



#### Studies on Pneumococcal vaccines in Colombia

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#### Introduction



- Colombia introduced pneumococcal conjugate vaccines (PCV) in 2007.
- Heptavalent (7v) was used first and then changed to decavalent (10v)
- Initially, PCV7 targeted geographical areas with high infant mortality rate.
- In 2012, 10v vaccine was universally adopted for newborns all around the country.

#### Introduction process



- Studies conducted:
  - Several economical and epidemiological analysis were carried out before a decission was made.
  - The first one was conducted in 2008 focusing on the use of the 7v.
  - The second one was conducted in 2010 estimating the economical and epidemiological impact of the 13v, 10v, and 7v.

## Introduction process



#### • Burden of disease before vaccine introduction:



## Cost-effectiveness of pneumococcal conjugate vaccines of 7, 10, and 13 valences in Colombian children

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# Burden of disease before vaccine introduction. Children < 2 years.

Events	No vaccination (NV)
AOM Outpatient RX pneumonias	767,133 (701,326-812,796) 17,548 (14,670-24,020)
Inpatient RX pneumonias	31,197 (26,080-42,702)
Pneumococcal meningitis	464 (380-633)
Deaths by outpatient pneumonia	526 (293-1441)
Deaths by inpatient pneumonia	936 (522-2562)
Deaths by meningitis	172 (125-342)
Total deaths	1634 (940-4345)
YLL by Pneumonias	107,519 (59,922-294,351)
Total YLL	12,619 (9209–25,115) 120,138 (69,130–319,465)



### Burden of disease before vaccine introduction. Children < 2 years.

Cost	No vaccination
Vaccination's costs	US\$ 0
AOM's cost	US\$ 94,295,343 (US\$ 68,965,093-119,889,796)
RX	US\$ 38,503,110 (US\$ 25,750,079-63,243,236)
pneumonias' cost	
Pneumococcal meningitis' cost	US\$ 666,000 (US\$ 435,928-1,089,844)
Total cost (included vaccination)	US\$ 133,464,453 (US\$ 95,151,100-184,222,876)
YLL	44,911 (13,429-319,465)
(discounted)	
Cost	US\$ 124,421,317 (US\$ 82,464,435-184,222,876)
(discounted)	

Castañeda C et al 2012. Vaccine 30:1936-1943

## Impact of 7v pneumococcal vaccine introduction in Bogota



	Vaccine 31 (2013) 4033-4038	
	Contents lists available at SciVerse ScienceDirect	<b>x</b>
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Changes in *Streptococcus pneumoniae* serotype distribution in invasive disease and nasopharyngeal carriage after the heptavalent pneumococcal conjugate vaccine introduction in Bogotá, Colombia<sup>\*</sup>

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- Cross sectional surveys before and after 7v vaccine universal introduction in Bogota.
- Children < 2 years
- Nasopharyngeal swabs to detect *S pneumoniae* carriers.
- First survey May 2005 November 2005 (n=246). Prevaccine
- Second survey June 2011- November 2011 (n=197).
   Postvaccine

## Impact of 7v pneumococcal vaccine introduction in Bogota



 Analysis of serotype distribution among invasive isolates of S pneumoniae collected from national surveillance by the Colombian National Institute of Health.

• 2005-2009. Prevaccination period

#### • 2010-2011. Postvaccination period.

Parra E et al 2013. Vaccine 31:4033-4038



## Impact of 7v pneumococcal vaccine introduction.

#### Table 2 Frequency of PCV and non-PCV serotypes found in carriage and IPD in children.

Serotypes	NP Carriage		Р	IPD		Р
	Unvaccinated n - 246	Vaccinated n <del>-</del> 197		2005/09 n <b>-</b> 340	2010/11 n <b>-</b> 84	
	n (%)			n (%)		
PCV7						
4	2 (0.8)	0	-	5 (1.5)	1(1.2)	0.66
6B	12 (4.9)	2(1.0)	0.02	26 (7.6)	4 (4.8)	0.49
9V	0	0	0	8 (2.4)	2 (2.4)	0.62
14	9(3.7)	3(1.5)	0.16	163 (48)	14 (16.7)	< 0.001
18C	3 (1.2)	2(1.0)	0.83	11 (3.2)	5 (6.0)	0.19
19F	25 (10.2)	3 (1.5)	<0.01	13 (3.8)	2 (2.4)	0.40
23F	7 (2.8)	5 (2.5)	0.84	9 (2.6)	4 (4.8)	0.24
All PCV7	58 (23.6)	15 (7.5)	< 0.001	235 (69.1)	32 (38)	< 0.001

#### Parra E et al 2013. Vaccine 31:4033-4038



#### Table 2 Frequency of PCV and non-PCV serotypes found in carriage and IPD in children.

Serotypes	NP Carriage		Р	IPD		Р	
	Unvaccinated Vaccinated n-246 n-197		2005/09 n <b>-</b> 340	2010/11 n <b>-</b> 84			
	n (%)			n (%)			
PCV13							
3	1 (0.4)	1 (0.5)		6(1.8)	12(14.3)	< 0.001	
6A	23 (9.3)	6 (3.0)	<0.001	16 (4.7)	1 (1.2)	0.12	
19A	3 (1.2)	3 (1.5)	0.78	10 (2.9)	5 (6.0)	0.15	
6A/B	1 (0.4)	2(1.0)		0	0	-	
All PCV13	86 (34.9)	27 (13.5)	<0.001	304 (89.4)	59 (70.2)	0.002	
Non-PCV							
All Non-PCV	59 (24.0)	67 (34.0)		38 (8.2)	25 (29.8)	<0.00	
19A 6A/B All PCV13 Non-PCV All Non-PCV	3 (1.2) 1 (0.4) 86 (34.9) 59 (24.0)	3 (1.5) 2 (1.0) 27 (13.5) 67 (34.0)	0.78 <0.001	10 (2.9) 0 304 (89.4) 38 (8.2)	5 (6.0) 0 59 (70.2) 25 (29.8)		

Parra E et al 2013. Vaccine 31:4033-4038





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Procedia in Vaccinology 6 (2012) 150 - 155

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www.elsevier.com/locate/procedia

5th Vaccine and ISV Global Annual Congress

## Evaluating the effectiveness of conjugated pneumococcal vaccines in Colombia

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- Retrospective cohort study among children <2 years vaccinated in the private market.
- Cohort ensambled before 7v vaccine was available in EPI.
- Cross comparation between:

   children with complete 7v vaccination schedule
   vs
  - children not vaccinated.



 2631 children < 2 years affiliated to a major Health Management Organization (HMO-EPS in Spanish)

#### • Outcomes:

- Hospitalization due to acute respiratory infections (bronchitis, bronchiolitis, and pneumonia).
- Hospitalization due to sepsis or meningitis
- Hospitalization due to otitis media.



• Vaccine Effectiveness was adjusted by:

- Age
- Sex
- Previous hospitalization.
- Economical level



#### • Results:

- -877 children vaccinated
- -1,754 children unvaccinated
- -42 children hospitalized among vaccinated
- -114 children hospitalized among unvaccinated.



#### • Results:

- -877 children vaccinated. 42 hospitalized during follow up
- 1,754 children unvaccinated. 114 hospitalized during follow up.
- Hazzard Ratio (HR)=0.58 (0.41-0.82) for all hospitalization. - 42% effectiveness against hospitalization.



#### • Results:

#### Hazzard Ratio (HR)=0.51 (0.31-0.85) for pneumonia hospitalization.

#### -49% effectiveness against pneumonia hospitalization.





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Fig. 1. (a) Function of survival for all events potentially related with infection by Streptococcus pneumoniae. (b) Function of survival for pneumonia admission.



• Case control study conducted in 2013 and 2014:

- -MOH
- -CDC
- PAHO
- Cases: patients with invasive disease by S pneumoniae confirmed by isolate from sterile sites.



• Controls: 4 Healthy children per case.

• Controls selected from the same communities than cases.

• 32 cases and 132 controles were identified.

• Meningitis (n=12), neumonía (n=11), sepsis (n=9).



- 21 cases had serotypes contained in PCV10.
  - 23F. N=6
  - 14. N=4
  - 19F. N=4
- Frequency of vaccination was similar for cases and controls:
  - 94% cases
  - 95% controls



 However, timely vaccination with PCV10 was more frequent among controls:

- 25% among cases.
- 45% among controls.
- OR=0.16 (0,05-0,5)

## Surveillance of serotypes changes after vaccine introduction.



## Surveillance of serotypes changes after vaccine introduction.





Camacho G, Leal AL, Patiño J, Moreno V et al. Caracterización clinica, microbiologica y epidemiologica de la enfermedad neumococica invasiva en población pediatrica de 10 hospitales de Bogota, Colombia 2008-2017. 11- ISPPD, Melbourne 2018

### Conclusions



- Colombia has successfully introduced PCV7 first and PCV10 lately.
- Both vaccines have had an impact on distribution of serotypes, carriage and invasive disease.
- Effectiveness against clinical outcomes have been demonstrated for both of them.
- Changes in the distribution of serotypes have been demonstrated after PCV7 and PCV10 introduction.
- Still, efforts to improve opportune access to timely vaccination should be reinforced.

### Conclusions



- Potential Research questions:
  - How ecological changes on decrease of mortality on respiratory diseases can be attributed to pneumococcal vaccine alone or to a compound of actions inclouding other vaccines introduction?
  - What is the efficacy of PCV10 in older adults?
  - Changes in serotype 19 A are permanent? Cyclic?
  - Is really outside there an effectiveness of PCV10 on 19<sup>ª</sup> invasive infections?

## **Gracias-Thanks**

