

Seasonal influenza vaccination in the Americas: update on current policies, use and progress in estimating vaccine effectiveness

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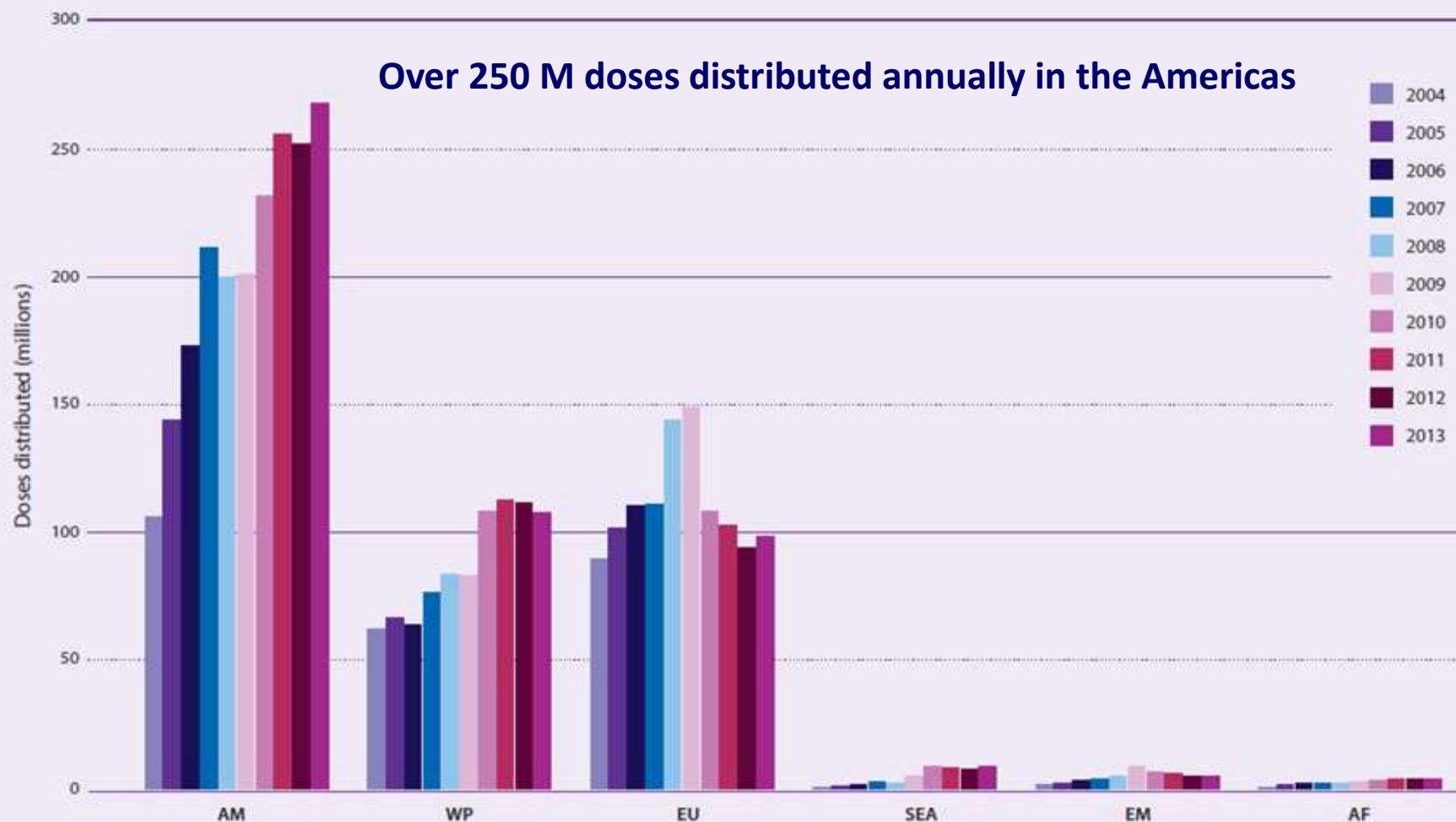
Red para la Evaluación de Vacunas En Latino América y el Caribe – *influenza*

Countries and Territories in the Americas with Policies for seasonal influenza vaccination, 2004-16

Number (%) of countries that have:	2004	2008	2016
Policies for influenza vaccination	13	35	40/52 (77%)
Vaccination of healthy children	6	22	25 (48%)
Vaccination of children with chronic diseases			5 (9%)
Vaccination of the elderly	12	33	39 (75%)
Vaccination of persons with chronic diseases	9	24	35 (67%)
Vaccination of health care workers	3	32	38 (73%)
Vaccination of pregnant women	3	7	31 (60%)

Distribution of influenza vaccines per WHO region, 2004-13

Over 250 M doses distributed annually in the Americas

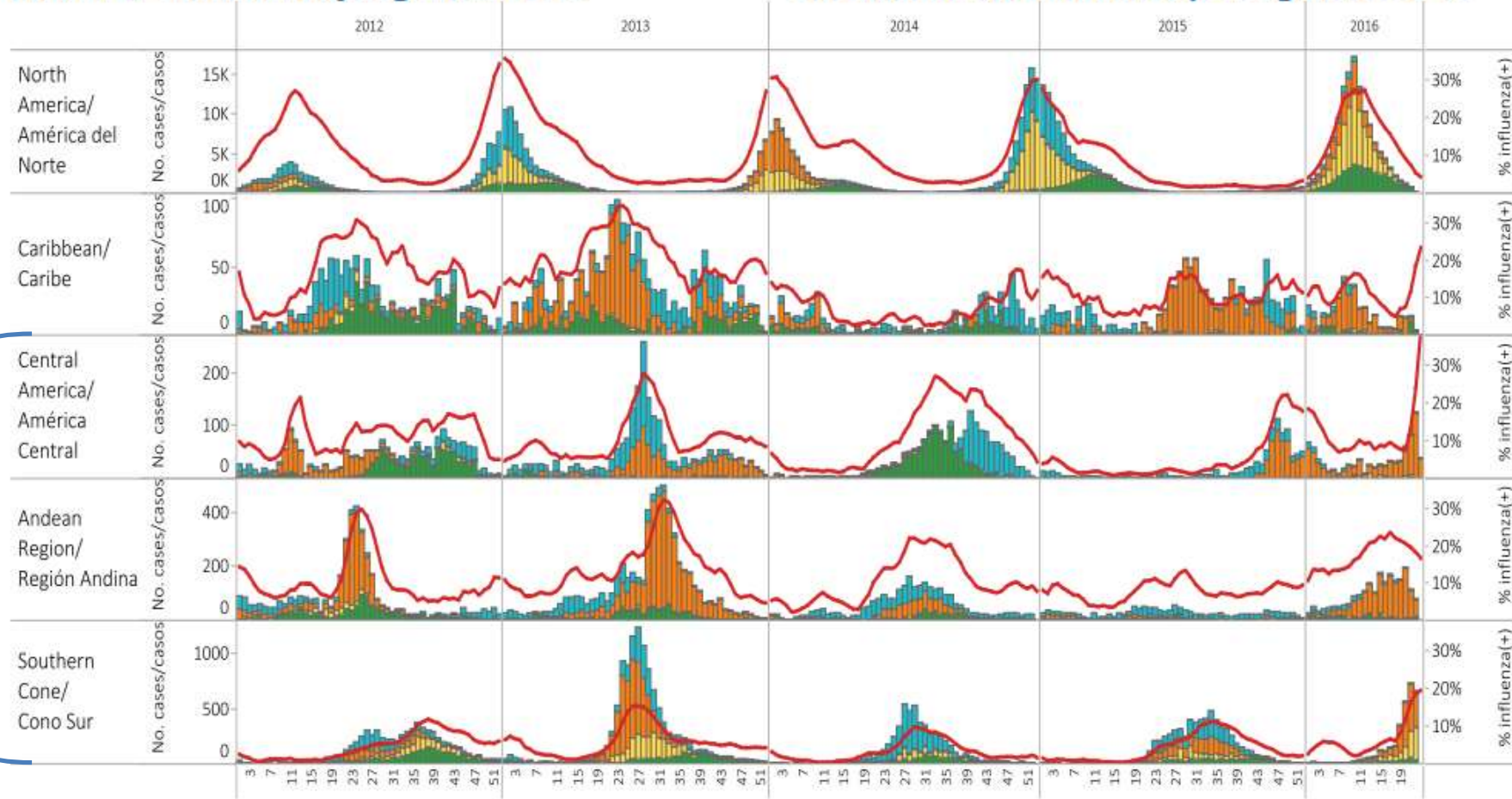


AF = Africa, SEA = South East Asian, EM = Eastern Mediterranean,
WP = Western Pacific, EU = European, AM = Americas

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Influenza circulation by region. 2012-16

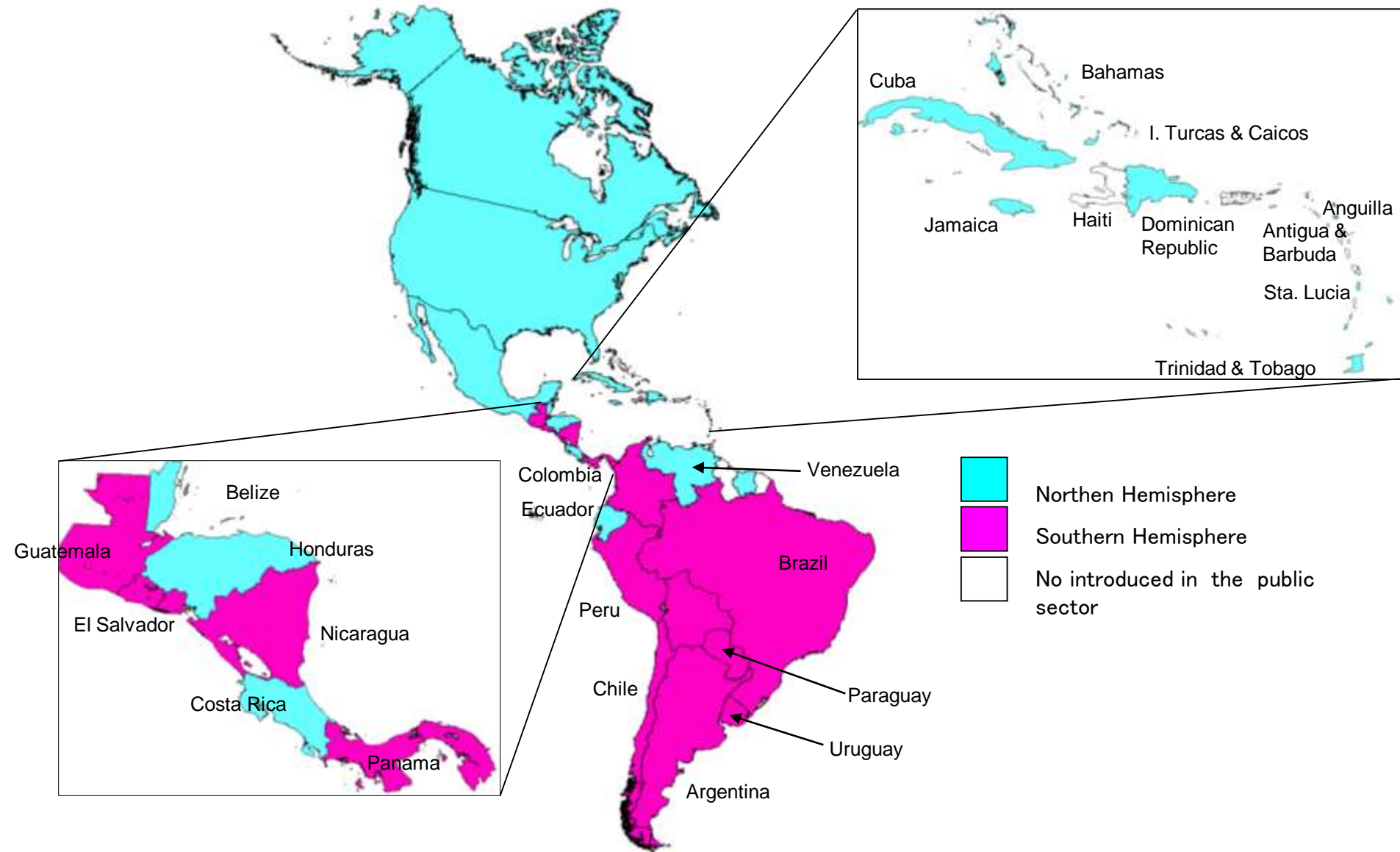
Circulación virus influenza por región. 2012-16



Influenza viruses/Virus influenza

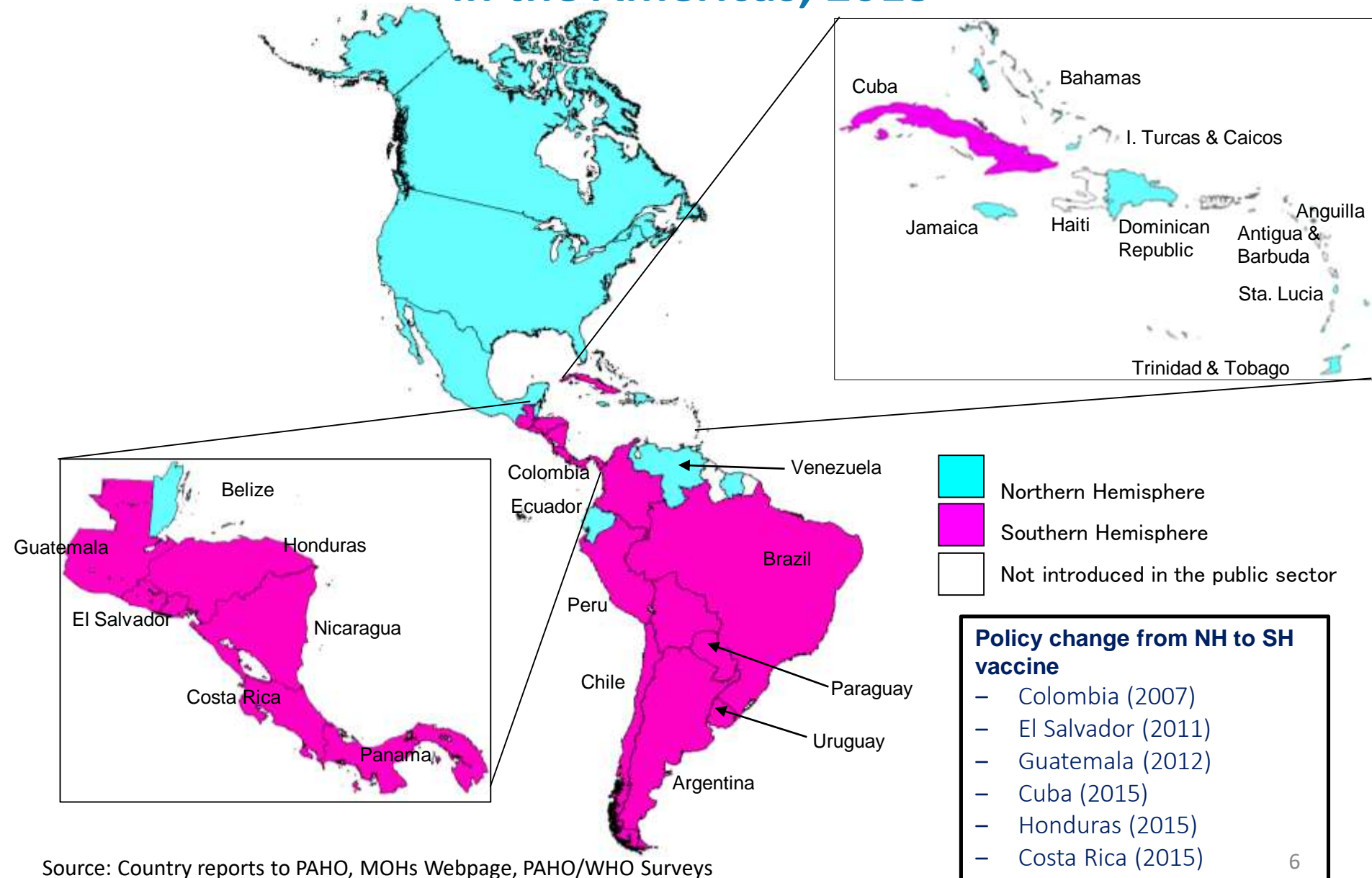
- Influenza A(H3N2)
- Influenza A(H1N1)pdm09
- Influenza A Not Subtyped/No subtipificado
- Influenza B
- Influenza A not subtypeable/no subtipifica...
- % Influenza

Use of seasonal influenza vaccine & formulation in the Americas, 2014



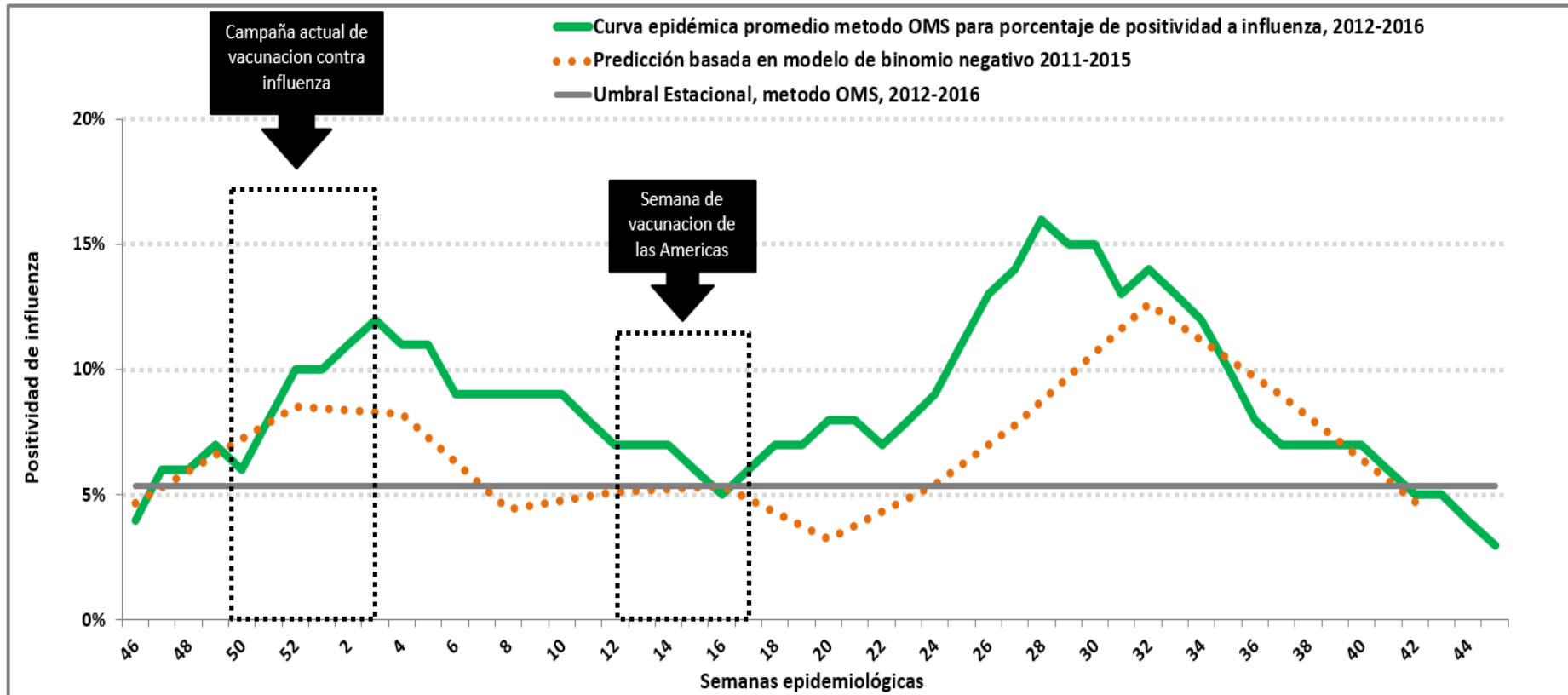
Source: Country reports to PAHO, MOHs Webpage, PAHO/WHO Surveys

Use and formulation of seasonal influenza vaccines in the Americas, 2015



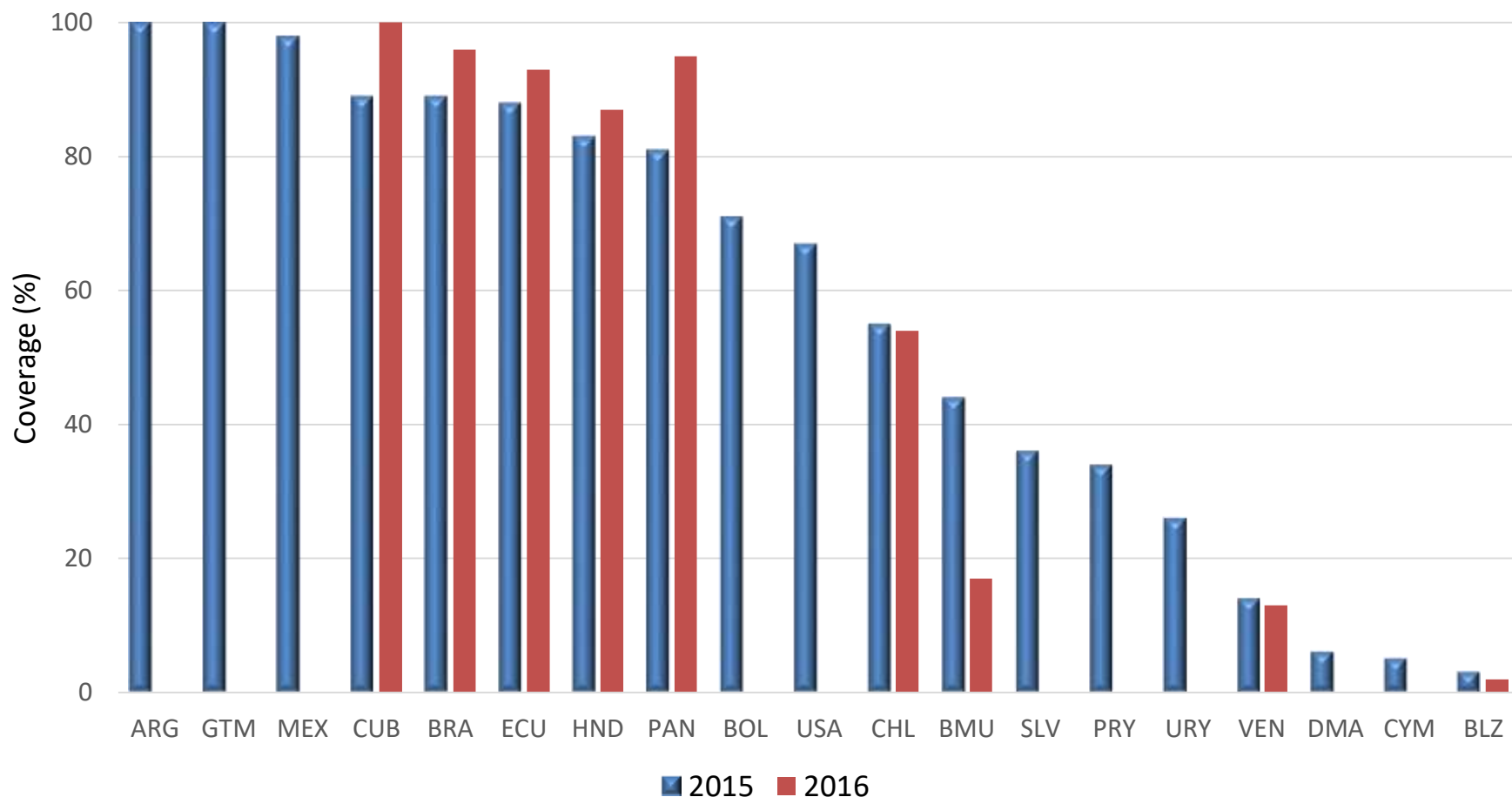
Source: Country reports to PAHO, MOHs Webpage, PAHO/WHO Surveys

Seasonality of influenza and vaccination, Ecuador 2011-16 (n=2,075)



Fuente: Ecuador. Vigilancia centinela de IRAG, datos virológicos reportados a FluNet 2011-2016.

Influenza vaccine coverage among the elderly Latin America and the Caribbean, 2015 and 2016*



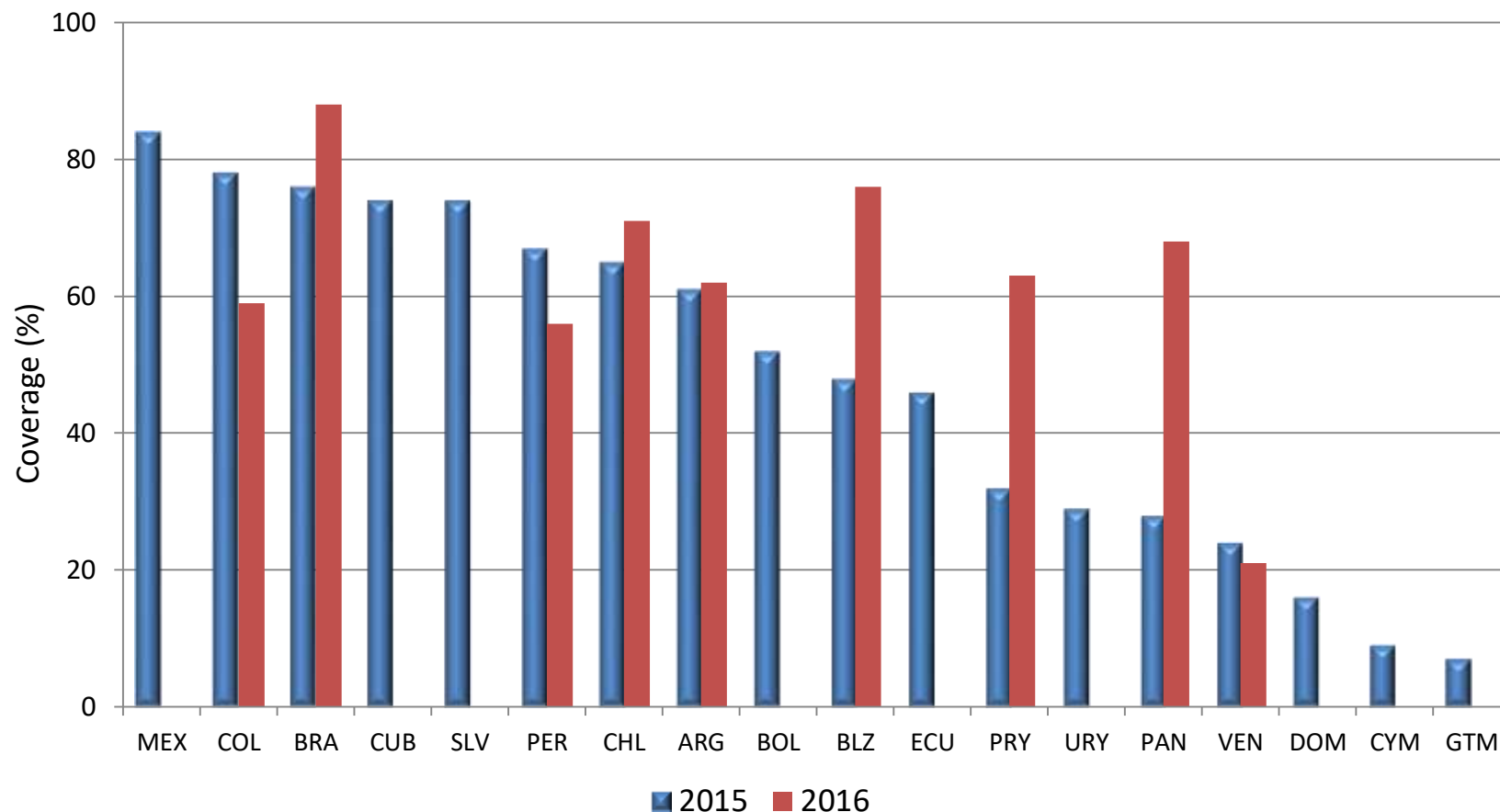
Source: Country reports through the PAHO-WHO Joint Reporting Forms (JRF), 2016 and 2017.

* Provisional data

Influenza vaccine coverage among children less than 5 years old

Latin America and the Caribbean, 2015-2016*

(full vaccination only)



Source: Country reports through the PAHO-WHO Joint Reporting Forms (JRF), 2016 and 2017.

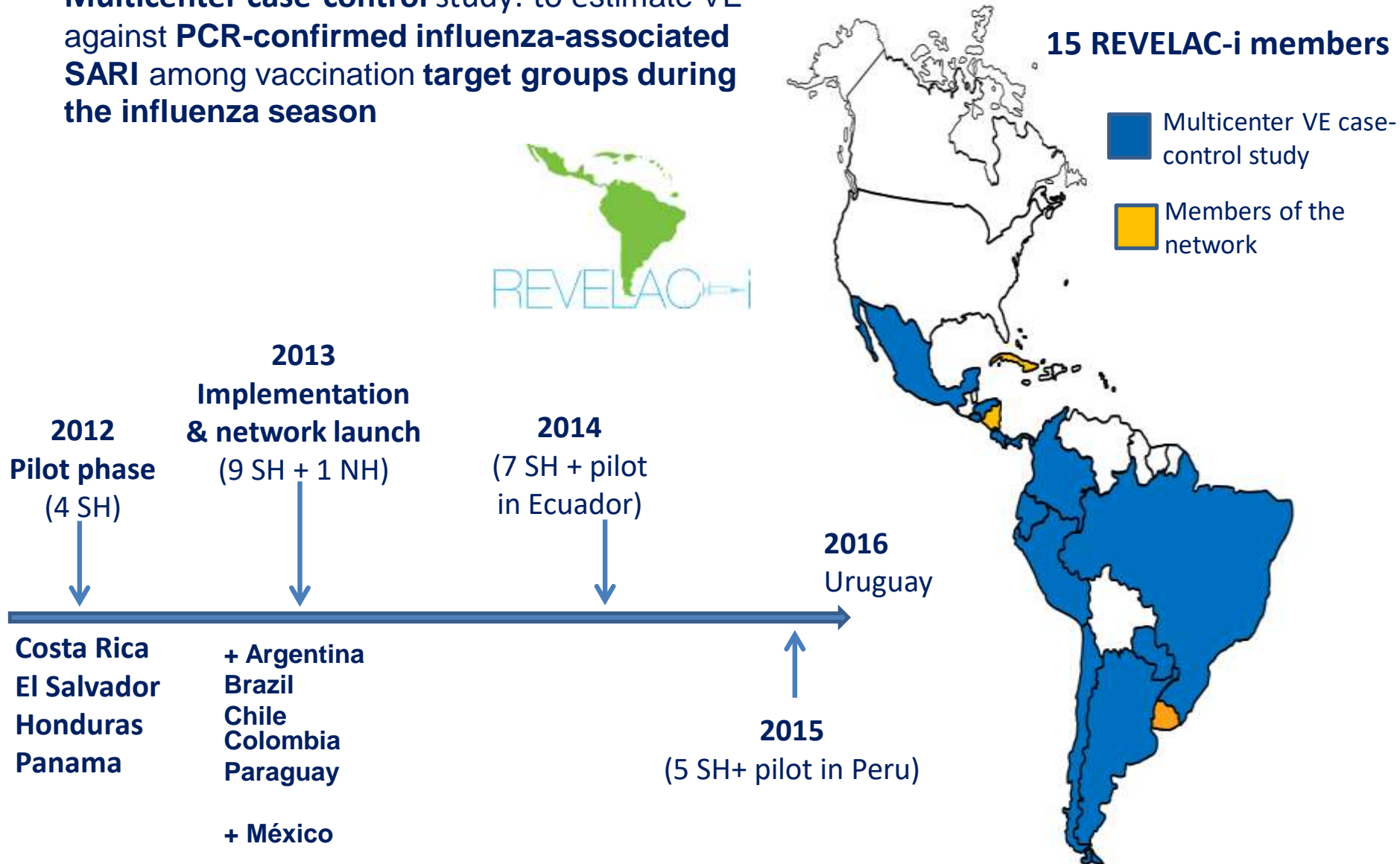
Note Pediatric coverage formula: $((2\text{nd dose} + \text{single dose}) / \text{denominator}) * 100$

* Provisional data

REVELAC-i network evolution

(Red para la Evaluación de Vacunas de Influenza en América Latina y el Caribe)

- **Multicenter case-control study:** to estimate VE against **PCR-confirmed influenza-associated SARI** among vaccination **target groups** during the **influenza season**



Network for Evaluation of Influenza Vaccine Effectiveness in Latin America and the Caribbean

REVELAC-i

Objectives:

Estimate the effectiveness of influenza TIV in preventing severe acute respiratory infections (SARI) laboratory-confirmed for influenza among EPI target groups during influenza seasons.



Building upon the existing regional SARI surveillance platform.



Using a common protocol, case-control (test-negative design).



RT-PCR laboratory confirmation for influenza.



Multidisciplinary efforts integrating influenza surveillance teams, reference laboratories and immunization programs

Revelac-i

Use of VE estimates in LAC

- Differences in use of VE between developed and developing countries
 - EPI provides vaccine free of charge and need to sustain investment in the vaccine (flu vaccine first target of budget cuts).
 - Show benefit of vaccination with current strategies as averted deaths and hospitalizations
- Not yet information for action, although REVELAC-i contributes to GIVE

Clinical, Epidemiological and Laboratory Investigation Form for Sentinel SARI Surveillance

Case Definition: An acute respiratory infection with history of fever or measured fever of $\geq 38^{\circ}\text{C}$, and cough, with onset within the last ten days, and requires hospitalization

1. CAPTURE			
1. Today's Date: / / <small>Day/month/year</small>		2. Case Code: _____	
4. Clinical History Number: _____		3. Establishment Name: _____	
6. Patient Name: _____		5. Capture Date: / / <small>Day/month/year</small>	
7. ID #: _____		8. Birth Date: / / <small>Day/month/year</small>	
10. Age Group: <input type="checkbox"/> <2y <input type="checkbox"/> 2-4y <input type="checkbox"/> 5-19y <input type="checkbox"/> 20-39y <input type="checkbox"/> 40-59y <input type="checkbox"/> 60+ y		9. Age: _____ <small>Years Months Days</small>	
11. Sex: <input type="checkbox"/> Male <input type="checkbox"/> Female			
2. CONTACT INFORMATION			
12. Telephone Number: _____			
13. Place of Residence: _____ <small>Department/Province/State Municipality/City Neighborhood</small>			
3. CLINICAL HISTORY/RISK FACTORS			
14. Did patient receive influenza vaccine during current season: <input type="checkbox"/> Yes <input type="checkbox"/> No <input type="checkbox"/> Unknown			
If yes, date of vaccination: / / <small>(Day/month/year)</small>			
If child, < 9 years: Specify number of doses and dates received: <input type="checkbox"/> 1: / / <small>(Day/month/year)</small> <input type="checkbox"/> 2: / / <small>(Day/month/year)</small>			
If child, < 6 months: Did mother receive influenza vaccine? <input type="checkbox"/> Yes <input type="checkbox"/> No <input type="checkbox"/> Unknown			
If yes, date of vaccination: / / <small>(Day/month/year)</small>			
Did mother breastfeed child? <input type="checkbox"/> Yes <input type="checkbox"/> No			
17. Risk Factors: <input type="checkbox"/> Yes <input type="checkbox"/> No			
Other Factors ¹		Comorbidities	
<input type="checkbox"/> Pregnant (<input type="checkbox"/> 1T <input type="checkbox"/> 2T <input type="checkbox"/> 3T)		<input type="checkbox"/> Asthma	
<input type="checkbox"/> Postpartum		<input type="checkbox"/> Chronic heart disease	
		<input type="checkbox"/> Chronic liver disease	
		<input type="checkbox"/> Chronic kidney disease	
		<input type="checkbox"/> Hematological disorders	
		<input type="checkbox"/> Obesity	
		<input type="checkbox"/> Other chronic pulmonary disease	
		<input type="checkbox"/> Diabetes	
		<input type="checkbox"/> Neurological or neuromuscular disorder	
		<input type="checkbox"/> Immunodeficiency (due to illness or treatment)	
		<input type="checkbox"/> Other: _____	
4. HOSPITALIZATION			
18. Fever onset date: / / <small>Day/month/year</small> EW _____		19. Hospitalization Date: / / <small>Day/month/year</small> EW _____	
20. Antiviral Use: <input type="checkbox"/> Not used <input type="checkbox"/> Oseltamivir <input type="checkbox"/> Zanamivir <input type="checkbox"/> Other		Antiviral Start Date: / / <small>Day/month/year</small>	
21. ICU Admission: <input type="checkbox"/> Yes <input type="checkbox"/> No		ICU Admission: / / <small>Day/month/year</small> EW _____	
22. Sample Collection: <input type="checkbox"/> Yes <input type="checkbox"/> No		ICU Discharge: / / <small>Day/month/year</small> EW _____	
1 st <input type="checkbox"/> Type: <input type="checkbox"/> Swab <input type="checkbox"/> Aspirate <input type="checkbox"/> Bronchial wash <input type="checkbox"/> Tissue <input type="checkbox"/> Serum <input type="checkbox"/> Other		Collection date: / / <small>Day/month/year</small>	
2 nd <input type="checkbox"/> Type: <input type="checkbox"/> Swab <input type="checkbox"/> Aspirate <input type="checkbox"/> Bronchial wash <input type="checkbox"/> Tissue <input type="checkbox"/> Serum <input type="checkbox"/> Other		Collection date: / / <small>Day/month/year</small>	
5. LABORATORY DATA			
23. Processing: _____		Receipt Date: / / <small>Day/month/year</small>	
1 st Sample: _____		Processed: <input type="checkbox"/> Yes (<input type="checkbox"/> PCR <input type="checkbox"/> IFA) <input type="checkbox"/> No, reason: / /	
2 nd Sample: _____		Processed: <input type="checkbox"/> Yes (<input type="checkbox"/> PCR <input type="checkbox"/> IFA) <input type="checkbox"/> No, reason: / /	
24. Results ² : <input type="checkbox"/> Positive <input type="checkbox"/> Negative		Delivery Date: (Day/month/year): / /	
<input type="checkbox"/> A, not subtyped <input type="checkbox"/> A(H1N1)pdm09 <input type="checkbox"/> A/H1N1 <input type="checkbox"/> A/H3N2			
<input type="checkbox"/> Influenza B <input type="checkbox"/> B(Victoria) <input type="checkbox"/> Influenza B(Yamagata)			
<input type="checkbox"/> RSV <input type="checkbox"/> Adenovirus <input type="checkbox"/> Parainfluenza I <input type="checkbox"/> Parainfluenza II <input type="checkbox"/> Parainfluenza III <input type="checkbox"/> Other			
6. DISCHARGE			
25. Discharge Date: / / <small>(Day/month/year)</small> EW _____		26. Outcome: <input type="checkbox"/> Discharged <input type="checkbox"/> Deceased <input type="checkbox"/> Transferred	
27. Date Case Closed: / / <small>(Day/month/year)</small>		Responsible Signature: _____	

¹ You may include other factor based on your specific circumstances, for example: ethnicity, HIV/AIDS, tuberculosis, obesity, alcoholism, smoking

² You may include other viruses depending on your laboratory capacity, for example, adenovirus, bocavirus, coronavirus, metapneumovirus, rhinovirus, etc.

Vaccination status ascertainment

SARI surveillance forms/databases

14. Tarjeta de vacuna: Si: <input type="checkbox"/> No: <input type="checkbox"/> No aplica: <input type="checkbox"/>	Aplicada		Nº de dosis aplicada	Fecha de última dosis		
15. Tipo de vacuna	Si	No		Día	Mes	Año
15.1 Anti Influenza						
15.2 Anti H. Influenzae b						
15.3 Anti Neumocócica						
15.4 23 valente						
15.5 Anti meningocócica (*)						
(*)Especifique nombre de la vacuna:						

1. Improve completeness at hospital level and collect additional variables if necessary (ex.doses)



2. Complete vaccination data a posteriori
Matching patients by ID, name, age, address

Electronic national nominal vaccination registry

Chile, Costa-Rica, Uruguay,
Colombia and partly in Panama

EPI records at local level (nominal in paper, or Excel, vaccination cards reviewed through household visits/calls)

Other countries

Study population

Country	Vaccination Target groups			N SARI hospitals
	Children	Elderly	Individuals with chronic conditions	
Argentina	6–24 months	≥65 years		4
Brazil	6–23 months	≥60 years		29
Chile	6–23 months	≥65 years	X	6*
Colombia	6–23 months	≥60 years		7
Costa Rica	6 months–10 years with chronic cond.	≥65 years		6*
Cuba	6–23 months	≥65 years		TBD
El Salvador	6–59 months	≥60 years		4*
Ecuador	6–59 months	≥50 years	X	3
Honduras	6–35 months with chronic cond.	≥60 years		3*
México	6–59 months; 3–9 years with chronic cond.	≥65 years	X	46
Panamá	6–59 months	≥60 years		10*
Paraguay	6–35 months	≥60 years		2
Peru	6–24 months	≥60 years		3
Uruguay	6–48 months	≥65 years		7*
TOTAL	6–59 months	≥60 years		130

*All SARI surveillance sentinel hospitals included

Influenza cases and controls included in regional VE analysis, REVELAC-i

2013

	Argentina	Brazil	Chile	Colombia	Costa-Rica	Honduras	Panama	Paraguay	El Salvador	Total
Influenza	36	277	133	69	71	6	20	75	7	694
(%)	5	40	19	10	10	0.9	3	11	1	100
Controles	54	728	616	165	79	3	49	222	10	1,926
(%)	2.8	37.8	31.98	8.57	4.1	0.16	2.54	11.53	0.52	100
Total	90	1,005	749	234	150	9	69	297	17	2,620
(%)	3	38	29	9	6	0.3	3	11	0.7	100

2014

	Argentina	Brazil	Chile	Colombia	Honduras	Paraguay	El Salvador	Total
Influenza	16	145	194	24	8	43	4	434
(%)	4	33	45	6	2	10	0.9	100
Controles	54	413	1,027	97	20	128	12	1,751
(%)	3	24	59	6	1	7	0.7	100
Total	70	558	1,221	121	28	171	16	2,185
(%)	3	26	56	6	1	8	0.7	100

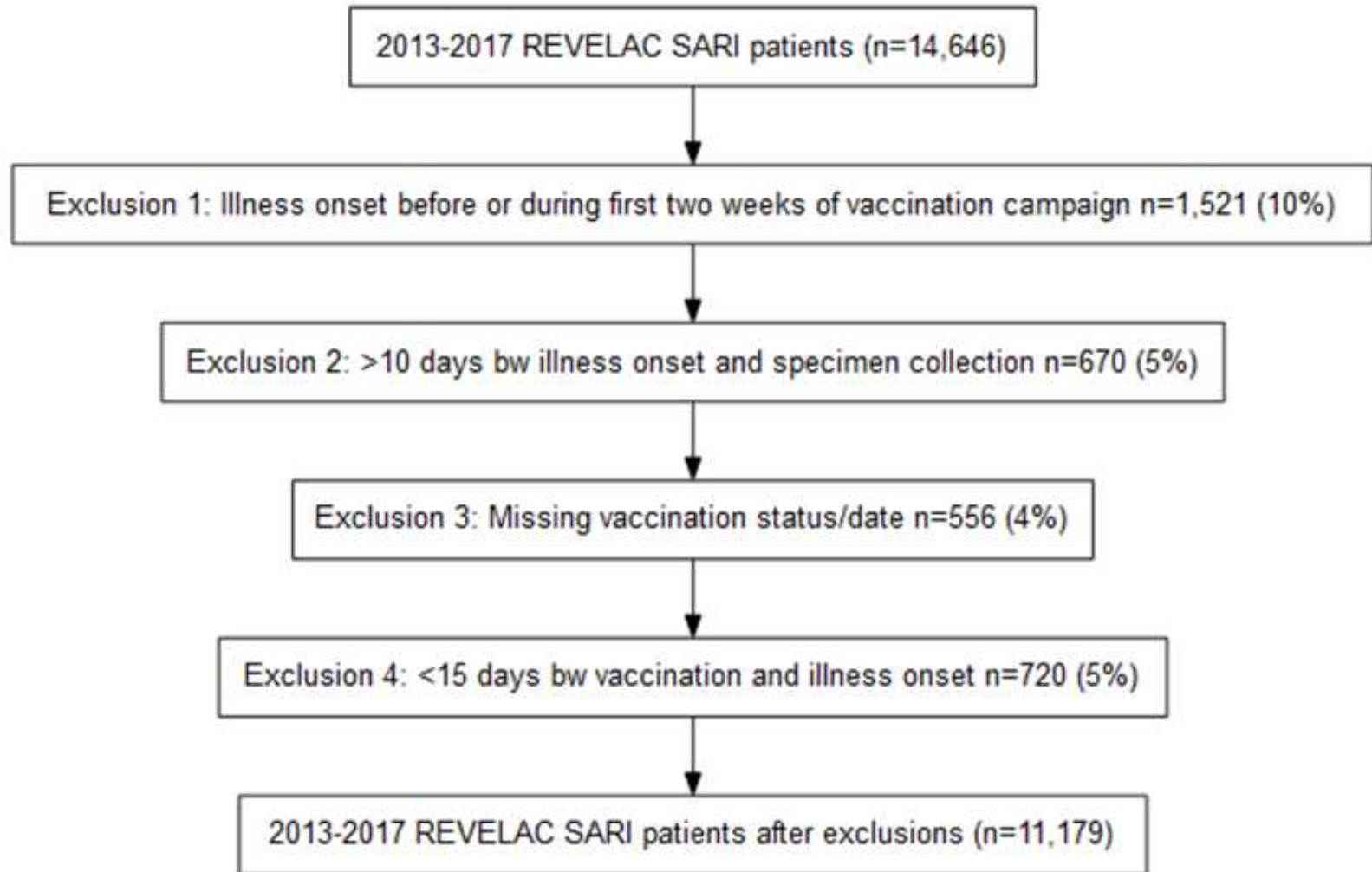
2015

2017: 2851 SARI patients, 587 flu(+) (ARG, CHI, PAR)

2016

	Chile	Colombia	Paraguay	Peru	Total		Argentina	Chile	Colombia	Paraguay	Uruguay	Total
Influenza	394	33	76	17	520	Influenza	37	226	11	77	67	418
(%)	75.8	6.3	14.6	3.3	100	(%)	8.9	54.1	2.6	18.4	16.0	100
Controles	1,705	90	222	149	2,166	Controles	209	1,048	93	224		1,574
(%)	79	4	10	7	100	(%)	13	67	6	14	0	100
Total	2,099	123	298	166	2,686	Total	246	1,274	104	301	67	1,992
(%)	78	5	11	6	100	(%)	12	64	5	15	3	100

REVELAC-I 2013-2017



REVELAC-I 2013-2017

Variable	Cases n=2,409 n(%)	Controls n=8,770 n(%)	p-value
Country			
Argentina	166 (7)	627 (7)	<0.001
Brazil	421 (17)	1148 (13)	
Chile	1300 (54)	5401 (62)	
Colombia	152 (6)	390 (4)	
Paraguay	328 (14)	954 (11)	
Uruguay	42 (2)	250 (3)	
Year			
2013	583 (24)	1855 (21)	<0.001
2014	415 (17)	1687 (19)	
2015	521 (22)	1995 (23)	
2016	397 (16)	1644 (19)	
2017	493 (20)	1589 (18)	
Gender			
Male	1116 (46)	4375 (50)	0.002
Female	1293 (54)	4395 (50)	
Age group			
0-18 years	679 (28)	3338 (38)	<0.001
19-64	399 (17)	1224 (14)	
65+	1331 (56)	4208 (48)	

REVELAC-I 2013-2017

Variable	Cases n=2,409 n(%)	Controls n=8,770 n(%)	p-value
≥one chronic condition	1573 (65)	5518 (63)	<0.001
Asthma	197 (8)	732 (8)	0.25
Diabetes	416 (17)	1176 (13)	<0.001
Respiratory disease	407 (16)	1595 (18)	0.31
Cardiovascular disease	323 (13)	1037 (12)	0.006
Liver disease	37 (2)	126 (1)	0.91
Renal disease	164 (7)	536 (6)	0.37
Obesity	113 (5)	323 (4)	0.05
Immune disease	94 (4)	479 (5)	0.001
Seasonal vaccination	897 (37)	4220 (48)	<0.001
Admitted to ICU	424 (18)	1433 (16)	0.02
Deceased	220 (9)	625 (7)	0.004
Days from illness onset to specimen collection			
0 to 2	576 (24)	2633 (30)	<0.001
3 to 5	57 (2)	393 (4)	
6 to 10	1776 (74)	5744 (65)	

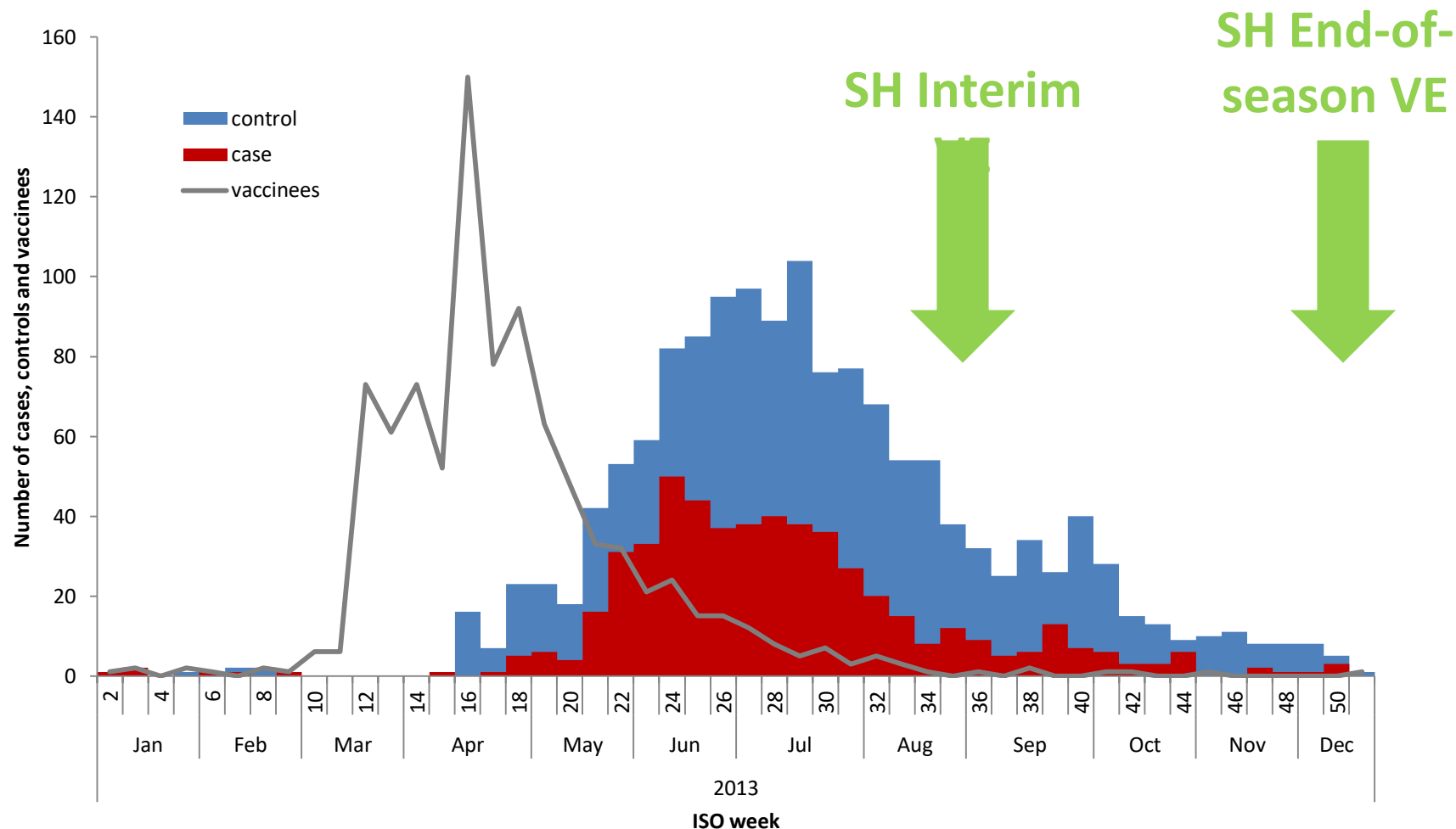
REVELAC Flu VE estimates 2013-2017

Table 2. Multiple Logistic Regression for VE

	Cases % vaccinated (n/N) %	Controls % vaccinated (n/N) %	aVE
Overall	897/2409 (37)	4220/8770 (48)	37 (30, 42)
2013	255/583 (44)	1070/1855 (58)	43 (21, 53)
2014	176/415 (42)	876/1687 (52)	37 (20, 51)
2015	169/521 (32)	876/1995 (44)	41 (27, 52)
2016	120/397 (30)	729/1644 (44)	50 (37, 61)
2017	177/493 (36)	669/1589 (42)	16 (-0.07, 34)
H3N2	431/1101 (39)	-	38 (30, 46)
H1N1p	276/769 (36)	-	45 (36, 53)
B	162/379 (43)	-	38 (25, 50)
Children	281/679 (41)	1709/3338 (51)	33 (20, 44)
Adults 19 to 64	100/399 (25)	423/1224 (35)	41 (22, 55)
Adults 65+	516/1331 (39)	2088/4208 (50)	37 (29, 45)

Adjusted for vaccination status, month of illness onset, pre-existing conditions, and age category/years.

Distribution of cases, controls per week of symptoms onset and vaccinees per week of vaccination, in 9 LAC countries, 2013



Influenza Vaccine Impact

Number of averted SARI hospitalizations by vaccination among children <5 years and the elderly, Chile 2013-14

Impacto Vacuna influenza en casos de IRA grave, niños de 6 a 23 meses, Chile, 2013



2013

Impacto vacuna Influenza en IRA grave, mayores de 65 años, Chile, 2013



Impacto Vacuna influenza en casos de IRA grave, niños de 6 a 23 meses, Chile, 2013				Impacto vacuna Influenza en IRA grave, mayores de 65 años, Chile, 2013			
Mes	Observados	Evitados	Total	Mes	Observados	Evitados	Total
Abril	0	0	0	Abril	50	0	50
Mayo	40	33	73	Mayo	250	75	325
Junio	20	16	36	Junio	550	181	731
Julio	150	160	310	Julio	450	145	595
Agosto	130	125	255	Agosto	180	58	238
Septiembre	10	0	10	Septiembre	250	77	327
Octubre	50	37	87	Octubre	120	10	130
Noviembre	0	0	0	Noviembre	150	45	195
Diciembre	20	10	30	Diciembre	150	0	150

Impacto vacuna Influenza en IRA grave, mayores de 65 años, Chile, 2014

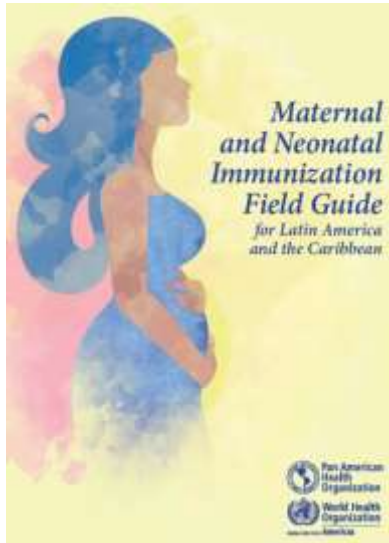


2014

Challenges for REVELAC-i

- Sustainability of the platform
 - High turnover of surveillance and EPI staff,
 - Uses existing surveillance platform (no additional resources for staff or research groups)
 - Information systems/integration
 - Political changes
- Vaccination status retrieval and timeliness of interim VE estimates
- Sample size
- Translation of evidence for decision makers

31/35 countries that vaccinate pregnant women against influenza are located in the Americas



Technical guidelines for Maternal and neonatal immunization:

- ✓ Vaccination recommendations
- ✓ Available vaccines
- ✓ Current evidence
- ✓ Operational aspects
- ✓ Communication
- ✓ Best practices



Factors associated with a successful expansion of influenza vaccination among pregnant women in Nicaragua

Carmen S. Arriola^{a,b,*}, Nancy Vasconez^c, Mark Thompson^b, Sara Mirza^b, Ann C. Moen^b, Joseph Bresee^b, Ivy Talavera^c, Alba María Roperio^d

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In 2013, Nicaragua expanded recommendations to include influenza vaccination to all pregnant women in Managua:

- Survey among 1,807 pregnant women who delivered at public hospitals in Managua to evaluate the uptake of influenza vaccination and factors associated with vaccination.
- Four antenatal visits were associated with receipt of influenza vaccination (OR=2.6 [1.15; 5.81]).
- Receipt of influenza vaccination recommendation from a health care provider was positively associated with receipt of influenza vaccination (OR=14 [10; 19]).

**1st network meeting
La Antigua Guatemala, 2013**



**2nd network meeting,
Cartagena de Indias, 2014**



**3rd network meeting,
Santiago de Chile, March 2016**

GRACIAS!

<http://www.paho.org/revelac-i/>