

## Congenital toxoplasmosis in the state of Minas Gerais, Brazil: a neglected infectious disease?

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### SUMMARY

This study aimed to investigate the distribution of congenital toxoplasmosis in the state of Minas Gerais, Southeastern Brazil and describe the demographic and socioeconomic profile of the municipalities associated with the disease. An ecological study was conducted using socioeconomic indicators of a database (MGSSRI) created by Fundação João Pinheiro (a government technical support agency of Minas Gerais), in order to show the development of the municipalities in the state. The prevalence of toxoplasmosis was the outcome and the items of the MGSSRI were the explanatory variables. Of 146 307 newborns screened (November 2006 to May 2007), 190 had congenital toxoplasmosis, yielding a prevalence of 1·3/1000, ranging from 0 to 76·9/1000 in the municipalities. The multivariate model indicated a higher occurrence of toxoplasmosis in municipalities with smaller populations and worse indexes of tax performance. Congenital toxoplasmosis appears to be a neglected disease in the state of Minas Gerais, given the high prevalence found and its concentration in municipalities with worse socioeconomic indexes.

**Key words:** Infectious disease epidemiology, *Toxoplasma gondii*.

### INTRODUCTION

*Toxoplasma gondii* is a worldwide protozoan that causes benign and self-limited disease in immuno-

competent individuals infected after birth, but severe complications in immunocompromised individuals or in cases of congenital infection. The infected newborns can manifest a wide range of symptoms depending on the gestational age, parasite load, parasite strain, and immunological status of both mother and fetus [1].

The prevalence of congenital toxoplasmosis (CT) in humans ranges across regions and according to a number of factors, such as: number of susceptible pregnant women, parasite presence and persistence in the environment, cultural habits, food preparation

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and cooking practices, as well as sanitary practices [2]. In Europe, high prevalence of the disease has been usually associated with the ingestion of raw or undercooked meat. In Central America and other developing countries, including Brazil, infection is usually associated with ingestion of oocysts and particularly affects young, poorly educated people living under low socioeconomic conditions. However, low levels of education and low socioeconomic conditions are presumably inter-related and possibly associated with other risk factors [3].

It is well known that determinants based on individual measurements are insufficient to explain the distribution of diseases affecting populations [4]. This implies that it is necessary to analyse a wide range of risk factors associated with toxoplasmosis, including biological specificities, lifestyles, as well as socioeconomic and demographic characteristics of the surrounding environment. Ecological studies make use of aggregated data, assuming a given geographical region as the smallest unit of analysis. This type of study can reveal highly predictive demographic, socioeconomic and environmental risk factors associated with the diseases, as people within a community tend to gather together in a systematic way and be influenced by their local environment [5]. With regard to toxoplasmosis, the sources of infection are numerous and range according to region. In such cases, ecological studies can be very informative.

In November 2006 a multidisciplinary research group (UFMG Congenital Toxoplasmosis Brazilian Group) started a survey named 'Newborn Screening of Congenital Toxoplasmosis in the State of Minas Gerais'. Under the coordination of one of the authors of this paper (G.M.Q.-A.), the research group found high prevalence of the disease in the state [6]. This result motivated the present study, aimed at assessing the distribution of CT in the state of Minas Gerais, and describing the demographic and socioeconomic profiles of the municipalities associated with the occurrence of the disease.

## METHODS

### Design

This is an ecological study using the prevalence of CT in municipalities in the state of Minas Gerais as an outcome variable and municipal demographic and socioeconomic characteristics as explanatory variables.

### Study setting

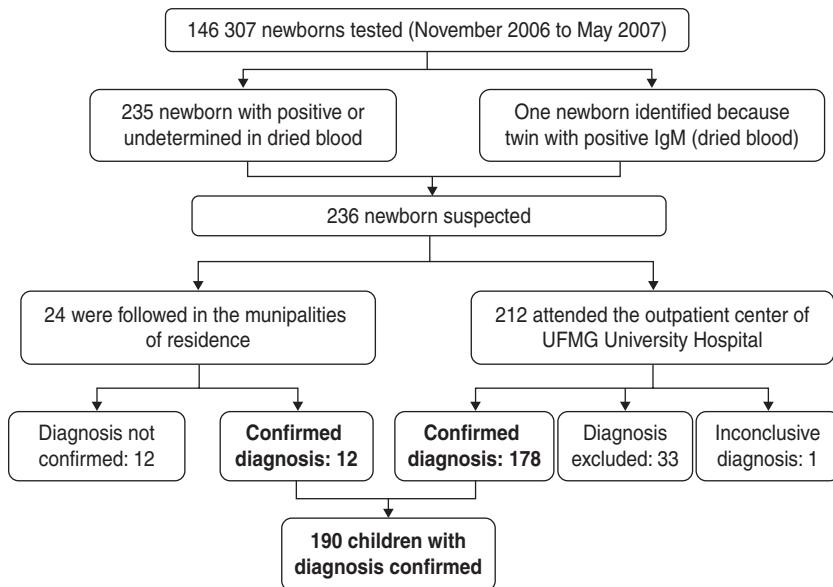
Minas Gerais is a southeastern state in Brazil, comprising 853 municipalities with a total of 19 497 330 inhabitants spread over an area of 586 520 km<sup>2</sup> [7]. According to a very recent review about the burden of toxoplasmosis in Brazil [8], the most accurate estimate on CT prevalence was provided by a study conducted in Minas Gerais by our group. A total of 146 307 live newborns were screened from 1 November 2006 to 31 May 2007, within the scope of the Minas Gerais State Programme of Newborn Screening (PETN-MG). CT was confirmed in 190 children corresponding to a prevalence of 1·3/1000 live births [6]. At the time of the study, this programme regularly provided free testing for four diseases (i.e. congenital hypothyroidism, sickle cell disease, phenylketonuria, cystic fibrosis) and covered about 95% of the live newborns in the state. The programme is coordinated by the Center for Newborn Screening and Genetic Diagnosis, a research centre of the School of Medicine at the Federal University of Minas Gerais (UFMG; Portuguese acronym) [9].

### Diagnosis of CT

Under parental consent all live newborn infants participating in PETN-MG in the period under scrutiny had dried blood samples collected on filter paper for analysis of anti-*T. gondii* IgM (Toxo IgM Q-Preven®, Symbiosis, Brazil). Infants with positive or undetermined results were tested with anti-*T. gondii* IgA (enzyme-linked immunosorbent assay) and with anti-*T. gondii* IgG and IgM (enzyme-linked fluorometric assay, bioMérieux SA, France), and their results were further compared with their mothers. The infants were also followed up in the outpatient units of UFMG University Hospital until diagnosis confirmation. The data relating to infants followed up in their home municipalities (7%) were later forwarded from the clinics to the research group.

The criteria for confirmed CT were: (1) positive anti-*T. gondii* IgM and/or IgA and positive IgG up to age 6 months; (2) negative anti-*T. gondii* IgM/IgA and positive IgG associated with retinochoroidal lesions within the first 3 months of life; (3) persistence of positive anti-*T. gondii* IgG results up to age 12 months.

All confirmed cases of CT were included in the study (Fig. 1). The UFMG Review Board Committee gave approval for the study (ETIC 510/07).



**Fig. 1.** Flowchart of the serological survey of congenital toxoplasmosis conducted in the state of Minas Gerais, Brazil.

### Demographic and socioeconomic profile of the municipalities

Minas Gerais, the fourth largest state in Brazil, has the third largest gross domestic product (GDP) of the country, but is hampered by stressed socioeconomic and geographical heterogeneity [10].

In order to verify the possible association of CT in Minas Gerais with socioeconomic indicators, a preliminary analysis was performed using the human development index (HDI). We found that toxoplasmosis was more prevalent in municipalities with the worst performance in global HDI and its components – life expectancy, literacy and educational attainment, and *per capita* GDP.

Based on these findings, we opted to assess the vulnerability to CT in the socioeconomic sphere of municipalities of Minas Gerais, using indicators built on the 2006 software package Minas Gerais State Index of Social Responsibility (MGSSRI) [11], which consists of a database developed by the Centre for Public Policy Studies at Fundação João Pinheiro with a view to depicting the level of development of the municipalities in the state. The software package includes 47 social, administration and health indicators of all 853 municipalities in the state. These indicators were converted into indexes, considering the standards of weights and references, resulting in values from 0 to 1, where they represent, respectively, the worst and best situation of the municipality in relation to the indicator. The indexes were grouped

in seven dimensions that constitute the synthetic global index (global MGSSRI): healthcare; educational attainment; housing and environment; employment and income; culture, sports and leisure; public security; and financial performance. The issues selected to constitute the dimensions represent both municipality conditions and public administration [11].

The demographic data correspond to the census carried out by the Brazilian Institute of Geography and Statistics (IBGE) in 2000 [7].

### Statistical analysis

To analyse the correlation between prevalence of CT and municipality performance as measured through the MGSSRI indexes, Spearman's correlation coefficient was used for those municipalities with at least one case of the disease. As the response variable, prevalence of CT, represents a count in a given time-frame, Poisson's regression was regarded as the most adequate model [12]. Nevertheless, because 712/853 (83.5%) municipalities registered no cases of the disease in the period under scrutiny, there was over-dispersion that violated the assumptions of the Poisson model. Some developments of the Poisson model have been suggested for such circumstances, such as the negative binomial model or the zero-inflated Poisson model [13]. The latter is the most adequate for the present study and has been widely reported in the literature as effective in accounting for the distribution of additional zeros [12, 13].

The theory suggests that the excess zeros are generated by a separate process from the count values and that the excess zeros can be modelled independently. Thus, the zero-inflated Poisson (ZIP) model has two parts, a Poisson count model and the logit model for predicting excess zeros. Therefore, in fitting the ZIP model, two separate models were generated and then combined. First, a logit model was generated for municipalities with zero prevalence rates, predicting whether or not a municipality would be in this group. Then, a Poisson model was generated to predict the counts for those municipalities with no zero prevalence rates. Finally, the two models were combined. This model is performed by the command 'ZIP' of the statistical package Stata v. 10.0 [14].

Therefore, the ZIP model was used as a means of including all municipalities for univariate and multivariate analyses. For multivariate analysis, we included variables with  $P \leq 0.20$  in the univariate analysis, adopting a model in blocks, with the construction of models for the items of each of the seven dimensions separately. Significant variables were entered one by one, according to the 'forward' criterion. After adjusting the seven models, a multivariate analysis was performed with new input variables in a hierarchical way. The order in which groups were defined by the significance of each dimension on the MGSSRI in the univariate analysis were: (1) culture, sports, and leisure; (2) public security; (3) employment and income; (4) housing and environment; (5) financial performance of municipalities; (6) educational attainment; and (7) health.

The final model was adjusted according to municipality population, given the strong association of this variable with the response variable and the other explanatory variables. The final model included only the variables significant at  $P \leq 0.05$ , and quality of fit was assessed through deviance [12].

The software package MapInfo 8.5 was used to build a choropleth map with the prevalence distribution of CT over the state of Minas Gerais. Both univariate and multivariate analyses were run by statistical software package Stata v. 10.0.

## RESULTS

### Toxoplasmosis distribution over the state of Minas Gerais

The overall prevalence of CT in Minas Gerais state of 1.3/1000, ranged from 0.78 to 2.77 by 13 macro-region

divisions (Fig. 2) and from 0 to 76.9/1000 according to the 853 municipalities. Figure 3a shows the distribution of CT in the municipalities of Minas Gerais according to quartiles of prevalence, and Figure 3b shows the performance of municipalities in quartiles of MGSSRI. When stratifying the municipalities according to the number of inhabitants, it was observed that the prevalence increased as the population declined, reaching 1.9/1000 live births in municipalities with <20000 inhabitants (Table 1).

### Correlation of prevalence of CT with demographic and socioeconomic characteristics of the municipalities

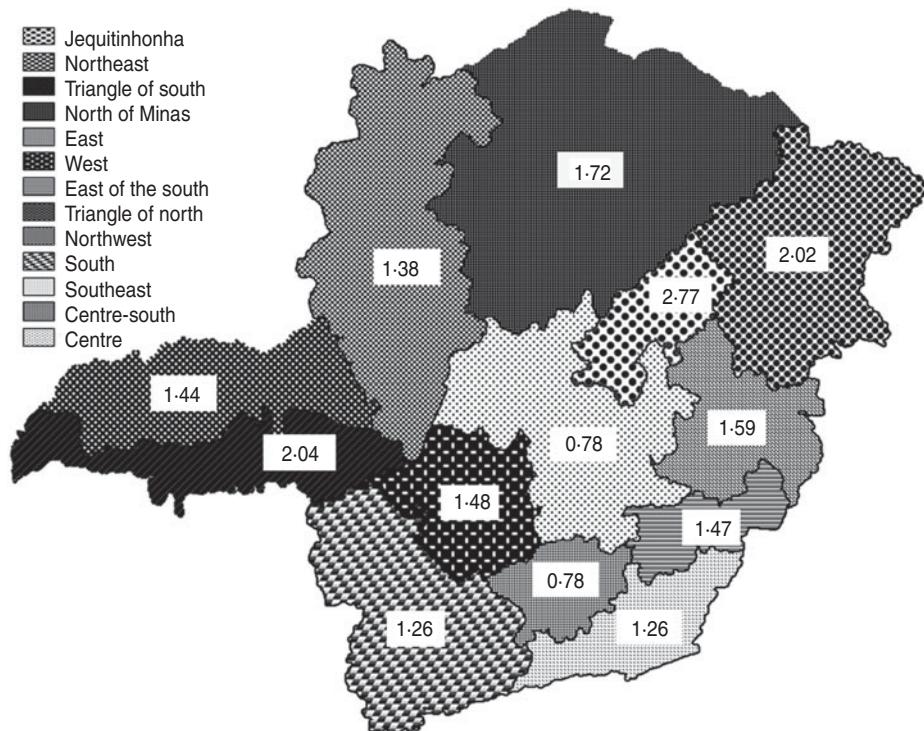
The prevalence of CT correlated inversely with the global MGSSRI, i.e. highest rates were found in the municipalities with the worst performance. Concerning the dimensions that compound the global MGSSRI, the correlation was direct only for 'public security', i.e. highest rates of CT were found in the municipalities with better performance. An inverse correlation was found for others dimensions such as: 'educational attainment', 'employment and income', 'housing and environment', 'culture, sports and leisure', and 'financial performance of municipalities'. No correlation was found for the health dimension. Considering the indexes of dimensions, several correlations were observed with prevalence of toxoplasmosis in the municipalities (Table 2).

A multivariate analysis performed to model the nature of the several correlations observed, resulted in a final model with nine parameters (Table 3). The parsimonious final model showed a good fitness with a deviance test ( $P=0.999$ ).

To assess the correlation between the variable municipality population and the others that remained in the final model, the municipalities were categorized in four strata according to the number of inhabitants. The strata with the more populous municipalities tended to have better mean values for the indexes 'tax performance', but worse mean values for the indexes 'violent crimes' (data not shown).

## DISCUSSION

The prevalence of CT in the state of Minas Gerais was high, and was higher in municipalities with worse indexes of economic development. Despite several ecological studies showing associations between infectious disease and local vulnerability indexes [15, 16], no study has, to the best of our knowledge,



**Fig. 2.** Map showing the prevalence of congenital toxoplasmosis (per 1000 live births), determined from November 2006 to May 2007 by screening of newborn infants, over the macro-regions of Minas Gerais, Brazil.

investigated the association between prevalence of CT and socioeconomic indexes at an ecological level.

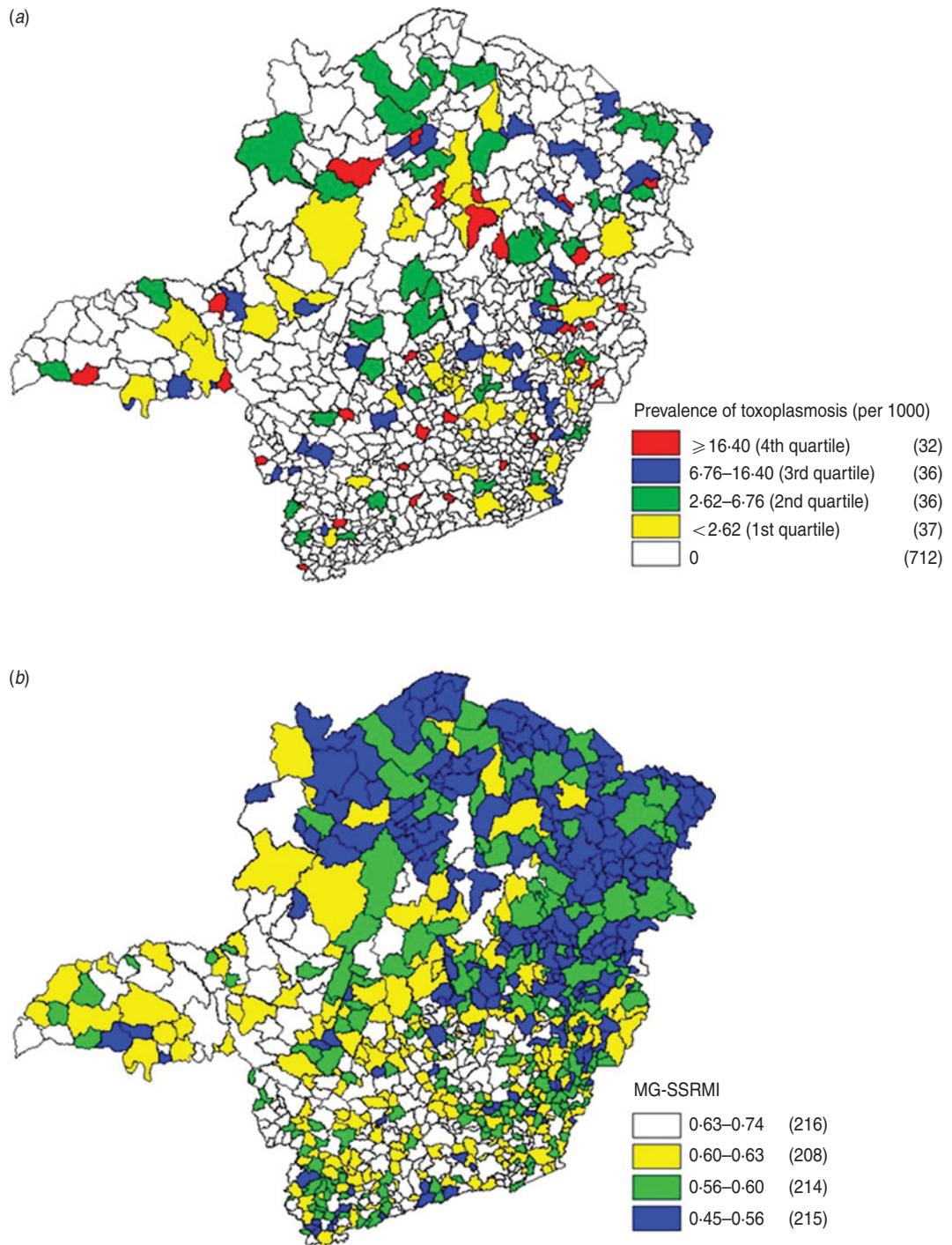
The variables associated with the prevalence of congenital disease that remained in the final model can be analysed from two perspectives. First, from the perspective of the association between the indexes and the parasite transmission cycle, as evident in the municipality variables. The percentage of population affected with disease was related to poor sanitation. Second, from the perspective of the indexes that reflect the socioeconomic disadvantages of the municipalities and inequalities of access to healthcare services. However, inter-relationship should be assumed in all variables.

Poor housing and sanitation conditions expose the population to several infectious diseases, including toxoplasmosis [3]. In Natal, capital of the state of Rio Grande do Norte, Brazil, a study conducted to assess risk factors associated with toxoplasmosis in students at public and private schools from ages 5–21 years revealed a higher rate of seropositivity in those living in areas without access to treated water and sewage [17]. In the present study, the univariate analysis indicated higher prevalence of CT in municipalities with poorer access to treated water, sewage and garbage collection services. The multivariate

analysis indicated higher prevalence of CT in the municipalities with a lower budgetary commitment to environment, sanitation and housing, which suggests low investment in this area.

Notwithstanding that toxoplasmosis is a disease able to be transmitted through ingestion of contaminated water, and whose transmission is facilitated by poor hygiene [3], the present study highlighted its highest prevalence in municipalities with a better performance in the index ‘percentage of population affected with diseases related to poor sanitation’. This indicator measures the percentage of hospitalizations due to these diseases [11]. People who live in regions with poorer sanitary infrastructure coexist endemically with several infectious agents, but do not necessarily require hospitalization. This is a possible explanation for the association observed in this study.

Another intriguing result is the higher prevalence of CT in municipalities with smaller native vegetation coverage. A plausible explanation for this association may be that deforestation and demographic expansion could have caused climate change, ecosystem disturbance and eventually promoted adequate conditions for parasite survival and dissemination, as described previously [18]. Although plausible, it is far from the



**Fig. 3.** Choropleth map showing (a) the prevalence distribution of congenital toxoplasmosis over the 853 municipalities in the state of Minas Gerais, Brazil, from November 2006 to May 2007 and (b) the global indexes of the municipalities.

scope of this work. We understand that this hypothesis should be explored in other studies with designs better suited to answer this question.

We found a higher prevalence of toxoplasmosis in municipalities with poor access to prenatal care and delivery assistance, which once again suggests negli-

gence in providing basic health services to the population. This finding is especially relevant, because prenatal care is a valuable opportunity for educative actions aimed at the prevention of health problems both to the pregnant women and their fetus. The literature has strong evidence that providing adequate

Table 1. Prevalence of congenital toxoplasmosis (CT) in the state of Minas Gerais from November 2006 to May 2007 stratified according to municipality population as IBGE census, reference year 2000

Municipality population	Municipalities of the state (%)	Municipalities with CT cases (%)	CT prevalence/1000 live newborns tested (95% CI)	Tendency $\chi^2$ (P)
$\leq 20000$	687 (80.5)	70 (10.2)	1.91 (1.48–3.39)	28.1 (<0.001)
20001–100000	143 (16.8)	51 (35.7)	1.42 (1.08–2.51)	
100001–500000	20 (2.3)	17 (85)	0.95 (0.61–1.56)	
>500000	3 (0.4)	3 (100)	0.49 (0.21–0.70)	
Total	853	141 (16.5)	1.30 (1.11–2.41)	

CI, Confidence interval.

guidance to pregnant women is a crucial prophylactic measure against CT [19, 20].

To understand the finding of higher prevalence of CT in municipalities with larger coverage in the Family Health Programme, it might be necessary to understand the implementation background of this programme in Brazil. The programme began in 1994, financing preferentially the small municipalities with lower income and higher epidemiological and social risk. The expansion of the programme to municipalities with >100 inhabitants started in 2003, and the programme is still developing in large urban centres [21]. Consequently, the higher prevalence of CT in municipalities with larger coverage of the Family Health Programme – which has historically catered for the more vulnerable groups – may be the reflection of the reverse causality underlying the cross-sectional nature of this study design.

With regard to the indexes related to the economic conditions of the municipalities, the univariate analysis indicated higher prevalence of the disease in those municipalities with lower average income, lower formal employment rates, and lower GDP. The final model pointed to higher prevalence of the disease in the municipalities with worse tax performance. This index is calculated by the sum of municipal revenue (rates, taxes, and contributions for improvements) with the value-added tax on sales and services transferred to municipalities by the state using two criteria: (1) proactive criteria – actions directed to public policies in education, health, environment, etc., and (2) reactive criteria – minimum quota, regardless of the proactive criteria. The higher the index the greater is the municipality development and public administrative capacity to manage its financial activities and revenues [11]. The higher prevalence of CT in municipalities with worse tax performance corroborates the results of other studies that also found an association

– at the individual level – between toxoplasmosis seropositivity and poor socioeconomic conditions [3, 17, 22]. For instance, in the municipality of Londrina, state of Paraná, a study of 489 pregnant women showed that women with *T. gondii* antibodies were usually from the lower income group [22]. Bahia-Oliveira *et al.* found that the best predictor for toxoplasma seropositivity in the 1436 people tested was ‘worst socioeconomic situation’, even after adjustment for participants’ age [3].

Total municipality expenses *per capita* correlated with prevalence of toxoplasmosis positively. This result attracted attention because the biggest expense observed was not accompanied by satisfactory results neither in those indexes concerning the quality of the services provided in the municipalities nor in the global MGSSRI index. Studies conducted to assess municipality administration of public resources have proved that efficient administration is not necessarily associated with large expense, but the adequate use of resources [23–25]. Similar to the situation in the state of Minas Gerais, Rezende *et al.* did not find direct correlation between investment and HDI in the 503 municipalities they assessed in the state of São Paulo [23].

Initially, the finding of higher prevalence of CT in municipalities with better performance in the index of violent crimes contrasts with the above-mentioned results, as the literature has consistently reported the association with criminality, violence, social disorganization, and poverty [26–28]. However, in relation to criminality, it is known that, social inequality contributes more than poverty to the growth of criminality in a community. Therefore, even though we found a higher prevalence of CT in smaller municipalities with lower levels of human development, they also had less inequality than the large urban centres, and consequently lower rates of criminality. Indeed, the

Table 2. *Univariate analysis to assess the correlation between prevalence of congenital toxoplasmosis in the state of Minas Gerais from November 2006 to May 2007 and the socioeconomic and demographic characteristics of the municipalities*

Minas Gerais state index of social responsibility (MGSSRI)*	Coefficient	P value
Global MGSSRI	-5.84	<0.01
Dimension – Health †	0.07	0.79
Index: Standardized mortality rate	-0.39	<0.01
Index: Access to delivery assistance	-1.80	<0.01
Index: % of live newborns' mothers who attended $\geq 7$ prenatal visits	-0.33	0.01
Index: % of death reports with poorly defined causes	-0.58	<0.01
Index: Population coverage in the Family Health Programme	1.31	<0.01
Dimension: Educational Attainment‡	-2.39	<0.01
Index: Rate of functional illiteracy of people aged <15 years	-4.43	<0.01
Index: Net rate of children aged 15–17 years attending secondary schools	-1.95	<0.01
Dimension: Employment and income‡	-3.43	<0.01
Index: Home consumption of electricity ( <i>per capita</i> )	-1.80	<0.01
Index: Average income in the formal economy	-5.79	<0.01
Index: Employment rate in the formal economy	-1.99	<0.01
Index: Gross domestic product ( <i>per capita</i> )	-2.34	<0.01
Index: Total municipality expenses ( <i>per capita</i> )	3.27	<0.01
Dimension: Public security‡	4.90	<0.01
Index: Violent crimes	7.01	<0.01
Index: Number of inhabitants per number of police officers	-0.19	0.04
Dimension: Housing and environment‡	-3.75	<0.01
Index: % of population with access to potable water and bathroom	-0.78	<0.01
Index: Population with access sewage treatment	-1.07	<0.01
Index: Population with access to household waste collection	-0.44	<0.01
Index: % of population affected with disease related to poor sanitation	0.88	0.04
Index: % of native vegetation coverage	-0.42	<0.01
Index: % of restored vegetation coverage	-2.31	<0.01
Index: Full environmental protection area	-0.82	<0.01
Index: Budgetary commitment to environment, sanitation, and housing	-0.82	<0.01
Dimension: Culture, sports and leisure‡	-2.49	<0.01
Index: Cultural heritage preservation and management	-1.80	<0.01
Index: Budgetary commitment to culture, heritage, and sports	0.64	<0.01
Index: Diversity of cultural facilities (except for libraries)	-0.92	<0.01
Index: Existence of a concert/marching band	-0.70	<0.01
Index: Existence of at least one sports facility	-0.79	<0.01
Dimension: Municipality management‡	-3.94	<0.01
Index: Tax performance	-3.83	<0.01
Index: Net income <i>per capita</i>	2.60	<0.01
Index: % of expenses in the legislative body (Emend. No. 25/2000)	53.41	<0.01
Index: Defrayment of money to sustain the machinery of government in relation to the municipality net income	-0.88	<0.01
% of population living in urban areas	-0.020	<0.01
Population density (total number of residents/total municipality area)	-0.0016	<0.01
Number of people living in the municipality (IBGE Census as of 2000)‡	-0.00003	<0.01

\* Indexes vary from 0 to 1, with closeness to 1 indicating the better situation of the municipality in relation to the indicator.

† The following indexes were not statistically significant: Coverage of tetravalent vaccine in infants aged <1 year; net rate of children aged 7–14 years attending primary schools; education quality index; investment efforts; area for sustainable use; expenses *per capita* on environment, sanitation and housing in the municipality; rate of public debt; and percentage of expenses on public servants.

‡ IBGE (Brazilian Institute of Geography and Statistics) (<http://www.ibge.gov.br/home/estatistica/populacao/censo2000/defaulttabmunic.shtml>).

Table 3. Description of variables associated with prevalence of congenital toxoplasmosis in the municipalities of Minas Gerais from November 2006 to May 2006 that remained in the final model after the multivariate analysis

Variables*	Coefficient	P value
Non-inflated part		
Municipality management: tax performance*	-2.31	<0.01
Housing and environment: % of population affected with disease related to poor sanitation*†	1.54	<0.01
Housing and environment: budgetary commitment to environment, sanitation, and housing*	-0.22	0.03
Housing and environment: % of native flora*	-0.43	<0.01
Income: total municipality expenses ( <i>per capita</i> )*	3.18	<0.01
Health: population coverage in the Family Health Programme*	0.25	0.01
Public security: violent crimes*‡	1.03	0.03
Number of people living in the municipality (IBGE Census, 2000)	-0.00001	<0.01
Inflated part		
Number of people living in the municipality (IBGE Census, 2000)	-0.00004	<0.01

\* Indexes vary from 0 to 1, with 0 being the worst and 1 being the best situation of the municipality concerning the indicator.

† The higher the index, the lower the percentage of population affected with disease related to poor sanitation.

‡ The higher the index, the lower the rate of violent crimes.

P value in deviance test = 0.998.

variable 'violent crimes' in the univariate analysis showed a coefficient of 7.01, which fell to 1.04 after multivariate analysis adjustment that included population size. This finding suggests the presence of a residual confounding because the population size does not cover the extent of poverty and inequalities in cities. An ecological study performed to identify the determinants of criminality in the municipalities in the state of Minas Gerais indicated higher rates of violent crimes in the most populous municipalities with better human development indexes, probably because of social inequality [29].

Several studies have shown an association between toxoplasmosis and mothers' level of education at the individual level [17, 30, 31]. In the state of Minas Gerais higher prevalence of CT was found in the municipalities with worst performance in the educational attainment dimension and some of its indexes. Nevertheless, this and other important epidemiological variables were not influential in the multivariate model.

In 2008 Brazil hosted a world meeting to celebrate the 100th anniversary of the discovery of *T. gondii*. Parallel to the advances in biological, genetical and immunological research, have been the highlighting of gaps in the areas of prevention and treatment of CT, which has been systematically underestimated in relation to the disability-adjusted life years (DALYs), and which should be included in the list of neglected diseases. From this conference a letter was drafted by a group of professionals dedicated to the study of disease, containing a proposal for the

control of toxoplasmosis in Brazil and suggesting the inclusion of toxoplasmosis in the list of neglected diseases in that country [32].

The World Health Organization (WHO) classifies diseases as neglected when they are prevalent in unprivileged people who have little political expression and priority in public health actions [33]. In September 2000 the United Nations members agreed on the Millennium Development Goals (MDG) Declaration, which gave a commitment to have extreme poverty eradicated by 2015 [34]. Following suit, the Pan American Health Organization Board issued a resolution in 2009 extending this commitment specifically to the eradication of neglected diseases [35]. One decade after the MDG declaration, a preliminary report on neglected tropical diseases estimated that 149 countries are endemically affected with such diseases. Currently, there is an international alliance (Global Network for Neglected Tropical Diseases) and a specific department within the WHO that is specifically engaged in solving the problems related to such diseases [36, 37].

The WHO and the Doctors without Borders have recently proposed a new classification categorizing the neglected tropical diseases as 'global' when they affect the world as whole; 'neglected' when they are more prevalent in developing countries, and 'more neglected' when they are exclusive of developing countries [33]. The neglected diseases are more common in underprivileged groups and affect children's growth, undermining their productive capacity and maintaining the cycle of both poverty and

infection [38]. Hotez and colleagues have published a series of papers analysing the geographical distribution of the major neglected infectious diseases worldwide and emphasized the economic burden of such diseases in the countries where they are prevalent [38–40]. Of such diseases, these authors particularly highlighted the congenital infections because of their potential to lead to cognitive, hearing and ophthalmological disorders in the long term. CT was particularly found to prevail in the poorest and most segregated groups living in regions of the Mississippi Delta, South America, Mexico–USA border and Central and Eastern Europe [38, 40].

According to the results reported herein, CT is a neglected disease in the state of Minas Gerais, as the prevalence of the infection is high and usually affects municipalities with poorer MGSSRI indexes, i.e. municipalities with higher social and economic vulnerability. We understand that the findings cannot be derived at the individual level, but point to the need of testing further hypothesis taking into account both individual and contextual hierarchical levels.

Some limitations of the present study should be noted. The first refers to the unit of analysis. Although the municipalities in general had some characteristics associated with more or less occurrence of toxoplasmosis, they do have internal heterogeneity that could not be captured in this study given its ecological design. However, this type of study is valid and crucial to assess the impact of environmental, demographic and socioeconomic determinants on how the population is likely to become ill [4].

Moreover, the database used for characterizing the municipalities, the MGSSRI index, was not originally created to measure risk factors related to the occurrence of any disease, infectious or not. The process of determining the indexes involved a number of management records of city halls with different levels of organization and therefore subject to inaccuracy [11]. Although acknowledging this limitation in the use of a public database, the MGSSRI index was used with the intent of approaching, as much as possible, the determinants under scrutiny with the actual conditions of the municipalities in the state of Minas Gerais. The year 2006 was chosen because that was the year when the serological survey took place, and the 2000 Census was chosen because it was the only demographic data available at the time of data analysis.

Despite the limitations outlined herein, the study shows the highest prevalence of CT in the municipalities for which the MGSSRI index attests the worst life

conditions. This suggests that policies should aim at human and socioeconomic development as a means of reducing the prevalence of the disease. The results also point to the need of broadening the surveillance system for this disease and to the importance of further studies focusing on both individual and contextual risk factors associated with the distribution of CT in the state of Minas Gerais.

## APPENDIX

### UFMG Congenital Toxoplasmosis Brazilian Group (UFMGCTBG)

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## DECLARATION OF INTEREST

None.

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