Supplementary Online Content

Neal EFG, Nguyen C, Ratu FT, et al. A comparison of pneumococcal nasopharyngeal carriage in very young Fijian infants born by vaginal or cesarean delivery. *JAMA Netw Open.* 2019;2(10):e1913650. doi:10.1001/jamanetworkopen.2019.13650

eFigure. Directed Acyclic Graph to Select Factors for Statistical Adjustment **eAppendix.** Dagity Code for eFigure, Directed Acyclic Graph to Identify a Minimal Set of Factors for Statistical Adjustment

eTable 1. Comparison of Serotype-Specific Carriage in Fijian Infants Aged 5 to 8 Weeks, by Infant Mode of Delivery

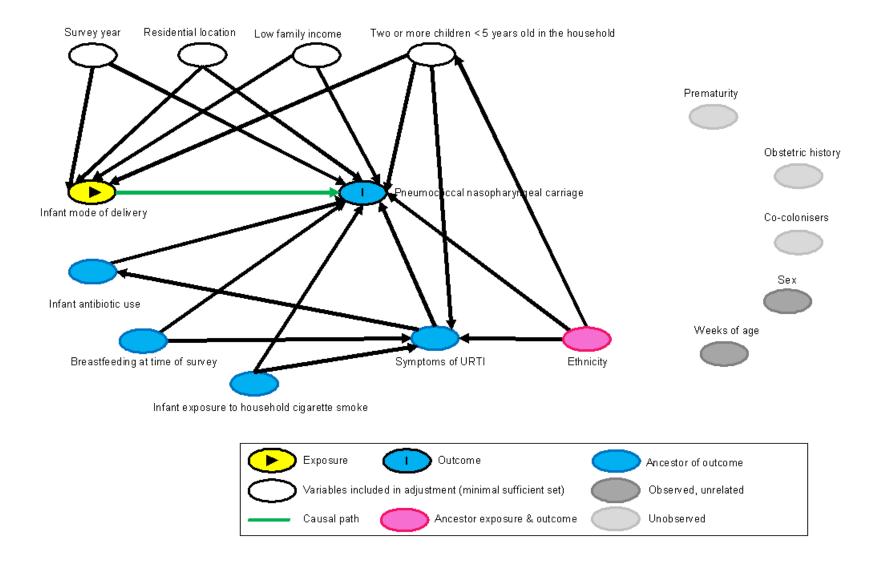
eTable 2. Correlation Between Exposures Included in Regression Models Assessed Using Pearson r

eTable 3. Associations Between Infant Mode of Delivery and Pneumococcal Carriage Density in Fijian Infants Aged 5 to 8 Weeks, in 4 Annual Pneumococcal Carriage Surveys

eReferences

This supplementary material has been provided by the authors to give readers additional information about their work.

eFigure. Directed Acyclic Graph to Select Factors for Statistical Adjustment



eAppendix. Dagity Code for eFigure, Directed Acyclic Graph to Identify a Minimal Set of Factors for Statistical Adjustment ¹

To recreate the directed acyclic graph in eFigure, please go to http://www.dagitty.net/dags.html# and copy and paste the code below in to the model code window on the bottom right of the screen, then select "Update Model", immediately below the model code. Finally, select "cancel" for the new variable option \(^1\).

Breastfeeding%20at%20time%20of%20survey 1 @0.309,0.771

Co-colonisers U @0.579,0.451

Infant%20antibiotic%20use 1 @0.258,0.727

Infant%20ethnicity 1 @0.498,0.813

Infant%20exposure%20to%20household%20cigarette%20smoke 1 @0.376,0.815

Infant%20mode%20of%20delivery E @0.264,0.603

Low%20family%20income A @0.395,0.270

Obstetric%20history U @0.568,0.388

Pneumococcal%20nasopharyngeal%20carriage O @0.410,0.595

Prematurity U @0.550,0.330

Residential%20location A @0.324,0.262

Sex 1 @0.575.0.528

Survey%20year A @0.262,0.260

Symptoms%20of%20URTI 1 @0.433,0.742

 $Two \% 20 or \% 20 more \% 20 children \% 20 \% 3 C\% 205\% 20 years \% 20 in \% 20 the \% 20 household\ A\ @0.482, 0.270$

Weeks%20of%20age 1 @0.552,0.593

Breastfeeding% 20at% 20time% 20of% 20survey Pneumococcal% 20nasopharyngeal% 20carriage Symptoms% 20of% 20URTI

Infant%20antibiotic%20use Pneumococcal%20nasopharyngeal%20carriage

Infant% 20ethnicity Pneumococcal% 20nasopharyngeal% 20carriage Symptoms% 20of% 20URTI

Two% 20or% 20more% 20children% 20% 3C% 205% 20years% 20in% 20the% 20household

Infant% 20exposure% 20to% 20household% 20cigarette% 20smoke

Pneumococcal%20nasopharyngeal%20carriage Symptoms%20of%20URTI

Infant%20mode%20of%20delivery Pneumococcal%20nasopharyngeal%20carriage

 $Low \%\,20 family \%\,20 income\,Infant \%\,20 mode \%\,20 of \%\,20 delivery\,Pneumococcal \%\,20 nasopharynge al \%\,20 carriage$

 $Residential \% 20 location\ Infant \% 20 mode \% 20 of \% 20 delivery\ Pneumococcal \% 20 nasophary ngeal \% 20 carriage$

 $Survey \% 20 year\ Infant \% 20 mode \% 20 of \% 20 delivery\ Pneumococcal \% 20 nasopharyngeal \% 20 carriage$

 $Symptoms \%\,20 of \%\,20 URTI\,In fant \%\,20 antibiotic \%\,20 use\,\,Pneumococcal \%\,20 nasophary ngeal \%\,20 carriage\,\,20 to 100 to 100$

Two % 20 or % 20 more % 20 children % 20 % 3 C% 205% 20 years % 20 in % 20 the % 20 household

Infant%20mode%20of%20delivery Pneumococcal%20nasopharyngeal%20carriage Symptoms%20of%20URTI

eTable 1. Comparison of Serotype-Specific Carriage in Fijian Infants Aged 5 to 8 Weeks, by Infant Mode of Delivery Vaginal delivery Cesarean delivery Serotypes n = 1,722n = 260P-value^a n, % [95% CI] n, % [95% CI] **PCV10** serotypes **1**b 2, 0.12 [0.01-0.42] **4**b 6, 0.35 [0.13-0.76] 6Bb 26, 1.51 [0.99-2.20] 2, 0.77 [0.09-2.75] .57 7Fb 7, 0.41 [0.16-0.84] 9Vb 6, 0.35 [0.13-0.76] -14^b 9, 0.52 [0.24-0.99] 18Cb 5, 0.29 [0.09-0.68] _ 19Fb 34, 1.97 [1.37-2.75] 3, 1.15 [0.24-3.33] .47 23Fb 23, 1.34 [0.85-2.00] 3, 1.15 [0.24-3.33] 1.0 Non-PCV10 serotypes 3 5, 0.29 [0.09-0.67] 6Ab 23, 1.34 [0.85-2.00] 3, 1.15 [0.24-3.33] 1.0 6C 18, 1.05 [0.62-1.65] 3, 1.15 [0.24-3.33] .74 6D 9, 0.52 [0.24-0.99] 1 0.38 [0.01-2.12] 1.0 7B 2, 0.12 [0.01-0.42] 7C 9, 0.52 [0.24-0.99] 2, 0.77 [0.09-2.75] .65 8 1, 0.06 [0.00-0.32] 10A 1, 0.06 [0.00-0.32] -10B 4, 0.23 [0.06-0.59] 1, 0.38 [0.01-2.12] .51 19, 1.10 [0.67–1.72] 11A 2, 0.77 [0.09-2.75] 1.0 12F 2, 0.12 [0.01-0.42] -2, 0.77 [0.09–2.75] 13 20, 1.16 [0.71–1.79] .75 15A 1, 0.06 [0.00-0.32] 15B/C 39, 2.26 [1.62–3.08] 6, 2.31 [0.85-4.95] 1.0 16F 31, 1.80 [1.23-2.55] 1, 0.38 [0.01-2.12] .11 17F 13, 0.75 [0.40-1.29] 2, 0.77 [0.09-2.75] 1.0 18A 5, 0.29 [0.09-0.68] 19Ab 19, 1.10 [0.67-1.72] 1, 0.38 [0.01-2.12] .50 19B 6, 0.35 [0.13-0.76] 20 3, 0.17 [0.04-0.51] _ _ 20B 2, 0.12 [0.01-0.42] _ 21 16, 0.93 [0.53-1.50] 5, 1.95 [0.63-4.43] .18 22F 8, 0.46 [0.20-0.91] 1, 0.38 [0.01-2.12] 1.0 14, 0.81 [0.45–1.36] 23A 23B 6, 0.35 [0.13-0.76] 1, 0.38 [0.01-2.12] 24B 2, 0.12 [0.01-0.42] .34 24F 5, 0.29 [0.09-0.68] -29 2, 0.12 [0.01-0.42] 31 5, 0.29 [0.09–0.68] _ _ 33A 1, 0.06 [0.00-0.32] _ 33B 5, 0.29 [0.09-0.68] 1, 0.38 [0.01-2.12] .57

33F

3, 0.17 [0.04-0.51]

	Vaginal delivery	Cesarean delivery	
Serotypes	(n = 1,722)	(n = 260)	<i>P</i> -value ^a
	n, (%) [95% Cl]	n, (%) [95% Cl]	
34	15, 0.87 [0.49–1.43]	2, 0.77 [0.09–2.75]	1.0
35A	7, 0.41 [0.16–0.84]	-	-
35B	23, 1.38 [0.85–2.00]	1, 0.38 [0.01–2.12]	.35
35F	1, 0.06 [0.00–0.32]	-	-
38	5, 0.29 [0.09–0.68]	1, 0.38 [0.01–2.12]	.57
39	4, 0.23 [0.06–0.59]	-	-
NT2	22, 1.28 [0.80–1.93]	2, 0.77 [0.09–2.75]	.76
NT2/NT3b	14, 0.81 [0.45–1.36]	2, 0.77 [0.09–2.75]	1.0
NT3b	16, 0.93 [0.53–1.50]	4, 1.54 [0.42–3.89]	.32
NT4	4, 0.23 [0.06–0.59]	-	-

Abbreviation: CI Confidence interval; ^a P-values were determined from Fisher's exact test; ^b Serotypes included in PCV13

eTable 2. Correlation Between Exposures Included in Regression Models									
Assessed Using Pearson r									
Characteristics	Infant mode of delivery	Survey year	Residential location	Low family income	≥ 2 children < 5 y in the household				
Infant mode of	N/A	-0.0887	0.1205	-0.0553	0.0794				
delivery									
Survey year	-0.0887	N/A	-0.0533	0.0913	-0.0342				
Residential location	0.1205	-0.0533	N/A	-0.0867	0.1569				
Low family income	-0.0553	0.0913	-0.0867	N/A	-0.0083				
≥ 2 children < 5 y in the household	0.0794	-0.0342	0.1569	-0.0083	N/A				

eTable 3. Associations Between Infant Mode of Delivery and Pneumococcal Carriage Density in Fijian Infants Aged 5 to 8 Weeks, in 4 Annual Pneumococcal Carriage Surveys									
	Unadjusted Model		Adjusted Model ^a						
Characteristics	Differences in medians (95% CI)	P Value	Differences in medians (95% CI)	P Value					
Overall pneumococcal density (log ₁₀ GE/ml) ^b (n=516)									
Infant mode of delivery									
Cesarean	1 [Reference]		1 [Reference]						
Vaginal	0.41 (-0.05, 0.87)	.08	0.27 (-0.16, 0.70)	.22					
Survey year	4 [D (1		4 ID (1						
Before introduction of PCV10 After introduction of PCV10	1 [Reference]		1 [Reference]						
1 y	-0.04 (-0.38, 0.29)		0.07 (-0.28, 0.43)						
-	, ,	<.001	` ,	<.001					
2 y	-0.20 (-0.58, 0.18)	<.001	-0.12 (-0.41, 0.37)	<.001					
3 y Residential location	-0.91 (-1.21, -0.60)		-0.68 (-1.00, -0.36)						
Residential location	1 [Reference]		1 [Reference]						
Urban	0.34 (0.10, 0.58)	.006	0.13 (-0.11, 0.38)	.13					
Low family income ^c	-0.05 (-0.33, 0.23)	.74	-0.04 (-0.28, 0.21)	.77					
≥ 2 children < 5 y in the household	0.25 (-0.01, 0.52)	.07	0.24 (-0.01, 0.50)	.06					
PCV10 seroty	pe pneumococcal dens	ity (log₁₀ GE/	/ml) ^d (n=121)						
Infant mode of delivery									
Cesarean	1 [Reference]		1 [Reference]						
Vaginal	0.44 (-0.64, 1.53)	.42	0.10 (-0.92, 1.11)	.85					
Survey year									
Before introduction of PCV10	1 [Reference]		1 [Reference]						
After introduction of PCV10									
Pre-PCV10	1 [Reference]		1 [Reference]						
1 y 2 y	-0.28 (-0.98, 0.43) -0.70 (-1.55, 0.16)	.06	-0.24 (-0.91, 0.43) -0.56 (-1.37, 0.25)	.15					
3 y	-0.70 (-1.55, 0.10)	00	-0.75 (-1.43, 007)	. 10					
Residential location	0.00 (1.07, 1.27)		0.70 (1.40, 0. 07)						
Rural	1 [Reference]		1 [Reference]						
Urban	0.31 (-0.23, 0.85)	.26	0.21 (-0.32, 0.75)	.43					
Low family income ^c	0.11 (-0.46, 0.67)	.71	-0.08 (-0.60, 0.44)	.77					
> 2 children < 5 y in the household	-0.13 (-0.70, 0.45)	.66	0.15 (-0.38, 0.69)	.57					
Non-PCV10 sero	type pneumococcal de	nsity (log ₁₀ C	SE/ml) ^e (n=393)						
Infant mode of delivery									
Cesarean	1 [Reference]		1 [Reference]						
Vaginal	0.51 (-0.01, 1.03)	.06	0.12 (-0.34, 0.60)	.60					
Survey year	4 [D=f=====1		4 [Dafanas - 1						
Before introduction of PCV10	1 [Reference]		1 [Reference]						
After introduction of PCV10 1 y	0.09 (0.29, 0.50)		0.17 (-0.25, 0.58)						
2 y	-0.03 (-0.45, 0.39)	<.001	0.08 (-0.38, 0.53)	<.001					
3 y	-0.86 (-1.20, -0.53)	1	-0.77 (-1.14, -0.39)						
Residential location									
Rural	1 [Reference]		1 [Reference]						
Urban	0.32 (0.05, 0.61)	.02	0.19 (-0.8, 0.46)	.17					
Low family income ^c	-0.05 (-0.39, 0.28)	.76	-0.11 (-0.39, 0.17)	.45					
> 2 children < 5 y in the household	0.19 (-0.14, 0.53)	.26	0.23 (-0.05, 0.52)	.11					

Abbreviations: PCV10, ten-valent pneumococcal conjugate vaccine; CI, confidence interval

- ^a Variables adjusted for were survey year, residential location, and low family income.
- ^b Only includes participants who were pneumococcal carriers, including non-encapsulated pneumococci
- ^c Weekly family income below the basic needs poverty line (<FJ\$175 per week)².
- ^d Only includes participants who were carriers of serotypes included in PCV10 (serotypes 1, 4, 5, 6B, 7F, 9V, 14, 18C, 19F, and 23F).
- ^e Only includes participants who were carriers of serotypes not included in PCV10, including non-encapsulated pneumococci.

eReferences

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 Narsey W, Raikoti T, Waqavonovono E. Preliminary Report: Poverty and Household Incomes in Fiji in 2008-09 (based on the 2008-09 Household Income and Expenditure Survey): Fiji Islands Bureau of Statistics; 2010.