

## (Network's) Models of epidemiology

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### **Black Death** (PDE)









H5N1 influenca - bird flu 2005 H1N1 influenca - swine flu 2009 (Networks)





Plant's diseases - rhisobia (CA)







Animal's diseases Mad Cow - BSE (CA, Networks)





# Differential equations vs Netwoks **Data!!!**



Wrocław – from the beginig of smallpox epidemiology modeling up to erudation of disease



In Poland smallpox last time appeared in Wroclaw in 1963, but it was stopped by the actions of the government and epidemiologists. Moreover smallpox was eradicated by WHO in 1979. Bernoulli (1760) actually used date provided from Wroclaw to estimate his model.



This Swiss mathematician **was the first to express the proportion of susceptible individuals of an endemic infection** in terms of the force of infection and life expectancy. His work describe smallpox, which cause a lot of epidemics in big European cities at his time. Smallpox devastated earlier the native Amerindian population and was an important factor in the conquest of the Aztecs and the Incas by the Spaniards.



I simply wish that, in a matter which so closely concerns the well-being of mankind, no decision shall be made without all the knowledge which a little analysis and calculation can provide.

**Daniel Bernoulli**, presenting his estimates of smallpox Royal Academy of Sciences-Paris, 30 April 1760



Bernoulii based at work of famous British **astronomer – Edmund Halley** "An Estimate of the Degrees of the Mortality of Mankind, drawn from curious Tables of the Births and Funerals at the City of Breslaw" published in Philosophical Transactions 196 (1692/1693).

In 17th century English Breslaw means Breslau-German name of Wrocław.



|              | Not      |          |           | Dying of |          | other    |
|--------------|----------|----------|-----------|----------|----------|----------|
| Survivors    | having   | Having   | Catching  | smallpox | Total    | diseases |
| according to | had      | had      | smallpox  | each     | smallpox | each     |
|              | omallnov | omallnov |           | Voor     | dootho   | voor     |
| папеу        | Smailpox | Smailpox | each year | year     | ueatiis  | year     |
| 1300         | 1300     | 0        |           |          |          |          |
| 1000         | 896      | 104      | 137       | 17,1     | 17,1     | 283      |
| 855          | 685      | 170      | 99        | 12,4     | 29,5     | 133      |
| 798          | 571      | 227      | 78        | 9,7      | 39,2     | 47       |
| 760          | 485      | 275      | 66        | 8,3      | 47,5     | 30       |
| 732          | 416      | 316      | 56        | 7        | 54,5     | 21       |
| 710          | 359      | 351      | 48        | 6        | 60,5     | 16       |
| 692          | 311      | 381      | 42        | 5,2      | 65,7     | 12,8     |
| 680          | 272      | 408      | 36        | 4,5      | 70,2     | 7,5      |
| 670          | 237      | 433      | 32        | 4        | 74,2     | 6        |
| 661          | 208      | 453      | 28        | 3,5      | 77,7     | 5,5      |
| 653          | 182      | 471      | 24,4      | 3        | 80,7     | 5        |
| 646          | 160      | 486      | 21,4      | 2,7      | 83,4     | 4,3      |
| 640          | 140      | 500      | 18,7      | 2,3      | 85,7     | 3,7      |
| 634          | 123      | 511      | 16,6      | 2,1      | 87,8     | 3,9      |
| 628          | 108      | 520      | 14,4      | 1,8      | 89,6     | 4,2      |
| 622          | 94       | 528      | 12,6      | 1,6      | 91,2     | 4,4      |
| 616          | 83       | 533      | 11        | 1,4      | 92,6     | 4,6      |
| 610          | 72       | 538      | 9,7       | 1,2      | 93,8     | 4,8      |
| 604          | 63       | 541      | 8,4       | 1        | 94,8     | 5        |
| 598          | 56       | 542      | 7,4       | 0,9      | 95,7     | 5,1      |
| 592          | 48,5     | 543      | 6,5       | 0,8      | 96,5     | 5,2      |

Table 1. Smallpox in Wroclaw [D. Bernoulli]



Bernoulli has constructed table with 'natural state with smallpox', in contradistinction to the third column, which shows the 'state without smallpox' and which gives the number of survivors each year assuming that nobody must die of smallpox. Difference between second and third column gave him **a gain in people's lives**. He introduced 'total quantity of life' of the whole generation.



#### **Differential equations**

Suppose that population is divided into three classes: the susceptibles (S) who can catch the disease; the infectives (I), who can transmit disease and have it; and the removed (R) who had the disease and are recovered (with immune) or isolated from society. Schema of transition can be represented:

# $S \to I \to R$

W. O. Kermack and A. G. McKendrick, 1927



#### **Differential equations**

$$\frac{dI}{dt} = rSI - aI,$$
$$\frac{dS}{dt} = -rSI,$$
$$\frac{dR}{dt} = aI$$

Where transmission rate is:  $r = \beta C/N$  $\beta$  – infectivity rate, C – contact rate, a – recovery rate



#### **Differential equations**

 $R_0 = \frac{r S_0}{r}$ 

where  $R_0$  is basic reproduction rate of the infection. This rate is crucial for dealing with and an epidemic which can be under control with vaccination for example. Action is needed if  $R_0>1$ , because epidemic clearly ensues then.



#### **Differential equations**



Influenza epidemic data  $(\bullet)$  for a boys boarding school as reported in UK in 1978. The continous curves for *I* and *S* were obtained from a best fit of numerical solution of SIR system with parameters:  $S_0=762, I_0=1, S_c=202,$ *r*=0,0022/day.



#### Agent-based models

#### Differential equation vs Agent-based models

#### Network of contacts:

In an SIR-type model, the population is split into three different groups and the majority of the population is placed in the susceptible compartment. All information about society is used in microsimulation, so it can give better prediction, then differential equations



#### Agent-based models vs DE

Differential equations were first applied to describe and predict those phenomena (in use since the 18th century when even the definition of "differential equation" has not been known). Epidemiological models that treat transmission from the perspective of differential equations do exist in older literature, but recently agent-based models appear even more often. Both approaches use variations of SIR (susceptibles - infectives removed) concept, so sometimes the same problem could be solved in both ways. Not always, because computer simulation has changed the world of mathematical modeling, agent-based models give better predictions and some hints for decision-makers even parallel development of numerical methods for differential equations. On the other hand, differential equations allow us to understand the core process, which could be missing in the agent-based approach. As a result, both perspectives are common among epidemiologists and depend on theoretical or applied aspect percentage representations differ.



#### **Celluar Automata**

Cellular Automata were invented in early 50<sup>th</sup>. There based on some designed topology of elements which have some possible number of states and they can evaluate in time. A cellular automaton may be also a collection of "colored" cells on a grid of specified shape that evolves through a number of discrete time steps according to a set of rules based on the states of neighboring cell.



#### **Celluar Automata**





#### **Celluar Automata**





#### **Celluar Automata**

#### No additional links



#### With additional links





#### H1N1 - introduction





#### H1N1 - MicroSim

Disease transmission is performed twice daily at 9 am and 5 pm. Programme checks where are all persons during that day hours and night hours.

We have for exmaple profiles of probability of sending infections



Profile of probability (sending H1N1) [adopted F. Carrat ]



The daily routines for the simulated persons. [L. Brouwers]



### H1N1 - Realization

The outbreak of pandemic influenza in Sweden starts depend of method in June or in September.  $R_0$  -value corresponding approximately to 1.4 in main model, because that was observed in New Zealand during their outbreak. The viral infectivity is markedly initially  $R_0$  value of approximately 2.1 in preliminary method.

 $R_{0} = \frac{-\ln(\frac{A}{S_{0}})}{1 - \frac{A}{S_{0}}}$ Immunity<sup>0</sup> was calibrated in model to obtain R<sub>0</sub>- 1.4 S<sub>0</sub>: Total number of susceptible individuals before the outbreak A: Total number of susceptible individuals after the outbreak

This formula is defined as is the average number of individuals a typical person infects under his/her full infectious period, in a fully susceptible population.



### H1N1 - Cost & Vaccination

To compare the societal costs of the scenarios, the following costs—obtained from health economists at the Swedish Institute for Infectious Disease Control —were used.

- •Cost of one day's absence from work, for a worker: SEK 900.
- •Cost of treatment by a doctor in primary care: SEK 2000.
- •Cost of one day's inpatient care: SEK 8000.
- •Cost of vaccine and administration of vaccination for one person: SEK 300.

The following scenarios were compared: no response, the vaccination coverage of 30%, 50%, 60%, 70% or 90% in simulation.



#### H1N1 - Results



No. of infections (upper graph) and cost of pandemia (lower graph) for different scenarios [by L. Brouwers].



#### H1N1 - Simulation & Reality



Vecka

No. of infections per week in simulation (upper graph by L. Brouwers) and cases collected by The Swedish Institute for Infectious Disease Control (lower graph by SMI).



#### **Network Theory - introduction**

## An agent/object's actions are affected by the actions of others around it.

## NETWORK

Actions, choices, etc. are not made in isolation i.e. they are contingent on others' actions, choices, etc.



#### Network Theory - terminology



Node = individual components of a network e.g. people, power stations, neurons, etc.

#### Edge = direct link

between components (referred to social networking as a relationship between two people)

#### Path = route taken across components to

connect two nodes



#### **Network Theory - clusters**



Clustering coefficient counts numer of triangles in networks



#### **Network Theory - paths**



Average path length



#### Network Theory - node's degree



**Degree distribution** 



#### Network Theory - randomness



#### Randomness



#### **Network Theory - communities**





#### Where's George





#### Where's George





#### Where's George





#### Sexually transmitted infections (STI)





#### STI - problem

#### Probability of transmission per contact (β) HIV 0,08-1,7% Chlamydia 2-60%

#### 30-34 years 2400 25-29 years 250 Heterosexual Probably infected abroad 20-24 years Probably infected in Sweden 2000 15-19 years Incidence per 100,000 in habitants 200 Females 1600 150 1200 100 800 50 400 0 0 87 88 89 90 91 92 93 94 95 96 97 98 99 00 01 2000 2001



#### **STI - networks**



Liljeros, 2004



#### STI - data

- Difficulties with access
- Sensitivity informations
- Bioethical comities



#### STI - data

- Difficulties with access
- Sensitivity informations
- Bioethical comities
  - sexuality of Gots
  - prostitution in Brazil



#### STI - data (Gothland)



Liljeros, 2001 (Nature)



#### STI - data (Brazil)





#### STI - data (Brazil)



Liljeros, 2010 (PNAS)



#### STI - model

#### Time and intensity of contacts does matter





STI - future...

#### Research in plans

