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Conditional Cash Transfer Program and Leprosy Incidence: Analysis of 12.9 Million Families from The 100 Million Brazilian Cohort

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Abstract

Leprosy is a neglected tropical disease predominately affecting poor and marginalized populations. To test the hypothesis that poverty-alleviating policies may be associated with reduced leprosy incidence, we evaluated the association between the Brazilian Bolsa Familia conditional cash transfer Program (BFP) and new leprosy case detection using linked records from 12,949,730 families in the 100 Million Brazilian Cohort (2007-2014). After propensity score matching BFP beneficiary to non-beneficiary families, we used Mantel-Haenszel tests and Poisson regressions to estimate incidence rate ratios (IRRs) for new leprosy case detection and secondary endpoints related to operational classification and leprosy-associated disabilities at diagnosis. Overall, cumulative leprosy incidence was 17.4/100,000 pyr (95%CI 17.1-17.7), and markediy ingher in "priority" (high-burden) versus "non-priority" (low-burden) municipalities (22.8/100,000 pyr, 95%CI 22.2-23.3 versus 14.3/100,000, 95%CI 14.0-14.7). After matching, BFP participation was not associated with leprosy incidence when restricted to families living in high-burden municipalities (IRR_{Poisson} 0.86, 95%CI 0.77-0.96). In high-burden municipalities, the association was particularly pronounced for paucibacillary cases (IRR_{Poisson} 0.82, 95%CI 0.68-0.98) and cases with leprosy-associated disabilities (IRR_{Poisson} 0.79, 95%CI 0.65-0.97). These findings provide policy-relevant evidence that social policies may contribute to on-going leprosy control efforts in high-burden communities.

Keywords: Bolsa Familia Program, cash transfers, poverty, inequality, infectious diseases, Hansen's disease.

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Abbreviations:

- CCT conditional cash-transfers
- BFP Bolsa Familia Program

IRR - incidence rate ratio

CadUnico – Brazilian National Registry for Social Programs Cadastro Único

SINAN - Brazilian Notifiable Disease Registry

- BRL Brazilian reals
- PS propensity score

PB - paucibacillary

MB - multibacillary

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Leprosy is a neglected tropical disease (NTD), which can lead to blindness and permanent disabilities if left untreated. While the prevalence of leprosy has declined over the last 30 years, leprosy continues to be an important cause of disability and stigma among the over 200,000 individuals diagnosed annually worldwide[1, 2]. There is an increasing recognition that leprosy and other neglected tropical diseases are strongly linked to poverty, being both attributable to and responsible for unfavorable economic conditions in affected populations[3-5].

Conditional cash transfer programs (CCTs) have been proposed as a promising cost-effective strategy for overcoming intergenerational poverty and ameliorating the social determinants of health[6]. However, there is limited evidence of their impact on neglected tropical diseases[7]. The Brazilian CCT, the Bolsa Familia Program (BFP), provides financial aid to low-income families, conditional on school attendance and preventive health check-ups, and has been linked to improvements in children's education, healthcare access, and food security[8-12]. Although leprosy in Brazil has been declining in the past decades, Brazil still register over 20 thousand new leprosy cases annually, accounting for over 14% of cases diagnosed globally[13]. Higher conditional cash transfer coverage has been associated with reductions in leprosy risk at the population level[14, 15]. However, no studies to date, have provided a robust assessment of the impact of BFP or any CCTs on the burden of leprosy using individual-level data. [11]

To address this gap, we tested the hypothesis that receiving BFP can reduce leprosy incidence, using prospective data that was routinely collected from families enrolled in the Brazilian National Registry for Social Programs (CadUnico), the BFP Payroll Database, and the Brazilian Notifiable Disease Registry (SINAN), and linked as part of the 100 Million Brazilian Cohort.

METHODS

Intervention

The BFP targets families registered in CadUnico who live in: i) extreme poverty (i.e., earning ≤ 60 Brazilian Real (BRL) per capita/month in 2007-2008 and ≤ 70 BRL per capita/month in 2009-2014) or ii) poverty (i.e., ≤ 120 BRL per capita/month in 2007-2008 and ≤ 140 BRL per capita/month in 2009-2014) with ≥ 1 child (i.e., <18 years old)

and/or with a woman who is pregnant or breastfeeding (Web Appendix 1, Web Table 1)[8] (1 BRL=approximately 0.25 USD). The BFP provides monthly payments to families conditional on compliance with i) children's attendance for \geq 80% of school days, ii) health monitoring of children \leq 6 years and breastfeeding women, and iii) prenatal care (See Web Appendix 1 for further details).

Data sources and linkage

The 100 Million Brazilian Cohort is a large-scale linked cohort that aim to evaluate the impact of the BFP and other social programs on health outcomes in Brazil[16]. For the current investigation, we linked the baseline of the 100 Million Brazilian Cohort, the BFP Payroll Database (2004-2015), and SINAN (2007-2014)[17] (See Web Appendix 2 for linkage details).

The 100 Million Brazilian Cohort baseline covariates comprised those from the first registry of families in CadUnico: sociodemographic variables (i.e., sex, age, self-identified race/ethnicity, education, and work) for the head of family (i.e., oldest member), the State and area of residence (urban vs rural), household living conditions (i.e., house ownership, housing material, water supply, electricity, sewage, and waste collection), per capita income, and individual-level identifiers for linkage (i.e., Social Identification Number (NIS), name, date of birth, sex, maternal name, and municipality). Exposure data extracted from the BFP Payroll Database included starting and end dates of BFP benefit receipt for each primary recipient per family, and the individual-level identifier for linkage (i.e., NIS). Outcome data extracted from SINAN included: date of leprosy diagnosis, clinical presentation (i.e., paucibacillary (PB): \leq 5 lesions; or multibacillary (MB): >5 lesions or positive slit skin smear), and disabilities at diagnosis (i.e., Grade 0 if no disabilities or Grade 1/2 with any sign of eye problems, visible deformity, damage or anesthesia in hands and feet)[2], and individual-level identifiers for linkage (i.e., name, date of birth, sex, maternal name, and municipality).

The 100 Million Brazilian Cohort baseline and BFP datasets were deterministically linked using a unique identifier (i.e., NIS). The cohort baseline and SINAN datasets were linked by the five individual-level identifiers in two steps using the CIDACS-RL tool (https://gitHub.com/gcgbarbosa/cidacs-rl)[16]. In the first step, entries were deterministically linked. In the second step, entries that were not linked deterministically were then linked based on a similarity score between all the pairwise comparisons (i.e., ranging from 0-1); entries with the highest similarity scores were considered to be linked pairs.

To assess the accuracy of the linkage procedures, we performed manual validation study. 10,000 pairs were randomly selected from all possible paired links. Manual verification was used to classify pairs as true or false links. Various cut-offs of the similarity score were used to declare pairs to be a link. These linkages were compared to the true link status to determine the sensitivity and specificity at each potential cut-off. The cut-off corresponding to the optimal sensitivity and specificity was chosen (a similarity score of 0.92) to determine links for this study (Web Appendix 2)[18]. Following linkage, individual identifiers were removed from the dataset.

Selection of the study population

The study population included individuals belonging to families who enrolled in the 100 Million Brazilian Cohort between January 2007 and December 2014. We excluded families who: (i) lacked at least one individual over 15 years of age at enrollment (i.e., children recorded separately from an adult caregiver were not considered to be a family), (ii) had a monthly per capita income exceeding 5000 BRL, and/or (iii) started receiving BFP benefits prior to enrollment. We defined as BFP beneficiary families those that started receiving BFP benefits within 6 months after enrollment in the cohort (i.e., reflecting the typical time-to-receipt for families who would eventually become beneficiaries) and non-beneficiary families (i.e., non-BFP) those that did not start receiving the benefit within 6 months after enrollment (Web Figure 1). We analyzed the overall sample and then stratified our study population by whether or not families resided in one of the 182 "priority" municipalities in Brazil as officially designated due to their high burden of leprosy (Figure 1)[19].

Statistical analysis

Propensity score matching

We used propensity score (PS) matching to compare BFP beneficiary (i.e., exposed) and non-beneficiary (i.e., unexposed) families. We estimated the PS by multiple logistic regression using baseline sociodemographic characteristics and year of application for each dataset (i.e., overall sample, high-burden and low-burden municipalities) (Web Figure 1). Missingness in PS covariates was considered as a category. We performed 1:1 nearest-neighbor matching with a caliper of 0.05, allowing a same non-beneficiary family to match with more than one beneficiary family (i.e., matching with replacement)[20]. We compared the difference in the distribution of PS covariates between beneficiary and non-beneficiary families using the standardized mean difference to

assess balance of potential confounders before and after matching (standardized mean difference>0.1 was taken to indicate potential confounding by that characteristic)[21](See Web Appendix 3 for matching details).

Primary and secondary outcome analysis

Incident cases were defined as the first newly detected case of leprosy occurring within family units after enrollment. Secondary endpoints for leprosy incidence included operational classification (i.e., PB and MB) and the presence of disabilities at diagnosis (i.e., Grade 0 and Grade 1/2). Families with a leprosy case diagnosed prior to or within the first 6 months after enrollment were not considered disease-free at baseline and were therefore excluded from the analyses. For family units with more than one case occurring during the study period, only the first case was considered in the analysis. Family-years at risk (fyr) began 6 months after enrollment (i.e. the time at which exposure status was determined) and ended on 31 December 2014 or at diagnosis of the first new leprosy case in the family. The total person-years at risk (pyr) for each family was defined as the contribution of each fyr multiplied by the number of individuals in the family. Unexposed families who later participated in BFP were censored at the time they started receiving BFP benefits. As the potential benefits of BFP participation (e.g., via behavior changes associated with the conditionalities) could persist after families stopped receiving the cash transfer benefit itself, **BFP** exposed families remained in the exposed group during the full study period. For analyses of secondary endpoints, families were censored at the first new leprosy diagnosis if that diagnosis was a operational classification/grade of disabilities other than the one being considered or if it was missing.

We estimated the incidence rate ratios (IRRs) of leprosy new case detection rate in the family (i.e., familial detection rate, FDR) in the matched cohort using Mantel-Haenszel tests and Poisson regressions with further adjustment for per capita income and robust standard errors clustered by family to account for matching with replacement. The pyr were included in the model as an offset variable. We estimated the cumulative incidence rate ratio for beneficiary and non-beneficiary families of BFP over time using the Nelson-Aalen estimator[22, 23]. In addition, we estimated the association of BFP participation and FDR of leprosy using Poisson regression models stratified by duration of follow-up (i.e., 0-6 months of exposure, 6-12 months, 1-2 years, 2-3 years, and 3+ years).

To account for the possibility that some individuals may have started receiving BFP after the sixth month, we also analysed BFP as a time-varying exposure. In this analysis, families that started receiving BFP between 6 months and 1 year after registration in CadUnico switched to the exposed group from 1 year on and were matched to families who had not received BFP by 1 year; similarly, families receiving between 1 and 2 years, and between 2 and 3 years switched to the exposed group and were matched to families remaining unexposed (see Web Appendix 4, Web Figure 1). To explore the robustness of our results to the way income was accounted for, since this is an important factor due to being the main eligibility criteria for BFP, we (i) excluded income from the Poisson regression model and (ii) adjusted for income using restricted cubic splines. Additionally, we estimated the association of BFP using inverse probability of treatment weighting and restricted the analysis to complete-cases (i.e., excluding participants with missing data for any covariate in the PS model) (Web Appendix 5). To test if there were potential biases due to differential loss of follow-up between BFP and non-BFP beneficiary families, we censored each matched pair by the smallest contribution of fyr for them to contribute to the same number of fyr. Finally, to test if there was competing risk bias due to lack of mortality information in our cohort, we limited the follow-up time of each matched pair to two years.

All analyses were performed using STATA version 15.0 (StataCorp LLC, College Station, Texas) and R version 3.5.2 (R Foundation for Statistical Computing, Vienna, Austria) (Packages: *dplyr, brmap, descr, ggplot2, ggthemes, gridExtra, grid, readxl, reshape2, and ggfortify*).

This study was performed under the international (Helsinki), Brazilian and UK research regulations and was approved by the three research ethics committee of the: (i) University of Brasília (1.822.125), (ii) Instituto Gonçalo Muniz-Fiocruz (1.612.302) and (iii) London School of Hygiene and Tropical Medicine (10580–1).

RESULTS

Of the 37,285,406 individuals in the 100 Million Brazilian Cohort who registered to CadUnico between 2007 and 2014, 31,613,355 individuals from 12,949,730 families were investigated in this study (Figure 2 From this sample, we identified 44,074 new leprosy cases among families in the cohort baseline between 2007 and 2014. This represents 94% (44,074/46,856) of the number of cases expected if the cohort had similar leprosy incidence

to the whole Brazilian population and 15.6% of the cases diagnosed in Brazil in the same period (Web Appendix 2, Web Table 2). After enrollment in the cohort, 4,328,630 commenced BFP participation within 6 months and additional 2,865,583 of the included families start received BFP benefits after that period. Among the 4,459,239 families living in high-burden leprosy municipalities, 41.9% (1,868,116/4,459,239) started benefiting from BFP within 6 months; and among the 8,490,491 families living in low-burden municipalities, 29.0% (2,460,514/8,490,491) start receiving BFP benefits within the same period.

Overall, leprosy cases were detected in 43,651 families in the cohort, of which 22,301 occurred after enrolment. Over the study period, from 2007-2014, leprosy incidence rates remained constant in BFP beneficiary and non-BFP beneficiary families from our cohort (Web Appendix 6, Web Figure 2). Out of the 22,301 incident cases, 8,622 (38.7%) cases were classified as PB, 13,661 (61.3%) as MB, and 18 (0.1%) as missing data on operational classification. 13,777 (61.8%) were diagnosed without disabilities, 6,290 (28.2%) were diagnosed with leprosy-associated disabilities (Grade 1/2), and for 2,234 cases (10.0%) grade of disabilities was not recorded. Overall, the FDR was 17.4/100,000 pyr (95%CI 17.1-17.7), and substantially higher in "priority" (high-burden) compared to "non-priority" (low-burden) municipalities (FDR=22,8/100,000, 95%CI 22.2-23.3 versus FDR=14.3/100,000, 95%CI 14.0-14.7). Crude cumulative leprosy incidence among families was markedly lower among BFP than among non-BFP beneficiaries (crude RR_{MH} =6.70, 95%CI= 0.68-0.73), with similar differences in high-burden (crude RR_{MH} =0.62, 95%CI= 0.59-0.65) and and in low-burden municipalities (crude RR_{MH} =0.69, 95%CI=0.66-0.72). (Figure 3A, 3B and 3C).

At baseline, there were significant differences between families who received BFP benefit (hereafter, BFP) and those that did not (hereafter, non-BFP) (Table 1). Relative to non-BFP participants, BFP family heads were more likely to be female (60.7% versus 53.4%) and were younger (median age 32.6 versus 40.2 years). BFP families also had relatively higher median numbers of individuals per family (3 versus 2), and lower median monthly per capita income (50.0BRL versus 177.7BRL, equivalent to 6.9% and 24.5% of the 2014 minimum wage). PS matching successfully matched >99.9% of the BFP families with similar non-BFP families in all matched samples (See Web Appendix 3, Web Figure 3, Web Figure 4, Web Table 4 and Web Table 5 for details of the PS analysis).

After matching, using Mantel-Haneszel tests or Poisson regression models with further adjustment for income, BFP was not associated with lower FDR of leprosy in our overall sample (IRR_{MH}=0.96, 95% CI=0.92-1.00; IRR_{Poisson}=0.97, 95% CI=0.90-1.04), but BFP beneficiary families living in high-burden municipalities had a substantially lower FDR of leprosy (IRR_{Poisson}=0.86, 95% CI=0.77-0.96) (Table 2). In high-burden municipalities, the point estimates for the association between BFP and leprosy was more extreme for the detection of leprosyassociated disabilities (IRR_{Poisson}=0.79, 95% CI=0.65-0.97) and paucibacillary cases (IRR_{Poisson}=0.82, 95% CI=0.68-0.98).

The cumulative FDR of leprosy was initially similar between beneficiary and non-beneficiary families, respectively (Figure 3C and 3D). However, after two years, the accrual of new cases detected was markedly lower among beneficiary families, and the difference in FDR by exposure status was larger among the families living in high-burden municipalities (Figure 3D). Also, by using Poisson models stratified temporally, the point estimate for the association between BFP receipt and leprosy FDR indicated slightly higher detection in the first 6 months on benefits, but lower FDR thereafter (Web Table 5 and Web Figure 5). Similar trends were observed for secondary endpoints related to grade of disabilities, but no differences over time were observed according to operational classification (Web Table 5 and Web Figure 5).

Sensitivity analysis

When allowing treatment vary over time, we obtained similar but less extreme point estimates for the association between BFP and leprosy primary endpoints (Web Table 6). We also obtained similar point estimates for the association between receiving BFP and FDR of leprosy when using Poisson regression without further adjusting for income (IRR_{Poisson}=0.96, 95% CI=0.89-1.03), and when further adjusting for income using spline (IRR_{Poisson}=0.93, 95%CI=0.87-1.00) (Web Table 7). Inverse probability of treatment weighting also generated similar estimates to the primary analysis, suggestive of slightly lower leprosy incidence among BFP beneficiary families (IRR_{Poisson}=0.95, 95%CI=0.90-1.00) and stronger point estimates among cases with disabilities (IRR_{Poisson}=0.87, 95%CI=0.79-0.96) (Web Table 8). The complete case analysis included 2,695,543 (63%) of the original BFP beneficiary families and yielded similar results to the primary analysis (IRR_{Poisson}=0.99, 95% CI=0.90-1.10) (Web Table 9). When considering the same number of fyr for each matched pair or restricting the

follow-up to 2 years, we also obtained similar or more extreme point estimates (Web Table 10 and Web Table 11).

DISCUSSION

This study investigated the impact of the Bolsa Familia conditional cash transfer program on new case detection of leprosy in a subset of the 100 Million Brazilian Cohort, which included 31.6 million individuals from over 12.9 million families. Our findings suggested that BFP was associated with lower incidence of leprosy among families living in high-burden municipalities for the disease in Brazil. We also obtained stronger point estimate for the association between BFP and lower incidence of paucibacillary forms and leprosy associated disabilities. These findings underscore the potential value of CCTs for the control of leprosy in low- and middle-income countries.

Our results indicate that families enrolled in the BFP between 2007 and 2014 who resided in high-burden municipalities for leprosy, had a 14% lower leprosy FDR relative to non-beneficiary families. These results point to a similar magnitude of the association between BFP and leprosy fisk to that previously described in ecological studies[14, 24]. These ecological studies have reported 15% lower leprosy new case detection rate in the general population and in children under 15 in Brazilian municipalities with high BFP coverage (\geq 48% coverage) compared to municipalities with low coverage (<28%)[14, 24]. Additionally, our study suggests that BFP is associated with fewer leprosy cases with PB presentations and among cases with disabilities, although point estimates were consistent across the other clinical presentations (i.e., MB forms and cases without disabilities). Due to the importance of reducing leprosy and related disabilities, these findings are of particular relevance to leprosy control strategies[2, 25].

CCTs are designed to have both short- and long-term impacts on beneficiary families[26]. By following up families for up to 7.5 years, our study provides new evidence that the association between BFP and lower leprosy incidence was more prominent after a minimum of two years in the program. The delayed association of the BFP may be partially explained by the chronic nature of leprosy, which has an incubation period of up to 10 years[27]. It is plausible that in the short term, BFP may increase food availability and bolster host immunity, while cumulative exposure to BFP may influence leprosy risk through longer-term mediators, such as education, crowding, and other social determinants of health[4, 11, 28]. Stronger point estimates for the association between

BFP and incidence of PB leprosy forms and leprosy-associated disabilities in high-burden leprosy municipalities deserve further consideration. As leprosy-associated disabilities can be prevented by early detection, enhanced healthcare utilization rates among beneficiary families could mediate the observed lower incidence of cases with disabilities[11]. Nevertheless, there is poor knowledge on the factors that mediate different immune response in PB and MB leprosy that explain more pronounced associations between BFP and PB leprosy forms. Also, increased access to healthcare among beneficiary families may increase leprosy detection and, therefore, it is likely that our results represent an underestimate of the causal effect of BFP on leprosy risk in high-burden municipalities is warranted, our results indicate that the impact of CCTs may have the greatest impact in scenarios where individuals face a higher and less heterogeneous disease risk[29].

The strengths and limitations of this study warrant consideration. The 100 Million Brazilian Cohort is a powerful resource of sociodemographic information covering the poorest half of Brazilian. Although previous studies have evaluated the association between BFP and leprosy or tuberculosis incidence in Brazil at the ecological level[14, 15], this is the first study to use linked administrative data to study the potential impact of a nationwide cash transfer program on infectious disease incidence at the individual level. Further, as leprosy is a rare disease, the large size of the analytical cohort provided unprecedented power to evaluate the associations between BFP and leprosy, as well as its understudied clinical manifestations. Finally, our analysis remained consistent, with similar point estimates in all sensitivity analyses conducted, including inverse probability of treatment weighting and restricted follow-up times. However, our study is also subject to limitations. First, although SINAN has national coverage, selection bias may have arisen due to the suboptimal linkage between the leprosy registry and the cohort baseline. This might be explained by potential heterogeneity in the quality of leprosy notification across Brazil, as individuals of mixed ethnicity and those living in the North and Northeast regions of the country appeared to be underrepresented among linked leprosy cases. Second, by defining exposure status at 6 months, we may have missed a very short-term impact of BPF participation in increasing leprosy diagnosis. Finally, residual confounding (e.g., distance to health clinics and/or access to primary health care) remains a concern even though key sociodemographic risk markers for leprosy were included in our propensity scores[4]. As this is a quasi-experiment, we are cautious regarding our causal claims. Nevertheless, as leprosy prevalence is low and the incubation period is long, and due to the nature of BFP as a nationwide social intervention, it would be very unrealistic to conceive an randomized control trial in this context.

This study has shown that the low-cost BFP (i.e., costing <0.4% of the Brazilian GDP in 2007) is associated with a significant reduction of leprosy in high-burden settings, including cases with G2D that are the focus of the WHO Global Leprosy Strategy 2016-2020[2]. We hypothesize that CCTs may reduce infectious disease morbidity, in part, by addressing some of the underlying determinants of health, such as poverty, education, healthcare access and nutrition[5, 7, 28, 30, 31]. A hundred years ago, it was stated that leprosy can be controlled with social development[32]. Now, a hundred years later, we have scientifically demonstrated that social policies, such as BFP could be a pillar for leprosy control, and perhaps contribute to its elimination. Although BFP has nationwide coverage, further efforts should be made to scale up the program to poor families that are just above the program eligibility threshold and living in municipalities with high leprosy risks. In conclusion, these findings indicate that relatively small cash transfer payments undertaken as part of long-term investment in social policies may have an important role in the control of a poverty-driven disease like leprosy.

Competing interests: None reported.

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Million Brazilian Cohort, 2007 to 2014 (N= 12,949,730		DED (9		BFP (4,328,630)			
Social and demographical variables	N	on-BFP (8	,621,100) Median (IQR)		BFP (4,2	Median (IQR)	- p-
	Ν	%	()	N	%	· · · · · · · · · · · · · · · · · · ·	value
Head of family characteristics							
Age			40.2 (26.1-59.2)			32.6 (26.4-42.1)	p<0.0 1
Sex							p<0.0 1
Males	4,017,128	46.6		1,699,12 4	39.3		
Females	4,603,972	53.4	^y y	2,629,50 6	60.7		
Ethnicity			\mathbf{N}				p<0.0
White	2,910,212	33.8		1,222,97 2	28.3		
Black	625,762	7.3		389,069	9.0		
Asian	43,379	0.5		1,8047	0.4		
Mixed/brown	4,566,436	53.0		2,465,30 4	57.0		
Indigenous	19,571	0.2		51,036	1.2		
Missing	455,740	5.3		182,202	4.2		
Literacy	\rangle						p<0.00
Yes	7,364,320	85.4		3,875,67 8	89.5		
No	1,220,161	14.2		422,482	9.8		
Missing	36,619	0.4		30,470	0.7		
Education							p<0.0 1
Primary school or less (≤5 years of education)	2,495,387	28.9		1,090,96 1	25.2		
Junior high school (≤ 9 years of education)	2069347	24.0		1,387,71	32.1		

Table 1 Description of non-beneficiary (non-BFP) and beneficiary families of the Bolsa Familia Program (BFP) within 6 months of registration to The 100 Million Brazilian Cohort, 2007 to 2014 (N= 12.949.730).

				/
High school (≥10 years of education)	2,055,301	23.8	5 990,945 22.9	
Missing	2,001,065	23.8	990,945 22.9 859,009 19.8	
Occupation	2,001,005	23.2		p<0.00
Currently not working	3,720,000	43.1	1,995,75 46.1	1
Working	3,996,326	46.4	1,773,20 41.0	
Missing	904,774	10.5	559,666 12.9	
Household characteristics				
Region of residence				p<0.00 1
North	890,499	10.3	558,683 12.9	
Northeast	2,810,584	32.6	$1,333,41 \\ 0 30.8$	
Southeast	2,969,267	34.4	1,777,09 41.1	
South	1,112,028	12.9	357,311 8.3	
Midwest	838,722	9.7	302,132 7.0	
Area of residence				p<0.00 1
Urban	7,055,818	81.8	3,548,22 4 82.0	
Rural	1,555,317	18.0	762,014 17.6	
Missing	9,965	0.1	18,392 0.4	
Leprosy high-burden municipality			2 1/2 21	
No	6,029,977	69.9	2,460,51 4 56.8	
Yes	2,591,123	30.1	$ \begin{array}{c} 1,868,11 \\ 6 \end{array} $ $ 43.2 \end{array} $	
Type of the household				p<0.00 1
Private	7,435,902	86.3	3,630,44 5 83.9	•
ORI		17		

Shared and informal housing	342,213	4.0	174,306 4.0	
Missing	842,985	9.8	523,879 12.1	
Construction material				p<0.00 1
Bricks/cement	6,894,374	80.0	3,402,24 78.6	
Wood, other vegetal materials and others	1,436,989	16.7	794,292 18.3	
Missing	289,737	3.4	132,097 3.1	
Water supply				p<0.00 1
Public network (tap water)	6,541,878	75.9	3,120,42 72.1	
Well, natural sources or others	1,789,487	20.8	1,076,09 24.9	
Missing	289,735	3.4	132,102 3.1	
Electricity				p<0.00 1
Electricity with counter	7,579,449	87.9	3,528,58 8 81.5	
Electricity without counter or no electricity	751,916	8.7	667,939 15.4	
Missing	289,735	3.4	132,103 3.1	
Sewage				p<0.00 1
Public network or septic tank	5,599,038	64.9	2,769,50 6 64.0	
Homemade septic tank, ditch or others	2,533,943	29.4	$ \begin{array}{c} 1,305,83 \\ 5 \\ 30.2 \end{array} $	
Missing	488,119	5.7	253,289 5.9	
Waste				p<0.00 1
Public collection system	6,898,397	80.0	3,430,66 3 79.3	-
Burned, buried, outdoor disposal or others	1,432,970	16.6	765,868 17.7	
Missing	289,733	3.4	132,099 3.1	
Basic services (water supply, electricicty, sewage and vaste)	l			p<0.00 1
OR		18		

					Å	
All adequate*	4,669,921	54.2	2,	146,96 49.6		
1 inadequate	1,832,776	21.3	93	35,173 21.6		
2 or 3 inadequate	799,949	9.3	48	36,692 11.2		
All inadequate	830,326	9.6	50	06,507 11.7		
Missing (all)	488,128	5.7	25	53,296 5.9		
Family members			2 (1-3)	\mathbf{N}	3 (2-4)	p<0.00
Residents per room			0.5 (0.3-0.8)		0.8 (0.5-1.0)	p<0.00
Number of children under 18 years old			0 (0-1)		1 (1-2)	p<0.00
Number of elders over 60 years old			0 (0-0)		0 (0-0)	p<0.00
Family income (BRL)		~	465 (110-724)		150 (30.0-300.0)	p<0.00 1
Per capita income (BRL)			177.7 (50-428.449)		50.0 (11.4-90.0)	p<0.00 1

^aTwo tailed t test used for comparison of continuous variables and Pearson chi-square for categorical variables; Missing data were considered a category.

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Table 2 Incidence rate ratio R) of leprosy (overall and according to grade of disabilities and operational classification of the second	cation) for BFP participation, in the
matched cohorts: Overall sample, high and lower leprosy burden municipalities.	

	1 0		*					/	
Brazil ^a	Leprosy cases	IR in BFP	95%CI	IR in non-BFP	95%CI	IRR ^b	95%CI	IRR ^c	95%CI
All new cases ^d	9,886	14.84	14.50, 15.18	15.48	14.91, 16.07	0.96	0.92, 1.00	0.97	0.90, 1.04
Grade 0	6,371	9.65	9.38, 9.93	9.73	9.28, 10.20	0.99	0.94, 1.05	1.00	0.92, 1.10
Grade 1 or 2 disabilities	2,534	3.74	3.57, 3.92	4.13	3,84, 4.45	0.91	0.83, 0.99	0.92	0.80, 1.05
Paucibacillary cases (PB)	4,022	6.09	5.87, 6.31	6.15	5.79, 6.52	0.99	0.92, 1.06	0.99	0.89, 1.10
Multibacillary cases (MB)	5,860	8.74	8.48, 9.00	9.33	8.89, 9.79	0.94	0.88, 0.99	0.96	0.87, 1.05
Leprosy high-burden municipalities ^e					/				
All new cases ^f	5,394	18.97	18.38, 19.58	22.26	21.17, 23.40	0.85	0.80, 0.90	0.86	0.77, 0.96
Grade 0	3,620	12.99	12.50, 13.49	14.19	13.33, 15.11	0.92	0.85, 0.98	0.91	0.80, 1.04
Grade 1 or 2 disabilities	1,251	4.29	4.01, 4.58	5.50	4.97, 6.08	0.78	0.69, 0.88	0.79	0.65, 0.97
Paucibacillary cases (PB)	2,415	8.43	8.04, 8.84	10.16	9.44, 10.94	0.82	0.76, 0.91	0.82	0.68, 0.98
Multibacillary cases (MB)	2,978	10.54	10.10, 11.00	12.09	11.29, 12.94	0.87	0.81, 0.94	0.89	0.77, 1.02
Leprosy low-burden municipalities ^g	$\langle \rangle$								
All new cases ^h	4,578	11.82	11.43, 12.76	12.08	11.43, 12.76	0.98	0.92, 1.05	0.99	0.90, 1.09
Grade 0	2,746	7.22	6.92, 7.55	6.90	6.41, 7.42	1.05	0.96, 1.14	1.06	0.94, 1.20
Grade 1 or 2 disabilities	1,319	3.35	3.14, 3.57	3.64	3.30, 4.03	0.92	0.82, 1.03	0.93	0.79, 1.11
Paucibacillary cases (PB)	1,672	4.39	4.15, 4.64	4.22	3.85, 4.63	1.04	0.93, 1.16	1.04	0.89, 1.21
Multibacillary cases (MB)	2,903	7.42	7.11, 7.75	7.85	7.34, 8.41	0.95	0.87, 1.02	0.96	0.86, 1.09
OF			20						

^a N=8,545,694 families; fyr=23,467,162.1; pyr=65,878,418.7.

^b Incidence rate ratio estimated estimated using Mantel-Haenszel method.

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^c Incidence rate ratio estimated using Poisson regression adjusting for income (continuous) and including robust standard errors clustered by family.

^d In the stratified analysis, missing in grade of disabilities at diagnosis (N=981) or operational classification (N=4) were censored at the time the leprosy case occurred.

^eN=3.674.130 families; fyr= 9,707,927; pyr=27,235,798.9.

^f In the stratified analysis, missingin grade of disabilities at diagnosis (N=523) or operational classification (N=1) were censored at the time the leprosy case occurred.

^g N=4.871.424 families; fyr=13,719,482.8; pyr=38,493,252.5.

^h In the stratified analysis, missing in grade of disabilities at diagnosis (N=513) or operational classification (N=3) were censored at the time the leprosy case occurred .

Figure 1 A) Priority municipalities for leprosy control in Brazil designated as high-burden and B) proportion of individuals residing in high-burden municipalities by State for the 26 Brazilian States and the Brazilian Federal district. High-burden municipalities include: all State capitals, municipalities in high risk areas for leprosy with a leprosy new case detection rate $\geq 20/100,000$ or ≥ 20 new cases or ≥ 10 new cases with at least 1 case in children under 15 years in 2010, and municipalities outside the geographical risk areas, with ≥ 50 new cases, with at least 5 cases in children under 15 years of age in 2010.

Figure 2 Flowchart of the study population.

Figure 3 Cumulative incidence of leprosy among families (per 100,000) defined as BFP (dashed line) and non-BFP (solid line) in the crude cohort: A) Overall; B) leprosy high-burden municipalities; C) leprosy low-burden municipalities; and in the matched cohort: D) Overall; E) leprosy high-burden municipalities; F) leprosy low-burden municipalities according to follow-up time. BFP+ Families who began receiving Bolsa Familia Program benefits within 6 months of enrollment in our cohort baseline; Non-BFP= Bolsa Familia Program non-beneficiary families.





