Stroke in Latin America: Systematic review of incidence, prevalence, and case-fatality in 1997–2021



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Abstract

Background: Stroke is a major global cause of death and disability. Most strokes occur in populations of low-middleincome country (LMIC); therefore, the subsequent disease burden is greater than in populations of high-income countries. Few epidemiological data exist for stroke in Latin America, composed primarily of LMIC.

Aims: To determine epidemiological measures of incidence, prevalence, and 1-month case-fatality for stroke in Latin America/Caribbean (LAC) during 1997–2021.

Summary of review: A structured search was conducted to identify relevant references from MEDLINE, WOS, and LILACS databases for prospective observational and cross-sectional studies in LAC populations from January 1997 to December 2021. A total of 9242 records were screened and 12 selected for analysis, seven incidence studies and five prevalence studies. **[AQ: 2]** Case-fatality rate was reported in six articles. Sub-group analysis by age, sex, and income countries was performed. A narrative synthesis of the findings was performed. Meta-analysis was performed using random-effect model to obtain pooled estimates with 95% confidence intervals (Cls). Studies quality was assessed according to the risk of bias criteria described in the Joanna Briggs Institute's guide. The overall crude annual incidence rate of first-ever stroke in LAC was 119.0 (95% Cl=95.9–142.1)/100,000 people per year (with high heterogeneity between studies (l^2 =98.1%)). The overall crude prevalence was 3060 (95% Cl: 95.9–142.1)/100,000 people (with high heterogeneity between studies (l^2 =98.8%)). The overall case-fatality rate at I month after the first stroke was 21.1% (95% Cl=18.6–23.7) (l^2 =49.40%).

Conclusion: This review contributes to our understanding regarding the burden caused by stroke in LAC. More studies with comparable designs are needed to generate reliable data and should include both standardized criteria, such as the World Health Organization clinical criteria and updated standard methods of case assurance, data collection, and reporting.

Keywords

Stroke, incidence, prevalence, case-fatality, Latin America, systematic review

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Introduction

Stroke is one of the leading global causes of death and disability.¹ Seventy percent of strokes occur in populations of low-middle-income countries (LMICs), and the subsequent burden of disease is higher than in populations of highincome countries.1 Some previous studies of stroke showed that incidence is increasing in low- and middle-income countries, by contrast with high-income countries where a 42% decrease in incidence has taken place in the past four decades.² It is unclear if this steadily declining trend has been sustained in recent years, particularly in view of the continuing aging of the population and the rise of diabetes mellitus and obesity.3 However, between 1990 and 2017, the absolute number of people with incident strokes has increased significantly by 81%, the number of people who survived by 95%, and number of deaths caused by stroke by 40%.²

Latin America, with an estimated population of 680 million inhabitants, has a marked ethnic, cultural, and socioeconomic heterogeneity.⁴ Despite the progress that has been made in recent decades, aging and population growth have led to an increase in the prevalence of cardiovascular risk factors, raising absolute numbers of death and disability due to stroke and cardiovascular diseases.⁵ In these countries, the burden of stroke was 5–14 disability-adjusted life years lost per 1000 population (age standardized to 2002), which is higher than in most high-income countries.⁶

Research about stroke risk factors and epidemiology is based mostly on studies from North America or Europe. Stroke epidemiology has been poorly studied in Latin America/Caribbean (LAC),^{7,8} which is primarily composed of LMIC. The most sensitive tool to study stroke is with standardized population-based registries, since the analysis limited to hospital cases or the variation of criteria and definitions can distort the results.⁹ The methodological differences between case ascertainment, classification, definitions, and reference populations for standardization are some of the problems in the region.¹⁰ Also, environmental, racial, and sociocultural factors can have a significant impact on the epidemiology of stroke in such a wide area.

The aim of this systematic review was to determine epidemiological measures of incidence, prevalence, and 1 month of case-fatality for stroke in LAC during 1997–2021.

Methods

The protocol for the review was registered in PROSPERO (CRD42022325247). A prevalence and incidence systematic review was developed according to the Metaanalysis of Observational Studies in Epidemiology (MOOSE) and Preferred Reporting Items for Systematic reviews and Meta-Analyses (PRISMA) checklist (Supplemental Table S.1).

Selection criteria

This review included prospective observational studies (e.g. community-based and cohort studies) reporting the incidence and case-fatality of stroke in 36 countries of LAC (Supplemental Table S.2). Cross-sectional studies for the assessment of the stroke prevalence were also included.

Study populations had to include adult participants with a confirmed stroke diagnosis, which was defined by the studies according to established clinical criteria, including cerebral infarction, intracerebral hemorrhage (ICH), subarachnoid hemorrhage (SAH), or uncertain pathological subtypes.

Studies using retrospective recruitment (e.g. case series and case–control), reviews, commentaries, and editorials were excluded.

Search strategy

A structured and comprehensive literature search was carried out to identify relevant studies from MEDLINE (Ovid), Lilacs and Science Citation Index Expanded (SCI— EXPANDED), Social Sciences Citation Index and Arts, and Humanities Citation Index within ISI Web of Science (WoS) (from inception to May 27, 2022).

The search strategy was similar to the one published by Jones et al.¹¹ Cochrane stroke strategy terms were combined and filtered by epidemiology studies, including terms for Latin America and Caribbean.^{12,13} Only studies published in English, Portuguese and Spanish from 1 January 1997 to 31 December 2021 were eligible for inclusion (Supplemental Tables S.3 to S.5). We chose 1997 as a starting year for this review as well as the Stroke Unit Trialists' Collaboration¹⁴ systematic review, considering this year as a landmark in global standardization for post-stroke care, a couple of years after the intravenous thrombolysis was proven effective treatment for acute ischemic stroke (IS).

Selection process

Before study selection, references obtained were imported into EndNote 20 and duplicates were removed using the automated function, and then manually by reviewer. All references were imported into JBI SUMARI software (http://www.jbisumari.org/) to carry out the selection process.¹⁵

Two independent reviewers (CD and MN) screened and selected all titles and abstracts potentially relevant. The full texts of these articles were independently assessed for eligibility. In cases of multiple publications based on the same patients' cohort, only those studies including the largest sample size and more complete data were selected to avoid overlapping. Disagreements were resolved by discussion and when it was necessary, a third reviewer was involved to reach consensus.

Outcomes measures

Main outcomes were the cumulative incidence, prevalence per 100,000 people and 30-day case-fatality of stroke in LAC between January 1997 and December 2021.

Data extraction

Two reviewers extracted data (CD and MN) independently using a standardized form piloted on a small number of included studies. Extracted data included: author and year of publication, study period, location, study design, sample size, mean age, and population type. Eligible studies were prepared for synthesis (tabulating the characteristics of each study) (Table 1).

Quality study and risk of bias assessment

The quality of studies was evaluated by two independent reviewers for each study design, using the Joanna Briggs Institute (JBI)²⁸ checklists for cohort or prevalence studies. Subsequently, disagreements were resolved by discussion and when it was necessary, a third reviewer was involved to facilitate consensus (Supplemental Table S.6).

Data synthesis and analysis

For incidence and prevalence rates, results were reported as presented in the original studies with the 95% CI. CIs were calculated if not available. Age-adjusted incidence was reported. Case-fatality data were reported as presented in the original studies, with the percentage of deaths within the first 30 days of the stroke. Analysis by type of stroke was reported in incidence, prevalence, and case-fatality data. The country income groups were classified according to the World Bank's country classification.⁴

Descriptive statistics were used to describe the studies characteristics; mean value with standard deviation (*SD*) for continuous data, and frequency in percentage and its 95% CI, for categorical variables.

The heterogeneity across the studies was assessed through visual inspection of the forest plot, and determining the χ^2 and I^2 statistics, and sub-group analysis. Metaanalysis was performed using *metaprop* command and forest plots were generated to summarize overall estimates. A random-effect model was used to obtain pooled estimates with 95% CIs.

Publication bias was assessed with Egger's test to determine a potential risk of reporting bias. *Metabias* command was formed for each analysis in incidence, prevalence, and case-fatality.

All analysis was performed in STATA v.17.1/SE software (College Station, TX: StataCorp LLC).

Results

Study selection

A total of 9242 articles were identified, of which 1941 duplicates were removed. After reading the titles and abstracts, 116 potential full texts were screened for eligibility. Finally, 12 articles were included.^{16–27} The PRISMA flow chart is shown in Figure 1.

Characteristics of the included studies

The review included 12 articles, seven cohort studies for incidence, 1-month case-fatality analyses,¹⁶⁻²² and five cross-sectional studies for prevalence analysis (Table 1).^{23–27} **[AQ: 3]** The articles included participants with stroke diagnosis confirmed by clinical criteria. Data from Brazil (n=4) followed by Argentina (n=2) and Chile (n=2) were the most frequent, the earliest study period was in 1998–1999,²² and the most recent in 2015–2016 (Figure 2).^{19,23}

The population denominator contemplated in the review was 4,405,227 inhabitants (mean=293,681.8, SD=343,974.2). All countries were classified as upper-middle income, except Chile²¹ and Martinique²² that were high-income countries according to the World Bank's classification.⁴ Information on the ethnic composition of the population could not be obtained.

Incidence reports

From seven articles that reported incidence, three of them reported two registration periods.^{18,19,22} Therefore, ten study periods were identified (Table 2). We identified a total number of 4835 patients with first-ever stroke in a sample of 4,369,876 persons, from which 2390 were women (49.8%). The mean age of the study periods that reported age data was 69.0 ± 3.5 years.^{16–18,20–22} Type of stroke was classified by computed tomography (CT) and/or magnetic resonance imaging (MRI). The mean of the percentage of brain images performed was 91.7% and ranged from 90%²¹ to 99.4%.²²

The crude annual incidence rate of first-ever stroke was heterogeneous, ranging from $73.6/100,000^{20}$ to $180.4/100,000^{21}$ people per year (Supplemental Figure S.1). The overall crude annual incidence rate of first-ever stroke in LAC was 119.0 (95% CI=95.9–142.1)/100,000 people per year (Figure 3). High heterogeneity was observed (I^2 =98.1%). The age-adjusted incidence reported on the basis of Segi's WORLD ranged from 77/100,000 in Martinique²² to 254.1/100,000 in Nuble, Chile.²¹

According to the stroke subtype, the crude annual incidence of first-ever IS rate ranged from 46.6^{20} to $127.8/100,000^{22}$ people per year, ICH rate ranged from 9.5^{18} to $22.9/100,000^{22}$ people per year, and SAH rate ranged from 1.3^{19} to $9.5/100,000^{21}$ people per year. High heterogeneity was observed in these analyses with I^2 of 97.6%, 91.8%, and 66.7%, respectively. The LAC overall crude

Location	Study period	Study registry design	Duration of surveillance (months)	Frequency of surveillance	Approach tool	Income level (according to the World Bank's classification)
Incidence and case-fat	ality studies					
I. Argentina (Tandil) ¹⁶	5 January 2013–30 April 2015	Overlapping sources	28	Continuous	Sudlow and Warlow criteria. WHO STEPS	Upper-middle
2. Barbados 17	15 October 2001–15 October 2002	Overlapping sources	12	Continuous	South London Stroke Register	Upper-middle
3. Brazil (Joinville) ¹⁸	1995 2005–2006 2012–2013	Overlapping sources	12	Continuous	WHO STEPS	Upper-middle
4. Brazil (Matao) ¹⁹	l November 2003–31 October 2004 l August 2015–31 July 2016	Overlapping sources	12	Continuous in each period	WHO definition	Upper-middle
5. Chile (Iquique) ²⁰	I July 2000–31 June 2001	Overlapping sources	24	Continuous	Sudlow and Warlow criteria	Upper-middle
6. Chile (Ñuble) ²¹	1 April 2015–30 March 2016	Overlapping sources	12	Continuous	WHO STEPS stroke	High income
7. Martinique ²²	June 1998–31 May 1999 November 2011–31 October 2012	Overlapping sources	12	Continuous	WHO	High income
Prevalence studies						
8. Argentina (General Villegas) ²³	1 August 2015–31 July 2016	Questionnaire	12	Annual	WHO STEP	Upper-middle
9. Brazil (Coari) ²⁴	May–October 2011	Questionnaire	5	Continuous	WHO STEPS	Upper Middle
10. Brazil (Sao Jorge) ²⁵	February–June 2008	Questionnaire	4	Annual	WHO STEPS	Upper-middle
II. Peru (Tumbes) ²⁶	April 2011–May 2012	Questionnaire	12	Continuous	WHO STEPS	Upper-middle
12. Suriname (Paramaribo) ²⁷	April 2013–August 2013	Questionnaire	5	Continuous	NS	Upper-middle

Table I. A summary of the characteristics of studies assessing the incidence, prevalence and fatality of stroke in Latin America and the Caribbean.

annual incidence was 91.8 (95% CI=72.5–111.1)/100,000 people per year, 16.6 (95% CI=12.2–21.1)/100,000 people per year and 4.9 (95% CI=3.7–6.1)/100,000 people per year for IS, ICH, and SAH, respectively (Figure 3).

Regarding age groups, the crude annual incidence rate of first-ever stroke was 3.8 (2.8–4.9)/100,000 people per year in the 0- to 34-year-old group (I^2 =32.6), 33.8 (25.8–41.8)/100,000 people per year in the 35- to 44-year-old group (I^2 =62.8), 127.8 (101.6–154.0)/100,000 people per year in the 45- to 54-year-old group (I^2 =83.9), 252.8 (210.9–294.7)/100,000 people per year in the 55- to

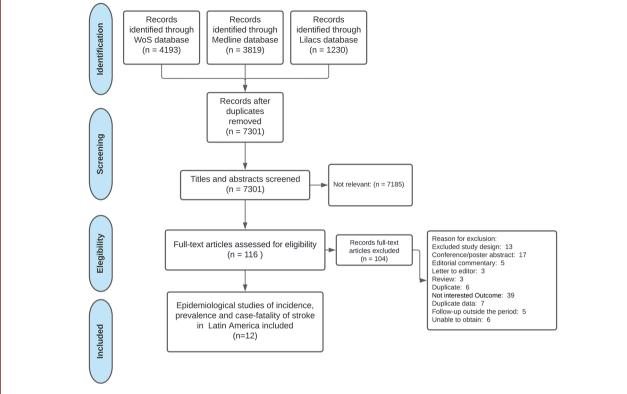
64-year-old group (I^2 =81.9), 580.3 (504.6–656.0)/100,000 people per year in the 65- to 74-year-old group (I^2 =78.5) and 1202.3 (1007.4–1391.1)/100,000 people per year in the 75 years old or older group (I^2 =91.2) (Supplemental Figure S.2)

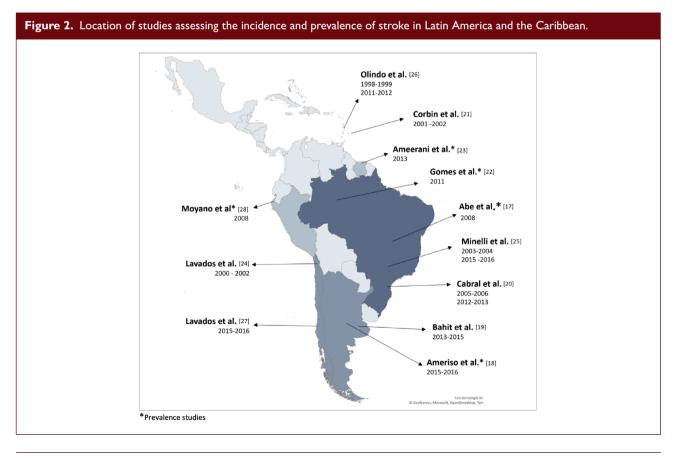
Prevalence reports

Five studies reported prevalence rates (Table 3).^{25,27} Of a total of 35,342 persons, 684 respondents were classified as stroke victims.

Figure 1. The PRISMA flow diagram.







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Study and publication year	Country (City)	Periods	Study period	No. Of cases of stroke identified	Population denominator	Mean (SD) age (years)	Female, N (%)	CT available (%)	Stroke type N (%)	Crude annual incidence/100,000, N (95% CI)	Sex-disaggregated incidence rate female/10 ⁵ , N (95% CI)	Sex-disaggregated incidence rate male/ 10 ⁵ , N (95% Cl)	Age-adjusted incidence/10 ⁵ , N (95% C1)*	One-month case-fatality (95% CI)
I. Bahit et al. ¹⁶	Argentina (Tandil)	-	2013-2015	334	261,182	72.2 (14.4)	175 (52.4)	1.76	IS 251 (75) ICH 54 (16) SAH 17 (5)	127.9 (114.5–142.3)	129.8 (111.2–150.5)	125.9 (61 <i>.7–</i> 86.1)	SN	20.1 (15.9–24.8)
2. Corbin et al. ¹⁷	Barbados	-	2001–2002	352	26,8792	72.5 (14.8)	210 (59.7)	92.6	IS, 288 (82) ICH. 42 (12) SAH, 7 (2)	131.0 (117.6–145.4)	163.2 (141.5–187.2) [*]	I I 5.0 (96.3–I 36.4) [*]	SN	SZ
3. Cabral et al. ¹⁸	Brazil (Joinville)	-	2005-2006	759	987,619	64.9 (13.9)	367 (48.4)	98.0	IS, 610 (80) ICH, 94 (12) SAH, 50 (7)	76.9 (71.5–82.5)	73.5 (66.2–81.4)	80.2 (72.5–88.5)	105.4 (98–113.2)	19.1 (16.0–22.4)
		2	2012-2013	922	1,073,318	63.7 (15.5)	450 (48.8)	0.66	IS, 786 (85) ICH, 84 (9) SAH, 52 (7)	85.9 (80.4–91.6)	83.3 (75.8–91.4)	88.6 (80.8–97.0)	90.9 (85.1–96.9)	SZ
4. Minelli et al. ¹⁹	Brazil (Matao)	-	2003–2004	8	76,786	65.2 (11.8)	30 (37.0)	S	IS, 69 (85) ICH, I1 (14) SAH, I (1)	105.5 (83.8–131.1)	66.4 (49.4–87.3)	39.0 (26.3–55.7)	SN	18.5 (10.8–28.7)
		7	2015–2016	8	78,890	71 (13.1)	40 (50.6)	SS	IS, 67 (83) ICH, 10 (12) SAH, 4 (5)	102.7 (81.5–127.6)	50.7 (36.2–69.0)	51.9 (37.3–70.5)	SN	17.3 (9.8–27.3)
5. Lavados et al. ²⁰	Chile (Iquique)	-	2000–2002	292	396,712	64.8 (5.1)	128 (44)	0.16	IS, 185 (63) ICH, 69 (24) SAH, 15 (5)	73.6 (65.4–82.5)	65.3 (53.9–76.6)	81.8 (69.3–94.3)	94.1 (83.3–104.9)	23.3 (18.6–28.6)
6. Lavados et al. ²¹	Chile (Ñuble)	-	2015–2016	890	49,3464	70.3 (14.1)	443 (49.8)	0.06	IS 639 (72) HIC 111 (12) SAH 47 (5)	180.4 (168.7–192.6)	175.2 (159.3–192.3)	185.8 (168.9–203.8)	254.1 (237.4–271.1)	24.6 (21.8–27.6)
7. Olindo et al. ²²	Martinique	-	1998–1999	580	362,259	71.2 (14)	295 (50.9)	92.8	IS, 463 (80) HIC, 83 (14) SAH, 20 (3)	160.1 (147.3–173.7)	154.9 (138–182)	165.8 (146–186)	111 (102–120)	SN
		7	2011–2012	544	370,854	72 (15)	252 (46.3)	99.4	IS, 439 (81) HIC, 84 (15) SAH, 14 (3)	146.7 (134.6–159.5)	125.7 (110–141)	171.2 (152–190	77 (70–84)	SN

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Fi	g. 3.1 The overall crude annual incider	nce rate					
10) Events	Total		te [95% CI]	SE (95%CI)	Weight
1	Argentina, Tandil (2013-2015)	334			[114.5-142.3]		10.01
2	2 Barbados (2001-2002)	352	268792	131.0	[117.6-145.4]		10.01
3		759	987619		[71.5-82.5]	+	10.32
4		922	1073318		[80.4-91.6]	+	10.32
5		81			[83.8-131.1]		9.41
6		81			[81.5-127.6]		9.46
7		292	396712		[65.4-82.5]	+	10.24
8		890			[168.7-192.6]	\rightarrow	10.10
2		580			[147.3-173.7]		10.05
1	0 Martinique (2011-2012) Total (95% CI)	544			[134.6-159.5]		10.08 100.00
		4025	4303870	119.0	[95.9-142.1]		100.00
	Total events Heterogenity: X ² = 481.22, d.f.=9	4835	- 08 120%		Estimate of het	ween-study variance T ² = 0.000	
F	g. 3.2 Ischemic stroke	(p= 0.000), 1	= 30.12376		estimate of bet	ween-study variance 1 = 0.000	
10) Events	Total	Ra	ite [95% CI]	SE (95%CI)	Weight
1	Argentina, Tandil (2013-2015)	251	261182	96.1	[84.6-108.7]		10.04
2	2 Barbados (2001-2002)	288	268792	107.1	[95.1-120.3]		10.01
3	Brazil, Joinville (2005-2006)	610	987619	61.7	[57.0-66.9]	+	10.41
4	Brazil, Joinville (2012-2013)	786	1073318	73.2	[68.2-78.5]	+	10.40
5	Brazil, Matao (2003-2004)	69	76786	89.8	[69.9-113.7]	- _	9.19
e	5 Brazil, Matao (2015-2016)	67	78890	84.9	[65.8-107.8]		9.28
7	Chile, Iquique (2000-2001)	185	396712	46.6	[40.2-53.9]	+	10.34
8	6 Chile, Ñuble (2015-2016)	639	493464	129.4	[119.7-139.9]	-	10.17
9	Martinique (1998-1999)	463	362259	127.8	[116.4-140.0]		10.06
1	0 Martinique (2011-2012)	439	370854	118.3	[107.6-130.0]		10.10
	Total (95% CI)		4369876	91.8	[72.5-111.1]		100.00
	Total events	3797				.001	
_	Heterogenity: X ² = 383.84, d.f.=9	(p= 0.000); 1	= 97.65%		Estimate of bet	ween-study variance T ² = 0.000	
_	g. 3.3 Intracerebral Hemorrhage		T + 1			55 (054(0))	
10	 Country, Location (study period) Argentina, Tandil (2013-2015)) Events 54	Total	Ra	ite [95% CI]	SE (95%CI)	Weight
	Algentina, fanun (2013-2013)		261102	20.7	[15 5-27 0]		
	Barbados (2001-2002)		261182		[15.5-27.0]		9.69
	Barbados (2001-2002)	42	268792	15.6	[11.3-21.1]		9.69 10.16
3	Brazil, Joinville (2005-2006)	42 94	268792 987619	15.6 9.5	[11.3-21.1] [7.7-11.6]		9.69 10.16 11.45
3	Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013)	42 94 84	268792 987619 1073318	15.6 9.5 7.8	[11.3-21.1] [7.7-11.6] [6.2-9.7]		9.69 10.16 11.45 11.52
3	Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Matao (2003-2004)	42 94 84 11	268792 987619 1073318 76786	15.6 9.5 7.8 14.3	[11.3-21.1] [7.7-11.6] [6.2-9.7] [7.2-25.6]		9.69 10.16 11.45 11.52 7.84
3 4 5	Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Matao (2003-2004) Brazil, Matao (2015-2016)	42 94 84 11 10	268792 987619 1073318 76786 78890	15.6 9.5 7.8 14.3 12.7	[11.3-21.1] [7.7-11.6] [6.2-9.7] [7.2-25.6] [6.1-23.3]		9.69 10.16 11.45 11.52 7.84 8.22
3 4 5 6 7	8 Brazil, Joinville (2005-2006) 8 Brazil, Joinville (2012-2013) 5 Brazil, Matao (2003-2004) 5 Brazil, Matao (2015-2016) 7 Chile, Iquique (2000-2001)	42 94 84 11 10 69	268792 987619 1073318 76786 78890 396712	15.6 9.5 7.8 14.3 12.7 17.4	[11.3-21.1] [7.7-11.6] [6.2-9.7] [7.2-25.6] [6.1-23.3] [13.5-22.0]		9.69 10.16 11.45 11.52 7.84 8.22 10.51
3 4 5 7 8	 Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Matao (2003-2004) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Ñuble (2015-2016) 	42 94 84 11 10 69 111	268792 987619 1073318 76786 78890 396712 493464	15.6 9.5 7.8 14.3 12.7 17.4 22.5	[11.3-21.1] [7.7-11.6] [6.2-9.7] [7.2-25.6] [6.1-23.3] [13.5-22.0] [18.5-27.1]		9.69 10.16 11.45 11.52 7.84 8.22 10.51 10.47
3 4 5 6 7 7 8 8 9	 Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Matao (2003-2004) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Ñuble (2015-2016) Martinique (1998-1999) 	42 94 84 11 10 69 111 83	268792 987619 1073318 76786 78890 396712 493464 362259	15.6 9.5 7.8 14.3 12.7 17.4 22.5 22.9	[11.3-21.1] [7.7-11.6] [6.2-9.7] [7.2-25.6] [6.1-23.3] [13.5-22.0] [18.5-27.1] [18.2-28.4]		9.69 10.16 11.45 11.52 7.84 8.22 10.51 10.47 10.05
3 4 5 6 7 7 8 8 9	Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Joinville (2013-2004) Brazil, Matao (2003-2004) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Ruigue (2015-2016) Martinique (1998-1999) Martinique (2011-2012)	42 94 84 11 10 69 111	268792 987619 1073318 76786 78890 396712 493464 362259 370854	15.6 9.5 7.8 14.3 12.7 17.4 22.5 22.9 22.7	[11.3-21.1] [7.7-11.6] [6.2-9.7] [7.2-25.6] [6.1-23.3] [13.5-22.0] [18.5-27.1] [18.2-28.4] [18.1-28.0]		9.69 10.16 11.45 11.52 7.84 8.22 10.51 10.47 10.05 10.10
3 4 5 6 7 7 8 8 9	Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Matao (2003-2004) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Nuble (2015-2016) Martinique (1998-1999) Martinique (2011-2012) Total (95% CI)	42 94 84 11 10 69 111 83 83	268792 987619 1073318 76786 78890 396712 493464 362259	15.6 9.5 7.8 14.3 12.7 17.4 22.5 22.9 22.7	[11.3-21.1] [7.7-11.6] [6.2-9.7] [7.2-25.6] [6.1-23.3] [13.5-22.0] [18.5-27.1] [18.2-28.4]		9.69 10.16 11.45 11.52 7.84 8.22 10.51 10.47 10.05
3 4 5 6 7 7 8 8 9	Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Matao (2003-2004) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Ñuble (2015-2016) Martinique (1998-1999) Martinique (2011-2012) Total (95% CI) Total events	42 94 84 11 10 69 111 83 84 642	268792 987619 1073318 76786 78890 396712 493464 362259 370854 4369876	15.6 9.5 7.8 14.3 12.7 17.4 22.5 22.9 22.7	[11.3-21.1] [7.7-11.6] [6.2-9.7] [7.2-25.6] [6.1-23.3] [13.5-22.0] [18.5-27.1] [18.2-28.4] [18.1-28.0] [12.2-21.1]	vween-study variance T ² = 0.000	9.69 10.16 11.45 11.52 7.84 8.22 10.51 10.47 10.05 10.10
3 4 5 6 7 7 8 9 1	Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Matao (2003-2004) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Nuble (2015-2016) Martinique (1998-1999) Martinique (2011-2012) Total (95% CI)	42 94 84 11 10 69 111 83 84 642	268792 987619 1073318 76786 78890 396712 493464 362259 370854 4369876	15.6 9.5 7.8 14.3 12.7 17.4 22.5 22.9 22.7	[11.3-21.1] [7.7-11.6] [6.2-9.7] [7.2-25.6] [6.1-23.3] [13.5-22.0] [18.5-27.1] [18.2-28.4] [18.1-28.0] [12.2-21.1]	eveen-study variance T ² = 0.000	9.69 10.16 11.45 11.52 7.84 8.22 10.51 10.47 10.05 10.10
3 4 5 6 7 8 9 1 1	Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Matao (2003-2004) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Nuble (2015-2016) Martinique (1998-1999) O Martinique (2011-2012) Total (95% CI) Total events Heterogenity: X ² = 110.48, d.f.=9 g. 3.4 Subarachnoid Hemorrhage	42 94 84 11 10 69 111 83 84 642 (p= 0.000); I ^s	268792 987619 1073318 76786 78890 396712 493464 362259 370854 4369876 2 = 91.85%	15.6 9.5 7.8 14.3 12.7 17.4 22.5 22.9 22.7 16.6	[11.3-21.1] [7.7-11.6] [6.2-9.7] [7.2-25.6] [6.1-23.3] [13.5-22.0] [18.5-27.1] [18.2-28.4] [18.1-28.0] [12.2-21.1] 		9.69 10.16 11.45 11.52 7.84 8.22 10.51 10.47 10.05 10.10 100.00
3 4 5 6 7 8 9 1 1 1 1 1 1 1	Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Matao (2003-2004) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Ruble (2015-2016) Martinique (1998-1999) Martinique (2011-2012) Total (95% CI) Total events Heterogenity: X ² = 110.48, d.f.=9 S.4.5 Ubarachnold Hemorrhage Country, Location (study period)	42 94 84 11 10 69 111 83 84 (p= 0.000); F (p= 0.000); F	268792 987619 1073318 76786 78890 396712 493464 362259 370854 4369876 * = 91.85%	15.6 9.5 7.8 14.3 12.7 17.4 22.5 22.9 22.7 16.6	[11.3-21.1] [7.7-11.6] [6.2-9.7] [7.2-25.6] [6.1-23.3] [13.5-22.0] [18.5-27.1] [18.2-28.4] [18.1-28.0] [12.2-21.1] 	ween-study variance T ² = 0.000	9.69 10.16 11.45 11.52 7.84 8.22 10.51 10.47 10.05 10.10 100.00
3 4 5 7 8 5 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1	Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Matao (2003-2004) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Nuble (2015-2016) Martinique (1998-1999) Martinique (2011-2012) Total (95% CI) Total events Heterogenity: X ² = 110.48, d.f.=9 Country, Location (study period) Argentina, Tandil (2013-2015)	42 94 84 11 00 69 111 83 84 (p= 0.000); 12 (p= 0.000); 13) Events 17	268792 987619 1073318 76786 78890 396712 493464 362259 370854 4369876 * = 91.85%	15.6 9.5 7.8 14.3 12.7 17.4 22.5 22.9 22.7 16.6 Ra 6.5	[11.3-21.1] [7.7-11.6] [6.2-9.7] [7.2-25.6] [6.1-23.3] [13.5-22.0] [18.5-27.1] [18.2-28.4] [18.1-28.0] [12.2-21.1] 	ween-study variance T ² = 0.000	9.69 10.16 11.45 11.52 7.84 8.22 10.51 10.47 10.05 10.10 100.00 Weight 7.56
3 4 5 7 8 5 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1	Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Matao (2003-2004) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Nuble (2015-2016) Martinique (1998-1999) Martinique (2011-2012) Total (95% CI) Total events Heterogenity: X ² = 110.48, d.f.=9 O Courtry, Location (study period) Argentina, Tandil (2013-2015) Barbados (2001-2002)	42 94 84 11 10 69 111 83 84 (p= 0.000); I ² (p= 0.000); I ³ (p= 0.000); I ³ 7 7	268792 987619 1073318 76786 78890 36259 370854 4362259 370854 4369259 370854 4369876	15.6 9.5 7.8 14.3 12.7 17.4 22.5 22.9 22.7 16.6 Ra 6.5 2.6	[11.3-21.1] [7.7-11.6] [6.2-9.7] [7.2-25.6] [6.1-23.3] [13.5-22.0] [18.5-27.1] [18.2-28.4] [18.1-28.0] [12.2-21.1] [18.1-28.0] [12.2-21.1] [18.5-27.1] [18.1-28.0]	ween-study variance T ² = 0.000	9.69 10.16 11.45 11.52 7.84 8.22 10.51 10.47 10.05 10.10 100.00 Weight 7.56 11.33
3 4 5 7 8 5 1 1 1 1 1 1 1 1 1 1 2	Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Matao (2003-2004) Brazil, Matao (2003-2004) Brazil, Matao (2003-2004) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Rubique (2015-2016) Martinique (1998-1999) Martinique (2011-2012) Total (95% CI) Total events Heterogenity: X ² = 110.48, d.f.=9 Ocountry, Location (study period) Argentina, Tandil (2013-2015) Barbados (2001-2002) Barazil, Joinville (2005-2006)	42 94 84 11 10 69 111 83 84 (p= 0.000); I ² (p= 0.000); I ² 7 7 50	268792 987619 1073318 76786 78890 396712 493464 362259 370854 4369876 * = 91.85% * = 91.85% * = 91.85%	15.6 9.5 7.8 14.3 12.7 17.4 22.5 22.9 22.7 16.6	[11.3-21.1] [7.7-11.6] [6.2-9.7] [7.2-25.6] [6.1-23.3] [13.5-22.0] [18.5-27.1] [18.2-28.4] [18.1-28.0] [12.2-21.1] [18.1-28.0] [12.2-21.1] [18.5-25.6] [19.5.6] [10.5.4] [10.5.4] [10.5.4] [10.5.4]	ween-study variance T ² = 0.000	9.69 10.16 11.45 11.52 7.84 8.22 10.51 10.47 10.05 10.10 100.00 Weight 7.56 11.33 13.23
3 4 5 7 8 9 1 1 1 1 1 1 1 1 1 2 3	Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Matao (2003-2004) Brazil, Matao (2003-2004) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Ñuble (2015-2016) Martinique (1998-1999) Martinique (2011-2012) Total (95% CI) Total events Heterogenity: X ² = 110.48, d.f.=9 Country, Location (study period) Argentina, Tandil (2013-2015) Barbados (2001-2002) Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013)	42 94 84 11 10 69 111 83 84 (p= 0.000); F 17 7 50 52	268792 987619 1073318 76786 78890 396712 493464 362259 370854 4369876 * * 91.85% * * * * * * * * * * * * * * * * * * *	15.6 9.5 7.8 14.3 12.7 17.4 22.5 22.9 22.7 16.6 8 6.5 2.6 5.0 4.8	[11.3-21.1] [7,7-11.6] [6,2-9,7] [7,2-25.6] [6,1-23.3] [13.5-22.0] [18.5-27.1] [18.2-28.4] [18.1-28.0] [12.2-21.1] [12.2-21.1] [3.8-10.4] [3.8-10.4] [3.8-6.7] [3.8-6.4]	ween-study variance T ² = 0.000	9.69 10.16 11.45 11.52 7.84 8.22 10.51 10.47 10.05 10.10 100.00 Weight 7.56 11.33
3 4 5 7 8 8 9 9 9 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1	Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Matao (2003-2004) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Nuble (2015-2016) Martinique (1998-1999) Martinique (2011-2012) Total (95% CI) Total events Heterogenity: X ² = 110.48, d.f.=9 Country, Location (study period) Argentina, Tandil (2013-2015) Barbados (2001-2002) Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Joinville (2003-2004)	42 94 84 11 10 69 111 83 84 (p= 0.000); I ² (p= 0.000); I ² 7 7 50	268792 987619 1073318 76786 78890 396712 493464 362259 370854 4369876 2 91.85% 2 100 261182 266182 266792 987619 987619 1073318	15.6 9.5 7.8 14.3 12.7 17.4 22.5 22.9 22.7 16.6 8 5.0 4.8 1.3	[11.3-21.1] [7,7-11.6] [6,2-9,7] [7,2-25.6] [6,1-23.3] [13.5-22.0] [18.5-27.1] [18.5-27.1] [18.2-28.4] [18.2-28.4] [12.2-21.1] [12.2-21.1] [12.2-21.1] [12.2-21.1] [13.8-6.7] [3.8-6.7] [3.8-6.4] [0.0-7.3]	ween-study variance T ² = 0.000	9,69 10.16 11.45 11.52 7,84 8,22 10.51 10.47 10.05 10.10 100.00 Weight 7,56 11.33 13.23 13.53
3 4 5 6 6 7 7 7 8 8 5 5 1 1 1 1 1 1 1 2 3 8 4 5 5 6 6 6 6 6 7 7 7 7 7 7 7 7 7 7 7 7 7	Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Joinville (2012-2014) Brazil, Matao (2003-2004) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Iquique (2005-2016) Martinique (1998-1999) Martinique (2011-2012) Total (95% Cf) Total (95% Cf) Total vents Heterogenity: X ² = 110.48, d.f.=9 Country, Location (study period) Argentina, Tandil (2013-2015) Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Joinville (2012-2013) Brazil, Joinville (2012-2013) Brazil, Joinvilla (2013-2004) Brazil, Matao (2015-2016)	42 94 84 11 0 69 111 83 84 (p= 0.000); 1 (p= 0.000); 1 7 7 50 52 1 4	268792 987619 1073318 76786 78890 396712 493464 362259 370854 4369876 2493464 362259 370854 4369876 261182 261182 268792 987619 1073318 76786 78890	15.6 9.5 7.8 14.3 12.7 17.4 22.5 22.9 22.7 16.6 8 8 6.5 2.6 5.0 4.8 1.3 5.0	[11.3-21.1] [7.7-11.6] [6.2-9.7] [7.2-25.6] [6.1-23.3] [13.5-22.0] [18.5-27.1] [18.2-28.4] [18.2-28.4] [18.2-28.4] [12.2-21.1] [12.2-21.1] [12.2-21.1] [13.8-10.4] [1.0-5.4] [3.8-6.7] [3.8-6.7] [3.8-6.4] [0.0-7.3] [1.4-13.0]	ween-study variance T ² = 0.000	9.69 10.16 11.45 11.52 7.84 8.22 10.51 10.47 10.05 10.10 100.00 Weight 7.56 11.33 13.23 13.53 9.22
3 4 5 6 6 7 7 7 8 8 5 5 1 1 1 1 1 1 1 2 3 8 4 5 5 6 6 6 6 6 7 7 7 7 7 7 7 7 7 7 7 7 7	Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Matao (2003-2004) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Ruble (2015-2016) Martinique (1998-1999) Martinique (2011-2012) Total (95% CI) Total events Heterogenity: X ² = 110.48, d.f.=9 Country, Location (study period) Argentina, Tandil (2013-2015) Barbados (2001-2002) Brazil, Joinville (2005-2006) Brazil, Joinville (2003-2004) Brazil, Matao (2013-2013) Brazil, Joinville (2012-2013) Brazil, Joinville (2012-2013) Brazil, Joinville (2012-2013)	42 94 84 11 0 69 111 83 84 (p= 0.000); 17 7 50 52 1 4 52 1 4	268792 987619 1073318 76786 78890 396712 493464 362259 370854 4369876 * * 91.85% * * * * * * * * * * * * * * * * * * *	15.6 9.5 7.8 14.3 12.7 17.4 22.5 22.9 22.7 16.6 5.0 4.8 1.3 5.0 3.7	[11.3-21.1] [7.7-11.6] [6.2-9.7] [7.2-25.6] [6.1-23.3] [13.5-22.0] [18.5-27.1] [18.2-28.4]	ween-study variance T ² = 0.000	9.69 10.16 11.45 11.52 7.84 8.22 10.51 10.47 10.05 10.10 100.00 Weight 7.56 11.33 13.23 13.53 13.23 13.53 9.22 4.17 11.38
3 6 6 7 7 8 8 9 9 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1	Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Matao (2003-2004) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Nuble (2015-2016) Martinique (1998-1999) Martinique (2011-2012) Total (95% CI) Total events Heterogenity: X ² = 110.48, d.f.=9 Ocourtry, Location (study period) Argentina, Tandil (2013-2015) Barbados (2001-2002) Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Matao (2013-2014) Brazil, Matao (2015-2016) Chile, Ruique (2000-2001) Grazil, Matao (2015-2016)	42 94 84 11 00 69 111 83 84 (p= 0.000); H (p= 0.000); H (p	268792 987619 1073318 76786 78890 36725 370854 4362259 370854 4368276 370854 4369876 261182 268792 987619 1073318 7073318 78890 396712 493464	15.6 9.5 7.8 14.3 12.7 17.4 22.5 22.9 22.7 16.6 5.0 8.5 5.0 4.8 1.3 5.0 3.7 9.5	[11.3-21.1] [7.7-11.6] [6.2-9.7] [7.2-25.6] [6.1-23.3] [13.5-22.0] [18.5-27.1] [18.2-28.4] [18.1-28.0] [12.2-21.1] [18.2-28.4]	ween-study variance T ² = 0.000	9.69 10.16 11.45 11.52 7.84 8.22 10.51 10.47 10.05 10.10 100.00 Weight 7.56 11.33 13.23 13.53 9.22 4.17 11.38 8.70
3 5 6 7 7 8 8 5 5 5 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1	Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Matao (2003-2004) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Ruidue (2015-2016) Martinique (1998-1999) Martinique (2011-2012) Total (95% CI) Total events Heterogenity: X ² = 110.48, d.f.=9 Stabarachnoid Hemorrhage O Country, Location (study period) Brazil, Joinville (2012-2013) Brazil, Joinville (2012-2004) Brazil, Joinville (2012-2013) Brazil, Matao (2015-2016) Chile, Řubie (2015-2016) Chile, Ruique (2000-2001)	42 94 84 11 10 69 111 83 84 (p= 0.000); H (p= 0.000); H (p	268792 987619 1073318 76786 78890 396712 493464 362259 370854 4369259 370854 4369876 261182 268792 987619 1073318 76786 78890 396712 493464 362259	15.6 9.5 7.8 14.3 12.7 17.4 22.5 22.9 22.7 16.6 8 5.0 4.8 1.3 5.0 3.7 9.5 5.5	[11.3-21.1] [7,7-11.6] [6,2-9,7] [7,2-25.6] [6,1-23.3] [13.5-22.0] [18.5-27.1] [18.2-28.4] [18.1-28.0] [12.2-21.1] [18.2-28.4] [19.2-28.4]	ween-study variance T ² = 0.000	9.69 10.16 11.45 11.52 7.84 8.22 10.51 10.47 10.05 10.10 100.00 Weight 7.56 11.33 13.23 13.53 9.22 4.17 11.38 8.8.70 9.64
3 5 6 7 7 8 8 5 5 5 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1	Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Matao (2003-2004) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Nuble (2015-2016) Martinique (1998-1999) Martinique (2011-2012) Total (95% CI) Total events Heterogenity: X ² = 110.48, d.f.=9 Ocourtry, Location (study period) Argentina, Tandil (2013-2015) Barbados (2001-2002) Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Matao (2013-2014) Brazil, Matao (2015-2016) Chile, Ruique (2000-2001) Grazil, Matao (2015-2016)	42 94 84 11 00 69 111 83 84 (p= 0.000); H (p= 0.000); H (p	268792 987619 1073318 76786 78890 36725 370854 4362259 370854 4368276 370854 4369876 261182 268792 987619 1073318 7073318 78890 396712 493464	15.6 9.5 7.8 14.3 12.7 22.5 22.9 22.7 16.6 5.0 4.8 1.3 5.0 4.8 1.3 5.0 3.7 9.5 5.5 3.7	[11.3-21.1] [7.7-11.6] [6.2-9.7] [7.2-25.6] [6.1-23.3] [13.5-22.0] [18.5-27.1] [18.2-28.4] [18.1-28.0] [12.2-21.1] [18.2-28.4]	ween-study variance T ² = 0.000	9.69 10.16 11.45 11.52 7.84 8.22 10.51 10.47 10.05 10.10 100.00 Weight 7.56 11.33 13.23 13.53 9.22 4.17 11.38 8.70
3 5 6 7 7 8 8 5 5 5 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1	Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Matao (2003-2004) Brazil, Matao (2003-2004) Brazil, Matao (2003-2004) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Nuble (2015-2016) Martinique (1998-1999) Martinique (2011-2012) Total events Heterogenity: X ² = 110.48, d.f.=9 Country, Location (study period) Aratoini (2013-2015) Barabados (2001-2002) Brazil, Joinville (2005-2006) Brazil, Joinville (2012-2013) Brazil, Joinville (2012-2013) Brazil, Joinville (2015-2016) Chile, Ruique (200-2001) Brazil, Joinville (2015-2016) Chile, Ruique (2015-2016) Chile, Ruique (2015-2016) Martinique (1998-1999) Martinique (1998-1999)	42 94 84 11 10 69 111 83 84 (p= 0.000); H (p= 0.000); H (p	268792 987619 1073318 76786 78890 396712 493464 362259 370854 4369876 26182 268792 987619 987619 987619 1073318 76786 78890 396712 493464 362259 370854	15.6 9.5 7.8 14.3 12.7 22.5 22.9 22.7 16.6 5.0 4.8 1.3 5.0 4.8 1.3 5.0 3.7 9.5 5.5 3.7	[11.3-21.1] [7,7-11.6] [6,2-9,7] [7,2-25.6] [6,1-23.3] [13.5-22.0] [18,5-27.1] [18,5-27.1] [18,2-28.4] [18,1-28.0] [12,2-21.1] [2,2-21.1] [1,0-5,4] [3,8-6,7] [3,8-6,7] [3,8-6,7] [3,8-6,7] [3,8-6,7] [3,8-6,7] [3,8-6,7] [3,8-6,7] [3,8-6,7] [3,8-6,7] [3,8-6,7] [3,8-6,7] [3,8-6,7] [3,8-6,7] [3,8-6,7] [3,8-6,7] [3,4-8,5] [3,4-8,5]	ween-study variance T ² = 0.000	9.69 10.16 11.45 11.52 7.84 8.22 10.51 10.47 10.05 10.10 100.00 Weight 7.56 11.33 13.23 13.53 9.22 4.17 11.38 8.870 9.64 11.15

The crude prevalence rate ranged from 704.4/100,000 to 5271.7/100,000 people per year.^{25,26} High data heterogeneity was found (I^2 =98.8%). The overall LAC crude prevalence was 3060.1 (95% CI=1090.6–5023.6)/100,000 people (Figure 4).

There was not enough information about stroke subtype

prevalence.

One-month case-fatality reports

Six registry periods reported 1-month case-fatality rates.^{16,19–21} A moderate heterogeneity was observed between studies (I^2 =49.4%). The overall case-fatality rate at 1 month after the first stroke was 21.1% (95% CI=18.6–23.7). The reported 1-month case-fatality ratio ranged from 17.3%

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	•										
Study and publication year	Country (city)	Study period	No. of cases of stroke identified	Population denominator	Mean (SD) Female, N age (years) (%)	Female, <i>N</i> (%)	Image available (%)	Stroke type, N (%)	Crude prevalence/100,000, N (95 % CI)	Sex-disaggregated prevalence rate female/100,000 (95% C1)	Sex-disaggregated prevalence rate male/100,000 (95% CI)
I. Ameriso et al. ²³	Argentina (General Villegas)	2015-2016	41	2000	68.9 (9)	14 (34.1) 95	95	IS, 78% ICH, 17%	2050 (1475–2770)	2050 (1475–2770) 13,72.5 (752–2292)	27,55.1 (1823–3983)
2. Fernandes et al. ²⁴ Brazil (Coari)	* Brazil (Coari)	2011	246	5925	57.8 (4)	141 (57.3) NS	NS	NS	4151.8 (3658-4691)	4151.8 (3658-4691) 4555.7 (3848-5350) 3710.2 (3044-4473)	3710.2 (3044-4473)
3. Abe et al. ²⁵	Brazil (Sao Jorge)	2008	193	3661	NS	123 (63.7) NS	NS	NS	5271.7 (4570–6045)	5271.7 (4570–6045) 5444.8 (4545–6462) 4992.8 (3912–6266)	4992.8 (3912–6266)
4. Moyano et al. ²⁶	Peru (Tumbes)	2011-2012 157	157	22,278	NS	76 (48.3) NS	NS	NS	704.4 (598–823)	686.2 (541–858)	722.9 (574–897)
5. Jarbandhan et al. ^{2.}	5. Jarbandhan et al. ²⁷ Surinam (Paramaribo)	2013	47	1478	58.9	21 (44.6) 87	87	NS	3179.9 (2345-4206)	3179.9 (2345–4206) 2267.8 (1409–3445) 4710.1 (3099–6825)	4710.1 (3099–6825)
NS: not stated; SD: stand	NS: not stated; SD: standard deviation; CI: confidence interval.										

(95% CI=9.8–28.6) in Matao, Brazil¹⁹ to 24.6% (95% CI=21.9–27.6) in $\tilde{N}uble^{21}$ (Figure 5).

The overall 1-month case-fatality rate in IS was 13.3% (95% CI=10.9–15.7) and for ICH 38.0% (95% CI=22.9–53.3). SAH was the most fatal type of stroke, with an overall 1-month case-fatality rate of 48.6% (95% CI=37.9–59.2) (Figure 5).

Risk of bias in studies

We used the JBI²⁸ Critical Appraisal Checklist for study reporting cohort and prevalence data to assess the 12 included studies that revealed incidence, case-fatality, and prevalence rate. The grade (yes, no, or unclear) is shown in Supplemental Table S.6. Most of the incidence studies indicated that statistical methods were adequate. Prevalence studies had insufficient coverage of the identified sample (Supplemental Table S.6).

Publication bias

Publication bias was ruled out with Egger's test in incidence, prevalence and case-fatality studies. All results had a p-value > 0.05 (Supplemental Table S.7).

Discussion

In this article, we present a systematic review of the incidence, prevalence, and case-fatality of stroke for LAC. Limited epidemiological data were found, representing only seven of the 36 LAC countries. Nevertheless, the valuable information available was derived from high-quality studies, representing the leading countries in stroke burden research in the region, such as Brazil and Chile. These results are a first approximation of quality epidemiological parameters, although collaborative academic research efforts in LAC are a priority to improve the information in the region, as it has been previously suggested.²⁹

Large variation was detected in the stroke crude incidence rates in the region, ranging from 73.6²⁰ to 180.4/100,000,²¹ with increasing rates in older age groups. Even though previous reports have considered LAC within the lowest stroke burden regions in the world,³⁰ this information contributes to evidencing wide differences between countries.

When the age-adjusted incidence was analyzed, the lowest values were found in Martinique (77/100,000) and the highest values were found in Ñuble, a region of southern Chile (254.1/100,000), being both high-income countries. The higher values for Ñuble, Chile may be related to the fact that the study was conducted in an underserved, lowincome population in Chile with a high prevalence of cardiovascular and environmental risk factors. In addition, the design was a population-based study that captured all strokes, even the mildest. It is also possible

Table 3. Summary of the epidemiological data from studies assessing the prevalence in Latin America and the Caribbean.

Figure 4. Overall crude prevalence rate per 100,000 people in Latin America and the Caribbean. The overall crude annual prevalence rate SE (95%CI) ID Country, Location (study period) Events Total Rate [95% CI] Weight 41 2000 2050.0 [1475.0-2770.8] 20.01 1 Argentina, Gnal Villegas (2015-2016) 2 Brazil, Coari (2011) 246 5925 4151.9 [3658.2-4691.3] 20.14 3661 5271.7 [4570.2-6045.8] 19.86 3 Brazil, Sao Jorge (2008) 193 4 Peru, Tumbes (2011-2012) 157 22278 704.4 [599.1-823.5] 20.40 1478 3179.9 [2345.6-4206.4] 19.59 5 Surinam (2013) 47 Total (95% CI) 35342 3060.1 [1090.6-5023.6] 100.00 Total events 684 Heterogenity: $X^2 = 343.16429$, d.f.=4 (p= 0.000); l² = 98.83% Estimate of between-study variance $T^2 = 0.0004$ SE: Standard error; CI: Confidence interval.

Figure 5. Overall case-fatality rate at I month after the first stroke in Latin America and the Caribbean and by stroke subgroup.[AQ: 4]

ID	Study period	Events	Total	Case-fatali	ty (%)[95% CI]	SE (95%CI)	Weight
1	Argentina, Tandil (2013-2015)	67	334	20.1	[15.9-24.8]		18.08
2	Brazil, Joinville (2005-2006)	145	759	19.1	[16.3-22.1]		25.68
3	Brazil, Matao (2003-2004)	15	81	18.5	[10.8-28.7]		7.29
4	Brazil, Matao (2015-2016)	14	81	17.3	[9.8-27.3]		7.61
5	Chile, Iquique (2000-2001)	68	292	23.3	[18.6-28.6]		15.84
6	Chile, Ñuble (2015-2016)	219	890	24.6	[21.8-27.6]		25.50
	Total (95% CI)		2437	21.1	[18.6-23.7]	\diamond	100.00
	Total events	528				0 2 .4	
	Heterogenity: X ² = 9.8822, d.f.	.=5 (p= 0.	.078); l ⁱ	² = 49.40%			
	Estimate of between-study vari	ance T ² = (0.000				
Fig	s. 5.2 Ischemic stroke						
ID	Study period	Events	Total	Case-fatali	ty (%)[95% CI]	SE (95%CI)	Weight
1	Argentina, Tandil (2013-2015)	37	251	14.7	[10.6-19.7]		22.04
2	Brazil, Matao (2003-2004)	9	69	13.0	[13.0-23.3]		8.09
3	Brazil, Matao (2015-2016)	7	67	10.4	[4.3-20.3]		9.37
4	Chile, Iquique (2000-2001)	33	185	17.8	[12.6-24.1]		- 15.31
5	Chile, Ñuble (2015-2016)	75	641	11.7	[9.83-14.4]		45.19
	Total (95% CI)		1213	13.3	[10.9-15.7]	\diamond	100.00
	Total events	161					
	Heterogenity: X ² = 5.073, d.f.=	=4 (p= 0.2	280); I ²	= 21.156%		.1 .2	-
_	Estimate of between-study vari	ance T ² = (0.000				
Fig	3. 5.3 Intracerebral Hemory	rhage					
ID	Study period	Events	Total	Case-fatali	ty (%)[95% CI]	SE (95%CI)	Weight
1	Argentina, Tandil (2013-2015)	13	54	24.1	[13.5-37.6]		23.95
2	Brazil, Matao (2003-2004)	5	11	45.5	[16.7-76.6]	······	13.52
3	Brazil, Matao (2015-2016)	4	10	40.0	[12.2-73.8]		13.08
4	Chile, Iquique (2000-2001)	20	69	29.0	[18.7-41.2]		24.34
5	Chile, Ñuble (2015-2016)	61	111	55.0	[45.2-64.4]		25.11
	Total (95% CI)		255	38.0	[22.9-53.3]	$\langle \rangle$	100.00
	Total events	103					
	Heterogenity: X ² = 21.540, d.f.			2 = 81.43%	•	5	
_	Estimate of between-study vari	_	0.021				
1210	3. 5.4 Subarachnoid hemor						
_		Events	Total	Case-fatali	ty (%)[95% CI]	SE (95%CI)	Weight
ID							20.06
ID 1	Argentina, Tandil (2013-2015)	8	17	47.1	[22.9-72.1]		
1D 1 2	Argentina, Tandil (2013-2015) Brazil, Matao (2015-2016)	8 3	4	75.0	[19.4-99.3]		6.27
1D 1 2 3	Argentina, Tandil (2013-2015) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001)	8 3 6	4 15	75.0 40.0	[19.4-99.3] [16.3-67.7]		6.27 18.37
ID 1 2 3 4	Argentina, Tandil (2013-2015) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Ñuble (2015-2016)	8 3 6 23	4 15 47	75.0 40.0 48.9	[19.4-99.3] [16.3-67.7] [34.1-63.9]		6.27
ID 1 2 3 4	Argentina, Tandil (2013-2015) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Ñuble (2015-2016) Brazil, Matao (2003-2004)	8 3 6	4 15 47 1	75.0 40.0 48.9 (excluded)	[19.4-99.3] [16.3-67.7] [34.1-63.9] [-]		6.27 18.37 55.29 -
ID 1 2 3 4	Argentina, Tandil (2013-2015) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Ñuble (2015-2016) Brazil, Matao (2003-2004) Total (95% CI)	8 3 6 23 1	4 15 47	75.0 40.0 48.9	[19.4-99.3] [16.3-67.7] [34.1-63.9]		6.27 18.37
ID 1 2 3 4	Argentina, Tandil (2013-2015) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Ñuble (2015-2016) Brazil, Matao (2003-2004) Total (95% CI) Total events	8 3 6 23 1 40	4 15 47 1 83	75.0 40.0 48.9 (excluded) 48.6	[19.4-99.3] [16.3-67.7] [34.1-63.9] [-] [37.9-59.2]		6.27 18.37 55.29
ID 1 2 3 4	Argentina, Tandil (2013-2015) Brazil, Matao (2015-2016) Chile, Iquique (2000-2001) Chile, Ñuble (2015-2016) Brazil, Matao (2003-2004) Total (95% CI)	8 3 6 23 1 40 .=3 (p= 0.	4 15 47 1 83 .5792);	75.0 40.0 48.9 (excluded) 48.6	[19.4-99.3] [16.3-67.7] [34.1-63.9] [-] [37.9-59.2]		6.27 18.37 55.29 -

that the population incidence of IS is increasing due to the prevalence of vascular risk factor in an aging population.²¹ The evidenced large variation in age-adjusted stroke incidence between LAC countries is similar to what was previously informed by the global burden diseases (GBDs) for 2019,¹ but the recently reported increased incidence in one of the high-income countries of the region (Chile) is a concern that will require a careful oversight.

Regarding stroke prevalence, only five studies were found,^{23–27} reporting data with large variation. Rates ranged from 705/100,000 in Perú²⁶ to 5271.7/100,000 in Brazil.²⁵All of them were cross-sectional studies, but they used different definitions and age groups, making comparisons difficult. Despite these limitations, the crude prevalence ranges in middle-income countries were in line to those reported by the GBD for LAC.^{31,32} Stroke experts and policy makers in Latin America have addressed the increase in stroke prevalence that has occurred in recent decades,³² but more research on stroke prevention, incidence, prevalence, and outcomes is needed to obtain a better approximation of the impact of these estimates.

Overall case-fatality rate at 1 month after the first stroke of 21.1% was low compared to other regions of the world.³⁰ One-month case-fatality rates had low-moderate heterogeneity found across studies. This finding may reflect the degree of development and local resources of countries included in this study, since its values were similar to those reported by other studies that analyzed mainly high- and middle-income countries.³²

Strengths and limitations

One of the main strengths of this systematic review is that we covered the last three decades, which made possible to estimate epidemiological information over a long period. In addition, the study was mostly based on population-based research that met the criteria for an ideal stroke revision,³³ ensuring that the data were complete and comparable and provided the most precise estimates of incidence, prevalence, and early case-fatality rate of stroke.

Our study has some limitations. First, meta-analysis techniques were used in data with high heterogeneity. This was decided because the interpretation of heterogeneity in systematic reviews of prevalence and incidence is controversial, since it is expected to find clinical, methodological, and statistical heterogeneity when studying a large area and different populations. It has been argued that I^2 may not be discriminative and should be interpreted with caution.³⁴ As most previous prevalence systematic reviews, we presented the results with ranges and also point estimates including prediction intervals for a correct interpretation. Second, restricting our analysis to published reports might have introduced publication bias; nevertheless, this bias is likely to be small because uncited publications may not be

relevant given the strict methodology required to diagnose and report stroke studies. Finally, this analysis can be limited by the fact that the analyzed countries had uppermiddle or high incomes, leaving other LAC countries underrepresented,²⁹ and because it is very difficult to compare countries in such a wide area with large differences even within countries over time in terms of income, level of education, access to health services, ethnicity, and sociocultural determinants.

Conclusion

In summary, the results of this review contribute to our understanding regarding the burden caused by stroke in LAC. More studies with comparable designs are needed to generate reliable data and should include both standardized criteria, such as the World Health Organization³³ clinical criteria and updated standard methods of case assurance, data collection, and reporting.³⁵ More research is required in middle-low- or low-income populations in LAC, given their low representation. This information should be examined further to determine whether gender differences exist in these populations.

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Data accessibility statement

The data that support the findings of this study are available from the corresponding author (P.M.) upon reasonable request.

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Supplemental material

Supplemental material for this article is available online.

References[AQ: 5]

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