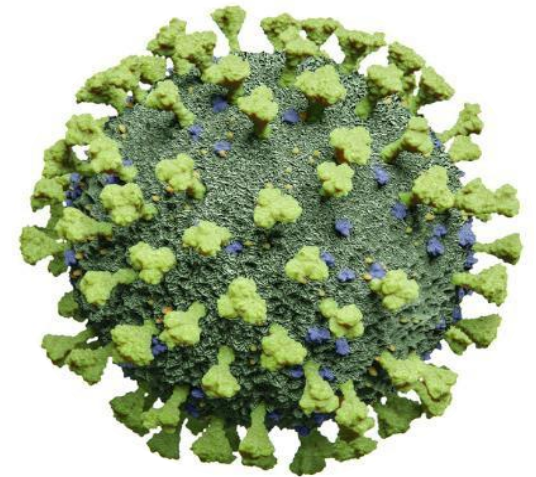


Metapneumovirus en todas las quinielas

Dr. Fernando González Romo

Servicio de Microbiología Clínica, Hospital Clínico San Carlos. Madrid

Profesor Asociado de Microbiología, Facultad de Medicina. UCM



Metapneumovirus humano

“Conoce a tu enemigo y saldrás victorioso en mil batallas”
(Sun Tzu, siglo V a. C.)

“Las comparaciones son odiosas”
(Leon Festinger, 1954)

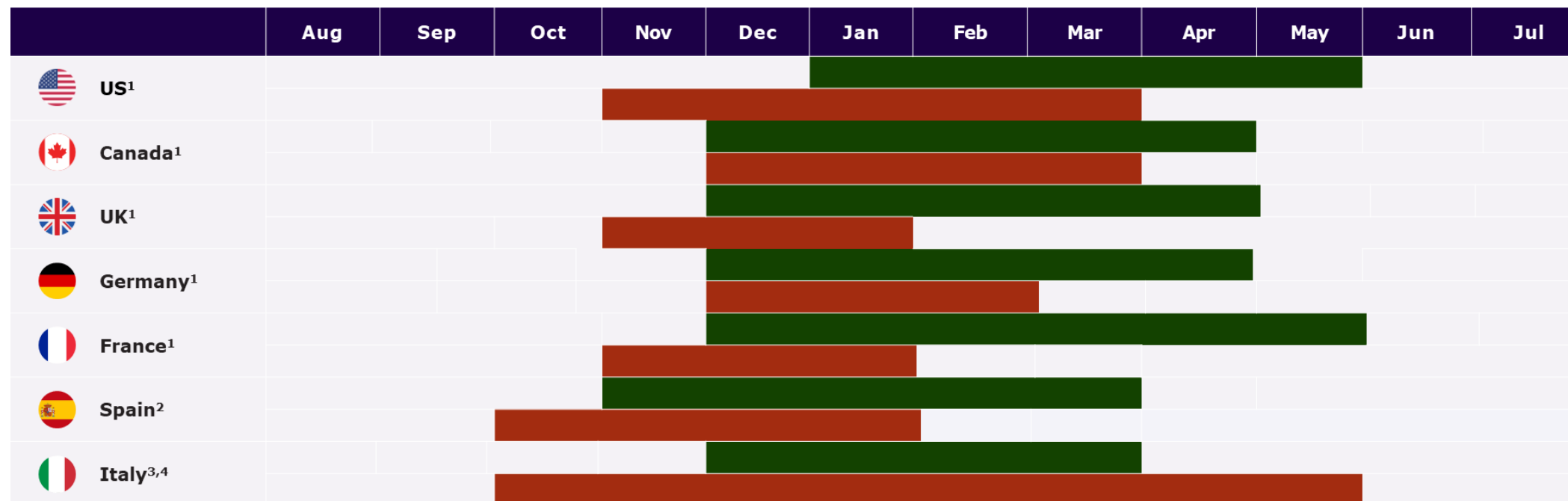
“Si algo funciona cópialo y, si puedes, mejóralo”
(Emilio Duró, 2013)



—■— RSV

—■— hMPV

Estacionalidad



Correlation between weekly positive rate of human metapneumovirus and meteorological variables.

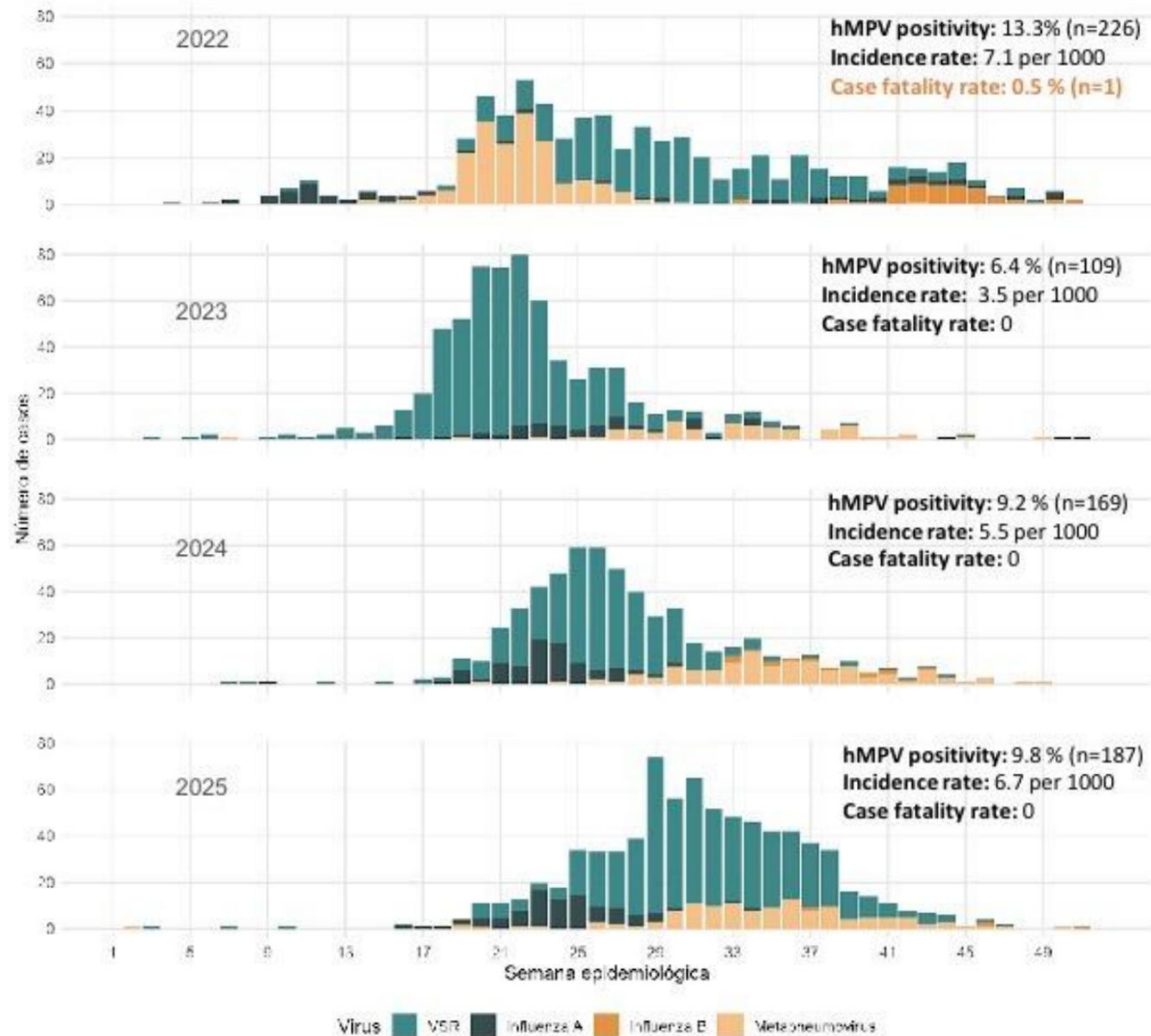
Meteorological variables	Univariate regression		Multivariate regression ^a	
	Coefficient	P	Coefficient	P
Temperature (°C)	-0.085	0.016	-0.013	0.813
Relative humidity (%)	0.047	0.054	-0.030	0.380
Rainfall (mm)	0.007	0.074	0.004	0.401
Sunshine duration (hour)	-0.067	<0.001	-0.070	<0.001
Atmosphere pressure (hPa)	0.008	0.268	0.006	0.376
Wind speed (m/s)	0.013	0.967	-0.261	0.449

Bold characters indicate statistically significant results.

1. Li Y, Nair H et al.. The Lancet Global Health 2019;7:e1031-e1045
2. Piñana M, et al. *J Clin Virol.* 2020;132:104590.
3. Obando-Pacheco P, et al. *J Infect Dis.* 2018;217(9):1356-1364.
- 4- Guido M, et al. *Virology.* 2011;417(1):64-70.
5. Ye H et al. *Int J Infect Dis* 2023; 137:40-47

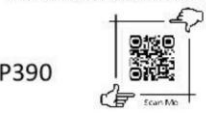


Estacionalidad



HUMAN METAPNEUMOVIRUS (hMPV): A PREVALENT CAUSE OF PEDIATRIC ACUTE LOWER RESPIRATORY INFECTIONS IN THE POST-PANDEMIC PERIOD. A MULTICENTER STUDY IN ARGENTINA.

ReSViNET
RESPIRATORY SYNCYTIAL VIRUS FOUNDATION



GENTILE, Angela¹; JUAREZ, María del Valle¹; GREGORIO, Gabriela²; GIOIOSA, Andres²; FERNANDEZ, Tatiana³; PEJITO, Natalia¹; ZUZEL, Paula¹; ENSINCK Gabriela³; LOBERTTI Silvina³; LOPEZ, Oscar⁴; LOPEZ, Leandro⁴; LUCION, María Florencia¹.

¹ "Dr Ricardo Gutiérrez" Children's Hospital, Buenos Aires City; ² "Dr. Victor J. Vilela" Children's Hospital, Rosario; ³ "Dr. Fernando Barreyro" Children's Hospital, Posadas; ⁴ "Prof. Alejandro Posadas" National Hospital, Buenos Aires Province.

Periodo 2022-2025
Menores de 18 años
Hospitalizaciones por ITRI (n=7021)

Variable	hMPV	ORV	p value
Age (months)	10 (5–19)	10 (4–24)	0.755
Age < 2 years	545 (79.1%)	3569 (74.9%)	0.018
Male sex	395 (57.2%)	2611 (54.8%)	0.252
Comorbidities	403 (58.3%)	2737 (57.4%)	0.681
Symptom duration at admission (days)	3 (2–5)	3 (2–5)	0.081
Bronchiolitis at admission	380 (55.0%)	2577 (54.1%)	0.683
Viral coinfection	188 (27.2%)	965 (20.2%)	0.001
Oxygen therapy	668 (96.7%)	4494 (94.3%)	0.009
Duration of oxygen therapy (days)	5 (4–8)	5 (3–8)	0.751
Total length of hospitalization (days)	7 (5–10)	7 (5–10)	0.047
ICU admission	115 (16.6%)	792 (16.6%)	1.000
Mechanical ventilation	102 (14.8%)	642 (13.5%)	0.373
Length of stay in ICU (days)	10 (6–16)	9 (6–15)	0.353

RSVWV'26 Rome, Italy, 16-20 February 2026



hMPV burden in infants attending hospital emergency departments in Iberia from 2021 to 2024: The RHEDI Prospective Study



Ana Dacosta-Urbieta ¹, Jacobo Pardo-Seco ¹, Narmeen Mallah ¹, Irene Rivero-Calle ¹, Ricardo Suárez Camacho ¹, Paula Vazquez López ³, Carlos Luaces Cubells ⁴, Pablo Ferrer González ⁵, Silvia Oliva Rodríguez Pastor ⁶, Pedro Flores ⁷, Teresa Bandeira ⁸, Fernanda Rodrigues ⁹, Mathieu Bangert ², Rolf Kramer ², Pierre Bourron ², Leticia Platero ², Catarina Gomes ², Gema Barbeito ², Alberto Gómez-Carballa ², Carmen Rodríguez-Tenreiro Sánchez ¹, Federico Martínón-Torres ¹, RHEDI CONSORTIUM

¹Grupo de Investigación en Genética, Vacunas, Infecciones y Pediatría (GENVIP), Hospital Clínico Universitario de Santiago de Compostela, Spain; ²Sanofi Vaccines; ³Hospital Gregorio Marañón, Madrid; ⁴Hospital Sant Joan de Déu, Cataluña; ⁵Hospital La Fe, Valencia; ⁶Hospital Regional Universitario de Málaga, Andalucía; ⁷CUF Descobertas Lisboa; ⁸Hospital de Santa Maria Lisboa; ⁹Hospital Pediátrico Coimbra

Prospectivo

Periodo 2021-2024

Menores de 5 años

Atención en Urgencias por ITR u otitis (n=1 654)

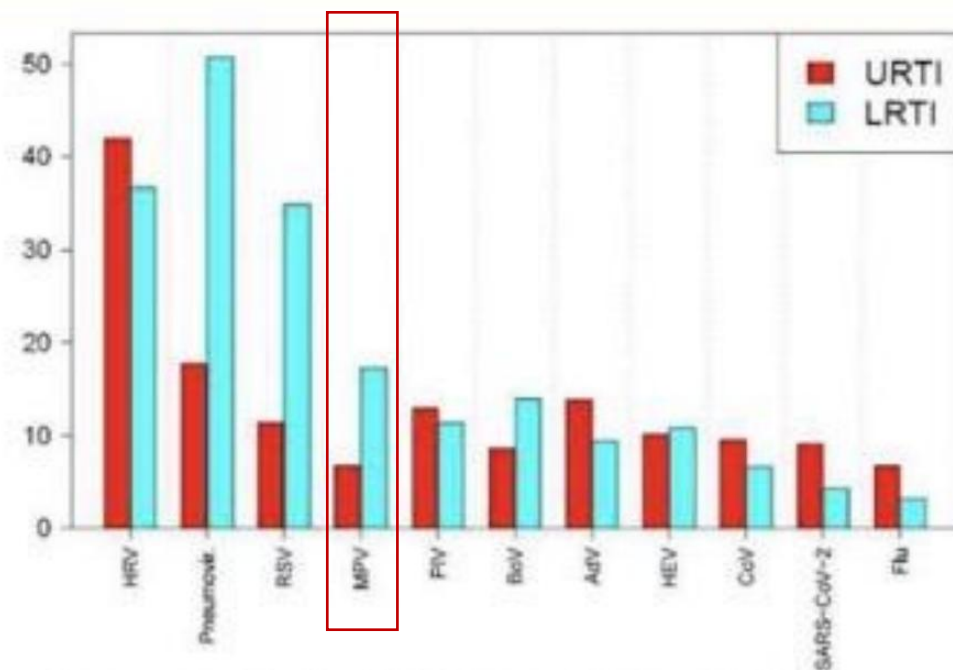
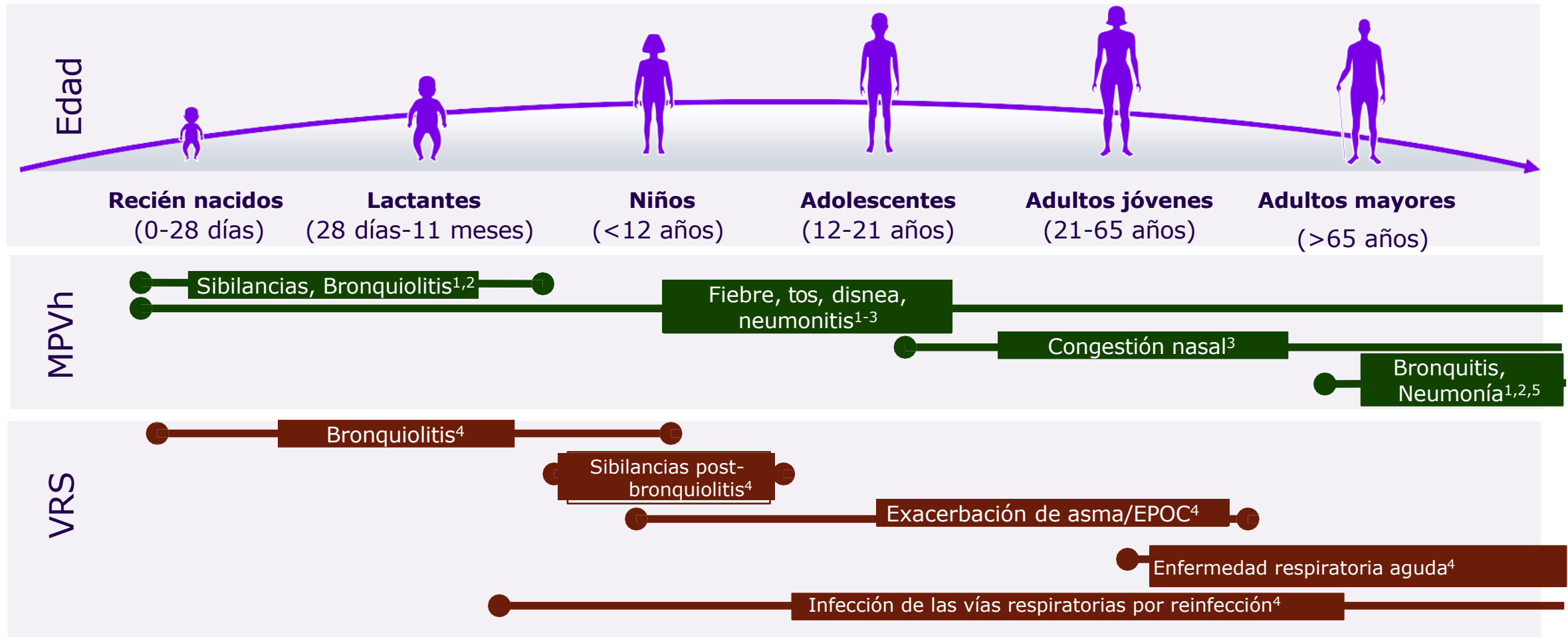


Figure 3: Positivity rate by type of infection (%)

Variable	hMPV	OVR	Valor de p
Bronquiolitis	46,3%	33,3%	<0,01
Sibilancias inducidas por virus	24,6%	14,5%	<0,01
Neumonía	10,3%	4,3%	<0,01
Tasa de hospitalización	34,5%	21,7%	<0,001
Estancia media (días)	10,3	4,7	<0,01
Necesidad de oxigenación	27,0%	13,1%	<0,001
Gravedad (escala ReSVinet)	5,5%	4,1%	<0,001



Espectro clínico



1. van den Hoogen BG, et al. *Pediatr Infect Dis J.* 2004;23(1 Suppl):S25-S32. 2. Hamelin M-E, et al. *Clin Infect Dis.* 2004;38(7):983-990. 3. Walsh EE, et al. *Arch Intern Med.* 2008;168(22):2489-2496. 4. Openshaw PJM, et al. *Annu Rev Immunol.* 2017;35:501-532. 5. Brochado O et al. *Infect Dis Ther* 2026; online 07 March



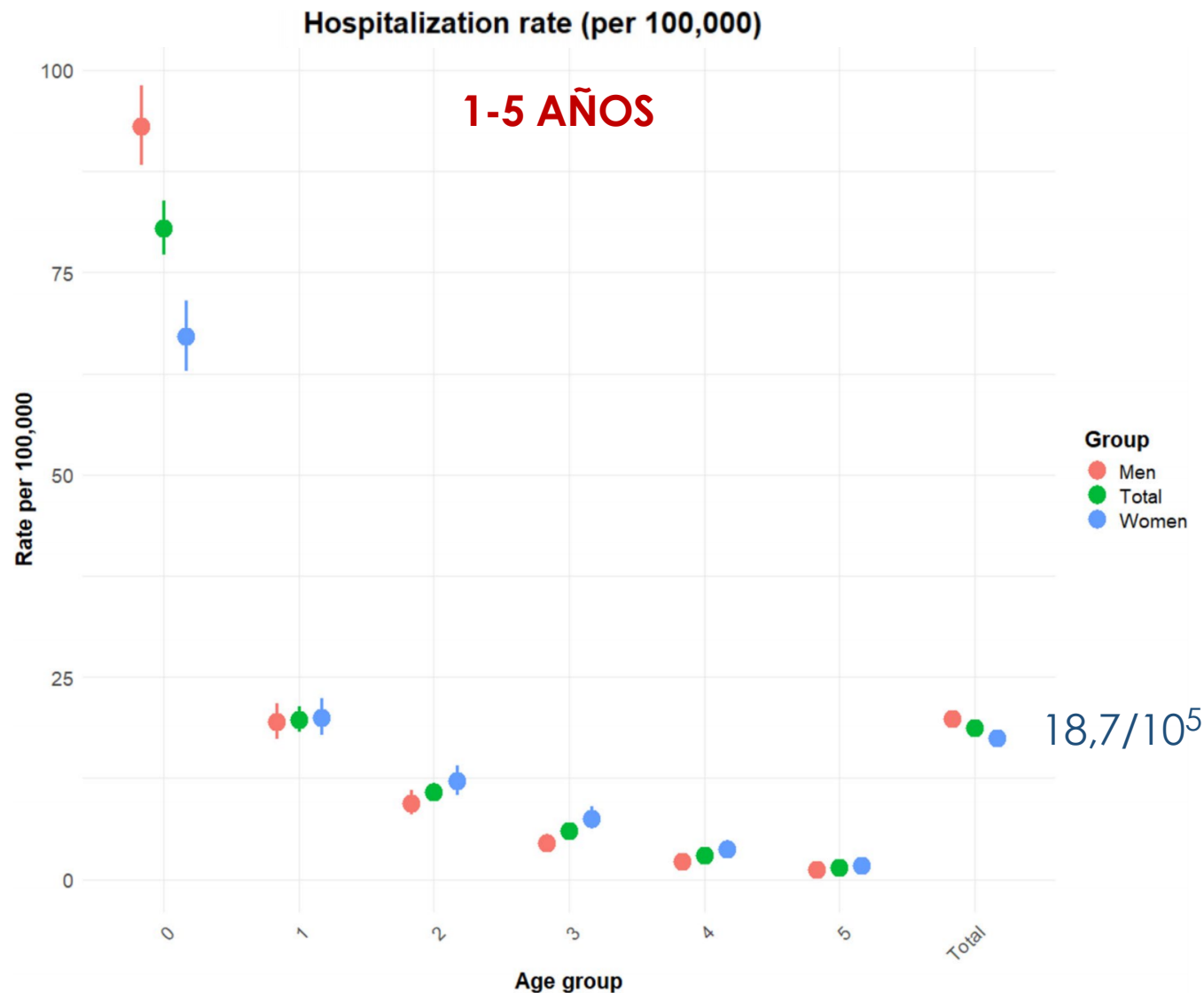


ORIGINAL RESEARCH

Epidemiological Burden and Hospitalization Trends of Human Metapneumovirus in Spain (2016–2023): A Retrospective Study

Oscar Brochado-Kith · Ruth Gil-Prieto · Angel Gil-de-Miguel

CMBD 2016-2023: 4824 hospitalizaciones
75% niños <6 años (18,7/10⁵) Bronquiolitis
16% adultos >59 años (0,79/10⁵) Neumonía



Brochado O et al. Infect Dis Ther 2026; online 07 March





ORIGINAL RESEARCH

Epidemiological Burden and Hospitalization Trends of Human Metapneumovirus in Spain (2016–2023): A Retrospective Study

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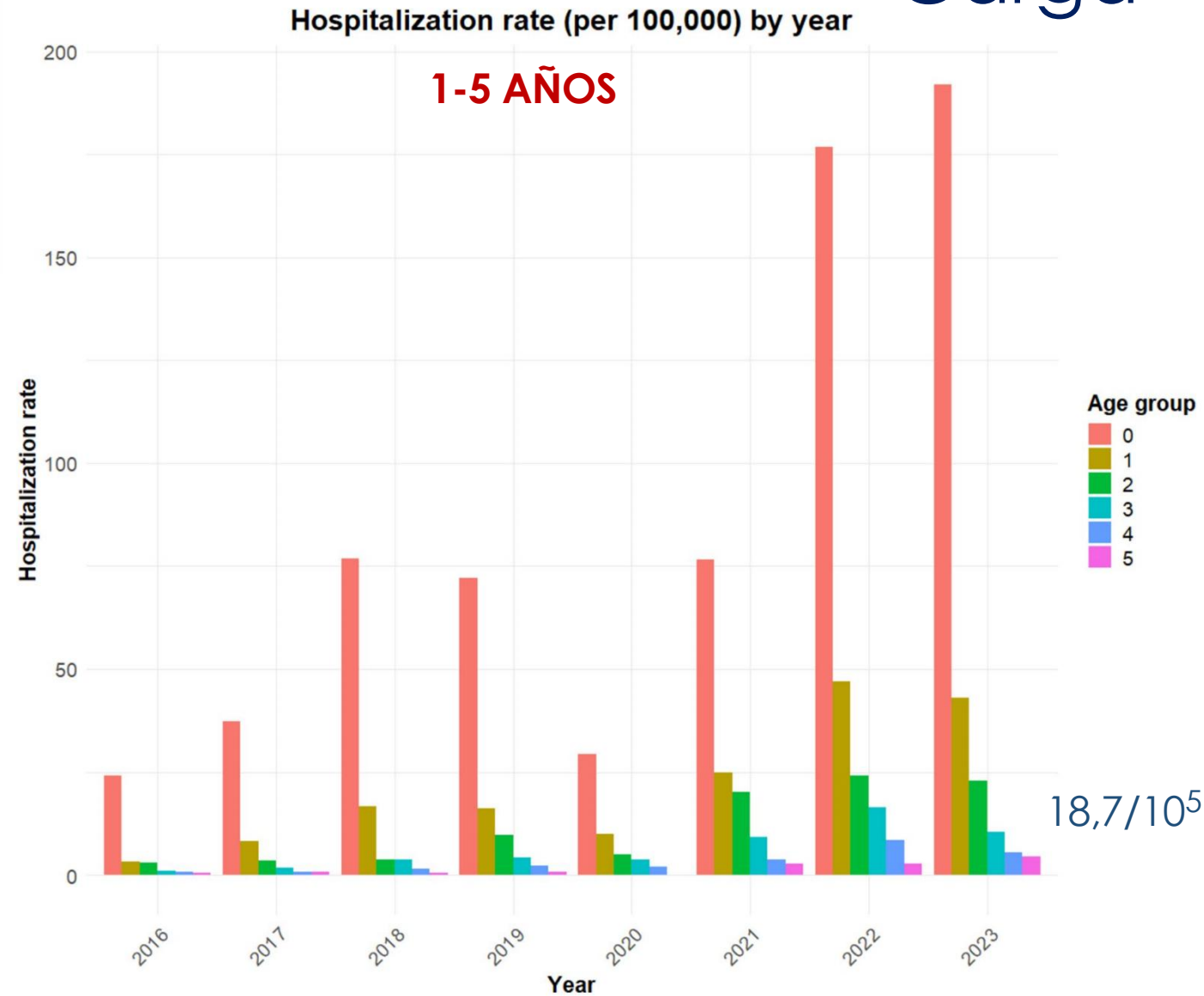
CMBD 2016-2023: 4824 hospitalizaciones
75% niños <6 años (18,7/10⁵) Bronquiolitis
16% adultos >59 años (0,79/10⁵) Neumonía

NIÑOS 1-5 AÑOS:

Estancia media: 5 días

Coste hospitalización: 3637,48€

Coste Total: 13 millones de euros



Brochado O et al. Infect Dis Ther 2026; online 07 March





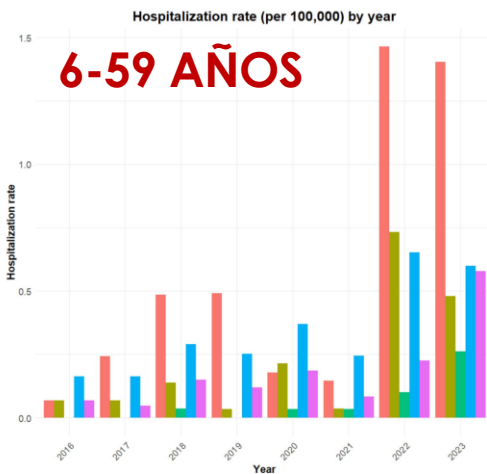
ORIGINAL RESEARCH

Epidemiological Burden and Hospitalization Trends of Human Metapneumovirus in Spain (2016–2023): A Retrospective Study

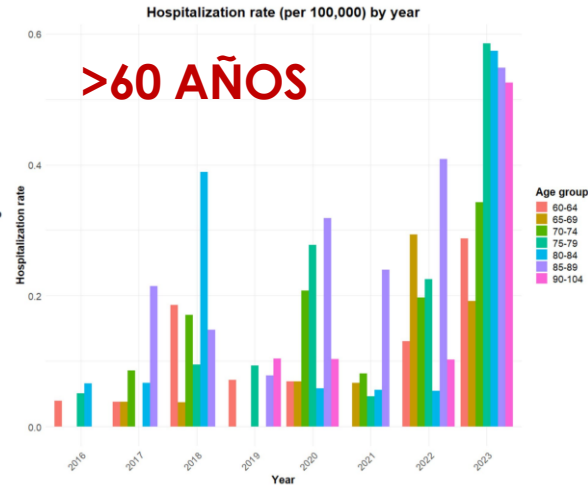
Oscar Brochado-Kith · Ruth Gil-Prieto · Angel Gil-de-Miguel

CMBD 2016-2023: 4824 hospitalizaciones
75% niños <6 años (18,7/10⁵) Bronquiolitis
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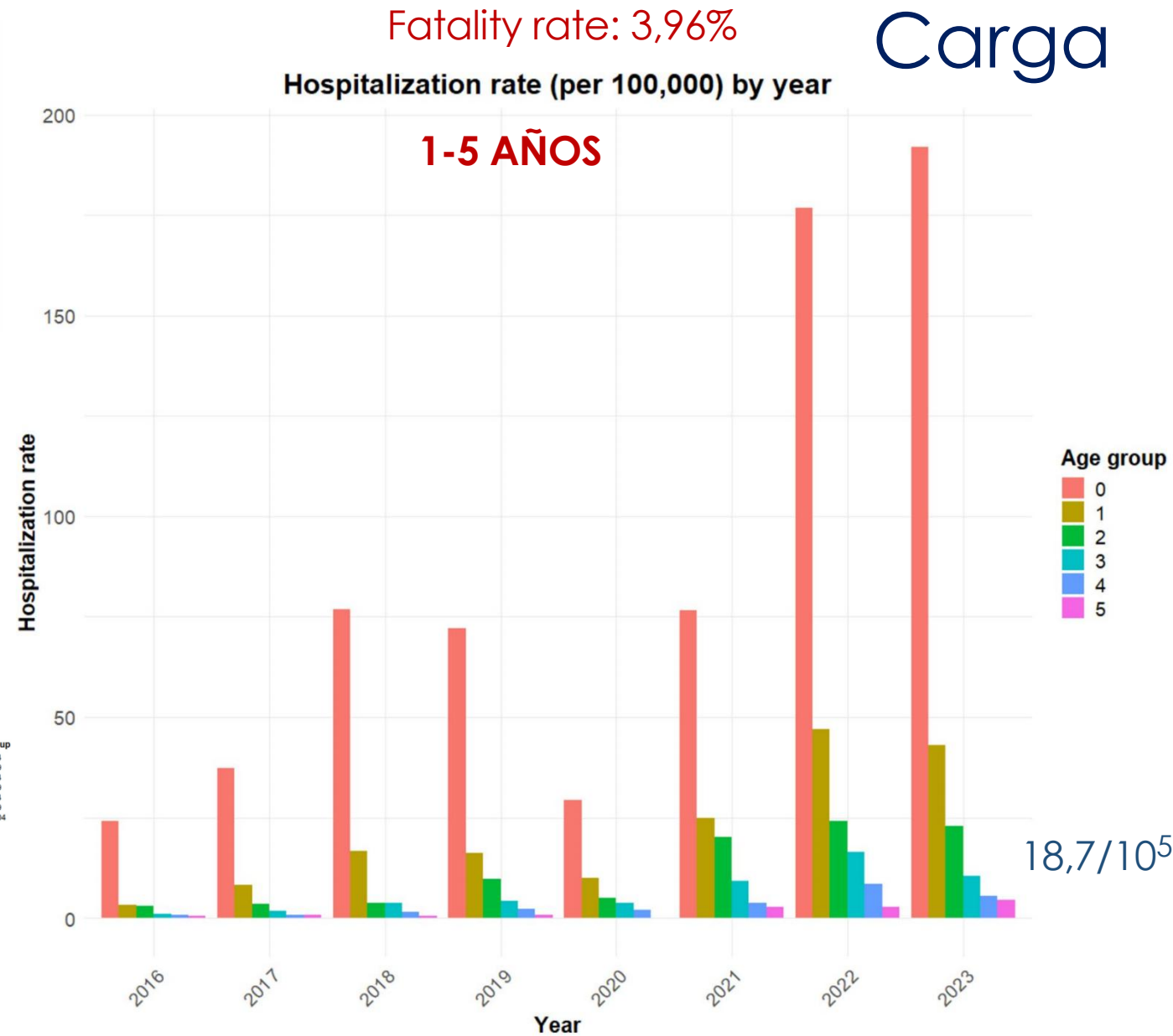
Fatality rate: 9,65%



0,17/10⁵



0,79/10⁵



Brochado O et al. Infect Dis Ther 2026; online 07 March



Generación Silenciosa
1920 - 1940



Baby Boomer
1946 - 1964



Generación X
1965 - 1979



Generación Y
1980 - 2000



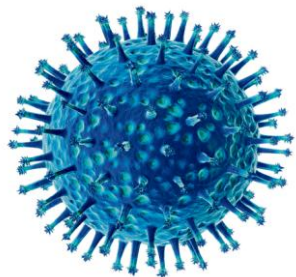
Generación Z
2001 - 2010



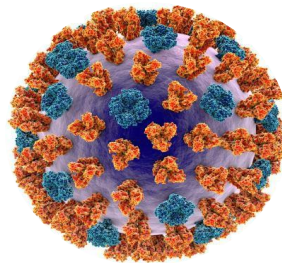
Generación Alfa
2010 - 2025



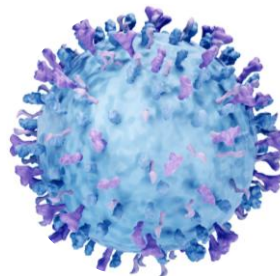
Gripe
1933



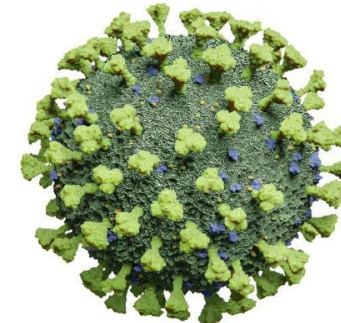
VRS
1957



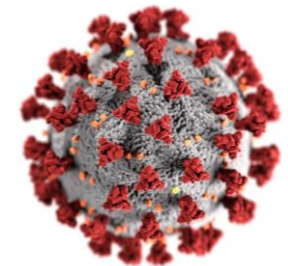
HCoV-229E
1966

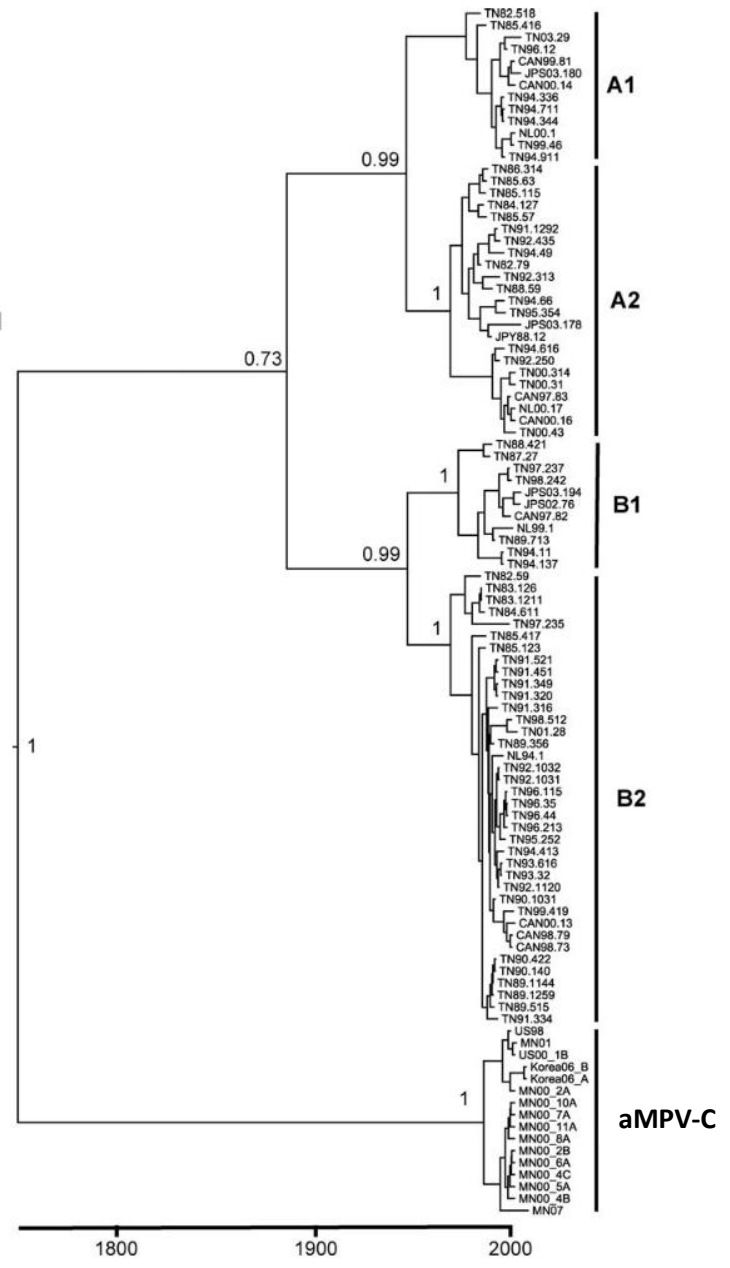
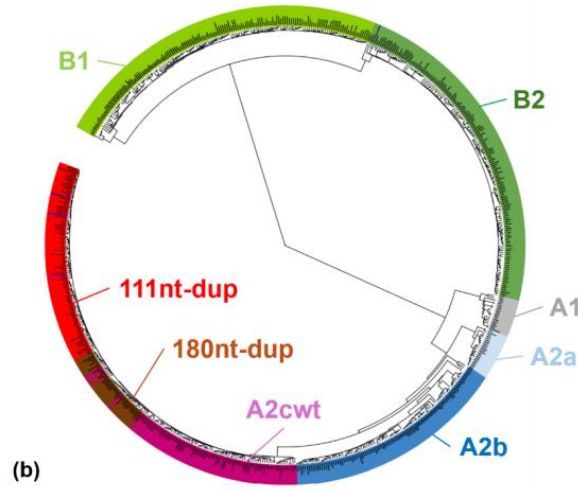
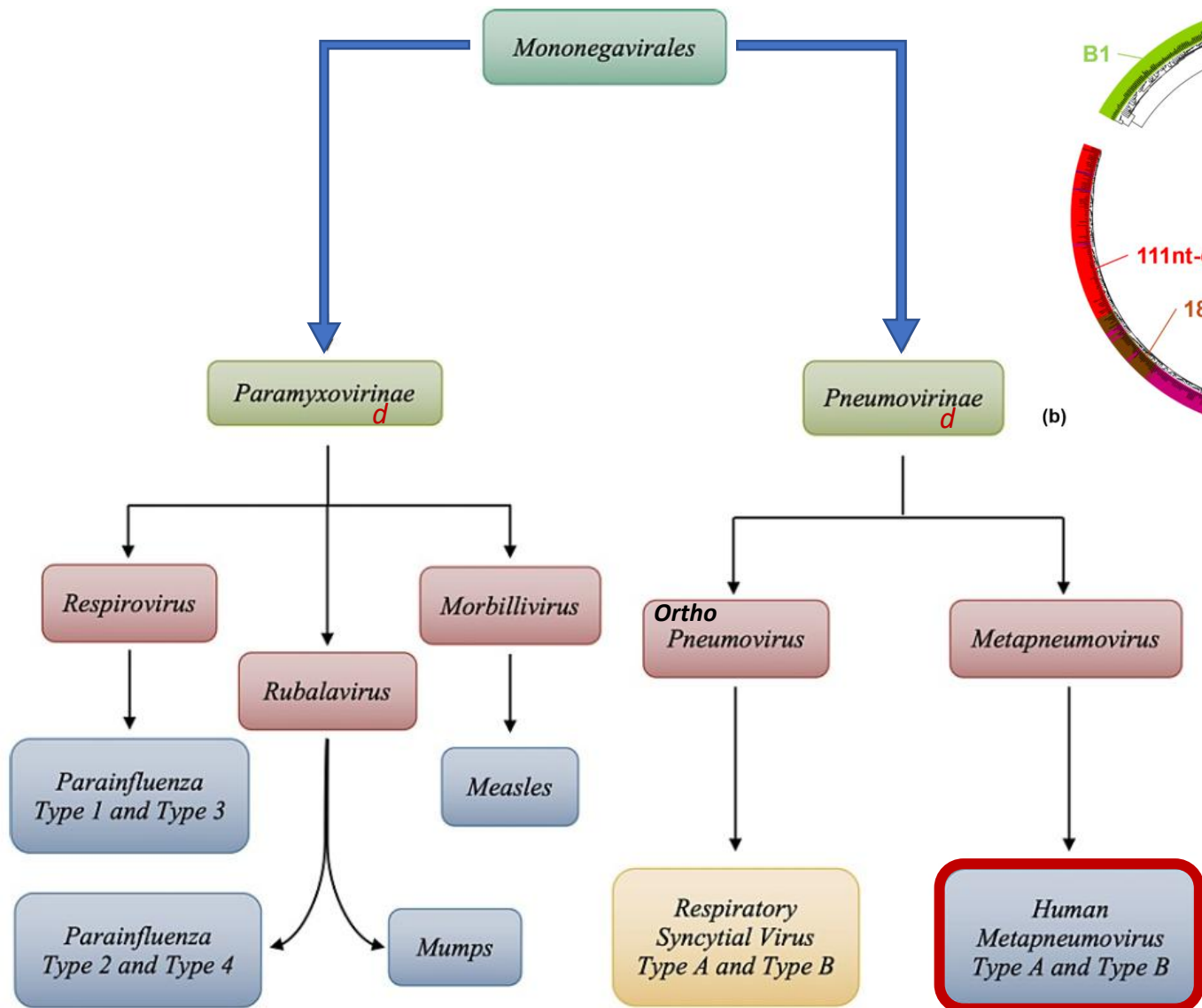


hMPV
2001



SARS-CoV2
2019

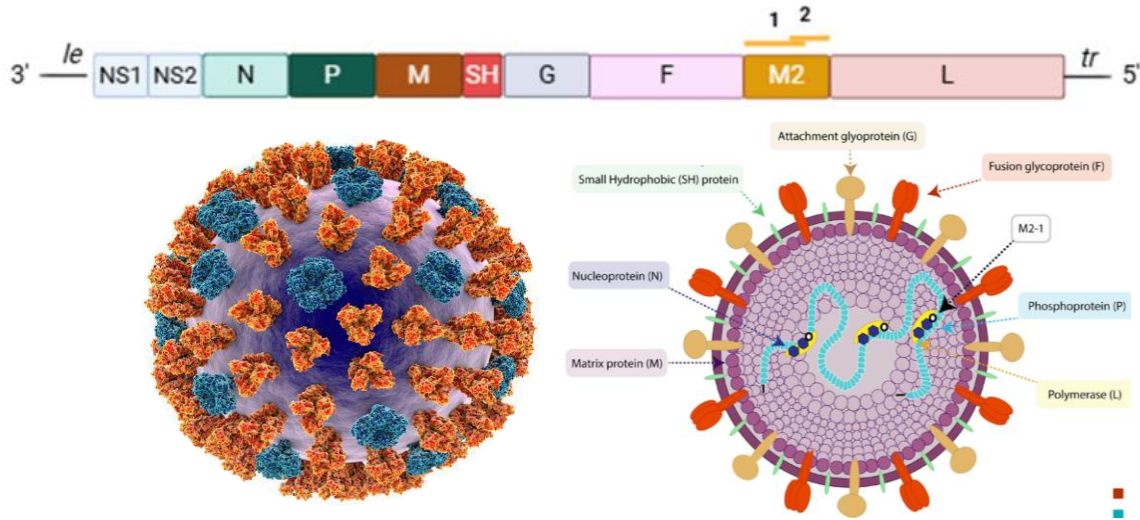




Chittipol N, Kandi V, Pinnelli V, et al. (June 02, 2025) The Re-emergence of Human Metapneumovirus: Virus Classification, Characteristics, Mechanisms of Infection, Clinical Features, Diagnosis, Epidemiology, Prevention, and Treatment. Cureus 17(6): e85259



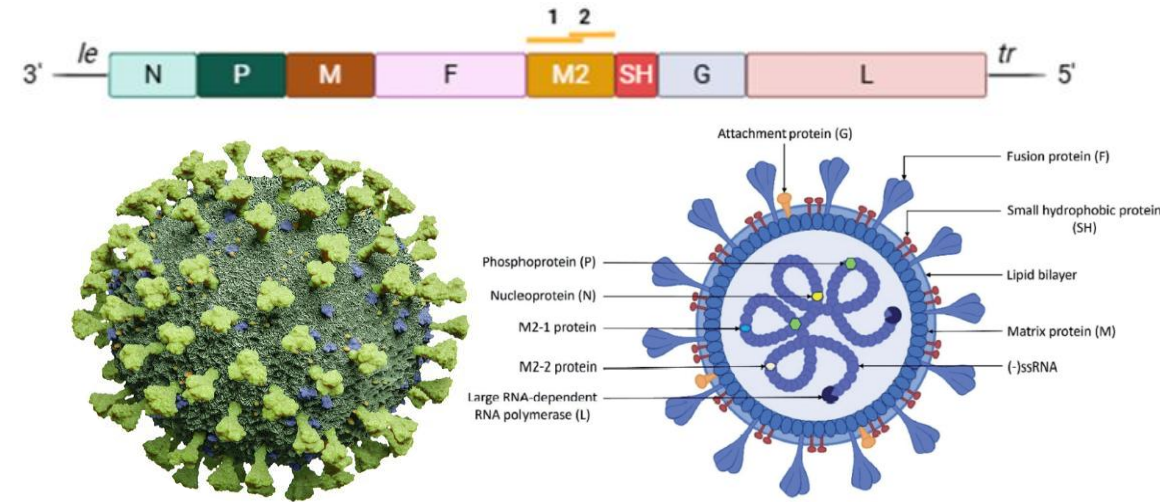
VRS



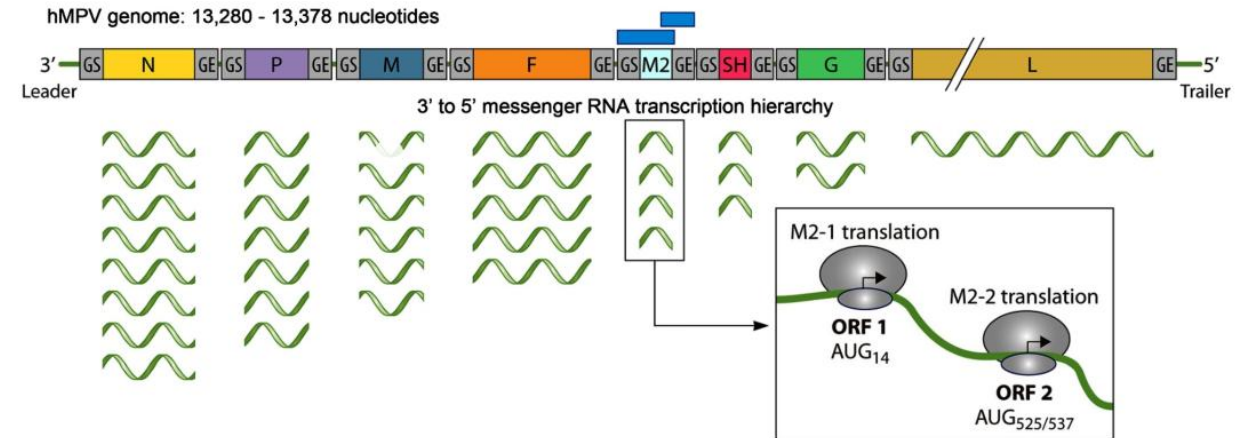
ARN monocatenario de sentido negativo de 15,2kb
Codifica 11 proteínas (Proteínas NS1 y NS2)

...y el orden de los genes es diferente al del VRS

hMPV

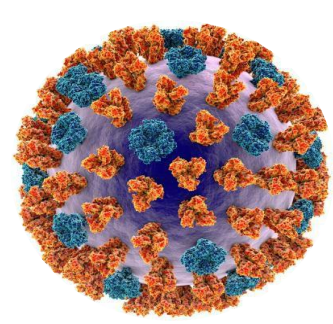


ARN monocatenario de sentido negativo de 13 kb
Codifica 9 proteínas



Twumasi S, et al. Reviews in Medical Virology 2025;35:e70043
Ríos Guzmán E, et al. Ther Adv Infect Dis 2022;9:1-21
van den Hoogen BG, et al. Virology 2002;295(1):119-132.
Kolli D, et al. Pathogen 2013;1;2(2):232-263

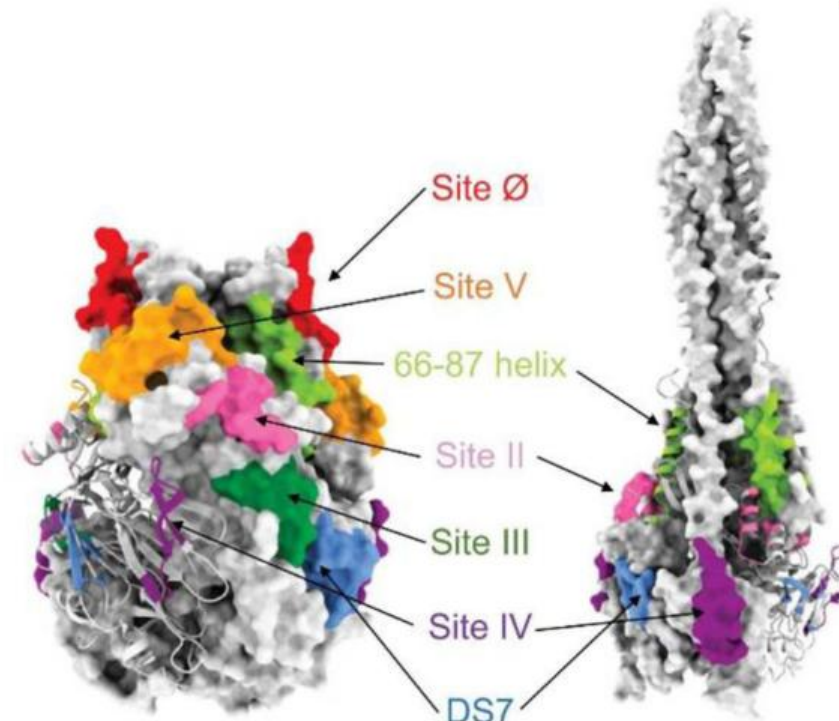
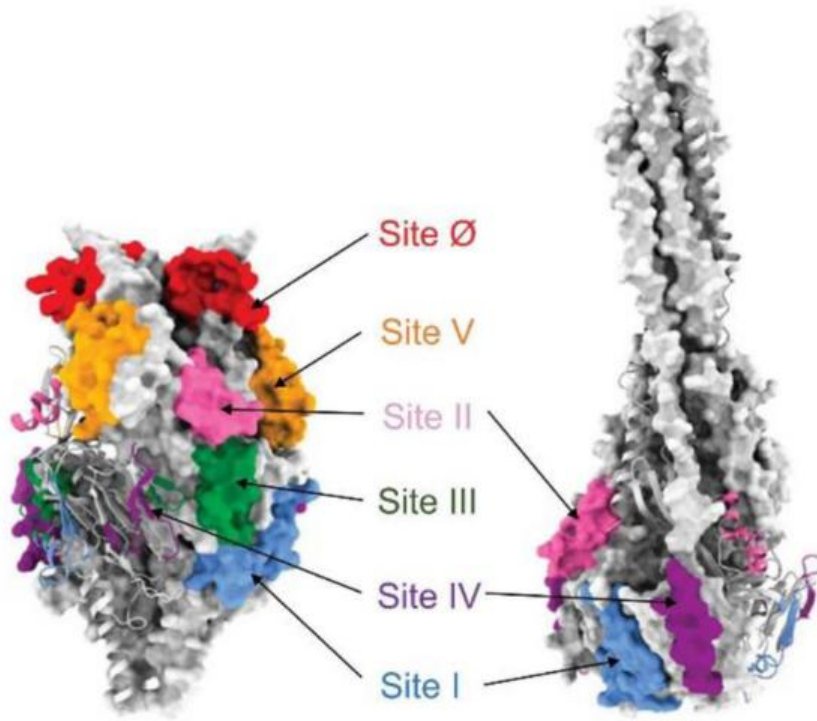
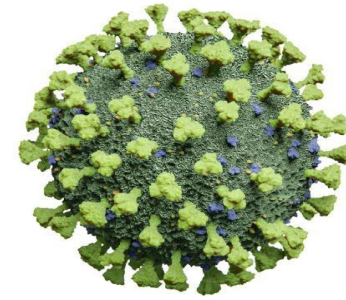




VRS

Glicoproteína F

hMPV



Principal determinante antigénico
Comparten 30% de identidad en la secuencia de aa
Forma pre y postfusión
Site Ø y V exclusivos de la forma prefusión

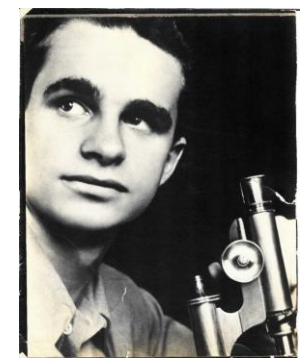
Muy conservada entre clados A y B
Doble función: Adhesión y Fusión
Genera nAb frente a formas pre y postfusión
Epítipo no presente en VRS: 66-87 hélix
Dianas más potentes para mAbs: Site IV, III y II

Miller RJ et al. Curr Opin Virol 2023;61:101337



Fracaso y abandono del ensayo clínico de la Vacuna VRS inactivada con formalina.

VRS 1968



Dr. R. Chanock

1956:
Chimpanzee
coryza agent
(CCA)

been reported that naturally acquired serum antibody at levels of 1:32 to 1:64 do not provide effective protection against infection(1).

We have noted that the respiratory tract illnesses associated with RS virus infection in RS vaccinees were more severe than those associated with RS virus infection in the group of infants who received the parainfluenza Type 1 virus

TABLE 6. EVIDENCE FOR FAILURE OF VACCINE-INDUCED VIRUS INFECTION AND B

SESSION I • Acute Respiratory Diseases

whom vaccine. Four of five RS vaccinees who were d did infected with RS virus required hospitalization, whereas none of five parainfluenza Type 1 vaccinees infected with RS virus was admitted to the hospital. These observations raise the question of a paradoxical vaccine effect. Possibly RS vaccines of low potency in some way "sensitize" an individual or potentiate the host's response to natural RS infection. Such a phenomenon has been noted previously with rickettsial vaccines (8), trachoma vaccine(9), and *M. pneumoniae* vaccine(10).

Thus our findings indicate that it is possible

hMPV 2007



Available online at www.sciencedirect.com

ScienceDirect

Vaccine 25 (2007) 5034–5040

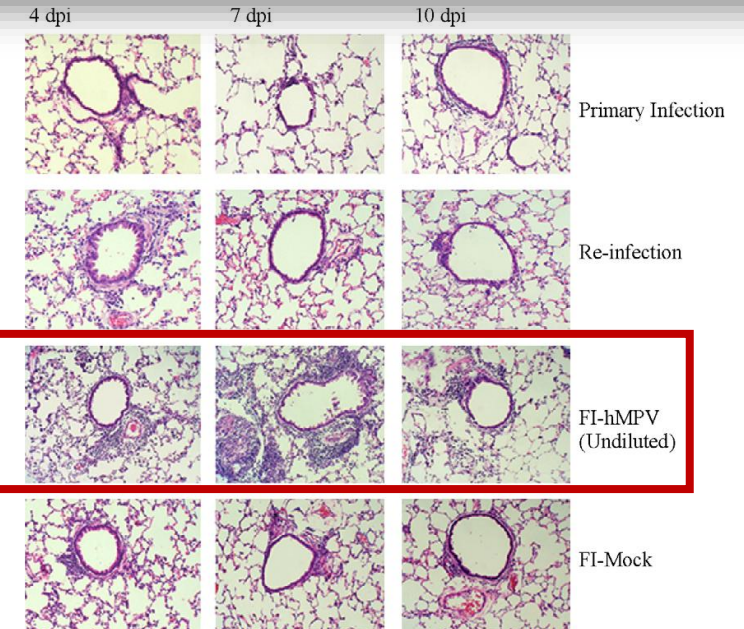
Vaccine

www.elsevier.com/locate/vaccine

Human metapneumovirus: Enhanced pulmonary disease in cotton rats immunized with formalin-inactivated virus vaccine and challenged

Kevin C. Yim^{a,*}, Rose P. Cragin^a, Marina S. Boukhvalova^a, Jorge C.G. Blanco^a, Marie-Ève Hamlin^b, Guy Boivin^b, David D. Porter^a, Gregory A. Prince^a

^a Virion Systems, Inc., 9610 Medical Center Drive, Suite 100, Rockville, MD 20850, United States
^b Research Center in Infectious Diseases of the Centre Hospitalier Universitaire de Quebec, and Laval University, Quebec City, Quebec, Canada

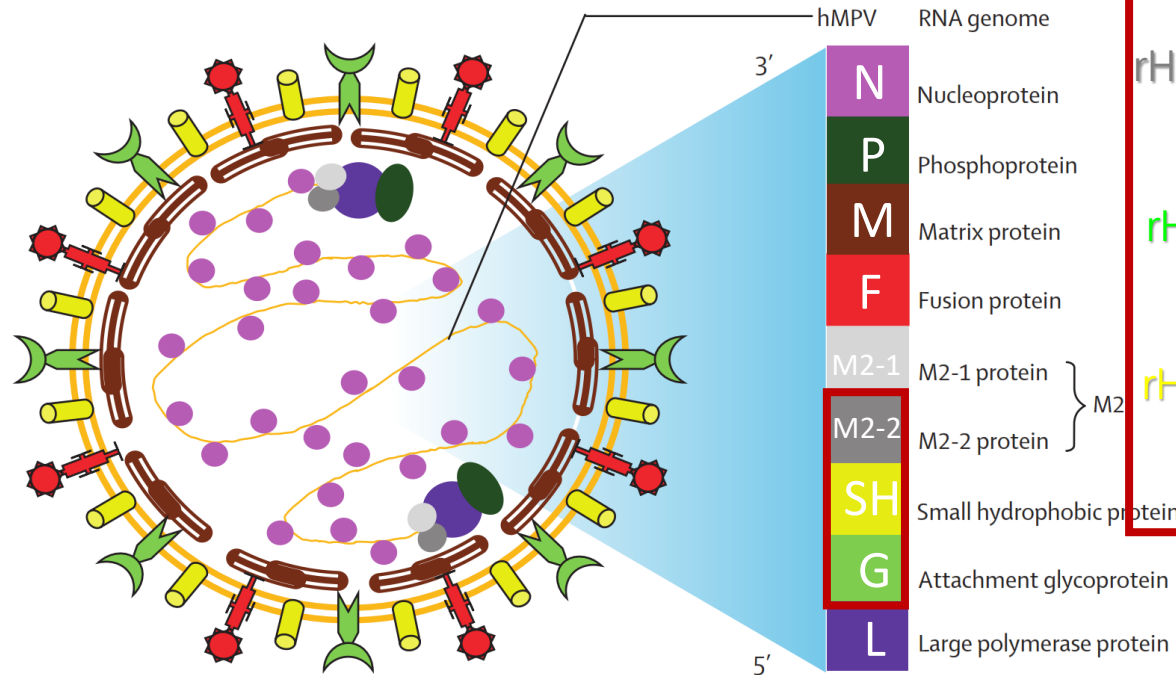


Yim KC, et al. Vaccine 2007;25: 5034–5040





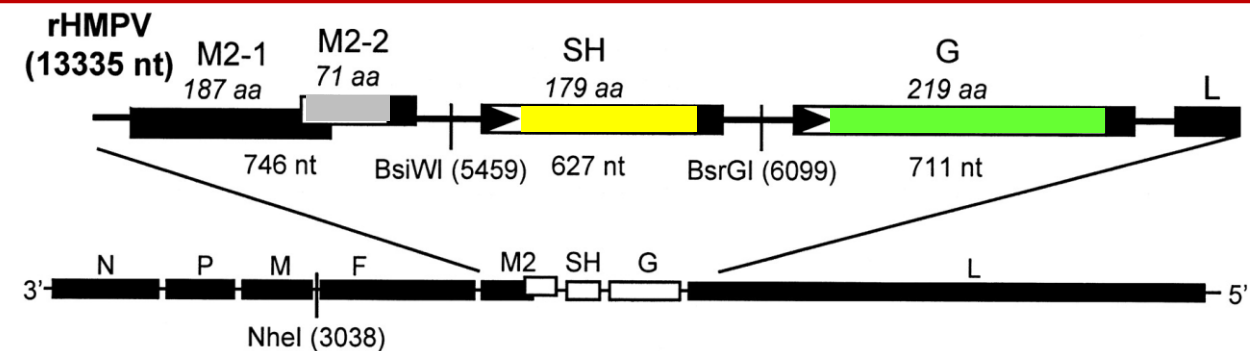
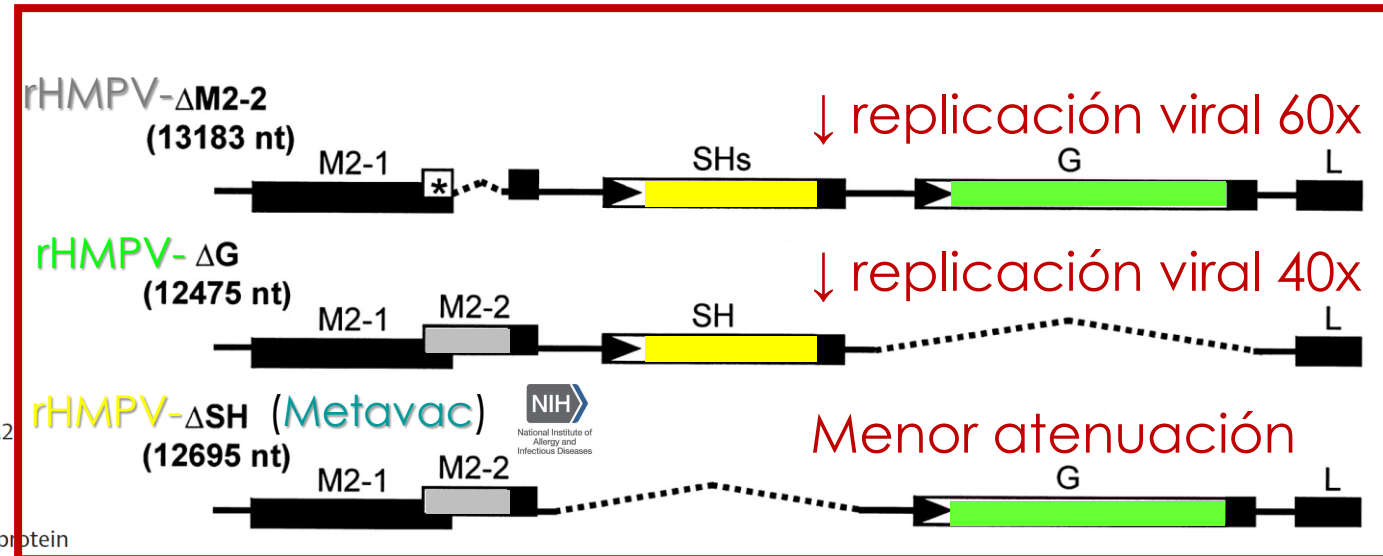
LAV: Delección molecular



hMPV

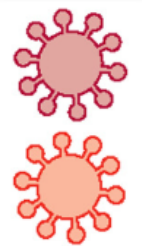
rHMPV- Δ G/SH

↓ replicación viral 600x



Biacchesi S, et al. J Virol 2005;79:12608–12613

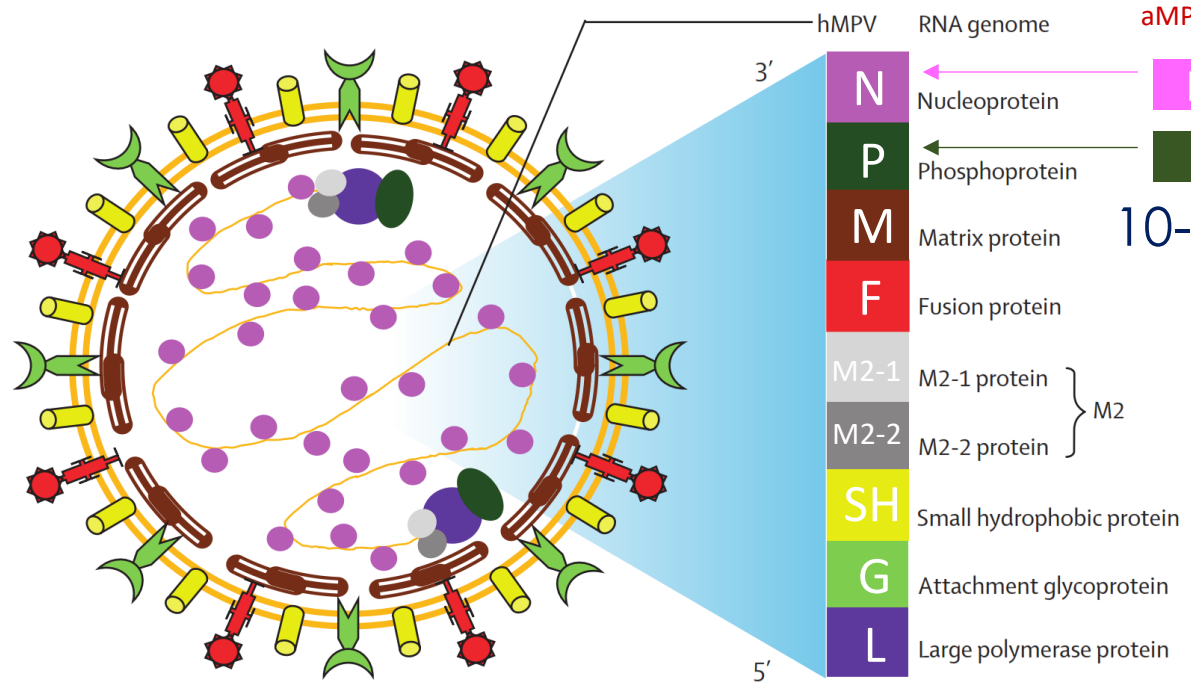




LAV: Delección molecular

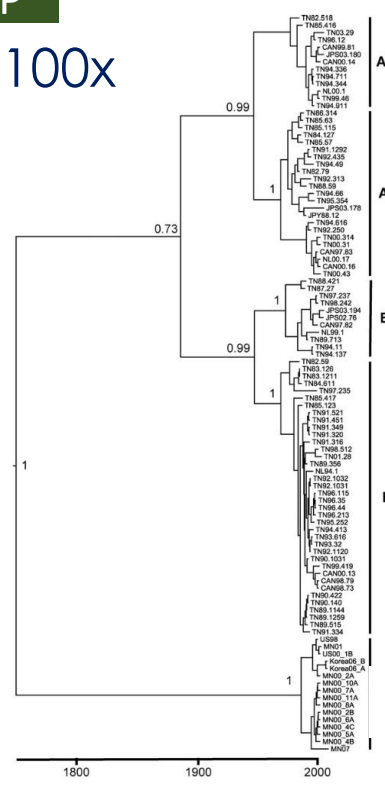


Recombinante



aMPV

10-100x



rHMPV-Pa

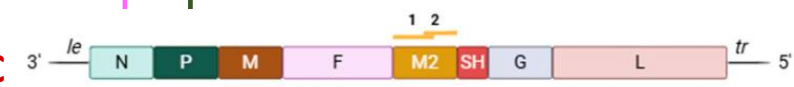


1 dosis
Intranasal (gotas)

Fase 1 (NCT01255410)

Adultos 18-49 años
Niños 12-59 m Seropositivos
Niños 6-59 m Seronegativos

Escasa inmunogenicidad en
niños 6-59 m Seronegativos
(Vacuna sobreatenuada)



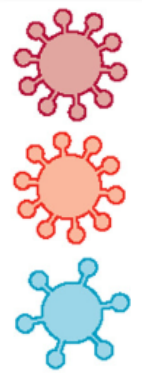
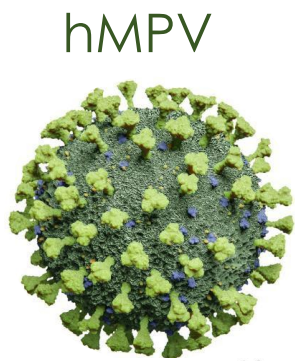
Karron R.A., et al. J Pediatr Infect Dis Soc 2018;7:86-89



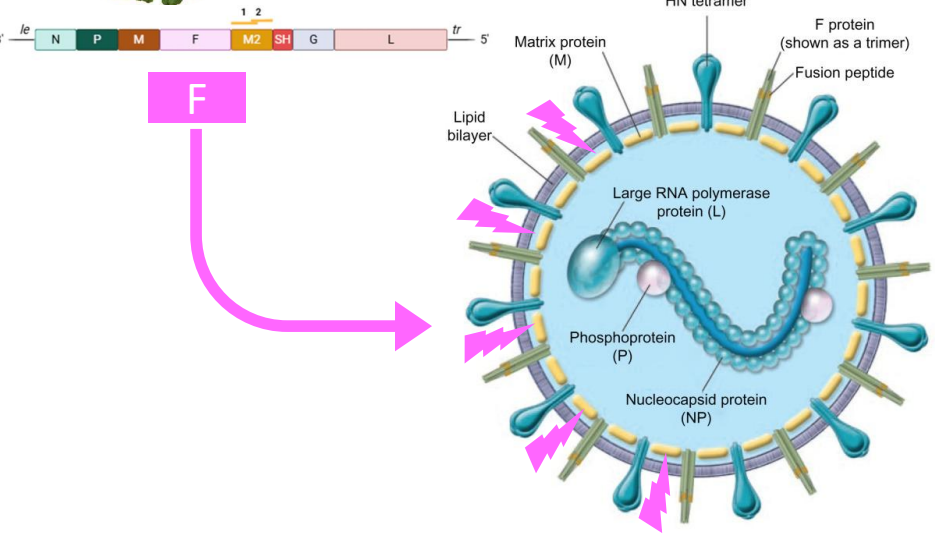
Asociación Española de Pediatría



Comité Asesor de Vacunas e Inmunizaciones

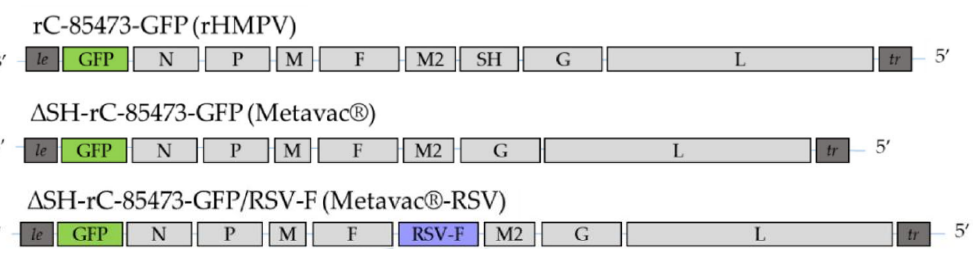


LAV: Delección molecular
Recombinante
Recombinante de vectores (quiméricas)



rB/HPIV3

PIV5
Sendei virus
Newcastle disease virus
Adenovirus



rHMPV-ΔSH (Metavac) →

<https://clinicaltrials.gov/study/NCT06546423>
Ogonczyk-Makowska D, et al. npj Vaccines 2024;9:111

B/HPIV3/HMPV-PreF-A
B/HPIV3/HMPV-PreF-B365



1 dosis
Intranasal (spray)

Fase 1 (NCT06546423)

Niños 24-59 m Seropositivos HPIV3

Activo (01/06/2025)

Bien tolerada
Buena inmunogenicidad frente a ambos virus

Metavac-RSV/hMPV



Intranasal. Preclínica (ratones)

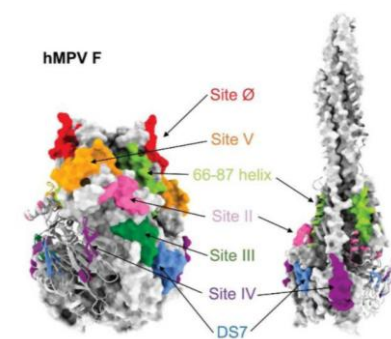
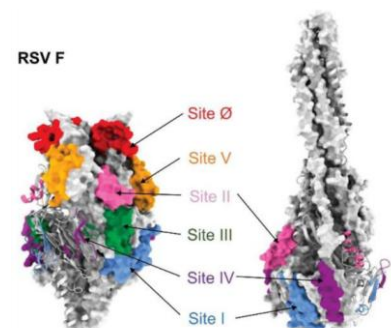
Fase 1 en Q3 2026





Fragmentos purificados solubles

Proteína Nativa



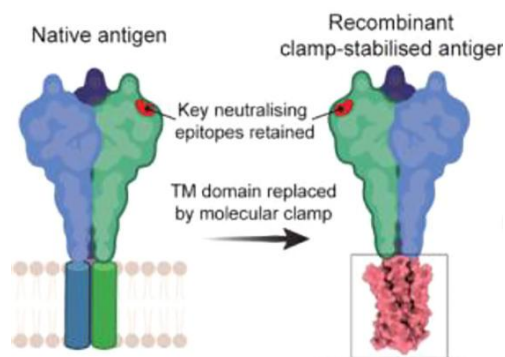
Estabilizar la
Proteína F



THE UNIVERSITY
OF QUEENSLAND
AUSTRALIA

Molecular Clamp:

Incorporación de un polipéptido
de trimerización muy estable.



Resp. inmune fuerte y específica
Fácil producción
Aplicable a numerosos virus
Purificación sin liofilización

VXB-241 (VRS/hMPV)

Dosis líquida
Refrigerada (2-8°C)
No adyuvada

Fase 1 (NCT06556147)

Adultos entre 18-40 y 60-83 años
1 dosis e impacto de booster 1 año

Activo (Fin 01/05/2027)

VXB-251 (VRS/hMPV/PIV3)

Fase 1 (NCT07295028)

Adultos entre 60-83 años; 1 dosis

Activo (Fin 24/04/2027)

sanofi

vicebio

RSVW'26:

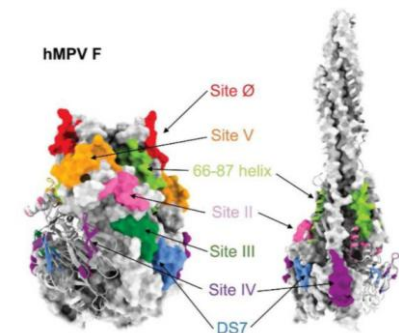
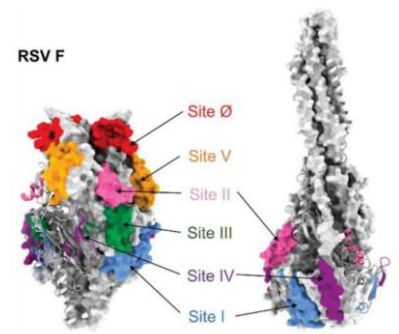
Segura
Inmunógena
Dosis 120+120





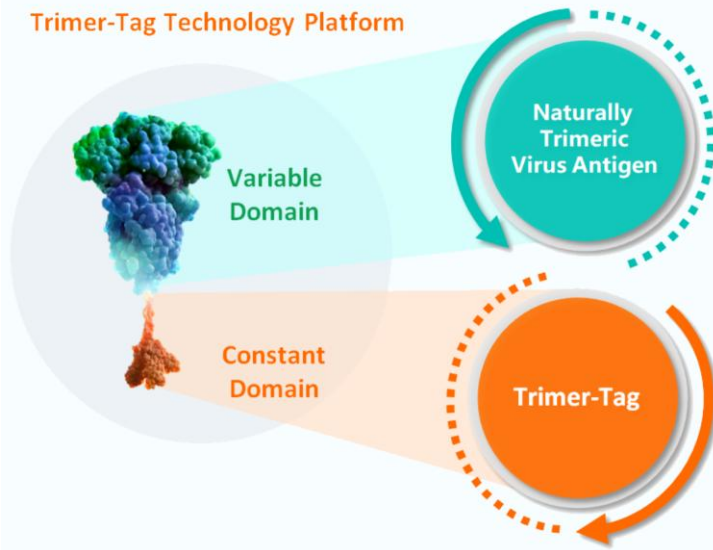
Fragmentos purificados solubles

Proteína Nativa



CLOVER
BIOPHARMACEUTICALS

Trimer-Tag Technology Platform



SCB-1022 (VRS/hMPV)

SCB-1033 (VRS/ hMPV/PIV3)

Fase 1 (NCT06984094)

Adultos entre 60-85 años

Activo (Fin 30/06/2026)

↑ nAbs x6-8 VRS
↑ nAbs x6-9 hMPV
↑ nAbs x4 PIV3

Fase 2

Programado en Q2 2026

Estabilizar la
Proteína F

<https://www.cloverbiopharma.com/science&technology/ourpipeline.html>

<https://clinicaltrials.gov/study/NCT06984094?term=NCT06984094&rank=1>

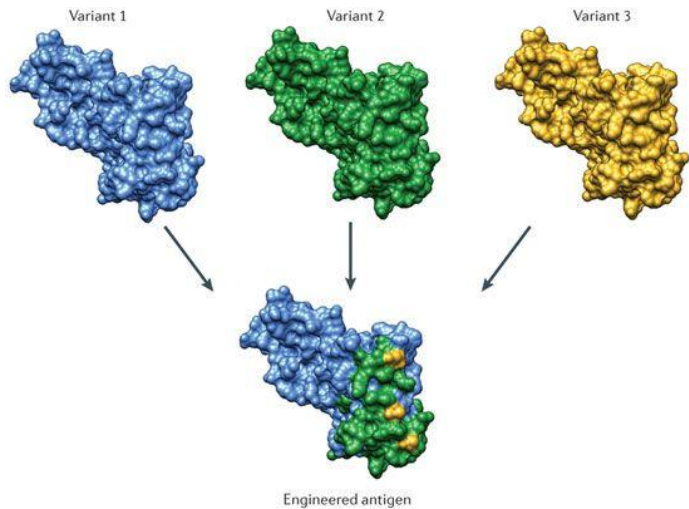




Fragmentos purificados solubles

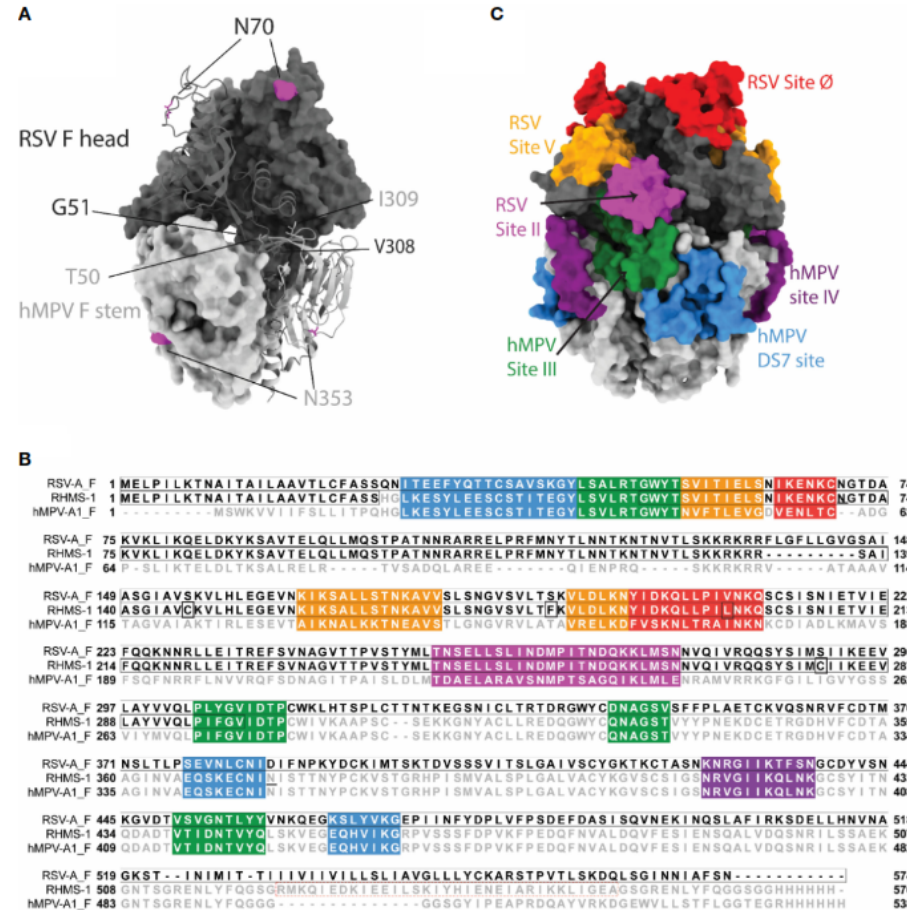
Proteína Nativa

Proteína Quimérica



Meningococo B

Proteína de unión al Factor H



VRS/hMPV

RHMS-1

Tallo hMPV y cabeza VRS

F-410, F-411

hMPV con varios epítopos VRS

RPM-1

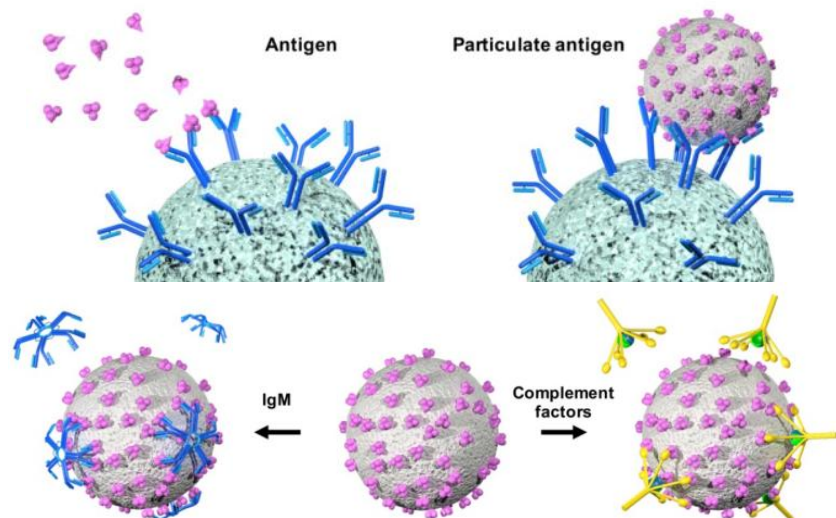
hMPV con un epítipo VRS





Fragmentos purificados solubles

VLP: Virus Like Particles



Bien tolerada

Fuerte respuesta a 28 días:

- VRS-A/VRS-B $\uparrow$ nAb x3-4
- hMPV-A/hMPV-B $\uparrow$ nAb x4-5

Niveles $\geq$ basales al año

IVX-A12

Fase 1 (NCT05664334)

Adultos entre 60-75 años, 1 dosis im

Completado (24/01/2024)

Fase 2 (NCT06481579)

Adultos $\geq$ 60 años, 1 dosis im

Completado (19/12/2024)

Fase 2a (NCT05903183)

Adultos entre 60-85 años, 1 dosis im

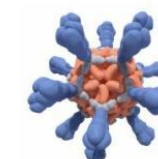
Completado (05/12/2025)

Fase 3

Preparado – Fast Track FDA

AstraZeneca

ICOSAVAX



IVX-121
RSV

+



IVX-241
hMPV

IVX-A12



RSV/hMPV
bivalent vaccine
candidate

+/- M59



mRNA-1365 (VRS/hMPV)

Fase 1 (NCT05743881) Niños entre 5-24 meses, 1 dosis im

Potenciales casos graves de RSV-LRTI en vacunados.
Menor inmunogenicidad tras Nirsevimab anterior.

PAUSADO
FDA (JUL 2024)

moderna

mRNA-1653 (hMPV/PIV3)

Fase 1 (NCT03392389) Adultos entre 18-49 años, 2 dosis im

Completado (19/07/2019)

Fase 1b (NCT04144348) Adultos y Niños de 12-59 meses

Completado (20/12/2022)

Bien tolerada e inmunógena (↑ nAb) de ambos virus.
No necesaria una segunda dosis.

<https://clinicaltrials.gov/study/NCT04144348>

Schnyder Ghamloush S et al. Pediatrics 2024;153:e2023064748

August A, et al. Open Forum Infect Dis 2022;9:ofac206

<https://clinicaltrials.gov/study/NCT06686654>

<https://clinicaltrials.gov/study/NCT06604767>

Ma S, et al. Hum Vaccines Immunother 2023;19:2293300

SP0256 (VRS/hMPV)

sanofi

Fase 1 (NCT06237296) Adultos entre 18-49 años y ≥ 60 años

Completado (11/02/2025)

Fase 1 (NCT06583031) Adultos entre 60-75 años

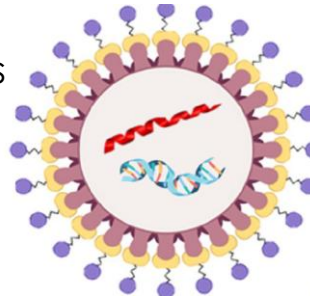
Completado (28/09/2025)

Fase 1/2a (NCT06134648) Adultos ≥ 60 años

Activo (Fin 10/03/2026)

Fase 1/2 (NCT06686654) Adultos ≥ 60 años

Activo (Fin 02/09/2027)



SP0291 (VRS/hMPV/PIV3)

Fase 1 (NCT06850051) Adultos entre 18-49 años (con TIV-HA)

Completado (31/12/2025)

Fase 1 (NCT06604767) Adultos ≥ 60 años

Activo (Fin 05/03/2026)



mRNA Multiepitopo

Incluir no solo genes de la proteína F sino también G, SH, M1, M2-1 o M2-2.
No generan nAb, pero favorecen citotoxicidad celular dependiente de Ac.

Metapneumovirus humano

CONCLUSIONES

Infrarreconocido, infraestimado e infradiagnosticado.

Potencial problema grave de salud pública.

Gran apoyo en plataformas vacunales conocidas.

La mayoría de candidatas son combinadas.

La mayoría en desarrollo orientadas a adultos.

Retos: Seguridad e inmunogenicidad.

