

# A comparison of least squares regression and geographically weighted regression modeling of West Nile virus risk based on environmental parameters

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**Background.** The primary aim of the study reported here was to determine the effectiveness of utilizing local spatial variations in environmental data to uncover the statistical relationships between West Nile Virus (WNV) risk and environmental factors. Because least squares regression methods do not account for spatial autocorrelation and non-stationarity of the type of spatial data analyzed for studies that explore the relationship between WNV and environmental determinants, we hypothesized that a geographically weighted regression model would help us better understand how environmental factors are related to WNV risk patterns without the confounding effects of spatial non-stationarity.

**Methods.** We examined commonly mapped environmental factors using both ordinary least squares regression (LSR) and geographically weighted regression (GWR). Both types of models were applied to examine the relationship between WNV-infected dead bird counts and various environmental factors for those locations. The goal was to determine which approach yielded a better predictive model.

**Results.** LSR efforts lead to identifying three environmental variables that were statistically significantly related to WNV infected dead birds (adjusted  $R^2=0.61$ ): stream density, road density, and land surface temperature. GWR efforts increased the explanatory value of these three environmental variables with better spatial precision (adjusted  $R^2 = 0.71$ ).

**Conclusions.** The spatial granularity resulting from the geographically weighted approach provides a better understanding of how environmental spatial heterogeneity is related to WNV risk as implied by WNV infected dead birds, which should allow improved planning of public health management strategies.

1 **A comparison of least squares regression and geographically weighted**  
2 **regression modeling of West Nile virus risk based on environmental**  
3 **parameters**

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17 **Abstract**

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37 risk as implied by WNV infected dead birds, which should allow improved planning of public  
38 health management strategies.

39

## 40 **Introduction**

41 West Nile Virus (WNV) is a vector-borne disease that was first detected in the United  
42 States in 1999 (Nash et al. 2001). Within a few years the virus had spread across the North  
43 American continent (Hayes et al. 2005). WNV has had important environmental and human  
44 impacts, including a decline in numerous bird species (CDC) and increased morbidity and  
45 mortality among humans. This has also resulted in increased economic burdens due to initial  
46 acute health care needs of infected individuals and subsequent long-term costs associates with  
47 infection, estimated at approximately \$56 million per year between 1999 and 2012 (Barrett  
48 2014). Because that study indicated how difficult predicting and planning for WNV outbreaks  
49 was, we became interested in developing a spatially explicit model using environmental factors  
50 in an attempt to improve WNV risk predictions.

51 There are two important considerations that should typically be examined when  
52 developing spatially explicit environmental disease risk models (Miller 2012). The first should  
53 be an examination of potential spatial autocorrelation (the degree to which a set of spatial  
54 features and their associated data values tend to be clustered together in space). This involves  
55 accounting for whether environmental factors and the corresponding disease rates in  
56 geographically proximate areas are more or less clustered together than they are in  
57 geographically distant areas. Second, data non-stationarity (changing means, variances and  
58 covariances in data across space) should be investigated and controlled when necessary  
59 (Fotheringham 2009a; Miller 2012). Geographically weighted regression (GWR) can be used for  
60 these two considerations and can often produce improved models that enable better spatial  
61 inference and prediction. Recent studies have applied GWR modeling to drug-resistant

62 tuberculosis versus risk factors (Liu et al. 2011); environmental factors versus typhoid fever  
63 (Dewan et al. 2013); local climate and population distribution versus hand, foot, and mouth  
64 disease (Hu et al. 2012); and environmental factors and tick-borne disease (Atkinson et al. 2012;  
65 Atkinson et al. 2014; Wimberly et al. 2008a; Wimberly et al. 2008b), all showing that predictor  
66 variables varied spatially across large geographic regions, implying that the results for such  
67 studies may be improved using GWR.

68         The spatially explicit model that is discussed in this paper uses GWR to account for  
69 spatial heterogeneity for two reasons: (a) WNV disease risk observed across space may be  
70 related to similar environmental variables that increase vector habitat suitability and (b)  
71 environmental variables that influence WNV risk are not typically uniformly distributed across  
72 geographic space. Although many epidemiological models of WNV risk have been developed, it  
73 appears that there has been little research to explicitly examine techniques that account for  
74 spatial heterogeneity. Most models assume that the impact of various environmental factors are  
75 constant across the study region, which is unrealistic as larger areas display substantial variations  
76 in distribution of environmental, socio-economic, and demographic conditions (Goovaerts 2008).

77         Due to the unavailability of reliable and complete data, developing models of WNV risk  
78 pose additional challenges. Human case data is lacking due to issues of under-reporting and  
79 limited surveillance. Our alternative strategy was to assess WNV infected dead bird counts as a  
80 surrogate measure of human risk because “infection rates” in dead birds can be more precise  
81 because of the genetic markers tested in dead birds may be more reliable than case data and/or  
82 surveillance data. Additionally, others have also used mosquito habitat suitability as a surrogate  
83 for estimating WNV risk for human infection (Cooke et al. 2006). For our study, we followed a  
84 similar approach and used a model of mosquito habitat suitability condition as a predictor of the

85 spatial distributions of infected birds, which in turn can be used to estimate WNV disease risk  
86 among human populations. Further, because the environmental variables considered in this study  
87 are known to vary across space, we account for spatial autocorrelation and non-stationarity using  
88 GWR following the approach of (DeGroot et al. 2008) in order to improve the predictability of  
89 a model.

90

91 WNV transmission and risk factors

92         The WNV transmission cycle was an important component of our modeling efforts. The  
93 first step in the WNV transmission cycle primarily occurs when a competent female mosquito  
94 vector bites an infected bird reservoir host, which in turn results in the virus being transmitted to  
95 the mosquito (Blair 2009). This occurs when the female mosquito is seeking a blood meal to  
96 obtain nutrients necessary for egg development. After taking an infectious blood meal, a  
97 mosquito may pick up a permanent infection. The infected mosquito now has the potential to  
98 transmit the virus to another bird or animal when it feeds again. Once infected, birds may fly to  
99 other locations where the virus can be transmitted to susceptible mosquitoes. Subsequently, the  
100 disease may be transmitted by infected mosquitoes to humans or other mammals that act as  
101 incidental hosts. Dead birds found to be infected with WNV are often the primary indicators for  
102 presence of the disease in a geographic region and have proven to be useful for disease  
103 prediction modeling and identifying areas for human infection risk (Cooke et al. 2006; Ruiz et al.  
104 2004; Valiakos et al. 2014). This relationship allows an assumption of a positive correlation  
105 between infected dead birds and WNV risk. Since the New York outbreak in 1999, WNV has  
106 been recovered from 26 mosquito species in North America, including *Culex. pipiens*, *Culex.*  
107 *salinarius*, *Culex. restuans*, *Ochlerotatus canadensis*, *Oc. japonicus*, *Aedes vexans*, and *Culiseta*

108 *melanura* (CDC 2000; Control & Prevention 2001). Results from a study (Goddard et al. 2002)  
109 assesses the vector competence of California mosquitoes. The results indicate that mosquitoes in  
110 the genus *Culex* (*Cx.*) are the principal hosts of WNV in California. The study also analyzed that  
111 on the basis of vector competence and host-feeding patterns, *Cx. tarsalis* may be the principal  
112 vector in rural agricultural ecosystems; and *Cx. pipiens* complex and *Cx. stigmatosoma* as  
113 important vectors in urban settings.

114         Vector and pathogen reservoirs overlap when certain environmental conditions are  
115 present (Rochlin et al. 2011). Table 1 provides an overview of the environmental conditions that  
116 are associated with WNV transmission, which were utilized for our research. These include  
117 characteristics of a place such as the mosquito species habitat: climatic conditions, topography  
118 and land use/land cover classes such as vegetation, water, and urbanized areas. Spectral indices  
119 acquired from satellite imagery provide information about environmental characteristics like  
120 temperature, vegetation cover, and moisture (Rodgers & Mather 2006). Liu and Weng (Liu &  
121 Weng 2012) in a study on WNV risk in southern California found that one of the main factors  
122 contributing to the WNV propagation included land surface temperature. They related higher  
123 temperature to viral replication in mosquitoes for WNV to be disseminated throughout the year.  
124 The results also show that areas with lower elevations tended to be more susceptible to WNV  
125 invasion as mosquito population propagates in the plain habitats with warmer temperatures  
126 compared to areas with higher elevation that have lower temperatures.

127         **Table 1 approximately here**

128         Statistical Considerations

129         Miller (2012) suggests that a ‘global’ model is the one that assumes that the parameters  
130 (commonly mean and variance) of some process are constant across geographic space

131 (commonly mean and variance), typically referred to as the spatial stationarity of a process.  
132 Miller suggests that in the case these parameters vary across geographic space (spatial  
133 heterogeneity), then such models may lead to inaccurate predictions and subsequent problems for  
134 decision-making. In an ecological context, spatial heterogeneity usually results from the  
135 interaction of various environmental processes that operate at different scales (Legendre 1993).  
136 Fotheringham (Fotheringham 2009b) used local statistics for linking the concepts of spatial  
137 autocorrelation and heterogeneity that are deemed important when developing spatial models.  
138 Local statistics disaggregate a global mean value into locally computed values for each spatial  
139 unit. It is based on a conceptualization of Tobler's first law in Geography (Tobler 1970) that  
140 specifies that "everything is related to everything else, but near things are more related than  
141 distant things." Spatial autocorrelation is a commonly used measure of the degree of spatial  
142 heterogeneity.

143 GWR is a local regression method that can be used for diagnosing spatial heterogeneity  
144 between dependent and explanatory variables over space (Fotheringham et al. 2003). It is  
145 performed within local windows centered on the nodes of a regular grid. Each observation within  
146 the local window is weighted based on its proximity to the center of that window. This approach  
147 has several advantages: it avoids abrupt changes in the local statistics computed for adjacent  
148 windows, helps visualize spatial variability within the geographic entity, and allows analysis of  
149 regionally aggregated data (Goovaerts 2008). A model's predictive ability, particularly in  
150 ecological modeling, is influenced not only by the strength of relationships between the species  
151 and its environment, but whether the model recognizes if the relationships are operating at  
152 multiple spatial scales. GWR provides a framework for exploring scale-dependent effects. It tests

153 the effect on a model's predictive ability by systematically increasing the size of the local  
154 window (Miller 2012).

155 GWR can be used for mapping the spatial distribution of a model's coefficient values in  
156 order to identify potential missing variables or to suggest other underlying factors associated  
157 with the observed non-stationarity (Miller 2012). GWR is also useful for exploratory data  
158 analysis and visualization; for example Kupfer and Farris (Kupfer & Farris 2007) used a 'leave-  
159 one-out' (jackknifing) methodology to compare residuals from GWR and ordinary least squares  
160 regression. They found that GWR often had more accurate predictions for sites that were  
161 difficult to predict (where both models had overall higher residuals), which is why we used a  
162 GWR framework for explicitly modeling the spatial relationships between WNV and its  
163 environmental risk factors.

164

165

166

## 167 **Materials & Methods**

### 168 Study area

169 The model was built for the state of California, which was the national epicenter of WNV  
170 activity in 2004 and 2005 (Jean et al. 2007). WNV was first detected there in July 2003 (Reisen  
171 et al. 2004). It is the third largest state by area in the United States and is made up of 58 counties.  
172 California has the largest population in the U.S., but it is unevenly distributed across the state.  
173 The state also has a variable landscape with a large valley in the middle, bounded by mountain  
174 ranges.

175

176 Environmental factors and data sources

177           Our model utilized various environmental factors (Table 1) that have been suggested as  
178 descriptive in local WNV risk distribution: surface slope, density of roads, density of streams,  
179 monthly mean temperature, monthly mean evapotranspiration, and land cover classes like  
180 vegetation, developed land, cultivated land, and open surface water. All environmental  
181 parameters except roads and streams (Table 2) were acquired in grid format and resampled to  
182 120 meter resolution as suggested by Cooke et al. (2006). Data resampling was done using the  
183 resampling tool available in ArcGIS software. The modeling method utilized in this study was  
184 based on analyzing data in raster format, and therefore road and stream vector data were  
185 converted to raster format using the ‘Kernel Density Estimation’ tool in ArcGIS to create road  
186 density and stream density grid files. The tool assumes a Gaussian distribution and thus assigns  
187 more importance towards the center of kernel in comparison to the features that are further apart.

188           Various dynamic environmental data including Normalized Difference Vegetation Index  
189 (NDVI), Land Surface Temperature (LST), and Evapotranspiration (ET) were downloaded from  
190 the Moderate Resolution Imaging Spectroradiometer (MODIS) toolbox incorporated in  
191 ArcGIS®. The Land Surface Temperature tool accesses MOD11-A1, the daily averaged LST  
192 product. The MOD11 product uses the algorithm which is optimally used to separate ranges of  
193 atmospheric column water vapor and lower boundary air surface temperatures into tractable sub-  
194 ranges. The NDVI is calculated according to the formula  $NDVI = (NIR - VIS) / (NIR + VIS)$   
195 where NIR is the near-infrared radiance and VIS is observed radiation in the visible spectrum.  
196 NDVI data is available from either satellite with MODIS (Aqua or Terra) as a monthly average.  
197 The time lag between the hatching of a mosquito egg to an adult mosquito taking blood meals  
198 and becoming infected with WNV to the subsequent infection of a human and the appearance of

199 WNV disease symptoms was taken into account and therefore environmental data used for this  
200 study was taken for the month of July, the month prior to peak WNV human incidence cases  
201 (Campbell et al. 2002).

202 **Table 2 approximately here**

203

204 Least Squares Regression (LSR) modeling

205 WNV disease annual incidence rate (cases per 100,000 populations) was used as the  
206 measure of disease severity in this study. Annual WNV-infected dead birds sentinel data,  
207 averaged for 2004-2010, was used as a surrogate of WNV risk and was the dependent variable  
208 for modeling purposes in this study because several other studies (Chaintoutis et al. 2014; Eidson  
209 et al. 2001a; Eidson et al. 2001b; Eidson et al. 2001c; Guptill et al. 2003; Johnson et al. 2006;  
210 Mostashari et al. 2003; Nielsen & Reisen 2007; Patnaik et al. 2007; Ruiz et al. 2004) have  
211 suggested links between infected dead birds and WNV human infection rates. Since wild birds  
212 are the primary reservoir hosts for WNV and indicator of human infection risk, we utilized this  
213 association to develop the disease prediction model. We determined the utility of this  
214 relationship by correlating the dead birds data with the human incidence rate ( $r^2 = 0.409$ ). While  
215 infected dead bird counts only explain about 40% of reported human cases in California, it is a  
216 highly significant predictor ( $p=0.01$ ). Hence we used a dead bird model, with infected dead birds  
217 as a dependent variable, to assess WNV risk among human population.

218 Interpretations of ordinary Least Squares Regression (LSR) model performance were  
219 based on assessing multi-collinearity, robust probability, adjusted  $R^2$ , and Akaike's information  
220 criterion (AIC) (Akaike 1974). Multi-collinearity was assessed through the variance inflation  
221 factor (VIF) statistic, which measures redundancy among explanatory variables. Explanatory  
222 variables associated with VIF values larger than about 7.5 indicate that these variables are

223 providing similar information, and they were removed one at a time from the model based on  
224 VIF value until the model became unbiased. Robust probability indicates the statistically  
225 significant variables that are important to the regression model. Examining VIF values and  
226 robust probability, we ran and re-ran LSR models until narrowing down to non-redundant and  
227 significant variables: land surface temperature; stream density, and; road density. Akaike's  
228 information criterion (AIC) was then used to determine the best LSR model.

229         The next step was to explore GWR models that might better explain the variation in  
230 infected dead bird counts based upon environmental data. Spatial autocorrelation (Global  
231 Moran's I) was utilized to assess whether the environmental factors exhibited a random spatial  
232 pattern (Goodchild 1986), and where adequate models have a random distribution of the  
233 residuals (Mitchell 2005).

234

235 Geographically Weighted Regression (GWR) modeling

236         Under conditions of non-stationarity in LSR modeling, geographically weighted  
237 regression (GWR) was explored to potentially improve modeling results. The same explanatory  
238 variables that were used in LSR modeling were used to run GWR rather than starting with the  
239 full global set of parameters so as to avoid introducing "improvement" that could not be  
240 attributed solely to which modeling approach was applied. In other words, if GWR modeling was  
241 not applied to the same variables as LSR modeling, but yielded better results, we would not  
242 know if the improvement was due to the modeling approach or the environmental data that was  
243 used to build each model.

244         Once key environmental factors were identified during LSR modeling, we proceeded to  
245 explore the spatial variability of local regression coefficients to determine whether the

246 underlying process exhibited spatial heterogeneity (Fotheringham et al. 2003). A GWR local  
247 model was applied to analyze how the relationship between infected dead bird counts and  
248 environmental factors changed from one county to another. Unlike conventional LSR regression  
249 modeling, which produces a single regression equation to summarize global relationships among  
250 the independent and dependent variables, GWR detects spatial variation within relationships in a  
251 model and produces information useful for exploring and interpreting spatial non-stationarity  
252 (Fotheringham et al. 2003).

253         A spatial kernel was used to provide geographic weighting for the local window centered  
254 on the grid nodes used in our model. There are two possible categories of spatial kernels:  
255 fixed/adaptive and bandwidth, which is a key coefficient that controls the size of the kernel  
256 (Akaike 1974). These kernels tend to be Gaussian or Gaussian-like which implies that distant  
257 samples are weighed lesser than the proximal ones. There are three potential bandwidth  
258 approaches: Akaike information criterion (AIC), cross validation (CV), and bandwidth  
259 parameter. For our GWR model, the AIC approach was chosen because the distribution of  
260 infected dead birds was not consistent in the study area. The following settings were used in  
261 ArcGIS GWR: Bandwidth method = AIC and Kernel type = Adaptive.

262         Finally, we examined independency and normality of residuals, to evaluate the fit of the  
263 model. Local collinearity, the square root of the largest eigenvalue divided by the smallest  
264 eigenvalue, of our GWR model was also assessed but no data points were removed as they  
265 compromised model diagnostics. The adjusted coefficient of determination (Adjusted  $R^2$ ) was  
266 used for comparing LSR and GWR models to determine which approach would provide a better  
267 understanding of the relationship between environmental conditions and West Nile Virus risk  
268 (Fotheringham et al. 2003).

269

270 **Results**

271 LSR modeling identified land surface temperature (VIF = 1.046), stream density (VIF =  
272 1.177), and road density (VIF = 1.143) as statistically significant ( $p < 0.05$ ) variables related to  
273 WNV risk:

274

$$275 \text{ WNV risk} = -75.87 + 595.60 (RD) + 1.89 (LST) - 146.89 (SD) \quad (1)$$

276

277 Where:

278 WNV risk = average infected dead bird count

279  $RD$  = road density280  $LST$  = land surface temperature281  $SD$  = stream density

282 The histogram of the LSR model's residuals approximates that of a normal curve, with a  
283 non-significant (0.134,  $p < 0.05$ ) Jarque-Bera statistic (Jarque & Bera 1980), and the Moran's I  
284 Index Z-score (1.23) all imply that the model is unbiased and significantly different than random.

285 However, the Koenker statistic (0.000007\*,  $p < 0.05$ ) confirmed non-stationarity in  
286 the LSR model indicating that there is not a consistent relationship between the explanatory  
287 variables and WNV risk across the study area. Further, the presence of mild heteroskedasticity  
288 was noted in the LSR model. We conclude that the LSR model is stable but non-stationary,  
289 suggesting that proceeding with GWR model was warranted.

290 The GWR model in this study was implemented using the following algorithm:

291

292 
$$\text{WNV risk}_{(i)} = \beta_{i0} + \beta_{(i1)} RD_{(i)} + \beta_{(i2)} LST_{(i)} - \beta_{(i3)} SD_{(i)} + \varepsilon_{(i)} \quad (2)$$

293 Where  $\beta$  coefficients are county (i) specific, and RD is road density, LST is land surface  
294 temperature, and SD is stream density.

295

296 Comparing the fit of the global LSR model (assumes homogeneity of variables across  
297 space) and local GWR model (makes no assumption of homogeneity), we found that the global  
298 LSR adjusted  $R^2$  was 0.61 ( $R^2$  was 0.66,  $P < 0.05$ , Fig. 1) with analysis run on all 58 counties. The  
299 local GWR adjusted  $R^2$  was 0.71 ( $R^2$  is 0.75,  $p < 0.05$ , Fig. 2) with a bandwidth of 54, which  
300 suggests that there has been some improvement by using a local modeling approach. Our  
301 preferred measure of model fit, AIC, gave a value of 567.7 for the global model and 551.4 for the  
302 local model. The difference of 16.3 is relatively strong evidence of an improvement in the model  
303 fit to the data. Further, the problem of heteroskedasticity that was noted in the OLS model was not  
304 observed in the GWR model.

305 We also tested the results using different bandwidth parameter. Several iterations were  
306 run but it was observed that although a smaller band-width criterion gave an improved  
307 combination of AICc and adjusted  $R^2$  values, it also compromised the model diagnostics by  
308 introducing local collinearity and thus instability in the model. Addressing local collinearity by  
309 removing the Counties having condition number greater than 30 affected the model's overall  
310 results. Thus, it is better to have a larger band-width rather than violating model assumptions and  
311 to avoid the unstable prediction (Charlton & Fotheringham ; Nakaya 2014).

312

313 **Figure 1 approximately here**

314 **Figure 2 approximately here**

315

316 Mapping the values of the standardized residual across California (Fig. 3a) provides a  
317 representation of: (a) areas with unusually high or low residuals and (b) whether the residuals  
318 were spatially autocorrelated. Counties with excessively large positive residuals would under-  
319 predict WNV risk, and counties with excessively large negative residuals would over-predict  
320 WNV risk. The spatial autocorrelation of GWR residuals for our model resulted in a Moran's I  
321 value of -0.11 ( $p=0.18$ ), implying little evidence of any autocorrelation in them.

322 **Figure 3 approximately here**

323 Local coefficient estimates for significant factors were mapped using quantile  
324 classification method. Figure 3b shows the variation in the model's coefficient estimates for the  
325 land surface temperature (LST) variable. The map for the local coefficients reveals that the  
326 influence of this variable in the model varies considerably over California, with a strong north-  
327 south direction. The range of the local coefficient is from 1.26 for the northernmost counties to  
328 3.06 for the southernmost counties – evidence that points to heterogeneity in the model structure  
329 within California. The global coefficient and all the local coefficients for this variable are  
330 positive – there is agreement between the two models on the direction of the influence of this  
331 variable. Figure 3c shows a similar distribution in north-south direction of positive road density  
332 coefficient. Figure 3d reveals the opposite for stream density coefficients, with larger values in  
333 the north and smaller values in south. Contrary to our initial thoughts, stream density  
334 demonstrated a negative relation to disease risk. This may reflect that flowing water is normally  
335 not suitable for larval development of the various species of mosquitos that commonly transmit  
336 WNV or that rasterizing the stream database into stream density introduces a component that is  
337 not yet fully understood.

338 Our best ordinary least squares model, the global LSR model (equation 1) produced an  
339 adjusted  $R^2$  of 0.61 ( $p < 0.05$ ) with a corresponding corrected AIC of 567.70. Utilizing the same  
340 environmental variables, our best local GWR model (equation 2) produced an adjusted  $R^2$  of  
341 0.71 ( $p < 0.05$ ) with a corresponding corrected AIC of 551.4. A 16 point decrease in the AIC and  
342 approximately 16% improvement in the model performance suggest that incorporating spatial  
343 data improves the predictive ability of WNV risk.

344

## 345 **Conclusions**

346 One of the frequent technical issues in modeling disease risk is to incorporate local rather  
347 than global associations in these models (Foley et al. 2009). In spatial regression models, a  
348 global model can be used to examine the relationship between disease risk and potential  
349 explanatory factors which are based on the assumption that the relationship is a stationary spatial  
350 process (Miller 2012). For a small and homogenous region of interest, it is reasonable to assume  
351 that the explanatory factors would not change significantly across the region, and the relationship  
352 between WNV risk and the potential factors would also be unchanged. However, important  
353 variables such as topography, climate, and population distribution change greatly when it comes  
354 to a large region like California with an area of over 163,000 square miles. California is  
355 geographically diverse and is equally varied in its range of climates with several climatic sub-  
356 regions recognized. It would be unexpected to find that the spatial stationarity assumption holds  
357 in such a large area.

358 The distinct north-south pattern revealed in our study could be attributable to typical  
359 latitudinal expressions of temperature and precipitation, especially since California has a north-  
360 south length of 1,350 kilometers. This environmental pattern is also a likely contributor to the

361 distribution of different mosquito species in the United States, especially notable in its  
362 manifestation in California. A recent report (CDC 2013) shows that while *Cx. tarsalis* is  
363 distributed throughout California, *Cx. pipiens* is a more important mosquito vector in northern  
364 California, while *Cx. quinquefasciatus* is more important in southern California. While WNV  
365 can be found in a wide variety of ecosystems, the north-south pattern of infected birds detected  
366 in this study may be expressed more noticeably in California due to the north-south differences  
367 in mosquito species distributions as observed in the Centers for Disease Control report.

368         Our results concur that understanding WNV risks is improved when considering spatial  
369 heterogeneity of the variables that affect the risk (Beck et al. 1994). Besides improving  
370 prediction accuracy, spatial heterogeneity can also provide insights into the underlying  
371 ecological processes controlling the distributions of vector populations and zoonotic pathogens  
372 (Wimberly et al. 2008a) because GWR models consider spatial heterogeneity by separating the  
373 large heterogeneous region into smaller, more homogeneous local regions. Fotheringham  
374 (Fotheringham 2009a), stated that an advantage of using GWR is that it accounts for much of the  
375 spatial autocorrelation in the residuals that is usually found in global modeling. Further, it is  
376 possible that a variable that is insignificant at the global level might be important locally.

377         There are several limitations of this study. First, it is assumed that factors suitable for  
378 mosquito habitat increase the likelihood of WNV spread in human populations. On the surface  
379 this seems to be reasonably apparent; however, we do not have specific evidence that this is true.  
380 Second, it is also assumed that the probability of human infection is higher in counties with  
381 multiple confirmed WNV bird cases, another reasonable conjecture with several references in the  
382 literature, but without direct confirmation. Potential problems with this assumption include  
383 varying human population density (e.g. two areas with the same number of infected dead birds

384 reported but one area's human population density is substantially different than the other),  
385 variations in level of public concern (as reports of infected dead birds increase, more people  
386 begin looking for dead birds), and resource availability might bias the reporting of dead birds  
387 (wealthy areas devote disproportionate resources to the issue). Thus, proper surveillance methods  
388 that take into consideration these limitations while collecting infected dead bird data will  
389 contribute to more meaningful results. Third, our approach assumes that people are infected  
390 within the county of their residence, ignoring the possibility of contracting an infection while  
391 traveling outside the county limits. Lastly, road density could also be correlated with dead bird  
392 surveillance effort and might be a potential bias for reporting dead birds. We recognize that if  
393 these assumptions do not hold, modeling WNV risks based on infected dead birds may yield  
394 biased results. However, if the assumptions do hold, the local modeling approaches should  
395 improve predictions of WNV risks.

396         The research described in this paper suggests that a spatially explicit local model using  
397 GWR approaches to adjust for spatial autocorrelation and non-stationarity can yield improved  
398 predictions compared to ordinary LSR modeling of WNV risk. A spatially explicit modeling  
399 technique may be useful in policy-making and decision-making depending on the granularity and  
400 resolution of available data. Identifying the spatial variations in relationships by estimating local  
401 regression parameters allows the spatial distribution and interaction of predictor variables to be  
402 explored. Analyzing local variations in relationships provides those concerned with public health  
403 policy the ability to target resources and to achieve improved outcomes through location-specific  
404 activities (Comber et al. 2011) because spatial heterogeneity can improve predictions by  
405 capturing geographic shifts in the ecological drivers (Wimberly et al. 2008a). While  
406 environmental data used in this research were of fine resolution, WNV disease human incidence

407 data and infected dead bird data that is used is available only at a coarser county scale. We had to  
408 assume that aggregating the environmental data up to the county adequately represented the  
409 environmental conditions presented in the county, but we knew that data aggregation was likely  
410 to introduce some uncertainty into the model. The dead bird model applied in this study can be  
411 used for better understanding of WNV risk and the techniques used could be replicated at finer  
412 spatial scales thus leading to better intervention efforts.

413 In summary, WNV, a globally emerging infectious disease, was found to be  
414 heterogeneously related to environmental factors at the county level throughout California during  
415 the time that our data were collected. Our findings may assist those conducting risk assessments  
416 for WNV transmission in local areas by helping local public health entities allocate resources and  
417 improve preparedness for an outbreak according to region-specific conditions.

418

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424

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- 559

**Table 1** (on next page)

Environmental conditions related to WNV transmission risk.

1

<b>Factors studied</b>	<b>Relation to WNV</b>	<b>References</b>
Streams	Sites for breeding and resting	Cooke et al. 2006; Curtis et al. 2014; Schurich et al. 2014
Temperature	Increases growth rate of vector, decreases egg development cycle and shortens extrinsic incubation period of vector	DeGroot et al. 2014; Kuehn 2012; Srivastava et al. 2001; Wimberly et al. 2008b
Surface slope	Water stagnation creating mosquito breeding ground	Cooke et al. 2006; Ozdenerol et al. 2008; Schurich et al. 2014; Srivastava et al. 2001
Cultivated land	Linkage between habitat used and human-commensal nature of WNV mosquito vectors	Kilpatrick 2011
Developed land	Linkage between habitat used and human-commensal nature of WNV mosquito vectors; warmer microclimates	Kilpatrick 2011
Roads	Sites for breeding and resting along roadsides	Cooke et al. 2006
Vegetation	Sites for breeding and resting.	Brownstein et al. 2002; Cooke et al. 2006; DeGroot et al. 2014; Ruiz et al. 2004a; Schurich et al. 2014; Srivastava et al. 2001
Evapotranspiration	Related to the amount of moisture that is related to mosquito abundance	Liu and Weng 2012; Trawinski and Mackay 2008

2

3

**Table 2** (on next page)

Data sources.

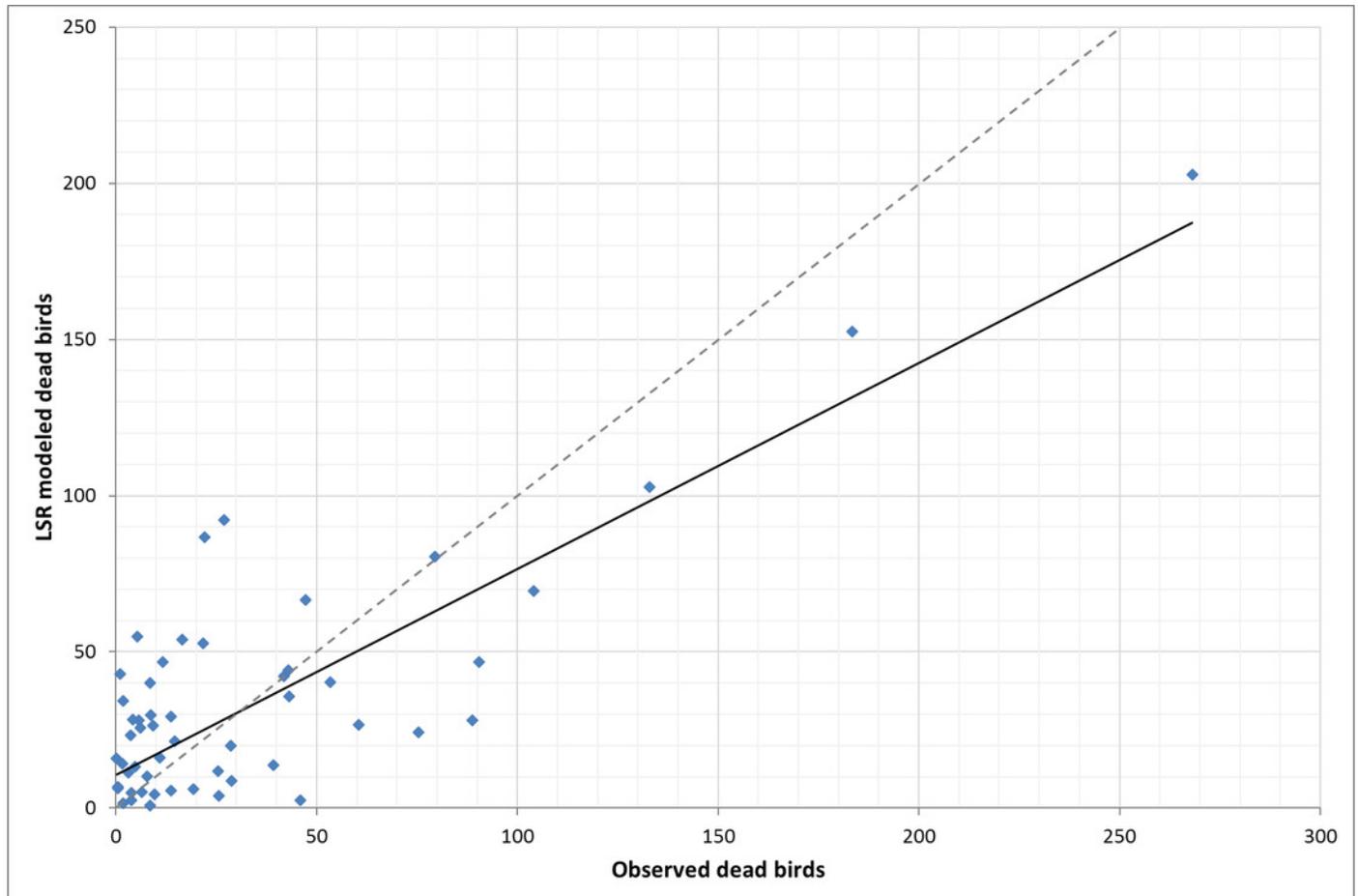
1

Data	Spatial resolution	Source
Elevation	10m	National Elevation Dataset (NED)
LST	1 Km	MODIS aboard the Terra and Aqua satellites
NDVI	250 m	MODIS aboard the Terra and Aqua satellites
Evapotranspiration (ET)	1 Km	MODIS aboard the Terra and Aqua satellites
Streams	available in vector format	U.S. bureau of reclamation
Roads	available in vector format	U.S. Census bureau
Cultivated land	30 m	National Land Cover Database
Developed land	30 m	National Land Cover Database
WNV infected dead birds count	County scale	U.S.G.S. National wildlife health center
WNV human incidence cases	County scale	U.S.G.S. National wildlife health center
Human population	County scale	U.S. Census bureau

2

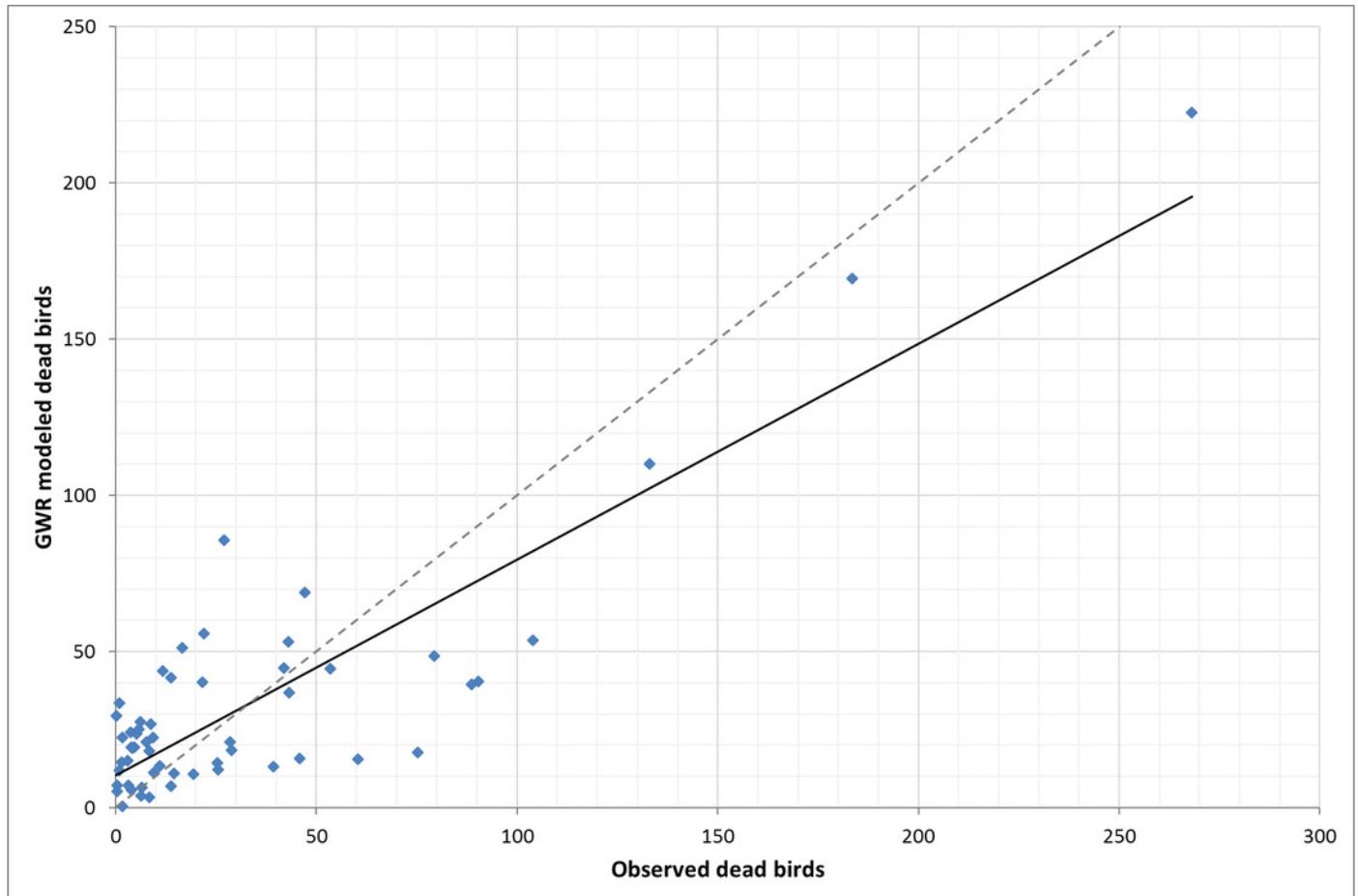
# Figure 1

Trendline plot for global LSR model (model:  $y = 0.6591x + 10.563$ ;  $r^2 = 0.66$ ), dashed line ideal 1:1 relationship.



## Figure 2

Trendline plot for local GWR model (model:  $y = 0.6911x + 10.259$ ;  $r^2 = 0.75$ ), dashed line ideal 1:1 relationship.



## Figure 3

Spatial distribution of (a) standardized residuals; (b) land surface temperature coefficients; (c) road density coefficients; (d) stream coefficients.

