

Spatio-temporal distribution analysis of TB in Xinjiang Uygur Autonomous Region, China

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Research article

Keywords: Tuberculosis, Spatial autocorrelation, Spatio-temporal scanning analysis

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Abstract

Background: Tuberculosis (TB) is a major global public health problem, which also affects economic and social development. China has the third largest burden of tuberculosis in the world. TB control made the slowest progress in western China while the highest prevalence of it showed up in Xinjiang. The study was conducted to investigate the spatial epidemiological features of pulmonary tuberculosis in Xinjiang Uygur Autonomous Region (referred to as Xinjiang) and compared the regional differences in the incidence of TB, for the 2013-2016 which can provide scientific reference for TB prevention and control.

Methods: Based on the TB monitored data, descriptive statistics was used to analyze the distribution characteristics of TB patients. Spatial correlation analysis and spatio-temporal scanning techniques were used to explore the clustering of TB in Xinjiang.

Results: A total of 178,674 TB cases were notified in Xinjiang from 2013 to 2016 with an average annual incidence of 195.32/100,000. The incidence of TB in Xinjiang showed an upward trend. Male and female patients accounted for 52.56% and 47.44% respectively with the sex ratio being 1.11:1. The number of cases continuously increased with the increasing age and the elderly TB patients aged 60 years and above accounted for 46.77%. Most of the patients with TB were farmers and shepherds, accounting for 72.11%. The incidence of TB presented an upward trend from east to west and from north to south. Obvious spatial aggregation was observed in the incidence of TB in 98 counties and districts from 2013 to 2016 and the global Moran's I was 0.5768 ($P < 0.001$). The reported incidence rate of TB showed remarkable seasonality. The hot spots of TB were mostly concentrated in the southern Xinjiang with Kashgar as the center, while the cold spots were in northern Xinjiang with Urumqi as the center.

Conclusion: The TB incidence displayed spatial and temporal aggregation at the levels of district and county in Xinjiang during 2013-2016, with high risk areas relatively concentrating in the southern Xinjiang. It is necessary to conduct targeted TB prevention and control in key areas and allocate health resources reasonably.

Keywords: Tuberculosis; Spatial autocorrelation; Spatio-temporal scanning analysis

Background

Tuberculosis (TB) is an airborne infectious disease caused by the mycobacterium tuberculosis, which not only typically infects the lungs, but also affects other parts of the body [1-2]. TB continues to be a significant public health problem in the world [3]. According to the report of the World Health Organization (WHO), TB ranks first among the most important infectious diseases in the world, though its incidence has decreased slowly by roughly 1.5% per year since 2000 [4-7]. Globally, there were 9.6 million incident cases of active TB disease in 2014, with 1.5 million TB related deaths, making TB the leading global infectious disease killer. According to the global TB report released by the WHO in 2016, China was the top third among 22 high-TB burden countries, with estimated 930,000 TB patients in 2015 [8-9]. Although China has been working hard to struggle against TB, it remains the third largest burden country in the world, after India and Indonesia.

TB control made the slowest progress in western China while the highest prevalence of it showed up in Xinjiang [10-11]. From 2010 to 2014, the Communicable Disease Network Direct Reporting System

showed that, 264,000 cases of TB were reported in Xinjiang, with 1,144 TB related deaths. The incidence of TB in Xinjiang showed an upward trend in 2013-2016. The prevalence of TB was rising rapidly from 157.83/100,000 in 2011 to 202.32/100,000 in 2016 among residents, which was far beyond the nation average, see Fig. 1. It was likely attributed to the implementation of the direct reporting network system launched in 2004 for infectious disease (include TB) and public health emergency to significantly improve disease monitoring and early warning [1]. The improvement of direct reporting network system might develop into a factor promoted the rise and change of TB case notifications in Xinjiang.

There were regional differences in the prevalence of TB in Xinjiang. Researches showed that TB had a specific spatial distribution pattern [11]. Understanding such spatial variations in TB prevalence and its determinants within a social, spatial, and temporal context is crucial for improved targeting of interventions and resources. Geospatial analytical methods, such as geographic information systems (GIS) and spatio-temporal scanning analysis, are effective tools for helping to achieve such understanding [12-14]. In China, there were some studies intended to reveal the spatio-temporal distribution characteristics of TB under province, or nationwide, however, less discussion on the temporal and spatial distribution of tuberculosis in Xinjiang in recent years [15-18]. Therefore, the main objectives of this study were to investigate the temporal trends and spatial patterns of the TB surveillance data of Xinjiang from 2013 to 2016 by epidemic characteristics analysis, spatial auto-correlation analysis and spatio-temporal scanning analysis.

Methods

1. Data source

District and county levels data of reported TB monthly cases and incidence from 2013 to 2016 were obtained from the Xinjiang Center for Disease Prevention and Control (Xinjiang CDC). Data from the Communicable Disease Network Direct Reporting System and the Xinjiang Statistical Yearbook were also collected.

2. Statistical analyses

(1) Descriptive statistics

Descriptive statistics include the distribution of year, career, gender and age groups were adopted to describe the epidemic characteristics of **morbidity** and reported cases. Chi-square test was used to analyze the trend of **incidence** annually. The spatio-temporal clustering analysis with a Poisson model was applied to identify country at high risk for TB during 1 January 2016 and 31 December 2016. To visualize the cluster pattern in a geographical context, the geographic information system (GIS) software ArcGIS 10.4.1, SaTScan 9.4, and GeoDa 1.6.0 were used.

(2) Spatial autocorrelation

The examination of spatial data is strongly affected by the location from which observations are made. Neighboring regions affect each other and proximate locations often share more similarities than widely-spaced locations spatial autocorrelation is measured using both global and local metrics. Spatial autocorrelation describes the correlation between variables in a spatial region and the same variable in its adjacent region, which could provide clues for some geographic regions to seek for the factors affecting diseases. Spatial autocorrelation analysis includes global spatial autocorrelation and local spatial autocorrelation [19-20]. To calculate spatial dependence, Moran's I (a statistical method that measures spatial autocorrelation) was employed. Moran's I is the most commonly used global autocorrelation index, whose value ranges from -1 to 1. Positive value means positive correlation; negative value means negative correlation, and 0 means non-correlation, namely spatial random distribution.

1) The global Moran's I

The global Moran's I quantified the similarity of observations among adjacent geographical units from a global perspective was used to analyze the overall spatial autocorrelation degree and spatial distribution pattern [1]. The global Moran's I is defined as

[Due to technical limitations this formula could not be inserted. It can be found in the supplemental file "Formulas.docx" Formula #1]

where n is the number of samples, x_i is the observed value in location i , \bar{x} is the mean of the observed value across all locations, and w_{ij} is an element value in the binary spatial weight matrix that describes the spatial relationship between location i and location j . The most frequently used spatial contiguity based weight matrices were applied here, when two countries or districts share a geographical border $w_{ij}= 1$, otherwise $w_{ij}= 0$.

In order to test the spatial autocorrelation between regions, the normalized statistic $Z(I)$ was used to test Moran's I index, corresponding to $P < 0.05$, indicating a significant correlation. The statistic $Z(I)$ is defined as follows

[Due to technical limitations this formula could not be inserted. It can be found in the supplemental file "Formulas.docx" Formula #2]

The global Moran's I ranges from -1 to 1 due to the use of the standardized spatial weight matrix.

2) The local Moran's I

Local spatial autocorrelation statistics could be used to identify different spatial patterns (or spatial aggregation patterns) that might exist in different spatial locations. This allows us to observe local non-stationary in different spatial locations and to find spatial heterogeneity among the data. In our study, the local Moran scatterplot is a two-dimensional scatterplot in which each point represents a country or a district, where

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Its standardized statistical formula $Z(I_i)$ is defined by

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The scatterplot is centered on 0 and is divided into four quadrants that represent different types of spatial association. The first quadrant is the High-High (HH) cluster quadrant, i.e., countries or districts with high TB incidence surrounded by countries or districts with high TB incidence. Similarly, the second, third and fourth quadrants are the Low-High (LH), Low-Low (LL) and High-Low (HL) cluster quadrants, respectively.

The registration data of TB merged with a vector map were used to build spatial databases by ArcGIS 10.4.1. Global and local Moran's I were calculated by GeoDa 4.6.0 software respectively, as well as spatio-temporal scanning analysis was studied by SaTScan 9.3 software to detect the spatial autocorrelation and cluster range of the distribution.

Ethical Review

The study protocol and utilization of TB related data were reviewed by Xinjiang Uygur Autonomous Region center for Disease Control and Prevention and no ethical issues were identified. Therefore, no ethics approval was required by our Investigation Review Board.

Discussion

The prevalence of TB in Xinjiang was mainly characterized by high infection rate, high prevalence, high mortality and low cure rate. The incidence of TB had obvious spatio-temporal distribution. Spatial scientists, practitioners, and policy makers are interested in understanding the spatial variation in TB incidence at various geographic scales and resolutions. This study identified the spatio-temporal patterns of TB at the district and country level in China from January 2013 to December 2016.

The geographic distribution of TB incidence tends to vary across a geographic landscape. Spatial correlation analysis and spatio-temporal analysis were used to detect spatial distribution characteristics and clustering patterns in the 98 districts of Xinjiang. Spatial correlation analysis revealed that the incidence of TB in the counties showed aggregated distribution with a positive global spatial autocorrelation in 2013-2016.

The retrospective space-time scan statistics, calculated by using the discrete Poisson probability model, was used to identify the temporal, spatial, and spatio-temporal clusters of TB at the county and district levels in Xinjiang. The thematic maps of newly reported TB cases and TB clusters provide a novel spatial understanding of the distribution of TB cases in Xinjiang. The most likely pure space cluster was mainly concentrated in the south of Xinjiang, covering seven counties such as Kashgar City, Aksu City, Minfeng

County and surrounding areas and clustered in the time frame from September 2014 to December 2016. Temporal scan result showed that the incidence of tuberculosis in Xinjiang had seasonal characteristics, with multiple occurrences in November and December. The space-time scanning results showed eighteen significant space-time clusters of TB in Xinjiang in 2016, which could be helpful in prioritizing resource assignment in high-risk areas for TB control and elimination in the future. The most likely spatio-temporal cluster was mainly concentrated in the south and west regions of Xinjiang, covering Kashi and Heshuo in southern Xinjiang and the Balikun Kazakh Autonomous County in eastern Xinjiang and clustered in the time frame from March 2016 to August 2016. The results of spatio-temporal scanning indicated that the capability and utility of the spatio-temporal approach in epidemiology and suggested that the high risk periods and areas of TB should be paid more attention to monitoring and early warning.

Conclusions

It is well known that the incidence of TB is related to demographic characteristics such as gender, age, occupation, and ethnicity, as well as geographical factors, economic development, and medical condition. We used spatial statistics to observe the TB epidemic in Xinjiang from 2013 to 2016. The research results showed that TB continues to be a significant public health problem in Xinjiang. Southern regions of Xinjiang have higher-TB burden, due in part to a less-developed economy and poverty. Analysis of the influencing factors of TB in Xinjiang is the direction of our further research.

Declarations

Ethics approval and consent to participate

Xinjiang Uygur Autonomous Region center for Disease Control and Prevention approved the use of an anonymised database of routinely collected TB data for this analysis.

Competing Interests

The authors declare that there is no conflict of interests.

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Authors' contributions

LPZ and YLZ analyzed the data. LPZ wrote this manuscript. All Authors read and approved the manuscript.

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Tables

Table 1

Table 1. Global Moran's I and test value of TB incidence between 2013 and 2016

Year	Moran's I	Z(I)	P
2013	0.5097	13.1823	0.001
2014	0.5376	12.9040	0.001
2015	0.5115	13.1406	0.001
2016	0.5500	13.2751	0.001

Table 2

Table 2. Pure space scan results of TB in 98 countries and districts in Xinjiang, 2016

CLUSTER	Cluster country	LLR	Observed cases	Expected cases	RR	P
1	Wuqia County, Shufu County, Kashgar City, Shule County, Artux City, Akto County, Yengisar County	1865.38	8438	4184.91	2.22	<0.001
2	Wushi County, Akqi County, Kalpin County, Wensu County, Aksu City, Awat County	1350.33	6469	3246.01	1.99	<0.001
3	Yutian County, Qira County, Minfeng County, Lop County, Hotan City	559.21	3785	2118.34	1.85	<0.001
4	Makit County, Shache County, Yopurga County, Zepu County, Jiashi County, Yecheng County	558.94	7126	4783.13	1.49	<0.001
5	Kuqa County, Xinhe County, Xayar County, Baicheng County	290.27	3724	2472.30	1.51	<0.001
6	Toksun	154.92	606	270	2.43	<0.001
7	Shanshan City	35.29	724	521.89	1.39	<0.001
8	Ruoqiang County, Qiemo County	28.72	342	20.57	1.55	<0.001

Table 3

Table 3. TB Pure time scan results of TB in Xinjiang, 2016

Year	Cluster time(month)	Observed cases	Expected cases	LLR	RR	P
2016	2016/11-2016/12	7417	6135.83	152.5264	1.26	<0.001

Table 4

Table 4. Space-time cluster analysis with monthly precision of TB incidence for Xinjiang 2016.

Cluster	Area (District and County Levels)	Coordinates /Radius[km]	Period	Observed cases	Expected cases	LLR	RR	P
1	Hoxud County[Kashgar City, Barkol Kazak Autonomous County	(42.818800 N, 85.187900 E) / 123.15 km	2016/3- 2016/8	2113	420.00	1760.776	5.28	<0.001
2	Xinyuan County	(41.697000 N, 85.700400 E) / 0 km	2016/3- 2016/8	1748	467.13	1048.655	3.88	<0.001
3	Huocheng County[Tacheng City] Toutunhe District[Bohu County	(38.928100 N, 89.743700 E) / 399.99 km	2016/1- 2016/6	866	225.08	531.610	3.92	<0.001
4	Yengisar County	(42.036500 N, 81.891300 E) / 0 km	2016/11- 2016/12	380	66.79	348.784	5.74	<0.001
5	Shule County[Gongliu County	(41.253400 N, 79.273300 E) / 92.59 km	2016/12	299	40.93	337.437	7.36	<0.001
6	Lop County	(40.051600 N, 80.431300 E) / 0 km	2016/11- 2016/12	343	72.81	262.429	4.75	<0.001
7	Artux City	(44.158500 N, 87.062400 E) / 0 km	2016/10- 2016/11	383	102.89	224.360	3.75	<0.001
8	Luntai County[Bachu County, Hejing County, Yecheng County	(48.305600 N, 86.398100 E) / 122.19 km	2016/5- 2016/10	806	359.50	206.985	2.27	<0.001
9	Wuqia County[Altay City][Zhaosu County	(44.835900 N, 81.864100 E) / 88.74 km	2016/8- 2016/12	628	332.15	105.362	1.91	<0.001
10	Emin County	(45.647200 N, 83.884600 E) / 0 km	2016/3- 2016/8	220	80.98	81.125	2.73	<0.001

11	Xinshi District, Wenquan County	(46.252000 N, 90.393800 E) /	2016/10- 2016/12	194	67.86	77.865	2.87	<0.001
		82.60 km						
12	Usu City	(38.367100 N, 80.035600 E) /	2016/12	179	81.29	43.713	2.21	<0.001
		0 km						
13	Jinghe County	(40.393800 N, 82.916300 E) /	2016/11- 2016/12	145	76.02	24.717	1.91	<0.001
		0 km						
14	Tianshan District	(43.369100 N, 83.549100 E) /	2016/10- 2016/11	153	89.12	18.863	1.72	<0.001
		0 km						
15	Qira County	(39.964300 N, 81.544600 E) /	2016/5- 2016/6	220	142.31	18.226	1.55	<0.001
		0 km						
16	Baicheng County	(37.147600 N, 79.907300 E) /	2016/3- 2016/8	395	291.06	16.824	1.36	<0.001
		0 km						
17	Shanshan City	(39.592300 N, 77.222100 E) /	2016/3- 2016/4	190	123.50	15.407	1.54	0.00017
		0 km						
18	Yining City	(44.340600 N, 85.465400 E) /	2016/8- 2016/12	209	143.35	13.211	1.46	0.0013
		0 km						
19	Burqin County∩Moyu County∩Wensu County	(45.981700 N, 85.593800 E) /	2016/3	26	13.63	4.425	1.91	0.988*
		55.70 km						

* No statistical significance.

Figures

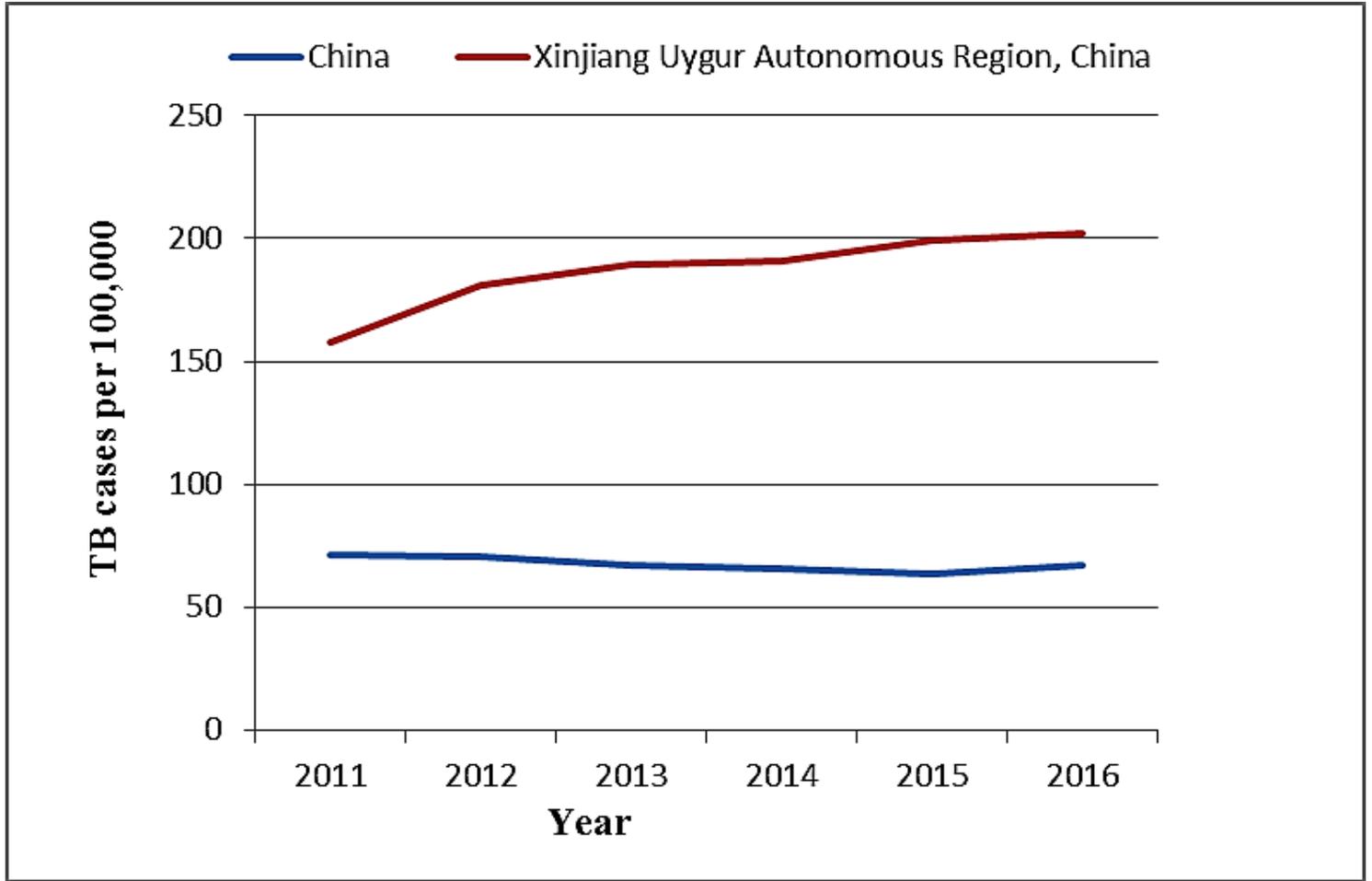


Figure 1

Comparison of TB incidence

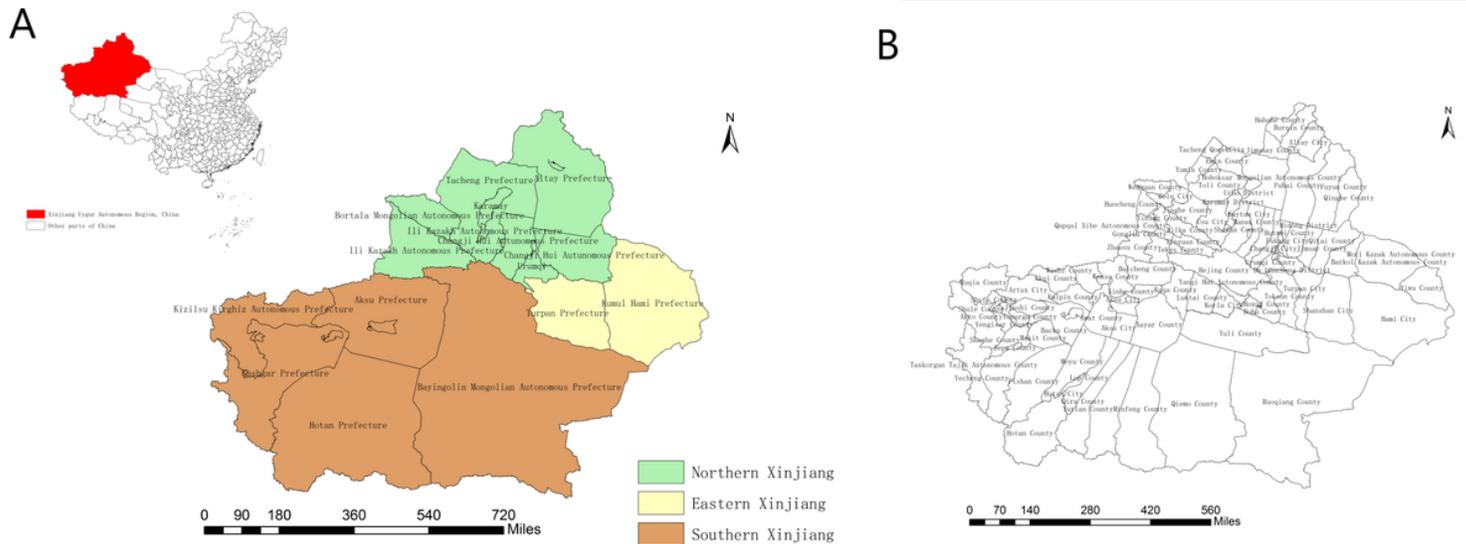


Figure 2

Geographical location of Xinjiang (a) 14 prefectures of Xinjiang, China (b) 98 districts and counties of Xinjiang, China. Note: The designations employed and the presentation of the material on this map do not imply the expression of any opinion whatsoever on the part of Research Square concerning the legal status of any country, territory, city or area or of its authorities, or concerning the delimitation of its frontiers or boundaries. This map has been provided by the authors.

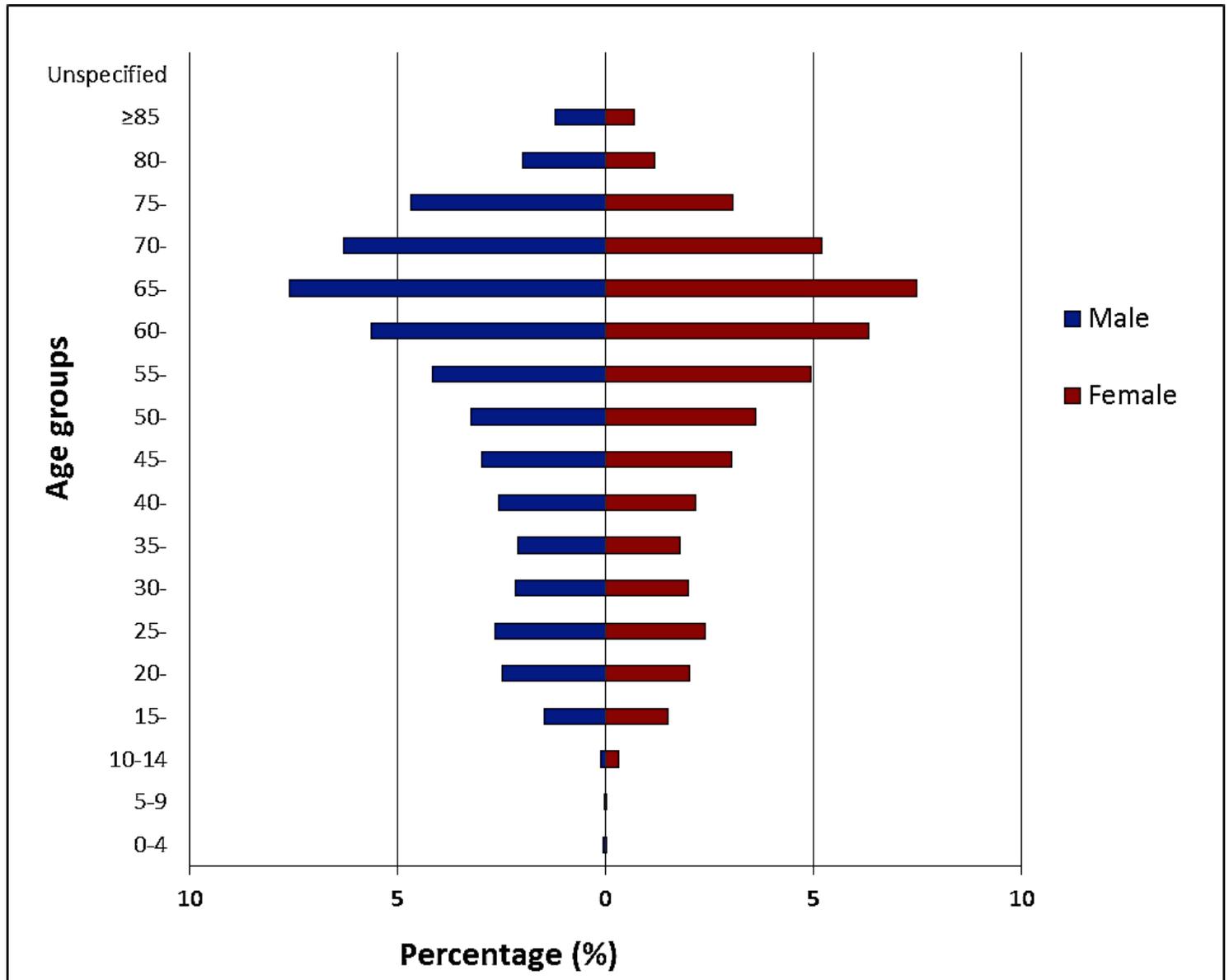


Figure 3

Comparison of TB incidence in terms of age groups and gender in Xinjiang, China 2016

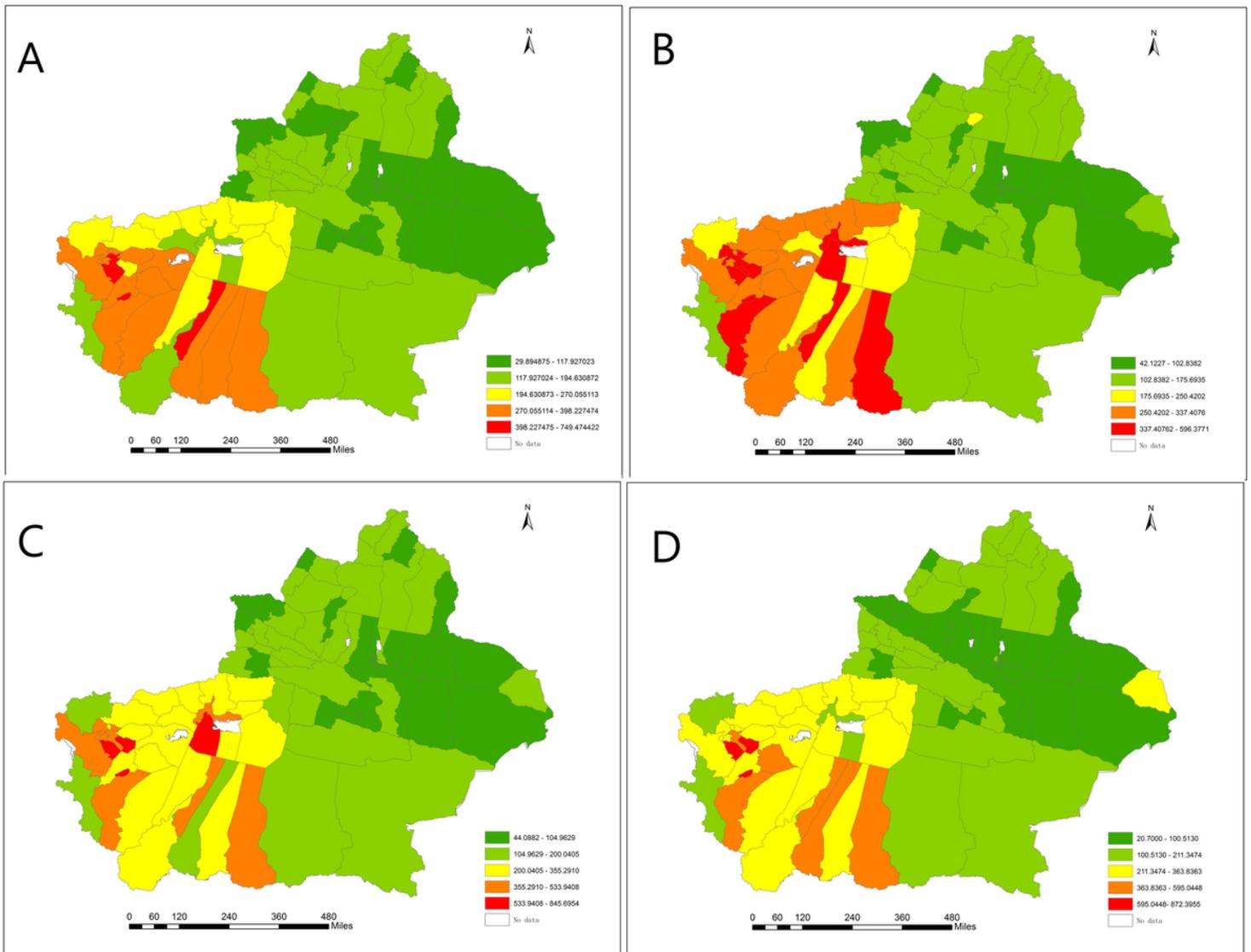


Figure 4

Comparison of TB incidence from 2013-2016 in Xinjiang, China (a) Tuberculosis cases rate per 100,000 residents, 2013 (b) Tuberculosis cases rate per 100,000 residents, 2014 (c) Tuberculosis cases rate per 100,000 residents, 2015 (d) Tuberculosis cases rate per 100,000 residents, 2016. Note: The designations employed and the presentation of the material on this map do not imply the expression of any opinion whatsoever on the part of Research Square concerning the legal status of any country, territory, city or area or of its authorities, or concerning the delimitation of its frontiers or boundaries. This map has been provided by the authors.

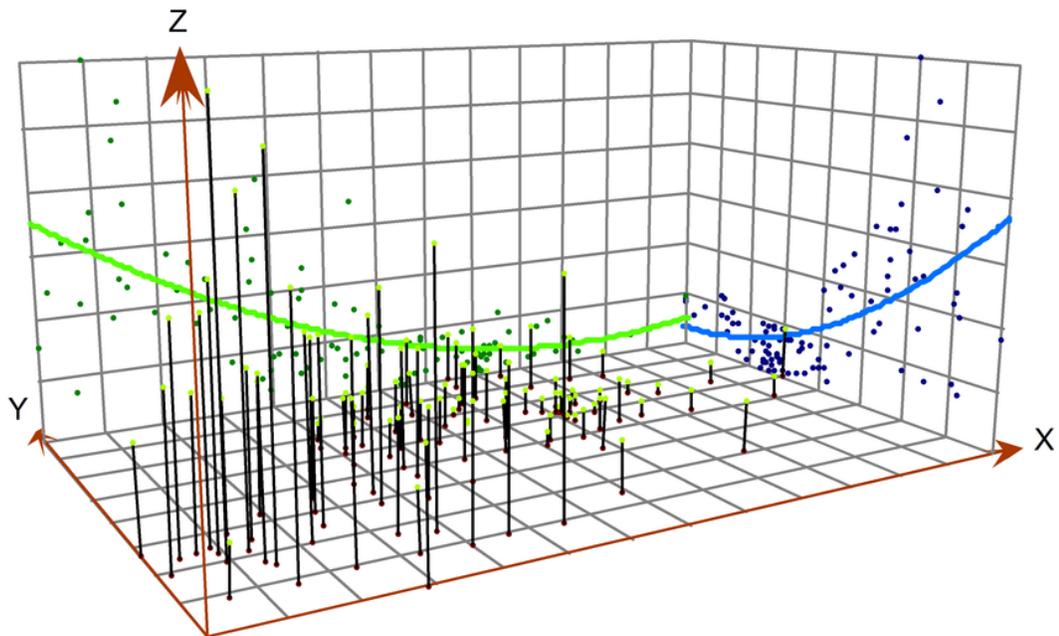


Figure 5

Trend analysis of TB incidence in Xinjiang, China (2013-2016)

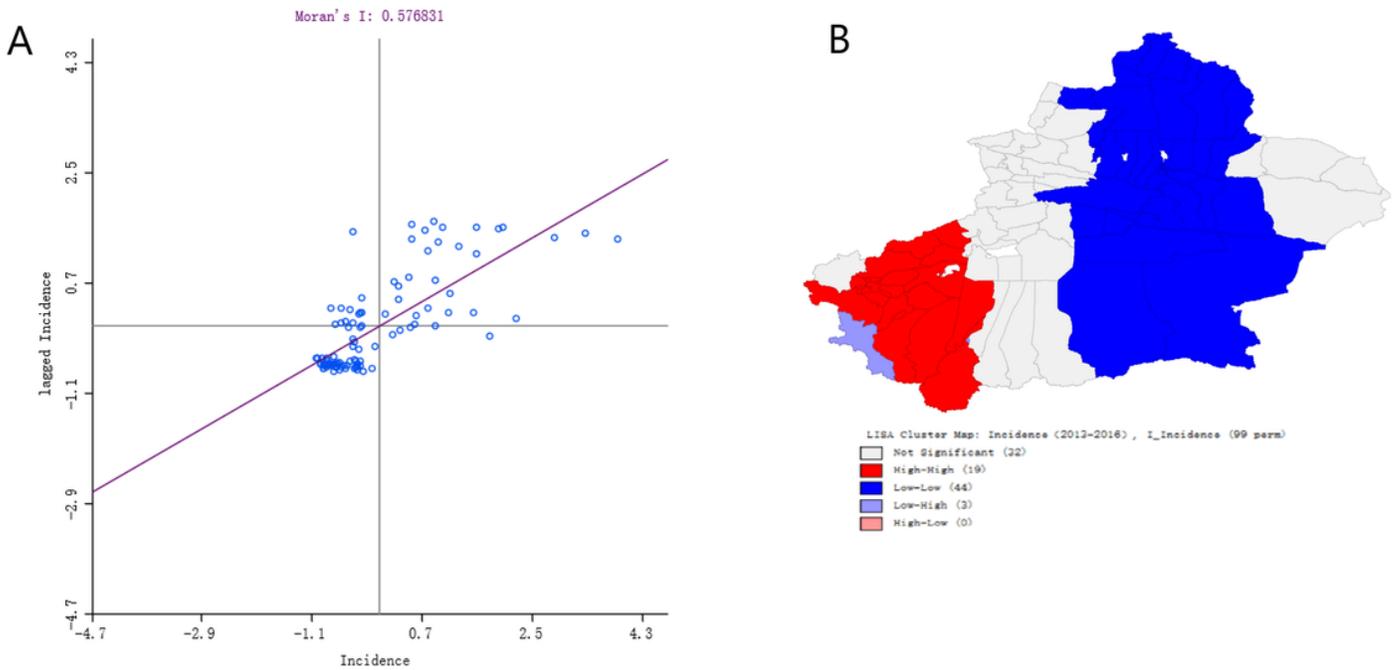


Figure 6

Moran scatter plot (a) and cluster map (b) of TB average annual incidence in 98 districts and countries of Xinjiang from 2013 to 2016. (a)Global Moran's I of TB annual incidence in Xinjiang from 2013 to 2016 (b)LISA cluster map of TB annual incidence in Xinjiang from 2013 to 2016 ($P < 0.01$). Note: The designations employed and the presentation of the material on this map do not imply the expression of any opinion whatsoever on the part of Research Square concerning the legal status of any country, territory, city or area or of its authorities, or concerning the delimitation of its frontiers or boundaries. This map has been provided by the authors.

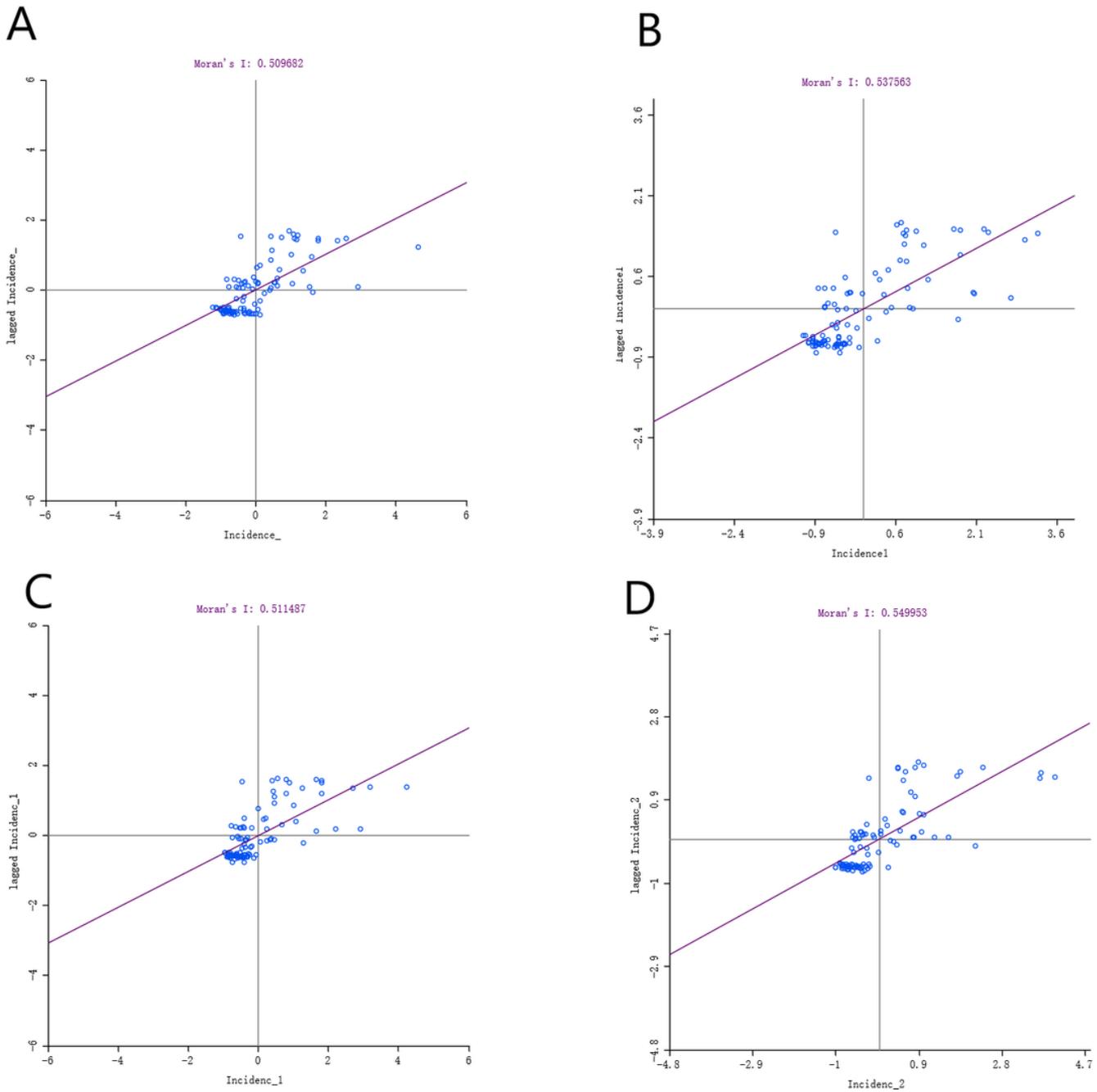


Figure 7

Moran scatter plot of TB incidence in 2013-2016 (a)Moran scatter plot of TB incidence in 2013 (b)Moran scatter plot of TB incidence in 2014 (c)Moran scatter plot of TB incidence in 2015 (d)Moran scatter plot

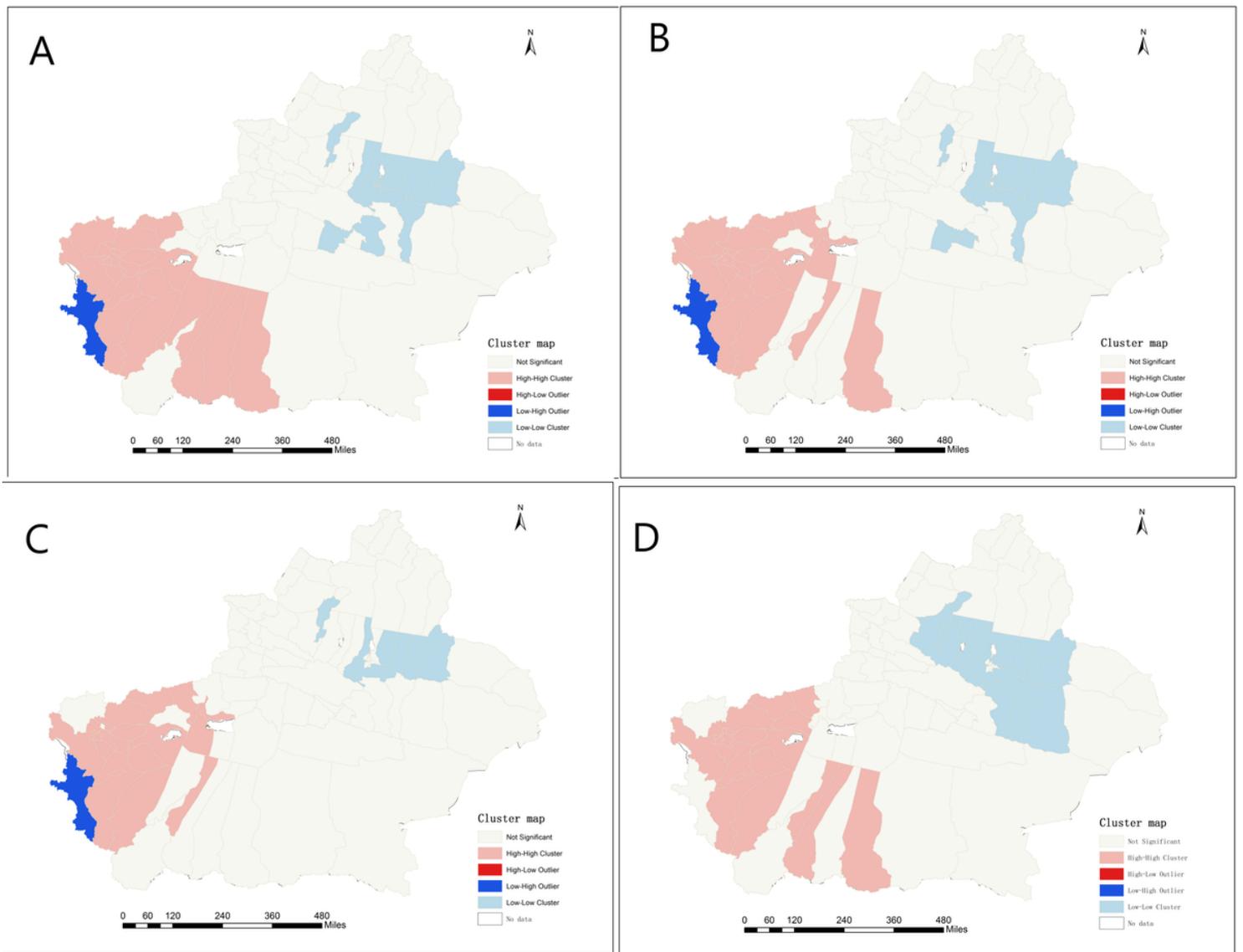


Figure 8

Cluster map of Xinjiang in China, 2013-2016 (a)Cluster map of 2013 (b)Cluster map of 2014 (c)Cluster map of 2015 (d)Cluster map of 2016. Note: The designations employed and the presentation of the material on this map do not imply the expression of any opinion whatsoever on the part of Research Square concerning the legal status of any country, territory, city or area or of its authorities, or concerning the delimitation of its frontiers or boundaries. This map has been provided by the authors.



Figure 9

Gini coefficient at spatial window stops

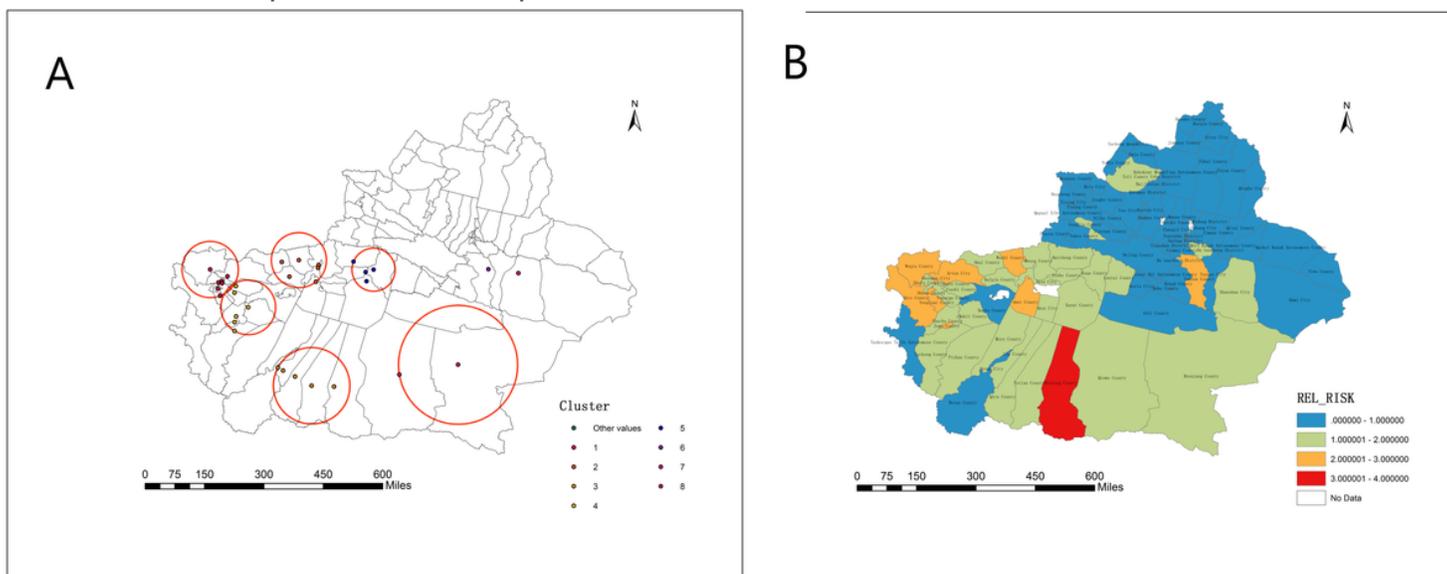


Figure 10

Pure space scan results of 92 countries and districts in Xinjiang, 2016 (a)Cluster results of pure space scan (b)Relative risk map. Note: The designations employed and the presentation of the material on this map do not imply the expression of any opinion whatsoever on the part of Research Square concerning the legal status of any country, territory, city or area or of its authorities, or concerning the delimitation of its frontiers or boundaries. This map has been provided by the authors.

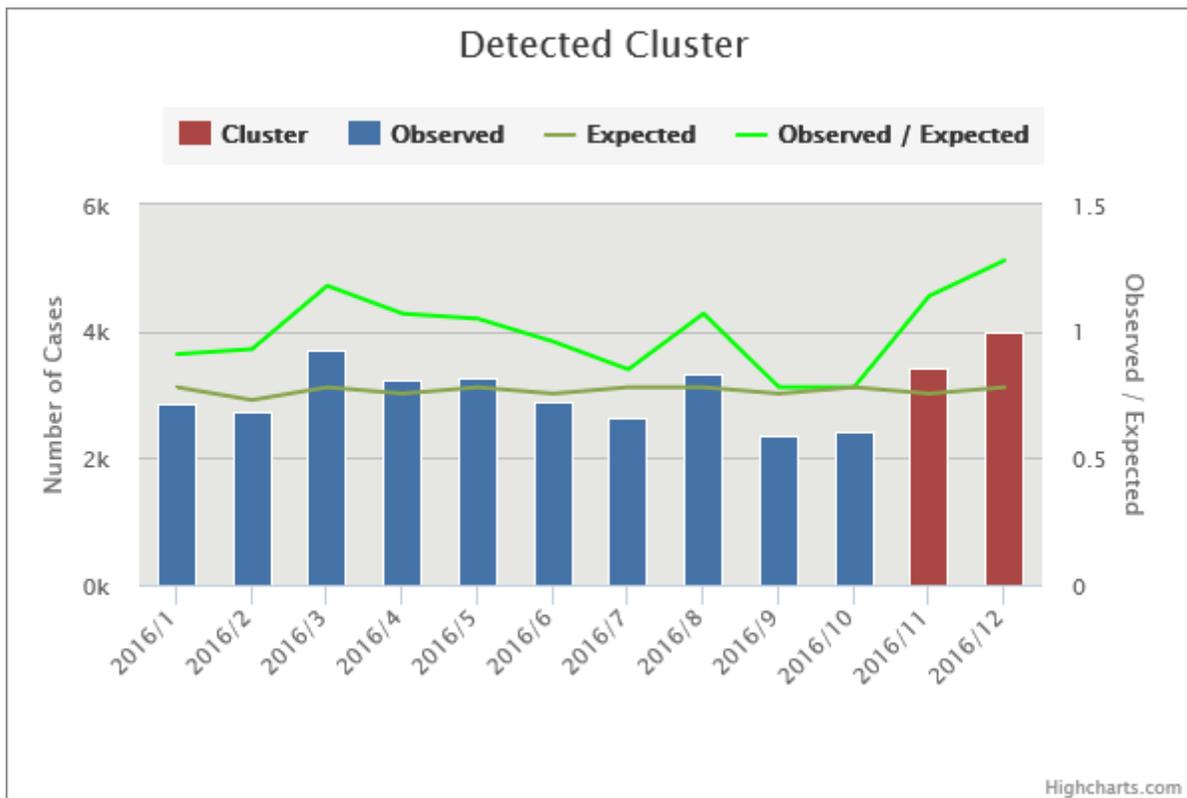


Figure 11

Time aggregation results of TB in Xinjiang, 2016

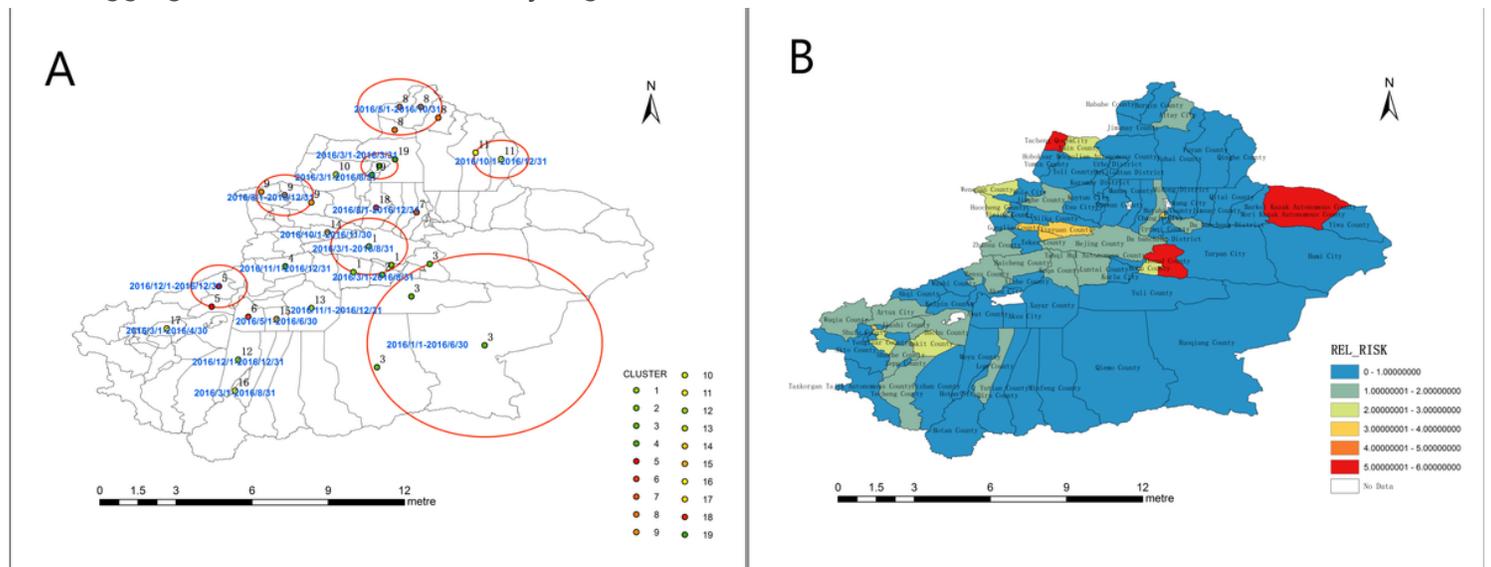


Figure 12

Temporal-space clustering area map of 98 districts and counties in Xinjiang. Note: The designations employed and the presentation of the material on this map do not imply the expression of any opinion whatsoever on the part of Research Square concerning the legal status of any country, territory, city or area or of its authorities, or concerning the delimitation of its frontiers or boundaries. This map has been provided by the authors.

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