

Environmental sanitation conditions and health impact: a case-control study

Condições de saneamento ambiental e impacto na saúde:
um estudo caso-controle

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Abstract *This epidemiological investigation examines the impact of several environmental sanitation conditions and hygiene practices on diarrhea occurrence among children under five years of age living in an urban area. The case-control design was employed; 997 cases and 999 controls were included in the investigation. Cases were defined as children with diarrhea and controls were randomly selected among children under five years of age. After logistic regression adjustment, the following variables were found to be significantly associated with diarrhea: washing and purifying fruit and vegetables; presence of wastewater in the street; refuse storage, collection and disposal; domestic water reservoir conditions; feces disposal from swaddles; presence of vectors in the house and flooding in the lot. The estimates of the relative risks reached values up to 2.87. The present study revealed the feasibility of developing and implementing an adequate model to establish intervention priorities in the field of environmental sanitation.*

Key-words: Diarrhea. Hygiene practices. Model for priority setting. Urban refuse. Water supply.

Resumo *Esta investigação epidemiológica estuda o impacto das condições de saneamento ambiental e de práticas higiênicas sobre a ocorrência de diarreia entre crianças menores de 5 anos, residentes em uma área urbana. O delineamento caso-controle foi utilizado; 997 casos e 999 controles foram incluídos na investigação. Casos foram definidos como crianças com diarreia e controles foram selecionados, aleatoriamente, entre crianças com menos de 5 anos. Após ajustamento (regressão logística), as seguintes variáveis foram detectadas como significativamente associadas à diarreia: lavar e higienizar frutas e vegetais; presença de água de esgoto na rua; coleta, armazenagem e disposição do lixo; condições dos reservatórios domésticos de água; disposição das fezes de fraldas, presença de vetores nas casas e inundação do lote da casa. As estimativas pontuais do risco relativo alcançaram valores até 2,87. O presente estudo mostra a factibilidade de desenvolvimento e implementação de modelos adequados para estabelecer prioridades de intervenção no campo do saneamento ambiental.*

Palavras-chaves: Abastecimento de água. Diarreia. Lixo urbano. Modelo para estabelecer prioridades. Práticas higiênicas.

Although the World Bank⁴⁶ had discouraged the development of investigations involving environmental sanitation conditions and health impact from the mid 1970's, studies concerning these associations have been receiving increased attention since the beginning of the 80's.

In 1983, Blum & Feachem⁶ stated that most of the studies published until then had methodological limitations. These constraints were identified as from one to more than eight methodological flaws and, in several of the 44 studies reviewed, the results obtained could not be considered unbiased.

In the same year, a workshop on this subject was held in Bangladesh⁸ and, as conclusion of the discussions, the implementation of epidemiological studies on water supply and sanitation exposure was again recommended, provided that some important methodological care was observed. In order to increase the applicability of those studies, the workshop suggested the case-control design as the most adequate epidemiological method and child diarrhea morbidity as the health variable to be measured.

More than 250 studies have been carried out to investigate the probable association of environmental

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sanitation and health conditions. The following main features were observed from the analysis of 256 epidemiological studies published in the literature²³: a) Fifty-seven (146) percent of the studies were developed on Asian³¹ or African³² scenarios and this tendency has not changed over the last decades; b) Seventy-seven (198) percent of the studies investigated exposure related to water supply³⁴ and forty-two (107) percent, aspects related to domestic wastewater disposal²². Few investigations analyzed other environmental sanitation conditions, such as refuse disposal¹⁸ (2%, 4), hygiene habits⁵ (17%, 44) or drainage, vector presence⁴² and other forms of exposure (5%, 12). In the majority of cases, only rural areas were investigated³⁴; c) Forty-one (105) percent of the studies adopted diarrhea morbidity⁹ as the health variable; (d) Case-control designs began to be employed in the last decade. In the universe of studies

analyzed, prospective² (25%, 64) and cross-sectional³⁰ (21%, 53) designs predominated.

The present paper describes an epidemiological case-control study, which attempts to explore some aspects of the epidemiological methodology insufficiently investigated, which could be applied to environmental sanitation exposure, such as: (a) the applicability of the *case-cohort* or *inclusive* design³⁸, in which controls are chosen, as a random sample, among the population from where the corresponding cases are identified; (b) an investigation conducted in an urban area outside African or Asian environment; (c) the inclusion of a large set of environmental conditions, as multicategorical variables and (d) a test of sample sizes adequacy, suitable for generalizing to other similar sanitary and environmental situations.

MATERIAL AND METHODS

Studied area. The study was conducted in the urban area of Betim, a city with about 160,000 inhabitants. Betim is an industrial city, located in the Metropolitan Region of Belo Horizonte, the capital of Minas Gerais State, southeast Brazil, with a population of nearly 3.5 million inhabitants. A public concessionaire is responsible for the water supply and sanitation services. Other environmental sanitation services, for instance refuse collection and disposal, urban drainage and vector control are directly provided by the municipality.

Sample size. Considering the methods of sample size determination for independent case-control designs⁴⁰ and for multicategorical exposures⁷, a sample size of about 1,000 cases and equal numbers of controls was considered as adequate, assuming: 1) probability of type I error (alpha value) = 0.05; 2) power of the test = 0.90 (probability of type II error (beta value) = 0.10); 3) the prevalence of the exposure factors among the controls (p_c) to be equal 0.30, the lowest among the various factors being analyzed and 4) the minimum significant difference (Δ) between exposure factors prevalence among cases (p_c) and controls (p_c) equal to 0.10.

Questionnaires. A standardized protocol was developed, with the technical assistance of Universidade Federal de Minas Gerais faculty members, specialists in sanitary engineering, epidemiology, biostatistics and sociology. The questionnaire was based on information derived from other published investigations. In addition, a large spectrum of variables related to environmental sanitation was included. After a pilot test, the final version of the protocol was defined, including 80 closed questions, organized through the following sections: 1) informed consent; 2) identification of residents in the selected houses; 3) participant identification; 4) socioeconomic status of selected families; 5) household characteristics; 6) water supply and individual hygiene habits; 7) wastewater disposal and existence of nearby streams; 8) domestic refuse storage and disposal; 9) rainwater flooding and pounding; 10) vector presence;

and 11) validation of collected information (*in loco* observation).

Case selection. A case was defined as a child under five years of age, resident in Betim urban area, attended at a local health institution, whether public or private, with a report of diarrhea. The attendant physician diagnosis of diarrhea was assumed as the case definition. All local health institutions, including 15 public and 14 private health centers, were investigated. All cases diagnosed between December 20, 1993 and April 4, 1994 were included in the study, comprising a final sample of 997 cases.

Control selection. In coherence with the *case-cohort* or *inclusive* design definition, control was selected as a child under five years of age, randomly chosen from the resident population universe of the urban area of Betim. The control selection was based on a random allocation of houses, taken from a register used by the municipality with the purpose of housing taxes. For the allocation, algorithms for the register pages and lines were employed, using random numbers generated by a TurboBasic compiler.

While conducting the study, when the assigned house did not have a resident child under five years of age, displacement to the house on the left was adopted. For other situations, when the selection was not possible, other standardized criteria were established. For instance, when 1) the assigned house was selected twice from the allocation lists; 2) the assigned house was located outside of the studied area, or 3) no houses in the selected city block had a child under five years even after adoption of the left displacement criterion, another address was randomly chosen from the original register list.

The 999 selected controls were interviewed from November 23, 1993 to April 22, 1994; the majority of the data was collected before December 18, 1993.

Interviews. The interviews were carried out by a team of ten trained interviewers, recruited among local residents familiar with this kind of activity. The conduction

of Interviews in the same region in which the interviewer resides was avoided. Double-masked interviews were planned, but in some situations the participant status was obvious for the respondent. Information from the questionnaires was coded and introduced in a database, developed with the aid of the software MS-Access for Windows²⁶. All data was double-entered.

Reliability test. In a sub-sample (10% from the original sample), reliability tests were performed, through re-interviews. Four groups were defined: a) *Group 1*, cases/same interviewer; b) *Group 2*, cases/other interviewer; c) *Group 3*, controls/same interviewer and (d) *Group 4*, controls/other interviewer. The statistical analysis considered the values of the *kappa* statistics²⁰.

Data analysis. The data analysis was developed through a sequence of steps, when several associations and confounding factors had progressively been identified. The data set was organized using the software MS-Excel 4 for Windows⁴¹ and statistically analyzed through the software SYSTAT²¹, EPIINFO²² and MULTLR²³. The sequence of the statistical analysis

followed these steps: 1) frequency distribution; 2) univariate analysis, including: a) point estimate and confidence interval for the relative risk (Cornfield method⁴⁰), b) trend analysis (Mantel method⁴⁰) and c) point estimate and confidence interval for the attributable risk⁴⁰; 3) bivariate analysis, with inspection of potential confounders and effect modifiers (Mantel-Haenszel method³³); 4) multivariate analysis, using the logistic regression model²⁵ following the sequence: a) preliminary selection of variables, from the univariate analysis ($p < 0.25$)³⁶; b) intermediate logistic models construction, using 8 different homogeneous subgroups (familiar structure, socioeconomic variables, hygiene practices, water supply, sanitation, urban refuse disposal, drainage, vectors presence). Variables attaining a significance level of $p < 0.15$ were kept in these models; c) final model construction, maintaining only those variables reaching significance of $p < 0.05$ and d) effect modification analysis, under the multiplicative model. Variables known to be associated with diarrhea were kept in the model throughout the analysis, even when they did not reach the established significance levels.

RESULTS

Approximately 29% of identified cases were lost for interviewing; the main reason was the impossibility of localizing the address given at the participating clinics. Regarding the temporal pattern of cases distribution, it was found not to be associated with any meteorological event, such as air temperature or daily precipitation. Controls were found to be uniformly spread throughout the sixty-six Betim metropolitan regions; the proportion of controls per occupied house was also found to be evenly distributed in Betim.

Table 1 shows the frequency distribution, as well as the results of the univariate analysis for the socioeconomic and familial structure qualitative variables, with the respective relative risk (RR) and its 95% confidence interval. Trend analysis results for the polychotomous variables are also presented. Except for *gender* and for *person who takes care of the child*, all other variables analyzed, reflecting a lower socioeconomic family status or a disrupted familiar structure, were statistically associated with diarrhea. All polychotomous variables showed a strong linear increase in the risk of disease with increasing levels of exposure.

Table 2 shows the comparison of quantitative variables between cases and controls. Student's t-test demonstrated differences between these groups, except for the variable *duration of breast feeding*. Younger mothers and children, and variables reflecting lower

socioeconomic status were found to be associated with diarrhea occurrence.

In Table 3, crude RRs for the main exposures (converted to dichotomous variables), as determined in the univariate analysis, are presented, together with the respective 95% confidence interval.

After the multivariate adjustment, several of the variables significantly associated with diarrhea, based on the crude RR, lost their significant effect. The remaining variables, in general, showed a smaller point estimate of that risk as can be seen on Table 4.

Twenty-eight controls were later selected as cases. This identification allowed a verification of the differences between the estimates of the relative odds (RO), obtained by simulation of a traditional case-control study and the correspondent RR. The simulation was done through the exclusion of these 28 cases from the control group. The results showed a rigorous similarity between both risk measures.

Finally, the results of the reliability test indicated that 46% of the questions presented an almost perfect or substantial concordance; the remaining questions had regular, poor, or no concordance at all, according to Landis and Koch criteria for interpreting the kappa index²⁹. In general, questions related to personal habits or daily observations had the worst index of reliability than those regarding house and family descriptions.

DISCUSSION

Although the case-control design was used, the risk measure used throughout the study was the RR instead of the RO. This is conceptually supported by the sampling scheme employed for control identification,

characterizing the case-cohort or the inclusive variant of the case-control method^{38,27}.

Cases and controls were identified from the same population. Residence in Betim urban area was an

Table 1 - Frequency distribution, crude RR and O² test of socioeconomic and familial structure (qualitative variables)

Variable ^{*a}	Case		Control		RR (95% CI) ^b	p ^d
	n ^c	%	n ^c	%		
Mother lives with the child						
no	23	2.31	10	1.00	2.33	<0.05
yes	974	97.69	989	99.00	(1.05 – 5.26)	
Does the father live with child?						
no	161	16.16	125	12.53	1.35	<0.05
yes	835	83.84	873	87.47	(1.04 – 1.75)	
Number of children under 5						
1	604	60.58	757	75.78	1.00[1]	
2	331	33.20	219	21.92	1.89[2]	<0.001
3	52	5.22	21	2.10	3.10[3]	(3)
4 or more	10	1.00	2	0.20	6.27[4]	
Birth order						
first	730	73.22	886	88.69	1.00[1]	<0.001
second	235	23.57	102	10.21	2.80[2]	(3)
third or more	32	3.21	11	1.10	3.53[3]	
Gender						
male	527	52.86	502	50.25	1.11	>0.05
female	470	47.14	497	49.75	(0.93 – 1.33)	
Place where child spends the day						
at home	929	93.18	909	91.08	2.23 (1.38 – 3.61)	
at nursery, partial time ⁽²⁾	28	2.81	61	6.11	1.00	0.001
at nursery, full time	40	4.01	28	2.81	3.11 (1.53 – 6.36)	
Person who takes care of the child						
mother	843	84.55	829	83.07		
father	11	1.10	13	1.30	-	>0.05
other	143	14.34	156	15.63		
Mother education						
complete high school or higher	88	8.97	184	18.79	1.00[11]	
incomplete high school	36	3.67	54	5.52	1.39[7]	
complete primary school	87	8.87	144	14.71	1.26[4]	<0.001
incomplete primary school	731	74.52	583	59.55	2.62[2]	(3)
read and or write	14	1.43	7	0.72	4.18[1]	
does not read nor write	25	2.55	7	0.72	7.47[0]	
Father education						
complete high school or higher	69	8.32	167	19.49	1.00[11]	
incomplete high school	29	3.50	56	6.53	1.25[7]	
complete primary school	101	12.18	147	17.15	1.66[4]	<0.001
incomplete primary school	599	72.26	481	56.13	3.01[2]	(3)
read and or write	15	1.81	4	0.47	9.08 [1]	
does not read nor write	16	1.93	2	0.23	19.36 [0]	

Table 1 - Continue.

Variable* ^a	Case		Control		RR (95% CI) ^b	p ^c
	N	%	N	%		
Mother religion						
any religion	940	94.95	960	98.87	4.64	<0.001
no religion	50	5.05	11	1.13	(2.32 – 9.50)	
Father religion						
any religion	801	94.12	845	98.14	3.30	<0.001
no religion	50	5.88	16	1.86	(1.81 – 6.08)	
House ownership ^d						
score 3	35	3.56	78	8.20	1.00	
owner (score 2)	345	35.06	385	40.48	2.00	
bonded (score 1)	105	10.67	152	15.98	1.54	<0.001
invasion, arrested or rented (score 0)	499	50.71	336	35.33	3.31	(3)
Toilet existence						
more than one (one inside)	68	6.82	163	16.32	1.00[2]	
one, inside	680	68.20	702	70.27	2.32[1]	<0.001
one, outside	201	20.16	119	11.91	4.05[0.5]	(3)
does not have	48	4.81	15	1.50	7.67[0]	
Kitchen existence						
no	259	25.98	104	10.43	3.01	<0.001
yes	738	74.02	893	89.57	(2.34 – 3.89)	
TV ownership						
more than one (one in colors)	48	4.82	121	12.11	1.00[3]	
one in colors	381	38.25	510	51.05	1.88[2]	<0.001
one black and white	337	33.84	236	23.62	3.60[1]	(3)
no	230	23.09	132	13.21	4.39[0]	

*did not know the answer and refusals, when comprising less than 10% of answers, were excluded from Table

^a (1) category with proportion of cases significantly higher. (2) category with proportion of cases significantly lower.

^b [] attributed score, trend analysis.

^c (3) p value, trend analysis.

^d one unity added to score, when family owned another house.

Table 2 - Frequency distribution and mean differences (Student t-test) of socioeconomic and familial structure (quantitative variables).

Variables	Cases		Controls		t	p
	n ^o	mean	n ^o	mean		
Mother's age (years)	997	26.99	999	29.21	-6.83	< 0.001
Number of children under 5 living in the house	997	1.47	999	1.27	7.72	< 0.001
Child's age (years)	997	1.72	999	2.63	-15.86	< 0.001
Mean age of children under 5 (years)	997	1.89	999	2.53	-12.24	< 0.001
Number of persons living in the house	996	4.97	998	4.71	2.93	<0.01
Number of rooms	996	4.67	996	5.68	-11.28	< 0.001
Person/room	995	0.95	995	0.75	10.60	< 0.001
Income (minimum salary)	728	2.56	646	3.59	- 8.59	<0.001
Income/person (minimum salary/person)	727	0.61	645	0.85	- 7.36	< 0.001
Breast feeding period(months)	977	3.84	939	3.95	- 1.07	> 0.05

Table 3 - Crude RRs for main exposures, converted to dichotomous variables.

Variable	Comparison	RR (95% CI)
Fruits and greens hygiene	other care x disinfected	4.75(2.84 - 8.05)
Refrigerator ownership	no x yes	3.39(2.71 - 4.24)
Water domestic reservoir existence	no x yes	3.29(2.62 - 4.13)
Way water withdrawn from dug well	manual x pump	3.00 (0.74 - 13.16)
Feces disposal from swaddle	other x toilet/latrine	2.94(2.19 - 3.94)
Water supply source	other source x public network system	2.78(1.51 - 5.18)
Superficial presence of wastewater in street	yes x no	2.74(2.27 - 3.32)
Refuse storage	other x refuse package	2.51(2.05 - 3.06)
Hand hygiene after defecation	never/low frequency x frequent	2.34(1.84 - 3.06)
Bathroom existence	no/outside x one or more	2.15(1.69 - 2.73)
Flooding in lot	yes x no	2.11(1.75 - 2.56)
Rat presence	> once a semester x < once a year	2.08(1.72 - 2.52)
Refuse disposal	other x public collection	1.99(1.61 - 2.48)
Wastewater disposal	other x public collection system	1.97(1.63 - 2.37)
Hand hygiene before eating	never/low frequency x frequent	1.92(1.48 - 2.50)
Cockroach presence	>3 months a year x < one month a year	1.74(1.45 - 2.09)
Well water quality complaint	yes x no	1.67(0.35 - 7.22)
Refuse collection frequency	2 times a week x 3 times a week	1.66(1.30 - 2.11)
Fly presence	> 3 months a year x < 1 month a year	1.59(1.29 - 1.96)
Drinking water care	no x yes	1.55(1.33 - 2.14)
Domestic reservoir coverage	no x yes	1.52(1.00 - 2.31)
Mosquito presence	all time x < 6 months a year	1.48(1.23 - 1.78)
Pooling in lot	yes x no	1.46(1.19 - 1.80)
Chlorination in dug well	no x yes	1.29(0.18 - 6.94)
Public network water shortage	yes x no	1.24(1.03 - 1.49)
Contact with nearby stream water	yes x no	1.22(0.54 - 2.79)
Domestic reservoir cleaning	never x sometimes	1.19(0.88 - 1.61)
Near stream existence	yes x no	1.14(0.94 - 1.37)
Wastewater network blockage	yes x no	1.11(0.75 - 1.65)
Public network water complaint	yes x no	1.03(0.66 - 1.61)
Dug well coverage	no/inadequate x adequate	0.96(0.24 - 4.36)
Fruit and greens hygiene	no x yes	0.64(0.30 - 1.36)

inclusion criteria for cases, in order to allow a house visit. Cases identified in participating clinics whose domicile was not located by the interviewers, were excluded from the investigation. In consequence, it is unlikely that a squatter or a child living in an unregistered house would be included in the studied sample. Selection criteria for controls required permanent address in Betim.

Among the several exposures and confounding factors studied, after the multiple adjustment by logistic regression, only 16 dichotomous comparisons showed significant values for the relative risk, reaching up to 2.87 of magnitude (point estimate), as present in Table 4. This fact suggests a strong co-linearity between the environmental sanitation and the hygienic variables and the presence of several confounding factors.

It should be noted that the effect modification term is included only in a model that has both of the corresponding main effects. This is because these terms can be interpreted as effect modifiers only when the

corresponding main effect terms are contained in the model. This is the general rule in model building: higher-order terms are included in a model only when the corresponding lower-order terms are present¹⁴.

Some of the results are in accordance with the literature. Superficial presence of wastewater in street as risk for diarrhea can be seen as an analogous result to studies concerning lack of latrines^{2 32 22 16 35}. An inadequate management of domestic refuse showed an odds ratio of 2.48 for infantile diarrhea in Nigeria¹⁸ and a similar result was also observed in Brazil²¹. Vector presence, mainly flies, was associated to diarrhea in studies carried out in Thailand⁴³ and in Myanmar⁴². Relationship between hygiene practices and infantile health was identified in several investigations, like those developed in Bangladesh^{46 13 24}, USA²⁸, Brazil⁵ and Philippines⁴. Moreover, inadequate feces disposal from swaddle was found to be significantly associated with infantile diarrhea in studies developed in the Philippines³ and Bangladesh⁴³.

Table 4 - Variables remaining in the logistic model: RR and respective confidence interval, without and with effect modification.

Variable	Comparison	RRAdjusted (model without effect modification)	RRAdjusted (model with effect modification)
(1) Fruit and greens hygiene	other care x disinfected	2.87(1.61 - 5.10)	2.79(1.57 - 4.96)
(2) Mother's religion	no x yes	2.58(1.18 - 5.65)	2.68(1.21 - 5.92)
(3) Superficial presence of wastewater in street	yes x no	2.38(1.87 - 3.03)	1.47(1.00 - 2.16)
(4) Refuse storage	other x no storage	1.97(1.55 - 2.50)	1.46(1.08 - 1.97)
(5) Domestic reservoir	no storage x covered and cleaning reservoir	1.91(1.37 - 2.67)	1.43(0.88 - 2.35)
(6) Domestic reservoir	vessel storage x covered and cleaning reservoir	1.91(1.01 - 3.60)	1.62(0.58 - 4.50)
(7) Child's age	continuous variable	1.81(1.63 - 2.02)	1.83(1.64 - 2.03)
(8) Feces disposal from swaddle	no swaddle use x toilet/latrine disposal	1.65(1.21 - 2.24)	1.63(1.20 - 2.22)
(9) Refuse disposal	vacant lot/stream disposal x frequent public collection	1.61(1.11 - 2.34)	1.57(1.07 - 2.29)
(10) Children number	continuous variable	1.58(1.28 - 1.95)	1.61(1.30 - 1.99)
(11) Near stream existence	no x yes	1.57(1.22 - 2.01)	1.56(1.21 - 2.01)
(12) Feces disposal from swaddle	other x toilet / latrine disposal	1.50(1.04 - 2.19)	1.45(0.99 - 2.12)
(13) Refrigerator ownership	no x yes	1.41(1.12 - 1.76)	1.38(1.03 - 1.86)
(14) Cockroach presence	>3 months a year x < one month a year	1.40(1.12 - 1.76)	0.93(0.59 - 1.47)
(15) Flooding in lot	yes x no	1.39(1.09 - 1.76)	1.40(1.10 - 1.79)
(16) Mosquito presence	all time x < 6 months a year	1.37(1.08 - 1.73)	1.05(0.75 - 1.48)
(17) Refuse collection frequency	2 times a week x 3 times a week	1.33(0.99 - 1.79)	1.32(0.98 - 1.79)
(18) Domestic water reservoir	covered and not cleaning x covered and cleaning	1.07(0.82 - 1.40)	1.19(0.81 - 1.76)
(19) Domestic water reservoir	uncovered and cleaning x covered and cleaning	1.02(0.56 - 1.88)	0.35(0.12 - 1.04)
(20) Domestic water reservoir	uncovered and not cleaned x covered and cleaned	0.94(0.40 - 2.20)	0.52(0.12 - 2.18)
(3)*(4)	-	-	2.17
(5)*(14)	-	-	1.71
(6)*(14)	-	-	1.30
(18)*(14)	-	-	0.80
(19)*(14)	-	-	5.60
(20)*(14)	-	-	2.89
(14)*(16)	-	-	1.67

There are several descriptions in the literature of water supply association with health^{11 12 15 44}, while other studies do not show any association, for example an investigation in Panama³⁹. The importance of quantity of water consumption on health conditions has also been demonstrated^{44 19}. In this study, the lack of association between several aspects of water supply and health can be explained by: 1) the very low population exposure to the absence of public water supply (1.6%), due to the high population coverage and 2) to the practice, among Betim inhabitants, of clandestine connection to the public network, observed in the study. This situation reveals an effective non-existence of exposure.

Albeit references of the health importance of domestic water storage and recommendation for its improvement can be found in the literature³⁷, studies that quantify these effects were not identified. Similarly, previous references regarding a health effect of a nearby stream and flooding of rainwater in the lot were also not identified.

The possible limitations of this study findings include: 1) the fact that 29% of the total cases identified were excluded from interview; however, there is no evidence of any relationship between these exclusions and the exposures studied. A chi-square test, comparing the proportion of cases at participating Health Institutions, showed that the proportion of exclusions was statistically different only for two of them. Both were small Institutions, responsible for a low proportion of cases (1.8 and 4.1%). Indeed, this limitation would result in an underestimation of the established risks; 2) the source of controls, represented by the register used by the municipality for housing taxes, could exclude the *informal city* that is supposedly more exposed to the lack of environmental sanitation measures. This effect was minimized by the updated municipal file used in this study and the strategy of displacement to the house at the left when the assigned house did not have a child under five years old. This fact was very frequent and allowed the inclusion of the unregistered houses, since slums are very integrated to the urban design of the formal city, in Betim. This possible limitation implies in

an overestimation of the risks; and 3) the lag of about three months between cases and controls interview. As control selection did not presume disease definition and the environmental and behavioral exposures studied have a long duration pattern, this time difference probably did not imply in bias in the disease or exposure information. Besides, cases and control interview were conducted in the rainy season.

According to the results of the reliability test, variables related to public environmental sanitation conditions and house characterization – such as reservoir existence and conditions - are more reliable, since direct observation for validation of the answers was carried out. As a consequence, information related to personal and domestic habits were less reliable.

Generalization of the study results seems to be possible for similar urban areas, analogous in size, socioeconomic conditions and public services. It is also possible to visualize that a priority setting for intervention, based on the adopted design, can be a feasible approach. From this point of view, generalization of the present method, adjusted to a specific situation, reveals an important issue: the epidemiological design used - inclusive case-control or case-cohort - proves to be valid, since some potential bias on the control group selection, frequent in traditional case-control studies, can be avoided. However, some simplifications, like a smaller sample size, the investigation of a smaller number of confounding variables and the dichotomization of variables in the analysis phase, can be utilized.

The main conclusion of this investigation suggests that an important impact on health status of Betim's children can be achieved by implementation of environmental sanitation measures and hygiene education programs.

Finally, this study also enables the conclusion that infantile diarrhea has multiple and complex determinants. Environmental factors, associated to the lack of appropriate public urban services, poor hygiene practices and social determinants play an important role in transmission of this disease.

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