

Dramatic changes in society and environment . .

Explosive population growth, spreading poverty, global warming, and urban migration

– new pathogens, old pathogens in new places

Dramatic changes in society and environment . .

Increased international travel and trade: we live in a global village

- rapid spread of disease

Dramatic changes in society and environment . .

Loss of control of national borders

– ineffective quarantine laws

Dramatic changes in society and environment . .

Rise of antibiotic resistant pathogens

1946 Penicillin became available

- Strep and staph (strep throat, pneumonia, septicemia, skin infections, wound infections, scarlet fever, toxic shock syndrome)

Today, 80 % of all strains of staph are resistant to penicillin

1950 Streptomycin, chloramphenicol, tetracycline

- Multiple infections

1953 Shigella outbreak in Japan resulted in the appearance of a strain of dysentery bacillus with multiple drug resistances

1982 Last new class of antibiotics – quinolones

- Resistance rising

1998 Vancomycin is antibiotic of last resort for staph and other gram positive pathogens

We are now seeing emergence of vancomycin resistance!

Why Antibiotic Resistance is Growing Rapidly

- **Antibiotics in animal feed and in aerosols for fruits and vegetables**

–Of the 50-million lbs of antibiotics produced annually in the U.S., 40% go to livestock

Why Antibiotic Resistance is Growing Rapidly

- **Growing numbers of immuno-compromised people**

- Chemotherapy Patients
- Transplant patients
- AIDS patients
- Aging population

Why Antibiotic Resistance is Growing Rapidly

- **Excessive use of antibiotics**

- Over prescription
- Unregulated over-the-counter sales

Why Antibiotic Resistance is Growing Rapidly

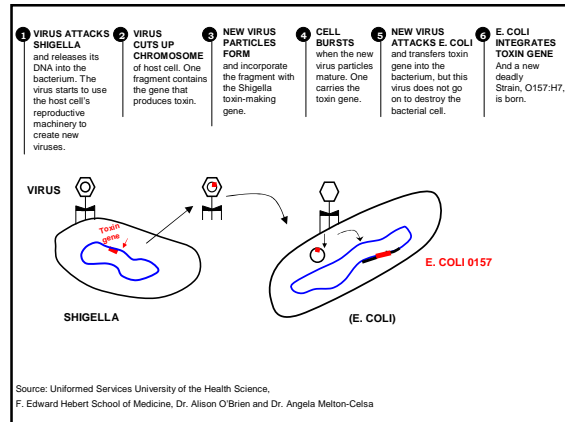
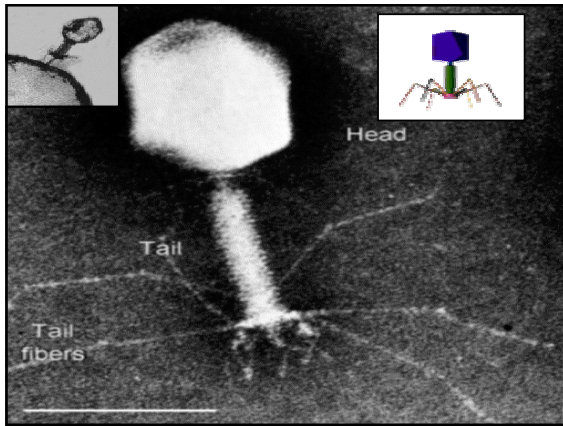
- **International travel**

- Multidrug-resistant streptomycin:

Rapid spread: Spain → UK
→ US → So. Africa

E. Coli 0157: H7
Where did this come from?

- A new and pervasive pathogen
- A food contaminant that is now the leading cause of kidney failure in children



West Nile Virus

SARS

Severe Acute Respiratory Syndrome

- Corona Virus – the same type of virus that causes the common cold
- High potential for natural evolution

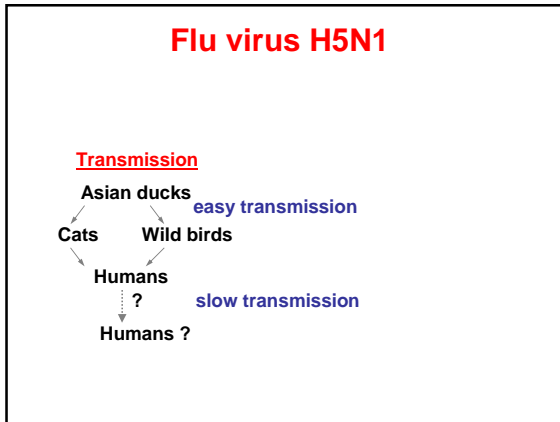
Influenza

Asian Bird Flu H5N1

- No strong evidence of human to human transmission, yet
- The concern is that H5N1 mutates rapidly

Flu virus H5N1

- Single strand of RNA containing 8 genes, each encoding a single protein
- High mutagenicity rate due to:
 - Reassortment of the genes
 - Single base mutations



Influenza viruses

H = haemagglutinin
 - binds to host cells to aid virus entry

N = Neuraminidase
 - allows newly formed viruses to escape and infect other cells (Tamiflu & Relenza target)

Influenza viruses

1918 flu (H1 N1) Killed ~40 million people

1957 flu (H2 N2) Killed ~2 million people

1968 flu (H3 N2) Killed ~1 million people

Current Asian bird flu (H5 N1) 50% death rate

Humans have no immunity against H5, only against H1, H2, H3

What Needs to be Done?

- Increase basic research to understand viral and bacterial pathogens
 - Identify genes essential for pathogen's survival
 - Sequence and compare viral and bacterial genomes
 - Identify "virulence" and "resistance" genes and how they work
- Design and stockpile new vaccine strategies and "combination" antibiotics

What Needs to be Done?

• **For epidemic control:**

- Develop techniques for fast (hours, not days) identification of causative agents
 - Exploit viral and bacterial DNA sequence-based technologies
- Increased network of surveillance and reporting protocols
- Return to historical use of quarantine
 - For resistant and virulent infections it is the only effective containment strategy