Assessment of epidemiological risk related to pathogen spread on the network of French cattle movements

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ModStatSAP 2014, Paris 11 December

Context: cattle trade as a pathway for pathogen spread

- Important to understand the contact structure among herds underlying pathogen transmission
 - In particular for infectious diseases endemic in livestock: animal health and welfare; economic losses in animal productions; zoonoses
 - Animal trade networks characteristics impact infection spread
- Network representation: farms as nodes, trade relationships as links

Animal trade as networks in the literature

- Aims of published studies
 - Demographic structure and pathogen dynamics
 - Dynamical patterns in longitudinal data, surveillance optimization
- Methods
 - Static networks of aggregated data
 - Continuous increments in steps of small time windows
 - Time-stamped networks
- Different countries and several animal markets
 - Cattle: Canada, Denmark, France, Italy, Sweden, UK, US
 - Pigs: Canada, France, Germany, Sweden

[Keeling and Eames, 2005; Bigras-Poulin *et al.*, 2006; Kao *et al.*, 2006; Kiss *et al.*, 2006, Natale *et al.*, 2009; Dube *et al.*, 2010; Danon *et al.*, 2011; Bajardi *et al.*, 2011, 2012; Rautureau *et al.*, 2011, 2012; Noremark *et al.*, 2011; Vernon and Keeling 2009; Vernon 2011; Buttner *et al.*, 2011; Dorjee *et al.*, 2013; Mweu *et al.*, 2011; Buttner *et al.*, 2013; Buhnerkempe *et al.*, 2013; Konschake *et al.*, 2013; Noremark and Widgren, 2014]

Objectives

• Study of the French animal trade network over several years¹

- To identify if and which structural patterns underlying pathogen spread are stable in time
- To explore control measures based on network-related characteristics of nodes (rely on previous data if real-time information not available)
- Joint economic and epidemiologic perspective²

¹B.L. Dutta, P. Ezanno, E. Vergu. (2014) Characteristics of the spatio-temporal network of cattle movements in France over a 5-year period. *Prev. Vet. Med.* 117(1):79-94.

²M. Moslonka-Lefebvre, C. Gilligan, H. Monod, C. Belloc, P. Ezanno, J.A.N. Filipe, E. Vergu. (2014) Market analyses of livestock trade networks to inform the prevention of joint economic and epidemiological risk. *Submitted*.

Some basic descriptors for the animal movement data in France (2005 - 2009)

 The data include all individual information for cattle (animal ID, date of birth, date, origin and destination holdings - farms (F), markets (M) and assembly centers (AC) - for each movement, etc).

| | 2005 | 2006 | 2007 | 2008 | 2009 |
|-------------------------------|------------|------|-------|-------|-------|
| Total holdings (V) | 243,324 | -3.7 | -7.5 | -11.7 | -14.7 |
| (M; AC) | (84; 1547) | | | | |
| No of movements (W) | 8,636,018 | -0.1 | -9.2 | -11.7 | -12.2 |
| No of links (A) | 1,279,576 | -9.3 | -13.1 | -19.8 | -22.8 |
| No of batches | 2,791,261 | -4.7 | -12.8 | -17.7 | -20.4 |
| No of cattle | 5,533,854 | -0.8 | -6.1 | -7.6 | -9.6 |
| No of holdings as origin | 228,400 | -3.6 | -7.8 | -12.3 | -15.2 |
| No of holdings as destination | 141,249 | -3.4 | -7.7 | -12.3 | -16.1 |

The yearly values are shown as % in change with respect to observations in 2005.

• Main flows: $F \longrightarrow AC$ (40%); $F \longrightarrow F$ (18%); $AC \longrightarrow F$ (15%),

Dutta et al. (2014) Prev. Vet. Med.

Sub-network of dairy herds in Finistère as an example

Subnetwork (2005-2009) of dairy herds trading non beef animals > 6 months

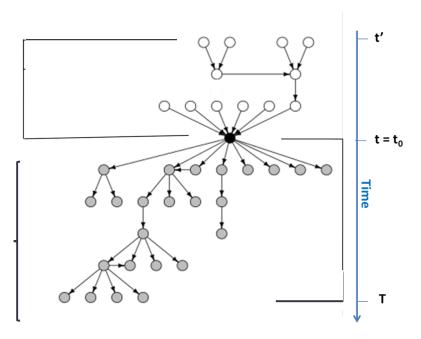
- 1945 herds
- 89,281 movements

Some elements for network analysis (1)

- The explored data are represented by directed, weighted and time-varying networks.
- Basics for directed weighted networks $W = \{w_{ij}\}, w_{ij} \ge 0$ the no. of mvts. $i \rightarrow j$ (unweighted $A = \{a_{ij}\}, a_{ij} = 1$ if $w_{ij} > 0$ and 0 otherwise)
 - degree (k): $k_i^{out} = \sum_j a_{ij}, k_i^{in} = \sum_j a_{ji}$
 - strength (s): $s_i^{out} = \sum_j w_{ij}, s_i^{in} = \sum_j w_{ji}$
 - betweenness centrality: C_B(i) = ∑_{j≠i≠l} σ_{jl}(i)/σ_{jl}, where σ_{jl}(i) is the no. of shortest paths between j and l passing through i, and σ_{jl} is their total no.
- Economic-like descriptors
 - activity (flow-share) (x): $x_i = (s_i^{in} + s_i^{out}) / \sum_j (s_j^{in} + s_j^{out})$ by definition, $0 \le x_i \le 1$ and $\sum_i x_i = 1$; for given $\delta_1, \delta_2 > 0$ agents *i* for which $x_i < \delta_1, x_i \in [\delta_1, \delta_2]$ and $x_i > \delta_2$ are denoted *nichers*, followers and leaders
 - ► flow polarity (δ): $\delta_i = (k_i^{in} k_i^{out})/(k_i^{in} + k_i^{out})$ by construction $-1 \le \delta_i \le 1$; for a given $\epsilon > 0$ agents *i* for which $\delta_i < -\epsilon$,
 - $\delta_i \in [-\epsilon, \epsilon]$ and $\delta_i > \epsilon$ correspond to suppliers, wholesalers and demanders

Some elements for network analysis (2)

- Measures for assessing time-varying characteristics of networks
 - ▶ short range similarity (SRS): $SRS_V = |V_t \cap V_{t+\Delta t}|/|V_t \cup V_{t+\Delta t}|$, $SRS_E = |E_t \cap E_{t+\Delta t}|/E_t \cup E_{t+\Delta t}|$ (where network $G_t = (V_t, E_t)$, with V, E sets of vertices and edges; $G_t, G_{t+\Delta t}$ are snapshots of the same network at two consecutive time intervals)
 - reachability ratio (RR): fractional size of set of influence (for node i, it includes nodes reachable from i by time respecting paths) averaged over all nodes with out-movements
- Proxies for pathogen spread and its control
 - giant strongly connected component (GSCC)
 - percolation: fractional change in the GSCC or RR size with respect to removal of nodes

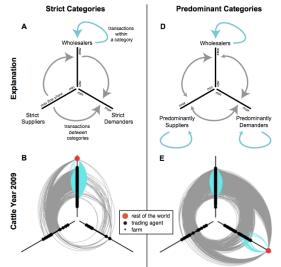


Distribution of flows by main categories based on flow share (x) and polarity (δ)

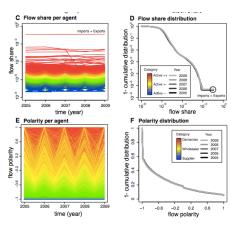
- each axis corresponds to a strict (A,B) or predominant (D,E) flow polarity (δ_i)
- distribution along axes according to flow share (x_i)
- only links with ≥ 12 transactions/yr are represented

Mid-wholesalers (intermediary levels of δ_i and x_i) play the central role in terms of flows \rightarrow potentially in terms of pathogen transmission also.

Moslonka-Lefebvre et al. (2014) Submitted.



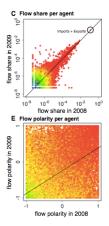
Stability over time: flow share (x) and polarity (δ)



Moslonka-Lefebvre et al. (2014) Submitted.

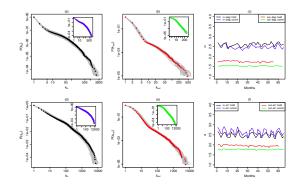
Individual trajectories of herds (all years - left, comparison 2008 vs 2009 - right)

- quite stable over time for x (C, left and right)
- more variable for δ , especially for wholesalers (E, left and right)



Topological stability over time: monthly degree (k) and strength (s) distributions (2005-2009)

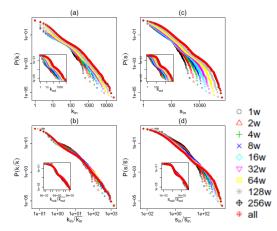
No statistically significant difference (K-S test) between monthly observed distrib. Not "far" from power-law distributions



 k^{in} (a), k^{out} (b), s^{in} (d), s^{out} (e); holdings as nodes (except for insets, commune as nodes); estimated exponents for k (c), for s (f)

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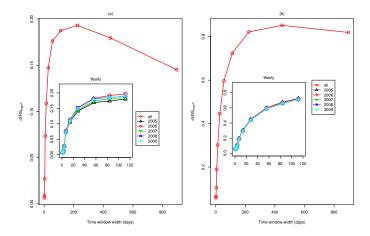
Topological stability over time: degree (k) and strength (s) distributions over increased time windows (2005-2009) For small time windows the distributions differ (K-S test, mult. testing correction). For increasing time windows distributions approach stability.



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Stability over time: common bakbone of consecutive periods assessed through *SRS*

Nodes remain active over two consecutive years ($\sim 80\%$), whereas links are much more prone to variation (less then 20% common to two consecutive years).

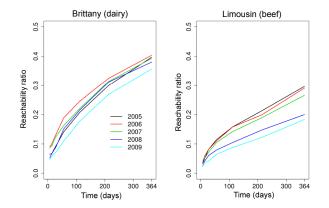


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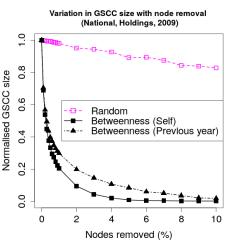
Reachability ratio (RR)

Dairy network related to a potentially larger epidemic size (expressed through the proxy RR) than beef netork in the temporal formalism (contrary to time-aggregated analysis: GSCC greater for beef than for dairy netw.)



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Percolation wrt GSCC: based on degree (k), strength (s)and betweeness centrality C_B

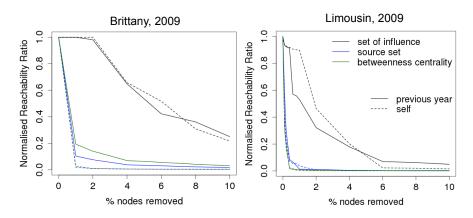


Site percolation based on nodes C_B is the most efficient strategy.

High resilience to random removal c nodes (scale-free)

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Percolation wrt RR: based on betweeness centrality C_B , set of influence, source set

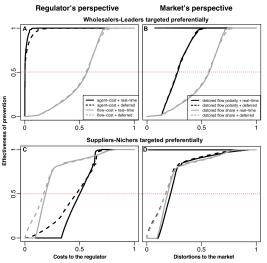


Node removal according to betweenness and source set are better strategies. Beef region seems easier to control than dairy region.

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Percolation wrt GSCC: based on flow share (x) and polarity (δ)



Targenting the WL is more efficient than SN if the cost is proportional to the number of agents to target (A-B, black curves).

The opposite occurrs if the cost is per flow unit (A-B grey curves).

SN strategies can induce less distortions than WL ones for most levels of prevention-effectiveness.

Moslonka et al. (2014) Submitted

Main implications of network analysis for infection spread

- Topological stability (of main indicators distributions) over time.
- Links related characteristics globally less stable over time than nodes related ones → important variation over time of the network backbone.
- Important to consider temporal formalism when assessing epidemiological risk and control.
- Maximizing the efficacy of interventions could lead to different targeting strategies based on criteria used (e.g. cost per node holding, vs cost per flows).
- Need of other appropriate indicators and computationally efficient programmes.



