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Bayesian spatio-temporal modeling for identifying unusual and unstable trends in mammography utilization

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ABSTRACT

Objectives: To develop and compare two Bayesian models capable of identifying unusual and unstable temporal patterns in spatio-temporal data.

Setting: Annual counts of mammography screening users from each statistical local area (SLA) in Brisbane, Australia, recorded between 1997 and 2008 inclusive.

Primary outcome measures: Mammography screening counts

Results: The temporal trends of 93 SLAs (59.24%) were dissimilar to the overall common temporal trend. SLAs which did follow the common temporal trend also tended to have stable temporal trends. SLAs with unstable temporal trends tended to be situated farther from the city and farther from mammography screening facilities.

Conclusions: This paper demonstrates the usefulness of the two models in identifying unusual and unstable temporal trends, and the synergy obtained when both models are applied to the same data set. Analysis of these models has provided interesting insights into the temporal trends of mammography screening counts, and has revealed several possible avenues for further research, such as extending the models to allow for multiple common temporal trends and accounting for additional spatio-temporal heterogeneity.

STRENGTHS AND LIMITATIONS OF THIS STUDY

- The models presented allow for the joint analysis of space and time, to provide useful insights on trends relating to a major health issue.
- The models fit the data well and provide good predictions.
- Some aspects of the model specification may be too restrictive.
- Additional data such as screening counts in rural areas were not available.

Keywords: mammography, public health, Bayesian, spatio-temporal, unusual trends.

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INTRODUCTION

Breast cancer is one of the most common types of cancer faced by women in Australia, and the second most common cause of cancer-related death, accounting for 2,914 deaths in 2011 alone.[1] Evidence suggests that regular mammography screening to be effective at detecting early breast cancer and increasing the chance of survival).[2-6] However, recent studies suggest that mammography screening services may be under-utilized due to geographical factors such as the travel distance to a mammography screening facility.[6-8]

Hyndman, Holman, and Dawes (2000)[6] conducted a study to investigate the effects of distance to a mammography screening facility and social disadvantage on service utilization. This study found that women with a lower socioeconomic status had more difficulty travelling to a mammography screening facility, and suggested that facilities should be located closer to disadvantaged communities to increase utilization. However, the influence of mammography screening facility location was less clear when socioeconomic factors play no significant role in service utilization. In other work, Legler et al. (2002)[9] modelled how state mammography screening rates depend on service user demographics, county-level socioeconomic factors, and previous mammography intervention research projects. The study found that states with one or more published intervention studies and states with higher levels of education tended to have higher rates of mammography screening service utilization. Zenk, Tarlov, and Sun (2006)[8] assessed the equitability of spatial accessibility to low or no-fee mammography screening services in Chicago by modeling distance and time to travel to a facility as a function of geographic and sociodemographic variables including race and poverty. The study concluded that travel time and distance was generally less for poorer neighborhoods, except for neighborhoods with a higher proportion of African-American residents. It is unclear from this study, however, if these barriers to access translated into lower screening utilization since mammography screening utilization rates were not considered.

The methodologies employed in the above studies vary greatly. Hyndman, Holman, and Dawes (2000)[6] used geographic information system (GIS) techniques and screening data from six mammography screening facilities in Perth, Australia. Legler et al. (2002)[9] opted for a hierarchical model to estimate the effects of education, occupation, and demographic group on mammography screening rates for each state. The model was fit to data collected at two time points, 1987 and 1993-1994, which mark a period during which numerous

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intervention studies were published. Although the model was applied to data at two time points, it only included spatial covariates, thus limiting inferences to differences between the two fitted models for each time point and spatial effects. Zenk, Tarlov, and Sun (2006)[8] used ordinary least squares regression models to estimate the effects of the covariates on the accessibility measures. However, the authors found that the residuals exhibited spatial autocorrelation, even when endogenous spatial lag regression was employed. Furthermore, the travel times and distances were estimated using GIS and other software rather than observed, and required numerous assumptions, adding to the uncertainty and potentially bias of the estimates.

The literature contains many other studies which analyze low- or no-fee mammography screening utilization rates and related data. The focus of these studies has typically been aimed at estimating the effect of one or more variables on screening rates. These variables are usually spatially dependent and include service user demographics, socioeconomic factors, accessibility factors, and variables relating to the spatial units of the study such as the degree of urbanization.[7, 10-13] Whilst estimation of covariate effects on mammography screening utilization is useful, little attention has been given to identifying trends in screening utilization rates, especially trends which vary over both space and time. Analysis of such trends permits a wider variety of statistical inferences, with implications for service management. Moreover, ignoring spatial and temporal correlation when present can lead to errors in prediction and inference.[14]

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This paper aims to build upon previous research in this field by presenting two spatiotemporal models, applied to no-fee mammography screening facility attendance data in Brisbane, Australia. In short, these models are designed to identify 'unusual' or 'unstable' temporal patterns. The use of the terms 'unusual' and 'unstable' are model specific, and their meanings are discussed in further detail in Methods.

By nature, spatio-temporal data can be clustered and/or autocorrelated, and are sometimes sparse whereby some regions exhibit relatively low numbers of observed and expected counts. Spatial models typically account for these issues by encoding neighborhood information as part of the wider model. This has the added advantage of reducing estimated risks with high uncertainty towards the mean risk. Bayesian methods naturally incorporate this information using prior distributions and hierarchical model structures, and allow for estimation of a full probability model for the unknown parameters.[8, 15-18]

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Both models considered in this paper have in common the specification of spatial and/or temporal random effects, albeit in different forms, and a model indicator as a means of differentiating SLAs that exhibit a common/stable temporal trend as opposed to an unusual/unstable temporal trend. Both models are estimated using Bayesian techniques and overcome the difficulties associated with autocorrelated data by explicitly including the spatial and temporal dependencies in the models.

METHODS

Data

The data employed in the current study consisted of the number of visits made to mammography screening facilities operated by BreastScreen Queensland in the Brisbane region per year, from 1997 to 2008 inclusive. For each year, the number of visits was recorded by statistical local area (SLA), with 158 SLAs included in the Brisbane region. The Moreton Island SLA was excluded from the analysis, however, since it has no neighboring SLAs. The eligible population was defined as women aged 40 years or over at time of screening, in line with the BreastScreen Australia Program eligibility criteria.[3]

The physical location and opening and closing dates of each mammography screening facility were also recorded. Some of the mammography screening facilities were mobile and therefore only available at a specific location for a shorter time period. Using these data, a covariate x_{ij} was created which represents the relative availability of services in a catchment area, defined as

$$x_{ij} = \frac{d_{ij}}{\max_{i} d_{ij}} \tag{2.1}$$

where d_{ij} is the cumulative number of days that each mammography screening facility was operating in SLA *i* or any SLA that shares a border with SLA *i*, {*i* = 1, ..., 157} during year *j*, {*j* = 1, ..., 12}. This catchment area service availability for odd years only is depicted graphically in Figure 1.

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Socioeconomic status was also considered as a covariate. However, a preliminary analysis of socioeconomic data did not indicate evidence of an effect. For this reason, it was excluded from the final models.

Model Formulation

Two models are proposed for spatio-temporal analysis of the data. Both models are examples of Bayesian spatial generalized linear mixed models (GLMMs) that fall within the wider class of linear models.[19-21]

Let y_{ij} denote the observed count of visits to mammography screening facilities in SLA *i* during year *j*. Given a population at risk R_{ij} , the corresponding expected number of visits is given by

 $E_{ij} = r_j R_{ij}$ $r_j = \frac{\sum_i y_{ij}}{\sum_i R_{ij}}$

where r_i is the reference screening rate in year *j*.[22]

The first model considered was the BaySTDetect model proposed by Li et al. (2012).[23] This model consists of two competing models: a common trend model where the temporal trend is the same for each SLA, and an area-specific model where the temporal trends are allowed to depart from the common trend. The two competing models are hierarchical in structure and are related to the likelihood via a model selection step, given by Equation (2.2). The BaySTDetect model assumes a Poisson random variable Y_{ij} to model the counts:

$$Y_{ij} \sim \text{Po}(E_{ij}\mu_{ij})$$

where

$$\log(\mu_{ij}) = \begin{cases} \alpha + \eta_i + \gamma_j + \beta_1 x_{ij} & \text{if } p_i = 1\\ u_i + \xi_{ii} + \beta_2 x_{ij} & \text{if } p_i = 0 \end{cases}$$
(2.2)

$$p_i \sim \text{Bern}(\delta_i).$$
 (2.3)

The components of Equation (2.2) are as follows: α is the common intercept, η_i and γ_j are random effects of space and time respectively, u_i is the area-specific intercept, ξ_{ij} is the area-

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specific random effect, and x_{ij} is the covariate defined by Equation (2.1). The parameters α , η_i , and u_i are assumed to be Normally distributed as follows,

$$\alpha \sim N(0, 1000)$$
$$\eta_i \sim N(0, \sigma_\eta^2)$$
$$u_i \sim N(0, 1000)$$

Regarding the prior for p_i , it is expected that temporal trends are fairly homogenous for most SLAs, and thus the hyper-parameter δ_i is chosen to be 0.95. To incorporate spatial and temporal smoothing, intrinsic conditional autoregressive (ICAR) priors[24] are assigned to the random effects γ_i and ξ_{ij}

$$\gamma_j | \boldsymbol{\gamma}_{ij} \sim N\left(\frac{1}{\sum_j w_{ij}} \sum_j w_{ij} \gamma_j, \frac{\sigma_{\gamma}^2}{\sum_j w_{ij}}\right)$$
(2.4)

$$\xi_{ij} | \xi_{i \setminus j} \sim N\left(\frac{1}{\sum_{j} w_{ij}} \sum_{j} w_{ij} \xi_{ij}, \frac{\sigma_{\xi,i}^2}{\sum_{j} w_{ij}}\right)$$
(2.5)

where j denotes all years excluding *j*, and w_{ij} is the *i*-*j*th element of a symmetric 'adjacency weight' matrix **W** with elements $w_{ij} = 1$ if the *i*th and *j*th years are neighbors, and zero otherwise. Note that the temporal adjacency information is the same for each of the *i* terms of ξ_{ij} .

The hyper-parameters σ_{η} and σ_{γ} were assigned weakly informative half-Normal priors to reflect a lack of prior knowledge about these parameters but restrict their values to be strictly positive and yet not too large.[25] The prior for the hyper-parameter $\sigma_{\xi,i}^2$ is log-Normal

$$\log(\sigma_{\xi,i}^2) \sim N(a, b^2)$$

where the variance is given an informative prior relative to the data,

$$b \sim N(0, 2.5^2)$$

to reflect prior expectations about the temporal variability.[23]

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The second model considered was based on the mixture model approach proposed by Abellan, Richardson, and Best(2008).[26] This model estimates the common spatial and temporal trends based on the data and identifies SLAs whose residual temporal patterns show volatility, that is, are unstable. In this hierarchical model, a Poisson random variable is also assumed to model the count data:

$$Y_{ij} \sim \text{Po}(\text{E}_{ij}\pi_{ij})$$
$$\log(\pi_{ij}) = \tau + \lambda_i + \psi_j + \nu_{ij} + \beta x_{ij}.$$
(2.6)

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Here the term τ is the common intercept, λ_i and ψ_j represent random effects for space and time respectively, and v_{ij} represents space-time interaction. Normal priors are defined for the first three terms,

$$\tau \sim N(0, 1000)$$
$$\lambda_i \sim N(0, \sigma_{\lambda}^2)$$
$$\psi_j \sim N(\omega_j, \sigma_{\psi}^2)$$

while the prior distribution for v_{ij} is described by a mixture of two Normal distributions with different variances, one representing stable patterns and the other unstable patterns:

$$\nu_{ij} \sim \mathrm{N}\left(0, \sigma_{\nu_{z_{ij}}}^{2}\right).$$
(2.7)

The variance is determined by a latent model indicator variable z_{ij} , specified in the model by the multinomial distribution consisting of a single draw,

$$z_{ij} \sim Mult(1, \boldsymbol{q})$$

where the prior for the mixture weights q is a Dirichlet distribution

$$(q_1, q_2) \sim \mathcal{D}(1, 1).$$

The latent indicators take the value 1 if v_{ij} is modelled by N(0, $\sigma_{v_1}^2$) or 2 if v_{ij} is modelled by N(0, $\sigma_{v_2}^2$), with $\sigma_{v_1}^2 < \sigma_{v_2}^2$. To avoid the issue of label-switching,[27, 28] and in line with the model specification, the priors for the two variances are specified as

$$\sigma_{\nu_1} \sim \mathrm{N}(0, 0.01) \mathbb{I}_{(0, +\infty)}$$

$$\sigma_{\nu_2} = \sigma_{\nu_1} + \kappa$$
$$\kappa \sim \mathrm{N}(0, 100) \mathbb{I}_{(0, +\infty)}$$

where \mathbb{I} denotes the indicator function $\mathbb{I}_{(a,b)} = 1$ if $a < \kappa < b$.

The second set of temporal random effects, $\omega_1, ..., \omega_{12}$, are modelled jointly using an ICAR prior

$$\omega_j | \boldsymbol{\omega}_{\backslash j} \sim N\left(\frac{1}{\sum_j w_{ij}} \sum_j w_{ij} \omega_j, \frac{\sigma_{\omega}^2}{\sum_j w_{ij}}\right)$$
(2.8)

where w_{ij} is as defined earlier (see Equations 2.4 and 2.5).

By analyzing the posterior frequencies of the latent indicator variables z_{ij} , this mixture model can be used to identify SLAs with unstable temporal trends. For example, let

$$P_{ij} = \Pr(z_{ij} = 2|y_{ij}).$$
(2.9)

represent the posterior probability that v_{ij} follows N(0, $\sigma_{v_2}^2$), that is, the posterior probability that the space-time interaction has a large variance. Thus the closer the P_{ij} values are to 2, and the more P_{ij} values that are close to 2 for j = 1, ..., 12, the more unstable the temporal patterns are for the ith SLA. Abellan, Richardson, and Best (2008)[26] propose two rules for classifying SLAs as unstable. The first rule considers the i^{th} SLA to be unstable if $P_{ij} > P_{cut}$ for at least one j, where P_{cut} is some specified threshold. The second rule classifies the i^{th} SLA as unstable if the average of the three largest P_{ij} values > P_{cut} . Rule 2 is slightly more conservative since it averages the P_{ij} values over three years. Both of these rules were employed.

Comparing the Two Models

While the distinction between unusual and unstable temporal trends may seem trivial, these two models aim to address two very different questions relating to spatio-temporal patterns, and hence each model may provide unique insights.

The BaySTDetect model assumes one common temporal trend, γ_j , across all areas and uses a model choice step to fit a competing model with independent random temporal effects for each area if there is considerable departure from the common trend. This allows identification of SLAs which have an unusual temporal trend. For example, assuming a constant mammography screening utilization rate on average (the common trend), then SLAs which exhibit a high screening rate one year followed by a low rate the next year would be considered to have an unusual temporal trend, and would therefore most likely be modelled by the area-specific model.

In contrast, the space-time mixture tries to estimate the overall spatio-temporal trend. If the annual screening counts for a given SLA are quite different from that which is predicted by the model, then this apparent departure from the overall spatio-temporal trend suggests that the screening rate for this SLA is unstable.

Implementation

Both models were estimated using Markov Chain Monte Carlo (MCMC) techniques, implemented in WinBUGS through R using the R2WinBUGS package.[29-31] The results are based on 25,000 iterations after discarding an initial 100,000 iterations as burn-in. Convergence was assessed informally via visual checks of trace and density plots, as well as formally using the Geweke convergence diagnostic.[32]

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Initially, both models were implemented with the hierarchical structure and priors as specified by the respective authors as described above. To resolve issues relating to computation, correlation between parameters, and insufficient adjacency information, it was necessary to modify these models to achieve convergence.

The main changes to the BaySTDetect model involved replacing the hyper-parameters for the means of η_i with zero due to a lack of spatial adjacency information provided by neighboring SLAs, and making the uninformative (hyper)priors for u_i and α less vague by decreasing the respective variances by a factor of 10. In the mixture model, the hyper-parameters for the means of λ_i were also replaced by zero for the same reason. Both models were also extended to include the covariate given by Equation (2.1).

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The WinBUGS code and schematic diagrams of the BaySTDetect and mixture models, after taking into account the changes outlined above, are provided as supplementary material available online. The posterior distributions of the key model parameters for each model are summarized in Figures 3 and 4.

Assessment of Model Fit and Predictive Performance

Posterior predictive checks (PPCs) were performed to assess the goodness-of-fit and predictive performance of the models given by Equations (2.2) and (2.6). In brief, PPCs aim to assess the consistency between predictions from the model and the observed data.[33-35] If y_{ij}^{pred} is a prediction of y_{ij} from the specified model, PPCs involve draws from the posterior predictive distribution

$$p(Y_{ij}^{\text{pred}}|y_{ij}) = \int p(Y_{ij}^{\text{pred}}|\boldsymbol{\theta}) p(\boldsymbol{\theta}|y_{ij}) d\boldsymbol{\theta}$$

where θ denotes all the parameters in the model.[34] These predictions were formed by sampling 200 times from the joint posterior distribution and using each posterior sample to generate $y_{ij}^{\text{pred}(k)}$, k = 1, ..., 200. The consistency between the predicted and observed counts for each SLA-year was evaluated using the L-criterion, [33] defined as the square root of the mean squared prediction error,

$$L_{ij} = \sqrt{E\left(\sum_{k=1}^{200} \left(Y_{ij}^{\text{pred}(k)} - y_{ij}\right)^2 \middle| y_{ij}\right)}.$$
 (2.10)

The estimate \hat{L}_{ij} of this quantity is easily computed from the MCMC estimate of the posterior predictive distribution. The results of this PPC are discussed below.

RESULTS

Predictive Performance of the Models

As a summary of the differences between the predicted and observed counts for each SLA, Figure 2 shows the L-criterion estimates \hat{L}_{ij} averaged over time for the BaySTDetect model.

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(The spatial composition of average \hat{L}_{ij} values for the mixture model is almost identical and thus omitted). The \hat{L}_{ij} values were about 19.28 and 18.73 counts on average for the BaySTDetect and mixture models respectively, suggesting acceptable and comparable predictive performance. While there were two regions of SLAs with predominantly larger \hat{L}_{ij} values (the north and south east), there did not appear to be any correlation between SLAs with larger \hat{L}_{ij} values and service availability (compare Figures 1 and 2).

BaySTDetect Model

The first three subplots in Figure 3 show the posterior means of the three spatially indexed parameters, p_i , η_i , and u_i , ordered by the average population at risk, $\sum_j R_{ij}/12$ from smallest to largest, left to right.

The first subplot in Figure 3 shows the posterior means for the model indicator parameter p_i which represent the posterior probabilities of selecting the common trend model for the i^{th} SLA (refer to Equation (2.3)). For those SLAs whose posterior mean of p_i was close to zero, the visits to mammography screening facilities Y_{ij} were better modelled by the area-specific model because these SLAs had temporal trends that differed considerably from the common trend γ_j . This was the case for most SLAs, with 93 (59%) SLAs having a posterior mean p_i less than or equal to 0.05, where 61 (39%) SLAs actually had a posterior mean p_i equal to zero. The p_i values for SLAs with a larger average population at risk tended to have a smaller posterior mean, indicating that the temporal trend for SLAs with a larger population at risk tended to be less similar to the common temporal trend. The spatial formation of these posterior means is provided in Figure 4.

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The next two subplots in the top row of Figure 3 show the posterior means of the parameters for the effects of space (on the logarithm scale) for the common-trend and area-specific models respectively. While the majority of the posterior means of η_i were close to zero, zero was only included in 32 of these 95% credible intervals (CIs), and a small quantity of these means were quite far from zero. In particular, eight SLAs had a posterior mean of less than -5 which correspond to SLAs with zero observed counts. The posterior distributions for u_i in the area-specific model were similar to those for η_i in that the majority of posterior means were close to zero, and it is the same eight SLAs which had a large negative posterior mean.

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Incidentally, these eight SLAs have the eight smallest aggregated L-criterion estimates, and can easily be identified in Figure 2 as the white regions.

The posterior densities of the parameters for the effects of the covariate x_{ij} in the two competing models are shown in the fourth and fifth subplots of Figure 3. The densities of these parameters indicate a positive marginal effect of the catchment covariate on service utilization, that is, a tendency for service utilization to be higher in SLAs that fall within the catchment area of a mammography screening facility, as would be expected.

The sixth subplot in Figure 3 shows the posterior means of the parameters for the effects of time for the common-trend model. This time effect exhibits an initial downward trend, then a consistent upward trend from 1999. Since the observed count of visits to mammography screening facilities also generally increase over time, this is not surprising. However, there are few SLAs for which the temporal trend agrees with this common trend, as indicated by the posterior means of p_i . This is partly explained by the variety of space-time trends in the area-specific model. The posterior means of these space-time trends ξ_{ij} for 6 selected SLAs are shown in the last two rows of subplots in Figure 3.

Space-Time Mixture Model

While the BaySTDetect model aims to determine SLAs with unusual temporal trends, the space-time mixture model is designed to identify SLAs whose residual temporal trend exhibits volatility. The first subplot in Figure 5 shows the posterior density of the parameter for the covariate effect, whose estimation and interpretation is comparable to that of β_2 in the BaySTDetect model. The second subplot shows the posterior means of λ_i , which are almost identical to those of the spatial effect term in the BaySTDetect model. (The eight smallest values of λ_i correspond to the SLAs with zero observed counts.) The temporal trend ψ_j shown in the third subplot is similar to the common temporal trend in the BaySTDetect model; its effect is stronger but estimated with less certainty, a result likely due to the dominance of the space-time interaction effects v_{ij} exhibit a similar variety of SLA specific temporal trends to ξ_{ij} in the BaySTDetect model, as shown in the second and third rows of subplots in Figure 5. The last two rows of subplots contain the posterior means of the latent indicator variables z_{ij} associated with the space-time interaction parameters (equal

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to P_{ij} + 1). By analyzing the posterior probabilities P_{ij} given by Equation (2.9), SLAs with unstable residual temporal trends can be identified. The two rules for classifying unstable SLAs proposed by Abellan, Richardson, and Best (2008)[26] were employed using a variety of different values for P_{cut} ; the results are summarized graphically in Figure 6.

DISCUSSION

This paper has presented two Bayesian hierarchical spatio-temporal models developed to model the utilization patterns of no-fee mammography screening services in Brisbane over twelve years. In contrast to previous studies, the models sought to identify SLAs with unusual or unstable temporal patterns as an initial step in improving management of these services. The results from both the BaySTDetect and space-time mixture models provide useful insight into the spatial and temporal patterns in mammography screening service utilization.

Firstly, the BaySTDetect model highlighted a large number of SLAs which had unusual temporal trends relative to the common trend. Although a covariate for the relative availability of services was included in the model to account for mobile facility relocations and facility operating times, service utilization rates for these SLAs changed from year to year in a way that differs to the common trend.

Secondly, although the BaySTD tect model estimates a common temporal trend γ_i , it is not common in the sense that very few SLAs exhibit this particular temporal trend. To understand why this is the case, consider the space-time trend ξ_{ii} in the area-specific model. The last two rows of subplots in Figure 3 show the area-specific temporal trend residuals for 6 selected SLAs. For SLA 9, ξ_{ij} exhibits a fairly stable upward trend, SLA 21 shows a downward trend, SLA 56 has a distinctive oscillating pattern, SLA 67 exhibits an oscillating pattern for the first 7 years followed by a relatively stable upward trend, SLA 91 shows no discernable pattern, and SLA 157 has a constant trend. This variety of trends suggests that there is not one but several common temporal trends, and explains why the area-specific model is favored by the Bayesian model choice.

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Thirdly, there is an apparent correlation between SLAs which follow the common temporal trends (lighter regions in Figure 4) and SLAs with stable temporal trends (white regions in Figure 6). This is most noticeable for smaller values of P_{cut} , especially when Rule 1 is used to classify unstable SLAs. This is unsurprising since the common temporal trend is itself fairly flat (only ranges between -0.01 and 0.01 approximately), i.e. stable. However, there also exist SLAs whose temporal trends are unusual but stable, and SLAs whose temporal Fourthly, the analysis of the space-time mixture model identified a number of SLAs as unstable, depending on the value of P_{cut} and the classification rule used. Figure 6 indicates that unstable SLAs tend to be situated on the outskirts of the Brisbane region, that is, unstable SLAs tend to be more rural than urban, particularly for larger values of P_{cut} . Comparing

trends are usual but unstable.

Figures 1 and 5, these unstable SLAs also tend to be in regions outside of catchment areas. This suggests that distance to a screening facility has an impact on the consistency of clients accessing mammography screening services. Lastly, the predicted values for eight SLAs with zero observed counts were the most consistent with the data in both the BaySTDetect and mixture models, as evidenced by the smallest L-criterion estimates. This implies that the existing model components and covariates are adequate in accounting for the lack of service utilization from these SLAs.

Figure 1 shows that these SLAs tend to fall outside the catchment areas, which reinforces the

notion that service utilization is influenced by the distance to the nearest screening facility.

The two models presented in this paper are not without limitations. Li et al. (2012)[23] raised a concern about the number of time periods over which the BaySTDetect model detects changes in the temporal trend. The authors advise that a single model indicator p_i for each SLA may be 'too restrictive' when the number of time periods is greater than ten because the current design assumes only one common temporal trend for the whole period, which is less likely to be the case for longitudinal data collected over many time points. This could be addressed by changing the model indicator to apply to SLAs and years, say p_{ij} . Another potential issue with the BaySTDetect model is the a priori specification of the prior for the model indicator, p_i . Li et al. (2012)[23] use 0.95 for the Bernoulli probability in Equation (2.3) to reflect their belief that only a small proportion of areas are actually unusual. This rather informative prior may have been adequate for the chronic disease mortality data

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analysis performed by Li et al. (2012)[23], but based on the results in Section 3.2 which indicate a large proportion of unusual SLAs, it may be more appropriate to specify a hyperprior for the Bernoulli probability, perhaps using additional spatial covariate information if available.

Similarly, the Dirichlet prior for q in the mixture model could be extended to include additional effects of space and/or time. The change in the dimensionality of q to q_i , q_j , or q_{ij} should be straightforward since the space-time effect v_{ij} is already indexed by space and time. However, this may increase the computational burden significantly.

In both models, there are a large number of parameters to be estimated, some of which have posterior means close to zero (in particular some of the space-time trends ξ_{ij} and v_{ij}). It may be beneficial to zero out such parameters using appropriate spike and slab priors.

A possible extension to the work of Abellan, Richardson, and Best (2008)[26] concerns the rules used to classify SLAs as unusual or not. The methodology proposed by Abellan, Richardson, and Best (2008)[26] allows SLAs with unstable temporal trends to be identified, but methods to identify the *degree* to which an SLA is unstable would undoubtedly be more informative and comparable to the results from the BaySTDetect model. Another avenue for future research is the inclusion of additional covariates that vary in space and/or time, such as accessibility to public transport, which may improve inferences relating to the spatial and/or temporal trends.

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The L-criterion values also provide insight into the observed trends. It is speculated that the larger \hat{L}_{ij} values may be attributed to some unknown factor such as the influence of services offered by private mammography screening facilities. For both models, the predictive performance tended to decrease with time, with the annual average \hat{L}_{ij} values increasing by about 4 between 1997 and 2008. Inclusion of other temporal factors may improve predictive performance in later years.

Overall, this paper has shown that the BaySTDetect and space-time mixture models are useful in analyzing mammography screening service utilization data. In particular, the BaySTDetect model was able to identify SLAs which had temporal trends that differed from the overall temporal trend, and the space-time mixture model identified SLAs with unstable

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temporal trends. Analysis of these models has shown insight into patterns of the observed trends, and revealed potentially important factors not yet considered.

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Contributors

EWD developed the models and analyzed the data, prepared an initial draft of the manuscript, and collaborated in the process of revising the manuscript. NMW and KM made substantial contributions to the methodological design and development of the models, provided critical comments and suggestions on all drafts of the manuscript, and assisted in the analysis and interpretation of the models and data. All authors approved the final draft of the manuscript for publication.

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Competing Interests

We have read and understood BMJ policy on declaration of interests and declare that we have no competing interests.

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Figure 1: Map of the SLAs in the Brisbane region (excluding Moreton Island) depicting the relative availability of mammography screening services, based on the operating duration and location of mammography screening facilities in each SLA and neighboring SLAs over time (odd years shown only).

375x434mm (300 x 300 DPI)

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Figure 2: Map of SLAs in the Brisbane region depicting the closeness between y_{ij} and $E(Y_{ij}^{pred} | y_{ij})$ for each SLA for the final (modified) BaySTDetect model, as specified by the L-criterion defined in Equation (2.10). Lighter regions represent SLAs with a smaller aggregated L-criterion estimates.

956x722mm (96 x 96 DPI)



Figure 3: Posterior densities and means of model parameters for 1 chain of the final (modified) BaySTDetect model. One-dimensional parameters are summarized by their posterior density; multi-dimensional parameters are summarized by the posterior mean and 95% credible interval for each SLA or year. 375x347mm (300 x 300 DPI)

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Figure 4: Map of SLAs in the Brisbane region representing the degree to which SLAs follow the common temporal trend (lighter regions) or exhibit unusual temporal trends (darker regions). 956x722mm (96 x 96 DPI)





375x434mm (300 x 300 DPI)





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SUPPLEMENTARY MATERIAL

Supplementary code 1: WinBUGS code for the BaySTDetect model

```
model {
  for (i in 1:N) {
    for (j in 1:T) {
      y[i,j] ~ dpois(mu[i,j])
      log(mu[i,j]) <- log(E[i,j]) + mix[i,j]</pre>
      mix[i,j] <- p[i] * crw.mix[i,j] + (1-p[i]) * rw.mix[i,j]</pre>
    }
    p[i] ~ dbern(0.95)
  }
# Common trend model
  for (i in 1:N) {
    for (j in 1:T) {
      y1[i,j] ~ dpois(mu1[i,j])
      \log(mu1[i,j]) < -\log(E[i,j]) + temp[i,j]
      temp[i,j] <- eta[i] + gamma[j] + beta1 * x[i,j]
      crw.mix[i,j] <- cut(temp[i,j])</pre>
    }
    eta[i] ~ dnorm(alpha,tau.eta)
  }
  alpha ~ dnorm(0,0.001)
  gamma[1:T] ~ car.normal(adj.t[],weights.t[],num.t[],tau.gamma)
  betal ~ dnorm(0, 0.001)
  tau.gamma <- pow(sigma.gamma,-2)</pre>
  sigma.gamma ~ dnorm(0,1)I(0,)
  tau.eta <- pow(sigma.eta,-2)</pre>
  sigma.eta ~ dnorm(0,1)I(0,)
# Area-specific Model
  for (i in 1:N) {
    for (j in 1:T) {
      y2[i,j] \sim dpois(mu2[i,j])
      log(mu2[i,j]) <- log(E[i,j]) + temp1[i,j]</pre>
      temp1[i,j] <- u[i] + xi[i,j] + beta2 * x[i,j]</pre>
      rw.mix[i,j] <- cut(temp1[i,j])</pre>
    }
    u[i] \sim dnorm(0, 0.001)
    xi[i,1:T] ~ car.normal(adj.t[],weights.t[],num.t[],tau.xi[i])
    tau.xi[i] <- pow(var.xi[i],-1)</pre>
    var.xi[i] <- exp(log.var.xi[i])</pre>
    log.var.xi[i] ~ dnorm(a,c)
  }
  a \sim dnorm(0, 0.001)
  c < -pow(b, -2)
  b \sim dnorm(0,d)I(0,)
  d <- pow(2.5, -2)
  beta2 ~ dnorm(0,0.001)
}
```

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Supplementary code 2: WinBUGS code for the space-time mixture model

```
model {
  for (i in 1:N) {
    for (j in 1:T) {
      y[i,j] ~ dpois(pi[i,j])
      log(pi[i,j]) <- log(E[i,j]) + temp[i,j] + beta * x[i,j]</pre>
      temp[i,j] <- lambda[i] + psi[j] + nu[i,j]</pre>
    }
  }
  # Priors
  tau ~ dnorm(0, 0.001)
  beta ~ dnorm(0, 0.001)
  for (i in 1:N){
    lambda[i] ~ dnorm(tau, tau.lambda)
  }
  for(j in 1:T){
    psi[j] ~ dnorm(omega[j], tau.psi)
  }
  omega[1:T] ~ car.normal(adj.t[],weights.t[],num.t[],tau.omega)
  for(i in 1:N){
    for(j in 1:T){
      nu[i,j] ~ dnorm(0, tau.nu[z[i,j]])
    }
  }
  # Hyperpriors
  tau.lambda ~ dgamma(0.5, 0.0005)
  tau.psi ~ dgamma(0.5, 0.0005)
  tau.omega ~ dgamma(0.5, 0.0005)
  sigma.lambda <- 1/sqrt(tau.lambda)</pre>
  sigma.psi <- 1/sqrt(tau.psi)</pre>
  sigma.omega <- 1/sqrt(tau.omega)</pre>
  for(i in 1:N){
    for(j in 1:T){
      z[i,j] \sim dcat(q[])
    }
  }
  sigma.nu[1] ~ dnorm(0, 100)I(0.0,)
  kappa ~ dnorm(0, 0.01)I(0.0,)
  sigma.nu[2] <- sigma.nu[1] + kappa</pre>
  q[1:2] ~ ddirch(alpha.q[])
  for(i in 1:2){
    tau.nu[i] <- pow(sigma.nu[i], -2)</pre>
    alpha.q[i] <- 1
  }
}
```

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Supplementary figure 1: Schematic diagram of the BaySTDetect model after modifications to the original model. W denotes the temporal adjacency matrix, $\gamma_j \sim \text{CAR}(\mathbf{Q}, \sigma_{\gamma}^2)$ and $\xi_{ij} \sim \text{CAR}(\mathbf{Q}, \sigma_{\xi,i}^2)$ are shorthand notation for Equations (2.4) and (2.5) respectively, and E_{ij} is the expected number of visits in the *i*th SLA and *j*th year.

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Supplementary figure 2: Schematic diagram of the space-time mixture model after modifications to the original model. W denotes the temporal adjacency matrix, $\omega_j \sim \text{CAR}(\mathbf{Q}, \sigma_{\omega}^2)$ is shorthand notation for Equation (2.8), and E_{ij} is the expected number of visits in the *i*th SLA and *j*th year.

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Bayesian spatio-temporal modeling for identifying unusual and unstable trends in mammography utilization

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ABSTRACT

Objectives: To compare two Bayesian models capable of identifying unusual and unstable temporal patterns in spatio-temporal data.

Setting: Annual counts of mammography screening users from each statistical local area (SLA) in Brisbane, Australia, recorded between 1997 and 2008 inclusive.

Primary outcome measures: Mammography screening counts

Results: The temporal trends of 91 SLAs (58%) were dissimilar to the overall common temporal trend. SLAs which did follow the common temporal trend also tended to have stable temporal trends. SLAs with unstable temporal trends tended to be situated farther from the city and farther from mammography screening facilities.

Conclusions: This paper demonstrates the usefulness of the two models in identifying unusual and unstable temporal trends, and the synergy obtained when both models are applied to the same data set. Analysis of these models has provided interesting insights into the temporal trends of mammography screening counts, and has revealed several possible avenues for further research, such as extending the models to allow for multiple common temporal trends and accounting for additional spatio-temporal heterogeneity.

STRENGTHS AND LIMITATIONS OF THIS STUDY

- The models presented allow for the joint analysis of space and time, to provide useful insights on trends relating to a major health issue.
- The models fit the data well and provide good predictions.
- Some aspects of the model specification may be too restrictive.
- Additional data such as screening counts in rural areas were not available.

Keywords: mammography, public health, Bayesian, spatio-temporal, unusual trends.

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INTRODUCTION

Breast cancer is one of the most common types of cancer faced by women in Australia, and the second most common cause of cancer-related death, accounting for 2,914 deaths in 2011 alone.[1] Evidence suggests that regular mammography screening to be effective at detecting early breast cancer and increasing the chance of survival).[2-6] However, recent studies suggest that mammography screening services may be under-utilized due to geographical factors such as the travel distance to a mammography screening facility.[6-8]

Hyndman, Holman, and Dawes (2000)[6] conducted a study to investigate the effects of distance to a mammography screening facility and social disadvantage on service utilization. This study found that women with a lower socioeconomic status had more difficulty travelling to a mammography screening facility, and suggested that facilities should be located closer to disadvantaged communities to increase utilization. However, the influence of mammography screening facility location was less clear when socioeconomic factors play no significant role in service utilization. In other work, Legler et al. (2002)[9] modelled how state mammography screening rates depend on service user demographics, county-level socioeconomic factors, and previous mammography intervention research projects. The study found that states with one or more published intervention studies and states with higher levels of education tended to have higher rates of mammography screening service utilization. Zenk, Tarlov, and Sun (2006)[8] assessed the equitability of spatial accessibility to low or no-fee mammography screening services in Chicago by modeling distance and time to travel to a facility as a function of geographic and sociodemographic variables including race and poverty. The study concluded that travel time and distance was generally less for poorer neighborhoods, except for neighborhoods with a higher proportion of African-American residents. It is unclear from this study, however, if these barriers to access translated into lower screening utilization since mammography screening utilization rates were not considered.

The methodologies employed in the above studies vary greatly. Hyndman, Holman, and Dawes (2000)[6] used geographic information system (GIS) techniques and screening data from six mammography screening facilities in Perth, Australia. Legler et al. (2002)[9] opted for a hierarchical model to estimate the effects of education, occupation, and demographic group on mammography screening rates for each state. The model was fit to data collected at two time points, 1987 and 1993-1994, which mark a period during which numerous

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intervention studies were published. Although the model was applied to data at two time points, it only included spatial covariates, thus limiting inferences to differences between the two fitted models for each time point and spatial effects. Zenk, Tarlov, and Sun (2006)[8] used ordinary least squares regression models to estimate the effects of the covariates on the accessibility measures. However, the authors found that the residuals exhibited spatial autocorrelation, even when endogenous spatial lag regression was employed. Furthermore, the travel times and distances were estimated using GIS and other software rather than observed, and required numerous assumptions, adding to the uncertainty and potentially bias of the estimates.

The literature contains many other studies which analyze low- or no-fee mammography screening utilization rates and related data. The focus of these studies has typically been aimed at estimating the effect of one or more variables on screening rates. These variables are usually spatially dependent and include service user demographics, socioeconomic factors, accessibility factors, and variables relating to the spatial units of the study such as the degree of urbanization.[7, 10-13] Whilst estimation of covariate effects on mammography screening utilization is useful, little attention has been given to identifying trends in screening utilization rates, especially trends which vary over both space and time. Analysis of such trends permits a wider variety of statistical inferences, with implications for service management. Moreover, ignoring spatial and temporal correlation when present can lead to errors in prediction and inference.[14]

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This paper aims to build upon previous research in this field by presenting two spatiotemporal models, applied to no-fee mammography screening facility attendance data in Brisbane, Australia. In short, these models are designed to identify 'unusual' or 'unstable' temporal patterns. The use of the terms 'unusual' and 'unstable' are model specific, and their meanings are discussed in further detail in Methods.

By nature, spatio-temporal data can be clustered and/or autocorrelated, and are sometimes sparse whereby some regions exhibit relatively low numbers of observed and expected counts. Spatial models typically account for these issues by encoding neighborhood information as part of the wider model. This has the added advantage of reducing estimated risks with high uncertainty towards the mean risk. Bayesian methods naturally incorporate this information using prior distributions and hierarchical model structures, and allow for estimation of a full probability model for the unknown parameters.[8, 15-18]

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(1)

Both models considered in this paper have in common the specification of spatial and/or temporal random effects, albeit in different forms, and a model indicator as a means of differentiating SLAs that exhibit a common/stable temporal trend as opposed to an unusual/unstable temporal trend. Both models are estimated using Bayesian techniques and overcome the difficulties associated with autocorrelated data by explicitly including the spatial and temporal dependencies in the models.

METHODS

Data

The data employed in the current study consisted of the number of visits made to mammography screening facilities operated by BreastScreen Queensland in the Brisbane region per year, from 1997 to 2008 inclusive. For each year, the number of visits was recorded by statistical local area (SLA), with 158 SLAs included in the Brisbane region. The eligible population was defined as women aged 40 years or over at time of screening, in line with the BreastScreen Australia Program eligibility criteria.[3]

The physical location and opening and closing dates of each mammography screening facility were also recorded. Some of the mammography screening facilities were mobile and therefore only available at a specific location for a shorter time period. Using these data, a covariate x_{it} was created which represents the relative availability of services in a catchment area, defined as

$$x_{it} = \frac{d_{it}}{\max_{i} d_{it}}$$

where d_{it} is the cumulative number of days that each mammography screening facility was operating in SLA *i* or any SLA that shares a border with SLA *i*, {*i* = 1, ..., 158} during year *t*, {*t* = 1, ..., 12}. This catchment area service availability for odd years only is depicted graphically in Figure 1.

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Socioeconomic status was also considered as a covariate. However, a preliminary analysis of socioeconomic data did not indicate evidence of an effect. For this reason, it was excluded from the final models.

Model Formulation

Two models are proposed for spatio-temporal analysis of the data. Both models are examples of Bayesian spatial generalized linear mixed models (GLMMs) that fall within the wider class of linear models.[19-21]

Let y_{it} denote the observed count of visits to mammography screening facilities in SLA *i* during year *t*. Given a population at risk R_{it} , the corresponding expected number of visits is given by

 $E_{it} = r_t R_{it}$ $r_t = \frac{\sum_i y_{it}}{\sum_i R_{it}}$

where r_t is the reference screening rate in year t.[22]

The first model considered was the BaySTDetect model proposed by Li et al. (2012).[23] This model consists of two competing models: a common trend model where the temporal trend is the same for each SLA, and an area-specific model where the temporal trends are allowed to depart from the common trend. The two competing models are hierarchical in structure and are related to the likelihood via a model selection step, given by Equation (2). The BaySTDetect model assumes that the Y_{it} counts are a Poisson random variable, e.g.

$$Y_{it} \sim \text{Po}(E_{it}\mu_{it})$$

where

$$\log(\mu_{it}) = \begin{cases} \alpha + \eta_i + \gamma_t + \beta x_{it} & \text{if } p_i = 1\\ u_i + \xi_{it} + \beta' x_{it} & \text{if } p_i = 0 \end{cases}$$
(2)

$$p_i \sim \text{Bern}(\delta_i).$$
 (3)

The components of Equation (2) are as follows: α is the common intercept, η_i and γ_t are random effects of space and time respectively, u_i is the area-specific intercept, ξ_{it} is the area-

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specific random effect, and x_{it} is the covariate defined by Equation (1). Regarding the prior for p_i , it is expected that temporal trends are fairly homogenous for most SLAs, and thus the hyper-parameter δ_i is chosen to be 0.95. To incorporate spatial and temporal smoothing, intrinsic conditional autoregressive (ICAR) priors[24] are assigned to the random effects γ_t and ξ_{it}

$$\eta_i | \boldsymbol{\eta}_{\backslash i} \sim N\left(\frac{1}{\sum_j w_{ij}} \sum_{j=1}^N w_{ij} \eta_j, \frac{\sigma_{\eta}^2}{\sum_j w_{ij}}\right), \tag{4}$$

$$\gamma_t | \boldsymbol{\gamma}_{\setminus t} \sim N\left(\frac{1}{\sum_s w_{st}} \sum_{s=1}^T w_{st}' \gamma_s, \frac{\sigma_{\gamma}^2}{\sum_s w_{st}}\right), \tag{5}$$

$$\xi_{it} | \xi_{i \setminus t} \sim N\left(\frac{1}{\sum_{s} w_{st}} \sum_{s=1}^{T} w_{st}' \xi_{is}, \frac{\sigma_{\xi,i}^2}{\sum_{s} w_{st}}\right)$$
(6)

where i denotes all areas excluding *i*, and w_{ij} is the *i*-*j*th element of a symmetric spatial adjacency weight matrix **W** with elements $w_{ij} = 1$ if the *i*th and *j*th areas are neighbors, and zero otherwise. Similarly, t denotes all years excluding *t*, and w'_{st} is the *s*-*t*th element of a symmetric temporal adjacency weight matrix **W**' with elements $w'_{st} = 1$ if the *s*th and *t*th years are neighbors, and zero otherwise. Note that the temporal adjacency information is the same for each of the *i* terms of ξ_{it} .

The parameters α , β , u_i , and β' are assumed to be Normally distributed with mean 0 and variance 1000. The hyper-parameters σ_{η} and σ_{γ} were assigned weakly informative half-Normal priors to reflect a lack of prior knowledge about these parameters but restrict their values to be strictly positive and yet not too large.[25] The prior for the hyper-parameter $\sigma_{\xi,i}^2$ is log-Normal

$$\log(\sigma_{\xi,i}^2) \sim N(a, c^2)$$

where the variance is given an informative prior relative to the data,

$$c \sim \mathrm{N}(0, 2.5^2) \mathbb{I}_{(0, +\infty)}$$

to reflect prior expectations about the temporal variability.[23]

The second model considered was based on the mixture model approach proposed by Abellan, Richardson, and Best(2008).[26] This model estimates the common spatial and temporal trends based on the data and identifies SLAs whose residual temporal patterns show volatility, that is, are unstable. In this hierarchical model, the counts Y_{it} are modelled as Poisson random variables with mean $E_{it}\pi_{it}$, e.g.

$$Y_{it} \sim \text{Po}(\text{E}_{it}\pi_{it})$$
$$\log(\pi_{it}) = \tau + \lambda_i + \psi_t + \nu_{it} + bx_{it}.$$
(7)

Here the term τ is the common intercept, λ_i and ψ_t represent random effects for space and time respectively, and v_{it} represents space-time interaction. Like the BaySTDetect model, the spatial and temporal random effects are modelled jointly using ICAR priors,

$$\lambda_i | \boldsymbol{\lambda}_{\backslash i} \sim N\left(\frac{1}{\sum_j w_{ij}} \sum_{j=1}^N w_{ij} \lambda_j, \frac{\sigma_{\lambda}^2}{\sum_j w_{ij}}\right), \tag{8}$$

$$\psi_t | \boldsymbol{\psi}_{\backslash t} \sim N\left(\frac{1}{\sum_s w_{st}} \sum_{s=1}^T w_{st}' \psi_s, \frac{\sigma_{\psi}^2}{\sum_s w_{st}}\right)$$
(9)

where w_{ij} and w'_{st} are as defined earlier (see Equations 4 through 6). Normal priors are defined for the intercept and covariate effect terms,

$$\tau \sim N(0, 1000),$$

 $b \sim N(0, 1000),$

while the prior distribution for v_{it} is described by a mixture of two Normal distributions with different variances, one representing stable patterns and the other unstable patterns:

$$v_{it} \sim q N(0, \sigma_{v_1}^2) + (1 - q) N(0, \sigma_{v_2}^2).$$

The variance is determined by a latent model indicator variable z_{it} , specified in the model by the multinomial distribution consisting of a single draw,

$$z_{it} \sim \text{Mult}(1, \boldsymbol{q})$$

where the prior for the mixture weights q = (q, 1 - q) is a Dirichlet distribution

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$$\boldsymbol{q} \sim \mathcal{D}(1,1)$$

The latent indicators take the value 1 if v_{it} is modelled by N(0, $\sigma_{v_1}^2$) or 2 if v_{it} is modelled by N(0, $\sigma_{v_2}^2$), with $\sigma_{v_1}^2 < \sigma_{v_2}^2$. To avoid the issue of label-switching,[27, 28] and in line with the model specification, the priors for the two variances are specified as

 $\sigma_{\nu_1} \sim N(0, 0.01) \mathbb{I}_{(0, +\infty)}$ $\sigma_{\nu_2} = \sigma_{\nu_1} + \kappa$ $\kappa \sim N(0, 100) \mathbb{I}_{(0, +\infty)}$

where \mathbb{I} denotes the indicator function $\mathbb{I}_{(a,b)} = 1$ if $a < \kappa < b$.

By analyzing the posterior frequencies of the latent indicator variables z_{it} , this mixture model can be used to identify SLAs with unstable temporal trends. For example, let

$$P_{it} = \Pr(z_{it} = 2|y_{it}).$$
 (10)

represent the posterior probability that v_{it} follows N(0, $\sigma_{v_2}^2$), that is, the posterior probability that the space-time interaction has a large variance. Thus the closer the P_{it} values are to 1, and the more P_{it} values that are close to 1 for t = 1, ..., 12, the more unstable the temporal patterns are for the i^{th} SLA. Abellan, Richardson, and Best (2008)[26] propose two rules for classifying SLAs as unstable. The first rule considers the i^{th} SLA to be unstable if $P_{it} > P_{cut}$ for at least one t, where P_{cut} is some specified threshold. The second rule classifies the i^{th} SLA as unstable if the average of the three largest P_{it} values $> P_{cut}$. Rule 2 is slightly more conservative since it averages the P_{it} values over three years. Both of these rules were employed.

Comparing the Two Models

While the distinction between unusual and unstable temporal trends may seem trivial, these two models aim to address two very different questions relating to spatio-temporal patterns, and hence each model may provide unique insights.

The BaySTDetect model assumes one common temporal trend, γ_t , across all areas and uses a model choice step to fit a competing model with independent random temporal effects for

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each area if there is considerable departure from the common trend. This allows identification of SLAs which have an unusual temporal trend. For example, assuming a constant mammography screening utilization rate on average (the common trend), then SLAs which exhibit a high screening rate one year followed by a low rate the next year would be considered to have an unusual temporal trend, and would therefore most likely be modelled by the area-specific model.

In contrast, the space-time mixture tries to estimate the overall spatio-temporal trend. If the annual screening counts for a given SLA are quite different from that which is predicted by the model, then this apparent departure from the overall spatio-temporal trend suggests that the screening rate for this SLA is unstable.

Implementation

Both models were estimated using Markov Chain Monte Carlo (MCMC) techniques, implemented in WinBUGS through R using the R2WinBUGS package.[29-31] The results are based on 25,000 iterations after discarding an initial 100,000 iterations as burn-in. Convergence was assessed informally via visual checks of trace and density plots, as well as formally using the Geweke convergence diagnostic.[32]

Initially, both models were implemented with the hierarchical structure and priors as specified by the respective authors as described above. These models were then adapted to our scientific problem of interest through a number of modifications. The main changes to the models involved modelling the spatial and temporal random effects η_i , λ_i , and ψ_t using ICAR priors directly, rather than modelling their respective means, due to a lack of strong autocorrelation between parameters and issues with identifiability of parameters (results not shown). Both models were also extended to include the covariate given by Equation (.1).

Schematic diagrams of the BaySTDetect and mixture models, after taking into account the changes outlined above, are provided in Supplementary Figure 1 and Supplementary Figure 2 respectively, available online. The WinBUGS code is also provided, in Supplementary Code 1 and Supplementary Code 2. The posterior distributions of the key model parameters for each model are summarized in Figure 2a-l and Figure 3a-o.

Assessment of Model Fit and Predictive Performance

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Posterior predictive checks (PPCs) were performed to assess the goodness-of-fit and predictive performance of the models given by Equations (2) and (7). In brief, PPCs aim to assess the consistency between predictions from the model and the observed data.[33-35] If y_{it}^{pred} is a prediction of y_{it} from the specified model, PPCs involve draws from the posterior predictive distribution

$$p(Y_{it}^{\text{pred}} | y_{it}) = \int p\left(Y_{it}^{\text{pred}} | \boldsymbol{\theta}\right) p(\boldsymbol{\theta} | y_{it}) d\boldsymbol{\theta}$$

where θ denotes all the parameters in the model.[34] These predictions were formed by sampling 200 times from the joint posterior distribution and using each posterior sample to generate $y_{it}^{\text{pred}(k)}$, k = 1, ..., 200. The consistency between the predicted and observed counts for each SLA-year was evaluated using the L-criterion,[33] defined as the square root of the mean squared prediction error,

$$L_{it} = \sqrt{E\left(\sum_{k=1}^{200} \left(Y_{it}^{\text{pred}(k)} - y_{it}\right)^2 \,\middle|\, y_{it}\right)}.$$
 (11)

The estimate \hat{L}_{it} of this quantity is easily computed from the MCMC estimate of the posterior predictive distribution. The results of this PPC are discussed below.

RESULTS

Predictive Performance of the Models

As a summary of the differences between the predicted and observed counts for each SLA, Figure 4 shows the L-criterion estimates \hat{L}_{it} averaged over time for the BaySTDetect model. (The spatial composition of average \hat{L}_{it} values for the mixture model is almost identical and thus omitted). The \hat{L}_{it} values were about 19.21 and 18.70 counts on average for the BaySTDetect and mixture models respectively, suggesting acceptable and comparable predictive performance. While there were two regions of SLAs with predominantly larger \hat{L}_{it} values (the north and south east), there did not appear to be any correlation between SLAs with larger \hat{L}_{it} values and service availability (compare Figure 1 and Figure 4).

BaySTDetect Model

Figure 2a-c show the posterior means of the three spatially indexed parameters, p_i , η_i , and u_i respectively, ordered by the average population at risk, $\sum_t R_{it}/12$ from smallest to largest, left to right.

Figure 2a shows the posterior means for the model indicator parameter p_i which represent the posterior probabilities of selecting the common trend model for the *i*th SLA (refer to Equation (3)). For those SLAs whose posterior mean of p_i was close to zero, the visits to mammography screening facilities Y_{it} were better modelled by the area-specific model because these SLAs had temporal trends that differed considerably from the common trend γ_t . This was the case for most SLAs, with 91 (58%) SLAs having a posterior mean p_i less than or equal to 0.05, where 61 (39%) SLAs actually had a posterior mean p_i equal to zero. The p_i values for SLAs with a larger average population at risk tended to have a smaller posterior mean, indicating that the temporal trend for SLAs with a larger population at risk tended to be less similar to the common temporal trend. The spatial formation of these posterior means is provided in Figure 5.

Figure 2b and Figure 2c show the posterior means of the parameters for the effects of space (on the logarithm scale) for the common-trend and area-specific models respectively. While the majority of the posterior means of η_i were close to zero, zero was included in only 5 of these 95% credible intervals (CIs), and a small quantity of these means were quite far from zero. In particular, eight SLAs had a posterior mean of less than -5 which correspond to SLAs with zero observed counts. The posterior distributions for u_i in the area-specific model were similar to those for η_i in that the majority of posterior means were close to zero, and it is the same eight SLAs which had a large negative posterior mean. Incidentally, these eight SLAs have the eight smallest aggregated L-criterion estimates, and can easily be identified in Figure 4 as the white regions.

The posterior densities of the parameters for the effects of the covariate x_{it} in the two competing models are shown in Figure 2d and Figure 2e. The densities of these parameters indicate a positive marginal effect of the catchment covariate on service utilization, that is, a tendency for service utilization to be higher in SLAs that fall within the catchment area of a mammography screening facility, as would be expected.

Figure 2f shows the posterior means of the parameters for the effects of time for the commontrend model. The posterior means generally decrease with time, indicating a fairly consistent downward trend. The observed counts of visits to mammography screening facilities, however, generally increase over time. While this result is surprising, the temporal effect is very small. More interestingly, there are few SLAs for which their respective temporal trends agree with this common trend, as indicated by the posterior means of p_i . This is partly explained by the variety of space-time trends in the area-specific model. The posterior means of these space-time trends ξ_{it} for six selected SLAs are shown in Figure 2g-l.

Space-Time Mixture Model

While the BaySTDetect model aims to determine SLAs with unusual temporal trends, the space-time mixture model is designed to identify SLAs whose residual temporal trend exhibits volatility. Figure 3a shows the posterior density of the parameter for the covariate effect, whose estimation and interpretation is comparable to that of β' in the BaySTDetect model. Figure 3b shows the posterior means of λ_i , which are almost identical to those of the spatial effect term in the BaySTDetect model. (The eight smallest values of λ_i correspond to the SLAs with zero observed counts.)

The temporal effect ψ_t shown in Figure 3c indicates a slight, decreasing trend overall. While this differs to the common temporal trend in the BaySTDetect model, the effect size in both cases is small.

Figure 3d-i show the posterior means and 95% CIs for the space-time interaction effects v_{it} for the same six selected SLAs in Figure 2g-l respectively. They exhibit a similar variety of SLA specific temporal trends to ξ_{it} in the BaySTDetect model. Figure 3j-o show the posterior means of the latent indicator variables z_{it} associated with the space-time interaction parameters (equal to $P_{it} + 1$). By analyzing the posterior probabilities P_{it} given by Equation (10), SLAs with unstable residual temporal trends can be identified. The two rules for classifying unstable SLAs proposed by Abellan, Richardson, and Best (2008)[26] were employed using a variety of different values for P_{cut} ; the results are summarized graphically in Figure 6.

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This paper has presented two Bayesian hierarchical spatio-temporal models used to analyze the utilization patterns of no-fee mammography screening services in Brisbane over twelve years. In contrast to previous studies, the models sought to identify SLAs with unusual or unstable temporal patterns as an initial step in improving management of these services. The results from both the BaySTDetect and space-time mixture models provide useful insight into the spatial and temporal patterns in mammography screening service utilization.

Firstly, the BaySTDetect model highlighted a large number of SLAs which had unusual temporal trends relative to the common trend. Although a covariate for the relative availability of services was included in the model to account for mobile facility relocations and facility operating times, service utilization rates for these SLAs changed from year to year in a way that differs to the common trend.

Secondly, although the BaySTDetect model estimates a common temporal trend γ_t , it is not common in the sense that very few SLAs exhibit this particular temporal trend. To understand why this is the case, consider the space-time trend ξ_{it} in the area-specific model. Figure 2g-l show the area-specific temporal trend residuals for 6 selected SLAs. For SLA 10, ξ_{it} exhibits a fairly stable upward trend, SLA 22 shows a downward trend, SLA 57 has a distinctive oscillating pattern, SLA 68 exhibits an oscillating pattern for the first 7 years followed by a relatively stable upward trend, SLA 92 shows no discernable pattern, and SLA 158 has a constant trend. This variety of trends suggests that there is not one but several common temporal trends, and explains why the area-specific model is favored by the Bayesian model choice. Given the large proportion of SLAs which have a temporal trend which departs from the common trend, departures from this trend should be interpreted with care.

Thirdly, there is an apparent correlation between SLAs which follow the common temporal trends (lighter regions in Figure 5) and SLAs with stable temporal trends (white regions in Figure 6). This is most noticeable for smaller values of P_{cut} , especially when Rule 1 is used to classify unstable SLAs. This is unsurprising since the common temporal trend is itself fairly flat (only ranges between -0.02 and 0.02 approximately), i.e. stable. However, there also exist SLAs whose temporal trends are unusual but stable, and SLAs whose temporal trends are usual but unstable.

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Fourthly, the analysis of the space-time mixture model identified a number of SLAs as unstable, depending on the value of P_{cut} and the classification rule used. Figure 6 indicates that unstable SLAs tend to be situated on the outskirts of the Brisbane region, that is, unstable SLAs tend to be more rural than urban, particularly for larger values of P_{cut} . Comparing Figure 1 and Figure 6, these unstable SLAs also tend to be in regions outside of catchment areas. This suggests that distance to a screening facility has an impact on the consistency of clients accessing mammography screening services.

Lastly, the predicted values for eight SLAs with zero observed counts were the most consistent with the data in both the BaySTDetect and mixture models, as evidenced by the smallest L-criterion estimates. This implies that the existing model components and covariates are adequate in accounting for the lack of service utilization from these SLAs. Figure 1 shows that these SLAs tend to fall outside the catchment areas, which reinforces the notion that service utilization is influenced by the distance to the nearest screening facility.

The two models presented in this paper are not without limitations. Li et al. (2012)[23] raised a concern about the number of time periods over which the BaySTDetect model detects changes in the temporal trend. The authors advise that a single model indicator p_i for each SLA may be 'too restrictive' when the number of time periods is greater than ten because the current design assumes only one common temporal trend for the whole period, which is less likely to be the case for longitudinal data collected over many time points. This could be addressed by changing the model indicator to apply to SLAs and years, say p_{it} . Another potential issue with the BaySTDetect model is the a priori specification of the prior for the model indicator, p_i . Li et al. (2012)[23] use 0.95 for the Bernoulli probability in Equation (3) to reflect their belief that only a small proportion of areas are actually unusual. This rather informative prior may have been adequate for the chronic disease mortality data analysis performed by Li et al. (2012)[23], but based on the results which indicate a large proportion of unusual SLAs, it may be more appropriate to specify a hyperprior for the Bernoulli probability, perhaps using additional spatial covariate information if available.

Similarly, the Dirichlet prior for q in the mixture model could be extended to include additional effects of space and/or time. The change in the dimensionality of q to q_i , q_t , or q_{it} should be straightforward since the space-time effect v_{it} is already indexed by space and time. However, this may increase the computational burden significantly.

In both models, there are a large number of parameters to be estimated, some of which have posterior means close to zero (in particular some of the space-time trends ξ_{it} and v_{it}). It may be beneficial to zero out such parameters using appropriate spike and slab priors.

In this study, the spatial autocorrelation between the observed data for any given year appears to be weak, as indicated by the posterior means of η_i and λ_i , shown in Figure 2b and Figure 3b. However, measures of spatial autocorrelation such as Moran's I and Geary's C indicate the contrary (results not shown). Although Geary's C is more sensitive to local spatial autocorrelation, such statistics imply that spatial autocorrelation in this dataset is global rather than local, and thus perhaps not easily captured through ICAR priors. Results may be improved by changing the adjacency weight elements w_{ij} to be non-zero for second- and third-order neighbors, for example.

A possible extension to the work of Abellan, Richardson, and Best (2008)[26] concerns the rules used to classify SLAs as unusual or not. The methodology proposed by Abellan, Richardson, and Best (2008)[26] allows SLAs with unstable temporal trends to be identified, but methods to identify the *degree* to which an SLA is unstable would undoubtedly be more informative and comparable to the results from the BaySTDetect model. Another avenue for future research is the inclusion of additional covariates that vary in space and/or time, such as accessibility to public transport, which may improve inferences relating to the spatial and/or temporal trends.

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The L-criterion values also provide insight into the observed trends. It is speculated that the larger \hat{L}_{it} values may be attributed to some unknown factor such as the influence of services offered by private mammography screening facilities. For both models, the predictive performance tended to decrease with time, with the annual average \hat{L}_{it} values increasing by about 4 between 1997 and 2008. Inclusion of other temporal factors may improve predictive performance in later years.

Overall, this paper has shown that the BaySTDetect and space-time mixture models are useful in analyzing mammography screening service utilization data. In particular, the BaySTDetect model was able to identify SLAs which had temporal trends that differed from the overall temporal trend, and the space-time mixture model identified SLAs with unstable temporal trends. Analysis of these models has shown insight into patterns of the observed trends, and revealed potentially important factors not yet considered.

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Contributors

EWD applied the models and analyzed the data, prepared an initial draft of the manuscript, and collaborated in the process of revising the manuscript. NMW and KM made substantial contributions to the methodological design and development of the models, provided critical comments and suggestions on all drafts of the manuscript, and assisted in the analysis and interpretation of the models and data. All authors approved the final draft of the manuscript for publication.

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Competing Interests

We have read and understood BMJ policy on declaration of interests and declare that we have no competing interests.

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Figure 1: Map of the SLAs in the Brisbane region (Moreton Island not shown) depicting the relative availability of mammography screening services based on the operating duration and location of mammography screening facilities in each SLA and neighboring SLAs over time (odd years shown only), as defined by Equation (1). 1103x1332mm (96 x 96 DPI)



Figure 2: Posterior densities and means of model parameters for 1 chain of the final (modified) BaySTDetect model: (a) posterior mean and 95% CI for p_i , (b) posterior mean and 95% CI for η_i , (c) posterior mean and 95% CI for μ_i , (d) posterior density for β , (e) posterior density for β \', (f) posterior mean and 95% CI for γ_t , (g) posterior mean and 95% CI for ξ_{10t} , (h) posterior mean and 95% CI for ξ_{22t} , (i) posterior mean and 95% CI for ξ_{57t} , (j) posterior mean and 95% CI for ξ_{68t} , (k) posterior mean and 95% CI for ξ_{92t} , and (l) posterior mean and 95% CI for ξ_{158t} .

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Figure 3: Posterior summary of the main model parameters for 1 chain of the final (modified) space-time mixture model: (a) posterior density for *b*, (b) posterior mean and 95% CI for λ_i , (c) posterior mean and 95% CI for ψ_{t_i} (d) posterior mean and 95% CI for v_{10t_i} (e) posterior mean and 95% CI for v_{22t_i} (f) posterior mean and 95% CI for v_{57t_i} (g) posterior mean and 95% CI for v_{68t_i} (h) posterior mean and 95% CI for v_{92t_i} (i) posterior mean and 95% CI for v_{158t_i} (j) posterior mean and 95% CI for z_{10t_i} (k) posterior mean and 95% CI for z_{22t_i} (l) posterior mean and 95% CI for z_{22t_i} (n) posterior mean and 95% CI for z_{22t_i} (n) posterior mean and 95% CI for z_{22t_i} (n) posterior mean and 95% CI for z_{158t_i} (n) posterior mean and posterior

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Figure 4: Map of SLAs in the Brisbane region (Moreton Island not shown) depicting the closeness between y_{ij} and $E(Y_{ij}^{pred} | y_{ij})$ for each SLA for the final (modified) BaySTDetect model, as specified by the L-criterion defined in Equation (11). Lighter regions represent SLAs with a smaller aggregated L-criterion estimates.

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Figure 5: Map of SLAs in the Brisbane region (Moreton Island not shown) representing the degree to which SLAs follow the common temporal trend (lighter regions) or exhibit unusual temporal trends (darker regions).

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Figure 6: Map of SLAs in the Brisbane region (Moreton Island not shown) with unstable trends (shaded areas) determined by Rule 1 and Rule 2 using different values for the threshold, P_{cut} . 822x946mm (96 x 96 DPI)



Supplementary Figure 1: Schematic diagram of the BaySTDetect model after modifications to the original model. **W** and **W'** denote the spatial and temporal adjacency matrices respectively, and $\eta_i \sim \text{ICAR}(\mathbf{W}, \sigma_{\eta}^2)$, $\gamma_t \sim \text{ICAR}(\mathbf{W}', \sigma_{\gamma}^2)$, and $\xi_{it} \sim \text{ICAR}(\mathbf{W}', \sigma_{\xi,i}^2)$ are shorthand notation for Equations (4), (5) and (6) respectively.

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Supplementary Code 1: WinBUGS code for the BaySTDetect model

```
model {
  for (i in 1:N) {
    for (t in 1:T) {
      y[i,t] ~ dpois(E.mu[i,t])
      E.mu[i,t] <- E[i,t] * mu[i,t]
      log(mu[i,t]) <- p[i] * crw.mix[i,t] + (1-p[i]) * rw.mix[i,t]</pre>
    }
    p[i] ~ dbern(0.95)
  }
# Common trend model
  for (i in 1:N) {
    for (t in 1:T) {
      y1[i,t] ~ dpois(mu1[i,t])
      log(mu1[i,t]) <- log(E[i,t]) + temp1[i,t]</pre>
      temp1[i,t] <- alpha + eta[i] + gamma[t] + beta * x[i,t]</pre>
      crw.mix[i,t] <- cut(temp1[i,t])</pre>
    }
  }
  alpha ~ dnorm(0, 0.001)
  gamma[1:T] ~ car.normal(adj.t[],weights.t[],num.t[],tau.gamma)
  beta ~ dnorm(0, 0.001)
  eta[1:(N-1)] ~ car.normal(adj[],weights[],num[],tau.eta)
  eta[N] <- 0
                   # SLA with zero neighbours
  tau.gamma <- pow(sigma.gamma,-2)</pre>
  sigma.gamma ~ dnorm(0,1)I(0,)
  tau.eta <- pow(sigma.eta,-2)</pre>
  sigma.eta ~ dnorm(0,1)I(0,)
# Area-specific Model
  for (i in 1:N) {
    for (t in 1:T) {
      y2[i,t] ~ dpois(mu2[i,t])
      log(mu2[i,t]) <- log(E[i,t]) + temp2[i,t]</pre>
      temp2[i,t] <- u[i] + xi[i,t] + beta.dash * x[i,t]</pre>
      rw.mix[i,t] <- cut(temp2[i,t])</pre>
    }
    u[i] \sim dnorm(0, 0.001)
    xi[i,1:T] ~ car.normal(adj.t[],weights.t[],num.t[],tau.xi[i])
    tau.xi[i] <- pow(var.xi[i],-1)</pre>
    var.xi[i] <- exp(log.var.xi[i])</pre>
    log.var.xi[i] ~ dnorm(a,cc)
  }
  beta.dash ~ dnorm(0, 0.001)
  a \sim dnorm(0, 0.001)
  cc <- pow(c, -2)
  c \sim dnorm(0,d)I(0,)
  d <- pow(2.5,-2)
}
```

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```
Supplementary Code 2: WinBUGS code for the space-time mixture model
model {
  for (i in 1:N) {
    for (t in 1:T) {
      y[i,t] ~ dpois(E.pi[i,t])
      E.pi[i,t] <- E[i,t] * pi[i,t]
      log(pi[i,t]) <- tau + lambda[i] + psi[t] + nu[i,t] + b * x[i,t]</pre>
    }
  }
  # Priors
  tau ~ dnorm(0, 0.001)
  lambda[1:(N-1)] ~ car.normal(adj[],weights[],num[],tau.lambda)
  lambda[N] <- 0 # SLA with zero neighbours</pre>
  psi[1:T] ~ car.normal(adj.t[],weights.t[],num.t[],tau.psi)
  for(i in 1:N) {
    for(t in 1:T) {
      nu[i,t] ~ dnorm(0, tau.nu[z[i,t]])
    }
  }
  b \sim dnorm(0, 0.001)
  # Hyperpriors
  for(i in 1:N) {
    for(t in 1:T) {
      z[i,t] \sim dcat(q[])
    }
  }
  sigma.lambda.sq <- pow(tau.lambda,</pre>
                                       -1)
  sigma.psi.sq <- pow(tau.psi, -1)</pre>
  tau.lambda ~ dgamma(0.5, 0.0005)
  tau.psi ~ dgamma(0.5, 0.0005)
  sigma.nu[1] ~ dnorm(0, 100)I(0.0,)
  kappa ~ dnorm(0, 0.01)I(0.0,)
  sigma.nu[2] <- sigma.nu[1] + kappa</pre>
  q[1:2] ~ ddirch(alpha.q[])
  for(i in 1:2){
    tau.nu[i] <- pow(sigma.nu[i], -2)</pre>
    alpha.q[i] <- 1
  }
}
```