

# De Wijs Process and Modeling Disease Risk

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## Outline for the talk

- **overview**
- Two approaches to disease mapping
  - **Gaussian Markov random fields**
  - **Geostatistical**
- Combining the above two approaches via **de Wijs Process**
- End with an example on **disease mapping**

## Overview

Interest on **maps of risks** of non-infectious diseases over a **geographic region**.

These maps

- reveal heterogeneity
- indicate areas of high relative risk that may require screening and intervention
- help detect effects of socio-economic conditions and exposure to environmental hazards
- provide clues to the etiology of diseases
- use counts aggregated over zip-codes, counties or states to protect individual identities.

Examples: **Clayton and Kaldor (1987)**, **Besag et al. (1991)**, **Best et al. (2001)**, **Kelsall and Wakefield (2002)**, **Banerjee et al. (2004)** and **Rue and Held (2005)**....

Two statistical approaches are **Markov random fields** and **geostatistics**.

## Example of irregular regions



## Markov random field approach (Besag et al., 1991, Rue and Held, 2005)

Consider the **conditional Poisson model**

$$Y(A_i) \sim \text{independent Poisson}\left(c(A_i)R(A_i)\right), \quad i = 1, \dots, n, \quad (1)$$

with

$$\log R(A_i) = T(A_i)\beta + U(A_i) + \epsilon(A_i),$$

$y(A_i)$  = **observed number** of cases of disease in region (e.g., county)  $A_i$

$c(A_i)$  = expected number of cases of disease in region  $A_i$

$R(A_i)$  = **log relative risk** in region  $A_i$

where  $T(A_i)$  = area-specific covariate,

$\beta$  = covariate effect

$\epsilon(A_i)$  = **unstructured component** or residual

$U(A_i)$  = A Markov random field

Assumes two areas are **adjacent** or **neighbors** when they share a common boundary.

Conditionally,  $U(A_i)$  follows  $N\left(\sum_{j \sim i} U(A_j)/l_i, \sigma_U^2/l_i\right)$ .

$j \sim i$  implies that area  $A_j$  is adjacent to  $A_i$  and  $l_i$  is number of neighbors of  $A_i$ .

## Geostatistical approach (Kelsall and Wakefield (2002))

Consider the **conditional Poisson model** driven by a Cox process

$$Y(A_i) \sim \text{independent Poisson} \left( \sum_k \int_{A_i} c_k(s) R_k(s) ds \right), \quad i = 1, \dots, n. \quad (2)$$

with

$$R_k(s) = R(s), \quad R(A_i) = \int_{A_i} f_i(s) e^{T(s)\beta + U(s) + \epsilon(s)} ds, \quad (3)$$

where

- $k$  = population stratum,  $R_k(s)$  log relative risk in stratum  $k$ , location  $s$
- $f_i(s)$  = weighted average of the stratum-specific population density
- $c(A_i)$  =  $\sum_k \int_{A_i} c_k(s) ds$  is expected number of cases of disease in region  $A_i$
- $T(s)$  = location-wise covariate information
- $\beta$  = covariate effect
- $\epsilon(s)$  = Gaussian white noise process
- $U(s)$  = A continuum spatial process

Includes **area** and **population information**, but computationally challenging.

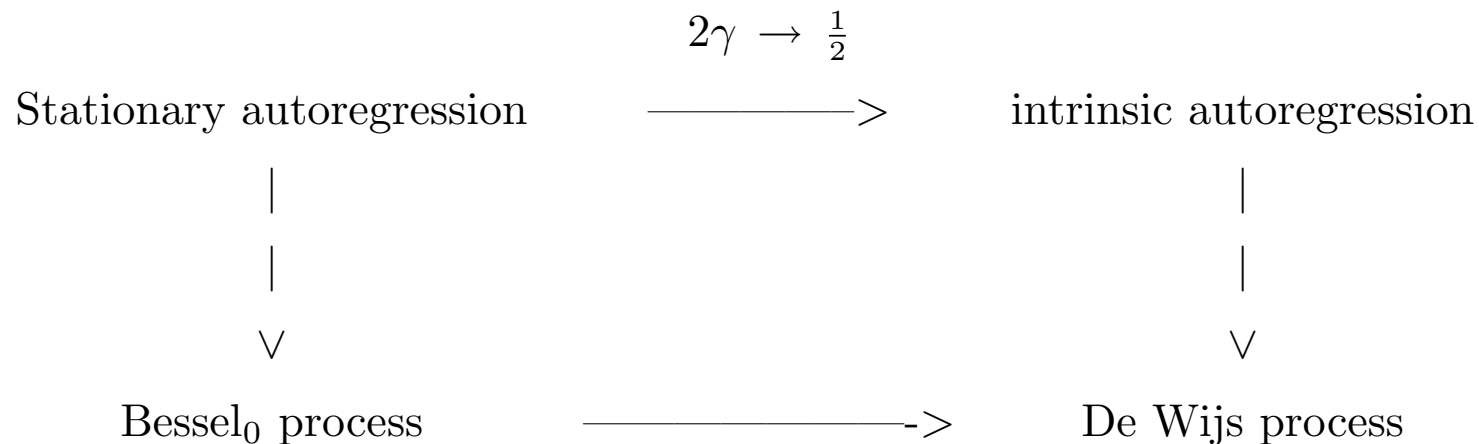
## Combining Markov random fields with geostatistical approach

Let  $\{X_{u,v} : (u,v) \in \mathcal{Z}^2\}$  be Gaussian with **conditional** means and variances

$$E(X_{u,v} | \dots) = \gamma(x_{u-1,v} + x_{u+1,v} + x_{u,v-1} + x_{u,v+1}), \quad \text{var}(X_{u,v} | \dots) = \sigma_X^2, \quad (4)$$

- (i)  $2\gamma < \frac{1}{2} \Rightarrow$  stationary autoregression e.g. Lévy (1948), Besag (1974).
- (ii)  $2\gamma = \frac{1}{2} \Rightarrow$  **intrinsic** case: Künsch (1987), Besag & Kooperberg (1995).

**Limit diagram** for stationary and intrinsic case



**Idea is to use exponential functional of the de Wijs process.**

In other words, use  $e^{X_{u,v}}$  to **approximate**  $U(s)$ !!!

## Combining Markov random fields with geostatistical approach

Embed study region in a fine mesh

$$\mathcal{L} = \{(u, v) : u = 1, \dots, I, v = 1, \dots, J\}, \quad (5)$$

Each grid point  $(u, v)$  is associated with a regular cell  $D(u, v)$ . Use

$$A_i = \cup_{(u,v) \in A_i} D(u, v). \quad (6)$$

Let  $m(A_i)$  be equal to the number of cells  $D(u, v)$  in  $A_i$ . Consider

$$f_i(u, v) = \int_{D(u,v)} f_i(s), ds.$$

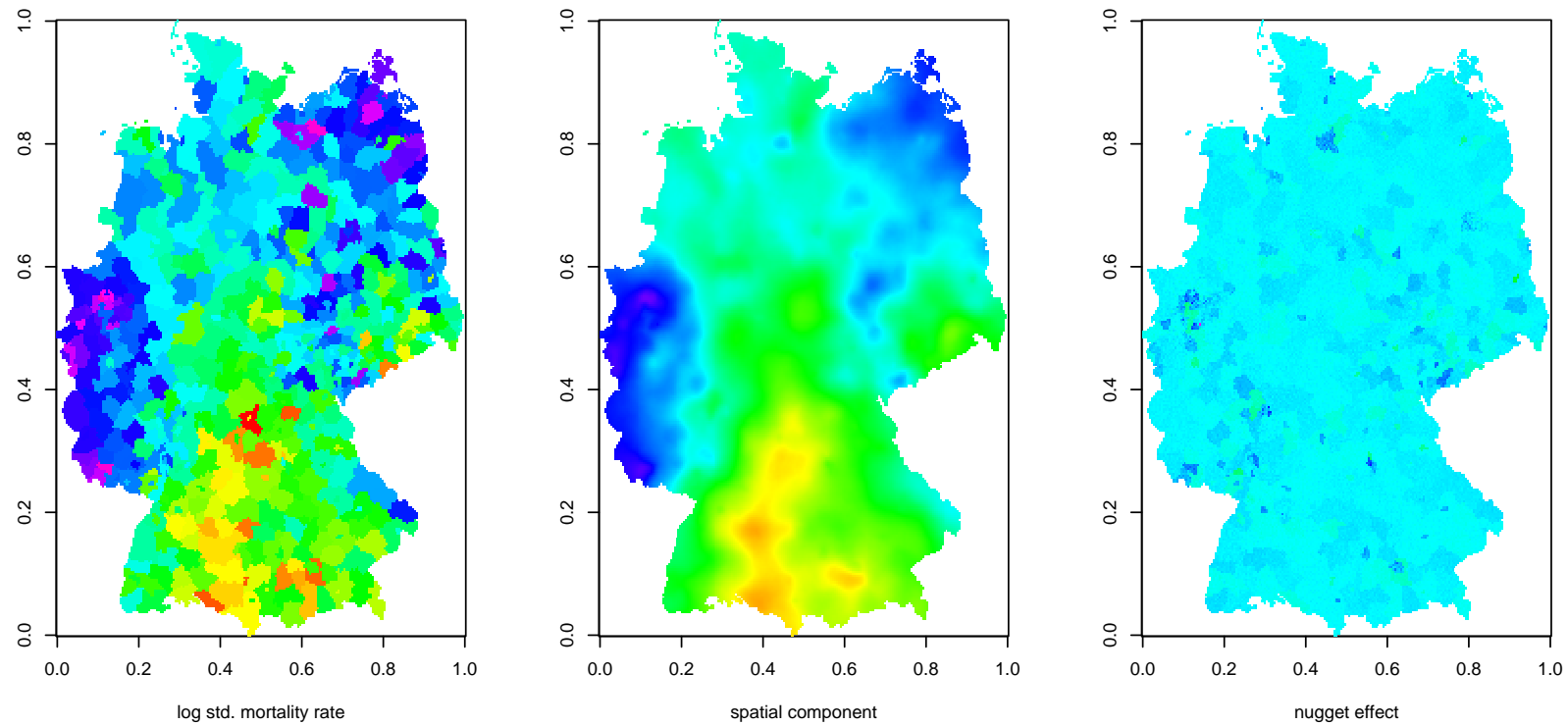
Finally, use

$$R(A_i) = \sum_{(u,v) \in A_i} f_i(u, v) e^{T(u,v)\beta + X_{u,v} + \epsilon_{u,v}}. \quad (7)$$

The **continuum random field theory** for another day!

This theory relies on **generalized Gaussian process**, **subordinate random fields** and the famous **Dobrushin's expansion**.

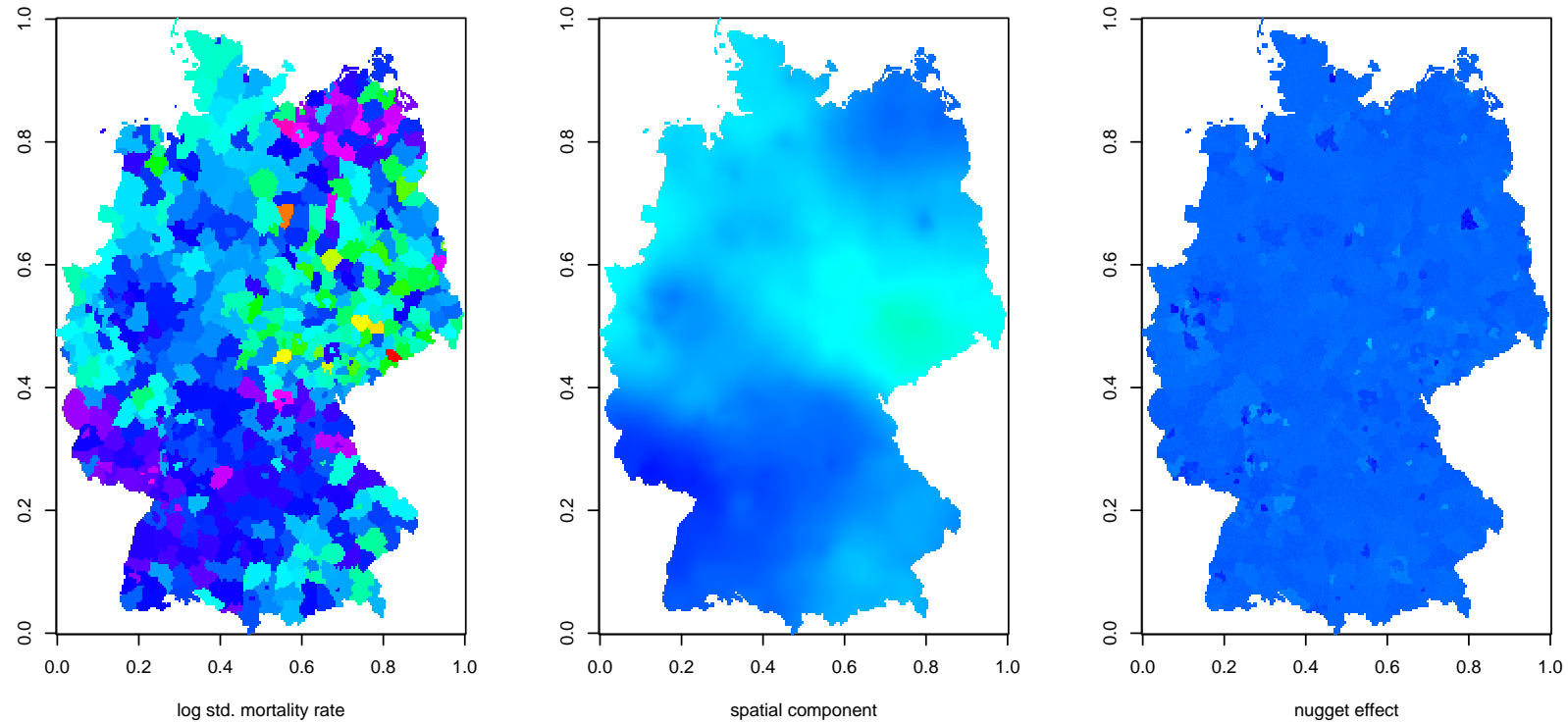
## Lung cancer in Germany



Data in the period 1986-1990. 544 districts embedded in  $289 \times 214$  grids.

See also Rue and Held, 2005.

## Oral cancer in Germany



Data during the years 1986-1990. 544 districts embedded in  $289 \times 214$  grids.

Joint disease mapping can also be done!

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