Model-Based Geostatistics for Prevalence Mapping in Low-Resource Settings

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CHICAS, Lancaster: Ole Christensen, Barry Rowlingson, Michelle Stanton, Ben Taylor, Rachel Tribbick

MLW, Blantyre, Malawi Sanie Sesay, Anja Terlouw

APOC, Ouagadougou: Hans Remme, Honorat Zoure, Sam Wanji

IRI, Columbia University: Madeleine Thomson

...and many others

Outline

- introduction
- general remarks on statistical modelling
- the standard binomial geostatistical model: Loa loa
- low-rank approximations: river blindness
- combining data from multiple surveys: malaria
- spatially structured zero-inflation: river blindness re-visited

- implementation
- closing remarks

Low resource settings



Single prevalence survey

Sample n individuals, observe Y positives

 $\textbf{Y} \sim \mathrm{Bin}(\textbf{n},\textbf{p})$

Multiple prevalence surveys

Sample n_i individuals, observe Y_i positives, i = 1, ..., m

 $Y_i \sim Bin(n_i, p_i)$?

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Extra-binomial variation

Sample n_i individuals, observe Y_i positives, i = 1, ..., m

 $Y_i | d_i, U_i \sim \operatorname{Bin}(n_i, p_i) \quad \log\{p_i/(1 - p_i)\} = d_i'\beta + U_i$

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This talk

What to do if the d_i and/or the U_i are spatially structured

- traditionally, a self-contained methodology for spatial prediction, developed at École des Mines, Fontainebleau, France
- nowadays, that part of spatial statistics which is concerned with data obtained by spatially discrete sampling of a spatially continuous process

A geostatistical data-set: Loa loa prevalence surveys



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Model-based Geostatistics (Diggle, Moyeed and Tawn, 1998)

- the application of general principles of statistical modelling and inference to geostatistical problems
 - formulate a model for the data
 - use likelihood-based methods of inference

- answer the scientific question

Statistical modelling principles

- models are devices to answer questions
- models should:
 - be not demonstrably inconsistent with the data;
 - incorporate the underlying science, where this is well understood
 - be as simple as possible, within the above constraints

"Too many notes, Mozart"

Emperor Joseph II

"Only as many as there needed to be"

Mozart (apochryphal?)

Empirical modelling: The AEGISS project (Diggle, Rowlingson and Su, 2005)

- early detection of anomalies in local incidence
- data on 3374 consecutive reports of non-specific gastro-intestinal illness
- log-Gaussian Cox process, space-time correlation ρ(u, v)



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Mechanistic modelling: the 2001 UK FMD epidemic (Diggle, 2006)

- Predominantly a classic epidemic pattern of spread from an initial source
- Occasional apparently spontaneous outbreaks remote from prevalent cases
- λ(x,t|H_t) =conditional intensity, given history H_t



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Onchocerciasis (River Blindness)



African Programme for Onchocerciasis Control

- "river blindness" endemic in wet tropical regions
- donation programme of mass treatment with ivermectin
- approximately 60 million treatments to date, in 19 countries
- serious adverse reactions experienced by some patients highly co-infected with *Loa loa* parasites
- precautionary measures put in place before mass treatment in areas of high *Loa loa* prevalence

http://www.who.int/pbd/blindness/onchocerciasis/en/



...and old



The Loa loa prediction problem

Ground-truth survey data

- random sample of subjects in each of a number of villages
- blood-samples test positive/negative for Loa loa

Environmental data (satellite images)

- measured on regular grid to cover region of interest
- elevation, green-ness of vegetation

Objectives

- predict local prevalence throughout study-region (Cameroon)
- compute local exceedance probabilities,

P(prevalence > 0.2|data)

Schematic representation of Loa loa model



location

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- use relationship between environmental variables and ground-truth prevalence to construct preliminary predictions via logistic regression
- use local deviations from regression model to estimate smooth residual spatial variation
- model-based approach acknowledges uncertainty in predictions

"The answer to any prediction problem is a probability distribution"

Peter McCullagh

Loa loa: a generalised linear model

• Latent spatially correlated process

$$\mathsf{S}(\mathsf{x}) \sim \mathrm{SGP}\{\mathsf{0}, \sigma^2,
ho(\mathsf{u}))\}
ho(\mathsf{u}) = \exp(-|\mathsf{u}|/\phi)$$

• Linear predictor (regression model)

$$\begin{split} d(\mathbf{x}) &= \text{environmental variables at location } \mathbf{x} \\ \eta(\mathbf{x}) &= d(\mathbf{x})'\beta + S(\mathbf{x}) \\ p(\mathbf{x}) &= \log[\eta(\mathbf{x})/\{1 - \eta(\mathbf{x})\}] \end{split}$$

• Conditional distribution for positive proportion Y_i/n_i $Y_i|S(\cdot) \sim Bin\{n_i, p(x_i)\}$ (binomial sampling)

Conditional dependence structure

Signal: S, S* (data-locations and prediction locations)

Data: Y (data-locations only)

Parameters: β (regression terms), θ (covariance structure)



logit prevalence vs elevation





How useful is the geostatistical modelling?



Predicted prevalence - 'without ground truth data'

Logistic regression



Predicted prevalence - 'with ground truth data' (%)

Model-based geostatistics

Probabilistic exceedance map for Cameroon (Diggle et al, 2007)



Jigure 6: PCM for /high risk/ in Cameroon based on ERMr with ground truth data.

- non-spatial extra-binomial variation
- low-rank approximations;
- combining data from multiple surveys
 - randomised and non-randomised

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- at different times
- spatially structured zero-inflation.

Non-spatial extra-binomial variation

Latent spatially correlated process

 $\mathsf{S}(\mathsf{x}) \sim \mathrm{SGP}\{\mathbf{0}, \sigma^2,
ho(\mathsf{u}))\} \quad
ho(\mathsf{u}) = \exp(-|\mathsf{u}|/\phi)$

- Latent spatially independent random effects $U_i \sim iidN(0, \nu^2)$
- Linear predictor (regression model)

$$\begin{split} & \mathsf{d}(x) = \mathrm{environmental variables at location } x \\ & \eta(x_i) = \mathsf{d}(x_i)'\beta + \mathsf{S}(x_i) + \mathsf{U}_i \\ & \mathsf{p}(x_i) = \mathsf{log}[\eta(x_i)/\{1 - \eta(x_i)\}] \end{split}$$

Conditional distribution for positive proportion Y_i/n_i

 $\mathbf{Y}_i | \mathbf{S}(\cdot) \sim Bin\{n_i, p(x_i)\}$ (binomial sampling)

Low-rank approximations (Rodrigues and Diggle, 2010)

$$\mathsf{S}(\mathsf{x}) pprox \mu + \sum_{\mathsf{j}=1}^{\mathsf{M}} \mathsf{w}(\mathsf{x} - \mathsf{k}_\mathsf{j}) \mathsf{Z}_\mathsf{j}$$

- w(u): kernel function
- $Z_j \sim \text{iid } N(0, \nu^2)$
- $k_j \in A \subset {\rm I\!R}^2$: fixed set of points

Choose $w(\cdot)$ to approximate to preferred family of correlation functions

Computation linear in number of prediction points

Application: onchocerciasis mapping Africa-wide (Zoure et al, 2014): 14,473 survey locations



Application: onchocerciasis mapping Africa-wide (Zoure et al, 2014): low-rank model

- M = 10734 points X_j in regular lattice at spacing 0.1 degrees
- to approximate Matérn correlation, $M(\phi,\kappa)$, $\kappa = 2$

$$\mathsf{w}(\mathsf{u}) = \phi^{-1} \exp(-2\sqrt{2}\,\mathsf{u}/\phi)$$

Parameter	estimate	95% confidence interval
μ	2:451	(2.469, 2.432)
$ u^2 $	31:570	(31.038, 32.112)
ϕ	65:208	(64.993, 66.301)

Application: onchocerciasis mapping Africa-wide (Zoure et al, 2014): prevalence estimates



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Application: onchocerciasis mapping Africa-wide (Zoure et al, 2014): exceedance probabilities



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Multiple surveys (Giorgi et al, 2015)

$$\begin{split} \text{Surveys: } i &= 1, \dots, r \quad \text{locations } x_{ij} : j = 1, \dots, n_i \\ \eta_{ij} &= d(x_{ij})^\top \beta_1 + S_i(x_{ij}) + I(i \in \mathcal{B})[B_i(x_{ij}) + d(x_{ij})'\beta_i] + U_{ij} \end{split}$$



Application: malaria mapping, Chikhwawa district, Malawi (Giorgi et al, 2015): rMIS individual locations



Longitude

Application: malaria mapping, Chikhwawa district, Malawi (Giorgi et al, 2015): eMIS individual locations



Longitude
Application: malaria mapping, Chikhwawa district, Malawi (Giorgi et al, 2015): EAG village locations and prevalences



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Application: malaria mapping, Chikhwawa district, Malawi (Giorgi et al, 2015): estimated prevalence map



Application: malaria mapping, Chikhwawa district, Malawi (Giorgi et al, 2015): estimated bias maps



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Continuous time: rolling malaria indicator surveys

Hotspots: P(prevalence > 20%)

Continuous time: rolling malaria indicator surveys

Coldspots: P(prevalence < 5%)

Spatially structured zero-inflation: river blindness re-visited

- public health experts have strong sense that some areas are fundamentally unsuitable for onchocerciasis transmission
- hence need to incorporate mix of structural and chance zeros

Non-spatial model

$$\label{eq:Yi} Y_i \sim \left\{ \begin{array}{rrr} 0 & : & \text{wp } q_i \\ \text{Bin}(n_i,p_i) & : & \text{wp } 1-q_i \end{array} \right.$$

Spatial model

 $\{q_i,p_i\} \to \{Q(x),P(x)\}: x \in \mathbb{R}^2 \sim \text{bivariate stochastic process}$

$$\mathsf{P}(\mathsf{Y} = \mathsf{y}|\mathsf{S}_1(\mathsf{x}),\mathsf{S}_2(\mathsf{x})) = \begin{cases} \mathsf{Q}(\mathsf{x}) + (1 - \mathsf{Q}(\mathsf{x})) \times \mathsf{Bin}(0;\mathsf{n},\mathsf{p}(\mathsf{x})) & : \mathsf{y} = 0\\ (1 - \mathsf{Q}(\mathsf{x})) \times \mathsf{Bin}(\mathsf{y};\mathsf{n},\mathsf{p}(\mathsf{x})) & : \mathsf{y} > 0 \end{cases}$$

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$$logit(Q(x)) = \mu_1 + S_1(x)$$

• logit(P(x)) =
$$\mu_2$$
 + S₂(x)

• $\{S_1(x),S_2(x)\}\sim$ bivariate Gaussian process

Sudan: probability exceedance map



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Sudan: non-transmissible probability map (Q(x))



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Mozambique/Malawi/Tanzania: probability exceedance map



Mozambique/Malawi/Tanzania: non-transmissible probability map (**Q(x)**)



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• Monte Carlo maximum likelihood

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- Plug-in prediction
- R package PrevMap
- Bayesian version?

Closing remarks

principled statistical methods

- make assumptions explicit
- deliver optimal estimation within the declared model
- make proper allowance for predictive uncertainty
- but there is no such thing as a free lunch

"We buy information with assumptions"

C H Coombs

- which is why statistics is at its most effective when conducted as a dialogue with substantive science
- and this should guide the way we teach statistics ...especially to science students