



0. Information-only slide

- **Contents:** Slides for the LDRD Day poster
- **Title:** Distributed micro-releases of bioterror pathogens: Threat characterization and epidemiology from uncertain patients observables
 - Project: 93505
 - People: Jaideep Ray (PI), Brian Adams, Karen Devine, Youssef Marzouk, Habib Najm.
 - LDRD IA: Enabling Predictive Simulations
- **Constraints :**
 - 8 slides
 - 20 pt font and higher
 - Problem/Approach/Results/Significance structure
- ***To the artist in Creative Arts: Please play around with font sizes etc, as needed.***



Problem definition

- **Aim: To characterize and infer the genesis of epidemics from partial information**
 - Estimate the size of a bioattack (for medical response purposes)
 - Estimate pathogenic characteristics (for tracking emerging infectious diseases)
 - Uncertainty quantification (UQ), due to partial observations
- **Technical challenges**
 - Stochastic model of disease spread
 - Separating social and pathogenic contributions to disease propagation
 - Inferring social parameters (social networks)
 - Formulating a Bayesian inverse problem to do so
 - MCMC methods for high-dimensional parameters spaces

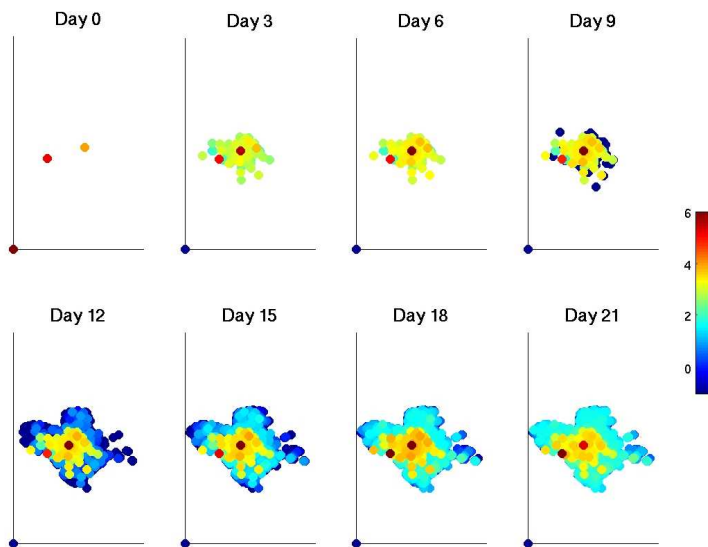


Overall approach

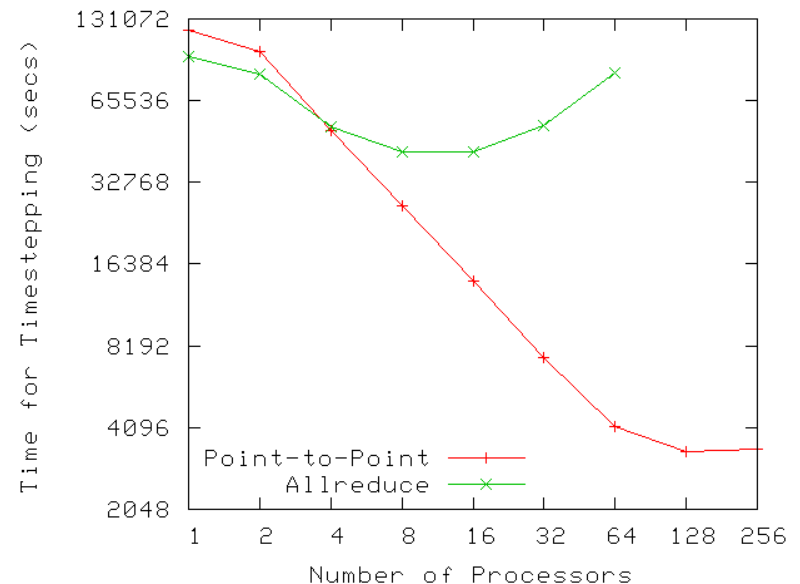
- **Create a stochastic, individual-based epidemiological model separating social and pathogenic contributions**
 - Start with an individual-based dynamic social contact network model (Network Forward Model, NFM)
 - Derive from it, an equivalent, but more efficient static network model
- **Inverse problem: Bayesian; parameters estimated as PDFs**
 - Estimation of bioattack parameters from partial observations of the epidemic
 - Estimation of pathogenic properties, e.g., transmissibility
 - MCMC algorithms to infer graphs
 - Prior beliefs of graph morphologies/parameters guided by previous item

Stochastic epidemiological model

- Dynamic model (EpiSims, Eubank, et al.) based on **bipartite, time varying graph** (transit network-based): people move among locations throughout a day; shed or absorb virus
- Each person or location has a disease “load”; when exceeds a threshold, progress through infection stages



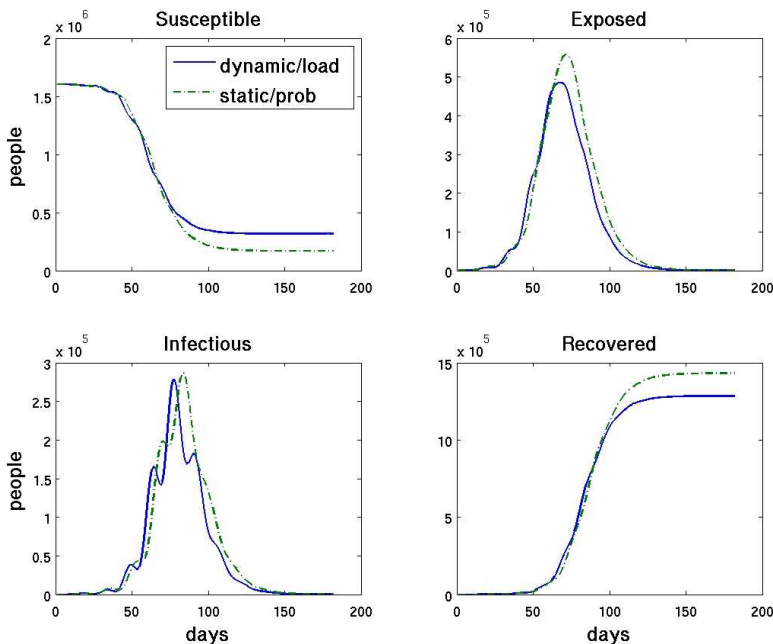
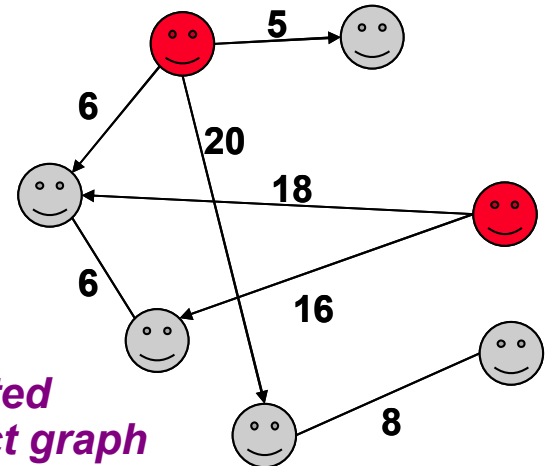
*simulated spread of smallpox in
an urban setting over 21 days*



*parallel scalability enables
rapid scenario analysis*

Epidemiological model reduction

- One reduced-order modeling approach: create static contact network based on person collocation in a 24 hour period
- Use a time-dependent probability of transmission model



- Similar epidemic prediction to dynamic, load-based model
- Also explored graph clustering and sampling techniques to reduce number of people/locations in simulation

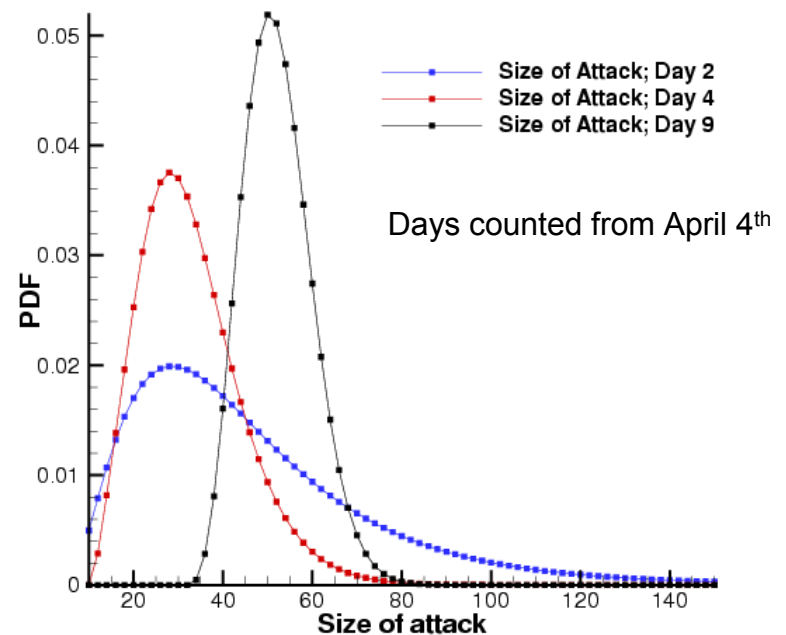
Estimation of bioattacks (anthrax)

- **Approach:**

- Disease model predicated on a dose-dependent incubation period distribution
- Analytical expression for the likelihood $\pi(N, \tau, D|S)$
 - N: number infected, τ : time of infection, D: average dose

- **Test/results: Sverdlovsk, 1979.**

- Accidental release; 80 victims; 70 deaths
- Release: 2nd April
- First symptoms: 4th April
- Outbreak lasted 42 days
- Medical response starts: April 14th
- People affected: 50,000-60,000

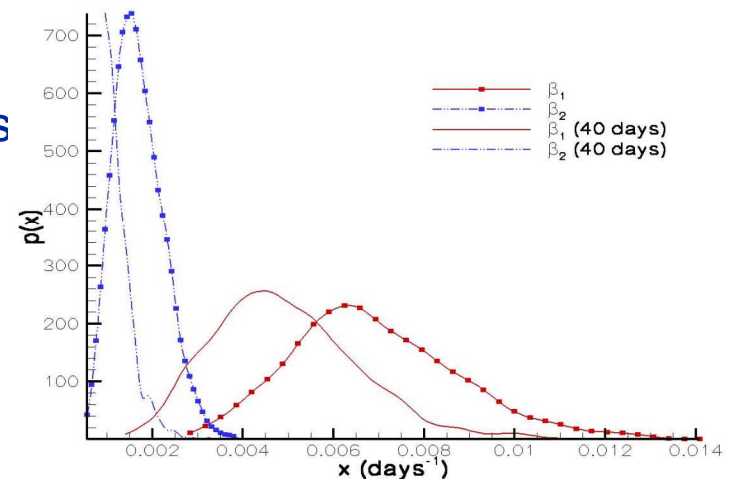


It was quite clear by April 13th that the outbreak would be small (less than 100 infected)

Estimation of pathogenic transmissibility

- **Approach:**

- Structured population, represented as a graph G , a collection of B -graphs
- Model disease transmission along links of G as a Poisson process, with rate β
- Differential transmission, inside and across groups
- MCMC to infer β , G and infection pathway \mathcal{P}

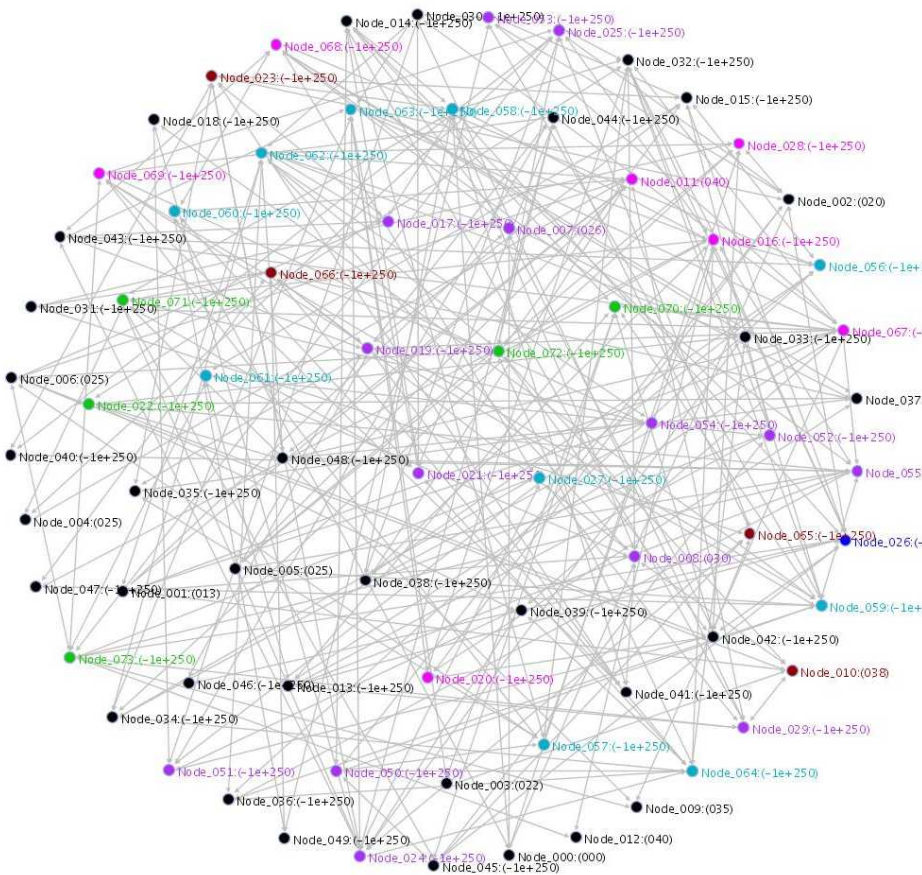


- **Test/results: Smallpox outbreak, Abakaliki, 1967. WHO/SE/68.3**

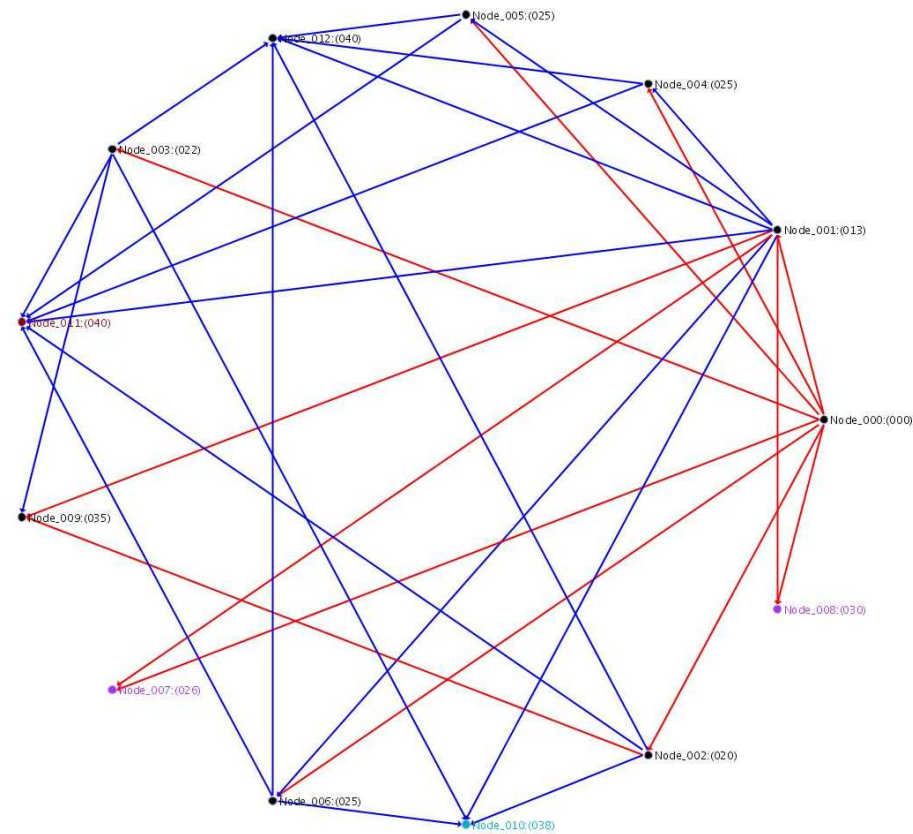
- Affected a closed population of 74, divided into 7 groups. 30 victims
- Differential rates of spread observed in data (“eyeball norm”)
- Outbreak lasted 3 months
- Only dates of appearance of symptoms recorded; victims’ fate unknown. No vaccinations.
- Inferences drawn from full data, and the first 40 days of data

Estimation of social characteristics

Expected social graph $\langle G \rangle$



Expected infection pathway $\langle P \rangle$



Results estimated from the first 40 days of the epidemic.



Significance

- **One can derive simplified, static network models of disease spread from detailed individual-based models**
- **Partial observations can yield significant information about an epidemic's genesis and pathogenic characteristics**
- **Requires:**
 - A good characterization of the social structure (to serve as a *prior* in the Bayesian inference method)
 - Powerful, multi-chain MCMC algorithms to traverse the parameter space
 - Mode-hopping MCMC to address multi-modality.
- **Few previously published works on both topics**
- **Potential for largest impact**
 - More realistic social networks, derived from individual-based models, used in the inference process
 - Advanced MCMC schemes for inferring networks