

On Spatial Disease Mapping Models Using Directed Acyclic Graphs

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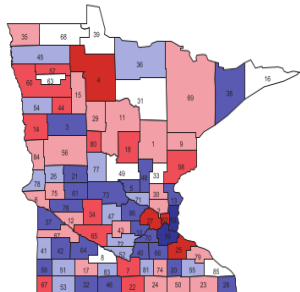


Disease Mapping: Mapping Random Effects

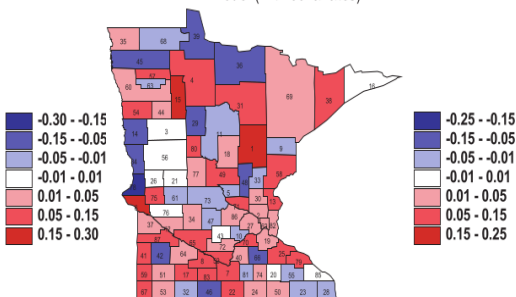
Model-based estimates of random effects across 87 counties in Minnesota

$$\begin{aligned} [\text{Infant Mortality Rates}] &= [\text{Intercept}] + [\text{Fixed Effects}] \\ &+ [\text{County-wise Random Effects}] \end{aligned}$$

I.I.D. model (without covariates)



I.I.D. model (with covariates)



- ▶ **Conditional autoregressive (CAR)** model (Besag, 1974; Clayton and Bernardinelli, 1992)
- ▶ Areal data modeled as a graph or network: V is the set of vertices (regions)
- ▶ $i \sim j$ if regions i and j share a common border
- ▶ **Adjacency matrix** $A = (a_{ij})$ such that $a_{ij} = I(i \sim j)$
- ▶ n_i is the number of neighbors of i
- ▶ CAR model:

$$w_i | w_{-i} \sim N \left(\frac{\rho}{n_i} \sum_{j | i \sim j} w_j, \tau_w n_i \right)$$

- ▶ At unit (region) i , we observe response y_i and covariate x_i
- ▶ $g(E(y_i)) = x_i^\top \beta + w_i$ where $g(\cdot)$ denotes a suitable link function

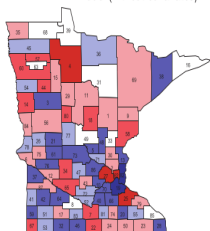
$$p_2(\beta, \tau_w, \rho) \times N(w | 0, \tau_w(D - \rho A)) \times \prod_{i=1}^k p_1(y_i | x_i^\top \beta + w_i)$$

- ▶ p_1 denotes the density corresponding to the link $g(\cdot)$

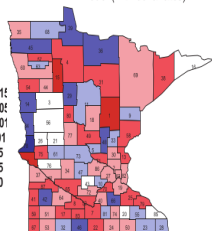
Disease Mapping: Mapping Random Effects

$$[\text{Infant Mortality Rates}] = [\text{Intercept}] + [\text{Fixed Effects}] + [\text{County-wise Random Effects}]$$

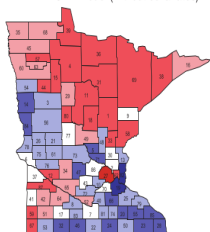
I.I.D. model (without covariates)



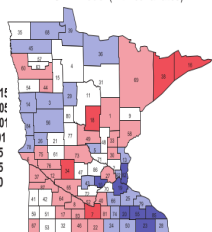
I.I.D. model (with covariates)



CAR model (without covariates)



CAR model (with covariates)



- ▶ **Simultaneous Autoregressive (SAR)** model (Whittle, 1954)
- ▶ Instead of taking the conditional route, SAR model proceeds by simultaneously modeling the random effects

$$w_i = \rho \sum_{i \neq j} b_{ij} w_j + \epsilon_i \text{ for } i = 1, 2, \dots, k$$

- ▶ $\epsilon_i \stackrel{ind}{\sim} N(0, \tau_i)$ are errors independent of w
- ▶ A common choice is to define $b_{ij} = I(i \sim j)/n_i$
- ▶ **Joint distribution:** $w \sim N(0, (I - \rho B)^\top F (I - \rho B))$, $B = (b_{ij})$ and $F = \text{diag}(\tau_1, \tau_2, \dots, \tau_k)$

- ▶ Calibration of ρ as a correlation, e.g., (as reported in Banerjee et al. 2014)

$\rho = 0.80$ yields $0.1 \leq \text{Moran's } I \leq 0.15$,

$\rho = 0.90$ yields $0.2 \leq \text{Moran's } I \leq 0.25$,

$\rho = 0.99$ yields Moran's $I \leq 0.5$

- ▶ So, used with random effects, scope of spatial pattern may be **limited**

- ▶ ρ cannot be interpreted as correlation between neighboring w_i 's (Wall, 2004; Assuncao and Krainski, 2009)

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M.M. Wall / Journal of Statistical Planning and Inference 121 (2004) 311–324

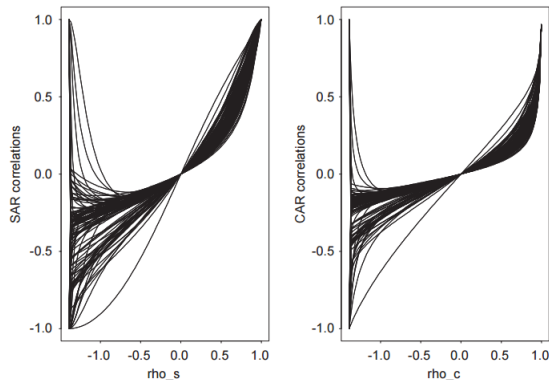


Figure: Neighbor pair correlations as a function of ρ for proper CAR and SAR models over the graph of US states

What if we construct a Directed Acyclic Graph

$$p(w) = p(w_1)p(w_2 | w_1)p(w_3 | w_1, w_2) \cdots p(w_n | w_1, w_2, \dots, w_{k-1})$$

$$w_1 = \epsilon_1 ; \quad \epsilon_1 \sim N(0, \tau_1)$$

$$w_2 = b_{21}w_1 + \epsilon_2 ; \quad N(0, \tau_2)$$

$$w_3 = b_{31}w_1 + b_{32}w_2 + \epsilon_3 ; \quad N(0, \tau_3)$$

\vdots

$$w_k = b_{k1}w_1 + b_{k2}w_2 + \dots + b_{k,k-1}w_{k-1} + \epsilon_k ; \quad N(0, \tau_k)$$

- ▶ $w = Bw + \epsilon$, where $\epsilon \sim N(0, F)$ and $F = \text{diag}(\tau_1, \dots, \tau_k)$

$B = (b_{ij})$ is now a strictly **lower triangular** matrix.

What if we construct a Directed Acyclic Graph

- ▶ **Advantages** of lower triangular B :
 - ▶ $w \sim N(0, (I - B)^\top F(I - B))$ is a **proper distribution** for any choice of lower triangular B
 - ▶ $\det(L^\top FL) = \prod_{i=1}^n \tau_i$ where $F = \text{diag}(\tau_1, \dots, \tau_k)$ and $L = I - B$
 - ▶ $w^\top L^\top FLw = \tau_1 w_1^2 + \sum_{i=2}^k \tau_i (w_i - \sum_{\{j < i\}} w_j b_{ij})^2$
 - ▶ Likelihood $N(w | 0, (I - B)^\top F(I - B))$ can be computed using **$O(k + s)$ flops** where s denotes the sparsity (number of non-zero entries) of B .
 - ▶ Even if k is large, evaluation of likelihood is fast if each region only shares border with a few others

- ▶ How to specify B and F ?
- ▶ Sparsity of B is desirable
- ▶ $b_{ij} = 0$ for j outside neighbor sets $N(i)$
 - ▶ **Pros:** For graphs with a fixed topological order, neighbor sets can be chosen: $N(i) = \{j \mid j \sim i, j < i\}$
 - ▶ **Cons:** There is no covariance function on arbitrary graphs from which we can obtain non-zero b_{ij} 's and F

- ▶ $D = (d_{ij})$ is the shortest distance matrix on the graph
- ▶ If the graph was a tree (no loops), then $\rho^D = \{\rho^{d_{ij}}\}$ is then a valid *autoregressive* correlation matrix (AR(1) model on a tree, Basseville et al., 2006).
- ▶ Areal graphs are *loopy* and are not usually trees

- ▶ **Embedded spanning trees (EST)** of a graph G is a subgraph of G which is a tree and spans all the vertices of G
- ▶ Note that to specify $w_i = \sum_{j \in N(i)} b_{ij} w_j + \epsilon_i$ we only need a joint distribution on $\{i\} \cup N(i)$
- ▶ Let G_i denote the subgraph of G which includes vertices $\{i\} \cup N(i)$ and the edges among them
- ▶ The subgraph T_i of G_i which only contains the edges $\{i \sim j \mid j \in N(i)\}$ is an embedded spanning tree of G_i
- ▶ Use the **local** embedded spanning trees T_i to specify the b_{ij} 's and τ_i

Directed acyclic graph autoregressive (DAGAR) model

- ▶ AR_i denotes the $AR(1)$ distribution on T_i
- ▶ Solve for b_{ij} and τ_i such that $E_{AR_i}(w_i | w_{N(i)}) = \sum_{j \in N(i)} b_{ij} w_j$ and $\tau_i = 1/Var_{AR_i}(w_i | w_{N(i)})$
- ▶ **No edge is left out !**

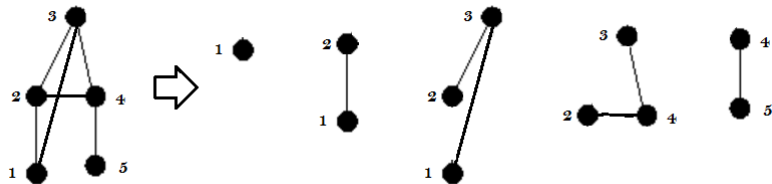


Figure: Decomposing a graph into a sequence of embedded spanning trees

- ▶ $b_{ij} = b_i = \rho / (1 + (|N(i)| - 1)\rho^2)$
- ▶ $\tau_i = (1 + (|N(i)| - 1)\rho^2) / (1 - \rho^2)$
- ▶ $\det(Q_{DAGAR}) = \prod_{i=1}^k \tau_i$
- ▶ **Positive definite** for any $0 \leq \rho \leq 1$
- ▶ **Interpretability of ρ :**
 - ▶ If the graph is a tree, then **DAGAR** model is same as the AR(1) model on the tree i.e. correlation between d^{th} order neighbors is ρ^d for $d = 1, 2, \dots$
 - ▶ If the graph is a closed two-dimensional grid, then each neighbor pair correlation is ρ
- ▶ $p_{DAGAR}(w)$ can be stored and evaluated using $O(e + k)$ flops where e is the total number of neighbor pairs

- ▶ Priors over the space of all $k!$ orderings/permutations
- ▶ Averaging over Cholesky decompositions of precision matrices:

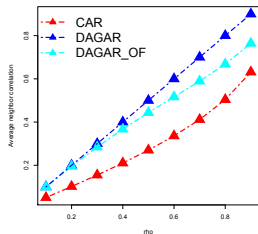
$$Q = \frac{1}{k!} \sum_{\pi} P_{\pi} L_{\pi} D_{\pi} L_{\pi}^T P_{\pi}^T$$

- ▶ The above produces closed-form expressions for Autoregressive models:

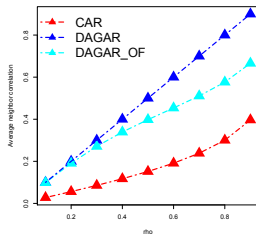
$$Q_{ii} = 1 + \frac{n_i \rho^2}{2(1 - \rho^2)} + \frac{\rho^2}{(1 - \rho^2)} \sum_{j \sim i} f(\rho, n_j)$$

$$Q_{ij} = \frac{\rho}{1 - \rho^2} I(i \sim j) + \frac{\rho^2}{1 - \rho^2} \sum_{k \in N(i) \cap N(j)} g(\rho, n_k) .$$

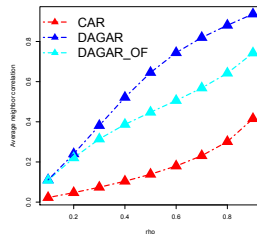
- ▶ New class of DAGAR models for areal/lattice data (Datta et al., 2017)



(a) path graph



(b) grid graph



(c) USA state map

Figure: Average neighbor pair correlations as a function of ρ for proper CAR and DAGAR models

Simulated data analysis

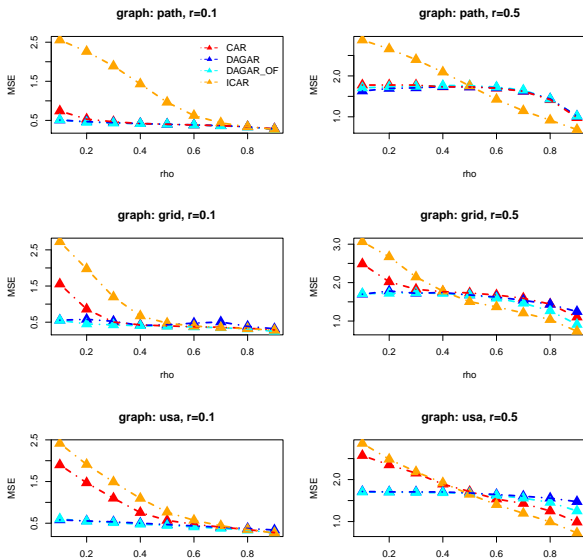


Figure: Mean square error as a function of ρ and $r = \tau^2/\sigma^2$ for DAGAR and CAR models

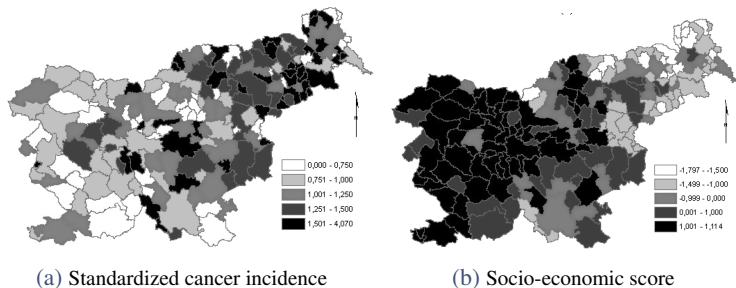


Figure: Slovenia stomach cancer data

- ▶ Observed (O_i) and expected (E_i) number of cancer counts for each of the 194 municipalities of the country
- ▶ $O_i \sim \text{Poisson}(E_i \exp(\alpha + \beta SE_i + w_i))$ where $w \sim N(0, \tau_w Q(\rho))$

Table: Parameter estimates with confidence intervals and model comparison metrics

	α	β	ρ	DIC	LPPD _{LOOCV} ¹
CAR	0.09 (0.02, 0.16)	-0.12 (-0.19, -0.04)	0.33 (0.02, 0.86)	1097	1170
DAGAR	0.11 (0.03, 0.18)	-0.12 (-0.19, -0.06)	0.08 (0.004, 0.24)	1091	1127
DAGAR _{OF}	0.11 (0.05, 0.17)	-0.12 (-0.18, -0.06)	0.06 (0.003, 0.2)	1090	1133

- ▶ Zadnik and Reich (2006) observed **spatial confounding** with ICAR model ($\hat{\beta}_{ICAR} = -0.02(-0.10, 0.06)$)
- ▶ Here for all three models the CIs for β lie outside zero
- ▶ Estimates of ρ are much smaller than 1
- ▶ Estimates of β here are closer to those obtained in the non-spatial (NS) analysis ($\hat{\beta}_{NS} = -1.4(-0.17, -0.10)$)

¹Log-predictive posterior density using Leave one out cross validation

- ▶ DAGAR models for areal data constructed from sparse Cholesky factors
- ▶ **Scalability** for large areal data
- ▶ Ordered vs order-free DAGAR
 - ▶ For all analysis, ordered model performed very similar to the order-free model
 - ▶ Ordered model is faster with theoretical results about interpretability of ρ
- ▶ DAGAR models are **positive definite** and can be directly used to model or simulate any multivariate data on graphs (like imaging or social network data)
- ▶ Better performance than CAR modes for many scenarios
- ▶ DAGAR available at
<https://arxiv.org/pdf/1704.07848.pdf>

Thank You!