# Articles

# Cervical cancer screening coverage in the Americas region: a synthetic analysis

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## Summary

Background The Americas region has the lowest (North America) and the second highest (Latin America and Caribbean) cervical cancer (CC) mortality worldwide. The lack of reliable data on screening coverage in the region hinders proper monitoring of the World Health Organization (WHO) CC elimination initiative.

Methods For this synthetic analysis, we searched data on CC screening coverage from official sources and national health surveys, supplemented with a formal WHO country consultation. Context data were obtained from official sources (income, health expenditure, inequality-adjusted human development index -IHDI-, universal health coverage, CC incidence/mortality). Country age-specific coverages for 2019 by screening interval were computed. Missing data were imputed through a multi-step algorithm. Beta-regression and Poisson-regression models were used to analyse associations between context variables, screening coverage, and CC mortality.

Findings We included data from 37 countries in the Americas. Data on coverage of HPV testing was scarce, and for many countries only Pap-smear coverage data was available. Overall, 78%, 34%, 60%, and 67% of women aged 25–65 years have been screened ever in their lifetime, and in the previous year, 3 years, and 5 years, respectively. By sub-region, 3-year coverage ranges from 48% (South America) to 72% (North America). Twenty-four countries showed screening coverage below 70%. Income and health system type were associated with screening coverage, but coverage was not associated with CC mortality.

Interpretation In the Americas region 35.1% and 56.8% of countries report 3-year and 5-year coverage over 70%, respectively. Inequalities remain a major challenge for screening programs in the region. The elimination campaign should reinforce the transition to HPV testing and strengthen surveillance systems.

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Keywords: Uterine cervical neoplasms; Early detection of cancer; Mass screening; Americas

#### Introduction

The World Health Organization (WHO) launched in 2020 an initiative for the elimination of cervical cancer aimed at reducing incidence below 4 per 100,000.<sup>1</sup> Target goals to achieve this objective include having, 90% of girls fully vaccinated against human papilloma-virus (HPV) at 15 years of age, 70% of women screened twice between 35 and 45 years old using a high-performance test, and 90% or more identified

precancerous lesions and invasive cancers treated by 2030.<sup>1</sup> Accordingly, the Pan American Health Organization (PAHO) considers strengthening information systems a critical component of cervical cancer prevention and control programmes to allow proper monitoring of the progress towards elimination targets.<sup>2</sup>

Significant disparities in the socioeconomic determinants of cervical cancer incidence and mortality have been described for the Americas region.<sup>3</sup> Although





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#### **Research in context**

#### Evidence before this study

The WHO launched a global initiative to eliminate cervical cancer (CC) as a public health problem. Screening coverage over 70% is a key elimination target; thus, reliable data are essential for monitoring the progress towards CC elimination. Most countries in the Americas region lack organized screening, and standardized and comparable coverage estimates are not yet available. We recently reported global CC screening coverage using an innovative methodology that allows for comparability; however, detailed data at the country level is not yet published. In March 2020, we carried out a search from 2009 to 2019 considering reports about national coverage of CC screening in the Americas region. The search was done in LILACS (DeCS terms "cáncer cervical" AND "tamizaje" AND "cobertura") and PubMed (terms "North America" [Mesh] OR "Caribbean Region" [Mesh] OR "West Indies" [Mesh] OR "Latin America" [Mesh] OR "South America" [Mesh] OR "Central America" [Mesh] AND "Mass Screening" [Mesh] OR "Early Detection of Cancer" [Mesh] OR "Diagnostic Screening Programs" [Mesh] OR "Early Diagnosis" [Mesh] AND \*Coverage OR Participation\* OR "Screening coverage" OR "Invitation coverage" OR "Up-to-date screening" OR "adherence" OR "uptake" OR "Effective screening" OR "Examination rate" AND "Uterine Cervical Neoplasms" [Mesh] OR "Uterine Cervical Dysplasia" [Mesh] OR "Cervical Intraepithelial Neoplasia" [Mesh] OR "cervical cancer" OR "cervical precancer" OR "cervical precancerous lesions"). We found 498 reports but only 27 articles used national data on CC screening coverage, mostly from the USA and Brazil. Only one study reported binational data (USA and Canada) and a narrative review reported coverage for 14 Latin American countries; however, no standard definition of coverage was used regarding target populations, screening intervals, or screening tests; in consequence, coverage data are highly heterogenous without any standardization or data processing.

#### Added value of this study

We used an innovative methodology to present baseline estimates of CC screening coverage for the 37 WHO member states in the Americas region. In addition, we collected and

the burden of disease has decreased in some countries, there are still significant differences between and within countries, without substantial changes for the most disadvantaged settings, where cervical cancer mortality has increased.<sup>3,4</sup> Accordingly, Latin America and the Caribbean is the region with the second highest cervical cancer mortality worldwide after Africa, whereas North America has the lowest incidence and mortality globally.<sup>5</sup>

Deficient organization of screening programmes challenges both the achievement of coverage targets and monitoring of programme performance.<sup>6</sup> Previous reported data on contextual factors with a potential influence on screening coverage and CC mortality. For every country, we report coverage data for the most common target populations in national programmes (women 25–65 years old), and for the target population defined in the WHO elimination strategy (women 35–45 years old). All data are reported for 1-, 3-, and 5-year screening intervals and ever in lifetime. Coverage estimates are analysed against contextual variables including income level, health expenditure, human development index, universal health coverage, programmatic approach, and predominant type of health system (segmented/not-segmented). The methodology enables the comparison of estimates between countries in the region with similar contextual conditions.

#### Implications of all the available evidence

CC control remains an unmet goal for most countries in the Americas region. Although low screening coverage has been suggested as a major cause for the lack of impact on CC mortality, no significant progress on programme monitoring is observed for the majority of countries. Previous reports on screening coverage are disconnected from screening guidelines and lack uniformity in terms of target populations and screening intervals. Using innovative methods to overcome these limitations, we estimate that 60% of women aged 25-65 years in the Americas region have been screened during the previous three years, as recommended by most national guidelines on cytology-based screening; however, coverage estimates increase to 67% with an extended screening interval (five years), suggesting that an improved transition to human papillomavirus (HPV) testing will bring the region closer to the WHO elimination target. Socioeconomic disparities remain a major challenge to increasing screening coverage, but we provide insights reqarding the characteristics of health systems and screening programmes to help countries understand the determinants of successful screening. Providing standardized information is not only a call to improve monitoring systems by using common indicators but also an opportunity to learn from neighbouring countries with similar cultural and socioeconomic backgrounds.

reports on screening coverage for the Americas region have failed to provide reliable data given not only the varied quality of information sources but also the lack of standardized reports regarding coverage definition, target populations, and screening intervals.<sup>7</sup> Indeed, several countries' reports do not provide information aligned with screening policies, making it difficult to properly assess the programme's performance.<sup>8</sup>

Recently, an accurate method for providing standardized information on screening coverage was reported by Bruni and colleagues.<sup>9</sup> This methodology enables comparability of the estimates despite the heterogeneity of screening policies and variability of available coverage data. However, the initial report does not provide detailed data for different screening intervals and age groups for individual countries.<sup>9</sup> Using this approach, we present detailed standardized estimates of cervical cancer screening coverage in the region, supplemented by a specific analysis to explore the association between contextual factors, screening coverage, and cervical cancer mortality.

# Methods

The methods for data extraction and the statistical analysis to provide standardized coverage estimates have been described elsewhere.<sup>9</sup>

A detailed description of the systematic search and inclusion and exclusion criteria is presented in the Supplementary Material.

#### Data sources

Briefly, from July 2019 to October 2020, we searched official websites and data from national surveys and governmental reports of different screening coverage intervals (previous one year, previous two years, previous three years, previous five years, and ever in lifetime). Complementary information from international data sources was also retrieved (WHO and USAID databases). After the first review, if no coverage information was available, we conducted a systematic search in the PubMed database (via Medline) without language or date restrictions (Supplementary Material, Appendix 1). If more than one source was found we consulted country experts about data reliability. Finally, to review the data collected and preliminary estimates on cervical cancer screening coverage, a formal consultation round with WHO member states was done from November 2020 to February 2021, resulting in the update of screening coverage data for the Bahamas, Brazil, Canada, Colombia, Jamaica, Peru, the United States, and Uruguay.

We examined changes in the screening programme (including screen-and-treat approaches),<sup>10</sup> income level, and health system reforms, in order to qualitatively assess the representativeness of coverage data for each country's situation by 2019 before the COVID-19 pandemic; thus, only national, population-based screening data representative of the country's situation in 2019 entered the final database and criteria for such representativeness have been previously described.<sup>9</sup>

We collected data on contextual variables from the United Nations (UN population prospects and UN inequality-adjusted human development index -IHDI-,<sup>11,12</sup> the World Bank (income level),<sup>13</sup> and the WHO Global Health Observatory (Universal Health Coverage and per capita health expenditure).<sup>14</sup> For the classification of health systems, we found an extensive number of typologies and the coexistence of multiple models; however,

no classification including all countries in the region was found.<sup>15,16</sup> Consequently, we classified health systems into four categories as proposed by Chung M17,18: national health systems with a unique payer and provider (NHS: Beveridge model), national health insurance systems (NHI: individual basis) based on public funding but with public or private providers, fragmented health systems with social security provision for workers and public health provision for the remaining population (SS: Bismarck model), and countries with predominantly out-ofpocket funding, including private insurance and health services provision (OP). Two senior investigators independently categorized health systems considering the main funding source and care provision model based on multiple sources of information; disagreements were solved by consensus between the two. There is no health system with a unique funding source or provision model; thus, we considered the models covering the majority of the population to be the predominant type.<sup>19-38</sup>

# Statistical analysis

Estimated data on cervical cancer incidence and mortality (2020) were retrieved from the WHO-IARC databases.<sup>5,39</sup> The Average Annual Percentage Change (AAPC) of mortality rates was estimated for the last 15 years of available data, with the corresponding 95% confidence intervals (95% CI).<sup>40</sup>

We extracted screening coverage data by age group for any available screening interval. Coverage rates were transformed into single-age data points by assigning the same coverage to all ages in the corresponding age group. Missing data were assumed at random and imputed through a multi-step algorithm using different statistical techniques,<sup>9</sup> and a sensitivity analysis was done to assess and validate the methodology. Briefly, we used linear interpolation between screening intervals, imputations per missing datapoint using predictive mean matching methods, last observation carried forward or next observation carried backwards, and ponderation rates based on coverage from countries with similar income and screening algorithms.<sup>9</sup>

Country-specific estimates were computed from the estimation of the number of screened women for each age group, screening interval, and the country as the numerator and the UN populations as the denominator. Bootstrap 95% confidence intervals were calculated using the percentile method with 3000 bootstrap replications. Country-specific estimates were aggregated by age group (25–65 years and 35–45 years) and contextual variables.

Additionally, in this paper, we used a Beta regression model to analyse the association of contextual variables with screening coverage as a dependent variable in the form of percentages, and we used a Poisson regression model to evaluate the association between screening coverage and contextual variables with cervical cancer mortality (Age Standardized Rate

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Country	Socioeconomi	ic characterist	ics		Health system	characteristics	Cervical cancer				
	Population 2019 <sup>(a)</sup>	Rural pop 2019 (%) <sup>(a)</sup>	Income level <sup>(b)</sup>	Inequality- adjusted HDI 2019 <sup>(c)</sup>	Predominant health system Type <sup>(d)</sup>	UHC index services capacity & access 2017 <sup>(e)</sup>	Health expenditure per capita (US \$) 2018 <sup>(e)</sup>	Incidence 2020 (ASR per 100,000) <sup>(f)</sup>	Mortality 2020 (ASR per	Average annual percentage change (AAPC) mortality (ASR)—15 years <sup>(9)</sup>	
									100,000)(")	Last year	AAPC (95% CI)
North America		_									
Bermuda	63,000	0.0	High	N/A	SS	N/A	N/A	N/A	N/A	N/A	N/A
Canada	37,411,000	18.5	High	Very high	NHI	100	4995	5.5	1.9	2015	-1.4 (-3.8, 1.1)
United States of America	329,065,000	17.5	High	Very high	OP	100	10,624	6.2	2.1	2016	-0.7 <sup>b</sup> (-0.9, -0.5)
Central America											
Belize	390,000	54.1	Upper-middle	Medium	NHI	56	286	19.1	14.8	2016	-0.7 (-3.7, 2.5)
Costa Rica	5,048,000	19.7	Upper-middle	Medium	NHI	77	910	11.7	5.4	2014	-3.4 <sup>b</sup> (-5.2, -1.6)
El Salvador	6,454,000	27.2	Lower-middle	Low	SS	79	289	13.1	7.4	2014	-2.9 <sup>b</sup> (-4.1, -1.7)
Guatemala	17,581,000	48.6	Upper-middle	Low	SS	32	260	20.3	11.9	2016	2.0 (-0.2, 4.2)
Honduras	9,746,000	41.5	Lower-middle	Low	SS	45	176	19.5	12.5	N/A	N/A
Mexico	127,576,000	20.3	Upper-middle	Medium	SS	80	520	12.6	5.7	2016	-3.9 <sup>b</sup> (-4.4, -3.3)
Nicaragua	6,546,000	40.0	Lower-middle	Low	SS	73	174	21.3	12.6	2017	-1.7 <sup>b</sup> (-2.7, -0.7)
Panama	4,246,000	31.8	High	Medium	SS	89	1132	14.0	7.5	2016	-1.9 <sup>b</sup> (-3.4, -0.4)
South America											
Argentina	44,781,000	8.1	Upper-middle	High	SS	89	1128	16.7	8.7	2016	-0.2 (-1.1, 0.6)
Bolivia	11,513,000	29.9	Lower-middle	Low	SS	82	224	36.6	18.0	N/A	N/A
Brazil	211,050,000	13.3	Upper-middle	Medium	NHS	99	848	12.7	6.3	2016	-0.7 <sup>b</sup> (-0.9, -0.6)
Chile	18,952,000	12.0	High	High	SS	94	1456	11.1	5.2	2016	-3.1 <sup>b</sup> (-4.6, -1.6)
Colombia	50,339,000	18.7	Upper-middle	Medium	NHI	85	513	14.9	7.4	2015	-3.0 <sup>b</sup> (-4.2, -1.8)
Ecuador	17,374,000	35.4	Upper-middle	Medium	SS	86	516	16.0	8.2	2016	1.1 <sup>b</sup> (0.2, 2.0)
Guyana	783,000	73.7	Upper-middle	Medium	NHI	70	296	29.5	15.1	N/A	N/A
Paraguay	7,045,000	37.8	Upper-middle	Medium	SS	63	400	34.1	19.0	2016	2.2 <sup>b</sup> (0.5, 4.0)
Peru	32,510,000	22.2	Upper-middle	Medium	SS	81	369	22.2	11.5	2015	0.6 (-1.0, 2.3)
Suriname	581,000	33.4	Upper-middle	Low	NHI	78	474	23.7	14.1	2014	-1.1 (-3.8, 1.7)
Uruguay	3,462,000	4.6	High	High	NHI	94	1590	11.7	5.6	2016	-2.4 <sup>b</sup> (-4.0, -0.9)
Venezuela	28,516,000	13.5	Upper-middle	Medium	SS	75	257	22.2	12.5	2013	-1.2 <sup>b</sup> (-1.9, -0.5)
Caribbean											
Antigua & Barbuda	97,000	81.4	High	N/A	NHI	70	875	NA	NA	N/A	N/A
Bahamas	389,000	17.5	High	N/A	NHI	81	2013	14.9	10.6	N/A	N/A
Barbados	287,000	69.0	High	Medium	NHS	78	1165	15.2	9.0	2013 <sup>a</sup>	-2.5 <sup>b</sup> (-0.8, -3.3)
Cuba	11,333,000	23.2	Upper-middle	N/A	NHS	100	987	13.9	6.9	2016	-1.5 <sup>b</sup> (-2.1, -1.0)
Dominica	72,000	30.6	Upper-middle	N/A	NHS	NA	491	NA	NA	N/A	N/A
Dominican Republic	10,739,000	18.6	Upper-middle	Medium	SS	75	462	17.9	11.7	2013	0.4 (-2.1, 2.9)
Grenada	112,000	61.6	Upper-middle	N/A	NHS	64	475	NA	NA	N/A	N/A
Haiti	11,263,000	43.7	Low	Low	NHS	30	64	11.6	9.0	N/A	N/A
Jamaica	2,948,000	43.4	Upper-middle	Medium	NHI	74	321	21.6	13.6	N/A	N/A
									(T	able 1 cont	inues on next page)

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Country	Socioeconomi	c characterist	ics		Health system o	characteristics		Cervical cancer			
	Population 2019 <sup>(a)</sup>	Rural pop 2019 (%) <sup>(a)</sup>	lncome level <sup>(b)</sup>	Inequality- adjusted HDI 2019 <sup>(c)</sup>	Predominant health system Type <sup>(d)</sup>	UHC index services capacity & access 2017 <sup>(e)</sup>	Health expenditure per capita (US \$) 2018 <sup>(e)</sup>	Incidence 2020 (ASR per 100,000) <sup>(f)</sup>	Mortality 2020 (ASR per	Average al change (A (ASR)—15	inual percentage APC) mortality years <sup>(g)</sup>
									100,000) <sup>(1)</sup>	Last year	AAPC (95% CI)
(Continued from previous page)											
Puerto Rico	2,933,000	8.0	High	N/A	SS	N/A	N/A	8.0	2.8	N/A	N/A
Saint Kitts & Nevis	53,000	73.6	High	N/A	NHS	N/A	993	N/A	N/A	N/A	N/A
Saint Lucia	183,000	80.3	Upper-middle	Medium	NHS	59	465	16.6	11.0	N/A	N/A
Saint Vincent & the Grenadines	111,000	46.8	Upper-middle	N/A	NHS	59	329	NA	NA	N/A	N/A
Trinidad & Tobago	1,395,000	46.2	High	N/A	SHN	78	1123	19.8	11.9	2012	-1.8 (-3.7, 0.1)
IHDI: Inequality-Adjusted Human Devel health provision for the remaining pop Available. <sup>4</sup> Includes only the last 14 yea <055, <sup>(6)</sup> Health system type, <sup>16-37</sup> <sup>(6)</sup> V	opment Index, UH ulation, NHI: natio rs. <sup>b</sup> Statistically sig VHO 2019, <sup>13 (f)</sup> IA	C: Universal Hea nal health insur Inificant. Source .RC 2020, <sup>5 (g)</sup> M	lith Coverage, ASR: ance (individual) bi s: <sup>(a)</sup> United Natior /HO cancer mortal	Age Standardize ased on public fu is Populational <sup>16</sup> ity database. <sup>38</sup>	ed Rate, Cl: Confide nding but public a Prospects 2019, <sup>(t</sup>	ence Interval, NHS: hea nd private providers, C <sup>3)</sup> World Bank, <sup>12 (d)</sup> Un	Ith systems with unique DP: predominantly out of ited Nations, IHDI classifi	payer, SS: health syst pocket health servic cation <sup>11</sup> : Very high (	:ems based on so es provision incl ≥80), High (0.7	ocial security Iuding private 0-0.79), Med	or workers and public insurance. N/A: Non- um (0.55-0.69), Low:
Table 1: Country's contextual char	acteristics.										

-ASR-per 100,000 women). For both models, we ran bivariate and multivariate analyses. For the multivariate analysis, only variables with p-values <0.05 were included. The contextual variables considered in the models include income level (four categories: high-, upper-middle-, lower-middle-, and low-) IHDI (four categories: very high, high, medium, and low), predominant health system model (two categories: universal financial coverage and integrated health care provision -NHS/NHI- and fragmented financial coverage and healthcare provision -SS/OP-), predominant health system financing (three categories: public revenues, mixed, and private), predominant health services provision (two categories: public and public/private), health system steering role (two categories: national and territorial), public health expenditure (as continuous and in two categories: <70% and  $\geq$ 70%), cervical cancer screening scheme (four categories: one approach, different approaches-same population, different approaches-different population, and no programme), screen-and-treat approach (two categories: yes and no/no programme),<sup>10</sup> rural population (as continuous in percentage), and Universal Health Coverage (UHC) service capacity index (as continuous).

## Role of the funding source

The funders of the study had no role in study design, data collection, data analysis, data interpretation, or writing of the report.

## Results

We collected data from 37 countries in North America (3), Central America (8), South America (12), and the Caribbean (14). In total, 12 countries were categorized as high-income, 20 as upper-middle-income, 4 as lowermiddle-income, and only Haiti as low-income; however, socioeconomic determinants largely varied within these income categories (for example, in high-income countries, the percentage of the rural population ranged from 0 to 69 and health expenditure per capita from USD\$875 to USD\$10,624). Accordingly, the countries' IHDI categorizations did not match their income levels (Table 1). Despite the differences in socioeconomic determinants, most countries showed a significant decrease in cervical cancer mortality rates in the last 15 years, with only Ecuador and Paraguay showing a significant increase for the period analysed (AAPC for Ecuador: 1.1 [95% CI, 0.2-2.0] and AAPC for Paraguay: 2.2 [0.5-4.0]; Table 1).

At the time of data collection, most countries' reports on coverage data corresponded to cytology-based screening and no specific data on coverage of screenand-treat approaches was identified. Only governments from Brazil, Colombia, and Uruguay reported data from administrative sources (Brazil: cancer information

Area	Previo	ous year			Previo	us 3 years			Previo	us 5 years			Ever in	lifetime		
	Numb wome N (95	er of screened n in millions % CI)	Cover (95%	age % Cl)	Numbe womer (95% C	er of screened in millions N [1]	Cover (95%	age % CI)	Numbe womer N (95%	er of screened n in millions 6 CI)	Cover (95%	age % Cl)	Numbe women N (95%	r of screened in millions 6 CI)	Covera (95%	age % CI)
Total	90.4	(78.6–103.4)	34%	(29–39%)	160.0	(140.1–182.0)	60%	(52–68%)	178.0	(156.1–202.0)	67%	(58–76%)	208.8	(183.1–236.6)	78%	(69–89%)
By subregion																
North America	42.0	(32.9–51.3)	43%	(34–52%)	70.8	(56.1-85.9)	72%	(57-88%)	76.6	(60.8–92.9)	78%	(62–95%)	85.6	(68.0–103.6)	87%	(69–100%)
Central America	16.8	(13.4–20.4)	37%	(30–46%)	29.1	(22.9–35.7)	65%	(51-80%)	33.2	(26.1-40.8)	74%	(58–91%)	37.9	(29.9–46.6)	85%	(67–100%)
South America	28.0	(25.1–30.9)	25%	(22–27%)	54.2	(48.1–60.5)	48%	(42-53%)	61.4	(54.3–69.0)	54%	(48-61%)	77.8	(67.2–89.6)	68%	(59–79%)
Caribbean	3.6	(3.1-4.2)	34%	(29-40%)	6.0	(5.1–7.0)	56%	(48-65%)	6.8	(5.8–7.8)	63%	(54–73%)	7.4	(6.4-8.5)	69%	(59–79%)
By income level																
High income	45.5	(34.7–57.3)	43%	(33-54%)	76.8	(59.0–96.1)	72%	(55-90%)	83.2	(64.0–104.1)	78%	(60–98%)	93.3	(71.8–116.6)	87%	(67–100%)
Low and middle income	44.8	(40.1-49.6)	28%	(25-31%)	83.3	(74.2–92.6)	52%	(46–58%)	94.8	(84.3–105.7)	59%	(52–66%)	115.5	(101.6–130.3)	72%	(63-81%)
Upper middle income	42.2	(37.7-47.0)	28%	(25-31%)	78,6	(69.8–88.0)	52%	(46–59%)	89.5	(79.3–100.5)	60%	(53–67%)	109.5	(95.9–124.2)	73%	(64-83%)
Lower middle income	2.6	(2.4–2.8)	33%	(30–36%)	4.6	(4.3-4.9)	58%	(54-62%)	5.1	(4.8–5.5)	65%	(60–69%)	5.8	(5.4–6.2)	73%	(68–78%)
Low income	52 k	(42 k-63 k)	2%	(2–3%)	122 k	(110 k–137 k)	5%	(4–5%)	185 k	(166 k–204 k)	7%	(7-8%)	245 k	(219 k–272 k)	10%	(9–11%)
By inequality-adjusted human develop	ment in	dex <sup>a</sup>														
Very high (≥ 0.80)	42.0	(34.2-49.4)	43%	(35-51%)	70.7	(58.4–82.7)	72%	(60-84%)	76.6	(63.3–89.4)	78%	(65–91%)	85.6	(70.8–99.7)	87%	(72–100%)
High (0.70–0.79)	8.1	(6.9–9.4)	46%	(39–54%)	13.0	(11.2–15.0)	75%	(64–86%)	13.8	(11.9–15.9)	79%	(68–91%)	15.7	(13.6–17.9)	90%	(78–100%)
Medium (0.55–0.69)	34.4	(30.1-38.9)	26%	(23–29%)	66.3	(57.8–75.1)	50%	(44-57%)	76.3	(66.4-86.7)	57%	(50–65%)	94.8	(81.7–109.0)	71%	(62-82%)
Low (<0.55)	3.8	(3.3-4.2)	26%	(23–30%)	6.5	(5.8–7.1)	45%	(41-50%)	7.4	(6.7-8.1)	51%	(47-56%)	8.5	(7.7–9.3)	60%	(54-65%)
By universal health coverage index $^{\mathrm{b}}$																
>82	68.0	(57.6–79.1)	34%	(29–40%)	119.2	(102.0–137.6)	60%	(51-69%)	131.1	(112.4–151.1)	66%	(57–76%)	154.9	(132.9–178.2)	78%	(67-90%)
74-81	18.7	(15.2–22.4)	34%	(27–40%)	34.6	(28.2–41.3)	62%	(51–74%)	39.8	(32.6–47.6)	71%	(58–85%)	46.0	(37.7-54.9)	83%	(68–98%)
<74	3.2	(2.8-3.8)	27%	(23–31%)	5.5	(4.8-6.3)	46%	(40-53%)	6.3	(5.5–7.1)	52%	(45-59%)	7.1	(6.2–8.0)	59%	(52-67%)
By predominant health system type																
National health system	8.8	(6.8–10.9)	14%	(10–17%)	23.0	(17.4–29.0)	35%	(27-44%)	27.2	(20.4–34.5)	42%	(31-53%)	38.3	(27.9–49.9)	59%	(43-77%)
National health insurance (individual)	5.2	(4.1-6.4)	37%	(29-45%)	10.1	(7.8–12.4)	72%	(56-88%)	10.9	(8.5–13.4)	77%	(60–95%)	12.0	(9.4–14.7)	85%	(67–100%)
Social security	38.0	(33.9-42.3)	38%	(34-42%)	63.6	(56.6–70.8)	64%	(57-71%)	71.2	(63.2–79.3)	71%	(63–80%)	81.3	(72.3–90.6)	82%	(73-91%)
Out of pocket (Private)	38.3	(29.8-46.5)	43%	(34–53%)	63.4	(49.5–76.8)	72%	(56–87%)	68.8	(53.8-83.1)	78%	(61–94%)	77.1	(60.5–93.0)	87%	(69–100%)
By public health expenditure (% Total	Health	Expenditure) <sup>c</sup>														
>65	14.5	(12.2–16.9)	44%	(37-51%)	22.8	(19.5–26.3)	69%	(59–79%)	24.7	(21.1–28.4)	75%	(64-86%)	27.1	(23.2–31.1)	82%	(70-94%)
50–64	62.7	(51.9–74.2)	41%	(34-48%)	107.2	(89.4–126.2)	70%	(58-82%)	118.2	(98.8–138.8)	77%	(64–90%)	134.1	(112.4–157.1)	87%	(73–100%)
<49	12.8	(10.7–14.9)	16%	(13–19%)	29.3	(23.6–35.3)	37%	(30–45%)	34.4	(27.5-41.7)	43%	(35-53%)	46.8	(36.3–58.4)	59%	(46-74%)

CI: Confidence Interval. <sup>a</sup>Countries of Antigua and Barbuda, Bahamas, Bermuda, Cuba, Dominica, Grenada, Puerto Rico, Saint Kitts and Nevis, Saint Vincent and the Grenadines, Trinidad and Tobago are excluded because of missing data. <sup>b</sup>Countries of Bermuda, Dominica, Puerto Rico and Saint Kitts and Nevis are excluded because of missing data. <sup>c</sup>Countries of Bermuda and Puerto Rico are excluded because of missing data.

Table 2: Estimated number of screened women and cervical cancer screening coverage by subgroups of analysis, 2019 (women aged 25-65 years).

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system, Colombia: registry of specific protection and early detection activities, and Uruguay: cervical cancer screening programme); data from the remaining countries were from population-based household surveys. For Antigua and Barbuda, Panama, Suriname, and Venezuela, we did not find representative data for 2019.

Overall, in the Americas region, 78% of women aged 25-65 years have been screened ever in their lifetime; 34% have been screened in the previous year, 60% in the previous three years, and 67% in the previous five years (Table 2). Among subregions, in a three-year screening interval, South America and the Caribbean showed the lowest coverage (48% and 56%, respectively) and North America and Central America the highest (72% and 65%, respectively) (Fig. 1), with high-income countries registering higher coverage estimates than low- and middle-income countries (LMIC) (Table 2). Moreover, countries with higher IHDI and higher public health expenditures had higher coverage. Similar trends were observed for coverage estimates at previous one-, five-year screening intervals and ever in their lifetime.

On the basis of individual countries, Haiti (Caribbean) and Guyana (South America) revealed the lowest coverage for a three-year screening interval (4% and 13%, respectively) in women aged 25-65 years, and among the remaining sub-regions, Belize (Central America) and the USA (North America) showed the lowest coverage (39% and 71%, respectively) (Table 3). For women 35-45 years old and a five-year screening interval, representing a proxy to the target population and the screening interval proposed for the WHO elimination strategy, the same countries revealed the lowest coverage in their corresponding subregions (8%, 18%, 50%, and 82%, respectively) and 12 countries (32.4%) remained under the target of 70% coverage (only 3 between 60% and 69%) (Supplementary Material, Appendix 2).

Income level, predominant health system type, public health expenditure, and UHC service capacity index were associated with screening coverage in the bivariate analysis (Supplementary Material, Appendix 3); however, only income level and health system type were significantly associated in the multivariate analyses (Table 4). In particular, the estimated odds of screening coverage decreased to 0.05 (95% CI: 0.01-0.29) for lowincome compared to high-income countries, and countries with an SS/OP health system had an estimated odds ratio (OR) of 0.59 (95% CI: 0.39-0.90) compared to countries with no-fragmented health system (NHS/ NHI). The factors associated with cervical cancer mortality in the bivariate analysis were income level, IHDI, predominant health system, public health expenditure, percentage of rural population, the presence of screenand-treat approaches, UHC service capacity index, and screening coverage (Supplementary Material, Appendix 3); however, only the IHDI and the presence of screen-and-treat approaches showed a significant effect on mortality in the multivariate analysis (Table 4). Specifically, countries with medium and low IHDI had an increase in mortality rate of 92% (rate ratio (RR): 1.92, 95% CI: 1.24–3.10) and 91% (RR: 1.91, 95% CI: 1.15–3.27), respectively, compared to countries with a high or very high index, and countries implementing a screen-and-treat approach had an increase in mortality rate of 38% (RR: 1.38, 95% CI: 1.03–1.84) compared to countries that did not implement this strategy.

# Discussion

To our knowledge, this is the first report to provide detailed and standardized estimates on cervical cancer screening coverage for every country in the Americas region, based on an innovative methodological approach.

Based on coverage estimates for the previous three years among women 25–65 years old, we found that all North American countries already report a screening coverage of over 70%; however, only 10 out of 34 countries in Latin America and the Caribbean meet this level of coverage for a three-year interval. Yet, according to the total number of women screened, we estimate that 60% of women 25–65 years old in the Americas have been screened in the previous three years, and 67% have been screened in the previous five years.

This shows that the Americas region, as a whole, is close to the WHO target to screen 70% women for CC; however, inequalities remain a major challenge for screening programmes, which are expressed not only as differences in screening determinants but also in screening rates and cervical cancer incidence and mortality. The highest mortality rate is nine times higher than the lowest (Paraguay vs Canada), and the difference in screening coverage between the lowest and the highest income levels is 67% (previous three years among women 25-65 years old). In addition, although cervical cancer mortality has decreased in several countries, inequality is evident within countries such as Colombia, Argentina, and Brazil, where mortality can be up to five times higher in the most disadvantaged regions compared to the regions with the lowest mortality.3,41-43

Indeed, we found an inverse association between income level and screening coverage and lower coverage for countries classified as having fragmented health systems (SS/OP) (Table 4). As indicated in the methods section, the classification of health systems was done by the research team, which might be a major source of bias for this estimate; however, fragmented health systems usually have different screening guidelines and levels of programme organization making it difficult to achieve large population impact at the country level. Despite the significant association observed between screening coverage and the type of health system, careful



Fig. 1: Three-year interval cervical cancer screening coverage for women aged 25-65 years in the Americas region, estimates until 2019.

interpretation is warranted since countries classified as having no-fragmented health systems could have differential services for population groups and this category is composed mainly of small countries from the Caribbean. In contrast, Brazil, a country with a large population which is in this category, reports one of the lowest coverages in the region. The Brazilian data are based on the official cancer information system (SISCAN)<sup>9</sup>; however, governmental institutions such as the Brazilian National Cancer Institute describe the lack of consolidation of the system as a weakness and use the National Health Survey as an alternative source of information, reporting 81.3% coverage for the previous three years.<sup>44</sup> In fact, the low coverage reported by Brazil does not correspond to the mortality rate as compared with the remaining uppermiddle-income countries in the region.

A lack of agreement between screening coverage and cervical cancer mortality has been previously reported in Latin America.<sup>45,46</sup> Although it could be the result of lag time, it also highlights the relevance of other programme components, such as proper follow-up for positive-screened women (diagnostic work-up and CIN2+ treatment rates), particularly in areas with low access to health care.<sup>47–50</sup> In agreement with previous data, we found no association between screening coverage and cervical cancer mortality but did find a direct association of mortality with high inequality levels and higher mortality with the presence of screen-and-

Country	Target population in thousands	Previous year % (95% CI)	Previous 3 years % (95% CI)	Previous 5 years % (95% CI)	Ever in lifetime % (95% CI)
North America					
Bermuda	19.4	63 (59–67)	86 (80-92)	89 (83–96)	93 (86–100)
Canada	10,457.9	38 (37-40)	76 (73-78)	81 (78-84)	88 (84-91)
United States of America	87,449.4	43 (41-45)	71 (68–74)	77 (75–80)	87 (86–88)
Central America					
Belize	91.1	23 (18-31)	39 (32-46)	45 (38–52)	60 (54-67)
Costa Rica	1378.1	49 (45-52)	74 (69–78)	75 (71-80)	77 (73-81)
El Salvador	1661.1	44 (38-49)	73 (65-81)	80 (71-89)	88 (79–97)
Guatemala	3743.6	28 (21-36)	46 (38-53)	54 (46-61)	64 (57-72)
Honduras	2151.2	34 (28-39)	55 (46-64)	62 (53-72)	70 (60–80)
Mexico	33,090.4	37 (34-40)	66 (59–73)	76 (68-84)	88 (79–96)
Nicaragua	1605.2	40 (34-46)	67 (58–76)	76 (65-86)	84 (73-95)
Panama	1043.6	44 (37-51)	68 (60–76)	77 (69-84)	87 (79-95)
South America					
Argentina	11,288.7	49 (45-54)	77 (72–82)	81 (76-87)	90 (83-96)
Bolivia	2513.2	19 (15–24)	42 (36-48)	47 (40-55)	57 (47-67)
Brazil	58,778.6	12 (11–12)	34 (31–36)	40 (37-44)	58 (50–67)
Chile	5278.8	43 (40-45)	72 (68–75)	76 (72–80)	89 (84-94)
Colombia	13,525.0	53 (50-57)	68 (62-73)	73 (67-79)	80 (72-88)
Ecuador	4165.3	26 (20–32)	45 (39-51)	52 (45-58)	66 (61-71)
Guyana	181.1	5 (2-8)	13 (11-15)	17 (15–29)	21 (19–23)
Paraguay	1579.4	41 (36-46)	80 (70–90)	82 (73–92)	85 (76–94)
Peru	8255.2	17 (16–19)	58 (51-64)	70 (62–79)	84 (74-95)
Suriname	144.0	15 (10–20)	26 (20-33)	31 (25–39)	40 (32–49)
Uruguay	897.5	23 (21–25)	53 (49-57)	63 (58–67)	89 (85-92)
Venezuela	7237.7	24 (18–32)	39 (31–48)	45 (37–54)	54 (45-65)
Caribbean					
Antigua & Barbuda	28.1	31 (25–37)	60 (53–67)	73 (66–79)	85 (78–91)
Bahamas	109.8	35 (29–41)	54 (49–59)	64 (59–68)	73 (70–77)
Barbados	81.3	50 (47-53)	77 (73–79)	85 (83–87)	94 (91–96)
Cuba	3261.5	47 (42–51)	77 (72–82)	85 (79–91)	89 (83-96)
Dominica	19.5	28 (26–31)	57 (52–61)	67 (62–73)	78 (71–84)
Dominican Republic	2594.4	48 (43–52)	73 (66–79)	79 (73–86)	86 (79–93)
Grenada	28.9	26 (24–29)	54 (50–59)	69 (64–74)	84 (79–89)
Haiti	2498.5	2 (1-2)	4 (4–5)	7 (6-8)	9 (8–10)
Jamaica	768.6	22 (18–26)	52 (46–58)	67 (62–73)	83 (77–88)
Puerto Rico	831.0	48 (43-53)	79 (74–84)	86 (82–91)	91 (87–96)
Saint Kitts & Nevis	15.1	46 (42–50)	71 (66–75)	78 (74-82)	85 (81–89)
Saint Lucia	52.6	31 (27–34)	63 (58–68)	76 (71-82)	90 (84–96)
Saint Vincent & the Grenadines	28.3	23 (21–25)	54 (50–58)	68 (64-73)	83 (77-88)
Trinidad & Tobago	401.3	22 (20–24)	44 (41-47)	54 (51–58)	65 (62–69)
CI: Confidence Interval. Countries in bold surveys except for Antigua & Barbuda, Pa	letters correspond to countries prov nama, Suriname, and Venezuela we	iding data based on administr re no available data were four	rative sources. Data from the rem nd.	aining countries correspond to p	opulation-based household

Table 3: Estimated cervical cancer screening coverage in 2019 for women aged 25-65 years.

treat programmes. The latter may be a result of the higher percentage of women residing in distant areas with low access to health care even in countries with nofragmented health systems such as Colombia<sup>10</sup>; however, due to the relatively low number of observations (37 countries) these results should be cautiously interpreted.

Previous data on cervical cancer screening coverage have been reported in different studies, but it reflects the difficulties for standardized measurements between countries: different target populations (age ranges), different screening intervals, different periods, and different reports for the same country. This makes it difficult to compare countries or benchmarks among them to identify progress or a lack of programme organization. A recent report by the International Agency for Research on Cancer shows data on cervical cancer screening coverage from selected countries globally, highlighting the use of program data exclusively; despite the relevant effort and complex methodology, the report

Independent variable	N countries	OR	95% CI	p-value
A. Screening coverage <sup>a</sup>				
Income level				<0.001
High	12	Ref	Ref	
Upper-middle	20	0.58	0.38-0.89	
Lower-middle	4	0.52	0.25-1.07	
Low	1	0.05	0.01-0.29	
Predominant health system type				0.02
No-fragmented (NHS/NHI)	19	Ref	Ref	
Fragmented (SS/OP)	18	0.59	0.39-0.90	
Independent variable	N countries	RR	95%Cl	p-value
B. Cervical cancer mortality <sup>b</sup>				
Screen and treat approach				0.03
No/no program	18	Ref	Ref	
Yes	12	1.38	1.03-1.84	
Missing	1	-	-	
Inequality-adjusted HDI				0.01
Very high/High	5	Ref	Ref	
Medium	7	1.92	1.24-3.10	
Low	4	1.91	1.15-3.27	
Missing	5	-	-	

OR: Odds Ratio, RR: Rate Ratio, CI: Confidence Interval, NHS: National Health system, NHI: National Health Insurance, SS: Social Security, OP: Out of pocket, HDI: Human Development Index. Data in bold correspond to statistically significant associations. <sup>a</sup>Beta-regression model. <sup>b</sup>Poisson-regression model. Coverage for women aged 25-65 years, 3-years interval. For cervical cancer mortality screening coverage is included as an independent variable and missing categories are not included in the regression models.

Table 4: Context variables associated with screening coverage and cervical cancer mortality in multivariate analyses.

makes evident the challenges of deficient program organization (no standard denominators), the potential limitations of program data (most data correspond to a number of tests per year rather than coverage by screening interval and data outside the program are considered overestimation rather than low program coverage), and consequently the scarcity of robust data (only five countries from the Americas region and three of them with fragmented health systems reporting only the public sector).<sup>51</sup> An additional source of data on screening coverage has been the WHO survey on cancer country profiles.52 This report was traditionally restricted to a single coverage value, with origins in nonstandardized sources and including data only from the public sector excluding the social security in countries with fragmented health systems. Recently, cervical cancer profiles have been updated to include data based on the same methodology used here, as previously published.<sup>9</sup>

Our report has several strengths including the systematic search, the supplement with governmental data, and the standardised estimates; however, it also bears several limitations. Only three countries provided administrative data, and only Uruguay's data came from the cervical cancer screening programme, as data from Brazil and Colombia were generated from other sources of administrative information, such as reimbursement claims. As a result, the sources of information are mainly the same as those used in the previous reports we have criticized (e.g. household surveys). However, for a more accurate estimate and prediction we filtered by quality of data, gathered data directly from the source whenever possible, and developed strong statistical algorithms for data validation and analysis if required due to missing data. To assess the reliability of the methodology, we did an exhaustive sensitivity analysis of the missing data treatment assuming over and underestimation of coverage, obtaining almost identical coverage estimates.9 This approach reduced, but did not eliminate, the recall bias from household surveys; therefore, the 1-year screening interval might provide more accurate and reliable data than longer intervals. Moreover, we cannot control consent bias or other respondent biases in household surveys.53 In consequence, coverage overestimation could be present, with variable impact among the different countries.

We reflect that improving the quality of program data is imperative as a critical component of program organization; however, due to the limited capacity and resources at present, this objective may take a long time to achieve in LMIC. Nevertheless, the improvement of program organization will not solve challenges regarding health systems fragmentation. Therefore, strengthening health surveys might be a complementary measure by reviewing methodologies to reduce recall and social desirability biases.<sup>54</sup> Despite the limitations described, we used the most accurate source of information currently available, which may be the case for a medium to long period of time given the lack of population-based programmes in most countries of the region. We consider our report to be a baseline measure, and we expect periodic updates to help countries develop stronger monitoring systems and build stronger cervical cancer screening programmes as an essential component of the WHO elimination strategy.

#### Contributors

LB and RM conceptualised the project. LB, BS, ER and RM designed the study and planned the analysis. BS designed the data-extraction form, did the literature search. BS, GFD, MCM and RM contributed to the collection of data. ER and JSC did the formal statistical analysis. LB and BS supervised the statistical analyses. LB, ER and BS did WHO country consultation. BS, ER, RM, JSC cross-checked data. GFD, JSC, RM and ER prepared the tables. GFD and RM prepared the first draft of the manuscript. All authors contributed to data interpretation, critically revised subsequent drafts, and read and approved the study and had final responsibility for the decision to submit for publication.

#### Data sharing statement

The study's findings are supported by data available in public online repositories and data available upon request from the data provider. Produced estimates are published on WHO Global Health Observatory Data Repository (https://www.who.int/data/gho). Upon request, computer code is available in the IDIBELL repository (https://repository.idibell.cat).

#### Editor note

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#### Appendix A. Supplementary data

Supplementary data related to this article can be found at https://doi. org/10.1016/j.lana.2024.100689.

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