We are facing a world without effective antibiotics-

The urgent need for collective action

Otto Cars

Professor Infectious Disease, Uppsala University
Director, ReAct – Action on Antibiotic Resistance
Uppsala University, Sweden

www.reactgroup.org



Antimicrobial resistance or Antibiotic Resistance - a semantical confusion?

Antimicrobial resistance (AMR)

Antiviral agents

(e.g drugs for **HIV**)

Antiparasitiic agents

(e.g drugs for malaria)

ABR

Antibacterial agents = Antibiotics (e.g. drugs for **tuberculosis**)

and

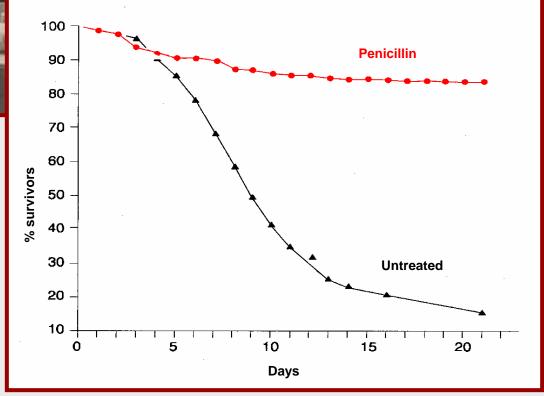
other bacterial diseases ...

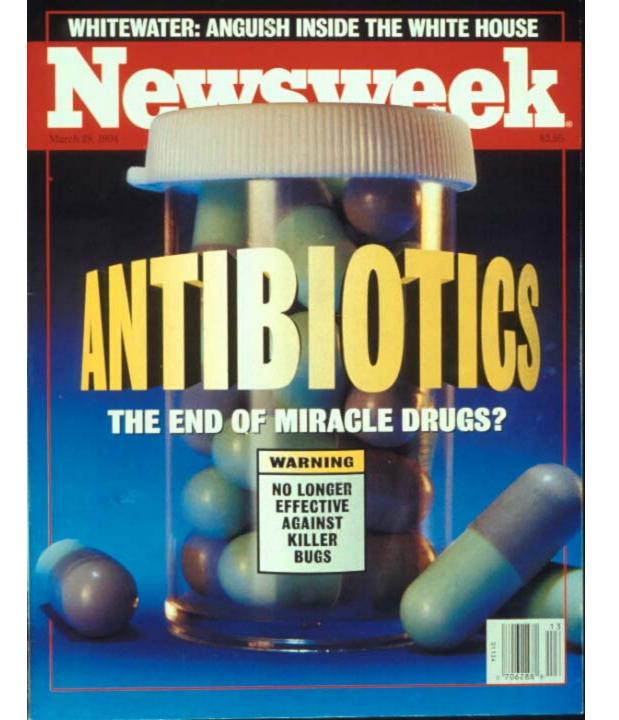


Patients with pneumonia and bacteria in the blood

Penicillin increased the chance of survival from 10% to 90%

Adapted from Austrian et al. Ann. Int. Med 1964; 60, 759





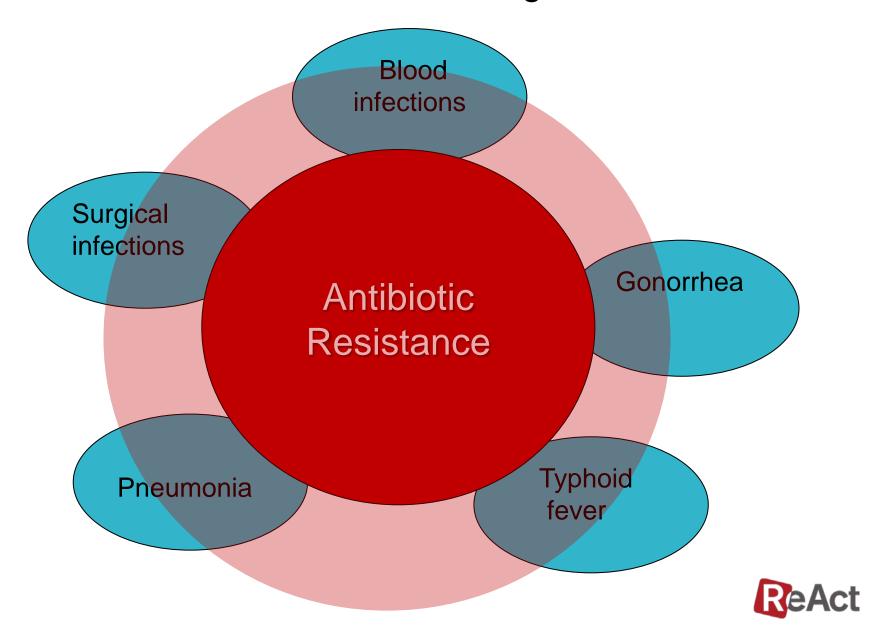


Antibiotic resistance - a rhetorical problem ?

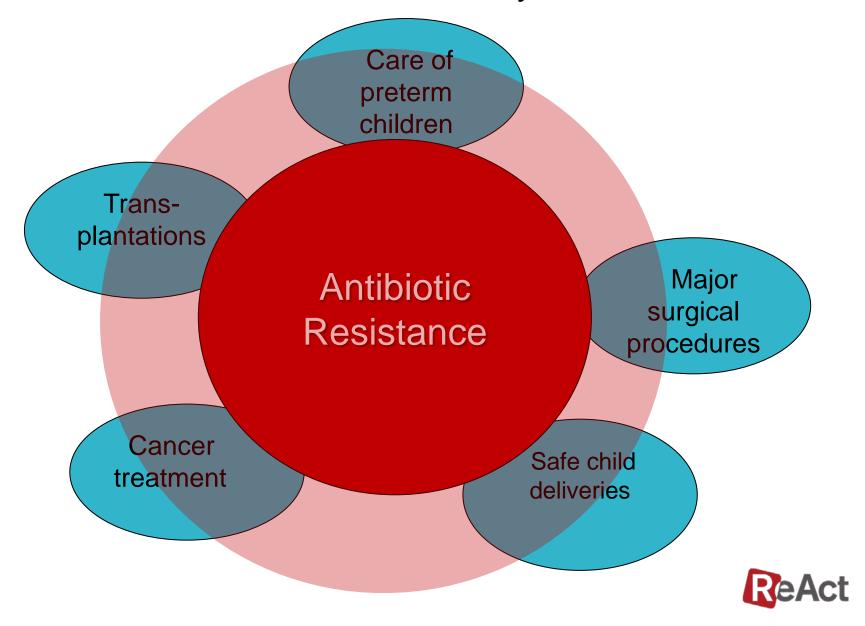
- Compared to other health problems,
 we are *not* talking about a specific disease
- Instead, we are talking about a medical product that is no longer working against bacteria causing many common infectious diseases



Common diseases- becoming non-treatable



In addition, many *medical procedures* are affected and threatened by ABR



Who is affected by ABR?

Everyone!

However

- →... vulnerable populations with the highest infectious disease burden pays the highest price
- →... these populations are also the ones that have least access to current and future interventions to manage antibiotic resistance



Some parts of the world have already run out of effective antibiotics





The mortality rate from Gram-negative bloodstream infection in children (43,5%) was more than double that of malaria.

Blomberg et al. BMC Infect Dis. 2007



About 70% of neonatal systemic infections can not be treated with the antibiotics recommended by WHO....

Lancet 2005; 365: 1175-88



COST IMPLICATIONS OF GROWING ANTIBIOTIC RESISTANCE ARE HUGE Typhoid fever

Cost of antibiotics for non-resistant cases

\$3-5

Cost of antibiotics for resistant cases

Azithromycin	\$35-42
/ Licin Only Onl	ψ 3 J- 1 Z

Oral cephalosporins (cefixime) \$37-42

Parenteral cephalosporins (ceftriaxone) \$84-104

Average Treatment Costs for Typhoid (US\$)
Child weighing 20 kg using standard treatment guidelines

Source: AKU Pharmindex 2004 & WHO guidelines 2003



The MDG:s

- Eradicate extreme poverty and hunger
- Achieve universal primary education
- Promote gender equality and empower women
- Reduce child mortality
- Improve maternal health
- Combat HIV/AIDS, malaria and other diseases
- Ensure environmental sustainability
- Develop a global partnership for development



No. 68421 THURSDAY JUNE 23 2005 www.timesonline.co.uk 55g



m-seekers nger protest

ls of Zimbabwean eekers in detention egan a hunger strike ain's decision to send ck to face torture bert Mugabe's regime.

cellor's call

Brown called on the night to end its onist ambitions and e radical economic needed to turn it from loc into a global force. age 2

New strains of superbug hit hospitals

At 65, she's the world's most beautiful woman

The incredible Sophia Loren • See Page 15



Vulnerable patients in danger from deadly bacteria running riot on wards

EXCLUSIVE BY ANTHONY BEYONS AND RACHEL ELLIS

THE spread of a deadly superbug in British hospitals is reaching epidemic proportions, a leading scientist warned

Yesterday.

Over the past nine years, there has been a 13-fold increase in the incidence of the bug known as MRSA, which is resistant to all but

the most powerful antibiotics.
In Scotland, Wales and the West Midlands, increases have been even more dramatic. Dr. David Liver-

more, of the Public Health Laboratory Service, told The Express
of Control of the Express
and official figures on the number of
deaths caused by the bug and it is
was due to epidemic strains 15 and
its particularly virtuent forms of
the hacteria which can be transmitted from patient to gatient simply
on the back of someone's hand.
Hospitals have been warred to
Hospitals have been warred to
10 37 per cent so far in 1996. Dr
Livermore said.



WEATHER 2 ● OPINION 10 ● LETTERS 39 ● DIARY 41 ● LIFE 43-48 ● GROSSWORD 58 ● TV 61-65 ● CITY 72-75

Burden of multidrug-resistant bacteria in the EU

Attributable deaths approx. 25,000 / year Extra hospital days approx. 2.5 million / year Total costs approx. € 1.5 billion / year

Limitation: these are underestimates.





A fact shiet from ReAct - Action on Antibiotic Resistance, www.reactgroup.org May 2012

Burden of Antibiotic Resistance

serious problem.

Gram-agative bacteria are particularly trouting these infections will increasingly worrisone. In the US, two thirds of deaths due to historial infections are caused by permanandowlys. Cirary-regative inscients."

recruing firest to berne health across bacteria are e.g. blood steam infections. affecting leadth care system and societies the world. No country can escape from the arrange tract infections, post-operative across the world. medical and communic impacts from this wound infections and intra-shoppinal infortage.

lend to serious health problems and

The consequences of authoric resistance offices patients' lives but also

Antibiotic resistance (AIIII) poses at Common diseases resulting from time resides for beyond the individual patient

Within just a few years, we may very well be found with untraginable softwales, Cortain bactoria and an each resistant. The tack of effective antificities for medically, socially and concentrally anims we macroose.

> The ongoing panderoic spread of restricted fracteria (Sudrates flut the problem one only be addressed through interpational cooperation.

The studies in this fact short were published. between 2003 and 2012.

PATIENT GROUPS ESPECIALLY AFFECTED BY ABR.

Newborn and children

- *ESBL-producing bacteria are frequently causing affections in newtorns. In an Indian bogital, Klehriella and E. coli were the most summer Gram-negative bacteria among infants with RSIs. About 13% of ESBL-infections were deadly in spite of available newer antibiotics and other supportive care.
- *In a studyfrom Pakistan, 37 of 78 newborns (less than 6 days old) with infactions due to Actionsform died within a short time frame. 71% of the bacteria were resistant to all antimities except polymyxis.
- + In an outfreak in India council by NDM-1 & rall, 4 sewhorn habites contracted blood stream infections (BSSs). All four died."
- *Three purnature babies in a German neonatal word died due to an outbreak of ESBL-producing Kletninks presentation from an unknown source.
- * In a study of Tananasian children, the BSI. rate was as high as 13.9%. One third of these chidnes died. The death rate from Gram-engative BSI (43.5%) was more than double test of restarts (20.2%). One



C.1709 Dayor Bussit, Misserisper, Courtoy of Pleasables.

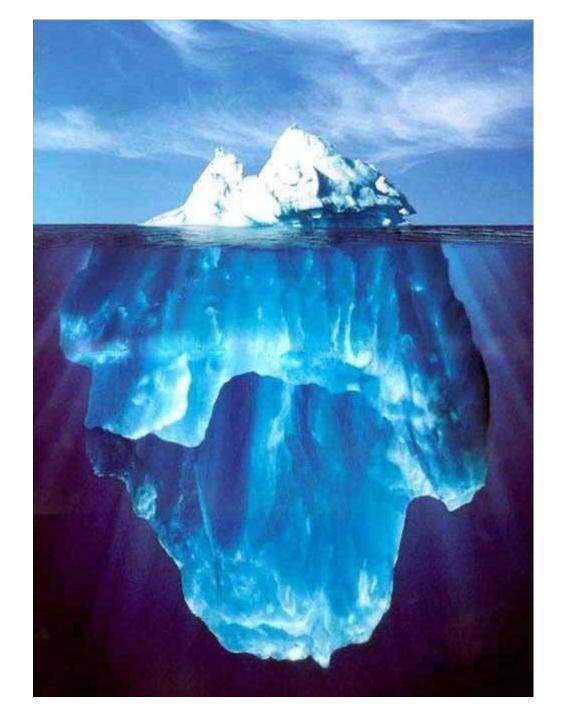
significant risk factor for death was treatment with ineffective antibiotics due to antihiotic resistance."

Transplantation and cancer patients

*In Spain, among 416 patients undergoing transplantations of kidneys, 58 were infected with realti drag resistant (MDR) hacteria, most often Gram-engetive. BSIs occurred in 14% of those. Death or graft loss was significantly more frequent among those with MDR infections (19% vs. 8%).

- *In the US, an outbrok of K. presumonice carbapenemase-producing infections among liver transplant recipients killed two patients. A larger orbreak involving 24 patients soon followed the two initial cases in the word.
- . In number study from the US, recent organ.







How did we end up here?

- The indiscriminate use of antibiotics
- The rapid dynamics of gene transfer between bacteria and the global spread of resistance
- Poor sanitation and hygiene
- Alarming decline in antibiotics development









Athens, Greece 2008 (174 pharmacies)

-100% of all visited pharmacies sold Amoxicillin/clavulanate acid OTC

- 53% sold Ciprofloxacin OTC, despite extra restrictions for fluoroquinolone prescriptions

Plachouras et al. Euro Surveill. 2010

2008:



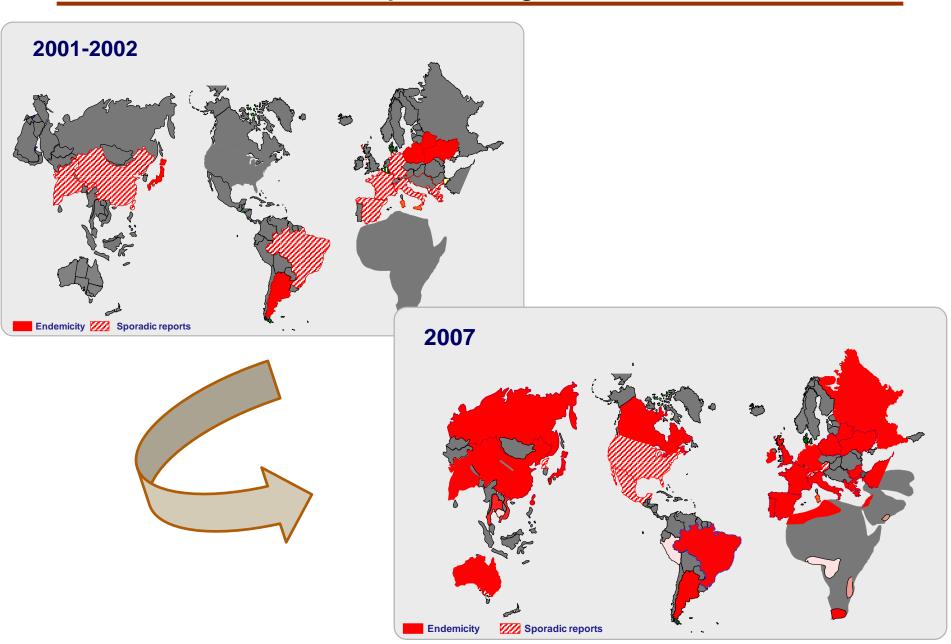








ESBL (CTX-M) producing *Enterobacteriaceae*



THE DRUG DEVELOPMENT PIPELINE FOR ANTIBIOTICS HAS GONE DRY



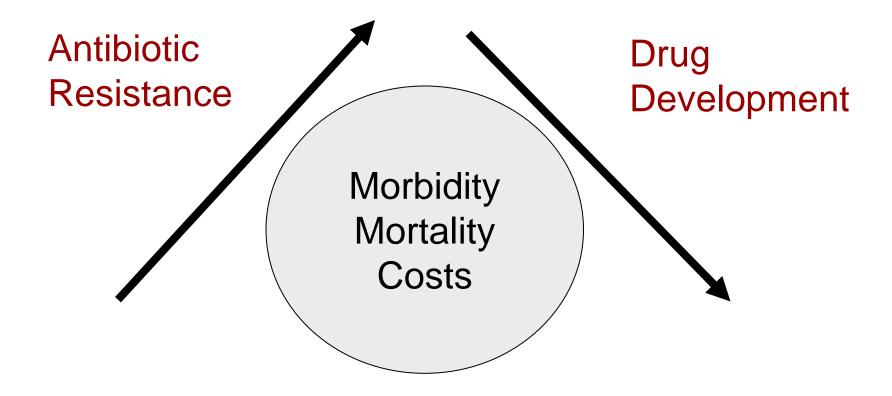


THE DRUG DEVELOPMENT PIPELINE FOR ANTIBIOTICS HAS GONE DRY

 The last overed in No new drugs for: 1987 - Typhoid fever - Shigella The pming - Gonorrhoea inef - Urinary tract infections - Blood infections either the The p nercial innovat interest to otics



We are facing a public health crisis





Strategies to stop manage antibiotic resistance

Prolong the lifespan of existing drugs

Rational Use Better diagnostics

Prevent the spread of resistant bacteria

Improved hygiene Infection control

Development of new antibiotics

New collaborative models

Controlled distribution and use of new future antibiotic

A new global system to preserve the effectiveness of antibiotics

Develop national action plans on antibiotic resistance!

Surveillance Decrease Use Coordinate the need antibiotics national for properly activities antibiotics Monitor: Reduce **Improve** Knowledge diagnostics antibiotic disease education, and usage research incidence usage and resistance and prevent (human & spread of veterinary patterns International bacteria sectors) collaboration



Antibiotic Resistance

Caused by human activity and by over- consumption of a gobal resource

A failure of public policy & global governance, research prioritization and the current market system

It is a collective responsibility by governments, supranational organizations and individuals to take action





