

#### Out of Sight, Out of Mind -Typhoid Fever Surveillance and Response in Africa

8th Symposium on Invasive Salmonelloses, March 2, 2013, Eric Mintz, MD, MPH



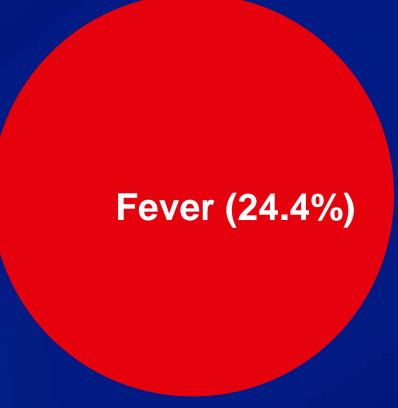
#### Outline

- Prevalence of fever in Africa
- Prevalence of typhoid fever in Africa
- Enteric fever among visitors to Africa
- Outbreaks of typhoid fever in Africa
- Antimicrobial susceptibility of S. Typhi in Africa

# Acute respiratory illness, diarrhea, and fever in the past 2 weeks, western Kenya, 2005 (n=6,223)

Acute respiratory illness (6.1%)

Diarrhea (6.3%)

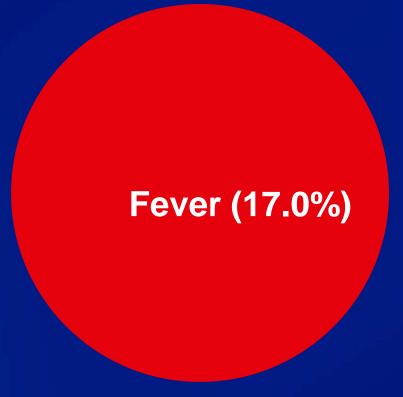


Burton DC, et al. J Health Popul Nutr 2011; 29: 61-70



Acute respiratory illness (7.7%)

Diarrhea (10.5%)



Breiman RF, et al. J Health Popul Nutr 2011; 29: 123-33

#### Typhoid Fever in Kibera Settlement, Nairobi, Kenya, 2007 - 2009

- Salmonella Typhi isolated by blood culture from
  - 10.4% of patients presenting with fever
  - 7.7% of patients presenting with ARI
- Crude rate = 247 cases per 1000,000 person years
- Highest rates in children
  - 5-9 years old (596 per 100,000 pyo)
  - 2-4 years old (521 cases per 100,000 pyo)
- Crude rate in Lwak only 29 per 100,000 pyo

Population-Based Incidence of Typhoid Fever in an Urban Informal Settlement and a Rural Area in Kenya: Implications for Typhoid Vaccine Use in Africa. Breiman R, et al. PLoS ONE. 2012. Vol 7, Issue 1, e29119.

#### Estimated Burden of Typhoid Fever, 2000



Crude incidence in Africa: 50 cases per 100,000 people per year

Global adjusted total: 21.5 million illnesses 200,000 deaths

The global burden of typhoid fever. Crump JA, Luby SP, Mintz ED. Bull WHO. 2004;82:346-353.

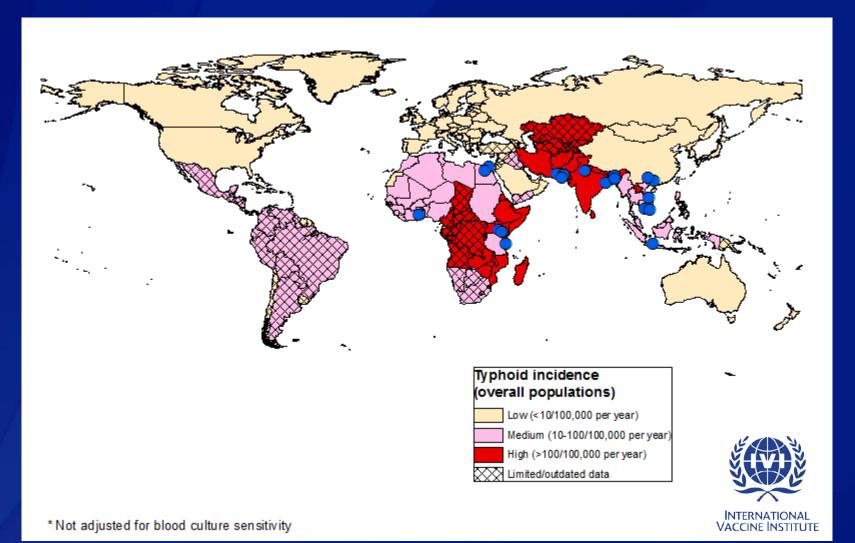
#### Estimated Burden of Typhoid Fever, 2010

- Global adjusted total
  - -26.9 million episodes of typhoid fever

- Africa
  - -Crude incidence of 362 cases per 100,000 persons per year

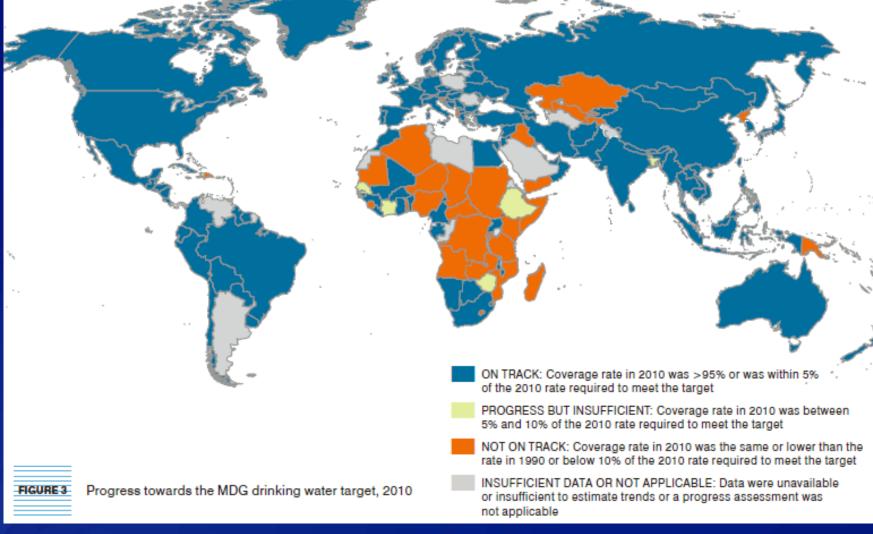
Typhoid fever and paratyphoid fever: systematic review to estimate global morbidity and mortality, for 2010 Buckle GC, Fischer Walker CL, Black RE. Journal of Global Health. 2012; 2: doi 10.7189/jogh.02.010401.

#### Revised Map of Typhoid Fever Incidence, 2010

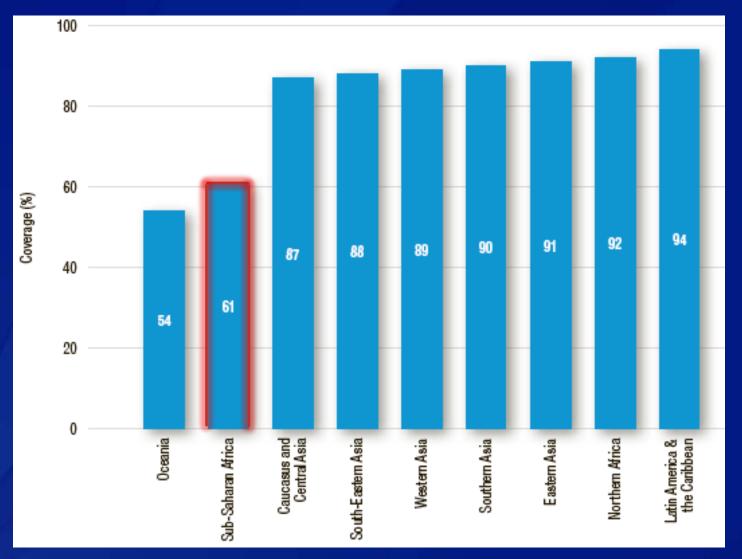


Brian Maskery and colleagues, International Vaccine Institute, Unpublished Data

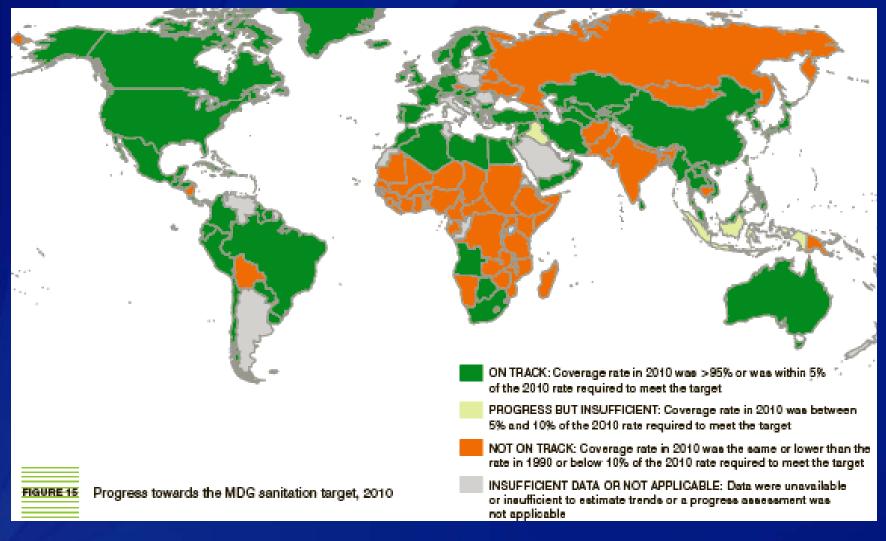
#### Progress towards MDG for Water by Country, 2010



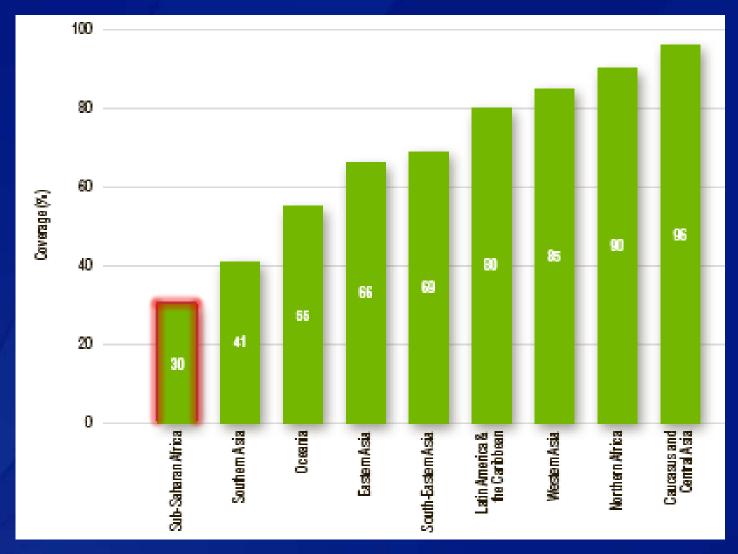
#### Coverage with Improved Water Sources by Region, 2010



#### Progress towards MDG for Sanitation by Country, 2010



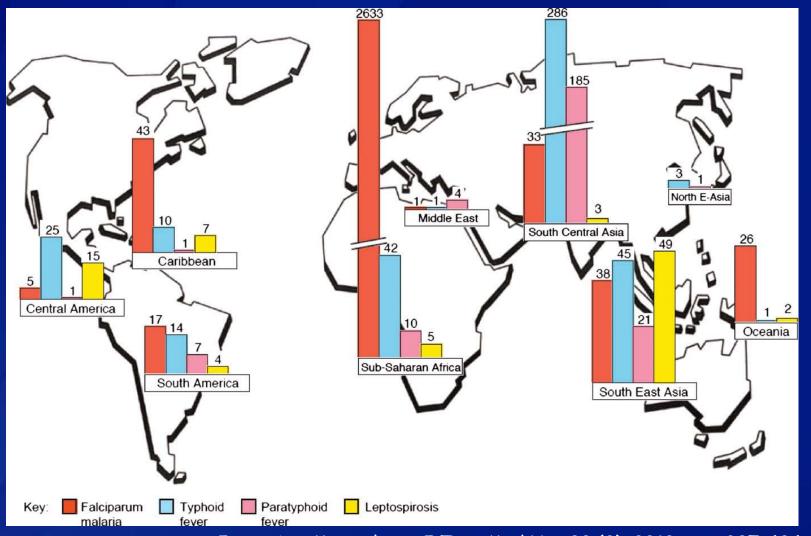
#### Coverage with Improved Sanitation by Region, 2010



#### Typhoid Fever in Travelers, 1996 -2011

- GeoSentinel network of 57 travel and tropical medicine clinics in 26 countries
- Identified 3,655 patients with acute, life-threatening febrile illnesses
  - 428 (11.7%) had typhoid fever
  - 233 (6.4%) had paratyphoid fever
- About half of all patients with enteric fever were hospitalized; none died

## Cases of selected acute life-threatening diseases by exposure region among travelers to the tropics, 1996-2011 (n=82,825)



Jensenius, M. et al. Am J Trop Med Hyg 88 (2), 2013, pps. 397-404

## Typhoid Fever Outbreaks in ProMED, 1994 - 2013

- Search yielded 223 matches \_\_\_\_\_\_\_ 106 events
- 38 (36%) reported typhoid outbreaks in Africa
  - 1994-2000 = 1 (1 country)
  - 2001-2007 = 23 (10 countries)
  - $\bullet$  2008-2013 = 13 (8 countries)
- Reported cases ranged from 1 to 42,564
- Reported deaths ranged from 0 50
- Recent outbreaks occurred in urban settings (Lusaka, Zambia and Harare, Zimbabwe)

# Typhoid Fever Outbreaks in ProMED, 1994 - 2013

## Characteristics of Selected Outbreaks of Typhoid Fever in Africa

Country	Dates	Cases	Severity
DRC	10/2004 - 1/2005	42,564 (ProMED)	Int. Perf. 4.6% CFR 1% (WHO)
Malawi	3/2009 - 11/2009	303 (46 confirmed)	Neurologic symptoms in 13%
Uganda	12/2007 – 7/2009	577 <b>-</b> 24,900	Int. Perf. ~ 5% - 8% Outbreak is ongoing

An outbreak of peritonitis caused by multidrug resistant *Salmonella* Typhi in Kinshasa, Democratic Republic of Congo. Muyembe-Tamfum J, et al. 2009. Travel Medicine and Infect Dis;7:40-43

Multidrug-Resistant Typhoid Fever With Neurologic Findings on the Malawi-Mozambique Border. Lutterloh E, et al. 2012 CID; 54:1100-6 and also Neurologic Manifestations Associated with an Outbreak of Typhoid Fever, Malawi - Mozambique, 2009. Sejvar J, et al. 2012. PLoS ONE; Vol 7 (12): e46099

A Large Outbreak of Typhoid Fever Associated With a High Rate of Intestinal Perforation in Kasese District, Uganda, 2008–2009. Neil K, et al. 2012. CID; 54:1091-9

### Global Burden of Typhoid Resistance

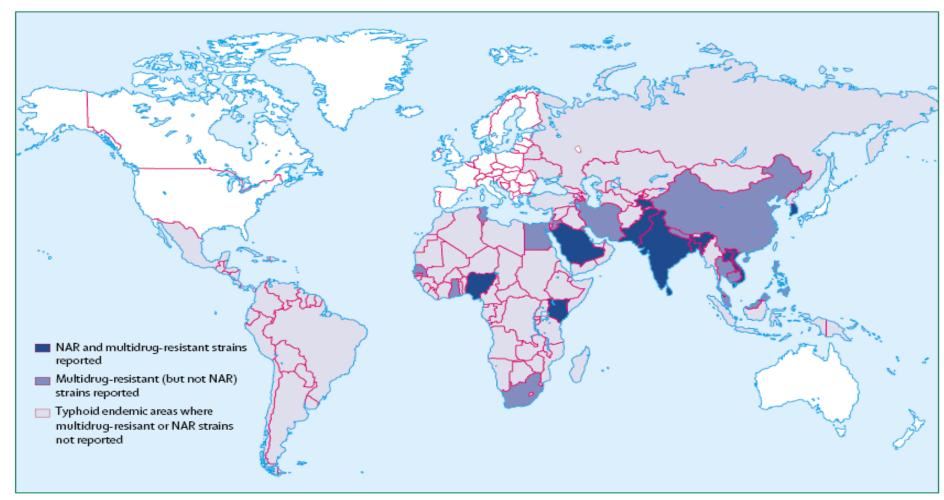


Figure: Global distribution of antimicrobial resistance in Styphi (1990–2004)

Adapted from Parry and colleagues<sup>55</sup> and updated on basis of data from past 3 years.

# Evolution of Antimicrobial Resistance among 5. Typhi Strains in Uganda

Dates	Number	% Pan Suscept	% MDR	% Nal R & Cip I
March-April, 2009	21	19	76	0
Oct – Dec, 2011	18	11	83	6
Aug – Oct, 2012	7	14	71	86

# Evolution of Antimicrobial Resistance among 5. Typhi Strains in Africa

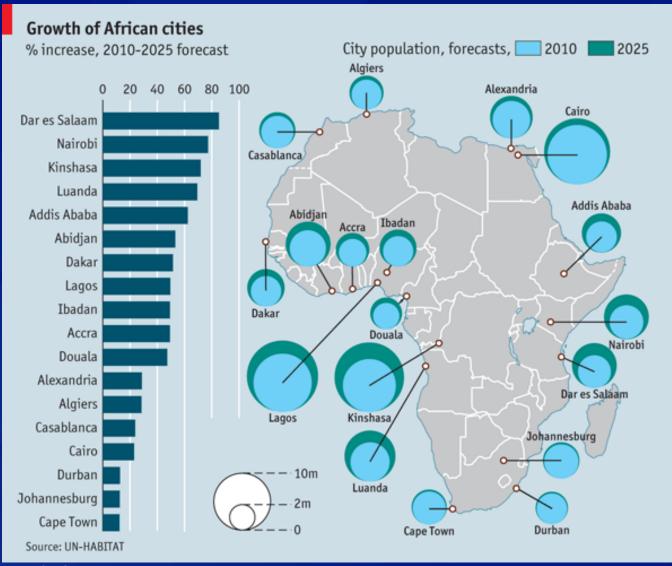
- No systematically collected data available
- Some data from individual sites
  - Nalidixic acid resistance and DCS in DRC
  - NAR and DCS in Kibera, Kenya
  - Abuja, Nigeria ~ 2-3% NAR
  - Agogo, Ghana NAR strains detected.
- Widespread FQ-resistance likely to follow

## Thank you

The findings and conclusions in this presentation are those of the author and do not necessarily represent the views of the Centers for Disease Control and Prevention



#### Projected Urbanization in Africa, 2010 - 2025



The urbanization of Africa: growth areas. Graphic Detail, The Economist, December 13, 2010