# Spatial statistics in public health research: methodological opportunities and computational challenges

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# Outline

- explosion of spatial data in health research
- examples of spatial health data
- modelling spatial risk in a case-control study
  - focus on computational efficiency
- methodological and research challenges

#### Increased attention to spatial analysis in public health

#### • areal data:

- public databases and geocoding of individuals to areas
- interest in health disparities and social science questions
- focus is on covariates, not spatial structure
- point data
  - geocoding and GPS are mainstream
    - \* health outcomes can be assigned point locations
  - GIS software
    - \* easy data management and manipulation
    - \* graphical presentation
    - \* spatially-varying covariate generation
  - strong applied interest in kriging and related smoothing methods
  - opportunities for more sophisticated spatio-temporal modelling, particularly Bayesian models

- environmental exposure modelling
  - \* spatial smoothing and additive modelling of monitoring data
- mixed point and area data
  - individual locations plus area-level covariates
- multivariate responses
  - multiple pollutants, multiple health endpoints
  - latent variable modelling, causal relationships

# Socioeconomic factors in health outcomes in NSW, Australia



- challenges
  - areal (postcode) units vary drastically in size
  - computational challenge
    - \* 650 units, 5 years daily data, 2 sexes, 9 age groups
  - spatial effect and spatially-varying covariates hard to tease apart
  - data misalignment
    - \* outcome at postcode, covariate at census analogue
- relate areal data to a latent smooth process (Kelsall & Wakefield, Rathouz)

### **Combining area and individual-level information**

- area-level covariates based on point process data
  - access to contraception at health clinics in Malawi
  - accessibility of liquor retail outlets in Chicago

Observed liquor density rates per census tract, Chicago (per 1,000 population)

- spatial scale of interest is based on outcome
- consider two-stage Bayesian model so smoothing is informed by the health outcome

#### Spatial variation in allergenic response

- geocoding of new mothers' residences
- measurement of blood serum IgE immune response
- interest in variance partitioning



# **Exposure estimation in the Nurses' Health Study**

- spatial estimation of individual environmental exposures
  - often air pollution
- particulate matter (PM) exposure in large cohort of nurses
  - estimate individual exposure, 1985-2003
  - EPA monitoring for large-scale spatio-temporal heterogeneity
  - spatially-varying covariates for local heterogenity
    - \* distance to roads, climate variables, local land use, ...
    - \* generated using GIS
  - geocoding of individual residences every two years
    - relate estimated exposure to health outcomes (chronic heart disease)

geocoding and GIS make this possible; spatial statistics provides a rigorous framework



 geocoding and GIS make this possible; spatial statistics provides a rigorous framework for estimation



#### Challenges for spatio-temporal exposure estimation

- computations: 50,000 monthly pollution measurements over 20 years at 500 monitoring sites
  - kriging is difficult, particularly Bayesian implementations
  - efficient, user-friendly computation is critical (gam() in R)
  - more complicated spatio-temporal structures for better prediction, but ...
    - \* Bayesian implementation would require a statistician
    - \* more computationally efficient methods needed
- non-standard measurement error results from smoothing
- multivariate, non-Gaussian modelling
  - modelling PM2.5 based on PM10 and on airport visibilility
  - simple multivariate normality not reasonable

#### Latent variable modelling

- exposure estimation for PM in the Boston area
- which pollutant sources are responsible for health outcomes?
  - traffic is locally heterogeneous, power plant pollutants (e.g., sulfates) are not
- estimate latent traffic exposure and relate to health outcomes
- two surrogates for traffic, elemental carbon and black carbon
- hierarchical Bayesian model with multiple data sources



#### Petrochemical exposure in Kaohsiung, Taiwan



#### Possible approaches for health analysis

- Explicitly estimate pollutant exposure difficult retrospectively
- Use distance to exposure source as covariate
- Use a moving window/multiple testing to detect clusters of cases
  - default approach software available
- Include space as a covariate to provide a map of risk

$$Y_i \sim \text{Ber}(p(\boldsymbol{x}_i, \boldsymbol{s}_i))$$
$$\log (p(\boldsymbol{x}_i, \boldsymbol{s}_i)) = \boldsymbol{x}_i^T \boldsymbol{\beta} + g_{\boldsymbol{\theta}}(\boldsymbol{s}_i)$$

# Modelling challenges from a Bayesian perspective

- thousands of case-control observations difficult for Bayesian kriging
- non-Gaussian spatial models particularly difficult
  - spatial process cannot be analytically integrated out of the likelihood/posterior
  - MCMC mixing is very slow because of high-level structure
    - \* correlation amongst process values and between process values and process hyperparameters



#### **Modelling Framework**

$$Y_i \sim \text{Ber}(p(\boldsymbol{x}_i, \boldsymbol{s}_i))$$
$$\log (p(\boldsymbol{x}_i, \boldsymbol{s}_i)) = \boldsymbol{x}_i^T \boldsymbol{\beta} + g_{\boldsymbol{\theta}}(\boldsymbol{s}_i)$$

- basic spatial model for  $\boldsymbol{g}_{\theta}^s = (g_{\theta}(\boldsymbol{s_1}), \dots, g_{\theta}(\boldsymbol{s_n}))$ 
  - GAM:  $g_{\theta}(\cdot)$  is a two-dimensional smooth term
    - \* basis representation

$$\boldsymbol{g}_{ heta}^{s} = Z \boldsymbol{u}$$

\* Gaussian process representation:

$$g(\cdot) \sim \mathsf{GP}(\mu(\cdot), C_{\theta}(\cdot, \cdot)) \Rightarrow \boldsymbol{g}_{\theta}^{s} \sim N(\boldsymbol{\mu}, C_{\theta})$$

- GLMM:  $oldsymbol{g}^s_ heta=Zoldsymbol{u}$ 
  - \* correlated random effects,  $\boldsymbol{u} \sim N(\boldsymbol{0},\boldsymbol{\Sigma})$

#### **Bayesian spectral basis function model**

- computationally efficient basis function construction (Wikle 2002)
- $g^{\#} = Zu$  and  $g^s = \sigma Pg^{\#}$ 
  - piecewise constant gridded surface on k by k grid
  - P maps observation locations to nearest grid point
- Z is the Fourier (spectral) basis and Zu is the inverse FFT
- Zu is approximately a Gaussian process (GP) when...
  - $\boldsymbol{u} \sim N(0, \operatorname{diag}(\pi_{\theta}(\boldsymbol{\omega})))$  for Fourier frequencies,  $\boldsymbol{\omega}$
  - spectral density,  $\pi_{\theta}(\cdot)$ , of GP covariance function defines V( $m{u})$

#### **Bayesian spectral basis functions**



#### **Comparison with usual GP specification**

- usual GP model:  $\boldsymbol{g}^s \sim N(\boldsymbol{\mu}, C_{\theta})$ 
  - $O(n^3)$  fitting:  $|C_{\theta}|$  and  $C_{\theta}^{-1}g$
- spectral basis uses FFT
  - $O\left((k^2)\log(k^2)\right)$
  - additional observations are essentially free for fixed grid
  - fast computation and prediction of surface given coefficients
  - a priori independent coefficients give fast computation of prior and help with mixing

#### **Other approaches**

- penalized likelihood based on mixed model (radial basis functions) with REML smoothing (Kammann and Wand, 2003; Ngo and Wand, 2004) [PL-PQL]
- penalized likelihood with GCV smoothing (Wood, 2001, 2003, 2004) [PL-GCV]
- Bayesian mixed model/radial basis functions fit by MCMC (Zhao and Wand 2004) [B-Geo]
- Bayesian neural network model fit by MCMC (R. Neal) [B-NN]

#### **Simulated datasets**

- 3 case-control scenarios:  $n_0 = 1,000$ ;  $n_1 = 200$ ;  $n_{\text{test}} = 2500$  on 50 by 50 grid
- 1 cohort scenario: n = 10,000;  $n_{\text{test}} = 2500$  on 50 by 50 grid



#### **Assessment on 50 simulated datasets**



#### Mixing and speed of Bayesian methods



#### **Taiwan revisited - assessment**



#### **Assessment on count simulations**

n = 225,  $n_{\text{test}} = 2500$  on 50 by 50 grid



# **Evaluation of methods**

- Effective process parameterization = effective Bayesian estimation
  - feasible for spatial models with thousands of observations
- Natural Bayesian complexity penalty works well
  - GP representation zeroes out high-frequency coefficients as appropriate
- Implementation requires MCMC, not very accessible to practicioners
- Power is a real issue with spatial data in general, but particularly with binary observations
- Focused cluster-hunting or distance-based assessment of health risk may provide more power, but without full spatial assessment

# Methodology challenges in spatial statistics related to public health

- design and power
  - how do we choose monitoring sites?
  - when we have enough power to estimate spatial features?
  - how do we model spatial processes when monitoring data is at lower resolution than the true surface?
- surveillance and hotspot detection
  - do Bayesian methods have a place in biosurveillance and cluster detection?
    \* current applied work focuses on testing not modelling
  - surveillance likely to benefit from a decision theoretic approach that carefully considers both false positives and false negatives
- assigning one location to an individual is problematic
- variance partitioning between spatial terms and spatially-varying covariates
- confidentiality restrictions with respect to point locations and individual privacy

# General challenges for spatial statistics in public health research

- computational: big datasets and fitting of complicated models
- collaborative: developing expertise among applied researchers
- leadership
  - statisticians should be at the forefront of analyzing geographicallyindexed health data
  - we shouldn't leave this area to GIS analysts/geographers
  - necessity of providing and publicizing software for rigorous statistical methods
    - \* e.g., success of mixed model software PROC MIXED, Ime()
    - evidence of mgcv: public health researchers will learn R if useful model-building tools exist

- reproducibility: difficult to replicate analyses with complicated models, particularly MCMC implementations
  - posting code and releasing software with papers
  - standardized MCMC in R
    - \* many models, particularly new methods, can't be implemented in BUGS
      - · e.g., complicated spatio-temporal models
    - \* library of MCMC sampling functions with random variable classes
      - Jouni Kerman (Columbia) has an initial implementation for Gibbs and Metropolis sampling (umacs)
      - contributed sampling functions (e.g., slice sampling, Langevin sampling) would make this very powerful
    - reduce bugs, increase portability and reproducibility, optimize mixing