

Disease mapping with Gaussian processes

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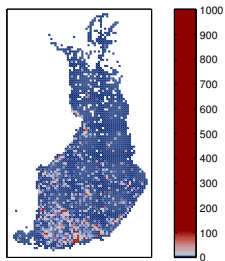


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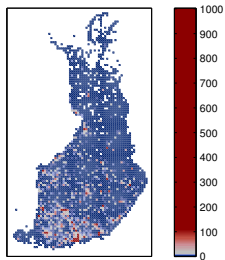
- Example: Alcohol related deaths in Finland
- Spatial priors and benefits of GP prior
- Computation and approximations
- Spatio-temporal
- Explanatory variables
- Integration over the latent space
- Hyperparameters

Example: deaths in Finland

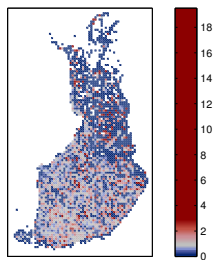


(a) Number of deaths

Example: deaths in Finland

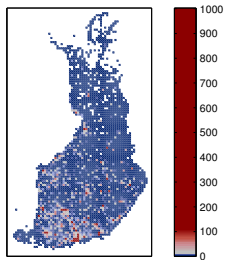


(d) Number of deaths

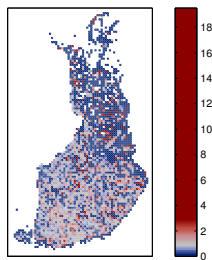


(e) Raw relative risk

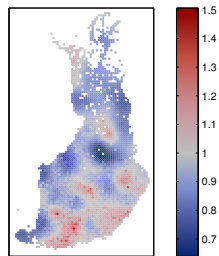
Example: deaths in Finland



(g) Number of deaths



(h) Raw relative risk



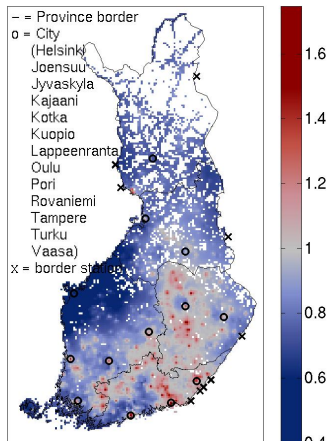
(i) Smoothed risk

Example: alcohol related diseases in Finland

- Collaboration: The National Institute for Health and Welfare
- Data: Statistics Finland
- Population of Finland: ≈ 5.3 million
- About 10 500 inhabited $5\text{km} \times 5\text{km}$ cells in Finland
 - many cells with no inhabited neighbors
- In 2001–2005 about 7 900 died due to alcohol diseases (more than five times compared to deaths due to traffic)
 - expected death count less than one per cell

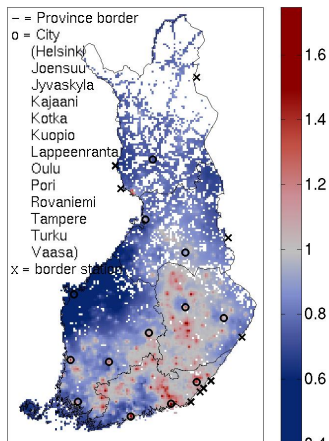
Example: alcohol related diseases in Finland

- Sex-age-education standardized expected death counts used to compute the raw risk
- Risk smoothed using GP with long and short length scale and negative-binomial observation model



Example: alcohol related diseases in Finland

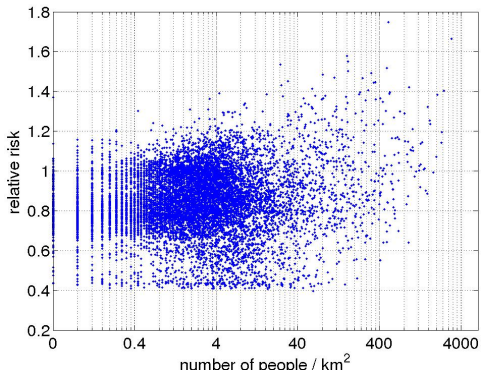
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Is the relative risk
higher in the population
centers?

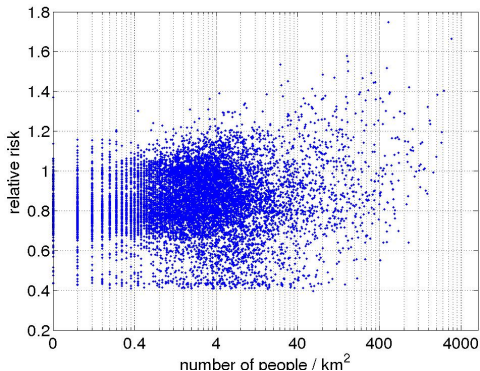
Example: alcohol related diseases in Finland

- The smoothed relative risks vs. the population density



Example: alcohol related diseases in Finland

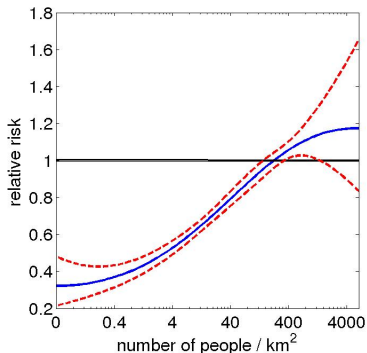
- The smoothed relative risks vs. the population density



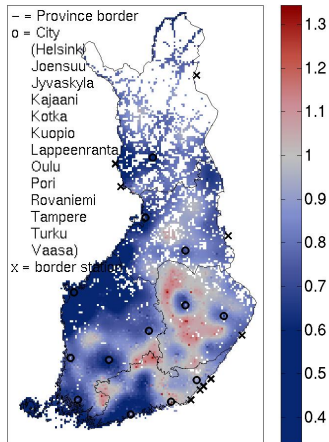
- Add population density as explanatory variable

Example: alcohol related diseases in Finland

- Population density and spatial variation explain the variation in the risk



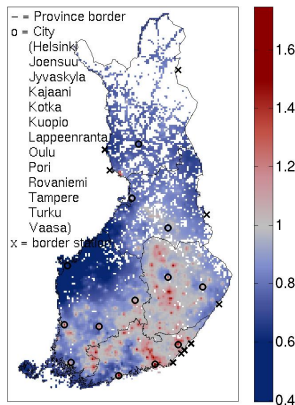
Population density effect



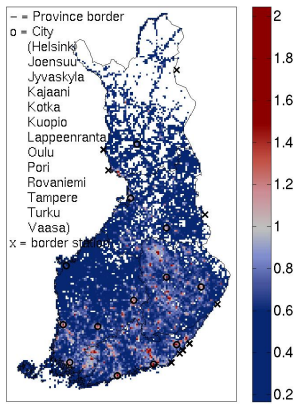
Spatial effect

Example: alcohol related diseases in Finland

- Adding explanatory covariate can change the picture



1) Spatial



2) Spatial+covariate

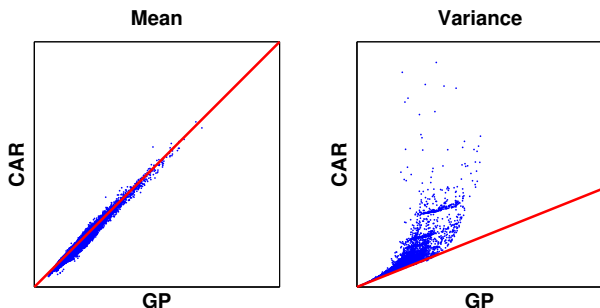
- Jarno Vanhatalo, Ville Pietiläinen and Aki Vehtari (2010). Approximate inference for disease mapping with sparse Gaussian processes. *Statistics in Medicine*, 29(15):1580-1607.
<http://dx.doi.org/10.1002/sim.3895>
- Jarno Vanhatalo, Pia Mäkelä and Aki Vehtari (2010). Regional differences in alcohol mortality in Finland in the early 2000s. http://becs.aalto.fi/en/research/bayes/publications/Vanhatalo_etal_Alcohol_mortality_in_Finland.pdf

- In spatial epidemiology CAR is most used model
- Correlation defined conditionally based on a neighborhood structure → discrete definition
 - + major computational speed-up if a precision matrix is sparse due to small neighborhoods
 - describes only local correlation
 - neighborhood definition may be difficult for irregularly spaced data and high dimensional data

Example: alcohol related diseases in Finland

Comparison to CAR

- Compared to CAR computed with INLA software
 - CAR model lacks long range correlation part
 - CAR model has much higher variance, especially for cells having no or few inhabited neighbors
 - GP has a better predictive performance



- Markov random field prior can be good
 - e.g. INLA-software can approximate Matérn covariance function with MRF
 - but precision matrix is not going to be sparse in high dimensional cases ($d \geq 3$), e.g. INLA-software doesn't support $d > 3$ and limited support for $d = 3$

- Full $O(n^3)$
- short range dependencies
 - Markov \rightarrow sparse precision matrix
 - compact support \rightarrow sparse covariance matrix
 - $O(p^3 n^3)$, where $0 < p < 1$ is the proportion of non-zeros
- long range dependencies
 - reduced rank (e.g. FIC) $O(nm^2)$
 - SVI-GP $O(m^3)$ (Hensman et al, 2013)

Reduced rank approximations and inducing points

- The correlation structure of FIC with different choices of inducing inputs

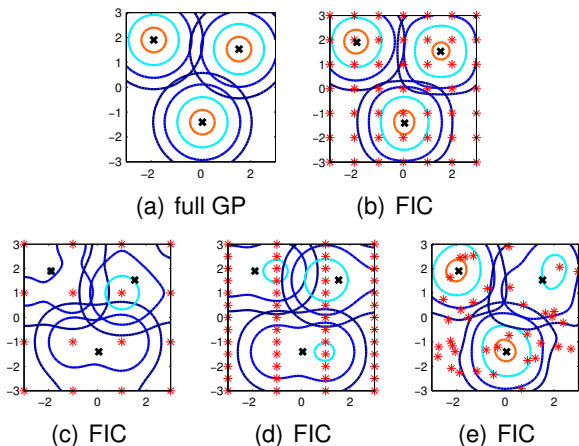


Figure: The correlation for 3 locations \mathbf{x} . Inducing inputs are marked with $*$.

- No single approximation which works efficiently for both short and long range dependencies

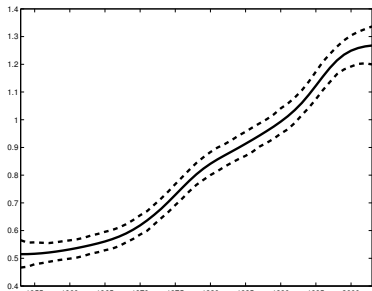
- No single approximation which works efficiently for both short and long range dependencies
- Short and long range dependencies
 - e.g. compact support + FIC (used in alcohol study)
Vanhatalo, Pietiläinen, Vehtari, Stat in med, 2010,
<http://dx.doi.org/10.1002/sim.3895>

- Full $O(n^3 T^3)$
- Markov / compact support / reduced rank
- INLA-software: unstructured interaction (ie. no model for spatio-temporal jointly)
- Cseke et al - discrete spatio-temporal model, sparse precision, restricted sparse messages
- infinite-dimensional filtering $O(n^3 T)$ ($O(nm^2 T)$)
Simo Särkkä talks about this tomorrow

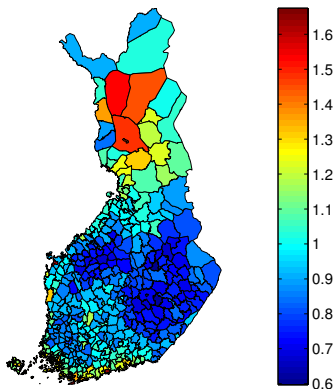
Example: lung cancer women

- County incidences and background population for years 1953–2003.
- 51 years, 431 counties → 21 981 observations
- Data: Finnish Cancer Registry
- Model: GP with temporal + spatial + spatiotemporal component

Example: lung cancer women

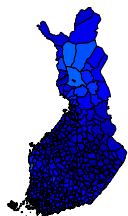


(a) Temporal

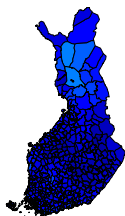


(b) Spatial

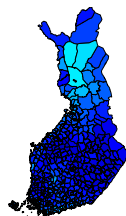
Example: lung cancer women



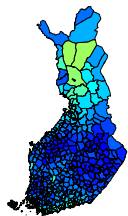
(a) 1953



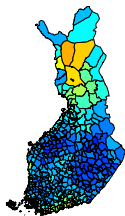
(b) 1963



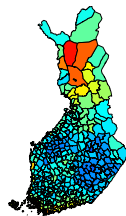
(c) 1973



(d) 1983



(e) 1993



(f) 2003

- Spatio-temporal GPs can be written as linear stochastic partial differential equations (SPDE)
- Reduces computational complexity from $O(n^3 T^3)$ to $O(n^3 T)$, i.e. method scales linearly in T
- SPDEs make it easier to specify non-stationary temporal dynamics, which are necessary, for example, when performing future predictions
- n limited as for spatial GP
 - few thousand with no sparse approximations
 - more than ten thousand with sparse approximations
- Has been tested with over million spatio-temporal points
- Simo Särkkä talks more about this tomorrow

Spatio-temporal malaria models?

- Spatio-temporal GPs can be written as linear stochastic partial differential equations (SPDE).
- SPDEs make it easier to specify non-stationary temporal dynamics, which are necessary, for example, when performing future predictions
 - seasonal variation
 - transmission dynamics with SPDEs?

- SPDEs make it easier to specify non-stationary temporal dynamics
- Spatial non-stationarity
 - deformations
 - additional GP for latent signal magnitude or length-scale

- Goal is to explain the spatial variation
- Spatial maps can be used to aid hypothesis generation
- Adding covariates hopefully makes the residual in spatial domain unstructured
- GP can model non-linearities and interactions implicitly

- 1043 cases of acute myeloid leukemia in adults
 - recorded between 1982 and 1998 in the North West Leukemia Register in the United Kingdom
 - log-logistic model for survival times (16% were censored)
 - predictors are
 - age
 - sex
 - white blood cell count (WBC) at diagnosis
 - the Townsend score which is a measure of deprivation for district of residence

Leukemia survival times

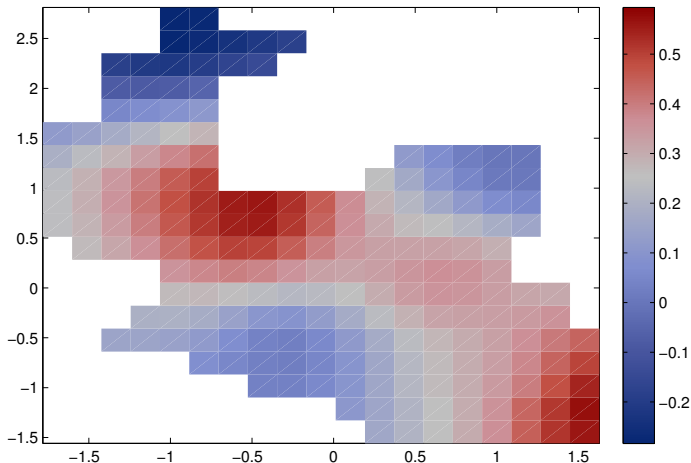
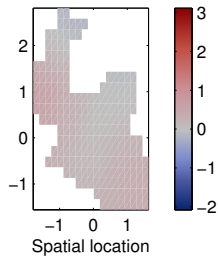
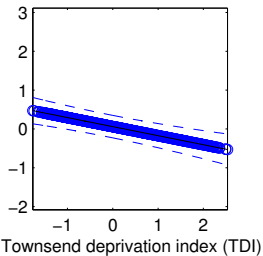
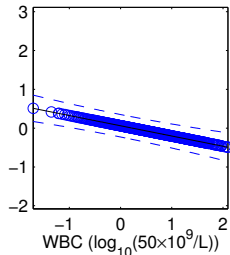
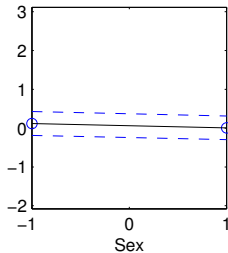
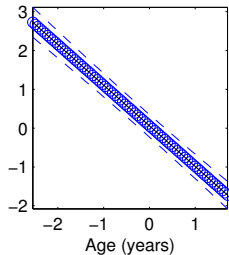
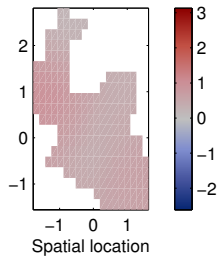
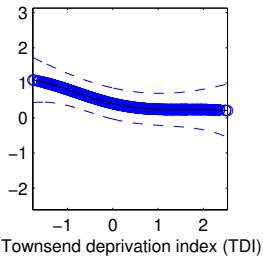
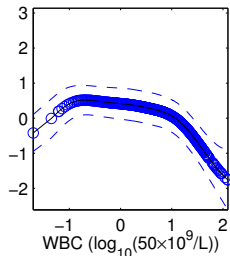
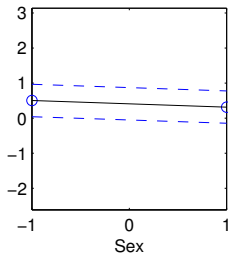
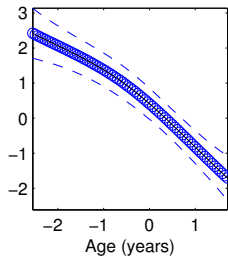


Figure: Posterior mean of the latent function

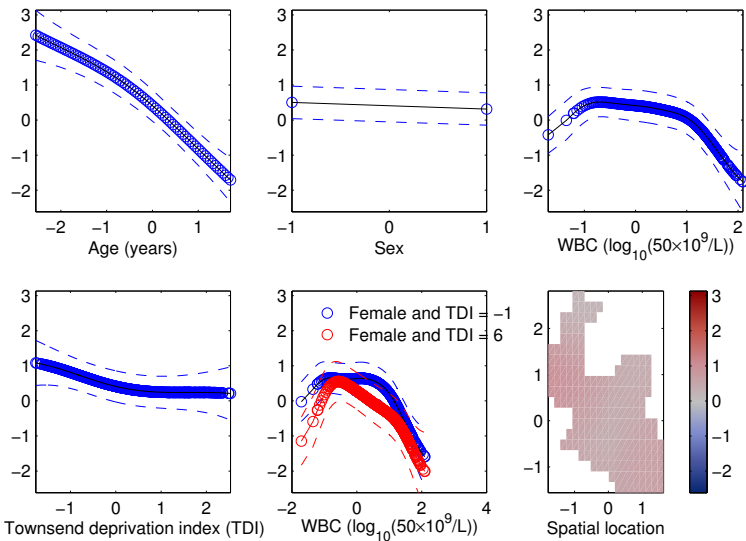
Leukemia survival times



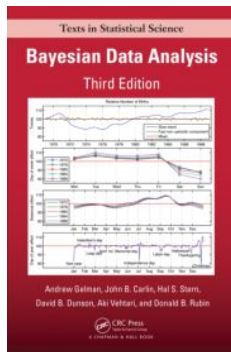
Leukemia survival times



Leukemia survival times



Analysis in GP chapter of



Andrew Gelman, John B. Carlin, Hal S. Stern, David B. Dunson, Aki Vehtari and Donald B. Rubin (2013). [Bayesian Data Analysis](#), Third Edition. Chapman and Hall/CRC.

- GP can model non-linearities and interactions implicitly
- INLA-software using MRFs allows additive effects and 2D interactions

- Multitask / multioutput GPs
 - just add the disease type as a covariate

- Non-Gaussian models, e.g., $y \sim \text{Poisson}(\alpha \exp(f(s, \theta)))$
- We are interested in predictions $p(y_i | s_i)$
- Integration over the latent variables f_i and hyperparameters θ required

- In our experiments
 - EP about as good as MCMC, but **much** faster
 - Laplace almost as good as EP, but somewhat faster
 - VB not as good as EP, but YMMV
 - difference is negligible for many likelihoods given larger datasets
 - differences in classification and with non-log-concave likelihoods

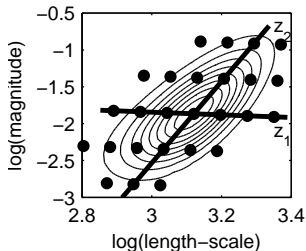
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- Mysterious Sheffield-method? (Hensman et al, submitted)
- I think that in most cases distributional approximations ok
 - If not, pseudo-marginal likelihood approach (Filippone & Girolami, 2013) might be the best choice for MCMC

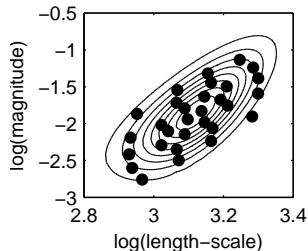
- Type II MAP
 - works well when the number of hyperparameters is small and n is big
- Adaptive grid 1–3 hyperparameters
- CCD
 - 1–15 hyperparameters \rightarrow 3–287 integration points
 - usually works well, but sometimes underestimates the uncertainty

- Type II MAP
 - works well when the number of hyperparameters is small and n is big
- Adaptive grid 1–3 hyperparameters
- CCD
 - 1–15 hyperparameters \rightarrow 3–287 integration points
 - usually works well, but sometimes underestimates the uncertainty
- Linear approximation (Garnett, Osborne, Hennig, 2013)
- EP can be used to integrate over noise and signal variances (other hyperparameters in theory, but not fast (yet?))
- MCMC

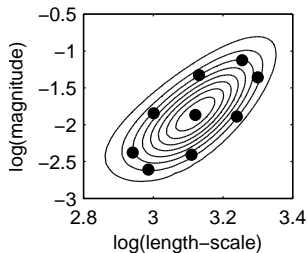
Hyperparameters



(a) Grid based



(b) Monte Carlo



Code available in Matlab/Octave (RccpOctave for R) toolbox
GPstuff

Jarno Vanhatalo, Jaakko Riihimäki, Jouni Hartikainen, Pasi Jylänki, Ville Tolvanen and Aki Vehtari (2013). GPstuff: Bayesian Modeling with Gaussian Processes. In Journal of Machine Learning Research, 14(Apr):1175-1179.

[http://www.jmlr.org/papers/volume12/
jylanki11a/jylanki11a.pdf](http://www.jmlr.org/papers/volume12/jylanki11a/jylanki11a.pdf)

GPstuff homepage: <http://becs.aalto.fi/en/research/bayes/gpstuff/>

- Sparse models
 - Compactly supported covariance functions
 - Fully and partially independent conditional (FIC, PIC)
 - Compactly supported plus FIC (CS+FIC)
 - Variational sparse (VAR), Deterministic training conditional (DTC), Subset of regressors (SOR)
- Latent inference
 - Laplace, EP, Parallel EP, Robust-EP
 - marginal posterior corrections (cm2 and fact)
 - Scaled Metropolis, Scaled HMC, Elliptical slice sampling
- Hyperparameter inference
 - Type II ML/MAP
 - Leave-one-out cross-validation (LOO-CV)
 - Metropolis, HMC, No-U-Turn-Sampler (NUTS), Slice Sampling (SLS), Surrogate SLS, Shrinking-rank and Cov-matching SLS
 - Grid, CCD, Importance sampling

- Researchers
 - Jaakko Riihimäki
 - Ville Tolvanen
 - Simo Särkkä
 - Jarno Vanhatalo
 - Jouni Hartikainen
 - Ville Pietiläinen
- Collaborators and data
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 - Helsinki University Central Hospital
 - The National Institute for Health and Welfare
 - Finnish Cancer Registry
 - Statistics Finland