# Overview of acute diarrheal disease at the dawn of the 21st century: The case of Mexico

Lina Sofía Palacio-Mejía, MA, PhD,(1) Maylen Rojas-Botero, MSc,(2) Diana Molina-Vélez, MPH Biostatist and Health Inf Syst,(3) Concepción García-Morales, MPH on Biostatist and Health Inf Syst,(3) Leonel González-González, PhD,(4) Ana Lidia Salgado-Salgado, MSc, (3) Juan Eugenio Hernández-Ávila, MSc, DSc in Epidemiol, (3) Mauricio Hernández-Ávila, MSc, DSc in Epidemiol. (5)

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#### Abstract

**Objective.** To provide an overview of morbidity and mortality due to acute diarrheal disease in Mexico in order to understand its magnitude, distribution, and evolution from 2000 to 2016. Materials and methods. We carried out a longitudinal ecological study with secondary sources of information. We used data from epidemiological surveillance, health services, and vital statistics. We calculated and mapped measures of utilization of health services rates and mortality due to diarrheal diseases. Results. Diarrhea morbidity decreased by 42.1% across the period. However, emergency department attendances increased by 50.7% in the Ministry of Health. The hospitalization rate and mortality among the general population decreased by 37.6 and 39.7%, respectively, and the infant mortality rate decreased by 72.3% among children under five years of age. Chiapas and Oaxaca had the highest mortality among the states of Mexico. Conclusions. Cases of diarrhea, including rotavirus, have decreased in Mexico. However, in 2016, 3.4 per 100 000 people died due to diarrhea, which could have been avoided with health promotion.

Keywords: diarrhea; rotavirus; communicable diseases; morbidity; mortality; spatial analysis; Mexico

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#### Resumen

**Objetivo.** Ofrecer un panorama de la morbimortalidad por enfermedad diarreica aguda (EDA) entre 2000 y 2016 en México, para entender su magnitud, distribución y evolución. Material y métodos. Estudio ecológico longitudinal, con fuentes de información secundarias. Se analizaron datos de vigilancia epidemiológica, prestación de servicios y estadísticas vitales. Se calcularon tasas de utilización de servicios y mortalidad. Resultados. La morbilidad por EDA disminuyó 42.1% en el periodo, sin embargo, la atención por urgencias aumentó 50.7% en SSa. La tasa de hospitalización descendió 37.6% y la mortalidad 39.7% en población general y 72.3% en menores de cinco años. Chiapas y Oaxaca fueron los estados con mayor tasa de mortalidad. **Conclusiones.** Los casos de diarrea, incluyendo los de rotavirus, han disminuido en el país. Sin embargo, en 2016 se encontró una tasa de 3.4 por 100 000 personas que mueren por EDA, lo cual podría evitarse con promoción de la salud.

Palabras clave: diarrea; rotavirus; enfermedades transmisibles; morbilidad; mortalidad; análisis espacial; México

- Instituto Nacional de Salud Pública. Cuernavaca, Mexico.
- Grupo de Investigación, Demografía y Salud, Universidad de Antioquia. Medellín, Colombia.
- Centro de Investigación en Evaluación y Encuestas, Instituto Nacional de Salud Pública. Cuernavaca, Mexico.
- Centro de Información para Decisiones en Salud Pública, Instituto Nacional de Salud Pública. Cuernavaca, Mexico.
- Instituto Mexicano del Seguro Social. Mexico City, Mexico.

Received on: August 1, 2018 • Accepted on: December 18, 2018 Corresponding author: Dr. Juan Eugenio Hernández-Ávila. Instituto Nacional de Salud Pública. Av. Universidad 655, col. Santa María Ahuacatitlán. 62100 Cuernavaca, Morelos, Mexico. E-mail: juan eugenio@insp.mx

Non-communicable diseases (NCDs) are now the main causes of morbidity and mortality world-wide. However, acute infectious diseases still occur in developing countries, predominantly affecting the most vulnerable social and demographic groups such as children under five years old and the elderly.<sup>2</sup>

Despite improvements in sanitation and hygiene interventions, safe water, promotion of exclusive breast-feeding, availability of oral rehydration salts (ORS), and vitamin A supplementation, acute diarrheal disease (DD) remains the second underlying cause of death in young children worldwide. 3-6 Each year, DD claims the lives of 525 000 children under five years, 3 80% of which occurs in the first two years of life. 7

Rotavirus is a leading etiologic cause of severe DD in young children, causing more than 215 000 deaths each year throughout the world<sup>8</sup> and 40% of DD hospitalizations.<sup>9,10</sup> As a result, the rotavirus vaccine was introduced in 2006.<sup>11,12</sup>

In Latin American countries, there was a marked decrease in DD mortality between 1980 and 1990.<sup>13</sup> In Mexico, a reduction of 81.5% was observed between 1990 and 2000 (125.3 and 23.2 per 100 000 children under five years, respectively).<sup>14</sup> Still, a significant number of DD cases are registered.<sup>7</sup>

According to National Health and Nutrition Survey 2012 (Ensanut 2012 by its Spanish acronym), 11% of children under five years had DD episodes in the two weeks prior to the application of the survey; 53.7% were male, 21.0% were infants, and 54.8% were between one and two years old. DD prevalence decreased from 13.1% in 2006 to 11.0% in 2012. 16

The aim of this paper is to provide an overview of morbidity and mortality due to acute DD in Mexico between 2000 and 2016, using secondary data sources, to understand its magnitude, distribution, and evolution.

# Materials and methods

We carried out a longitudinal ecological study

## The Mexican Health System

Access to healthcare in Mexico is conditioned by employment status. For the population with social security (PWSS), which includes employees in the formal economy, government employees, and their families (about 50% of the Mexican population), healthcare is provided by the social security institutions (SSI). Meanwhile, for the population without social security (PWOSS), which includes the self-employed or unemployed, healthcare is provided by the Ministry of Health and State Health Services (SSa/SESA). Starting in 2003, people affiliated

to Seguro Popular (SP) also received healthcare from the SSa/SESA.

#### **Sources of information**

We analyzed DD epidemiological surveillance data, health services records, outpatient care records, hospitalization discharges, emergency services records, and mortality records. All data included year of occurrence and cause. Additional data included age, which was divided into large groups (infants: under one year, one to four years, children under five years (inclusive of infants), five to 64 years, and 65 years and over); sex; state of residence (n=32); and institution providing health care, which we analyzed as a dichotomous variable: "1" indicating the public SSI and "0" the SSa/SESA. Inclusion criteria were: registers occurring between 2000 and 2016 and DD as the underlying cause coded by the International Classification of Diseases (ICD-10 codes A00–A09, including A09X).<sup>17</sup>

Epidemiological surveillance is carried out by the General Directorate of Epidemiological (DGE) Surveillance of the SSa, a Nationwide Health Information System to collect data on mandatory reporting health conditions, including DD. Data is collected on a predefined form at the point of services by health care providers according to ICD-10 classification. This information is published weekly on the Epidemiological Surveillance Bulletin (BVE). <sup>18</sup> We present here annual accumulated data.

Data were available for only some ICD-10 codes, place of residence (state), and sex (since 2004). Available tables included all DD ICD-10 codes: intestinal infectious disease (A00-A09 include A09X), cholera (A00), typhoid fever (A01.0), paratyphoid fever (A01.1-A01.4), other Salmonella infections (A02), shigellosis (A03), other bacterial foodborne intoxications (A05), intestinal amoebiasis (A06.0-A06.3, A06.9), amoebic liver abscess (A06.4), other protozoal intestinal diseases (A07.0-A07.2, A07.9), giardiasis (A07.1), rotaviral enteritis (A08, used since 2014), and intestinal infections by other organisms and the ill-defined (A04, A08, A09; except A08.0). Unfortunately, health care providers are not well trained in ICD-10 coding, and because this information is collected during the service and confirmatory tests are seldom available, physicians are prone to classify most DD as intestinal infections by other organisms and the ill-defined.<sup>17</sup>

Emergency data was only available for SSa/SESA managed health facilities for the period 2007 to 2016. Data on emergency services is produced by health care providers at the point of care and collected at the patient level through a nominal information system.

Data on outpatient care is produced by healthcare providers at the point of care and collected through

the Health Information System (SIS), an aggregated numerical system managed by the General Directorate for Health Information (DGIS). This database is publicly available through the DGIS data portal in dynamic tables for the years 2000-2016. 20,21

The *hospital discharge database* is managed by the DGIS and contains patient level information of all hospital services provided by the public health sector for the years 2004-2015. <sup>20,22</sup> A more complete data set (2000-2016) is available for hospitals managed by the SSa/SESA. <sup>19</sup>

Mortality data is collected through a nationwide system managed by the National Institute of Geography and Statistics (INEGI by its Spanish acronym) and published by the DGIS in dynamic tables;<sup>23</sup> data is available from 2000 to 2016.

*Population data* was obtained from the DGIS data portal and included: a) general population estimates made by the National Population Council (Conapo) from 1990–2030,<sup>24</sup> b) data on population access to Social Security<sup>25</sup> and c) births estimates made by Conapo.<sup>26</sup>

## **Data analysis**

We calculated measures of DD frequency; outpatient, emergency, and hospitalization attendances rates; as well as the DD mortality rate. Where possible, data was disaggregated by sex, age group, place of residence (state), and access to social security institutions. Corresponding subpopulations were used as the denominator. We present graphical and tabular results at the national level and maps at the state level.

#### **Emergency department data analysis**

We calculated the emergency department attendance rate by dividing the total number of emergency department attendances to SSa/SESA by the total PWOSS. Mortality rates in the emergency department were calculated using the number of DD deaths registered in the emergency department as the numerator and all DD emergency admissions as the denominator, annually. The rate of referral from the emergency department to hospitalization was calculated by dividing the number of patients referred to hospitalization by the total number of DD emergency admissions, per year.

#### **Outpatient attendances data analysis**

According to availability of information, it was only possible to calculate the outpatient attendances rate for children under five years. The rate was calculated by dividing the total DD outpatient attendances by the total population under five years. The ratio of hospi-

talization by consultation was constructed using the total number of DD hospitalizations among children under five as the numerator and the total number of DD outpatient attendances among children under five as the denominator.

#### Hospitalization data analysis

The hospitalization rate was calculated using the number of DD discharges as the numerator and the respective population (with and without social security) as the denominator. The hospital mortality rate was calculated using the number of deaths due to DD in hospitalization as the numerator and the total DD discharges as the denominator.

#### Mortality data analysis

Mortality rates were calculated for the general population and sub-groups using appropriate population denominators. Infant mortality was calculated using live births as the denominator. We calculated the mortality rate ratio for some sub-populations, with 95% CIs to test for differences in mortality. We used MS Excel,\* ArcGis‡ and Stata.§

# Results

## **Epidemiological surveillance**

Cases of DD in all ages in Mexico decreased from 6.9 to 4.8 million between 2000 and 2016; 54.9% were female. The rate declined 42.1% (table I) over this period. Causes of DD during the period were, primarily, intestinal infections by other organisms and the ill-defined (79.2%), intestinal amoebiasis (10.0%), typhoid and paratyphoid fevers (2.1%), other protozoal intestinal diseases (1.5%), other bacterial foodborne intoxications (0.5%), giardiasis (0.5%), shigellosis (0.3%), and other *Salmonella* infections (0.2%).

Data available for rotavirus (2014-2016) showed a 48.5% reduction in the number of reported cases, with 1 299 cases in 2016. The highest incidences occurred in Chiapas (290 cases) and Baja California (213). Durango had only one registered case in two years.

Excel 2016 [computer program]. Version 2016. Redmond, Washington: Microsoft Corporation, 2016.

<sup>&</sup>lt;sup>‡</sup> ArcGIS [computer program]. Version 10.0. Redlands, CA2010.

<sup>§</sup> StataCorp. Stata 1996-2018; https://www.stata.com/

Table I

Estimated indicators for Acute Diarrhea Disease by source information, Mexico 2000-2016

							Ger	neral										
Source	Indicator									Year								
		2000	2001	2002	2003	2004	2005	2006	2007	2008	2009	2010	2011	2012	2013	2014	2015	2016
Epidemiologi- cal Surveil- lance	Epidemiological Surveillance Incidence Rate <sup>c</sup>	683.0	676.5	660.6	591.2	561.8	551.8	531.8	504.0	494.2	493.1	499.4	520.9	519.2	498.5	455.2	447.1	399.6
Emergency Department <sup>a</sup>	Emergency Department Admission Rate <sup>c</sup>								40.4	47.8	59.3	69.4	78.3	87.6	80.9	80.5	83.3	81.8
	Emergency Department Mortality Rate <sup>c</sup>								2.7	3.6	1.4	1.1	1.5	1.3	1.4	1.2	1.3	1.3
	Emergency Department Reference to Hospitalization Rate <sup>b</sup>								12.4	9.6	6.0	4.4	4.1	4.1	4.3	4.5	3.9	4.1
Outpatient Care	Outpatient Care Rate <sup>b</sup>	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
	Hospitalization by Consultation <sup>e</sup>					-	-	-	-	-	-	-	-	-	-	-	-	
Hospital Discharges	Hospitalization Rate <sup>c</sup>					7.1	7.9	7.6	7.8	7.2	5.9	5.5	5.4	5.8	5.4	4.6	4.5	
	Hospital Mortality Rate <sup>b</sup>					0.6	0.6	0.5	0.6	0.6	0.6	0.6	0.7	0.6	0.7	0.9	1.0	
Mortality	Mortality rated	5.2	4.8	4.5	4.3	3.9	4.0	3.6	3.4	3.2	2.9	2.8	2.9	2.8	3.0	2.9	3.1	3.1
	Infant Mortality Rate (< I year old) <sup>d</sup>	77.9	66.5	63.3	58.3	50.5	52.3	43.8	42.7	31.7	27.8	26.1	25.0	25.2	27.9	23.3	20.0	21.6
						Chil	dren un	der five	years									
Source	Indicator									Year								
		2000	2001	2002	2003	2004	2005	2006	2007	2008	2009	2010	2011	2012	2013	2014	2015	2016
Epidemiologi- cal Surveil- lance	Epidemiological Surveillance Incidence Rate <sup>c</sup>	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
Emergency Department <sup>a</sup>	Emergency Department Admission Rate <sup>c</sup>								149.0	175.5	200.6	234.1	250.5	283.8	255.9	250.3	254.2	264.0
	Emergency Department Mortality Rate <sup>c</sup>								2.8	2.9	1.0	1.2	1.1	1.0	1.3	0.6	0.5	0.8
	Emergency Department Reference to Hospitalization Rate <sup>b</sup>								14.5	10.5	6.7	4.6	4.4	4.3	4.6	4.8	4.0	4.5
Outpatient Care	Outpatient Care Rate <sup>b</sup>	15.9	11.1	9.8	8.6	11.6	12.3	13.0	12.7	12.8	12.1	11.8	11.8	11.9	14.4	9.3	11.4	
	Hospitalization by Consultation <sup>e</sup>					3.0	3.3	2.9	3.4	2.9	2.2	2.0	1.9	2.1	1.6	2.1	1.6	
Hospital Discharges	Hospitalization Rate <sup>c</sup>					34.4	41.0	37.3	42.7	37.5	26.8	23.6	22.4	25.0	22.5	19.4	18.8	
	Hospital Mortality Rate <sup>b</sup>					0.5	0.5	0.4	0.4	0.3	0.4	0.4	0.5	0.3	0.5	0.5	0.4	
Mortality	Mortality rated	21.8	19.0	18.6	17.5	15.6	16.2	13.0	13.0	10.5	8.5	7.6	7.8	7.6	8.4	7.1	6.5	7.3

a Ssa; b Rate per 100; c Rate per 10 000; d Rate per 100 000; e Ratio

Source: DGE, SSa. Epidemiological Surveillance in Mexico, 2000-2016.18 DGIS, SSa. Dynamic table of Emergency departments, México, 2007-2016.22 DGIS, SSa. Dynamic table outpatient care services, Mexico, 2000-2016.23 DGIS, SSa. Dynamic table by hospital discharges, Mexico, 2000-2016.19 INEGI, DGIS. Dynamic table by mortality, Mexico 2000-2016.21 Birth's Projections in Mexico 1990 - 2030. CONAPO, DGIS. Population's Projections in Mexico, 1990 - 2030.26

#### **DD** attendance to emergency departments

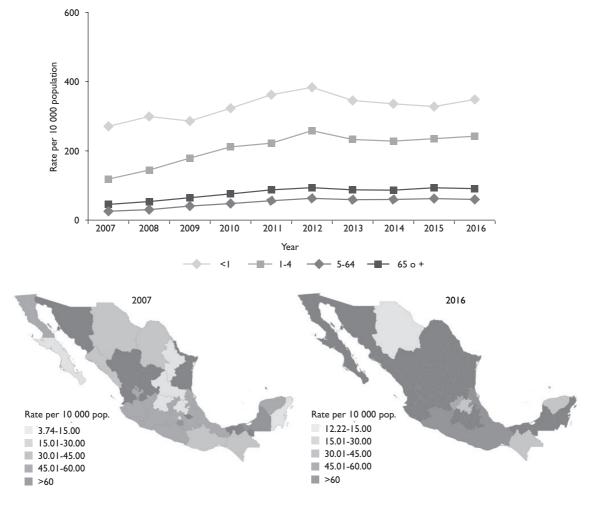
The Mexico SSa reported 547 790 DD emergency department attendances, equivalent to 81.8 per 10 000 PWOSS in 2016. Until 2012, there was a steady increase in the number of emergency department attendances, up to 87.6/10 000, and the rate remained stable around 81/10 000 thereafter. Again, a greater proportion of patients (56.1%) were female. When stratified by age group, emergency care was provided mainly to infants (an annual average of 32.8 attendances per 1 000 children under one year), followed by children 1-4 years of age (2.07 per 1 000 children 1-4). Elderly and adults were attended to a lesser extent (figure 1). Of all registered attendances, 5.2% were referred to hospitalization, while 1.5 out of 10 000 ended in death.

Maps in figure 1 showed no spatial pattern to DD attendance in emergency departments. In 2016, Colima reported the highest rate (46.6/1 000), followed by Coahuila (28.7/1 000) and Campeche (18.5/1 000). In contrast, the lowest rates were found in Chihuahua, Chiapas, Hidalgo, and Yucatan, where less than 3/1 000 population attended the emergency department for DD.

Emergency attendance due to rotavirus reduced by 77.7% during the period, from 1.8/100 000 population in 2007 to 0.4/100 000 in 2016, with no apparent spatiotemporal pattern.

#### **DD** attendance in outpatient care

Total DD outpatient attendance in children under five years decreased by 31.8% in the 2000-2015 period. In



Source: DGIS, SSa<sup>22</sup> and Conapo, DGIS.<sup>26</sup>

FIGURE 1. DD ATTENDANCE IN EMERGENCY DEPARTMENTS BY STATE OF RESIDENCE AND GROUP OF AGE. MÉXICO, 2007-2016

2000 there were 1.8 million attendances recorded, while in 2015 the attendances were 1.3 million. The outpatient attendance rate due to DD in children under five years decreased by 28.4% between 2000 and 2015. In 2011 we found a large inconsistency in data for IMSS-Prospera: the number of outpatient consultations was 3 287 323, which was clearly an outlier, so we imputed the data for this year by taking the average of the adjacent years (figure 2).

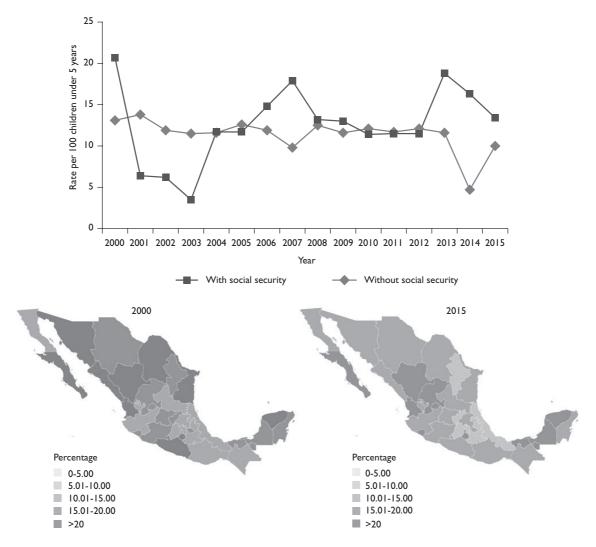
Data on outpatient attendance in SSI also showed inconsistencies throughout the study period (figure 2), therefore we decided not to interpret it and only present data on outpatient care services provided by SSa/SESA. Campeche, Durango, and Nayarit had the highest DD outpatient attendances in children (140.6, 139.5, and

134.1 attendances per 1 000 children under five years, respectively). There is an overall slight decreasing trend in outpatient services provided by SSa/SESA.

By cross-referencing outpatient services and hospitalization data in children under five years we found that, on average, there were 2.4 DD hospitalizations for every 100 DD outpatient consultations, with a decreasing trend over the study period.

#### **DD** hospitalization

The DD hospitalization rate reduced by 37.6% during the period; 50.3% were male. The DD hospitalization rate was 50.2/10 000 among infants, 24.1/10 000 among children



Source: DGIS, SSa<sup>20</sup> and Conapo, DGIS.<sup>26</sup>

FIGURE 2. DD OUTPATIENT ATTENDANCES RATE IN CHILDREN UNDER FIVE YEARS OLD BY STATE OF RESIDENCE AND SOCIAL SECURITY CONDITION. MÉXICO, 2000-2016

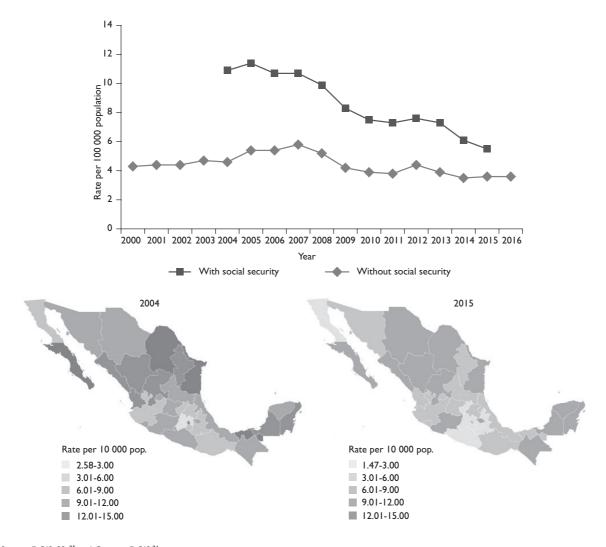
1-4 years, 3.1/10 000 among the population 5-64 years, and 11.1/10 000 among the population over 65 years. We observed a decreasing trend in all groups, starting in 2007.

The DD hospitalization rate was higher in PWSS (58.8%). The trend decreased by 49.5% from 2004 (10.9/10 000) to 2015 (5.5/10 000). In PWOSS, the magnitude of the decreasing trend was smaller (21.1%) from 2000 (4.6/10 000) to 2015 (3.6/10 000). Overall, Coahuila, Yucatán, and Sinaloa reported the highest DD hospitalization rate (8.8, 8.4, and 8.2 discharges per 10 000 population) (figure 3).

The DD hospital mortality rate increased by 53.3% between 2004 and 2015, from 0.6 to 1 deaths per 100 discharges.

Hospital discharges for Rotavirus decreased from 522 records in 2004 to 122 in 2015, representing 9.3 and 2.2% of all DD discharges, respectively. The higher percentage of deaths were among males (58.3%). Between 2004-2015, a total of 2 886 (51.2%) discharges were among children 1-4 years and 2 189 (38.8%) were among infants. These rates showed a decreasing trend over the period; in infants the rate decreased from 9.4/100 000 in 2004 to 2.1/100 000 in 2015, and in children 1-4 years the rate decreased from 2.7/100 000 in 2004 to 0.7/100 000 in 2015.

The median hospitalization rate due to rotavirus in children under-five years old decreased 59.1% after the introduction of the rotavirus vaccine. The median rate



Source: DGIS, SSa $^{25}$  and Conapo, DGIS. $^{26}$ 

FIGURE 3. HOSPITALIZATION RATE DUE TO DD BY STATE OF RESIDENCE AND SOCIAL SECURITY CONDITION. MÉXICO, 2000-2016

in the pre-vaccine era (2004-2006) was 5.2 hospitalizations per 100 000 children under-five years old while (table I), the rate was 2.1/100 000 in the postvaccine era (2007-2016).

Rotavirus hospitalizations were more common in PWOSS (76.4%). Chiapas and Guanajuato had the highest rate in 2015 (12.2 and 5.0 per million population, respectively) and 17 states did not report a case. In hospital mortality due to rotavirus was 0.3% (19).

## **DD Mortality**

Between 2000 and 2016, 66 849 people in Mexico died whose underlying cause of death was diarrhea. In this period, DD-specific mortality decreased by 39.7%. Mortality was higher in males at the beginning of the time series, however in 2008 the trend changed and females had the highest mortality rate from then until 2016 (figure 4). As the map shows, southern states had the highest mortality rates due to DD. Throughout the time series, Chiapas had the highest DD mortality rate; its peak value was in 2001 (14.4 deaths per 100 000) and its lowest was in 2011 (8.4/100 000).

According to age group, the highest risk of death by DD was in children under five years. In 2000, the risk of death by DD in infants was 63.9 times higher than in the population aged 5-64 (95%CI 59.2 - 68.9), though this ratio decreased to 23.5 in 2016 (95%CI: 21.0 - 26.2). The largest decrease occurred in infants; the infant mortality rate decreased by 72.3% between 2000 and 2016. As of 2010, the infant mortality rate due to DD became very close to the mortality rate of elderly people. The mortality rate in people over 65 years was the highest between 2014 and 2016.

There were differences in DD mortality according to the health coverage status. Between 2000 and 2016, PWSS had an average annual mortality rate due to DD of 2.3 deaths per 100 000 population, compared with PWOSS, who had an average annual mortality rate of 4.5 deaths per 100 000 population (almost twice, 95%CI 1.90 – 1.97). However, the gap was closed, such that the risk of dying due to DD decreased from 2.6 (95%CI 2.4 – 2.7) in 2000 to 1.2 (95%CI 1.1 – 1.2) in 2016, comparing people with and without social security. Among PWOSS, DD mortality decreased rapidly until 2009, after which the rate has remained almost unchanged.

There were 180 recorded deaths due to rotavirus during the study period. The average annual rate was one death per 100 000 population; this rate decreased by 76.4% over the period (from 1.4/100~000 in 2000 to 0.3/100~000 in 2016). The trend was similar by sex, though slightly higher among males (average annual rate 1.2~vs~0.8/100~000). The majority of rotavirus deaths

occurred in infants (65.6%), followed by children between 1-4 years (27.2%). Once again, PWOSS had 5.6 times the risk of dying by rotavirus compared to PWSS (95%CI: 3.6 - 8.7). According to the state of residence, Colima and Veracruz had the highest rotavirus mortality (23.1 and 13.0 /10 million population).

# Discussion

In this paper, we reviewed all sources of administrative records and population projections in order to analyze the magnitude, distribution and evolution of acute diarrheal diseases (DD) in Mexico from 2000-2016. Among our main results, we found an important decrease in the number of DD events reported in the Epidemiological Surveillance System, which was correlated to a decrease in the DD outpatient attendance rate in children under five years, as well as in the DD hospitalization rate. Although DD mortality decreased considerably, DD emergency department attendances doubled during the 2007-2016 period and hospital mortality due to DD increased.

Our study showed that DD mortality in children under five years of age has steadily and substantially decreased in Mexico between 2000 and 2010, and has remained almost constant until 2016. This trend could be attributed to the introduction of the rotavirus vaccine throughout the country between 2006 and 2007. The vaccine has had an efficacy of 85% against rotavirus-related disease and hospitalization, which is consistent with the decrease in hospitalizations and deaths due to rotavirus demonstrated in this study. In Mexico, the introduction of the rotavirus vaccine reduced overall DD hospitalizations in children under five years of age by 47%. Consistent with this previous research, we also observed a 59.1% reduction in the hospitalization rate due to rotavirus.

As reported by the literature, the factors that have contributed to the decrease in DD mortality in Mexico have been, mainly, the distribution and widespread use of ORS; the program *Piso firme*; 15,29 the increase of exclusive breastfeeding; an improvement in nutrition, sanitation, and hygiene conditions; vaccination against rotavirus and measles; and Zinc supplementation. 5,30,31

Our data suggest an important increase in emergency health services utilization, in concordance with official norms and programs launched by state and federal health authorities,<sup>32,33</sup> as well as an important decrease in the DD mortality rate in the emergency services. The observed decrease in the DD mortality rate in emergency departments reflects improvements in diarrheal management and treatment in this area.

Paradoxically, there was a 66.7% increase in hospital mortality due to DD. This could be the result of a

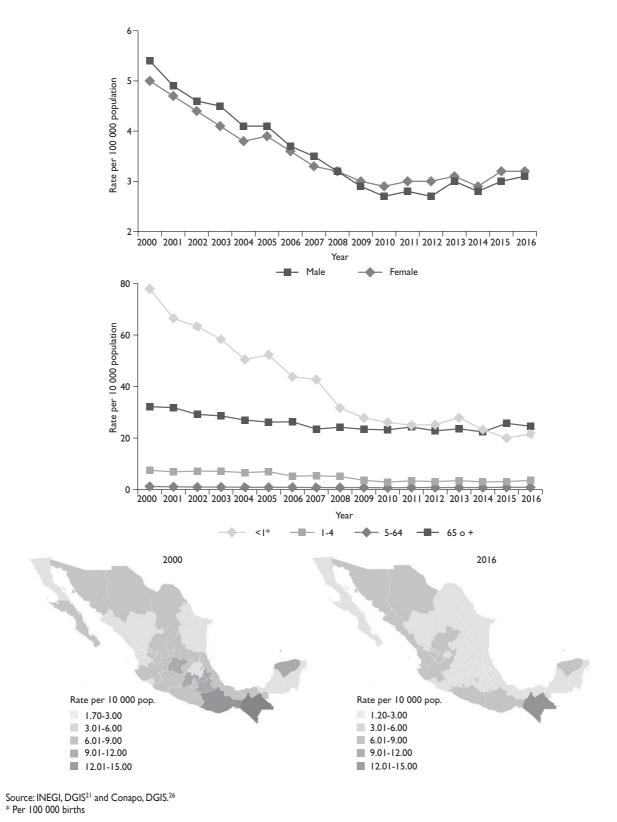


FIGURE 4. DD MORTALITY RATES BY SEX, AGE GROUP, AND STATE OF RESIDENCE. MÉXICO, 2000-2016

better handling of DD patients in the emergency room, such that only the most severe cases are referred to the hospitalization area. This had the effect of reducing the total number of hospitalized cases (denominator), but not the actual number of deaths (numerator).

Chiapas and Oaxaca had the highest diarrhea mortality rates among the states of Mexico during the entire period, which could be an indicator of gaps in the social determinants of health and a reflection of inequity. It should be mentioned that, although Guerrero does not appear among the states with the highest mortality due to DD, an analysis of under-registration carried out in 2018,<sup>34</sup> showed Guerrero as one of the territories with the highest DD mortality rates in the country. Additionally, the population without social security had a higher DD mortality compared to the population with social security, which points again to social inequalities in health.

Lack of accessibility to health services is still a barrier that can have a significant impact on occurrence of deaths due to diarrhea,35 so it is crucial to achieve universal access for the Mexican population. Future research should focus on analyzing the increase of emergency department attendances. A second issue was the increase in hospital mortality due to DD. Despite the substantial reduction in DD morbidity and mortality, 3 843 people in Mexico died due to DD in 2016, even though it corresponds to one of the Sustainable Development Goals,<sup>36</sup> and there are well-known and low-cost interventions for its treatment. DD mortality magnitude is an indicator of inequality in the social determinants of health, and the elimination of all avoidable differences among social groups should be an ethical imperative for the benefit of the Mexican population.

#### Limitations

This study has several limitations, including those of an observational study with secondary data sources. We used public information which, despite undergoing periodic quality assurance by each responsible institution, showed some inconsistencies. For example, data on outpatient consultation for Social Security Institutions had outliers which precluded its use. Another important limitation was the lack of available epidemiological data in usable format. Epidemiological data should be publicly available in database format, disaggregated by age group and sex. The available epidemiological surveillance data was further limited by the high number of cases classifed as intestinal infections by other organisms and the ill-defined, which could impact in the estimation of cause-specific DD. Another consideration may be the underreporting of deaths, which is about 5% in Mexico and most prevalent in infants and under five years in marginalized areas (unpublished data). In a study reported in 2012, researchers found 22.6% underreporting in children under five years old. However, this study was carried out in a representative sample of the 101 municipalities with the lowest Human Development Index,<sup>37</sup> where underreporting is more prevalent than in the current study.

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#### References

- I. World Health Organization. Global status report on noncommunicable diseases 2014. Attaining the nine global noncommunicable diseases targets; a shared responsibility. Geneva, Switzerland: WHO, 2014.
- 2. Bygbjerg IC. Double burden of noncommunicable and infectious diseases in developing countries. Science. 2012;337(6101):1499-501. https://doi.org/10.1126/science.1223466
- 3. World Health Organization. Geneva: WHO; c2018. Enfermedades diarreicas. Datos y cifras. WHO, 2017 [cited 2018 Jul]. Available from: http://www.who.int/es/news-room/fact-sheets/detail/diarrhoeal-disease
- 4. World Health Organization, United Nations Children's Fund. Ending preventable child deaths from pneumonia and diarrhoea by 2015: The integrated Global Action Plan for Pneumonia and Diarrhoea (GAPPD). France: WHO, 2013. https://doi.org/10.1163/2210-7975\_hrd-9841-3003
- 5. Richardson V, Hernandez-Pichardo J, Quintanar-Solares M, Esparza-Aguilar M, Johnson B, Gomez-Altamirano CM, et al. Effect of rotavirus vaccination on death from childhood diarrhea in Mexico. N Engl J Med. 2010;362(4):299-305. https://doi.org/10.1056/NEJMoa0905211
- 6. Sepúlveda J, Bustreo F, Tapia R, Rivera J, Lozano R, Oláiz G, et al. Improvement of child survival in Mexico: the diagonal approach. Lancet. 2006;368(9551):2017-27. https://doi.org/10.1016/s0140-6736(06)69569-x 7. Tomé P, Reyes H, Rodríguez L, Guiscafré H, Gutiérrez G. Muerte por diarrea en niños: un estudio de factores pronósticos. Salud Publica Mex. 1996;38(4):227-36.
- 8. World Health Organization. Immunization, vaccines and biologicals: rotavirus. Geneva: WHO, 2013.
- Cabrera-Gaytán DA, Maldonado-Burgos MA, Rojas-Mendoza T, Grajales-Muñiz C. Enfermedad diarreica aguda en niños menores de cinco años de edad: aportaciones de los núcleos trazadores de vigilancia epidemiológica 2012-2013. Arch Inv Mat Inf. 2013;(3):118-25.
- 10. World Health O. Weekly epidemiological record Relevé épidémiologique hebdomadaire. Wkly Epidemiol Rec. 2008;83(47):421-8.
- II. Reyna-Figueroa J, Vidal-Velázquez RP, Richardson López-Collada VL. Inmunización contra rotavirus en México con vacuna oral monovalente. Evaluación de los datos de dos años del sistema de reporte de eventos temporalmente asociados a vacunación. Rev Invest Clin. 2011;63(4):391-8.
- 12. Esparza-Aguilar M, Bautista-Márquez A, González-Andrade MC, Richardson-López-Collada VL. Mortalidad por enfermedad diarreica en menores, antes y después de la introducción de la vacuna contra el rotavirus. Salud Publica Mex. 2009;51(4):285-90. https://doi.org/10.1590/s0036-36342009000400004
- 13. Bern C, Martines J, de Zoysa I, Glass RI. The magnitude of the global problem of diarrhoeal disease: a ten-year update. Bull World Health Org. 1992;70(6):705-14.
- 14. Velásquez F, Garcia-Lozano H, Rodríguez E, Cervantes Y, Gómez A, Melo M, et al. Diarrhea morbidity and mortality in Mexican children: Impact of rotavirus disease. Pediatr Infect Dis J. 2004; 10(suppl): \$149-55. https://doi.org/10.1097/01.inf.0000142463.72442.91

- 15. Gutiérrez J, Rivera-Dommarco J, Shamah-Levy T, Villalpando-Hernández S, Franco A, Cuevas-Nasu L, et al. Encuesta Nacional de Salud y Nutrición 2012. Resultados nacionales. Cuernavaca, México: INSP, 2012.
- 16. Ferreira-Guerrero E, Mongua-Rodríguez N, Díaz-Ortega J, Delgado-Sánchez G, Báez-Saldaña R, Cruz-Hervert LP, et al. Diarreas agudas y prácticas de alimentación en niños menores de cinco años en México. Salud Publica Mex. 2013;55(supl 2):S314-22. https://doi.org/10.21149/spm.v55s2.5130 17. World Health Organization. International statistical classification of diseases and related health problems: ICD-10.Vol 2. 10th revision ed. Geneva. Switzerland: WHO. 2010.
- 18. Histórico Boletín Epidemiológico [internet]. México: Secretaría de Salud. c2018 [cited 2018 June]. Available from: https://www.gob.mx/salud/acciones-y-programas/historico-boletin-epidemiologico
- 19. Dirección General de Información en Salud [internet]. México: Secretaría de Salud; c2018 [cited June 2018]. Cubo dinámico: Defunciones (Mortalidad) 1979 2016. Available from: http://sinba08.salud.gob.mx/cubos/cmortalidadxp.html
- 20. Dirección General de Información en Salud [internet]. México: Secretaría de Salud; c2018 [cited June 2018]. Cubo dinámico: Servicios SIS. Servicios Otorgados y Programas Sustantivos 2000 2015 Sector Salud (SS, IMSS, IMSS-Oportunidades, ISSSTE, PEMEX, SEDENA, SEMAR, Otros). Available from: http://sinba08.salud.gob.mx/cubos/cboletin.html
- 21. Dirección General de Información en Salud [internet]. México: Secretaría de Salud; c2018 [cited June 2018]. Cubo dinámico: Proyecciones de la Población de México 1990 2030, CONAPO. Available from: http://sinba08.salud.gob.mx/cubos/cconapo2010proy.html
- 22. Dirección General de Información en Salud [internet]. México: Secretaría de Salud; c2018 [cited June 2018]. Cubo dinámico: Servicios SIS. Servicios de Salud: Concentrado estatal 2000-2006 y concentrado por unidad médica 2007-2016. Available from: http://www.dgis.salud.gob.mx/contenidos/basesdedatos/bdc serviciossis gobmx.html
- 23. Dirección General de Información en Salud [internet]. México: Secretaría de Salud; c2018 [cited June 2018]. Cubo dinámico: Egresos Hospitalarios 2004 2013 Instituciones Públicas morbilidad SECTOR SALUD (SS, IMSS, IMSS-Oportunidades, ISSSTE, PEMEX, SEDENA, SEMAR, Otros). Available from: http://sinba08.salud.gob.mx/cubos/ciNS-TITUmorbi.html
- 24. Dirección General de Información en Salud [internet]. México: Secretaría de Salud; c2018 [cited June 2018]. Cubo dinámico: Egresos Hospitalarios 2008-2016 Instituciones Públicas morbilidad por unidad médica SECTOR SALUD (SS, IMSS, IMSS-Oportunidades, ISSSTE, PEMEX, SEDENA, SEMAR, Otros). Available from: http://sinba08.salud.gob.mx/cubos/ciNSTITUmorbi sector.html
- 25. Dirección General de Información en Salud [internet]. México: Secretaría de Salud; c2018 [cited June 2018]. Cubo dinámico: Egresos Hospitalarios Secretaría de Salud 2000-2016. Available from: http://www.dgis.salud.gob.mx/contenidos/basesdedatos/bdc\_egresoshosp\_gobmx.html 26. Dirección General de Información en Salud [internet]. México: Secretaría de Salud; c2018 [cited June 2018]. Cubo dinámico: Proyecciones de

- los Nacimientos de México 1990 2030, CONAPO. Available from: http://sinba08.salud.gob.mx/cubos/cconapo\_nac\_edad2010.html
- 27. Ruiz-Palacios G, Pérez-Schael M, Velázquez R, Abate H, Breuer T, Clemens SC, et al. Safety and efficacy of an attenuated vaccine against severe rotavirus gastroenteritis. N Engl J Med. 2006;354(1):11-22. https://doi.org/10.1056/NEIMoa052434
- 28. Sánchez-Uribe E, Esparza-Aguilar M, Parashar UD, Richardson V. Sustained reduction of childhood diarrhea-related mortality and hospitalizations in Mexico after rotavirus vaccine universalization. Clin Infect Dis. 2016;62(suppl\_2):S133-9. https://doi.org/10.1093/cid/civ1205
- 29. Cordero-Arroyo E. Mejoramiento de la vivienda rural: impacto de la instalación de piso firme y estufas ecológicas en las condiciones de vida de los hogares. Estudios Agrarios. 2009;40: 143-52. Available from: http://www.pa.gob.mx/publica/rev\_40/NOTAS/Ernesto%20Cordero%20 Arroyo pdf
- 30. Mokomane M, Kasvosve I, de Melo E, Pernica JM, Goldfarb DM. The global problem of childhood diarrhoeal diseases: emerging strategies in prevention and management. Ther Adv Infect Dis. 2018;5(1):29-43. https://doi.org/10.1177/2049936117744429
- 31. United Nations International Children's Emergency Fund, World Health Organization. Diarrhoea: Why children are still dying and what can be done. Geneva, Switzerland: WHO/UNICEF, 2009.
- 32. Secretaría de Salud. PROYECTO de Norma Oficial Mexicana PROY-NOM-031-SSA2-2014, Para la atención a la salud de la infancia. Ciudad de México: SSa, 2015.
- 33. Secretaría de Salud. Programa de Acción Específico: Prevención de Enfermedades Diarreaicas Agudas y Cólera 2013-2018. Ciudad de México: SSa. 2014.
- 34. Hernández-Ávila JE, Palacio-Mejía LS, González-González L, García C, Molina-Vélez D, Quezada-Sánchez AD, Salgado AL. Desarrollo de un modelo que combina métodos probabilísticos, geográficos y demográficos para estimar y corregir el subregistro de las defunciones en México. 2018. Available from: http://www.beta.inegi.org.mx/eventos/2018/conacyt/doc/p\_JuanHdz.pdf
- 35. Reyes H, Tomé P, Gutiérrez G, Rodríguez L, Orozco M, Guiscafré H. La mortalidad por enfermedad diarreica en Mexico: ¿problema de acceso o de calidad de atención? Salud Publica Mex. 1998;40(4):316-23. https://doi.org/10.1590/s0036-36341998000400003
- 36. Secretaría de Gobernación, Instituto Nacional de Estadística y Geografía. 3.2.5 Tasa de mortalidad en niños menores de 5 años por enfermedades diarreicas (defunciones por cada 100 mil menores de 5 años). México: Secrertaría de Gobernación, INEGI, 2018.
- 37. Hernández B, Ramírez-Villalobos D, Duarte MB, Corcho A, Villarreal G, Jiménez A, Torres LM. Subregistro de defunciones de menores y certificación de nacimiento en una muestra representativa de los 101 municipios con más bajo índice de desarrollo humano en México. Salud Publica Mex. 2012;54:393-400. https://doi.org/10.1590/s0036-36342012000400009