VACCINATION IN OLDER PERSONS

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High council of Public Health

- Ad hoc workgroup
 - Pneumococcal vaccination for adults
 - Experts:
 - ID, microbiology, GP, geriatric medicine, vaccinology
 - Scientific literature, (inter)national reports
 - Expert opinion
 - Cost-effectiveness not incorporated
- Guideline approval
 - Permanent workgroup (NITAG) vaccination
 - College of the HCPH
 - Publication





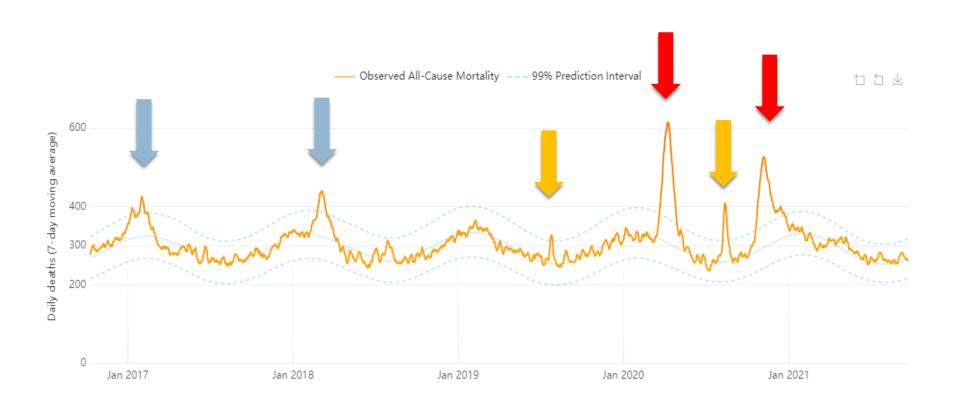
Overview

- Coronavirus in the older prerson
- Influenza in the older person
- Pneumococcal disease in the older person
- Frailty in older persons
 - Immunosenescence
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 - Age based vaccination
 - Risk based vaccination
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Excess mortality in Belgium

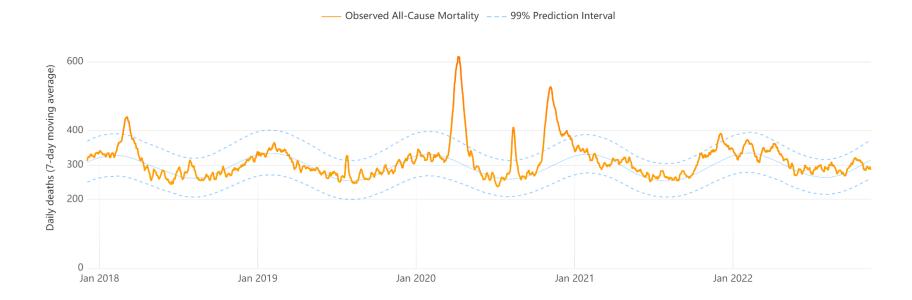


: RSV, Influenza, S. pneumoniae

: Heat

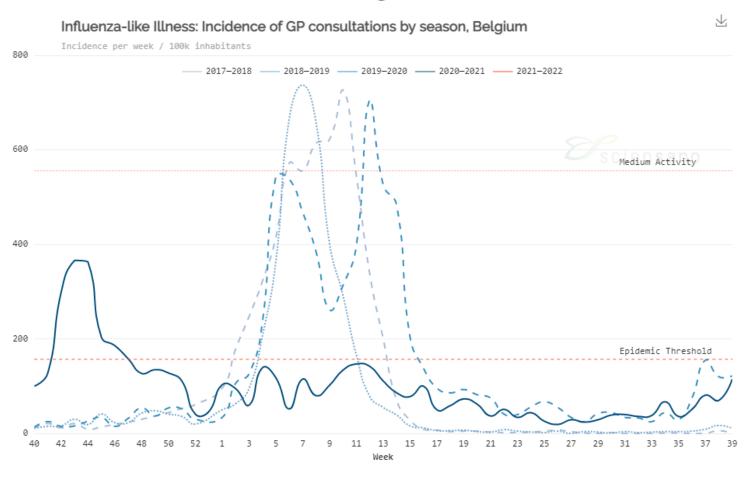
: SARS-Cov2

Excess mortality in Belgium



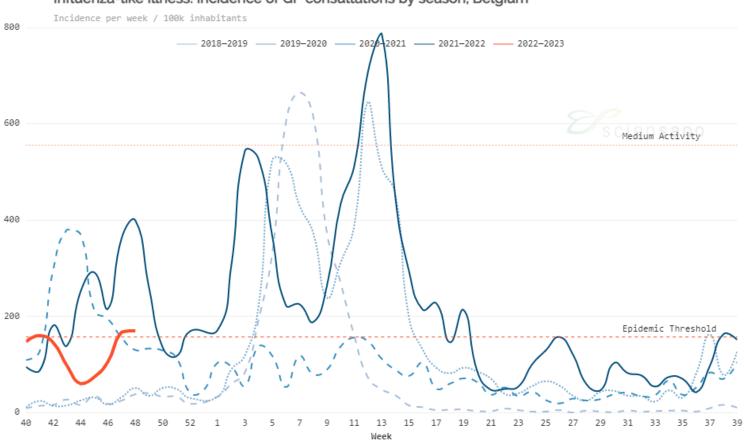
ILI in Belgium

Influenza-like Illness Monitoring



ILI in Belgium

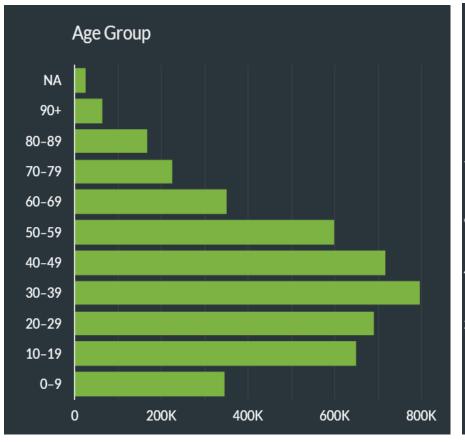
Influenza-like Illness: Incidence of GP consultations by season, Belgium

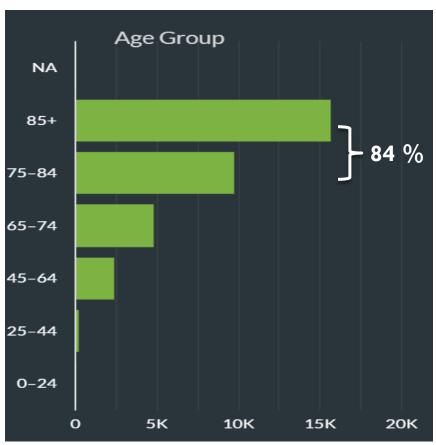


COVID-19 Belgium (7/12/2022)

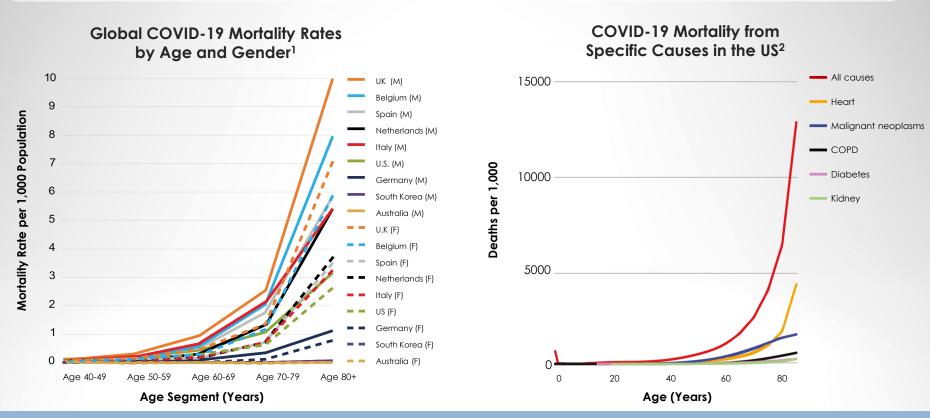
□ Cases: 4.644.478

□ Deaths: 33.086





Mortality Due to COVID-19 Increases With Age as Well as Comorbid Conditions



UK = United Kingdom; M = males; F = females; US = United States; COPD = chronic obstructive pulmonary disease.

1. RGA Web Site. COVID-19 Mortality Rates by Age and Gender: Why Is the Disease Killing More Men than Women? https://www.rgare.com/knowledge-center/media/research/covid-19-mortality-rates-by-age-and-gender-why-is-the-disease-killing-more-men-than-women. Accessed September 22, 2020. 2. Promislow DEL. J Gerontol A Biol Sci. 2020;75(9):e30-e33.

6 GOUDEN REGELS TEGEN CORONA 11 MILJOEN REDENEN OM VOL TE HOUDEN













11miljoenredenen.be

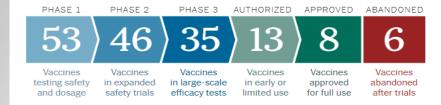


Een initiatief van de Belgische overheid



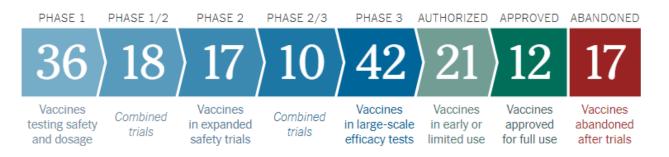
The New York Times

Coronavirus Vaccine Tracker



Coronavirus Vaccine Tracker

By Carl Zimmer, Jonathan Corum, Sui-Lee Wee and Matthew Kristoffersen Updated Aug. 31, 2022

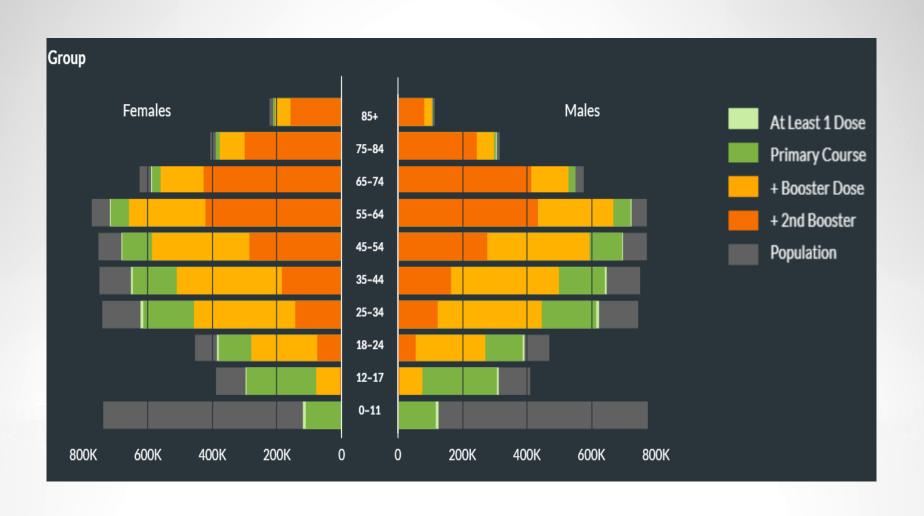


This tracker is no longer being updated. It followed the development of Covid vaccines from early 2020 through August 2022. More than 120 clinical trials were underway at that time.

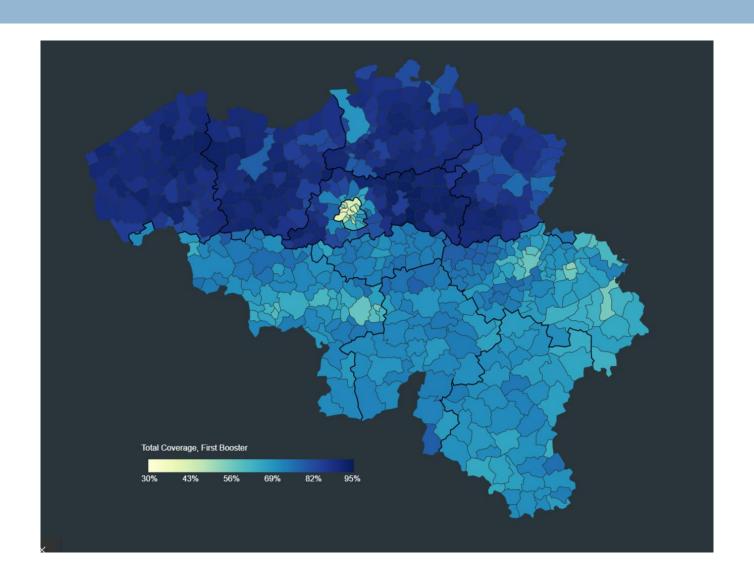
The Centers for Disease Control and Prevention maintains a list of <u>Covid</u> vaccines that are authorized or approved in the United States.



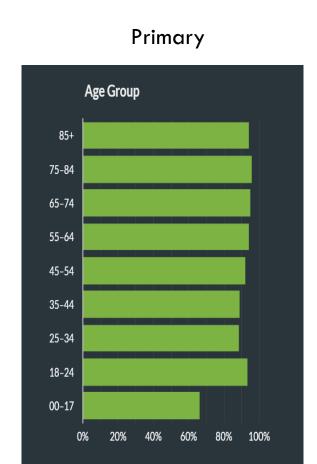
12/2022



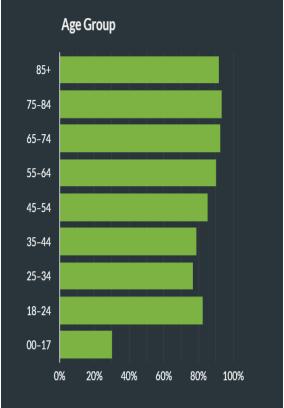
Coverage first booster Coronavaccine Belgium



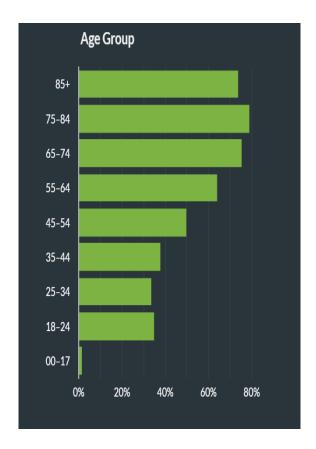
HCW vaccine coverage



Booster 1

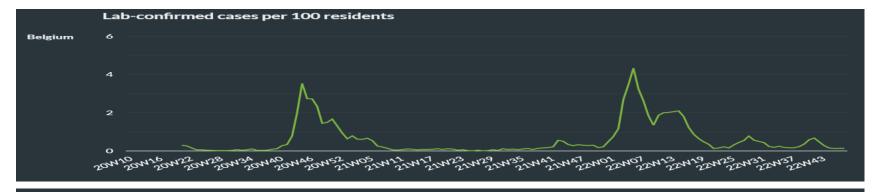


Booster 2

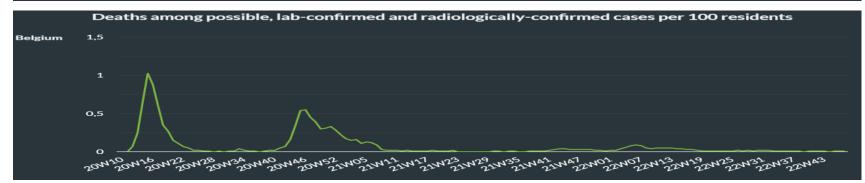


COVID-19 in nursing homes









Coronavirus vaccines effectiveness NH residents

TABLE 2. Estimated vaccine effectiveness* of a second COVID-19 vaccine booster dose relative to a first booster dose only, for four COVID-19–related outcomes in nursing home residents — 196 nursing homes, 19 states,† March, 29–July 25, 2022

	Cumulativ	e incidence§		
Outcome	Controls [¶] (n = 1,902)	Second booster dose recipients (n = 1,902)	Risk difference (per 1,000 residents)	Vaccine effectiveness % (95% CI)**
SARS-CoV-2 infection ^{††}	101	75	-26	25.8 (1.2 to 44.3)
Hospitalization ^{§§}	9	3	-5	60.1 (-18.8 to 91.5)
Death ^{¶¶}	8	1	-7	89.6 (45.0 to 100.0)
Severe outcomes***	16	4	-12	73.9 (36.1 to 92.2)

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Lung Disease & Respiratory Health > Coronavirus > News >

WEBMD NEWS BRIEF

107-Year-Old Who Beat Spanish Flu Beats COVID

By Allison Aulds



Aug. 19, 2020 -- Anna Del Priore beat the Spanish flu when she was just 6 years old. Now, more than 100 years later, she's successfully recovered from COVID-19.

Sunrise Senior Living shared a photo on Facebook of the 107-year-old from New Jersey smiling and dancing after beating the virus.

Coronavirus Outbreak 2020

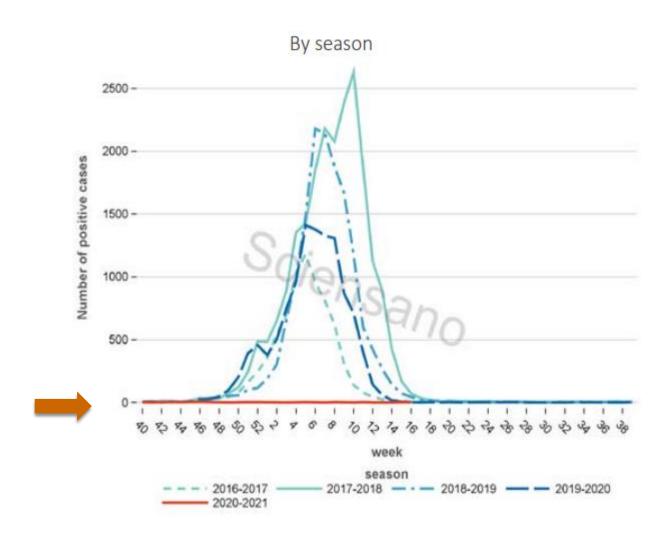


CORONAVIRUS HOME

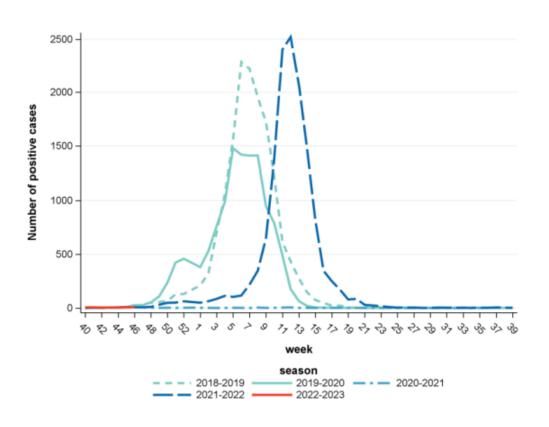
WebMD's Complete Coverage of the COVID-19 Outbreak

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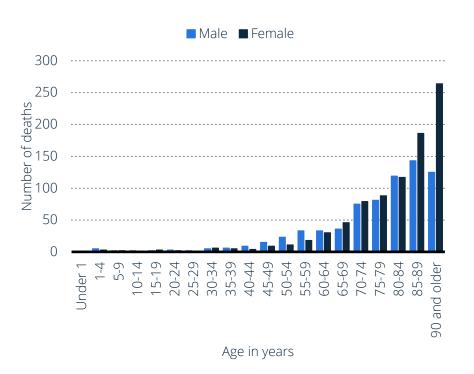
Influenza Belgium 2019 - 2020 - 2021

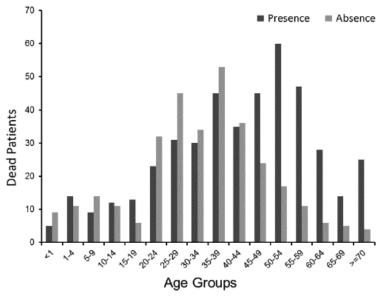


Influenza activity Belgium W47 2022



Influenza mortality by age and comorbidity



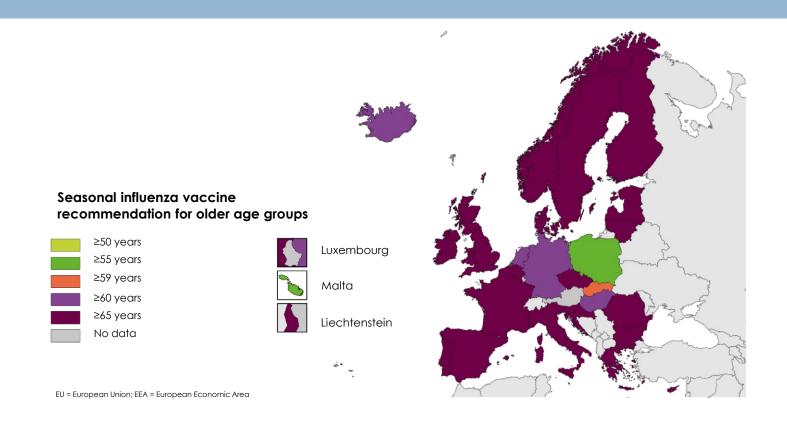


Note: United Kingdom (England, Wales); 2018

Further information regarding this statistic can be found on page 8.

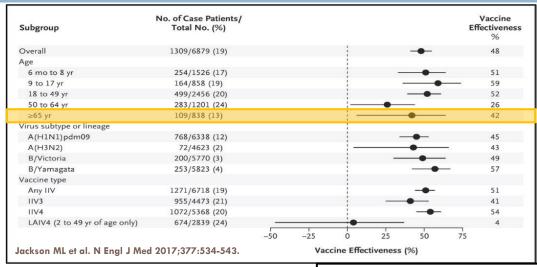
Source(s): Office for National Statistics (UK); ID 970800

Influenza vaccine recommendations for older adults



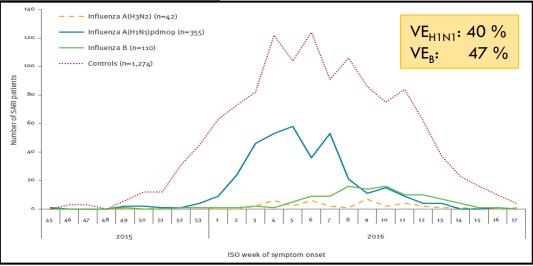
^{1.} European Centre for Disease Prevention and Control. Seasonal influenza vaccination and antiviral use in EU/EEA Member States – Overview of vaccine recommendations for 2017–2018 and vaccination coverage rates for 2015–2016 and 2016–2017 influenza seasons. Stockholm: ECDC; 2018.

Influenza Vaccine Effectiveness



Lab proven Influenza

Influenza Hospitalisation ≥ 65 y.



Influenza vaccination of HCW

Lab proven Influenza

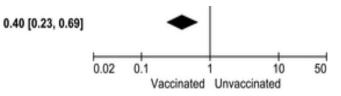
Total (95% CI) 962 502 100.0%

Total events 59 66

Heterogeneity: Tau² = 0.17; Chi² = 7.42, df = 4 (P = 0.12); I² = 46%

Test for overall effect: Z = 3.32 (P = 0.0009)

Test for subgroup differences: Chi2 = 5.14, df = 2 (P = 0.08), I2 = 61.1%



Absenteism incidence

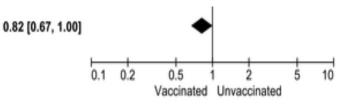
Total (95% CI) 1561 6512 100.0%

Total events 518 2633

Heterogeneity: Tau² = 0.02; Chi² = 7.85, df = 5 (P = 0.16); I² = 36%

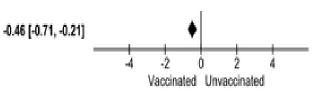
Test for overall effect: Z = 1.95 (P = 0.05)

Test for subgroup differences: Chi² = 5.85, df = 1 (P = 0.02), I² = 82.9%



Absenteism duration

Total (95% CI) 39 207 100.0% Heterogeneity: Tau² = 0.00; Chi² = 2.39, df = 3 (P = 0.49); l² = 0% Test for overall effect: Z = 3.57 (P = 0.0004) Test for subgroup differences: Chi² = 0.01, df = 1 (P = 0.91), l² = 0%



Influenza vaccination of **HCW** to protect residents of LTCF

Laboratory-proven influenza:

- little or no effect
- RD 0 (95% CI -0.03 to 0.03),
- 2 studies, 752 participants;
- low quality evidence

LRTI

- <u>- 6% to 4%</u>
- □ RD -0.02 (95% CI -0.04 to 0.01),
- 1 study of 3400 people;
- moderate quality evidence

Hospital admission for RTI:

- little or no effect
- □ (RD 0 (95% CI -0.02 to 0.02,
- one study of 1059 people;
- low quality evidence

Death from RTI & all cause:

- Varying direction and size of effect
- Very low quality evidence
- RTI death: two studies of 4459 people
- all cause deaths: four studies of 8468 people

Seasonal Influenza Vaccination

(2020-2021, HGR 9581)

A. Group 1:

- Pregnant women
- > 6 m. + chronic heart, lung, kidney, liver, metabolic, neuromuscular, immunological disease, BMI >35
- \geq 65 y.
- Institutionalized persons
- \blacksquare 6m 18 y. + chronic aspirine use

Group 2:

■ Health care workers

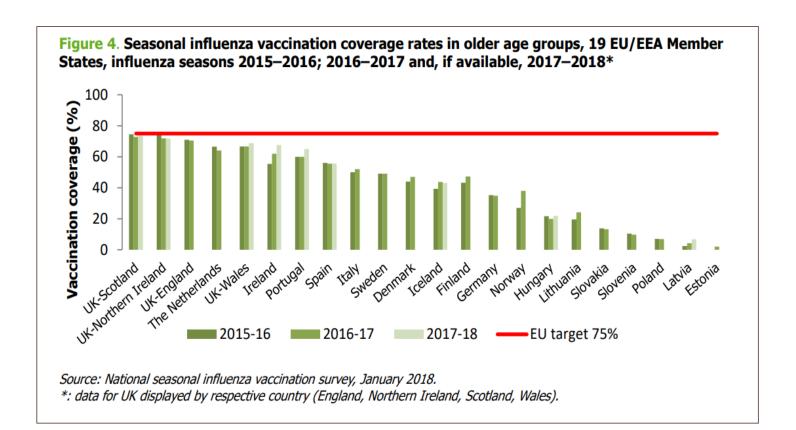
Group 3:

- Persons living with high risk person of group 1 or child < 6m.</p>
- B. Persons 50 65 y





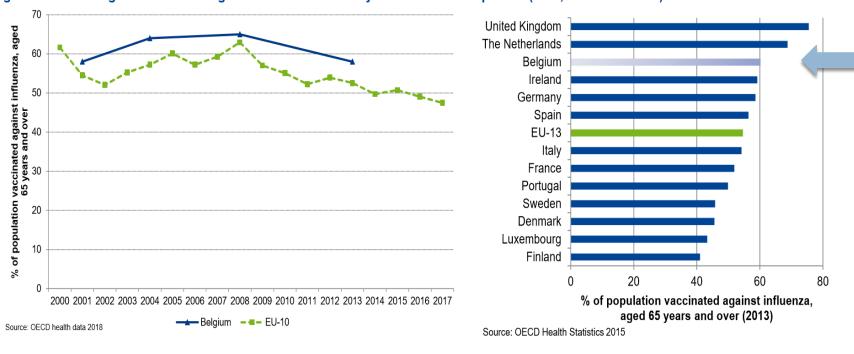
Influenza vaccination rates, EU



^{1.} European Centre for Disease Prevention and Control. Seasonal influenza vaccination and antiviral use in EU/EEA Member States – Overview of vaccine recommendations for 2017–2018 and vaccination coverage rates for 2015–2016 and 2016–2017 influenza seasons. Stockholm: ECDC; 2018.

Influenza vaccination rate ≥ 65 y. in Belgium and EU

Figure 146 – Coverage of vaccination against influenza for elderly: international comparison (2013, trend 2000-2017)



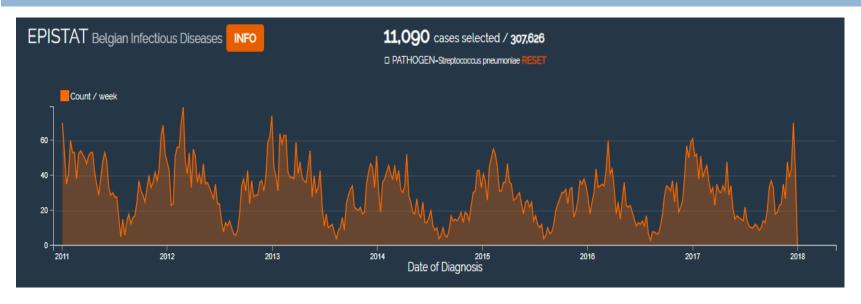
Source: Data from Belgium are based on the Health Interview Survey.

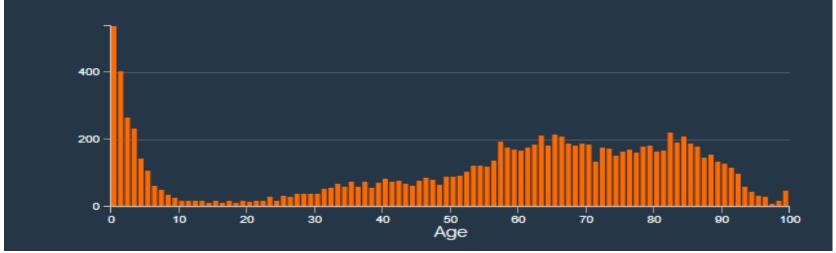
Note: fluctuation of the EU-15 is an artefact of the availability of the coverage data of the different countries.

Overview

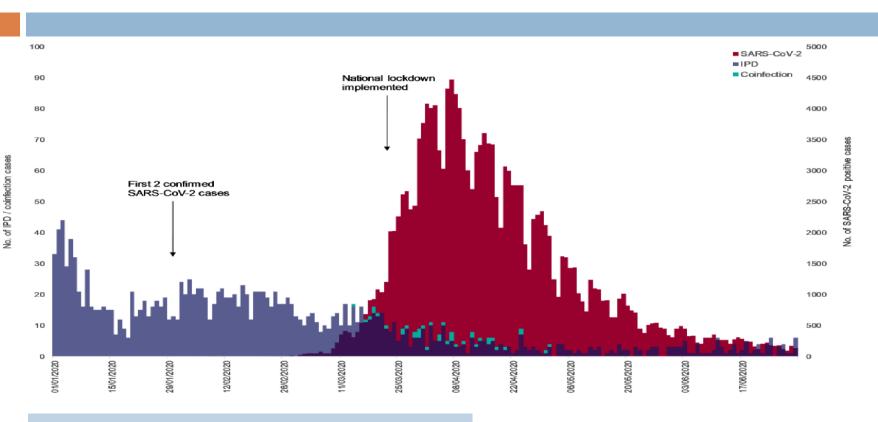
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Pneumococcal disease in Belgium





COVID-19 and IPD England



- IPD incidence after lockdown: 30 %
- Co-infection COVID + IPD:

• COVID-19: 0.025 %

• IPD: 3.5 %

CFR: x 7.8

Clinical Infectious Diseases, ciaa1728, https://doi.org/10.1093/cid/ciaa1728

IPD mortality

Global Mortality Associated With Pneumococcal Disease Remains High¹

Across all ages, number of deaths attributed to pneumococcal pneumonia is ≈1.5 million¹

Mortality Due to Pneumococcal Pneumonia²:

~1 out of 20

Mortality Due to Pneumococcal Bacteremia²:

~1 out of 6

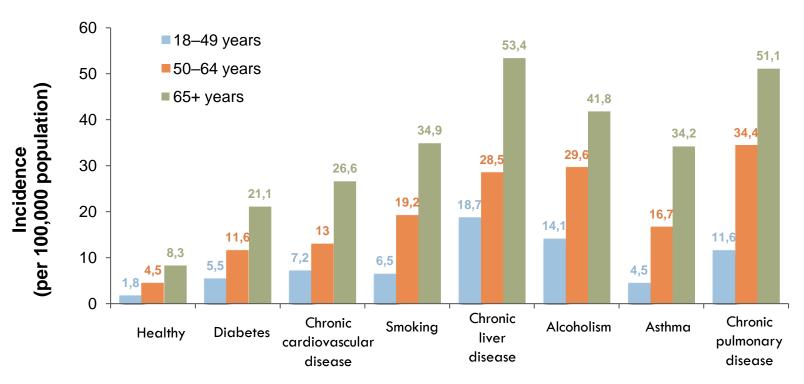
Mortality Due to Pneumococcal Meningitis²:

~1 out of 6



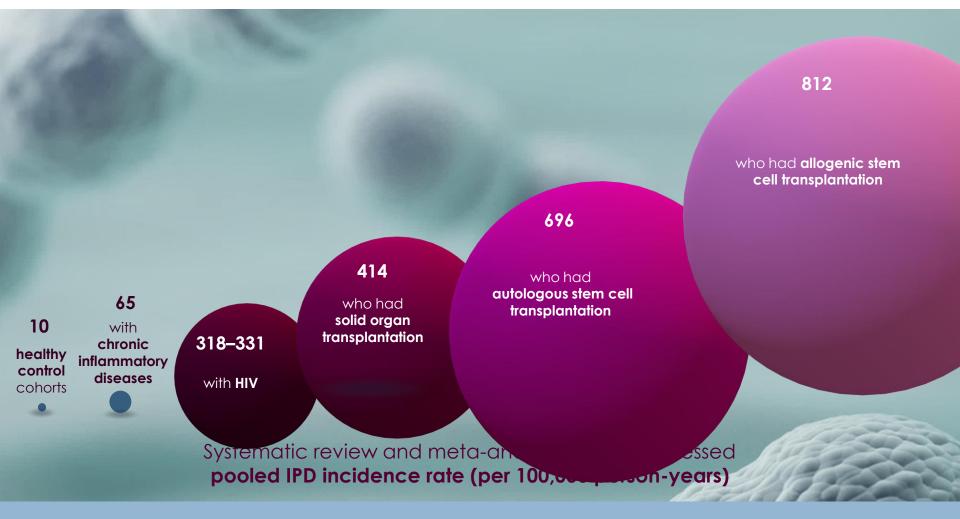
1. GBD 2015 Mortality and Causes of Death Collaborator. Lancet. 2016;388:1459–1544. 2. CDC. Adults: Protect Yourself with Pneumococcal Vaccines. 2019. www.cdc.gov/features/adult-pneumococcal/index.html. Accessed April 9, 2020.

IPD risk, age and comorbidities



Chronic risk condition or behavior

Risk for IPD in Immunocompromised Adults



HIV = human immunodeficiency virus; IPD = invasive pneumococcal disease 1. van Aalst M et al. *Travel Med Infect Dis*. 2018;24:89–100.

Risk groups for Pneumococcal disease

TARGET GROUPS:

Adults with high risk for PD

- Immunocompromised
- Asplenia (anatomic or functional)
- Sickle-cell disease and hemoglobinopathia
- CSF leakage or cochlear implant

Adults with comorbidity

- Chronic heart disease
- Chronic lung disease
- Chronic liver disease or ethylism
- Chronic kidney disease
- Chronic neurological conditions with aspiration risk
- Diabetes

July 2020, HGR 9562

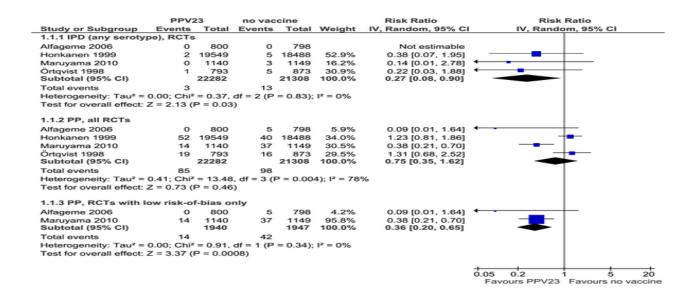




PPV23 effect on **IPD** and **PP** in elderly RCTs

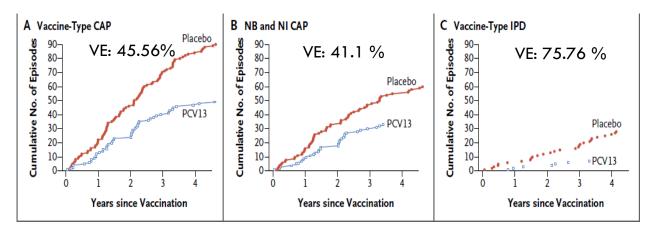
□ IPD: 73 % VE

□ PP: 64 % VE





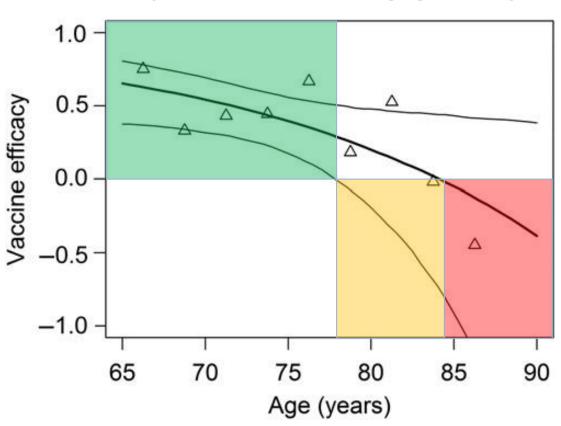
PCV13 in adults: CAPITA



	VE %	95 % CI	Р
VT CAP (PP)			
Total	45.56	21.82 - 62.49	< 0.001
Age groups			
< 75 y	52.54	24.09 - 70.99	0.001
75 – 84 y	46.43	-4.33 – 73.57	0.07
≥ 85 y	-100	-1156.63 - 57.78	0.51

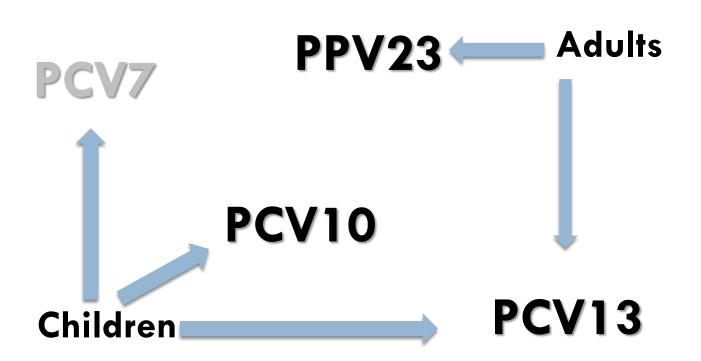
PCV13 vaccine efficacy and age

Model derived vaccine efficacy (VT-CAP-IPD in mITT population)

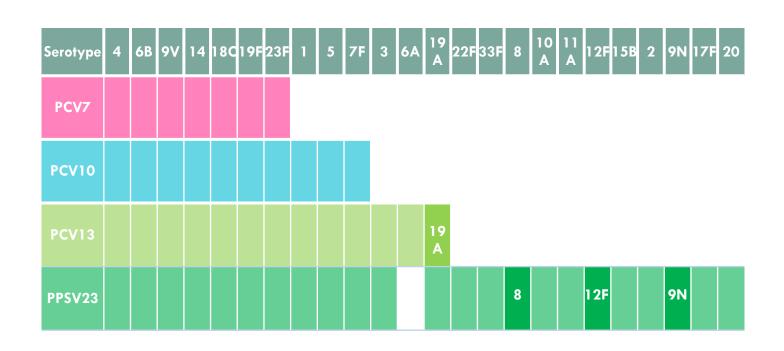


DOI: 10.1093/cid/civ686

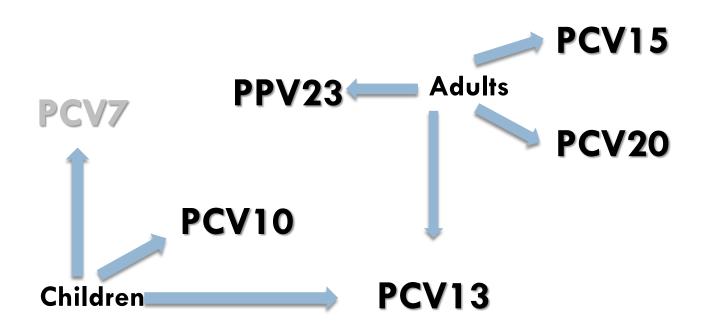
PCV7, PCV10, PCV13, PPV23



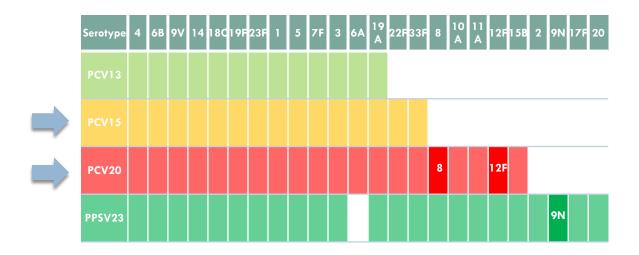
PCV7, PCV10, PCV13, PPV23



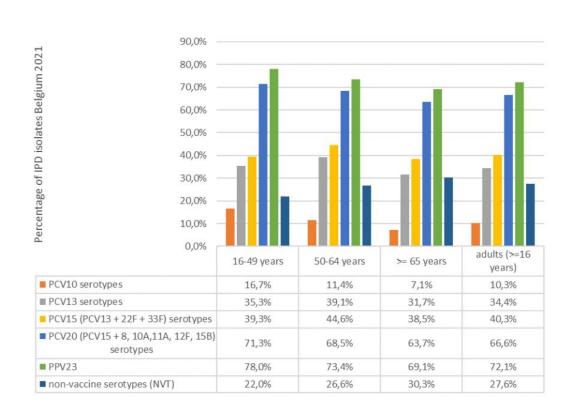
PCV7, PCV10, PCV13, PCV15, PCV20, PPV23



PCV15 and PCV20



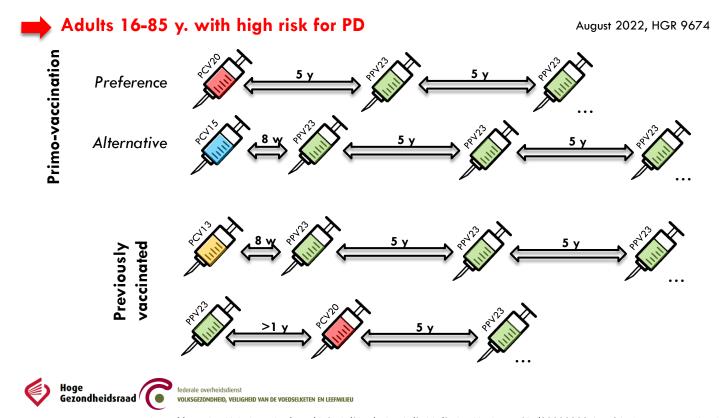
IPD serotypes included in PCV10, PCV13, PCV15, PCV20, and PPV23



Serotypes in IPD in adults Belgium, 2021

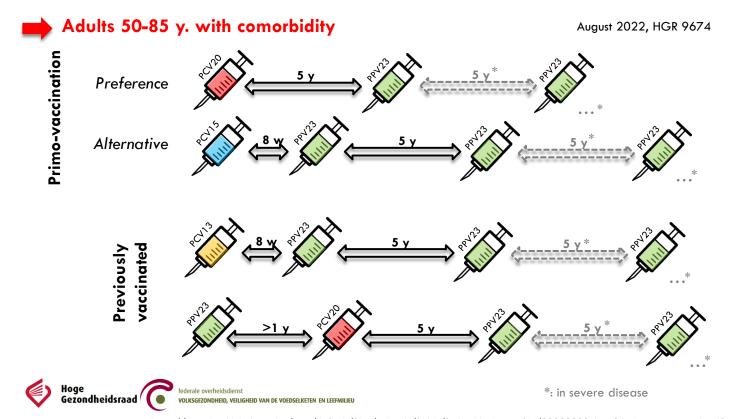
Serotype	Serotype geïncludeerd in vaccin	16-49 jaar (n=150)	50-64 jaar (n=184)	>= 65 jaar (n=366)
8	PCV20/PPV23	24.7%	17.4%	14.2%
19A	PCV13/PCV15/PCV20/PPV23	8.0%	12.5%	14.2%
3	PCV13/PCV15/PCV20/PPV23	10.7%	15.2%	9.8%
4	PCV13/PCV15/PCV20/PPV23	11.3%	8.2%	1.4%
6C	NVT	1.3%	3.3%	7.7%
9N	PPV23	4.0%	3.8%	4.4%
23B	NVT	4.0%	5.4%	3.0%
22F	PCV15/PCV20/PPV23	1.3%	2.7%	4.4%
15A	NVT	2.7%	3.3%	3.3%
11A	PCV20/PPV23	2.7%	2.2%	3.3%
16F	NVT	2.7%	4.3%	2.2%
10A	PCV20/PPV23	0.0%	2.2%	4.1%
33F	PCV15/PCV20/PPV23	2.7%	2.7%	2.5%
23A	NVT	4.0%	2.2%	2.2%
14	PCV13/PCV15/PCV20/PPV23	1.3%	1.6%	2.5%
12F	PCV20/PPV23	3.3%	1.6%	3.3%
35B	NVT	0.7%	0.5%	2.7%
31	NVT	0.7%	1.1%	1.9%
19F	PCV13/PCV15/PCV20/PPV23	2.0%	0.0%	1.4%
7B	NVT	0.7%	0.5%	1.6%
15B	PCV20/PPV23	1.3%	0.5%	1.1%
Andere serotypes		10.0%	8.7%	9.0%

Pneumococcal vaccination, Belgium



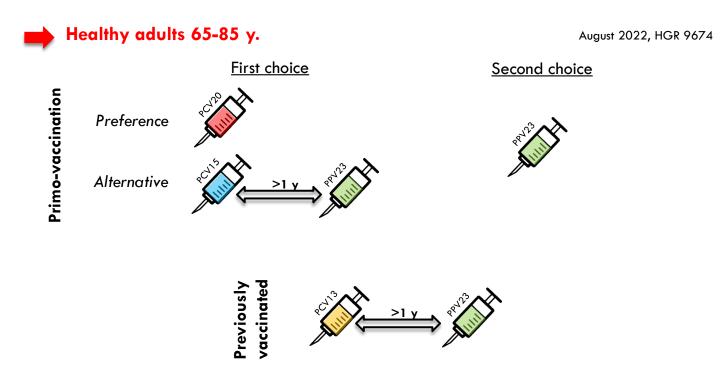
 $https://www.health.belgium.be/sites/default/files/uploads/fields/fpshealth_theme_file/20220908_hgr-9674_pneumo_vweb.pdf$

Pneumococcal vaccination, Belgium



https://www.health.belgium.be/sites/default/files/uploads/fields/fpshealth_theme_file/20220908_hgr-9674_pneumo_vweb.pdf

Pneumococcal vaccination, Belgium



Cost-effectiveness pneumococcal vaccination

Pfizer

Conclusions

Reimbursement of PCV13 in moderate/high-risk Belgian adults aged 65–84 years would be cost-effective from the Belgian healthcare perspective.

PLoS ONE 13(7): e0199427. https://doi.org/10.1371/journal. pone.0199427

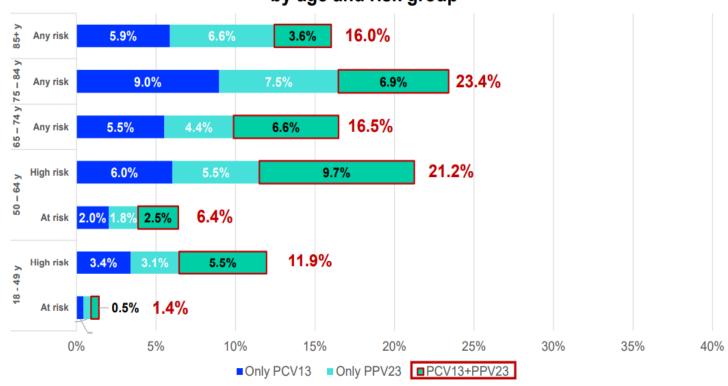
□ KCE

pneumococcal pneumonia. Pneumococcal vaccination would be most cost-effective in Belgium, when achieving high uptake with PPV23 in 75–84 year olds, as well as by negotiating a lower market-conform PPV23 price to improve uptake and cost-effectiveness.

Pneumococcal vaccination in Belgium

Implementation of Adult Pneumococcal Vaccination in Belgium – Low Uptake, Poor Adherence to Recommended Schedule

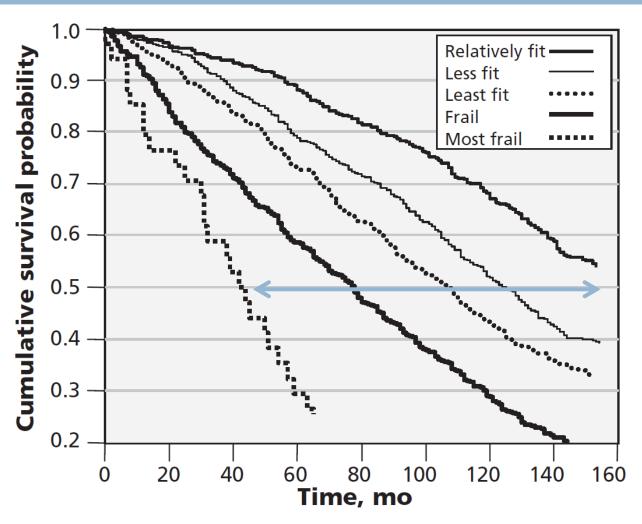
Percent of patients vaccinated for pneumococcal disase in the prior 5 years by age and risk group¹



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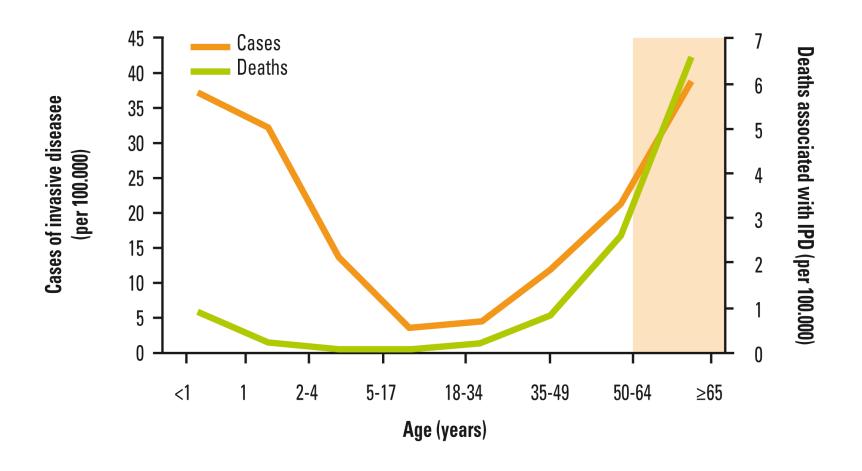
Frailty incidence and mortality



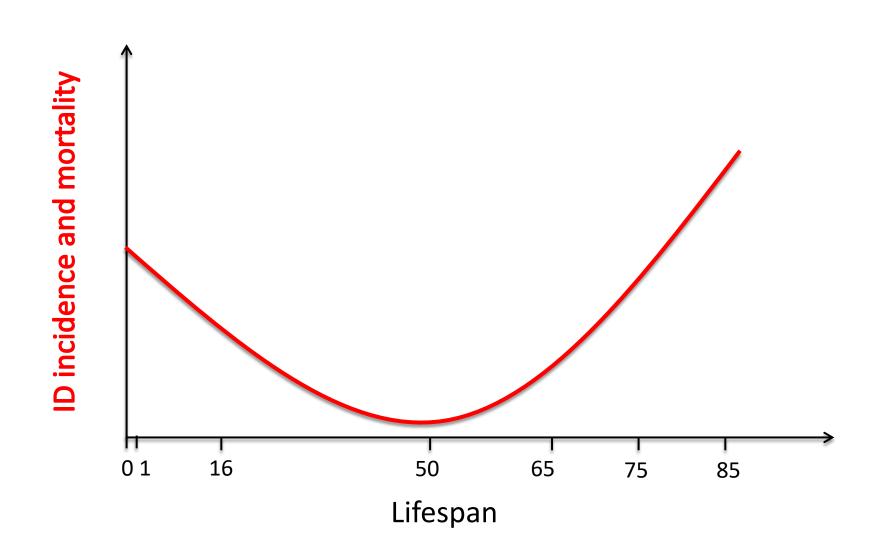
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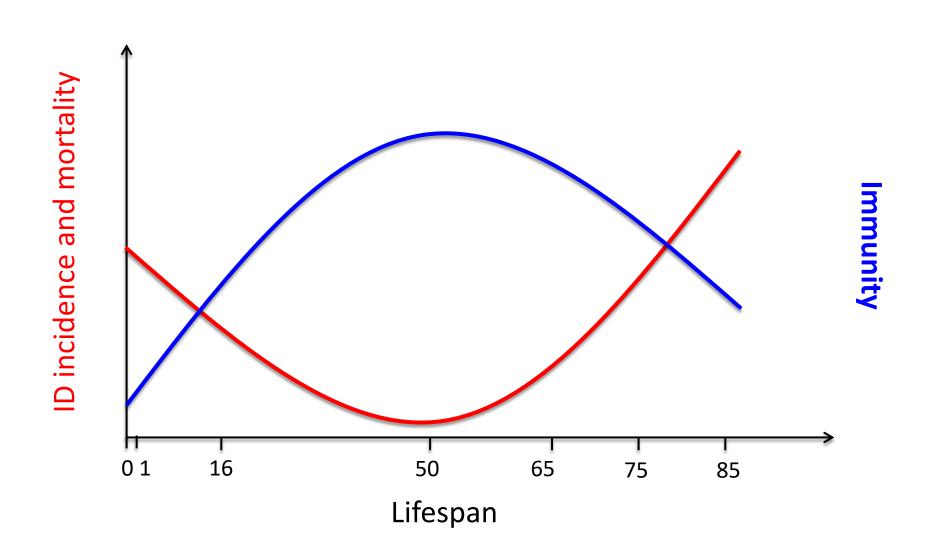
Incidence and mortality of infectious diseases



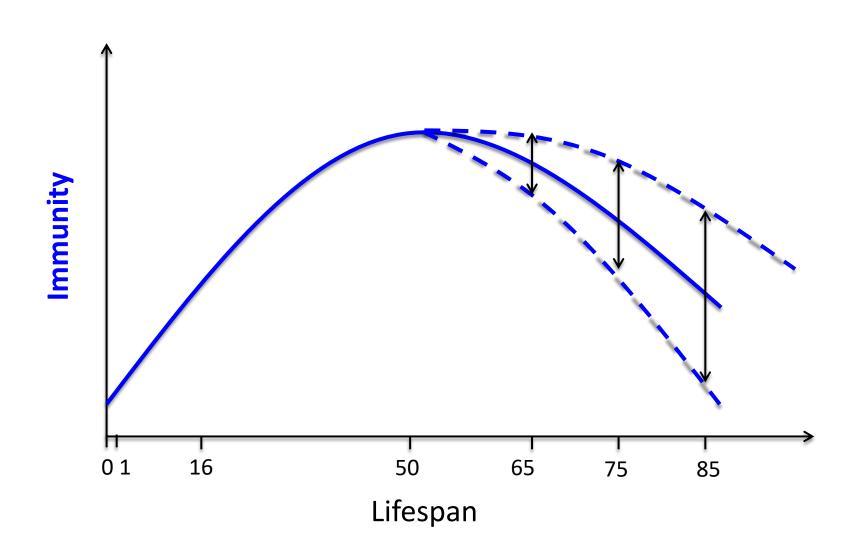
Incidence and mortality of infectious diseases



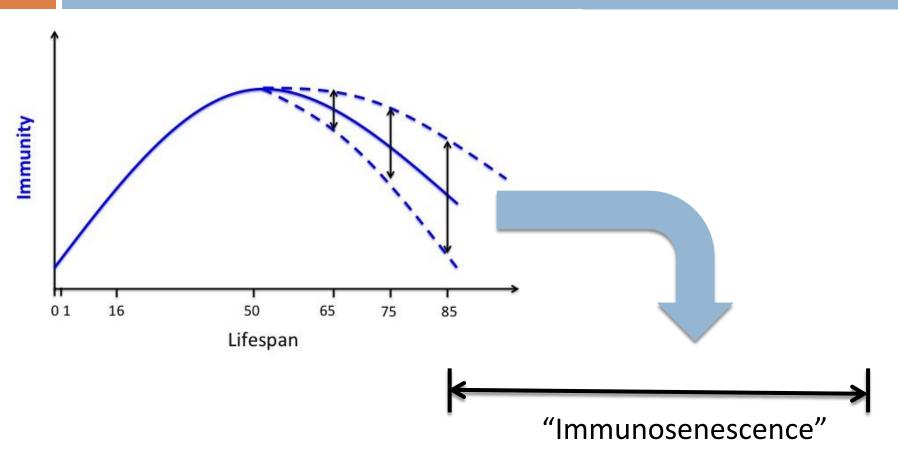
Infectious diseases and immunity



Immunity and aging



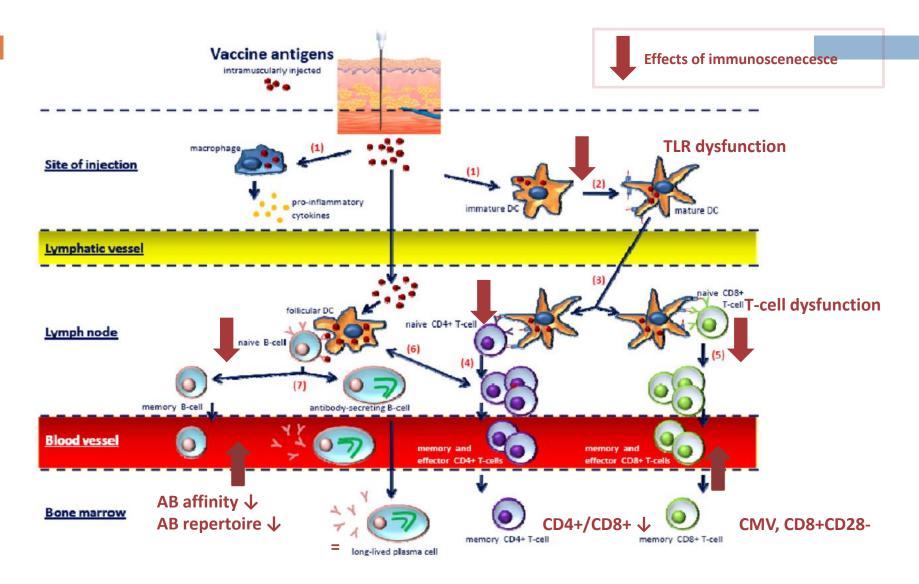
Immunity and aging



Poor immunity
Poor vaccine respons
VPD

Good immunity
Good vaccine respons
No/ less VPD

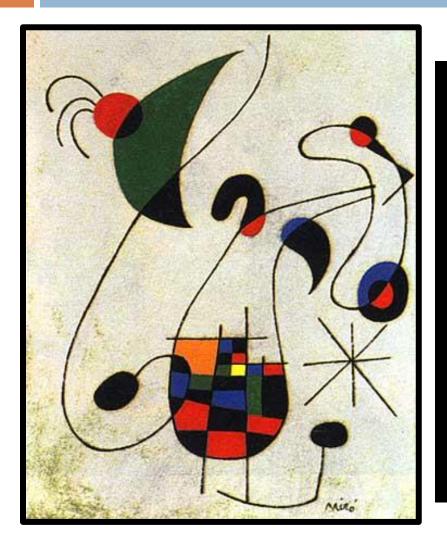
Immunosenescence affects the innate and adaptive immune response

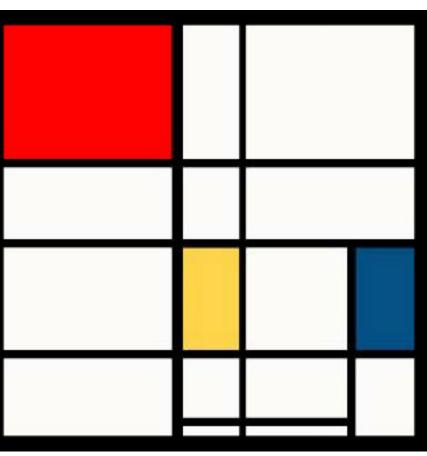


Overview

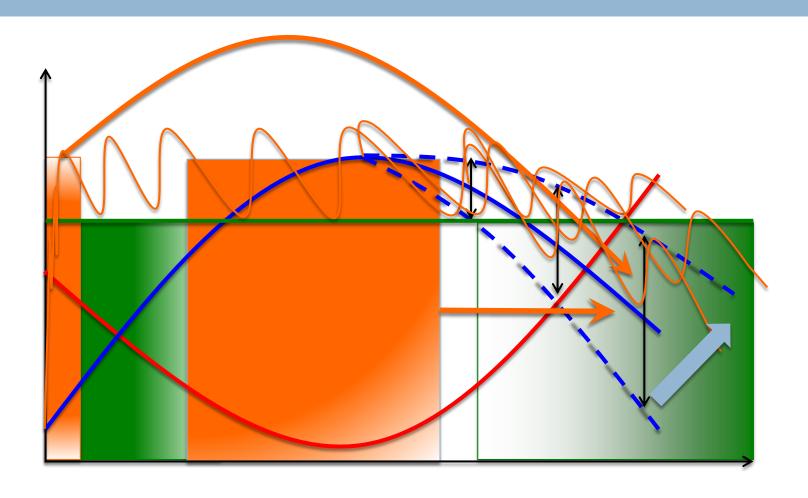
- Coronavirus in the older prerson
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 - Herd immunity and vaccination
- Immunosenescence and vaccination
- Ex. Herpes Zoster vaccination

The art: "Vaccination strategies"

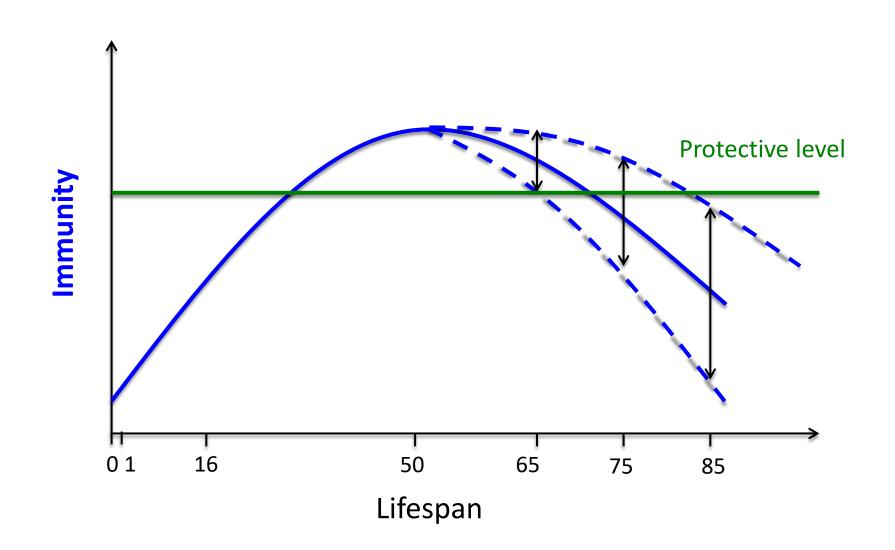




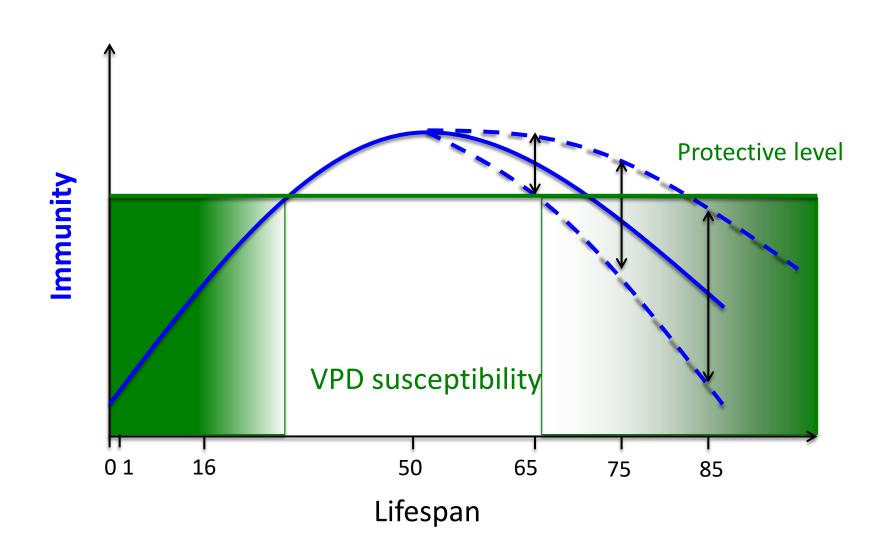
The art: "Vaccination strategies"



Immunity and VPD protection



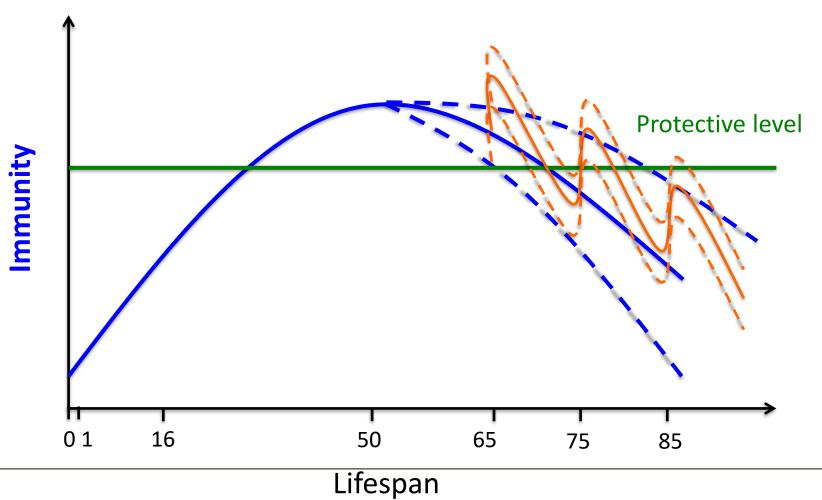
Immunity and VPD protection



Overview

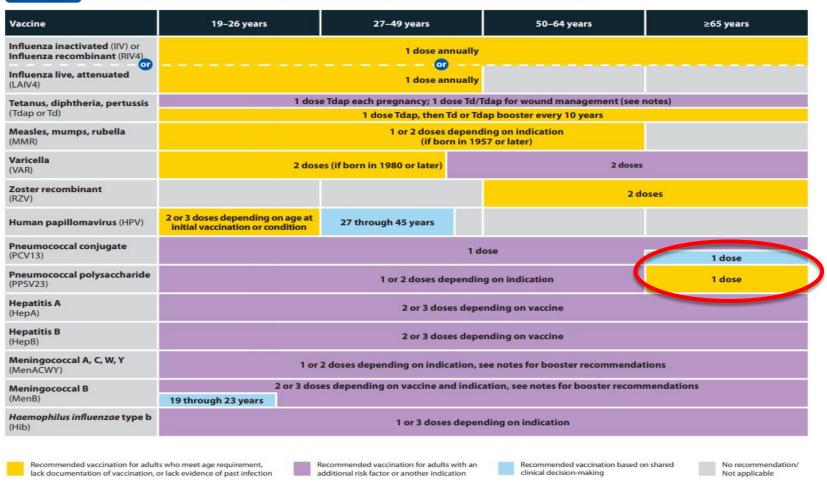
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Vaccination strategy \ge 65 y.

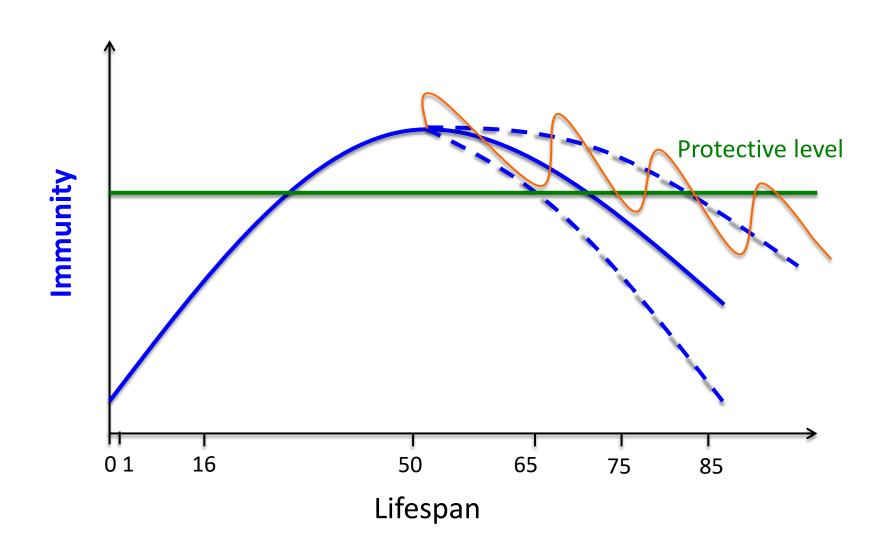


Vaccination strategy: Age based



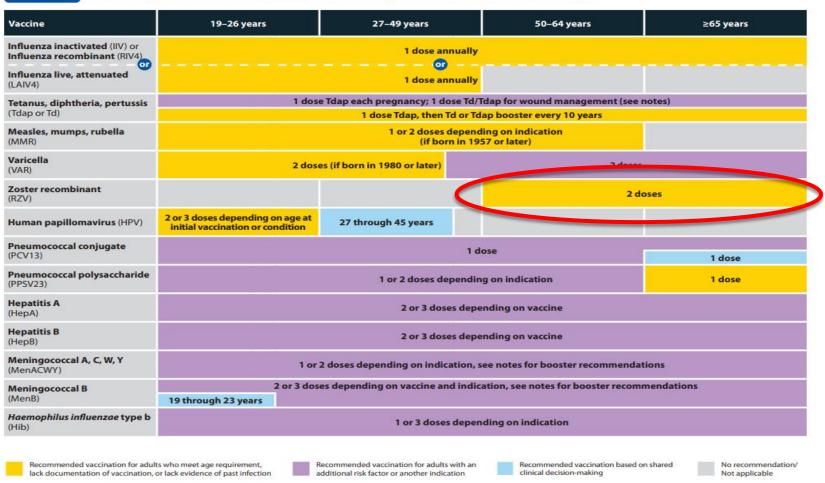


Vaccination strategy ≥ 50 y.

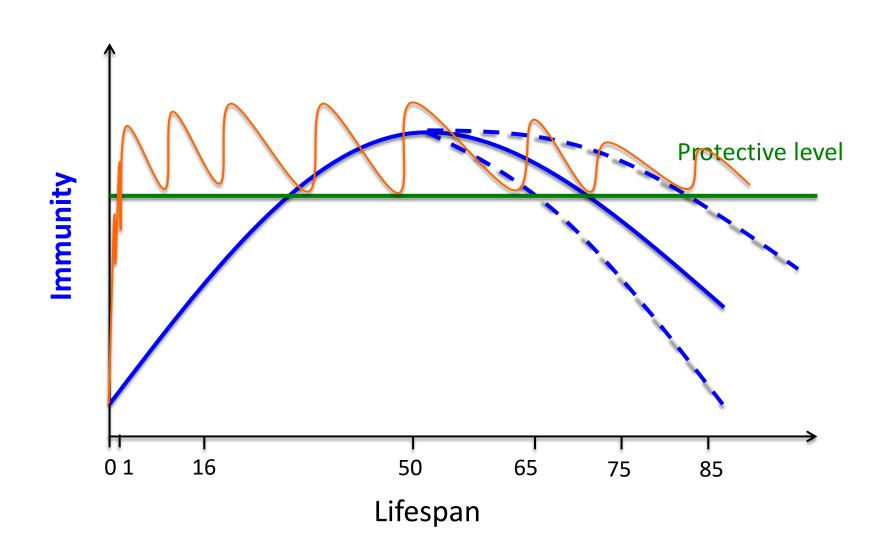


Vaccination strategy: Age based

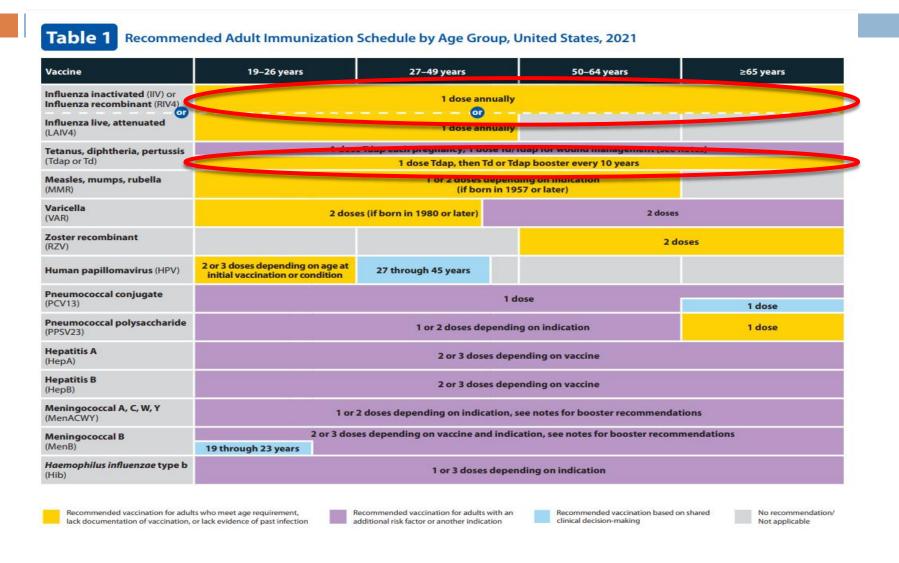




Lifelong vaccination strategy



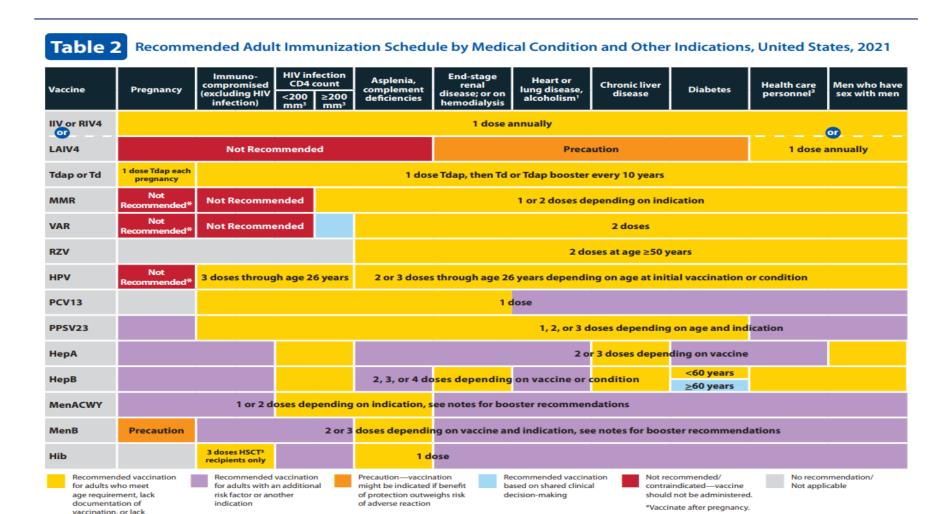
Vaccination strategy: Lifelong vaccination



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Vaccination strategy: Risk based



^{1.} Precaution for LAIV4 does not apply to alcoholism. 2. See notes for influenza; hepatitis B; measles, mumps, and rubella; and varicella vaccinations. 3. Hematopoietic stem cell transplant.

evidence of past infection

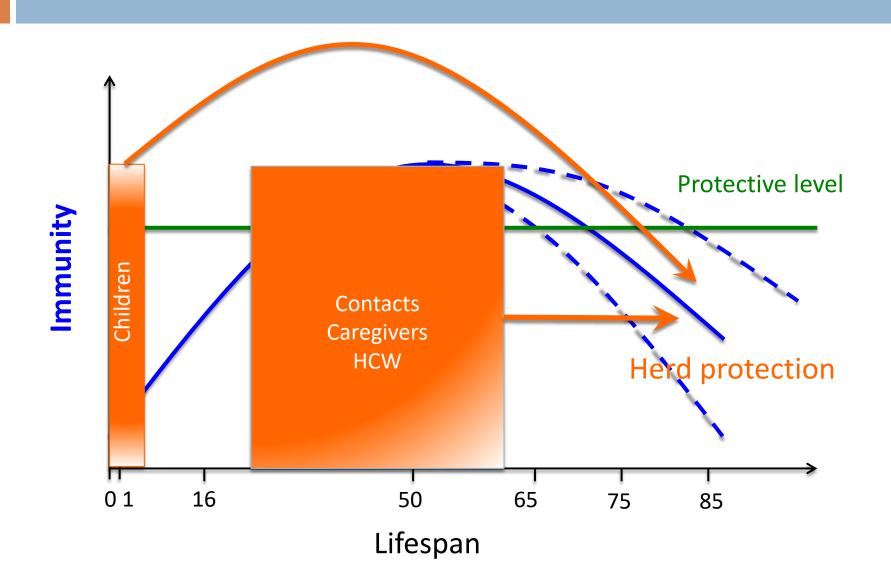
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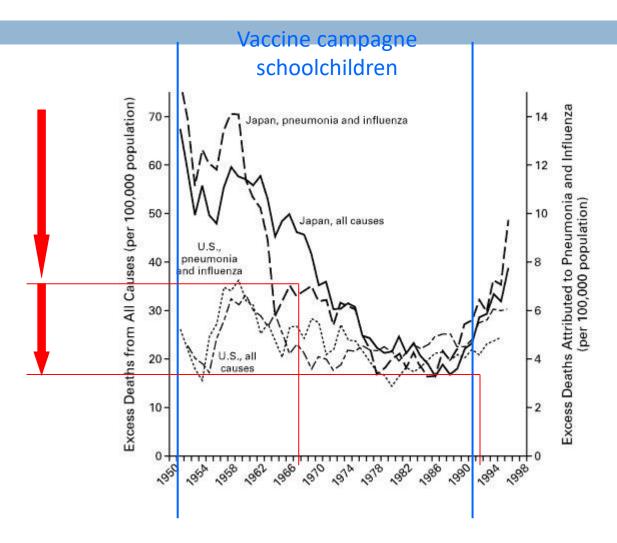
Herd immunity



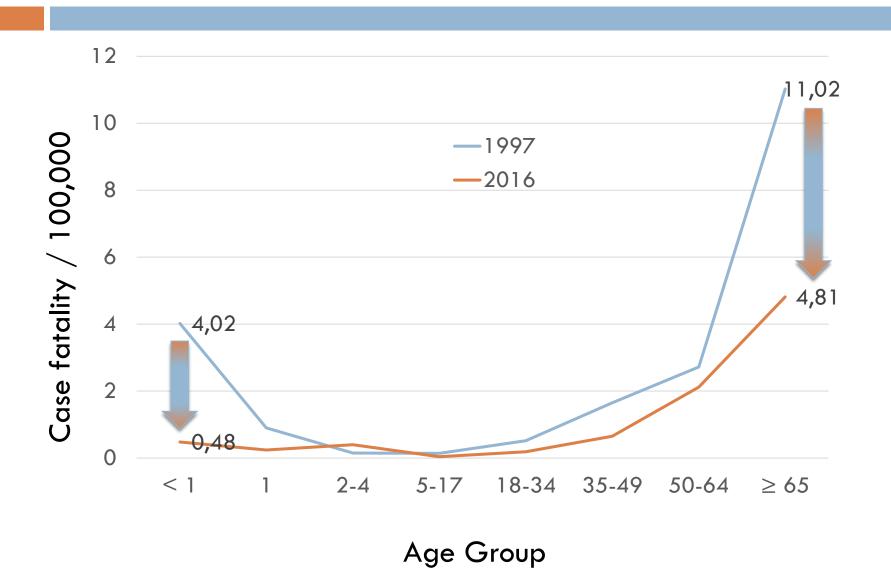
Herd vaccination strategy

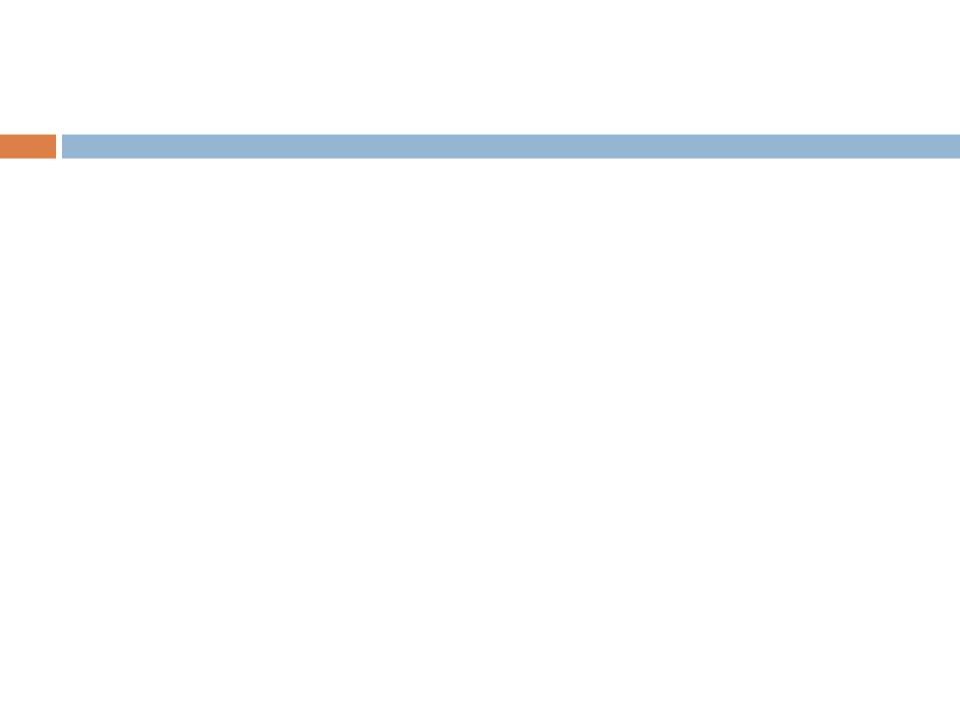


Herd vaccination strategy: Children and Influenza



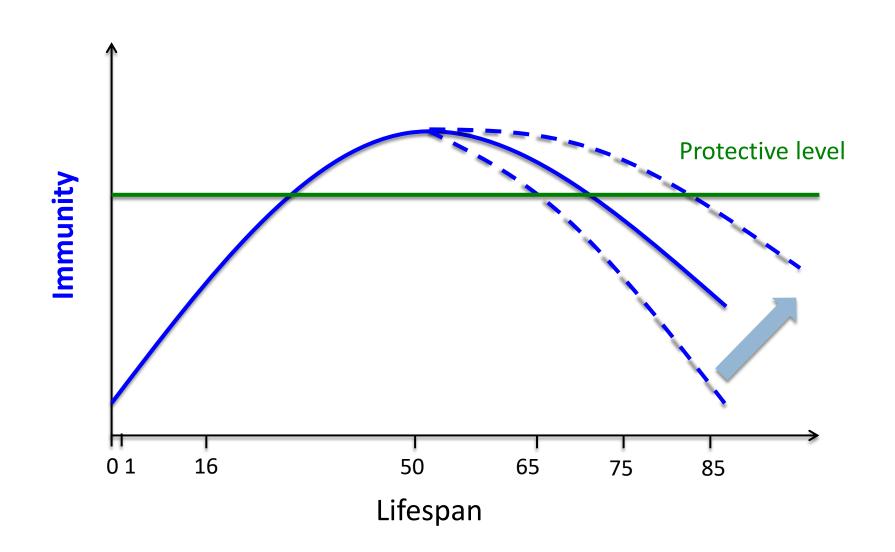
IPD case fatality, USA





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Tackling frailty

Tackling comorbidities

□ Immune - rejuvenation

3R's of REJUVENATION Restoration maintenance of a normal thymic environment by using cytokines, growth hormone, sexsteroids, growth factors and nutrients Replacement replacing the immune function lost by ex vivo generated cells Reprogramming regulating telomere length and stability Ability of the Ability of the cell-mediated immunity cell-mediated immunity **ADVANCING AGE** to respond to to respond to new antigens new antigens **IMMUNE RISK PROFILE** inverted CD4:CD8 ratio ≥ naïve T-cells 对 senescent T-cells CMV seropositivity INFLAMMAGING Switch Th1-like cytokine response to Th2-like cytokine response 7 IL6, TNF-α, IL-1β, IL-18, IL-12 **IMMUNOSENESCENCE**

Vaccines

Enhancing vaccine effect

- Vaccine coverage (epidemiology, valency)
- High dose vaccines
- Adjuvanted vaccines
- \square Vaccine formulations (virosome, ID, IN + SC/IM, vector, mRNA...)
- Booster vaccines

- More memory B & T cells
- Higher postvaccination Ab titers
- More functional Ab (OPA)
- Higher efficacy effectiveness

Vaccine coverage ≈ epidemiology

- Influenza vaccination
 - □ TIIV QIIV
 - H1N1, H3N2,
 - B victoria, B Yamagata
 - □ PCV 7 → PCV10 → PCV15 → PCV20 → PPV23
 - Coronavirus vaccine....??

Effectiveness **newer IV** against lab-confirmed influenza

MF-59 TIV vs. no vaccine \geq 65 y.

Outcomes	Vaccine effectiveness* (95% CI)
Influenza (any)	VE 44.9% (22.7 to 60.8)
Influenza A(H1N1)	VE 61.2% (43.7 to 73.3)
Influenza A(H3N2)	VE 10.6% (-24.5 to 35.7)
Influenza B	VE 28.5% (5.4 to 46.0)

HD-IIV vs. IIV

 \geq 65 y.

CB-TIV vs. no vaccine 18 - 49 y.

Outcomes	Relative effect (95% CI)
Influenza (any)	VE 70% (61% to 77%)
Influenza A(H1N1)	VE 82% (71% to 89%)
Influenza A(H3N2)	VE 72% (39% to 87%)
Influenza B	VE 52% (30% to 68%)

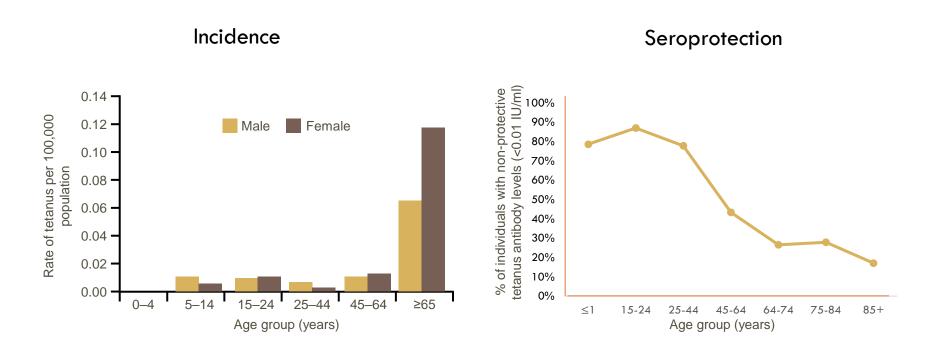
R-IIV vs. IIV

 \geq 50 y.

Outcomes [¥]	Relative effect (95% CI)
Influenza (any)^	VE 30% (10 to 47)

Booster vaccination

Tetanus

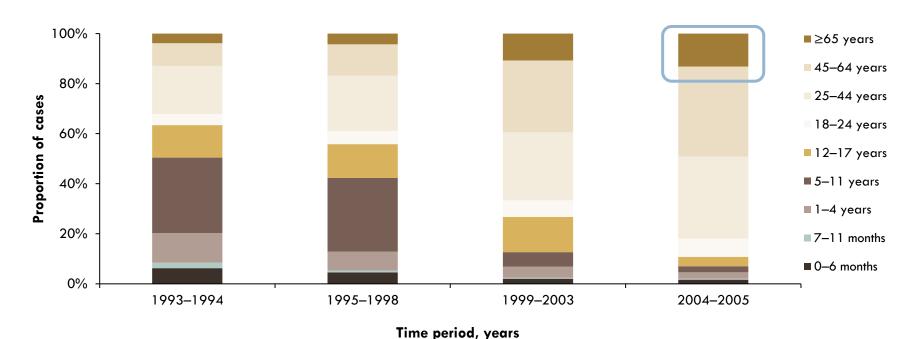


^{1.} EUROHEALTH, 2016. Demographics and diversity in Europe: new solutions for health. http://www.euro.who.int/en/about-us/partners/observatory/publications/eurohealth/full-list-of-past-issues/demographics-and-diversity-in-europe-newsolutions-for-health2 (accessed February 2021); 2. Filia A et al. Vaccine 2014;32:639–644; 3. World Health Organization (WHO). Wkly Epidemiol Rec 2017;6:53–76; 4. European Centre for Disease Prevention and Control. Annual Epidemiological report 2015. Tetanus. Stockholm: ECDC; 2016

Booster vaccination

Pertussis

Australia, 1993–2005 (n=35,695)

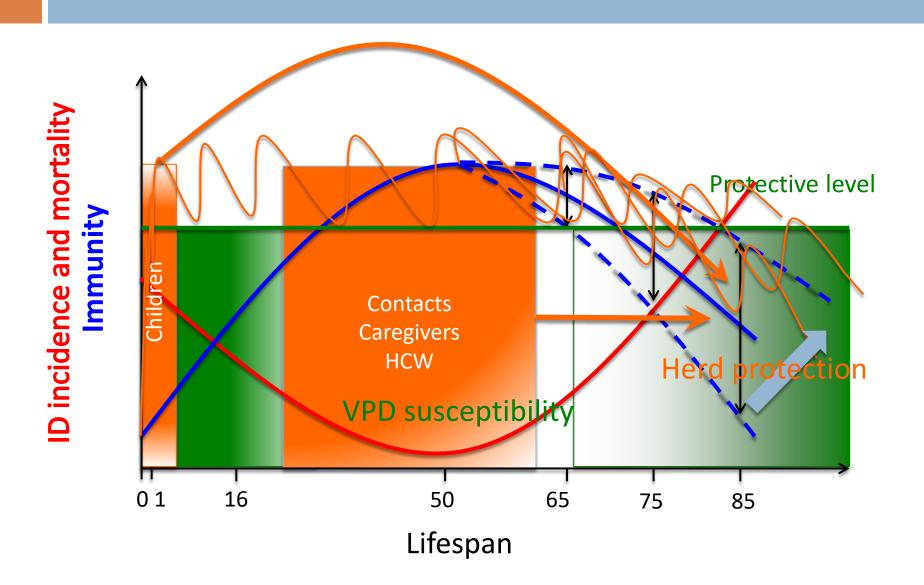


By 2005, nearly 90% of notified pertussis cases were in adults

Vaccination strategies in older persons

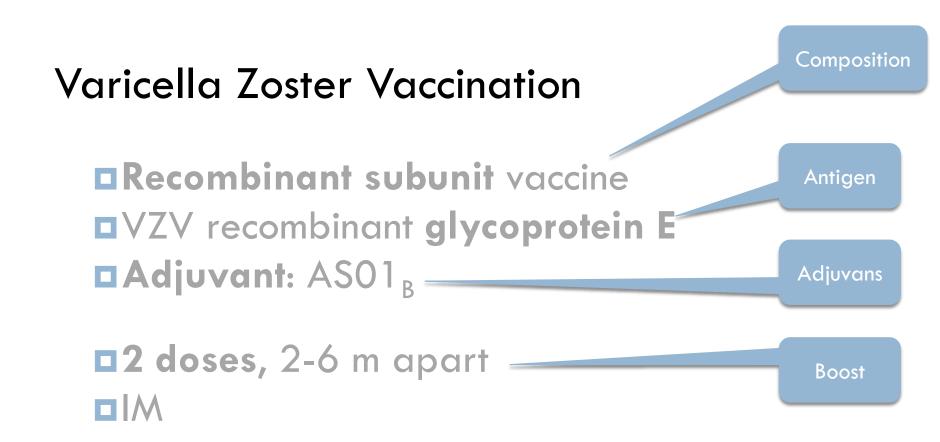
- Prevent/treat (pre)-frailty and comorbidities
 Vaccinate with most immunogenic vaccines
 Boost
 Vaccinate older (fit) adults
 Vaccinate (grand)(grand)children and contacts
 Vaccinate HCW
 - ■Lifelong vaccination strategy

The science: "Vaccination strategies"

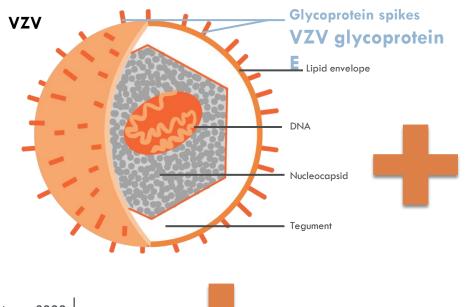


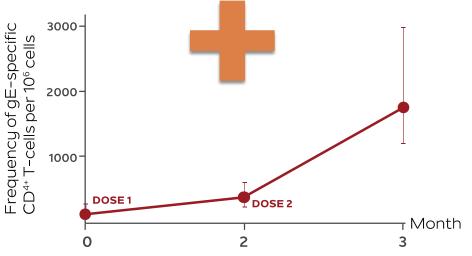
Overview

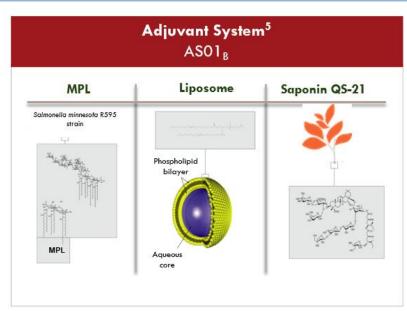
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Adjuvanted Recombinant Zoster Vaccine







References: 1. Dendouga N, et al. Vaccine. 2012 Apr;30(20):3126-35. 2. Lal H, et al. N Engl. J.Med. 2015 May;372(22):2087-96. 3. Lecrenier N, et al. Exp Rev Vaccine. 2018 Juri; 17(7):619-634. 4. Zerboni L, et al. Nat Rev Microbiol. 2014 Mar;12(3):197-210.

References: 1. Dendouga N, et al. Vaccine. 2012 Apr;30(20):3126-35. 2. Leroux-Roels G, et al. Clin Immunol. 2016 Aug;169:16-27. 3. Bharucha T, et al. Hum Vaccin Immunother. 2017 Aug;13(8):1789-97. 4. GlaxoSmithKline. Shingrix European public assessment report, Annex I: Summary of product characteristics: EMA; [updated August 2020; accessed October 2020]. 5. Garçon N, et al. Vaccine adjuvants. Amsterdam: Elsevier; 2011.

References: 1. Chlibek R, et al. Vaccine. 2014 Mar;32(15):1745-53. 2. Chlibek R, et al. 2014 Mar;32(15 Suppl):1745-53. 3. Schwarz TF, et al. Hum Vaccin Immunother. 2018 Jun;14(6):1370-7.

Varicella – Herpes zoster

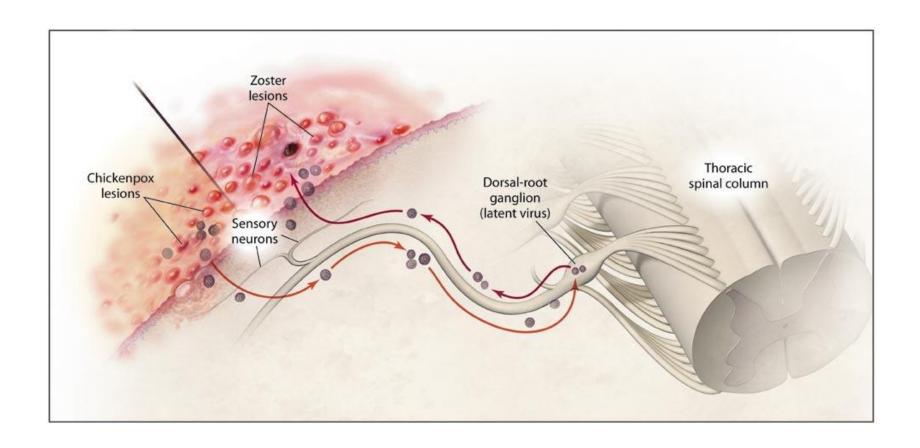


Windpokken Waterpokken

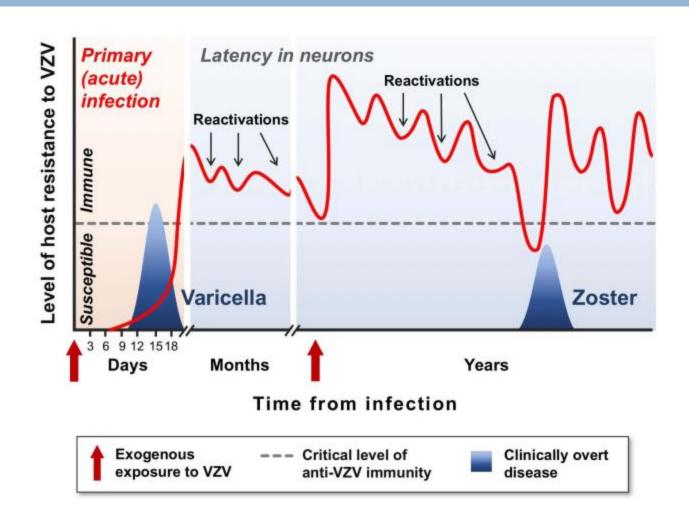


Zona Gordelroos

Varicella – Herpes zoster

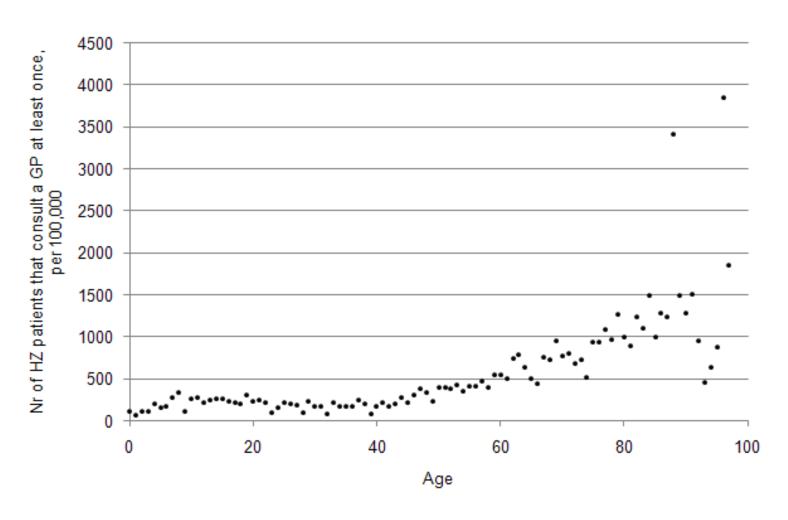


VZV reactivation

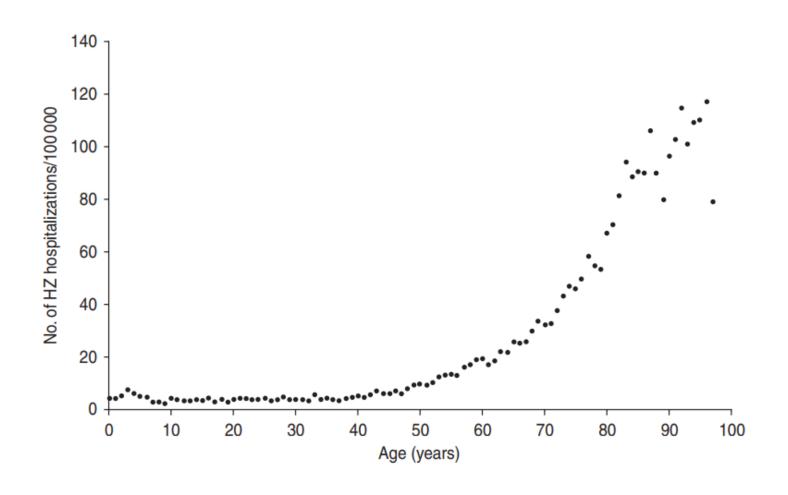


New Microbiologica, 41, 2, 95-105, 2018,

HZ incidence, GP visits (Be)



HZ incidencee, hospitalisations (Be)



Risk factors for HZ

Table 1. Common Risk Factors for Herpes Zoster

Older age (\geq 50 years of age)

Untreated depression

Inflammatory bowel disease^a

Rheumatologic disorder (eg, rheumatoid arthritis, SLE,

giant cell arteritis, dermatomyositis^a)

Chronic kidney disease and hemodialysis

HIV infection

Dermatologic disorder (eg, psoriasis treated with systemic corticosteroids)^a

Solid and hematologic malignancy, particularly when undergoing chemotherapy (eg, bortezomib)

PBSCT or SOT

Diabetes mellitus

Treatment with a systemic corticosteroid agent

Micronutrient deficiency

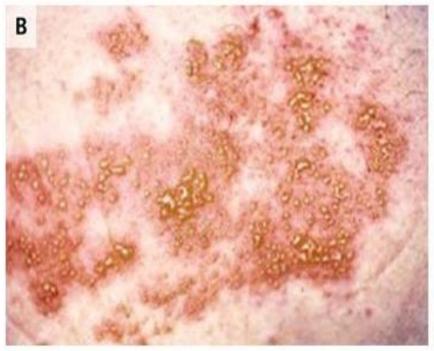
^aA recent study showed that treatment with anti-tumor necrosis factor agents does not increase the risk for herpes zoster.¹²

Abbreviations: HIV, human immunodeficiency virus; PBSCT, peripheral blood stem cell transplantation; SLE, systemic lupus erythematosus; SOT, solid organ transplantation.

DOI: 10.3810/pgm.2013.09.2703

Cutaneous HZ





Clinical manifestations of HZ

- Cutaneous HZ (localized or disseminated)
- Zoster sine herpete
- HZ ophthalmicus
- Ramsay Hunt syndrome
- Vasculopathy:
 - large vessel unifocal granulomatous arteritis
 - small vessel multifocal vasculopathy
- Meningitis/meningoencephalitis
- Myelopathy
- Focal motor weakness
- Acute retinal necrosis
- Progressive outer retinal necrosis

DOI: 10.3810/pgm.2013.09.2703

Disseminated HZ



HZ ophthalmicus





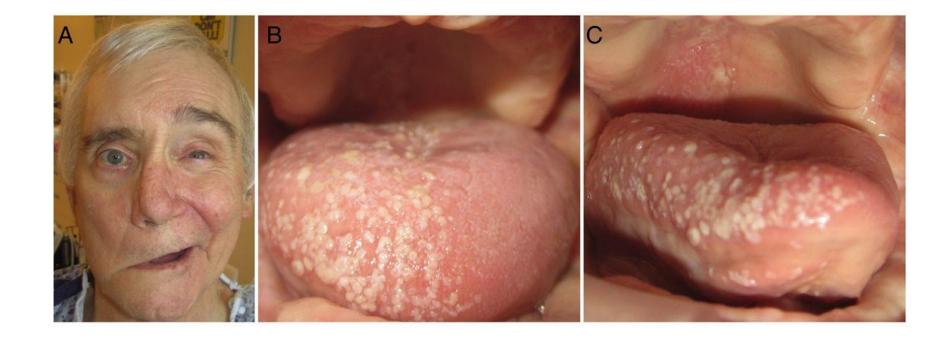
Hutchinson's teken

Zona ophthalmica

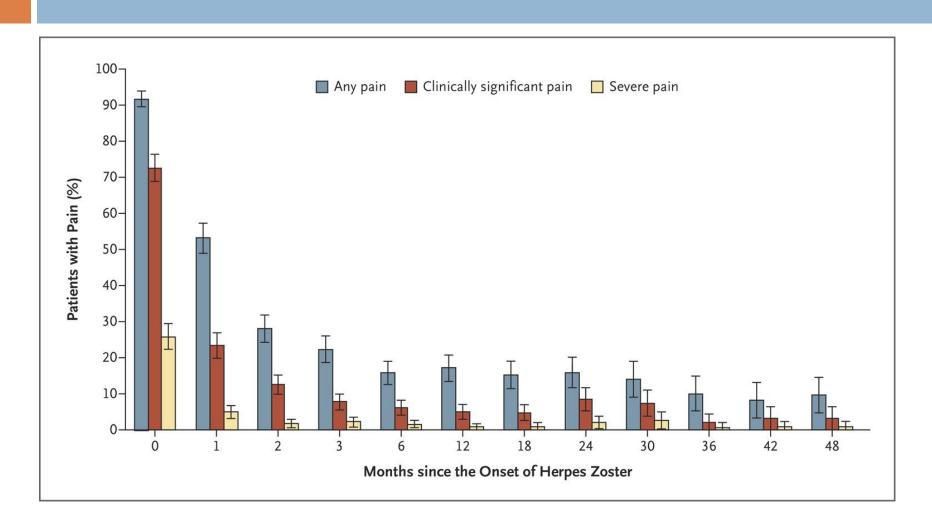
Ramsay-Hunt



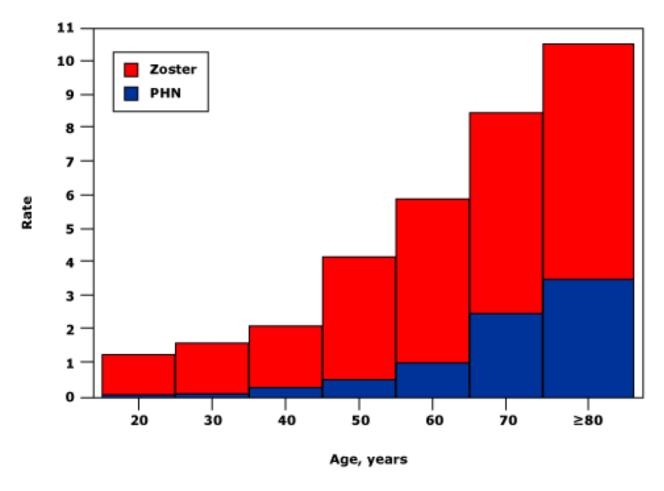
Ramsay-Hunt



PHN



HZ en PHN (USA)



Harpaz R, Ortega-Sanchez IR, Seward JF. MMWR Recomm Rep 2008; 57:1

Risk factors for PHN

Table 3. Patient Risk Factors for PHN

Older age (\geq 50 years of age)

Immunocompromised

Micronutrient deficiency

Low income

Involvement of trigeminal dermatomes or brachial plexus during the acute episode of Hz

Hz ophthalmicus with keratitis, conjunctivitis, or uveitis

Severe prodromal pain

Acute severe pain

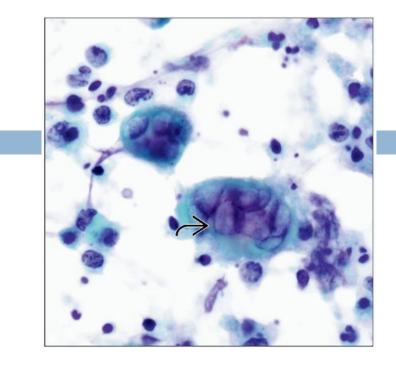
Involvement of larger surface area by skin lesions

Numerous lesions

Not receiving antiviral drugs

Abbreviations: Hz, herpes zoster; PHN, postherpetic neuralgia.

Dianosis of HZ



- Direct fluorescent antigen assay,
- Viral culture,
- Quantitative PCR for the detection of VZV DNA in skin lesion / visceral organ samples, CSF
- Antibody testing (immunoglobulin G)
- CSF and visceral samples: PCR and IgG

Antiviral treatment of HZ

DOI: 10.3810/pgm.2013.09.2703

- □ ≤ 72 h of lesion onset
 - shorter duration of viral shedding and new lesion formation,
 - reduction of the duration of zoster-associated acute pain
 - acceleration of cutaneous healing
- □ > 72 h of lesion onset
 - □ older adults (≥ 60 years of age) with severe pain and a large area of skin involvement
 - immunocompromised patients
 - continued new vesicle formation
 - visceral, neurologic, or ocular complications, including HZO
- Aciclovir, valaciclovir, brivudine
 - Bio-availability valacicovir (55 %) > acyclovir (10-20 %)

Prevention of PHN

- Acyclovir does not prevent PHN
 - □ 4m RR 0.75, (95% CI 0.51 to 1.11)
 - □ 6m RR 1.05, 9(5% CI 0.87 to 1.27).
 - □ 4w ?
- Prevention of PHN during acute illness
 - amitriptyline,
 - ganglion blockade with bupivacaine and dexamethasone,
 - repetitive epidural injection of methylprednisolone acetate and bupivacaine
- □ Corticosteroids (10 14 d tapering + antiviral)
 - No prevention of PHN
 - □ Acute neuritis ↓ (pain, activities, sleep)

PHN pharmacological treatment

N Engl J Med 2014; 371:1526-1533

Agent	Average Effective Dose in Clinical Trials	Starting Dose	Dose Adjustment	Number Needed to Treat (95% CI)†	Side Effects	Precautions
Topical treatments						
Lidocaine patch	5%; up to 3 patches/day	Maximum of 3 patches/day for a maximum of 12 hr		2.0 (1.4–3.3) ²⁰	Local erythema	
Capsaicin cream	0.075%; 4 applica- tions/day	NA		3.3 (2.3-5.8)20	Pain on application, local erythema, rash	Avoid eyes and nose
Capsaicin patch	8%; application time of 30–90 min	NA		11.0 (6.1–62.0) ²²	Pain on application, local erythema, rash; systemic adverse events in <5% of study participants;	
Oral treatments						
Gabapentin	2572 mg/day	2572 mg/day 100 mg Increase each of 3 times daily 100–300 mg ated; maxim day, but unl 3600 mg/da clinicians		4.4 (3.3–6.1) ²⁰	Sedation, dizziness, peripheral edema	Avoid in patients with renal insufficiency
Pregabalin	398 mg/day	50–75 mg twice daily	Increase to 300 mg daily after 3–7 days, then by an additional 150 mg daily every 3–7 days as tolerated, to a maximum dose of 600 mg daily	4.2 (3.4–5.4) ^{20,23}	Same as with gabapentin	Same as with gabapentin
Tricyclic antide- pressants (off- label use)	Amitriptyline, 95 mg/day; or nortriptyline, 122 mg/day	10–25 mg at bedtime	Increase by 10–25 mg every 3–7 days as tolerated to 75–150 mg/day with caution as side effects permit; if blood level of active drug and its metabolite is >100 ng/ml, continue dose adjustment very cautiously	2.6 (2.1–3.5) ²⁰	Sedation, dry mouth, blurred vision, weight gain, uri- nary retention	Avoid in patients with cardi- ac disease, glaucoma, o seizure disorder; avoid concomitant use of tra- madol
Morphine and oxy- codone	Morphine, 90 mg/day; oxycodone, 45 mg/day	5–15 mg every 4 hr as needed	After 1–2 wk, convert total daily dose to long-acting opioid and continue short-acting formulation as rescue medication	Morphine, 2.8 (2.0–4.6) ²⁰ ; oxycodone, 2.5 (1.7–4.4) ²⁰	Nausea, vomiting, constipa- tion, drowsiness, dizzi- ness, mood change, dis- orientation	There is risk of abuse and uncertainty over long-term effectiveness and safety§
Tramadol	298 mg/day	50 mg every 4–6 hr	Increase by 50–100 mg/day in divided doses every 3–7 days as tolerated, to maximum dose of 400 mg/day (300 mg/day in patients >75 yr of age)	4.8 (2.6–27.0) ²⁰	Nausea, vomiting, constipa- tion, drowsiness, dizzi- ness, seizures	Same as with morphine and oxycodone; also, avoid concomitant use of SSRIs, SSNRIs, tricyclic antidepressants

^{*} Data are primarily from Hempenstall et al.20 and Dworkin et al.21 NA denotes not available, SSNRIs selective serotonin- and norepinephrine-reuptake inhibitors, and SSRIs selective serotonin-reuptake inhibitors.

[†] This is the number needed to treat for one person to have at least 50% pain relief.

[±] Systemic adverse events include diarrhea, nausea, vomiting, fatigue, infections, musculoskeletal disorders, hypertension, dizziness, and headache.

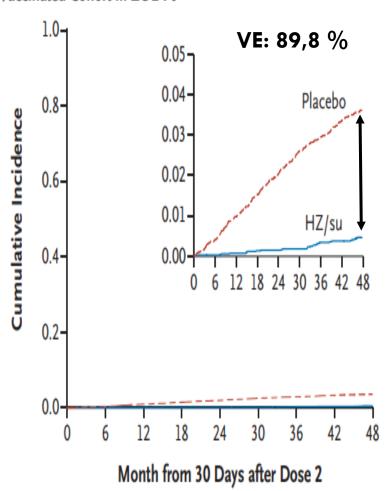
§ See also national guidelines on opioid use for chronic pain.^{24,25}

HZ/su (RCT)

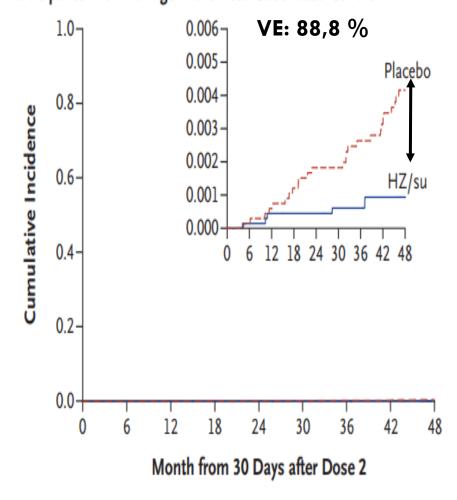
Herpes Zoster Incidence

Post-herpetic Neuralgia





C Participants ≥70 Yr of Age in the Total Vaccinated Cohort



HZ/su VE against HZ complications

- □ ZOE-50 & ZOE-70
- Non-PHN complications
 - Ophtalmic disease
 - Disseminated disease
 - Neurologic disease
 - HZ vasculitis
- □ **VE** ≥ **50 y. 93.7** % (95% CI 59.5-99.9 %)
- □ VE ≥ 70 y. 91.6 % (95% CI 43.3-99.8 %)
- 5 HZ related hospitalisations in placebo group
- No HZ related deaths

RZV long-term efficacy

Table 2. Vaccine Efficacy in the ZOE-50 and ZOE-70 Studies and the Current Long-Term Follow-up Study After at Least 2 Additional Years of Follow-up

	Ad	Adjuvanted Recombinant Zoster Vaccine				rical Co	ontrol ^a /Placebo Gr ZOE-70 ^b				
	N	n	Sum of Follow-up Years	Incidence (per 1000 Person-Years)	N	n	Sum of Follow-up Years	Incidence (per 1000 Person-Years)	Vaccine Efficacy, % (98 Confidence Interval)		
Vaccine efficac	y in the curre	nt foll	ow-up study: prim	ary objective (up to th	e data lock	c point	for the interim an	alysis in the current foll	ow-up study)		
Overall ^a	7277	27	19 621.7	1.4	7277	169	19 621.7	8.6	84.0 (75.9-89.8)		
Vaccine efficac	y from 1 mon	th po	st-dose 2: second	ary objective (up to th	e data lock	point	for the interim and	alysis in the current follo	ow-up study)		
Overall	13 881	59	72 744.6	0.8	13 881	651	72 744.6	8.9	90.9 (88.2–93.2)		
Year 1 ^b	13 881	3	13 744.5	0.2	14 035	130	13 823.3	9.4	97.7 (93.1–99.5)		
Year 2 ^b	13 569	10	13 415.6	0.7	13 564	136	13 332.5	10.2	92.7 (86.2–96.6)		
Year 3 ^b	13 185	9	13 016.1	0.7	13 074	116	12 834.0	9.0	92.4 (85.0–96.6)		
Year 4 ^b	12 757	10	12 946.7	0.8	12 517	95	12 637.4	7.5	89.8 (80.3–95.2)		
Year 6 ^a	7277	10	7208.8	1.4	7277	66	7208.8	9.2	84.9 (70.4–93.1)		
Year 7 ^a	7097	10	6993.1	1.4	7097	68	6993.1	9.7	85.3 (71.3–93.3)		
Year 8 a,c	6876	7	5160.2	1.4	6876	44	5160.2	8.5	84.1 (64.4–94.0)		

VZV recommendations



HGR HZ vaccination

recommendation

federale overheidsdienst

VOLKSGEZONDHEID, VEILIGHEID VAN DE VOEDSELKETEN EN LEEFMILIEU



- August 2022
- HGR 9684 Vaccination against Herpes Zoster
- Recommendation:
 - Recombinant HZ subunit vaccine (2 dose regimen) for:
 - Immunocompetent adults aged ≥ 60 years.
 - Patients under immunosuppressive therapy and JAK-inhibitors aged ≥ 16 * years



No. 9158 Vaccination of immunocompromised or chronically ill children and/or adults Chapter 5. LIST OF (POTENTIALLY) IMMUNOSUPPRESSIVE MEDICATIONS

Co-administration
 dTpa is safe**

*: RZV is indicated for prevention of herpes zoster (HZ) and post-herpetic neuralgia (PHN), in:

- adults 50 years of age or older;
- adults 18 years of age or older at increased risk of HZ.

The use of Shingrix should be in accordance with official recommendations. (SPC Shingrix)

**: Shingrix can be given concomitantly with unadjuvanted inactivated seasonal influenza vaccine, 23-valent pneumococcal polysaccharide vaccine (PPV23) or reduced antigen diphtheria-tetanus-acellular pertussis vaccine (dTpa)

(SPC Shingrix)

Costs HZ vaccination (Be)





BCFI.be

Terugbetaling RZV

Het vaccin tegen zona: doeltreffend maar te duur

KCE

25-10-2022

Het Federaal Kenniscentrum voor de Gezondheidszorg (KCE) stelt hierbij zijn resultaten voor: de prijs van dit vaccin is te hoog in vergelijking met het voordeel dat ervan verwacht kan worden. Deze vaccinatie zou bij terugbetaling ongeveer 602 miljoen euro kosten in het eerste jaar, gevolgd door 23 miljoen euro per jaar daarna. Het is de vraag of dergelijke zware uitgaven te verantwoorden zijn voor een ziekte die relatief mild blijft voor de meeste mensen, zeker in een tijd van budgettaire beperkingen.

Varicella zoster vaccine

- HZ burden in older population
- PHN burden in older population
- PHN treatment difficult unsuccesful
- □ Highly effective VZV vaccine

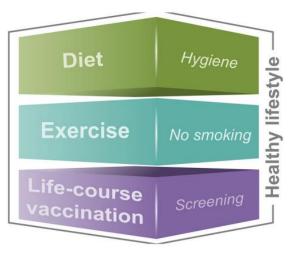


 The older population stays at risk for HZ and its complications in Belgium

Pneumococcal vaccination and COVID-19

Table 2. Probability of a SARS-CoV-2 positive nasopharyngeal swab according to influenza and pneumococcal vaccination status.²⁷

	All			18 to 64 years of age			65 to 104 years of age		
	OR	95% CI	<i>p</i> -Value	OR	95% CI	<i>p</i> -Value	OR	95% CI	<i>p</i> -Value
(a) Not adjusted model									
Flu vaccine during previous autumn	1.02	0.91-1.15	0.7387	0.86	0.75-0.99	0.0301	0.83	0.60-1.14	0.2440
Pneumococcal vaccination	0.77	0.58-1.02	0.0635	0.67	0.46-0.97	0.0342	0.42	0.26-0.66	0.0002
(b) Adjusted model*									
Flu vaccine during previous autumn	0.89	0.78-1.01	0.1408	0.85	0.74-0.98	0.0235	0.87	0.59-1.28	0.4826
Pneumococcal vaccination in previous 12 months	0.56	0.41-0.75	0.0001	0.61	0.41-0.91	0.0156	0.56	0.33-0.95	0.0313

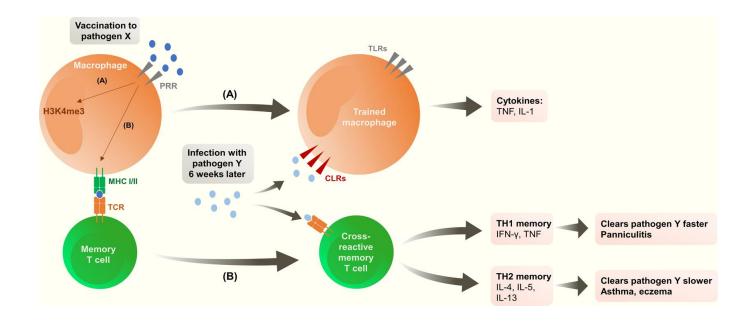




HUMAN VACCINES & IMMUNOTHERAPEUTICS https://doi.org/10.1080/21645515.2021.1957647

"Immune fitness"

Immune fitness



Exercise and pneumococcal immunity

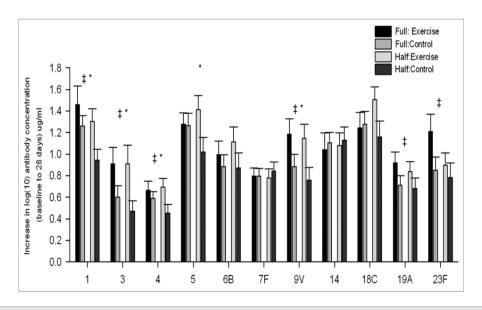
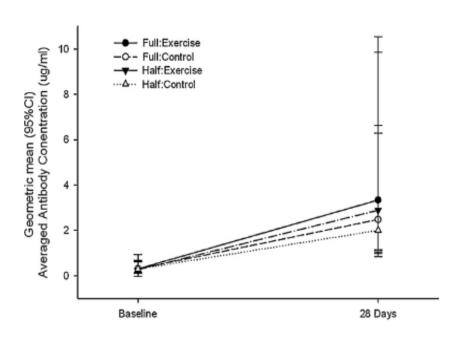
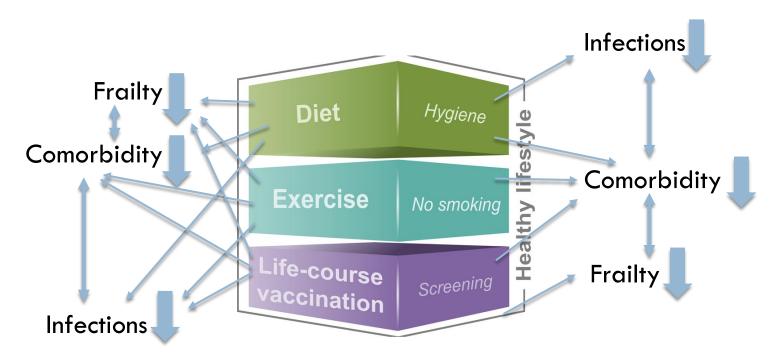


Figure 1. Change scores for pneumococcal strains from baseline to 28 d for all groups. ‡, significant effect of exercise over both full and half dose; *significant effect of exercise within half dose groups.



Vaccine 30 (2012) 6389–6395 Human Vaccines & Immunotherapeutics 2013 9:4, 907-910,

THOM



npj Vaccines (2021) 6:93; https://doi.org/10.1038/s41541-021-00354-z

Overview

- Coronavirus in the older prerson
- Influenza in the older person
- Pneumococcal disease in the older person
- Frailty in older persons
 - Immunosenescence
- Vaccination strategies for older persons
 - Age based vaccination
 - Risk based vaccination
 - Herd immunity and vaccination
- Immunosenescence and vaccination
- Ex. Herpes Zoster vaccination
- Addendum: future vaccines for older persons

RSV disease burden



RSV is a major cause of severe respiratory disease in older adults and adults with high-risk medical conditions^{1,2}



RSV reinfections occur frequently in adults,³ and RSV carries an annual disease burden similar to or greater than influenza in those infected⁴

60+ years old

Unmet need



Despite the high RSV disease burden, no licensed vaccine is available for prevention of RSV



Immunity to RSV following natural infection wanes quickly and re-exposure occurs frequently; thus, an effective vaccine needs to induce a durable immune response

Annual RSV disease burden in the US^{5,6}

64 M

ARIs

in adults and children

177 K

Hospitalizations in adults ≥65 years

14 K

Deaths

in adults ≥65 years

ARI, acute respiratory infection; K, thousand; M, million; RSV, respiratory syncytial virus; US, United States.

Confidential do not share

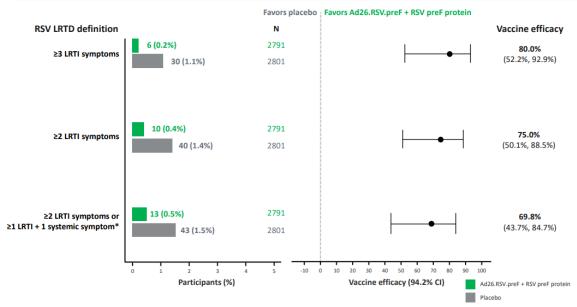
1. Nam HH, Ison MG. *BMJ*. 2019;366:I5021. 2. Walsh EE, et al. *J Infect Dis*. 2004;189(2):233-8. 3. Berbers G, et al. *J Infect Dis*. 2021;224(2):269-278. 4. Falsey AR, et al. *Open Forum Infect Dis*. 2021;8(11):ofab491. 5. National Institute of Allergy and Infectious Disease. 2008. (link). 6. CDC. Accessed July 2022. (link).

RSV vaccine - Janssen

Ad26.RSV.preF + RSV preF protein showed

PRE-READ

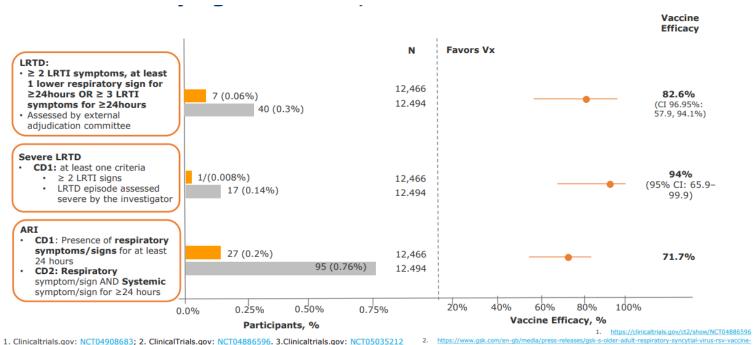
high vaccine efficacy against RSV LRTD in the first RSV season¹



^{*}This definition of RSV LRTD captured all RSV ARIs observed in the study in the first season

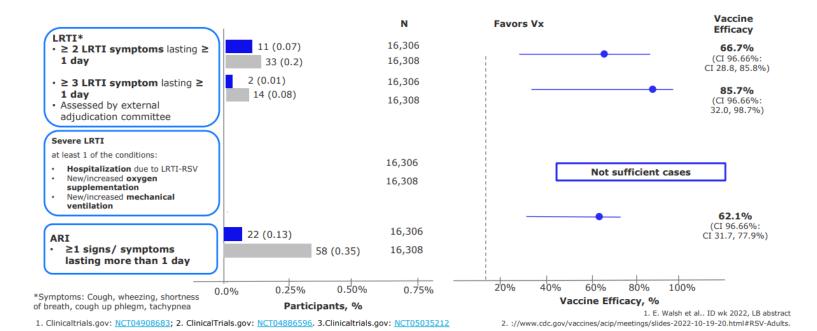
VE, vaccine efficacy; CI, confidence Interval

RSV vaccine - GSK

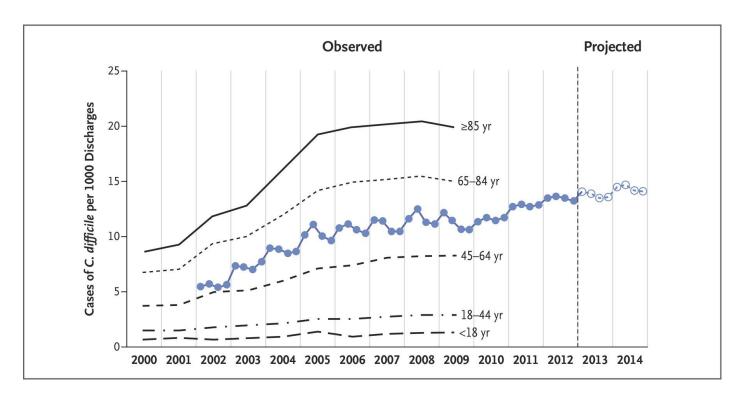


2. https://www.gsk.com/en-gb/media/press-release gsk-s-older-adult-respiratory-syncytial-virus-rsv-vaccine-

RSV vaccine - Pfizer



C. Difficile



Clostridium vaccine trial – CLOVER (Pfizer)

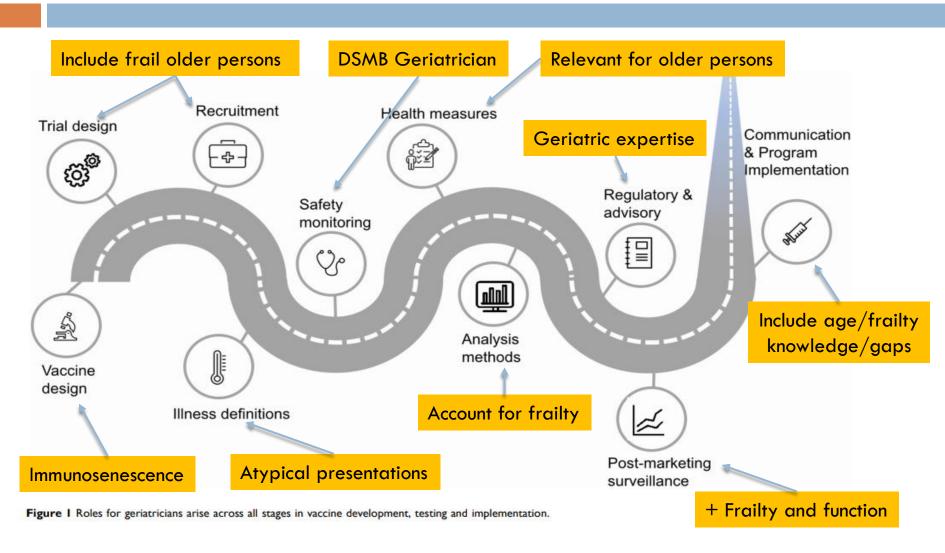
- □ 3 doses
- Primary efficacy endpoint: prevention of primaryCDI
 - □ 31 % (95 % CI: -38,7 66,6) after 3 doses
 - □ 28,6 % (95 % CI -28,4 61) after 2 doses
- Secondary endpoint:
 - No CDI with medical attention in vaccine group

Overview

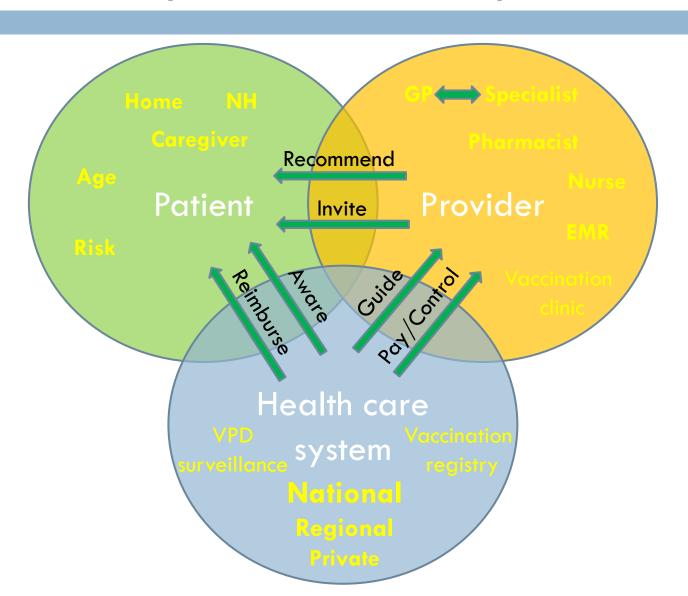
Conclusions

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Vaccine development: include older persons!



Multicomponent interventions to improve vaccine uptake





Vaccinate!

To celebrate!

Tiende editie Rimpelrock lokt 30.000 bezoekers, GVA