Research Article
Spatial and Temporal Distribution of Tuberculosis in the State of Mexico, Mexico

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Received 17 April 2012; Accepted 29 May 2012

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Tuberculosis (TB) is one of the oldest human diseases that still affects large population groups, mainly in marginal areas and comprising vulnerable groups impacted by extreme poverty, malnutrition, and crowded housing. These groups are prone to infection by the tuberculosis bacilli and to acquiring active TB [2, 3].

In Mexico, the National Population Commission (CONAPO) considers a marginal index based on a census of localities according to the global impact of several total measurements or items, such as lack of education (illiteracy, incomplete grade school) and inadequate housing conditions (without water, drainage, bathrooms, electricity, refrigerator [4], and other basic household equipment, houses with earth floors and crowded living conditions). These items can be used to identify geographic areas with greater marginalization (socioeconomic development) and a greater risk from TB infection.

The World Health Organization (WHO) reported in 2010 that there were an estimated 9.4 million incident cases (range 8.9 million–9.9 million) of TB globally, equivalent to 137 cases per 100,000 population, and that 1.1 million of those cases also tested positive for human immunodeficiency virus (HIV). The mortality of HIV-negative patients with TB was estimated at 1.3 million, this being equivalent to 20 deaths per 100,000 people. The incidence of TB patients in Asia was 55% and 30% in Africa; smaller proportions of cases occurred in the Eastern Mediterranean Region (7%), the European Region (4%), and the Americas Regions (3%) [5].

1. Introduction

Tuberculosis (TB) is an infectious disease, generally chronic, and caused by a group of bacteria: Mycobacterium tuberculosis, M. africanum, M. bovis, M. microti and M. canettii [1]. The main transmission is from person to person by microdrops generated by coughing or sneezing of a person with active TB [1, 2].

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In Mexico, 18,848 TB cases (in all clinical forms) were reported in 2010, which is a rate of 16.77 per 100,000 inhabitants. Of all the cases reported, 15,384 or 13.7 per 100,000 corresponded to lung TB, the mortality rate was 2.14 per 100,000 (in all clinical forms) [6], and the incidence rate was higher in the 60-year-old or 60+-year-old groups of patients, and males had a 1.5 times higher rate than females [3].

In the federal entities, TB presents a well-defined concentration pattern. High incidence rates are present in the western states and the Gulf of Mexico, and lower rates are found in the central states. Tamaulipas and Baja California have double the national rate (13.5 per 100,000 inhabitants). In these states, together with the Veracruz, Chiapas, Nuevo León, Jalisco, Sinaloa, Nayarita, Guerrero, Sonora, Oaxaca, and Chihuahua states, are concentrated 70% of all the identified cases of TB in the country [3].

At present, geographic information systems (GISs) are among the most useful tools in epidemiology, as they can be used to identify geographical areas and population groups with a higher risk of sickness or premature mortality and which therefore require higher preventive care or health information and monitoring of diseases in time and space [7, 8].

Spatial cluster of a disease grouping is one of the most important techniques in epidemiology. In 1995, Kulldorff and Nagarwalla developed a new space statistics method called SCAN for the detection and cluster inference in time-space under the likelihood ratio (LLR) hypothesis [8].

SCAN statistics is a spatial-temporal analysis defined by a large number of transposed cylinders; the circular base defines the geographic area with a radius that can vary from zero to a larger distance influenced by the percentage of the population at risk, while the height of the cylinder represents the temporal parameter. The LLR was calculated for each cylinder. The cylinder with the highest LLR was considered to constitute the most probable cluster. A Monte Carlo simulation was used to calculate the \( P \) value [9].

In the case of TB, various researchers have used GIS to study this infectious disease. Moonan et al. [10] used GIS to identify the geographic locations of TB transmission and incidence in the United States of America during 1993 to 2000. In India, Tiwarin et al. [11] carried out a geospatial investigation of TB occurrence in the Almora district using GIS and the SCAN statistics program. Nunes [12] in Portugal detected spatial and temporal clusters during 2000–2004 by using SCAN. The above-mentioned authors agree that GIS and SCAN are useful tools for vigilance against TB.

The SCAN statistics program has been used worldwide by various researchers with different lengths of search windows: the most frequently used are 25% and 50% of the population [11]. Sabel et al. [13] added some health determinants in their model, such as age, gender, and margination, as covariants as a form of correction or data adjustment.

The objective of this research was to use the spatial statistics program SCAN and GIS to identify the spatial and temporal distribution of TB during 2006 to 2010 and to determine the geographic locations of higher transmission and incidence of TB in the State of Mexico, Mexico.

### 2. Materials and Methods

#### 2.1. Study Location

This study used as its research area the State of Mexico, located in the centre of Mexico at 98°35’30”–100°37’00” west longitude and 18°21’15”–20°17’00” north latitude, and occupying 22,499.95 km², which represents 1.1% of the national territory. It is divided into 125 municipalities that include 4,341 localities (Figure 1).

According to the II Conteo de Población y Vivienda 2005 (II Population and Housing Census 2005), the State
of Mexico had 14,007,495 inhabitants, representing 13.6% of the entire Mexican population and making it the most populated state.

2.2. Data Collection

2.2.1. Population Data. In accordance with the Sistema Nacional de Vigilancia Epidemiológica (Epidemiology Vigilance National Surveillance), tuberculosis cases were reported from 499 localities during the period 2006–2010. Cases in each locality were categorized using a marginalization index [14]: Category 1: very low; Category 2: low; Category 3: medium; Category 4: high; Category 5: very high. The number, age and gender of the inhabitants in each locality were obtained from the II Population and Housing Census 2005 by the “Instituto Nacional de Estadística y Geografía” (INEGI) (National Institute of Statistics and Geography) [15]. Age was classified into five categories: Category 1: 0–4 years old; Category 2: 5–14 years old; Category 3: 15–17 years old; Category 4: 18–59 years old; Category 5: 60 and 60+ years of age. The data for gender was combined with the age categories, and therefore 10 groups of age-gender were created to describe the population of each locality.

2.2.2. Tuberculosis Case Data. The collected cases were diagnosed at several hospitals and healthcare centres and their demographic characteristics were reported to the TB National Registry during 2006–2010. Marginality was obtained from the 2005 marginality index [14] for the locality where each TB case was found. For the statistical strategy, the covariables were age-gender and marginalization.

2.2.3. Geographic Localization Data. For data analysis, locality with the TB case was used as a geographic unit. The georeference came from the latitude-longitude projection system obtained from the Instituto de Geografía Estadística y Catastral del Estado de México (IGECEM) (Institute of Statistical Geography and Tax Revenue of the State of Mexico) [16].

3. Results

There were 2,164 human TB cases detected from 2006 to 2010, distributed in 499 localities and in 125 municipalities. The municipalities with more than 100 TB cases are Ecatepec de Morelos with 365 (16.9%) cases; Netzahualcóyotl with 172 (7.9%); Naucalpan de Juárez with 153 (7%); Tlalnepantla de Baz with 123 (5.7%); Toluca county with 109 cases (5%). The distribution of the 2,164 cases is shown in Figure 2 with an increment from 405 cases in 2006 up to 449 in 2010.

As mentioned in the methodology section, the 10 “age-gender-margination” groups or categories were found in 499 locations with TB cases. A database with 4990 groups was created. Two analyses were performed using the collected information: the first analysis was spatial and the second was a space-time analysis.

3.1. Spatial Analysis. A search window with a maximum length of ≤50% population with high TB rates was used. A primary cluster and five secondary clusters were identified. The centre of the primary cluster was located in the municipality of Valle de Chalco Solidaridad in the locality of Santa Cruz. The relative risk (RR) was 337.68 with 14 TB cases versus the 0.04 expected cases. The second cluster was formed of the municipalities of Acolman, Atenco, Chiautla, Chiconcuac, Ecatepec de Morelos, Papalotla, Tecámac, Teotihuacán, Tepehuaxtoc, Texcoco, and Tezoyuca. The RR was 1.68 with 423 TB cases and the expected RR was 282.2 cases. The third cluster, in the Atizapán de Zaragoza, Villa Nicolás Romero, and Cuautitlán Izcalli municipalities, had a 215.58 RR with seven TB cases compared to 0.03 expected cases. The 10.75 RR for the fourth cluster with 20 TB cases, located in Huixquilucan and Lerma municipalities, had an expected number of cases of 1.88.

Cluster 5, located in the municipalities of Tlalnepantla de Baz, Tultitlán, Apizapán de Zaragoza, and Cuahutitlán Izcalli,
Table 1: TB clusters with a high RR identified in the State of Mexico using a space analysis and a search window maximum length of ≤50% and ≤25% population, adjusted for age-gender-margination.

<table>
<thead>
<tr>
<th>Cluster</th>
<th>Loc.</th>
<th>Centre coordinates</th>
<th>Radius (km)</th>
<th>Obsb</th>
<th>Expc</th>
<th>RRe</th>
<th>LLRc</th>
<th>P</th>
</tr>
</thead>
<tbody>
<tr>
<td>Primary cluster</td>
<td>1</td>
<td>19.287 N, 98.933 W</td>
<td>0</td>
<td>14</td>
<td>0.04</td>
<td>337.68</td>
<td>67.5</td>
<td>0.001</td>
</tr>
<tr>
<td>Secondary clusters</td>
<td>2</td>
<td>19.585 N, -98.945 W</td>
<td>13.11</td>
<td>423</td>
<td>282.20</td>
<td>1.62</td>
<td>35.8</td>
<td>0.001</td>
</tr>
<tr>
<td></td>
<td>3</td>
<td>19.596 N, -99.269 W</td>
<td>4.48</td>
<td>7</td>
<td>0.03</td>
<td>215.58</td>
<td>30.6</td>
<td>0.001</td>
</tr>
<tr>
<td></td>
<td>4</td>
<td>19.306 N, -99.386 W</td>
<td>6.88</td>
<td>20</td>
<td>1.88</td>
<td>10.75</td>
<td>29.3</td>
<td>0.001</td>
</tr>
<tr>
<td></td>
<td>5</td>
<td>19.610 N, -99.191 W</td>
<td>5.08</td>
<td>38</td>
<td>8.77</td>
<td>4.39</td>
<td>26.7</td>
<td>0.001</td>
</tr>
<tr>
<td></td>
<td>6</td>
<td>19.501 N, -99.284 W</td>
<td>5.57</td>
<td>8</td>
<td>0.78</td>
<td>10.24</td>
<td>11.4</td>
<td>0.001</td>
</tr>
</tbody>
</table>

aLocations with TB cases, bObserved number of cases in cluster, cExpected number of cases in cluster, dRelative risk of the cluster, eLog likelihood ratio.

Figure 3: Geography of the identified high TB rate clusters in the State of Mexico detected with a spatial analysis and a search window maximum length of ≤50% and ≤25% population, adjusted for age-gender-margination.

was found to have a 4.39 RR with 38 TB cases compared with the 8.77 expected cases.

The last identified secondary cluster, located in the Tlalnepantla de Baz, Jilotzingo, Atizapán de Zaragoza, and Naucalpan de Juárez, showed a 10.24 RR with eight TB cases against the 0.78 expected cases.

A maximum length of ≤25% search window was used for a population with high TB rates and the results of the analysis were similar to those obtained with a ≤50% search window. General information about the identified clusters is shown in Table 1, and their geography is shown in Figure 3.

3.2. Space-Time Analysis. A primary cluster was identified using a search window maximum of ≤50% population with high TB rates. The cluster centre was located in the Tultitlán, a municipality formed of the Tlalnepantla de Baz, Cuautitlán, Cuautitlán Izcalli, and Atizapán de Zaragoza municipalities. The relative risk (RR) during the 2007–2009
Table 2: A cluster with high rates of tuberculosis in the State of Mexico was detected with a spatial-temporal analysis and a search window maximum length of \( \leq 50\% \) and \( \leq 25\% \) risk population, adjusted for age-gender-margination.

<table>
<thead>
<tr>
<th>Cluster</th>
<th>Loc.</th>
<th>Centre coordinates</th>
<th>Radius (km)</th>
<th>Year</th>
<th>Obs(^b)</th>
<th>Exp(^c)</th>
<th>RR(^d)</th>
<th>LLR(^e)</th>
<th>P</th>
</tr>
</thead>
<tbody>
<tr>
<td>Primary cluster</td>
<td>1</td>
<td>6</td>
<td>19.315 N, −98.928 W</td>
<td>3.49</td>
<td>2006–2008</td>
<td>16</td>
<td>0.2</td>
<td>81.08</td>
<td>54.467248</td>
</tr>
<tr>
<td>Secondary clusters</td>
<td>2</td>
<td>19</td>
<td>19.585 N, −98.945 W</td>
<td>11.93</td>
<td>2006–2008</td>
<td>267</td>
<td>140.59</td>
<td>2.03</td>
<td>48.880307</td>
</tr>
</tbody>
</table>

\(^a\)Locations with TB cases, \(^b\)Observed number of cases in cluster, \(^c\)Expected number of cases in cluster, \(^d\)Relative risk of the cluster, \(^e\)Log likelihood ratio.

*Cluster detected with a search window maximum length of \( \leq 50\% \) risk population.

Figure 4: Geographic locations of the identified clusters with high rates using a space-time analysis and a search window maximum length of \( \leq 50\% \) and \( \leq 25\% \) population, adjusted for age-gender-margination.

A search window maximum length of \( \leq 25\% \) population was used to identify three significant clusters. The primary cluster was located in the Valle de Chalco Solidaridad, Ixtapaluca, and a small part of Chalco. The RR was 81.08, with 16 TB cases during the period 2008–2006, compared with 0.2 expected cases.

Two secondary clusters were identified. One was similar to that described above using the \( \leq 50\% \) population search window. The other was formed by 11 municipalities: Acolman, Atenco, Chiconcuac, Ecatepec de Morelos, Papalotla, Tecamac, Teotihuacan, Tepetlaoxtoc, Texcoco, and Tezoyuca. The RR was 2.03 with 267 cases of TB during the years 2006–2008 against expected cases of 140.50. General information about the identified clusters is shown in Table 2 and their geography in Figure 4.

4. Discussion

In the field of epidemiology one of the most important analyses is the detection in space-time of a disease cluster, since a non-control-spreading of the disease takes place and a social or economic factor favours cluster formation. The use of GIS plus spatial statistics has been used to determine spatial distribution patterns of various infectious and noninfectious diseases [18–22]. SCAN, available in
the SaTScan V8 software, has been used worldwide to detect various disease clusters, including TB.

GISs plus molecular diagnosis techniques have been used to locate TB clusters and to identify the *Mycobacterium* responsible for the outbreak [10]. Frequently, however, it is not possible to obtain a detailed and complete analysis since the information is mainly based on a positive bacilloscopy.

However, this study obtained data on TB according to age, gender, and marginality index, representing valuable information since the highest rate of this disease occurs in adults of 60 and 60+ years of age [3], in males 1.5 to 1 rate and in marginal groups [2].

With the SCAN statistical results and a 25% and 50% search window population and a spatial analysis ($P \leq 0.05$), six clusters were identified, showing that TB is not randomly distributed in the State of Mexico but in clusters in a spatial pattern. In similar studies by Navae et al. [23] in Acapulco, Mexico, using a different spatial method, clusters were not found. We used two search windows (50% and 25%) in order to find small clusters but identification differences were not seen in cluster identification. These results were in agreement with those reported by Tiwari et al. [11].

The six identified clusters are located close to and around Mexico City (Federal District), the largest city in the Mexico. The highest incidence rates of TB are found in urban zones [24] in comparison with rural areas. Our results agreed with those reported by Moonan et al. [10]. They found a strong association between the strains of TB clusters and the distance to the centre of urban zones.

Three TB clusters were identified with spatial and temporal analysis using the same search windows; two of these clusters grouped the TB cases from the 2006 to 2008 period, and the cases from 2007 to 2009 were in the third cluster. The three clusters all contained the 2008 TB cases, making it the year with the highest incidence.

Of the three identified clusters, only one is in agreement with those identified with the spatial-only analysis, but the others are located in areas close to the spatial clusters.

The results of the present study are of great importance in TB epidemiological surveys in the State of Mexico. The presence of clusters is valuable to the healthcare system since it is possible to find small clusters but identification differences were not seen in cluster identification. These results were in agreement with those reported by Moonan et al. [10]. They found a strong association between the strains of TB clusters and the distance to the centre of urban zones.

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The results of the present study are of great importance in TB epidemiological surveys in the State of Mexico. The presence of clusters is valuable to the healthcare system since strategy can be revised accordingly in those areas containing TB clusters.

5. Conclusions

TB is spatially clustered in the State of Mexico, Mexico. Nine clusters were identified using two types of analysis: spatial and spatial-temporal. Such clusters were mainly found in Valle de Chalco Solidaridad, Atenco, Ecatpec de Morelos, Tlahuapan de Baz, Acolma, Tultitlán, Cuautitlán Izcalli, Atizapán de Zaragoza, Huixquilucan, and Naucalpan de Juárez, all municipalities that are very close to Mexico City (Federal District).

Acknowledgments

The authors extend their thanks to The Health Institute of the State of Mexico for providing the necessary help and support for the study. This work is derived from the thesis for the degree of Doctorate in Sciences of Health, registered in the PNPC-CONACYT, and benefited from the economic contribution of Project SIEA-UAEM 2742/2008U.

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