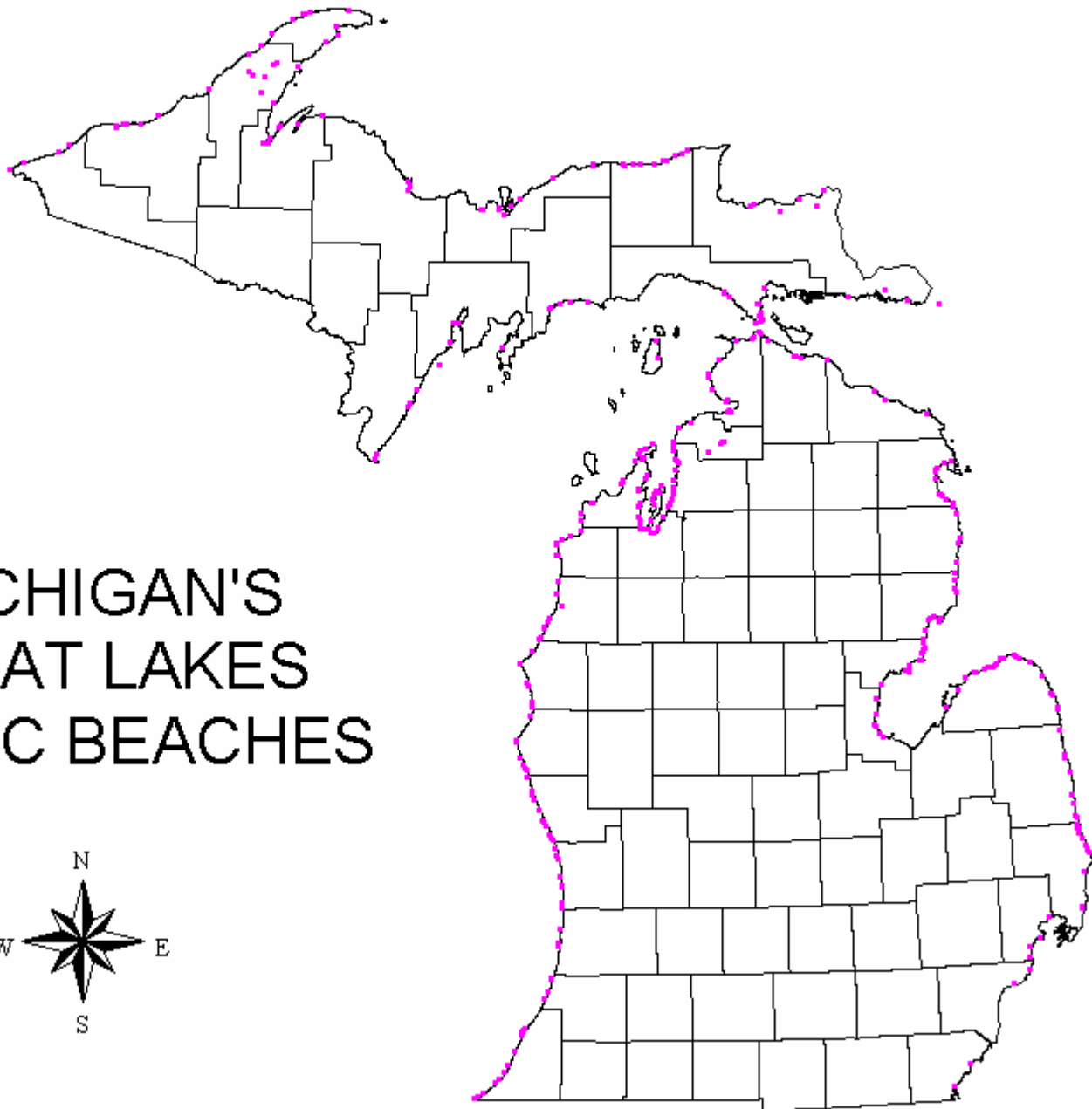
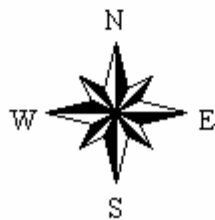
CSQs in the Great Lakes Region



- > 18,000 days of closings and advisories at USA beaches in 2003
- ~20,000 days of closings in 2004
- Fecal pollution indicators, give no indication as to the source of fecal pollution.
- In Michigan in 2004, 27 billion gallons of a sewage-storm water mix was dumped into the Great Lakes
- 2005 Michigan communities reported 338 sanitary sewer overflows 147 MG raw sewage into the GL



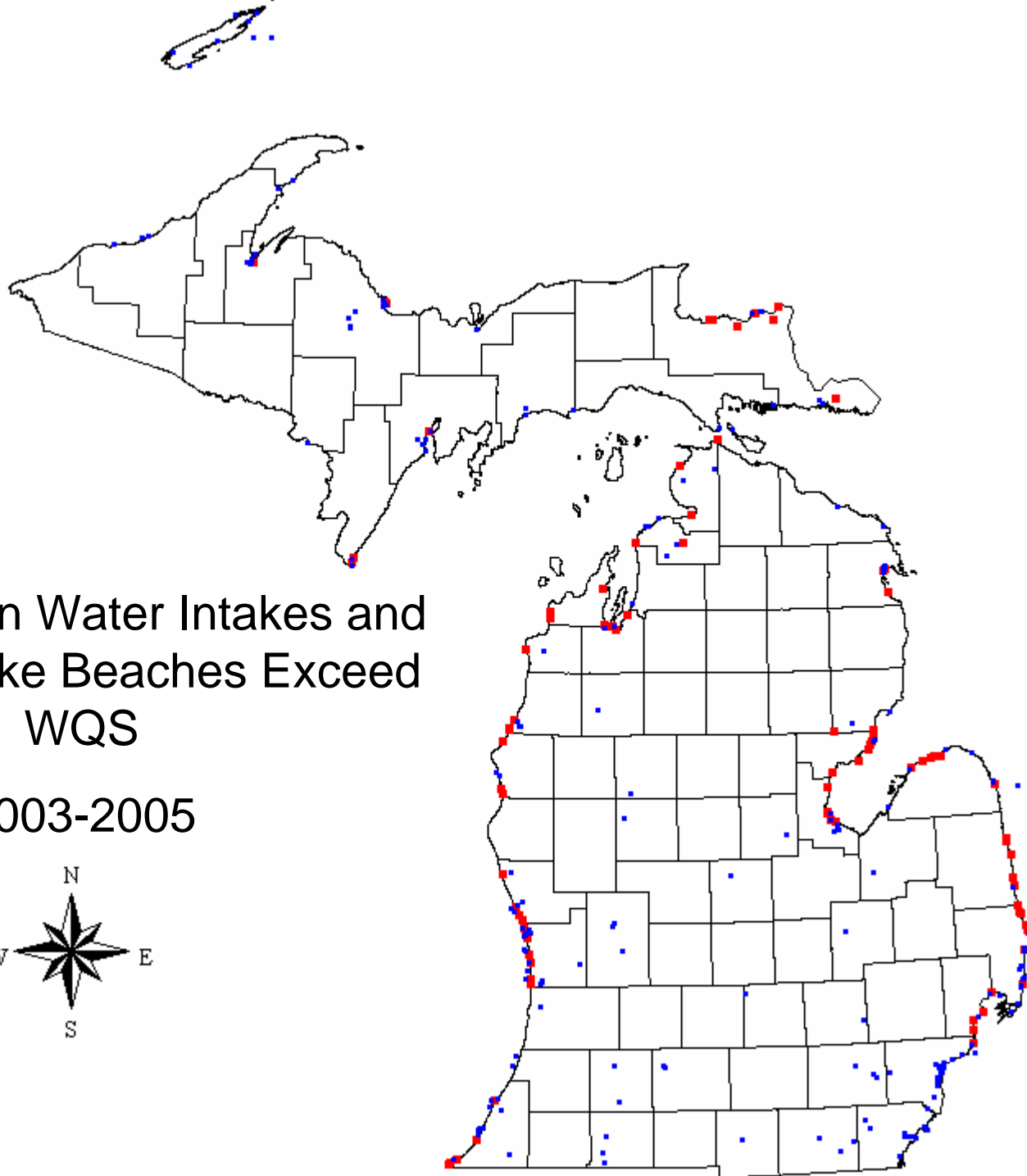
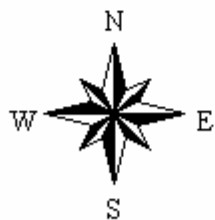
MICHIGAN'S GREAT LAKES PUBLIC BEACHES

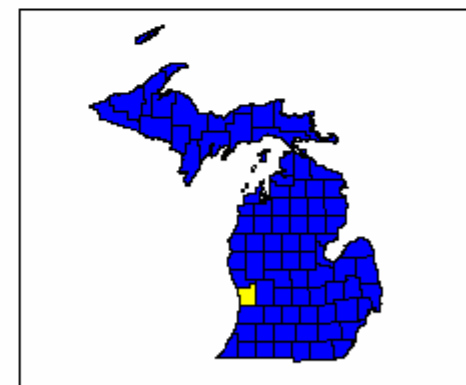
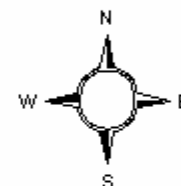
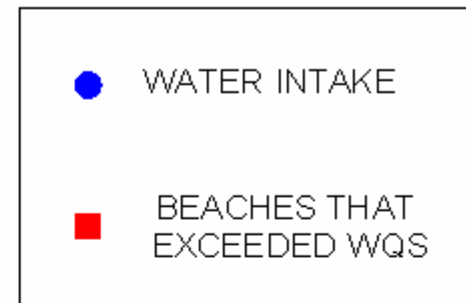
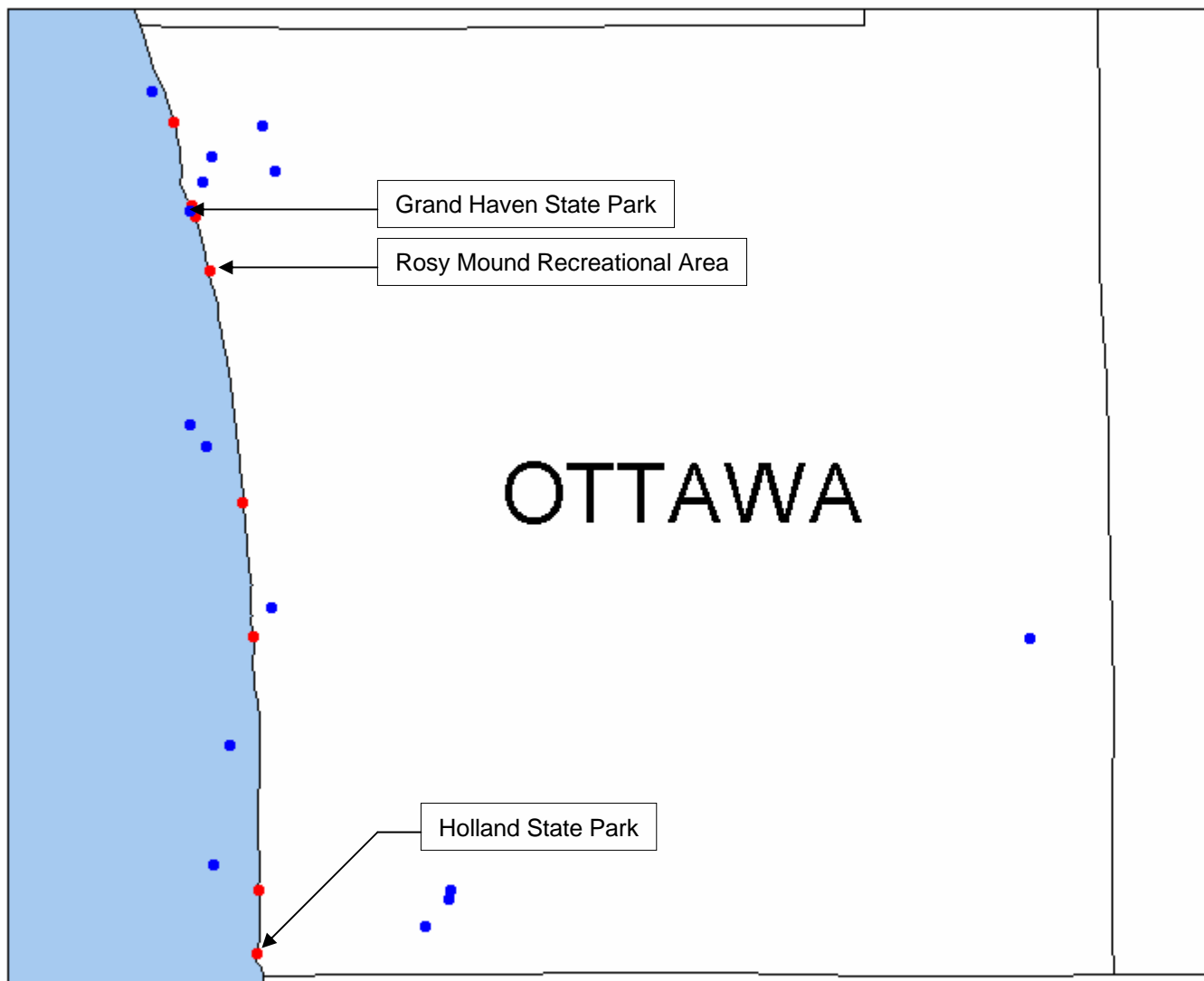


N=412 Beaches

219 Michigan Water Intakes and 96 Great Lake Beaches Exceed WQS

2003-2005





8 Exceedance sites
15 Water Intake sites

The HAZARDS

Emerging Biological Hazards

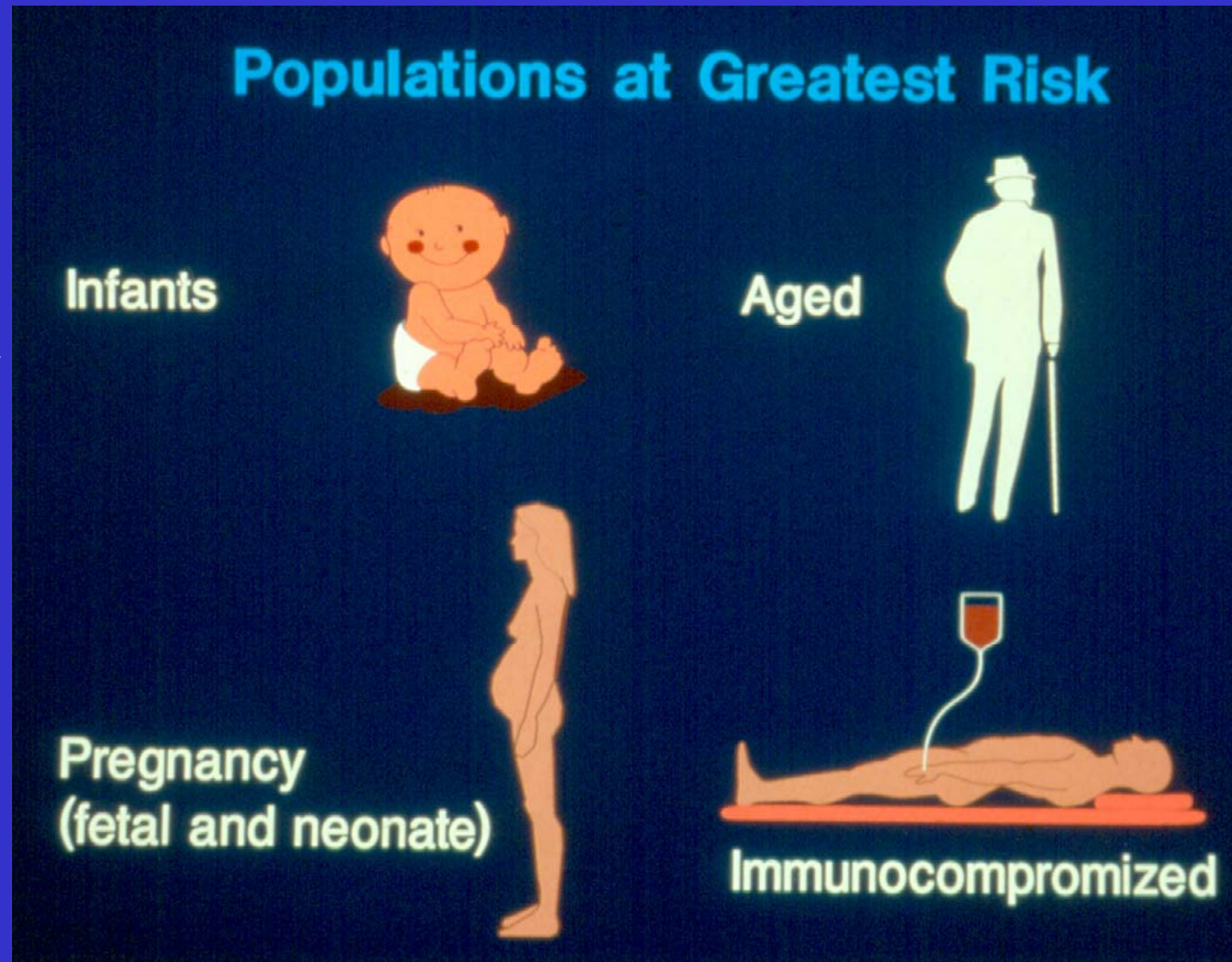
- Viruses, prions, bacteria, and protozoa are more likely than fungi or helminths to be associated with emerging infections.
- Zoonotic pathogens comprise 75% of emerging infectious diseases.
- Pathogens which are subject to relatively frequent mutation or genomic reassortment events (e.g. RNA viruses and viruses with segmented genomes) are more likely to emerge.
- Pathogens which infect multiple hosts or pathogens that infect species that can harbour multiply closely related agents providing an opportunity for reassortment or recombination (e.g. SARS in cats) are likely to emerge.
- Agents transmissible by more than one route or by indirect contact, e.g. water, food, environmental contamination, vectors, etc, are likely to emerge.

Acute and Chronic Outcome Associated with Microbial infections

| | Acute disease | Chronic disease |
|---|---|---|
| Microorganism | Outcomes | Outcomes |
| <i>Campylobacter</i> | Diarrhea | Gullain-Barre' syndrome |
| <i>E. Coli 015H7</i> | Diarrhea | Hemolytic uremic syndrome |
| <i>Helicobacter</i> | Gastritis | Ulcers and stomach cancer |
| <i>Salmonella,</i> <i>Shigella, & Yersinia</i> | Diarrhea | Reactive arthritis |
| <i>Coxsackievirus B</i> <i>Adenoviruses</i> | Encephalitis, aseptic Meningitis, diarrhea, respiratory disease | Diabetes Myocarditis Obesity |
| <i>Giardia</i> | Diarrhea | Failure to thrive, lactose intolerance, chronic joint pain |
| <i>Toxoplasma</i> | Newborn syndrome, hearing and visual loss | Mental retardation, dementia, seizures |

Morbidity and Mortality greater in the Sensitive Populations
30% of our populations Fall into one of the Sensitive Populations
at any one time.

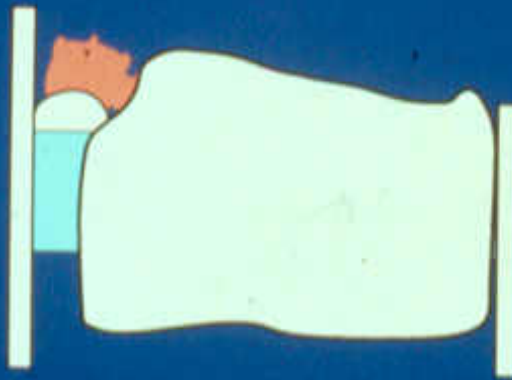
ZOONOTIC
AGENTS
OPPORTUNISTIC
AGENTS
EFFECT
THIS
GROUP



DOSE-RESPONSE

Outcomes of Microbial Exposure

Infection → Disease → Mortality

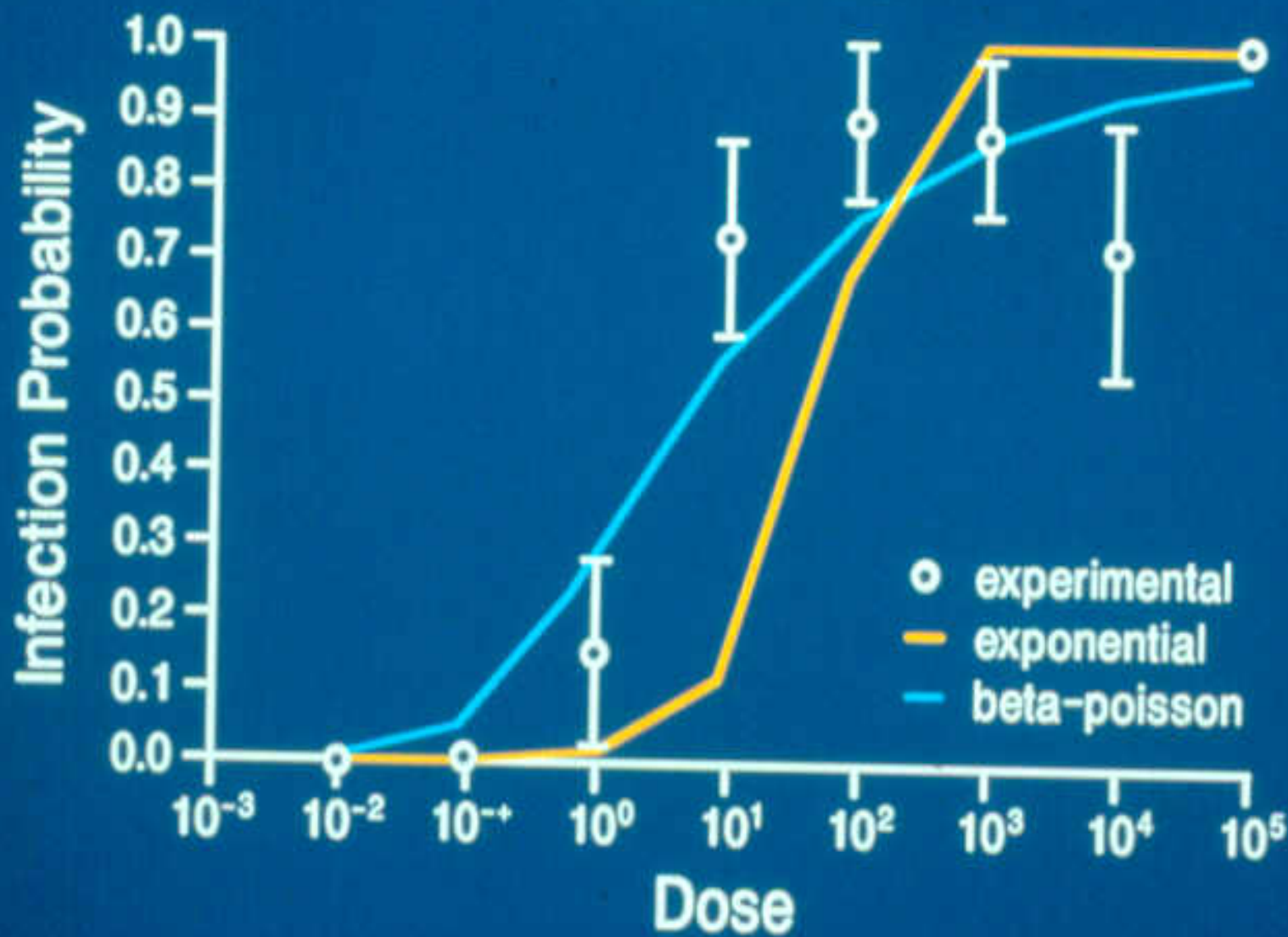


Dose~Response

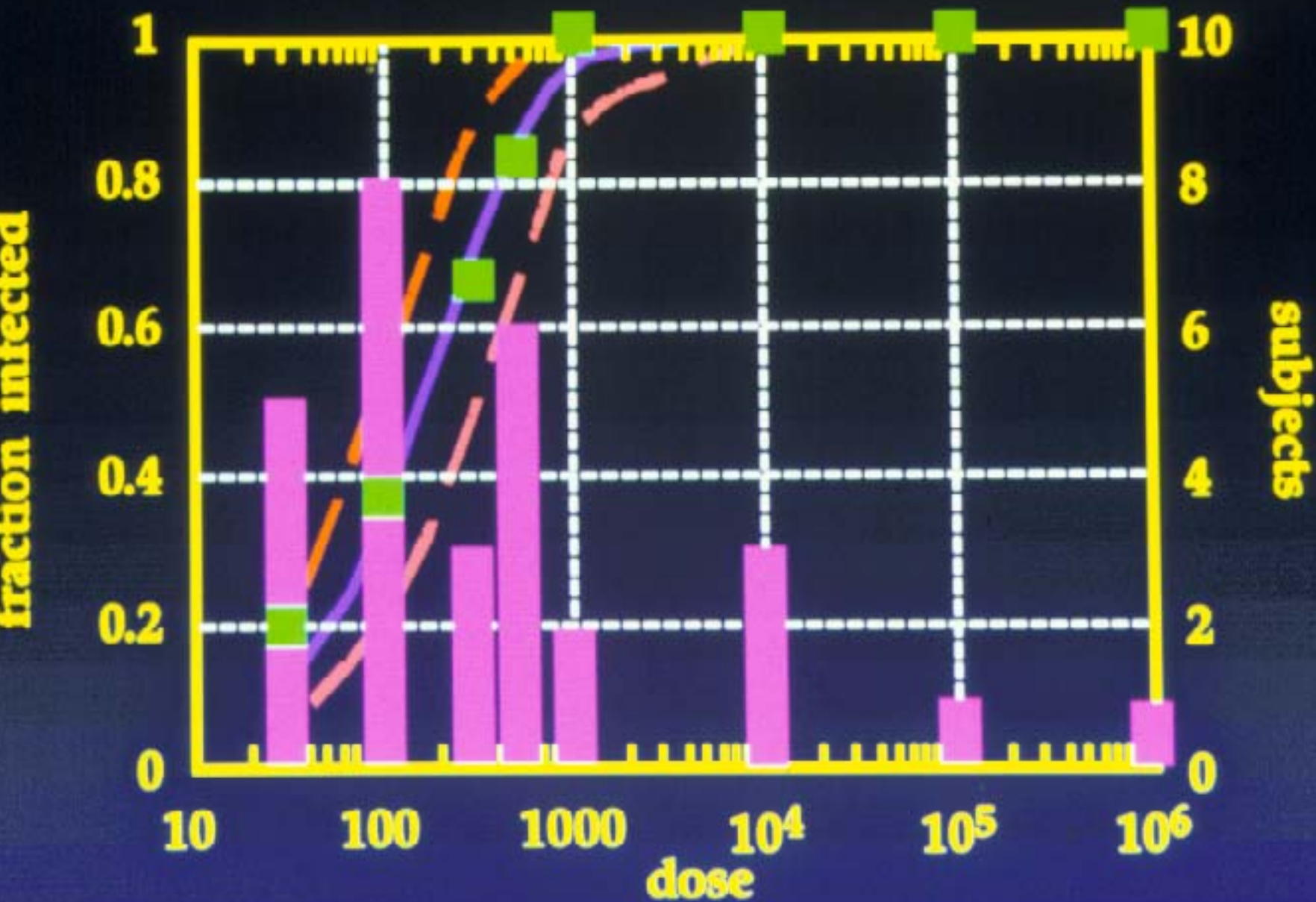
Dose~response data sets have been developed in human feeding studies for

- Dose measurements were by PFU/or by infectious titer, CFU or cysts or oocysts.
- End points of measurements were excretion of the pathogen and/or antibody response, rarely disease.
- Mathematically address the shape of the ratio of those affected/exposed.
- Need minimum of three doses. Must have doses which elicit effects different from 0% and 100%

Rotavirus



Dose - Response Information (DuPont)



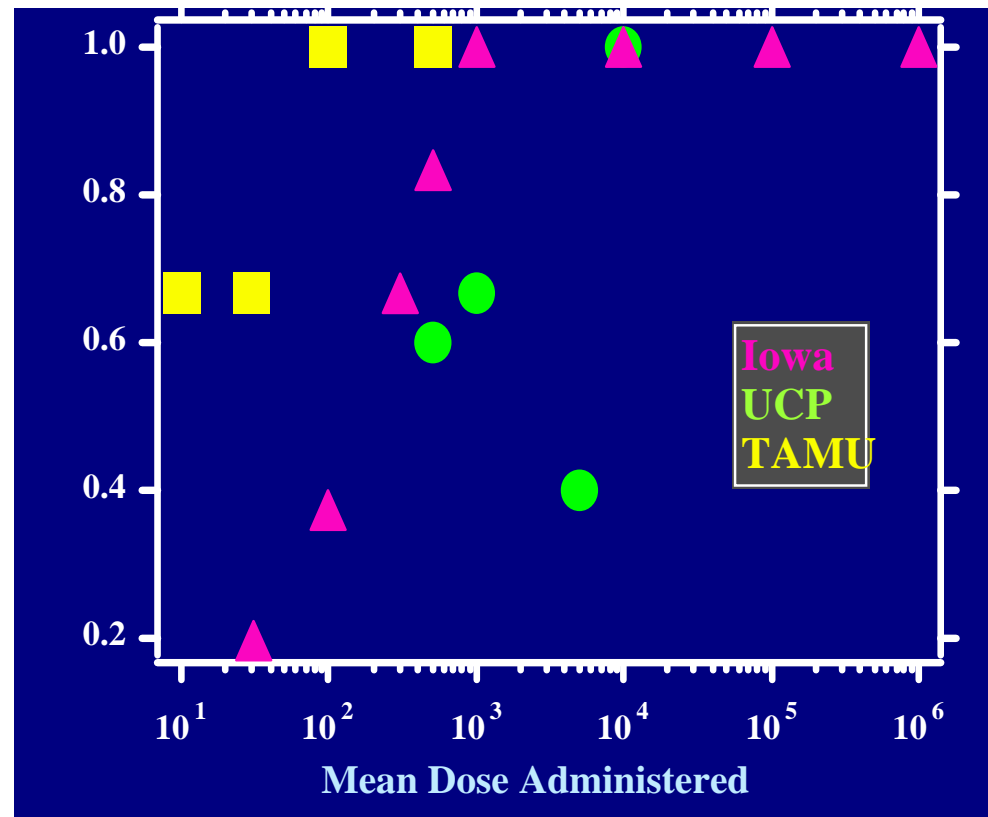
Strain Differences

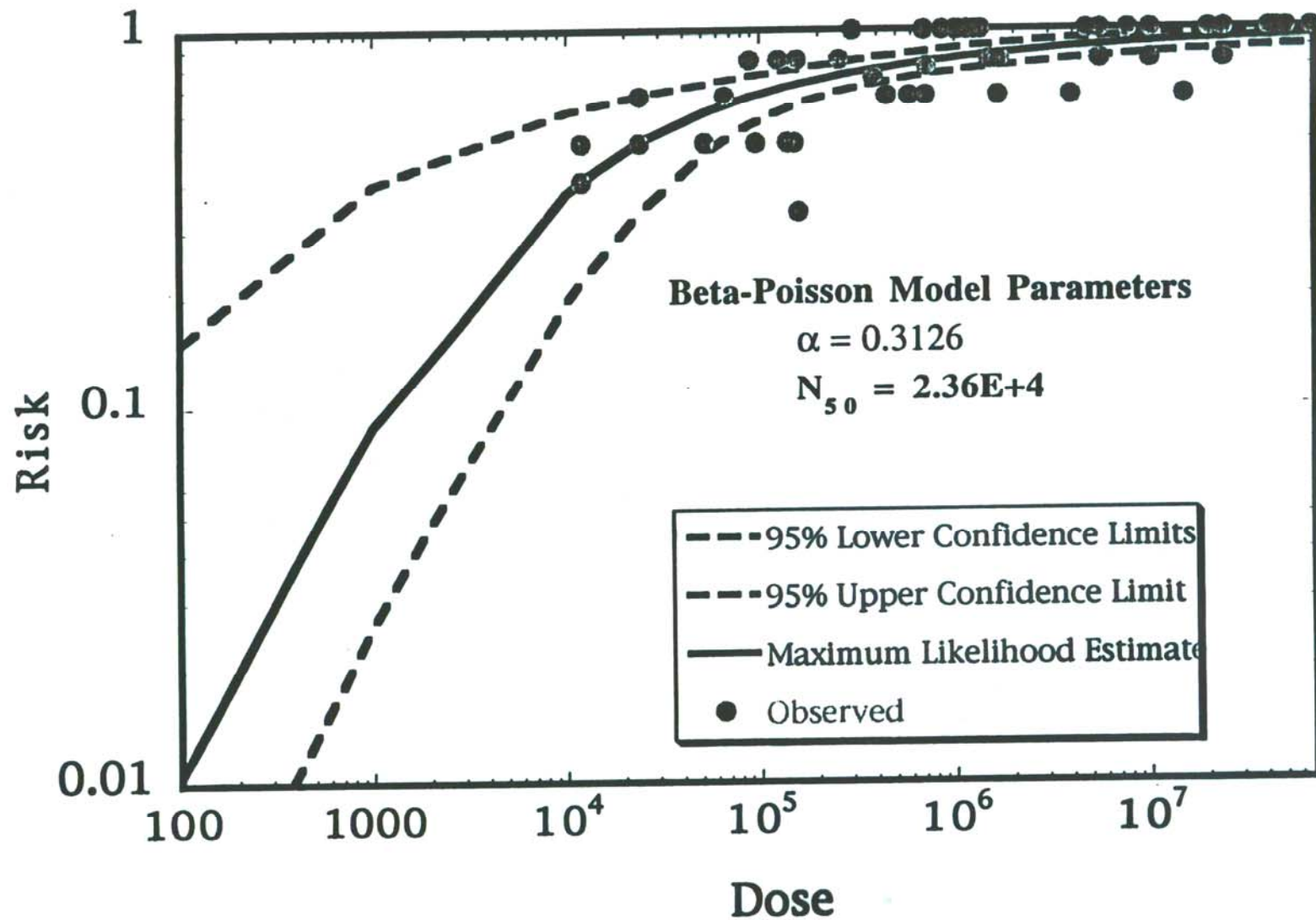
Human volunteers,
C. parvum,

DuPont et al.
(1995)

Okhuysen et al.
(1999)

Potential for probabilistic
modeling of inter-strain
variability (Teunis and
colleagues)





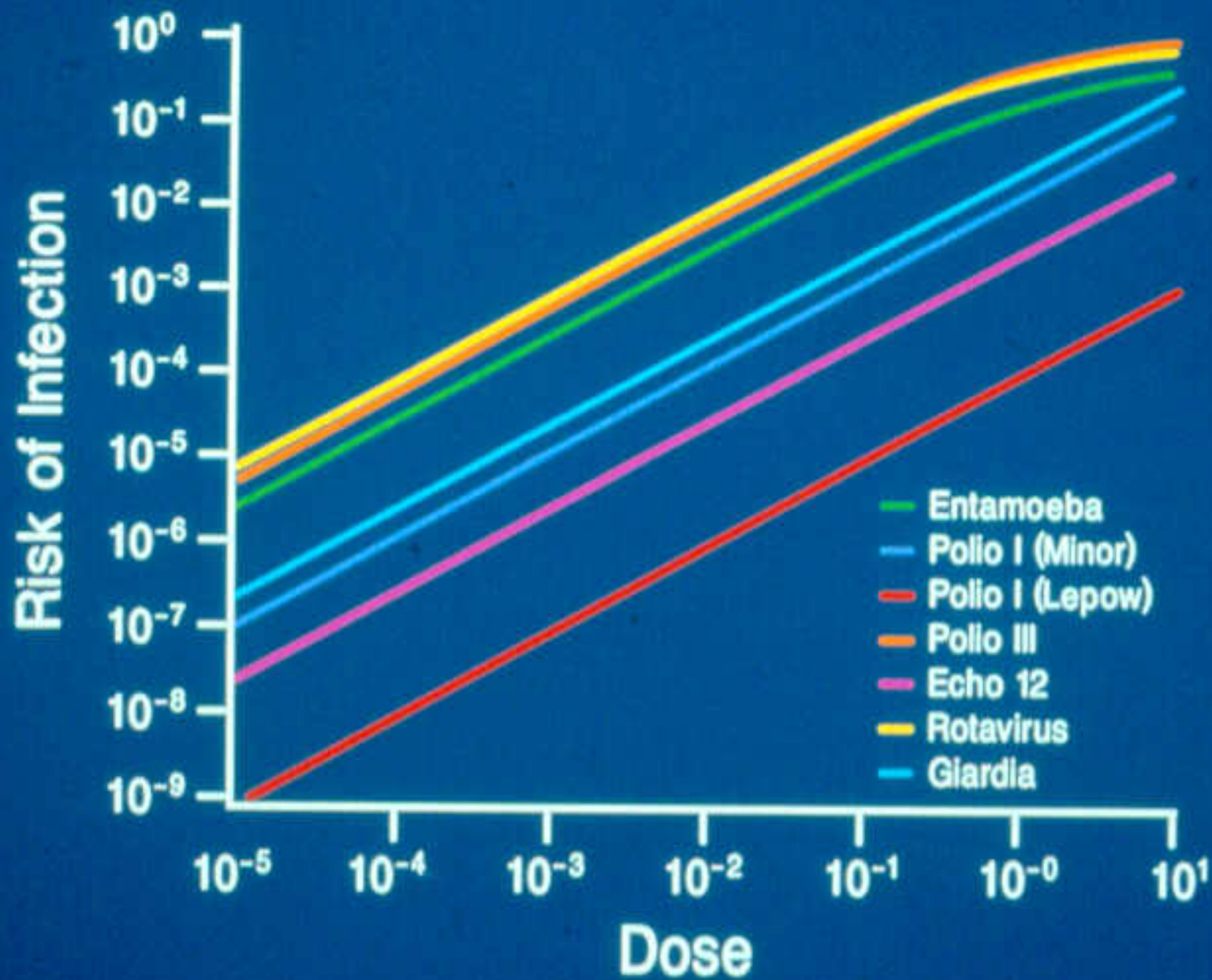
Probability of Infection

| Organism | Best Model | Model Parameters |
|------------------|--------------|--------------------------------------|
| <i>Echovirus</i> | beta-poisson | $\alpha = 0.374$ $\beta = 186.69$ |
| <i>Rotavirus</i> | beta-poisson | $\alpha = 0.26$ $\beta = 0.421$ |
| Adenovirus | exponential | $r = 0.4172$ |
| <i>Polio1</i> | beta-poisson | $\alpha = 0.1097$ $\beta = 1524$ |
| <i>Polio3</i> | beta-poisson | $\alpha = 0.409$ $\beta = 0.788$ |

Models: $P_i = 1 - (1 + N/\beta)^{-\alpha}$ (beta-poisson model)

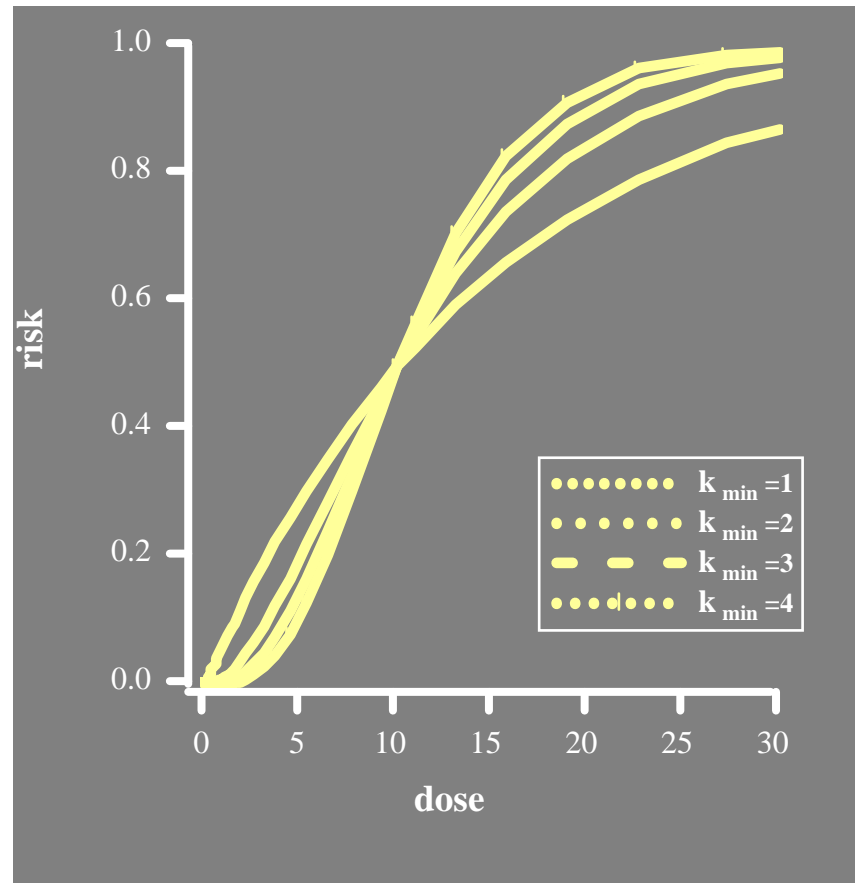
$P_i = 1 - \exp(-rN)$ (exponential)

N=exposure



“Threshold (>1)” Models

- threshold models ($k_{\min} > 1$) yield steeper slopes and non-linear low dose models
- no human data sets yet examined justify these models



Validation - Water, *Cryptosporidium*

- 1993 Milwaukee outbreak (drinking water)
 - 400,000 cases; (21 % attack rate)
 - 21 day duration
-

- estimated daily risk=1.12 %
 - from dose-response (infections), estimate daily intake=2.4 oocysts/day
 - from water consumption==> 1.2 oocysts/L
-

- best estimate(?) from ice measurements, corrected for freezing = 0.79/L

EXPOSURE ASSESSMENT

- Route of Exposure
- Duration of exposure
 - Seconds, hours, minutes
- Number of exposures
 - How many times in a day, month, year
- Degree of exposure
 - Liters of water ingested
 - Liters of air inhaled
 - Grams of food ingested

Watershed assessment, Flow, Transport, Integration with water quality and thus exposure.

Overland



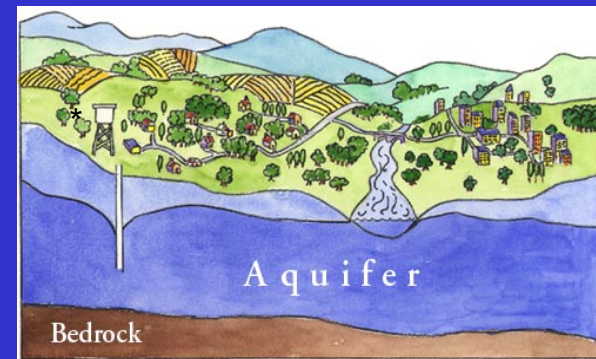
In-Stream



Near-Shore



Sub-Surface



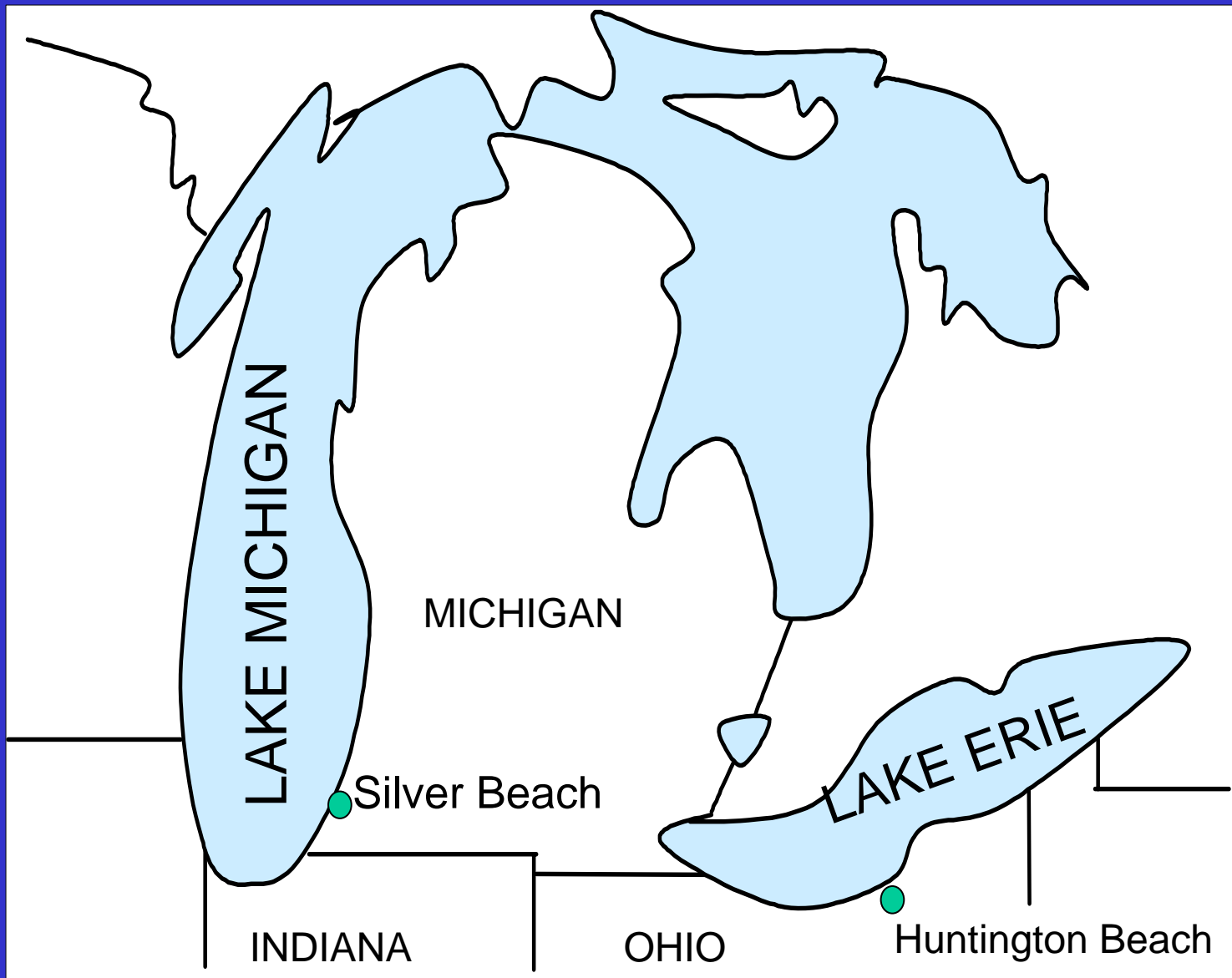
Microbial Source Tracking

- Tools are now available to determine the molecular fingerprint of the fecal pollution.
- Health risks
- Remediation
- Prioritization
- Responsibility



Host Specific Markers are Key to Source Tracking Future

- Bacteroides (genetic approaches PCR)
 - 4/4 sewage; 4/4 human; 4/5 cow (lowest concentration missed) 4/4 dogs however no marker for Birds: Missed 2 samples with dog and 2 with cow that were mixed.
 - E.coli Toxin genes able to detect sewage (4/4).
 - Enteroviruses and Adenoviruses found in 3 of 4 sewage samples.
 - Enterococci ESP marker found in 107/109 human sewage water samples and zero of 80 animal samples.
- USGS blind samples 0/10 animals; 6/8 Human





Location of sampling sites (1-3) at Silver Beach, St Joseph, Michigan (Lake Michigan).

Photo: US Army Corps of Engineers.

esp human marker in Great Lakes recreational beach waters

| | | | Silver Beach, MI | | |
|------------------|--|--|--------------------|------------------------------|-----------------------------------|
| Sampling time | | | Number of swimmers | Mean turbidity (NTU) [SD] | positive for <i>esp</i> marker |
| Morning | | | 17 | 3.2 [3.8] | 5/168 3% (n=276) |
| Afternoon | | | 805 | 3.7 [3.7] | 4/88 4.5% (n=138) |
| Total | | | 822 | | 9/256 |

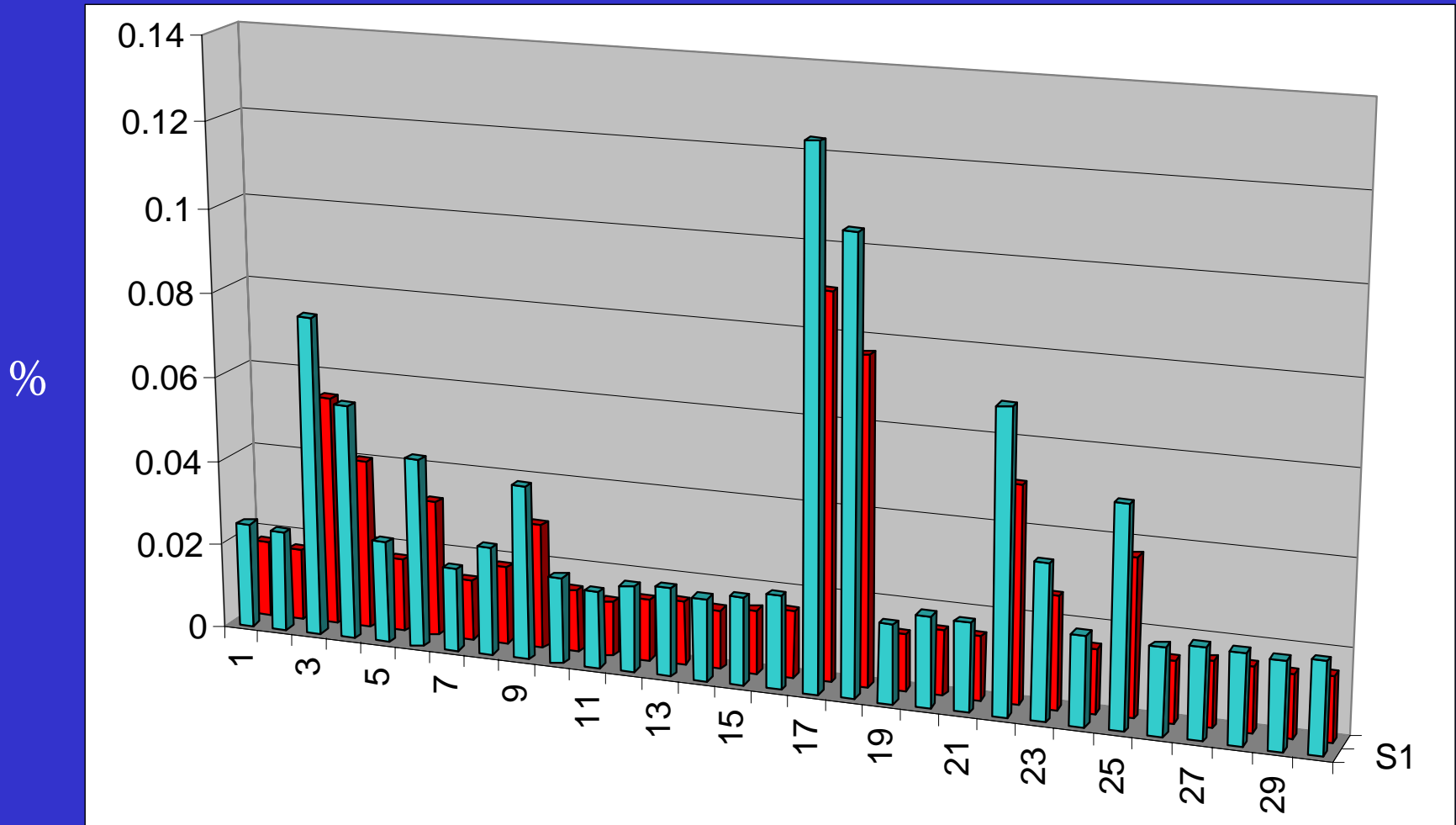
Presence of Enteric Viruses

| | | Silver Beach, MI | | | |
|----------------------|--|-------------------------|-------------------------|-------------|-------------|
| Sampling time | | cell culture +/n | Id by PCR/RT-PCR | | |
| | | | Ad | EV | RV |
| Morning | | 8/15 | 10/11 | 0/11 | 1/11 |
| Afternoon | | 8/15 | 6/9 | 0/9 | 2/9 |
| Total | | 16/30 | 16/20 | 0/20 | 3/20 |

- *esp* marker detected in
~5-9% Lake Michigan samples
- enteric viruses detected in 16/30 samples
from Silver Beach via cell culture
- adenovirus and rotavirus identified in
samples from these beaches by PCR
- EPA found that between 1 and 10% of all
swimmers became ill after swimming at GL
beaches.

Daily Risk of Viral Infections

Based on Rota and Adeno virus models



30 day risk is 7.5 and 14/1000 (0.7 to 1.4%)

Ohio blames groundwater for Lake Erie island outbreak

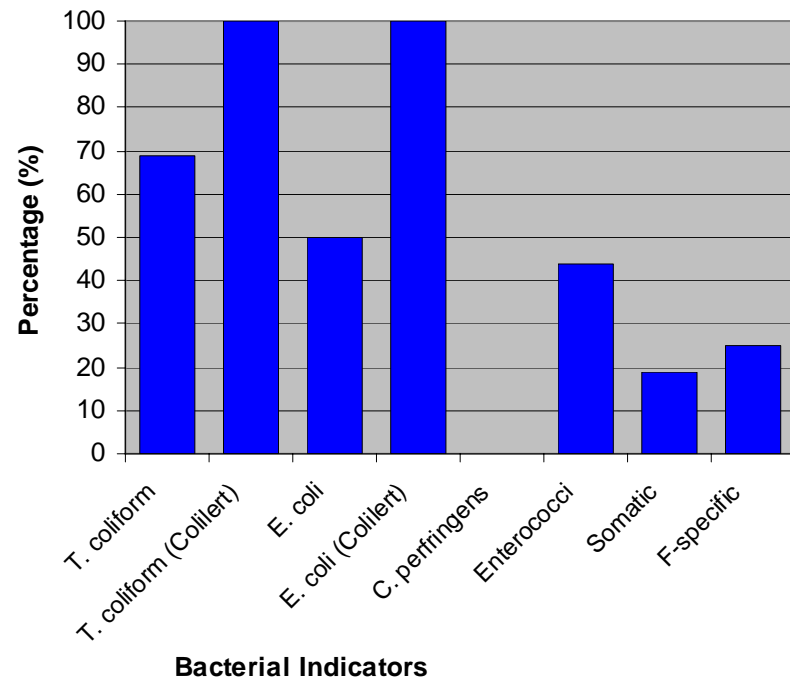
Tuesday, February 22, 2005

ASSOCIATED PRESS

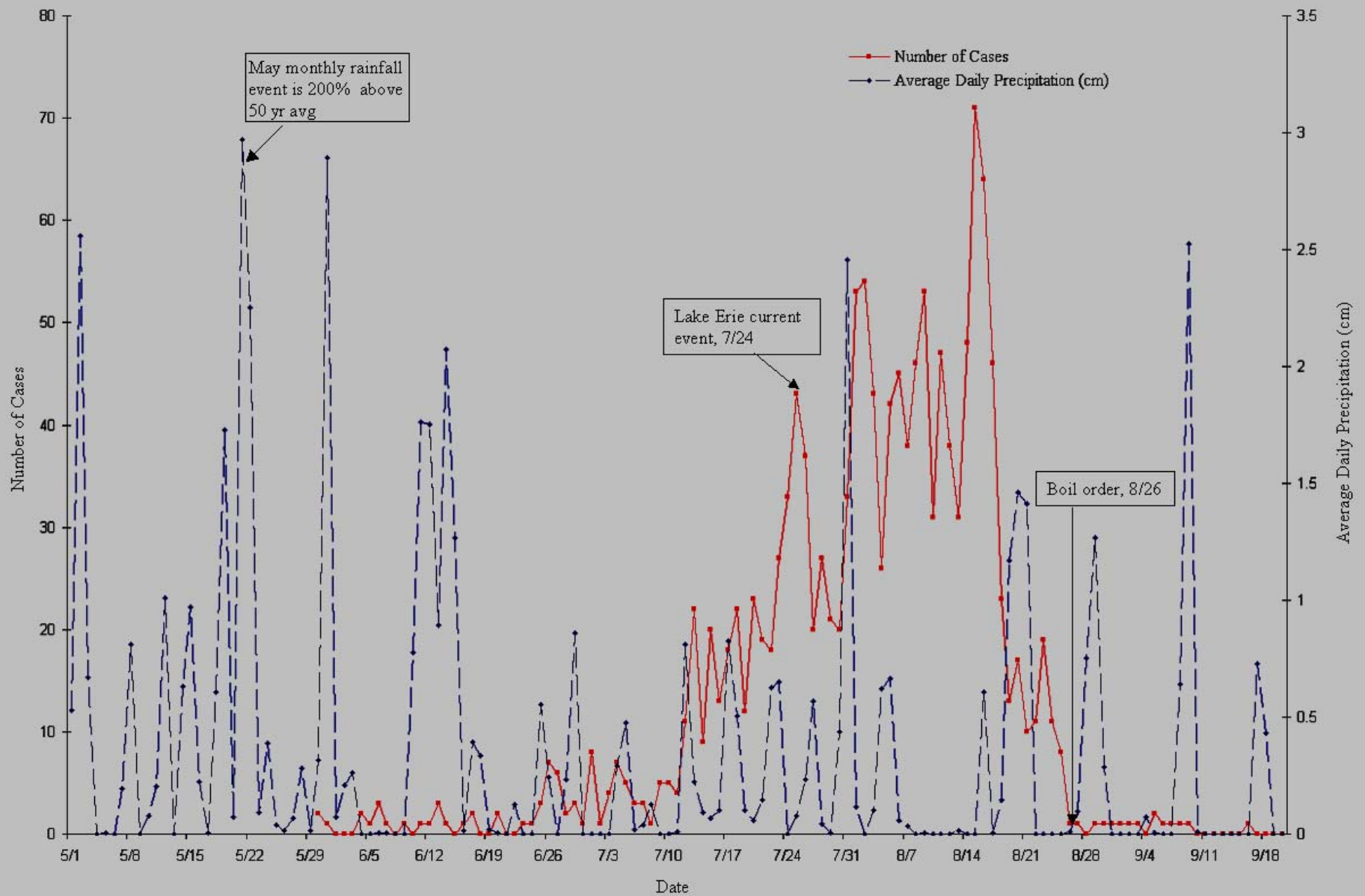
TOLEDO, Ohio -- Widespread groundwater contamination on a Lake Erie resort island was the likely source of illnesses that sickened hundreds last summer, the Ohio health department said Tuesday.

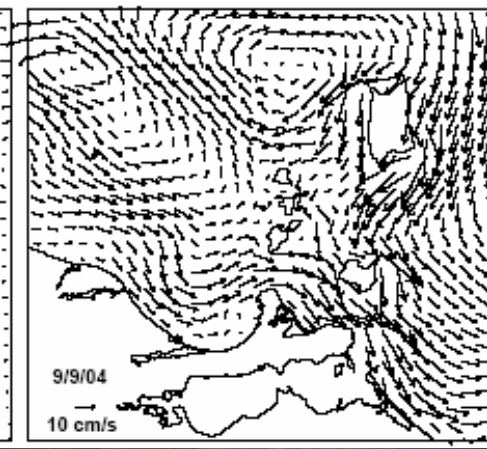
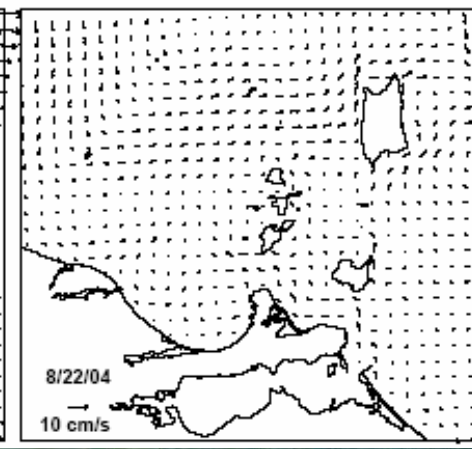
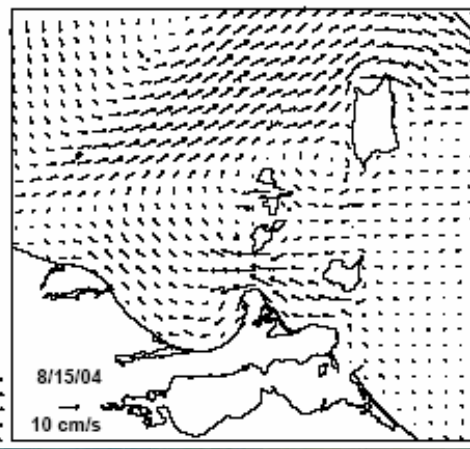
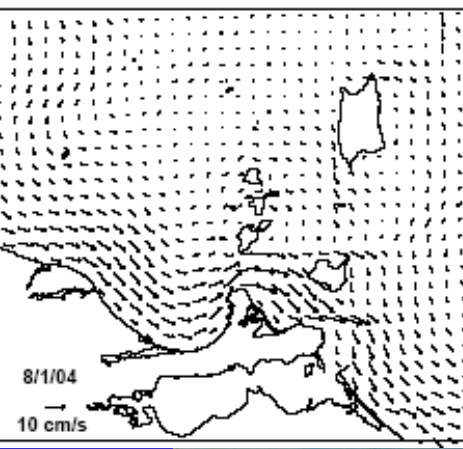
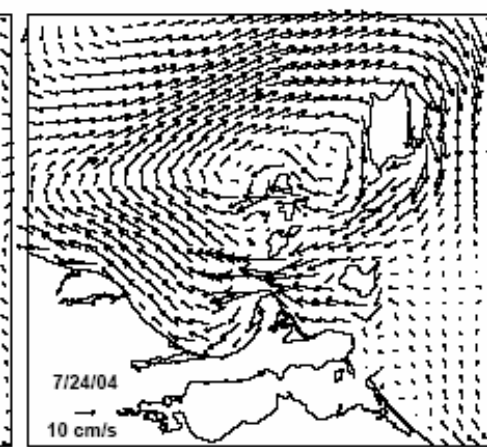
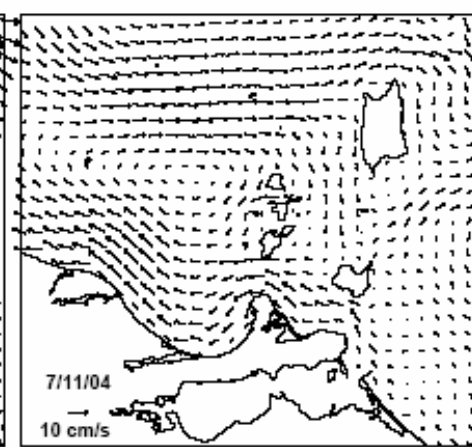
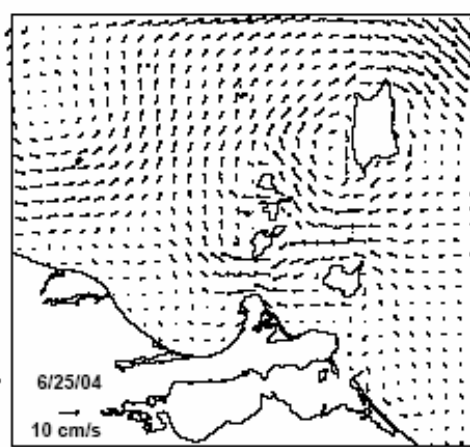
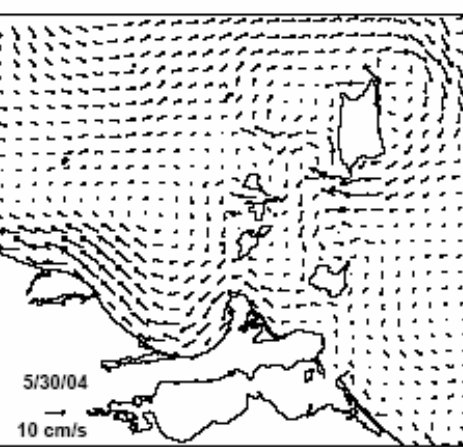
Several sources, including septic tanks, have tainted the South Bass Island's groundwater over a long period, and the contamination may have been worsened last summer because of a season of heavy rains, a health department report said.

The outbreak of gastrointestinal illness sickened about 1,400 tourists and residents, ending the tourist season early for many businesses.



**MSU assisted with the investigation
Identified virus contamination and
potentially a new and emerging bacteria**





South Bass Island
Lake Erie
Sources septic Tanks
& sewage discharges
Massive Ground Water
and Surface Water
Contamination



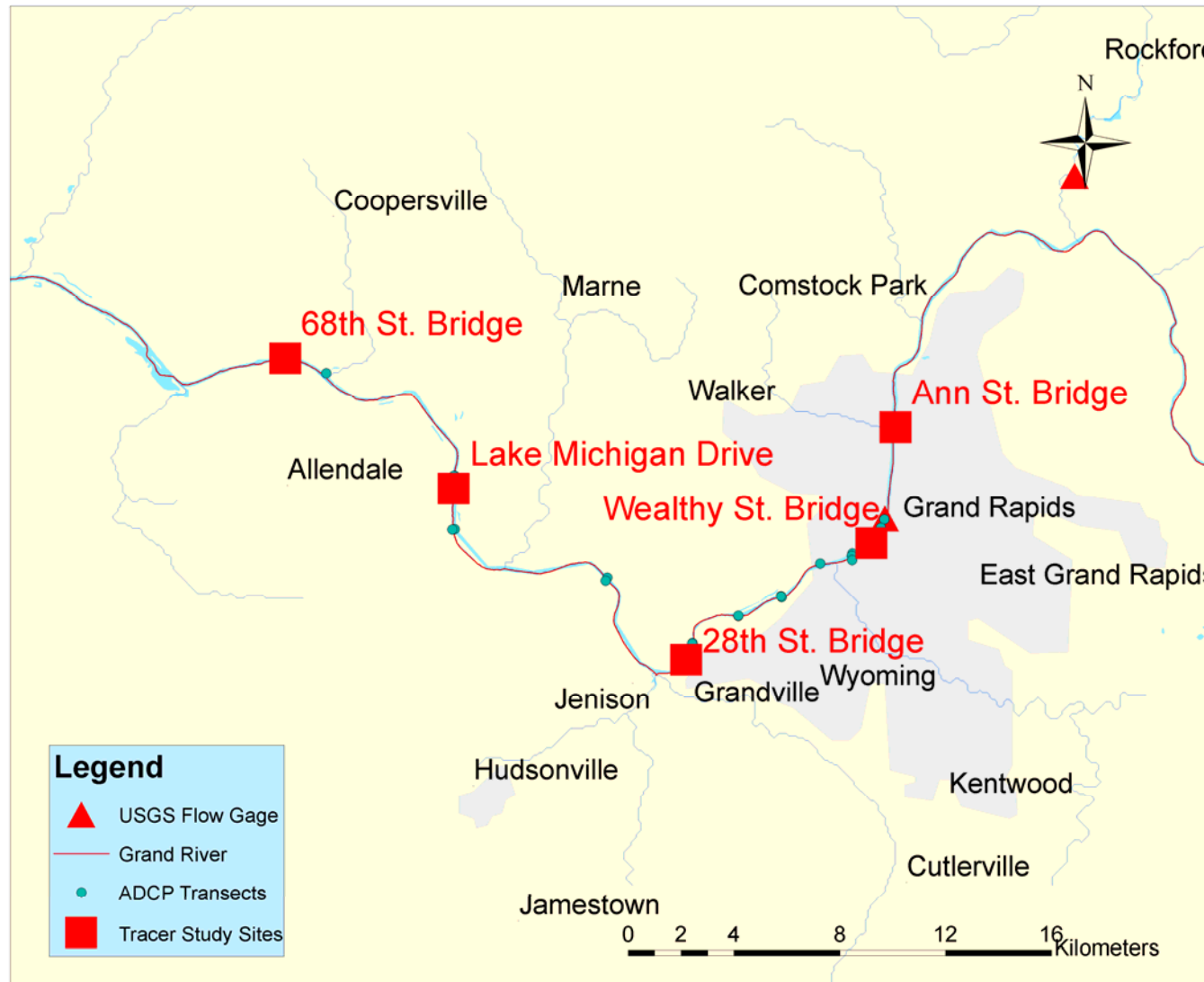
Study Reach

**Examine
Fecal Indicators**

**Examine
Sediments**

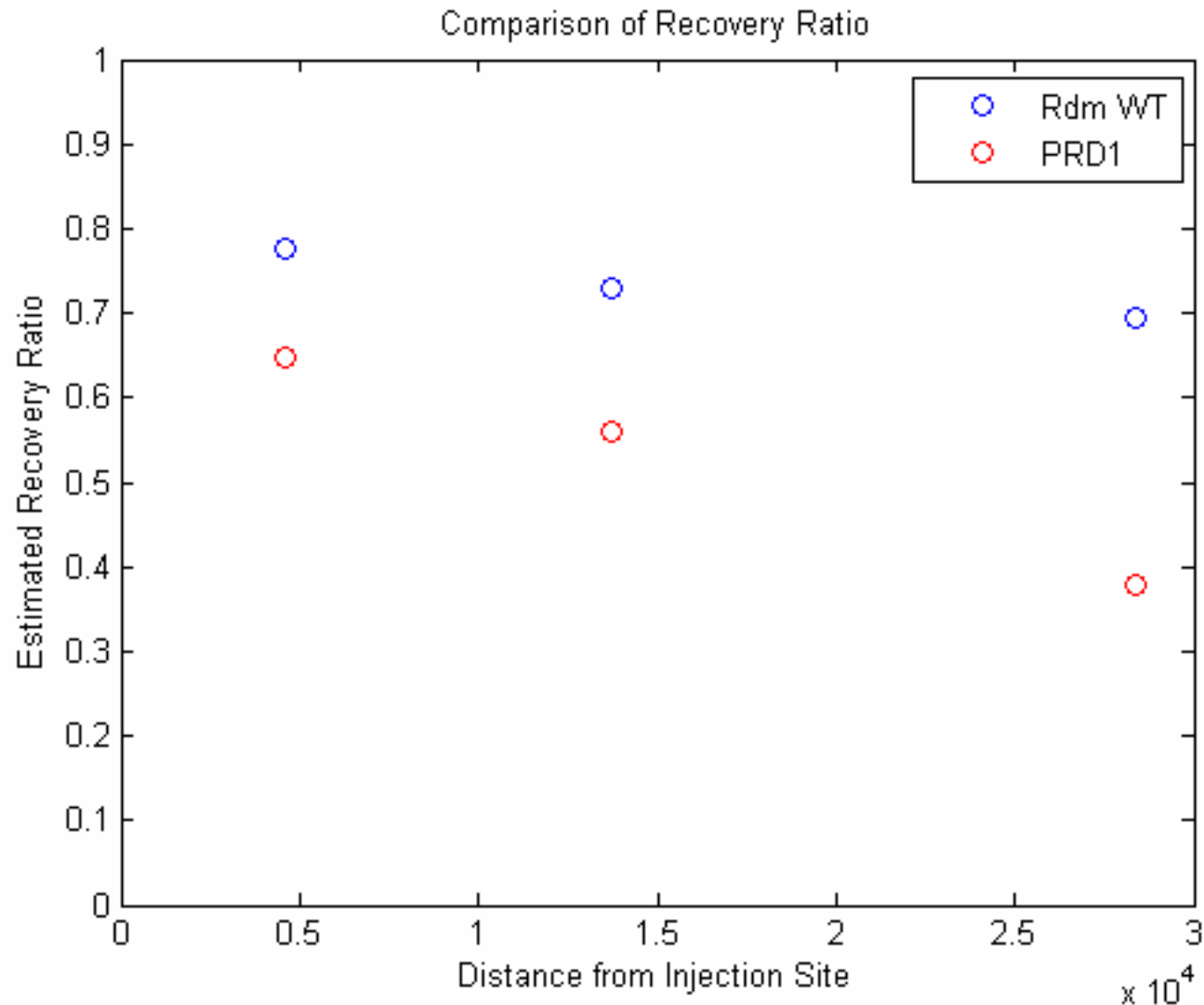
**Examine
Transport from
River to Beach**

**Monitoring &
Tracer Studies**

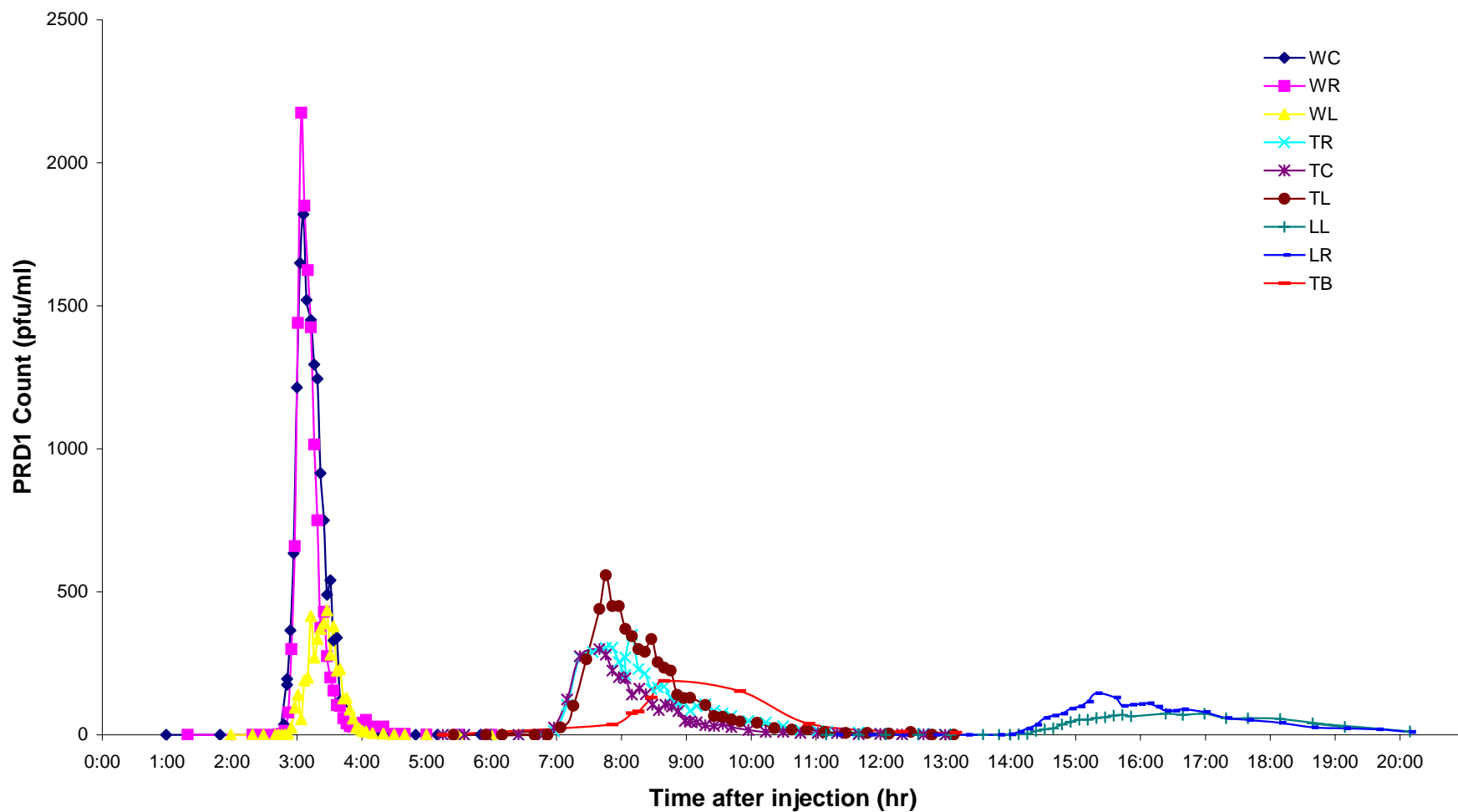


Rhodamine vs PRD1

- Recovery Ratio (Estimated from Average Breakthrough Curve)



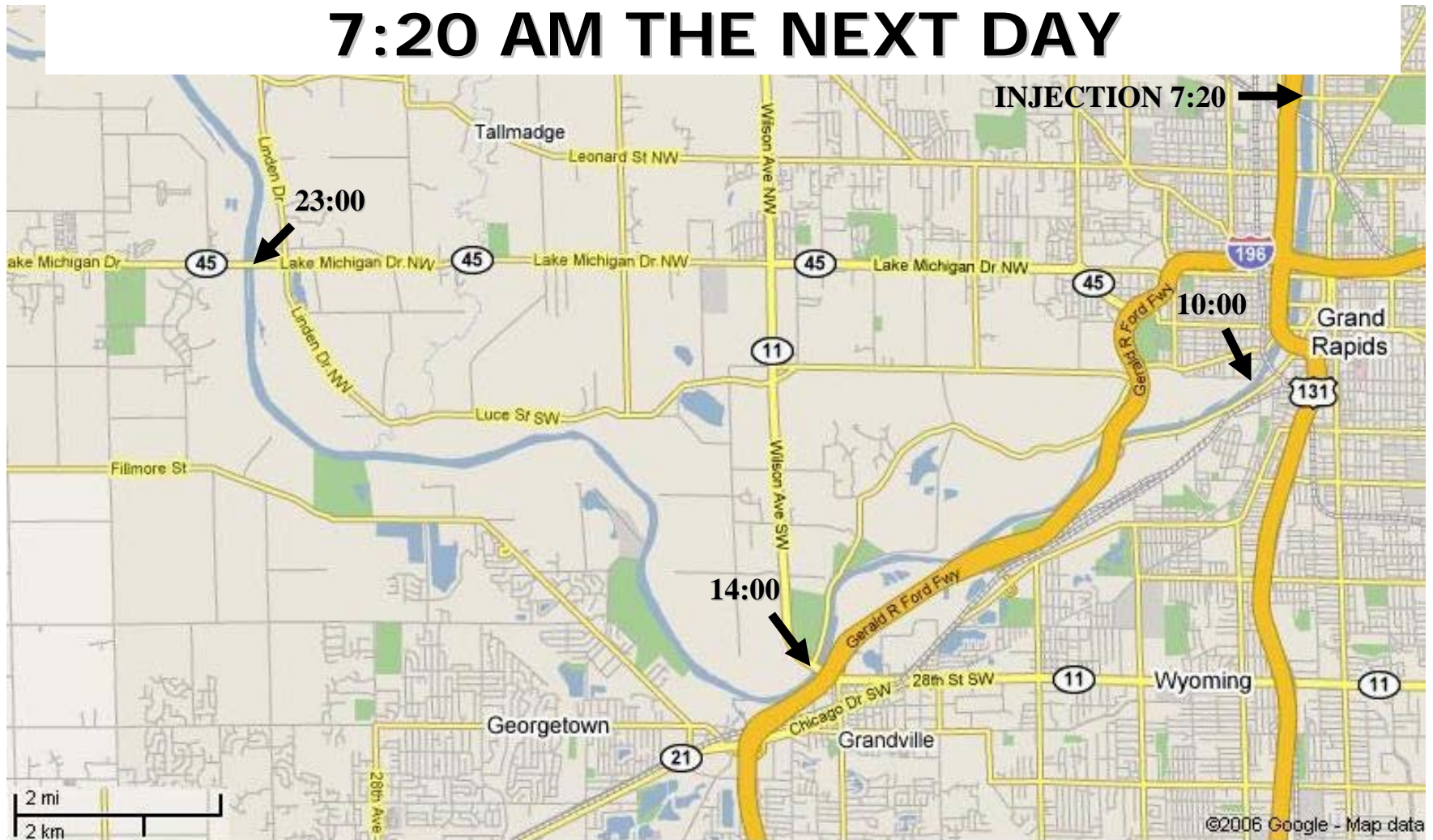
PRD-1



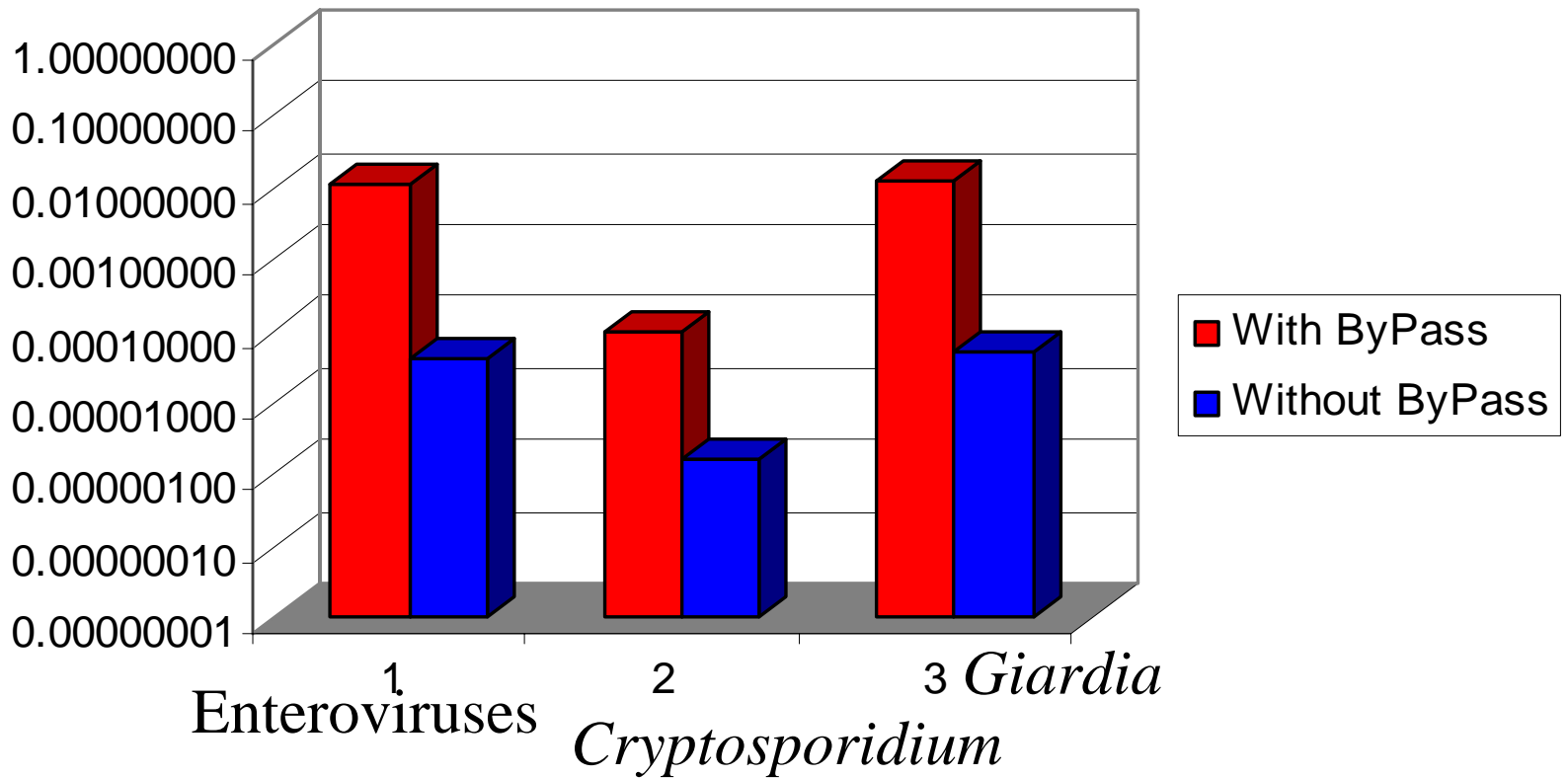
Arrival Time at Each Site

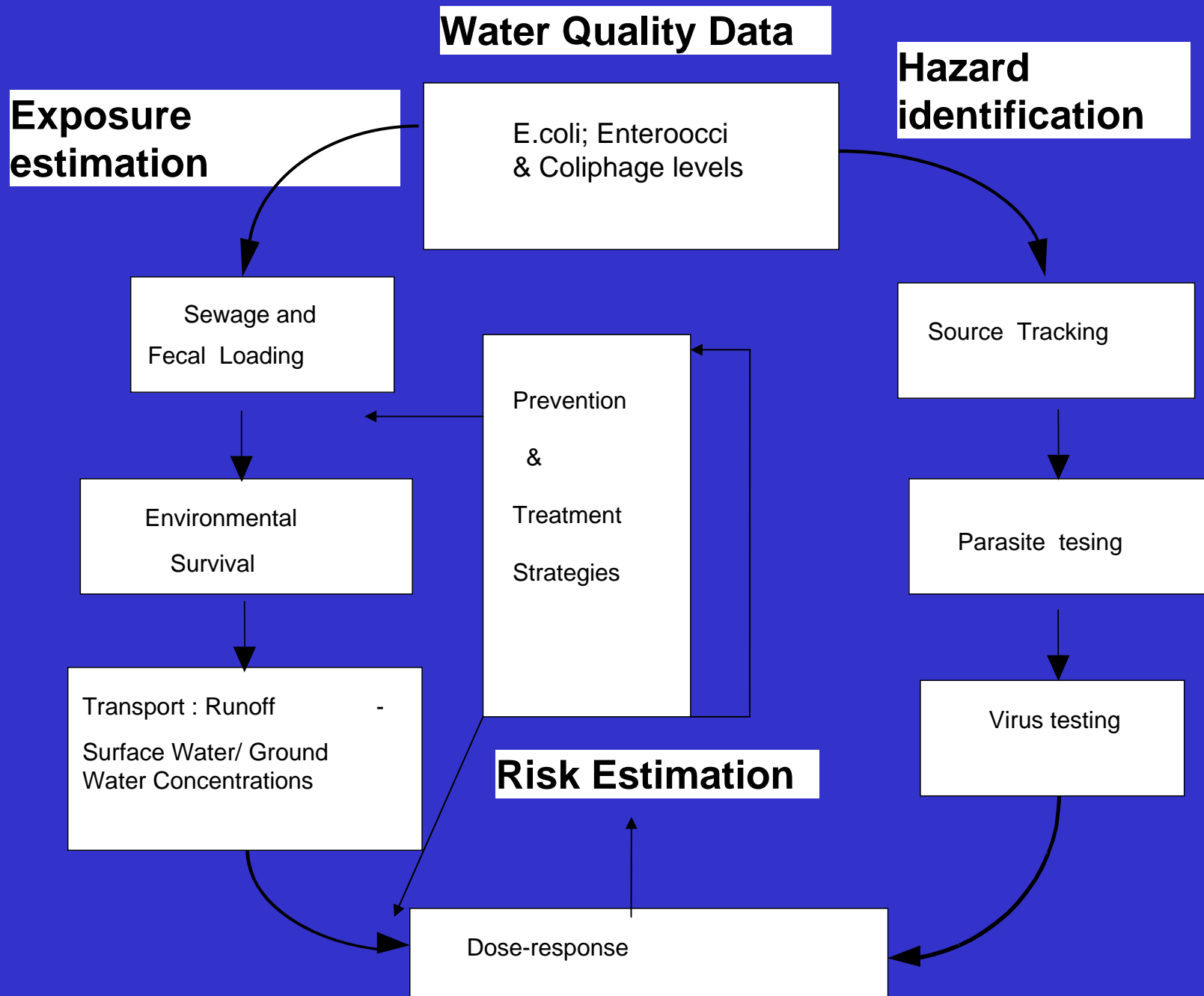
ESTIMATED ARRIVAL AT LK MICHIGAN

7:20 AM THE NEXT DAY



Risk Estimates



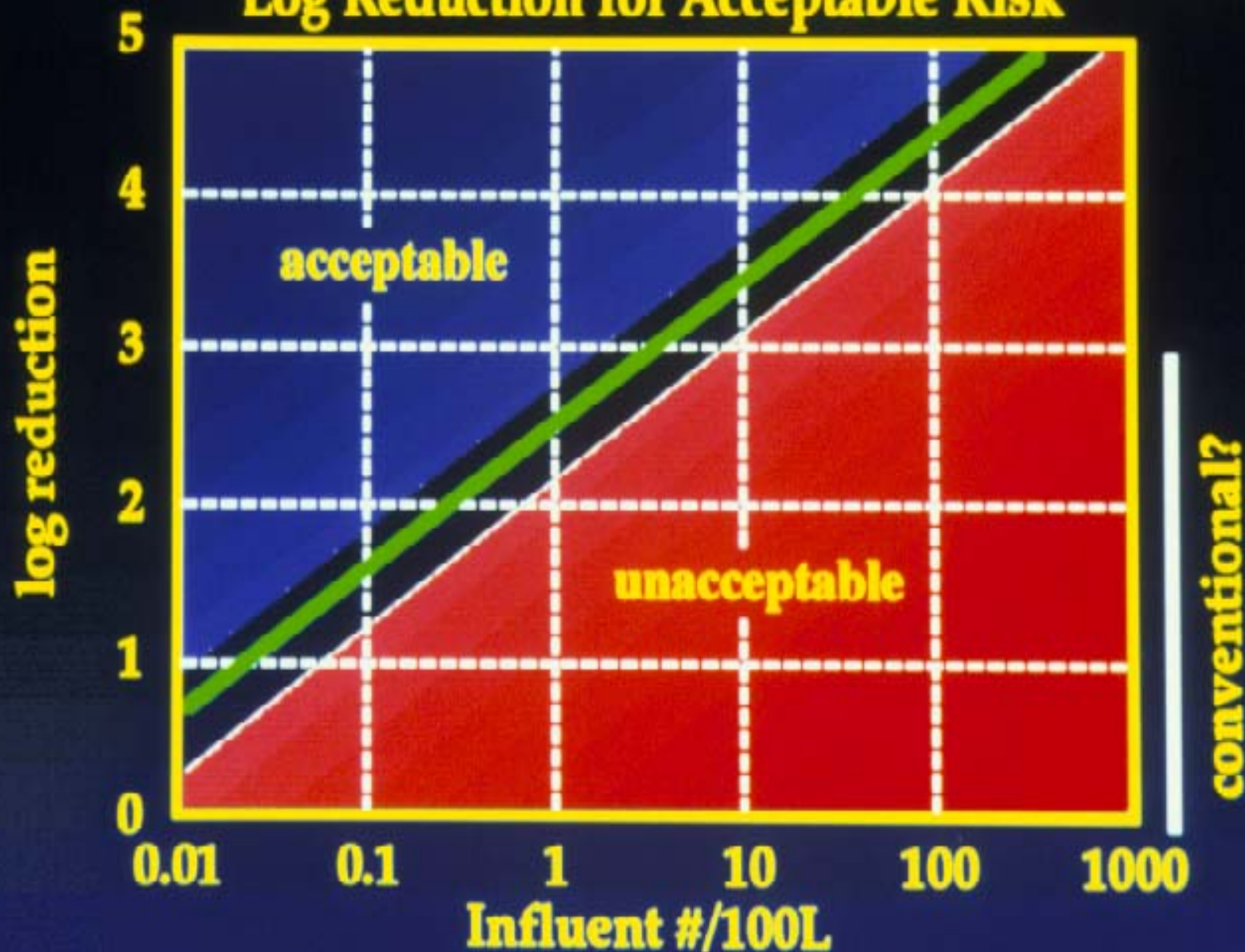


Applications for Microbial Risk Assessment

- Establish policies for protection of health using standards or performance based criteria
- Compare risks
- Evaluate alternative solutions
- Prioritize risks
- Identify scientific data gaps
- Develop protocols for monitoring

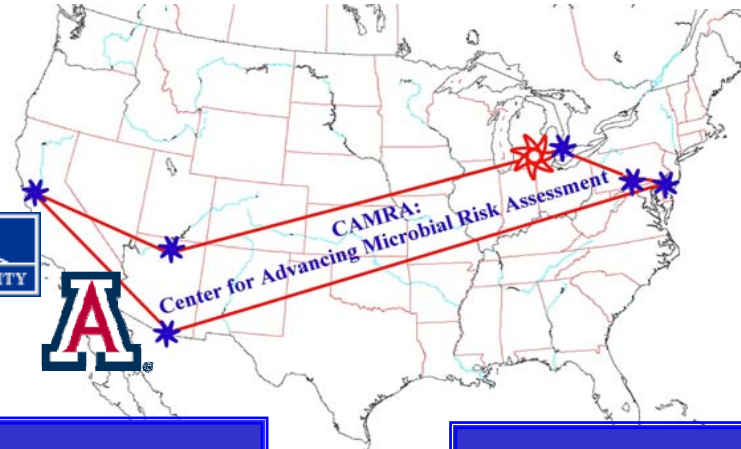
Treatment vs. Influent: Endemic Risk

Log Reduction for Acceptable Risk





Carnegie Mellon

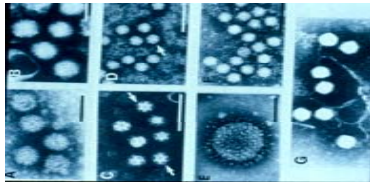


to build a national network for microbial risk knowledge management, learning and transfer, for the community of scientists, and students via educational programs and community of professionals in the field and in our communities.

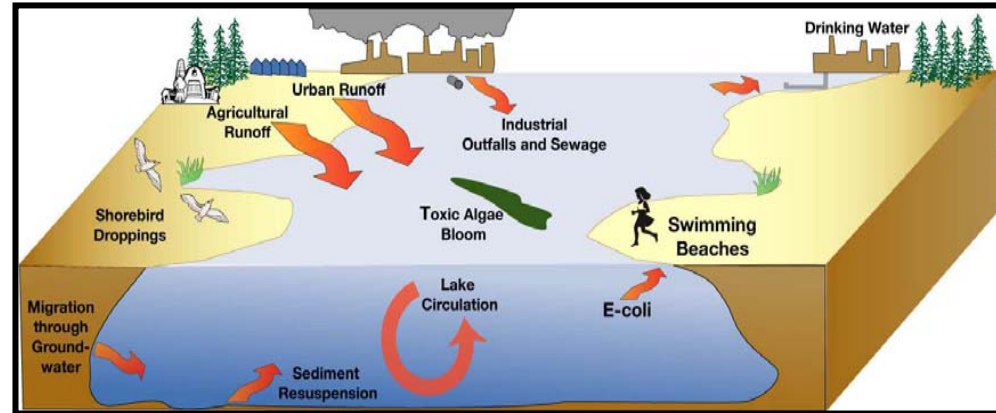
to develop models, tools and information that will be used in a credible risk assessment framework to reduce or eliminate health impacts from deliberate use of biological agents of concern in the indoor and outdoor environment.

ADVANCING MICROBIAL RISK ASSESSMENT

Hazards

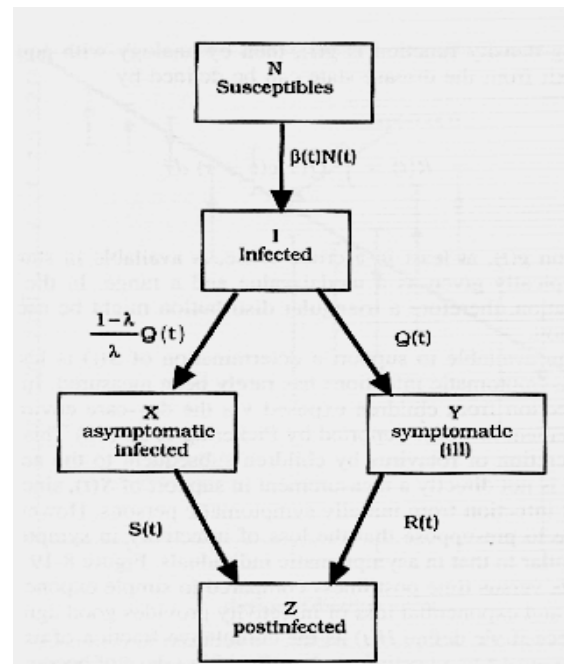
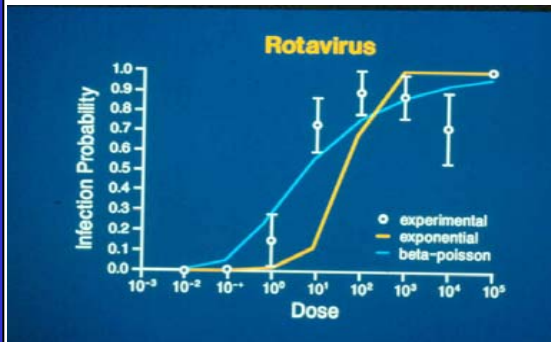


Exposure

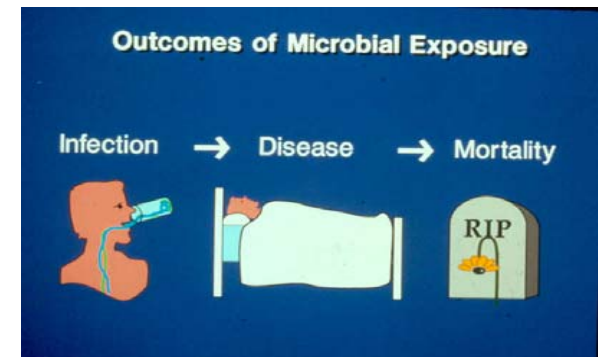


Disease Dynamics

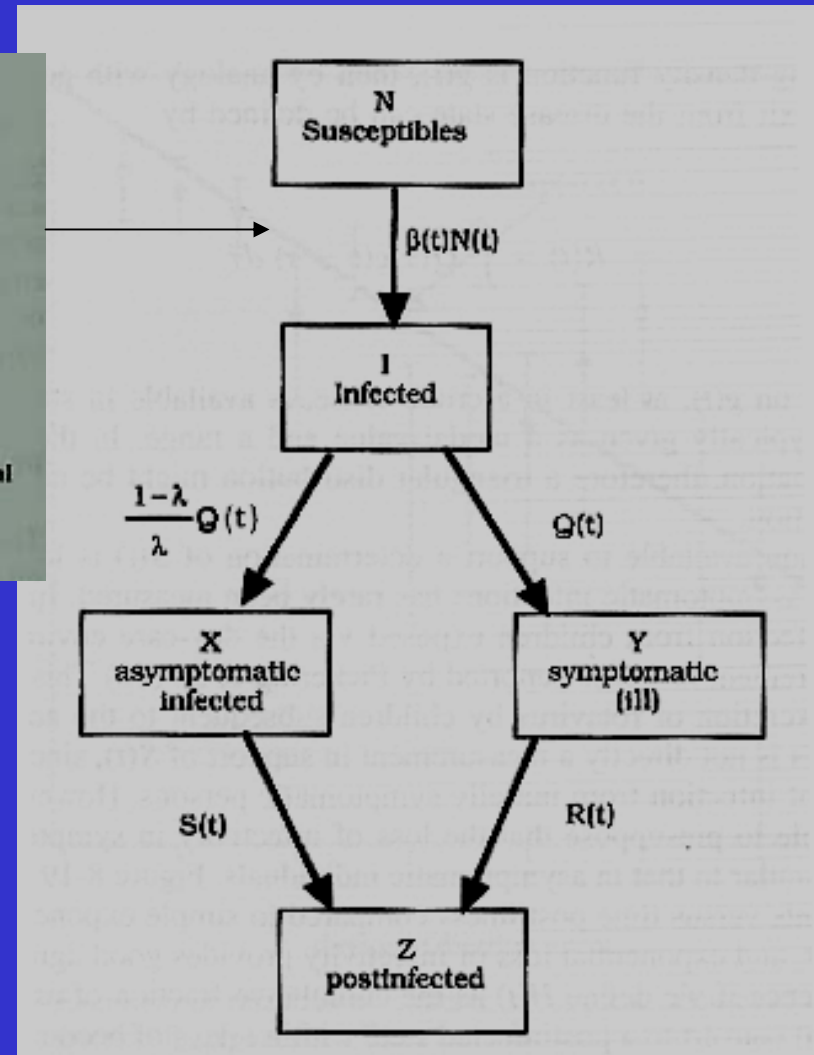
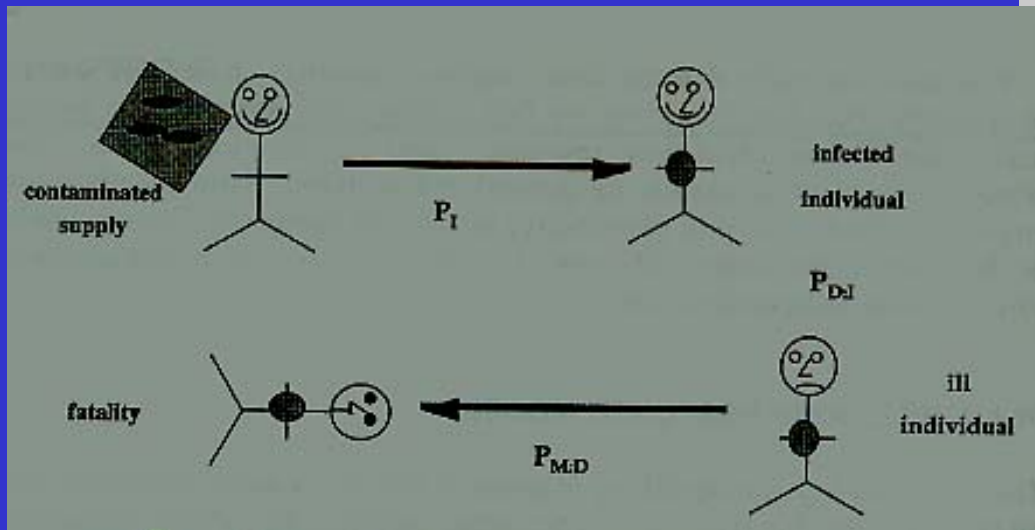
Dose-response



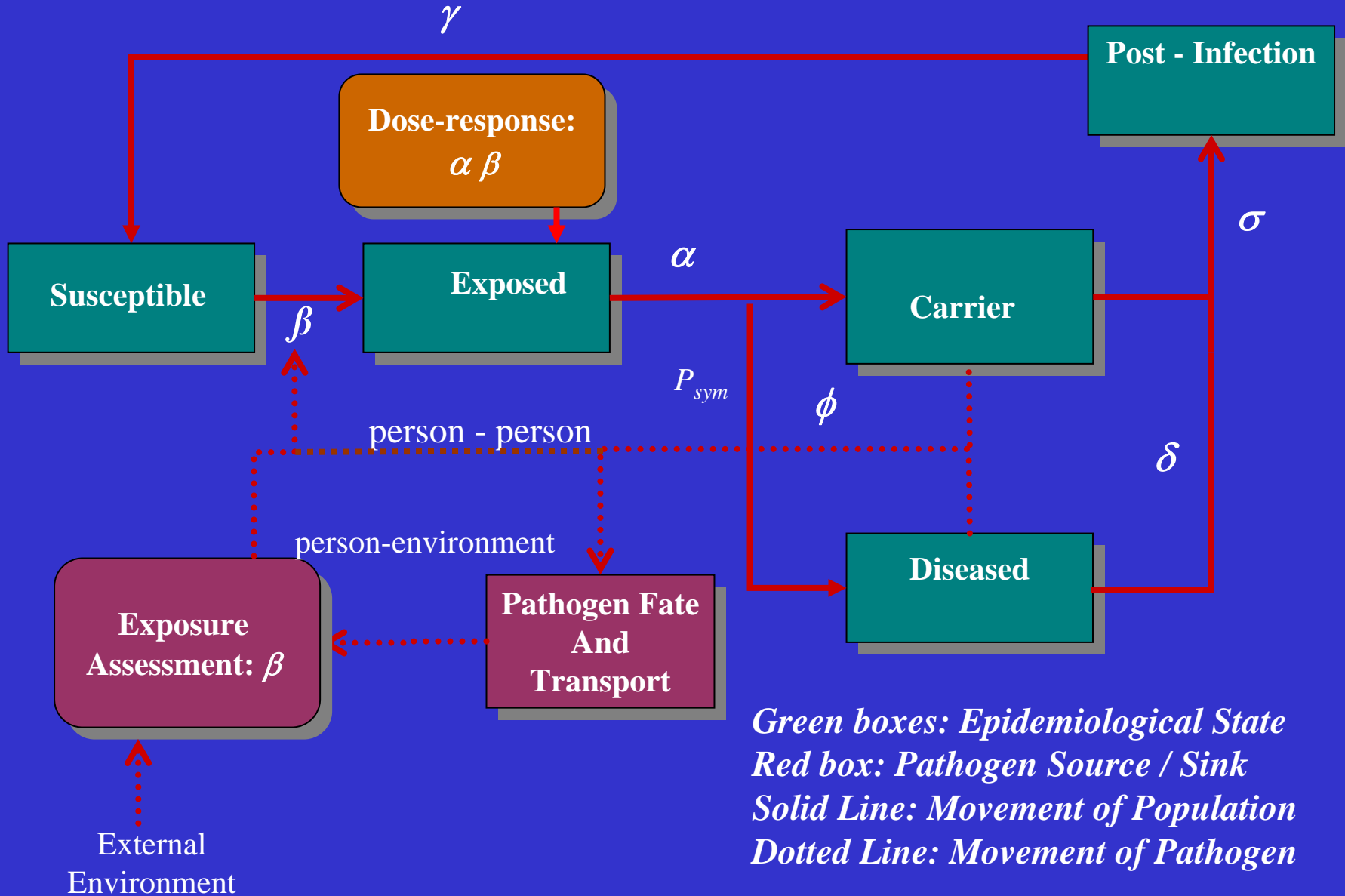
Risk Characterization

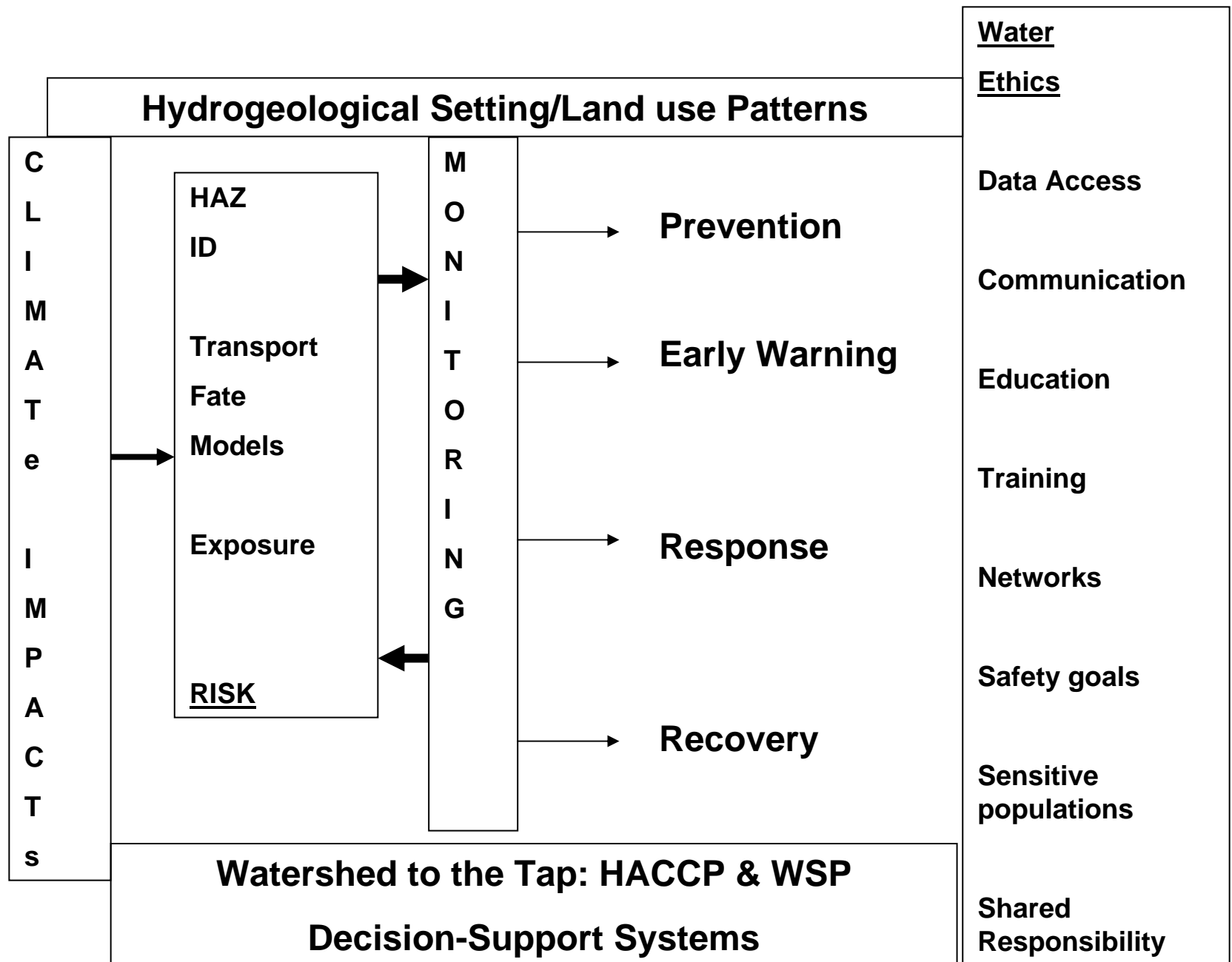


Must Link probability of infection to population models & address the environmental exposure



Interaction between Disease Transmission and the Environment





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