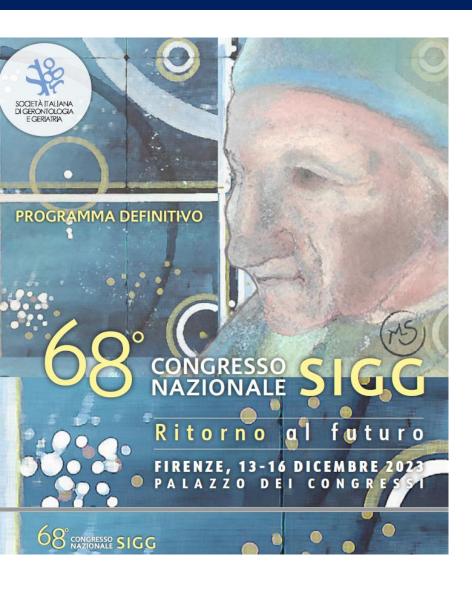


Renzo ROZZINI

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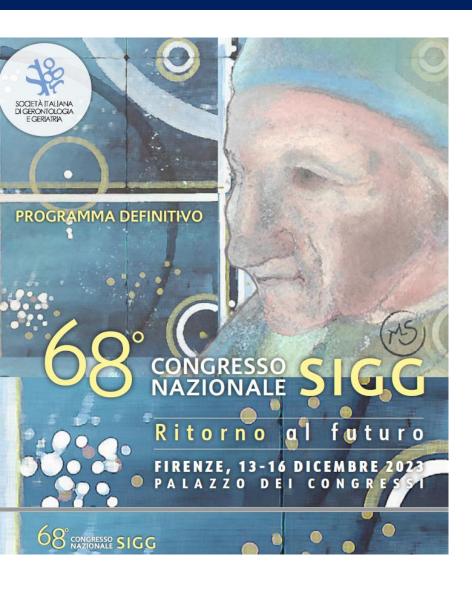
Introduzione: perché un geriatra deve occuparsi di vaccinazioni?

- Elevata suscettibilità alla infezioni (prevalenza)
- Mortalità
- Funzione
- Cognitività
- Uso dei servizi

Malattie prevenibili con la vaccinazione (Polmonite, COVID-19, HZ, Flu, RSV)

Benefici della vaccinazione

Cause di mancata vaccinazione: cosa si può fare (prospettive)



Introduzione: perché un geriatra deve occuparsi di vaccinazioni?

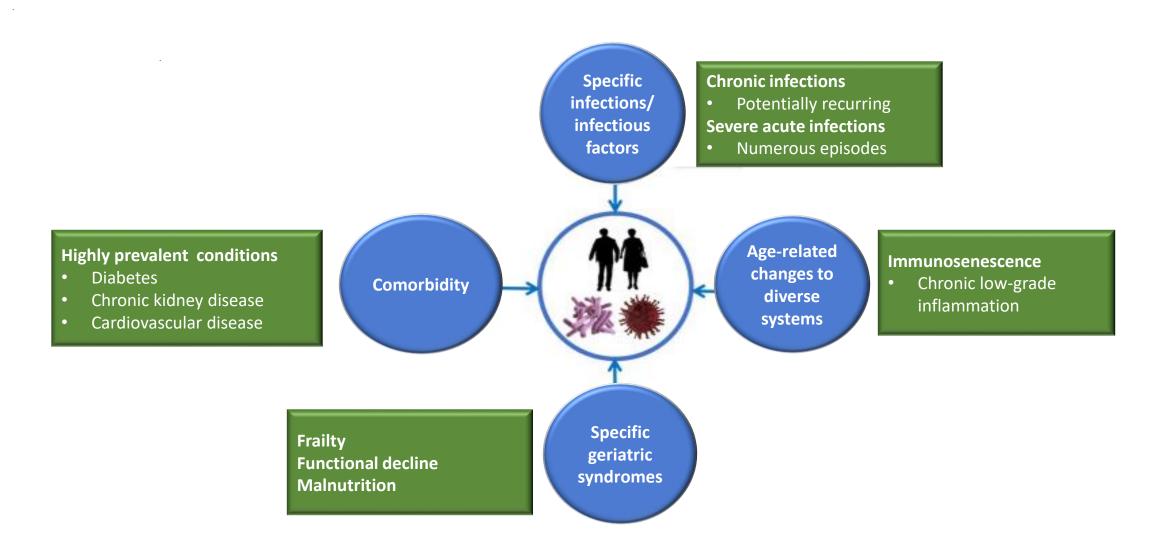
- Elevata suscettibilità alla infezioni (prevalenza)
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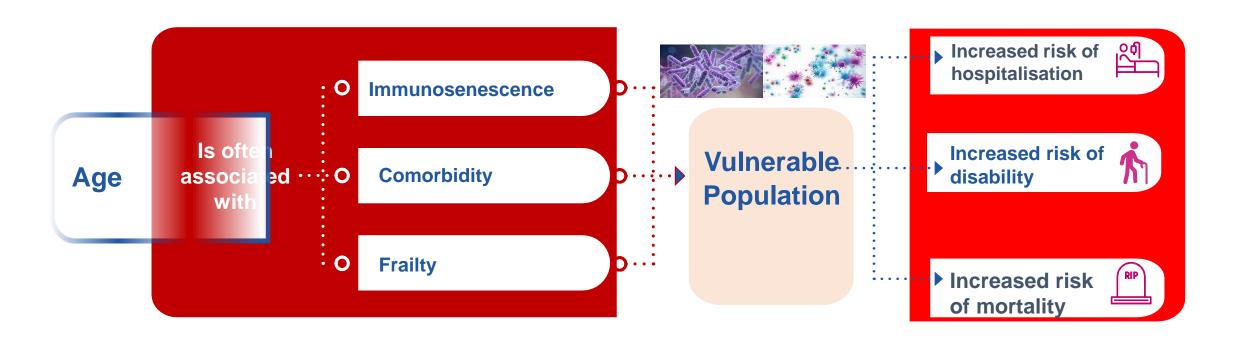
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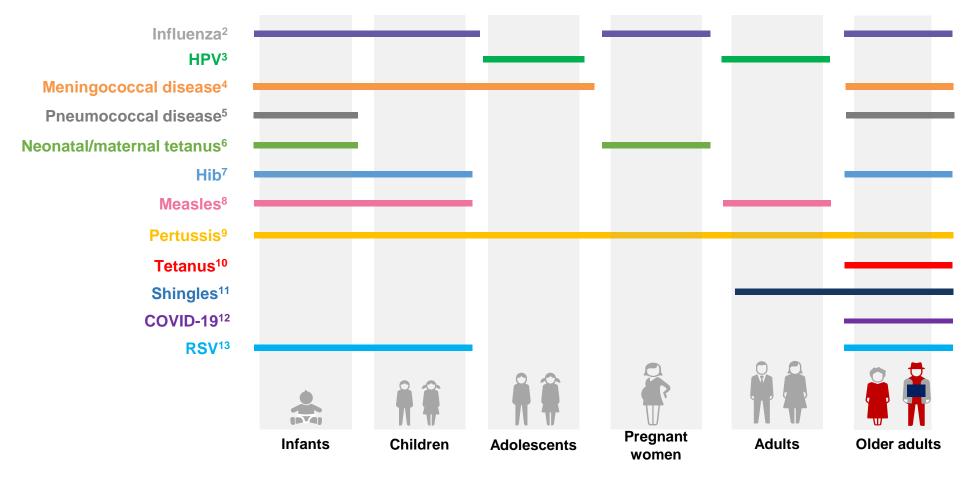
Older adults are more susceptible to infectious diseases and to a potential weakened immune system



Older adults are more vulnerable to vaccine-preventable diseases (VPDs) than younger populations*



Vaccine-Preventable Diseases-VPDs that commonly affect different age groups



^{*}People can be susceptible to any of these infections at any age – this illustration is only meant to show where the greatest burden of disease lies CDC, Centers for Disease Control and Prevention; COVID-19, Coronavirus Disease 2019; Hib, Haemophilus Influenza Type B; HPV, human papillomavirus; RSV, respiratory syncytial virus; VPD, vaccine-preventable disease; WHO, World Health Organisation

^{1.} Rappuoli R et al. Nat Rev Immunol 2011;11:865–872; 2. CDC, 2022. Key facts about influenza (flu). https://www.cdc.gov/flu/about/keyfacts.htm; 3. CDC. 2022. Vaccine Pink Book. https://www.cdc.gov/vaccines/pubs/pinkbook/hpv.html; 4. CDC. 2023. Meningococcal disease. Risk Factors. https://www.cdc.gov/meningococcal/about/risk-factors.html; 5. CDC. 2022. Pneumococcal disease. https://www.cdc.gov/pneumococcal/about/risk-transmission.html; 6. WHO. 2023. Tetanus. https://www.who.int/news-room/fact-sheets/detail/tetanus; 7. National Vaccine Information Center (NVIC). Who is at Highest Risk for Getting Hib? 2022. https://www.nvic.org/disease-vaccine/hib/hib-highest-risk; 8. CDC. 2021. Measles. Pink book. https://www.cdc.gov/vaccines/pubs/pinkbook/meas.html; 9. CDC, 2022. Pertussis. https://www.cdc.gov/vaccines/pubs/pinkbook/pert.html; 10. CDC. 2022. Tetanus. https://www.cdc.gov/tetanus/clinicians.html#risk-groups; 11. CDC. 2021. Zoster. In: Epidemiology and Prevention of Vaccine-Preventable Diseases Pink Book, Hall E et al. (Eds). 14th edn. Washington, DC: Public Health Foundation, 2021. pp. 349–358; 12. WHO. 2021. Coronavirus. https://www.who.int/health-topics/coronavirus#tab=tab_1; 13. CDC. 2022. Key facts about RSV. https://www.cdc.gov/rsv/index.html. URLs accessed October 2023



Impact of pre-admission functional status on in-hospital mortality of COVID-19 older patients – a cohort study

Table 2Cox Multivariable Regression Analysis of Determinants of In-Hospital Mortality

Model 1: Clinical and Laboratory Variables			Model 2				
Variables	HR	95% CI	P	Variables	HR	95% CI	P
Barthel Index, (\geq 75 vs < 75)	0.383	0.24-0.62	<.001	COVID-19 MRS, (for unitary increase)	1.49	1.33-1.69	<.001
Age (per year increase)	1.06	1.01-1.11	.015				
Dementia (no vs yes)	0.52	0.31-0.88	.015	Barthel Index (\geq 75 vs < 75)	0.35	0.22-0.57	<.001
RR (per breaths/min increase)	1.06	1.02-1.09	<.001	Frailty (no vs yes)	0.60	0.39-0.94	.024
Pao ₂ /Fio ₂ (per unit increase)	0.995	0.994-0.999	.019				
Creatinine (per mg/dL increase)	1.20	1.04-1.39	.012				
Platelets (109/L per unit increase)	0.997	0.992-0.998	.003				

CI, confidence interval; HR, hazard ratio; RR, respiratory rate.

Variables excluded (P > .10) from Model 1: frailty, number of drugs, C-reactive protein, and number of comorbidities.

<u>Frailty</u> was assessed based on the modified Frailty Index (mFI) created by Saxton and Velanovich by mapping 11 variables (nonindependent functional status, history of diabetes mellitus, chronic obstructive pulmonary disease or pneu- monia, heart failure, myocardial infarction, angina or coronary revascularization, hypertension, peripheral vascular disease, presence of impaired sensorium, TIA or cerebrovascular event without or with deficit) present in the Canadian Study of Health and Aging Frailty Index.



Table 2—Three-Month Mortality Risk in 2,948 Elderly Hospitalized Patients

Characteristic	Events/No. of Patients	RR*	95% CI	RR^b	95% CI
Age ≥90 y	58/269	2.5	1.8-3.4	1.7	1.2-2.4
Male	144/1,070	1.5	1.2-1.8		
Smoker	103/857	1.2	1.0-1.5	1.2	1.0-1.8
Disabled 2 wk before admission	145/642	3.5	2.7-4.4	1.6	1.2-2.2
APACHE II-APS subscore > 12	55/138	6.2	4.3-8.9	1.7	1.2-2.6
Urea/creatinine ratio > 60	110/770	1.7	1.3-2.2	1.4	1.1-1.9
Pneumonia	46/240	2.1	1.5-2.9	1.4	0.9 - 2.0
Delirium	71/322	2.6	1.9-3.5	1.5	1.1-2.1
Dementia	122/554	3.6	2.8-4.6	1.9	1.4-2.7
COPD	88/608	1.5	1.2-1.9	2.4	1.1-5.8
Renal failure	70/447	1.7	1.2-2.2		
Malnutrition	70/227	4.4	3.2-5.9	1.9	1.3-2.6
Stroke	65/362	1.9	1.4-2.6		
Metastatic cancer	50/173	3.7	2.6-5.3	1.2	1.1-1.3
Charlson index ≥5	217/1,067	4.2	3.3-5.4	1.8	1.3-2.5
Drugs ≥7	133/621	2.5	1.9-3.3	1.6	1.2-2.1

Cox proportional hazard ratio was used to model the time-of-death data to identify possible predictors of mortality. Variables significantly associated with mortality in crude analysis were included as potential confounders in a final Cox proportional hazards regression model, with CAP admission status as the main predictor and time to death as the outcome. RR = relative risk. See Table 1 for expansion of other abbreviations.

*Crude analysis.

Rozzini & Trabucchi, 2003, 2011

Is Pneumonia Still the Old Man's Friend?

Table 1. Characteristics and 6-Month Mortality Rate of 1803 Inpatients Consecutively Admitted in a Geriatric Ward for Pneumonia or Other Acute Noninfectious Diseases*

Characteristic	Pneumonia (n = 241)	Acute Noninfectious Diseases (n = 1562)	<i>P</i> Value
Age, y	83.3 ± 6.9	79.7 ± 7.0	.001
Male, %	24.5	19.3	.001
MMSE score	19.7 ± 9.1	22.9 ± 7.1	.001
GDS score	5.1 ± 3.2	5.1 ± 3.6	.98
Barthel Index (15 days before admission)	72.6 ± 31.5	83.8 ± 23.2	.001
Barthel Index (on admission)	55.3 ± 37.9	74.5 ± 30.0	.001
IADL (functions lost)	3.9 ± 3.0	3.1 ± 2.8	.001
Diseases, No.	6.1 ± 2.1	5.3 ± 2.0	.001
Charlson Index	8.3 ± 2.5	7.0 ± 2.6	.001
Drugs, No.	4.5 ± 2.3	4.3 ± 1.9	.19
APACHE II score	13.3 ± 6.3	7.9 ± 4.1	.001
APS-APACHE II subscore	3.8 ± 4.2	1.9 ± 2.7	.001
Serum albumin, g/dL	3.6 ± 1.3	3.9 ± 0.6	.001
Hemoglobin, g/dL	11.7 ± 2.3	12.3 ± 2.0	.02
Serum cholesterol, mg/dL	186.2 ± 51.9	204.7 ± 51.1	.001
CRP, mg/dL	7.5 ± 5.6	2.6 ± 7.8	.001
Creatinine, mg/dL	1.2 ± 0.8	1.1 ± 0.8	.20
Length of stay, d	8.1 ± 5.1	6.4 ± 3.3	.001
6-mo mortality, %	27.4	20	.001

Abbreviations: APACHE, Acute Physiology and Chronic Health Examination; APS, Acute Physiology Score; CRP, C-reactive protein; GDS, Geriatric Depression Scale; IADL, Instrumental Activities of Daily Living; MMSE, Mini-Mental State Examination.

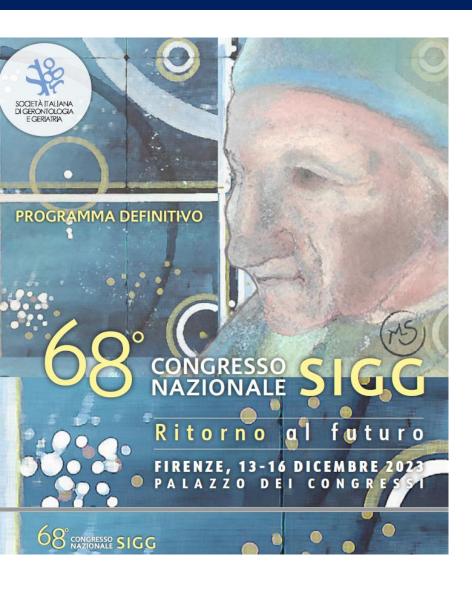
SI conversion factors: To convert cholesterol to millimoles per liter, multiply by 0.0259. To convert creatinine to micromoles per liter, multiply by 88.4.

*Data are mean ± SD value unless otherwise specified.

†P value derived from χ^2 test. Other P values were derived from the t test.



^bFinal Cox proportional hazards regression model.



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Cause di mancata vaccinazione: cosa si può fare (prospettive)

THE PRINCIPLES AND

PRACTICE OF MEDICINE

DESIGNED FOR THE USE OF PRACTITIONERS AND STUDENTS OF MEDICINE

11.30

WILLIAM OSLER, M. D.

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NEW YORK
D. APPLETON AND COMPANY
1892

Rijkers and Pelton Pneumonia (2018) 10:8 https://doi.org/10.1186/s41479-018-0052-7

EDITORIAL

The old man's friend

Ger T. Rijkers^{1*} and Stephen I. Pelton²

Editorial

The term "old man's friend" is often used when referring to pneumonia. Searching for it on Google yields 16,400 results in 0.33 s for this combination. The term is attributed to William Osler, who in the first edition of his book *The Principles and Practice of Medicine* (1892) wrote:

In children and in healthy adults the outlook is good. In the debilitated, in drunkards and in the aged the chances are against recovery. So fatal is it in the latter class [i.e. the elderly] that it has been termed the natural end of the old man [1].

Etiology.—Pneumonia is one of the most wide-spread of acute diseases. Hospital statistics show that the ratio to other admissions is in the proportion of twenty to thirty per thousand.

It prevails at all ages Children are quite as susceptible to it as adults, and it is the special enemy of old age. Males are more frequently affected than females. Dwellers in cities and persons whose occupations are associated with exposure, hardship, and cold are most liable to the disease. Contrary to the general rule in infectious diseases, newcomers and immigrants seem less susceptible than the native inhabitants. Debilitating causes of all sorts render individuals more susceptible. Alcoholism is perhaps the most potent predisposing factor. Persons weakened by disease are especially prone to it; thus we find many cases in connection with chronic Bright's disease, diabetes, the chronic affections of the nervous system, and protracted fevers. One important predisposing cause is a previous attack. No acute disease recurs with such frequency. Instances are on record of individuals who have had ten or more attacks.

INFLUENZA

FREQUENTLY COMPLICATED WIT

PNEUMONIA

THIS THEATRE IS CO-OPERATING WITH THE DEPARTMENT OF HEALTH

YOU MUST DO THE SAME

IF YOU HAVE A COLD AND ARE COUGHING AND SNEEZING DO NOT ENTER THIS THEATRE

GO HOME AND GO TO BED UNTIL YOU ARE WELL

Coughing, Sneezing or Spitting Will Not Be Permitted In The Theatre. In case you must cough or Sneeze. do so in your own handkerchief, and if the Coughing or Sneezing Persists Leave The Theatre At Once.

This Theatre has agreed to cooperate with the Department Of Health in disseminating the truth about Influenza. and thus serve a great educational purpose.

HELP US TO KEEP CHICAGO THE HEALTHIEST CITY IN THE WORLD

JOHN DILL ROBERTSON

The consequences of CAP in adults can be severe, particularly in those with comorbid conditions

In a prospective, population-based, cohort study among adults 18+ years of age hospitalised for CAP in Louisville, KY*



A total of **8284 hospitalisations** were due to CAP during the 2-year study period

~9%



were re-hospitalised for CAP due to a new episode during the same study year



Of the **3789 adults** hospitalised due to CAP during the first year of the study:

13%



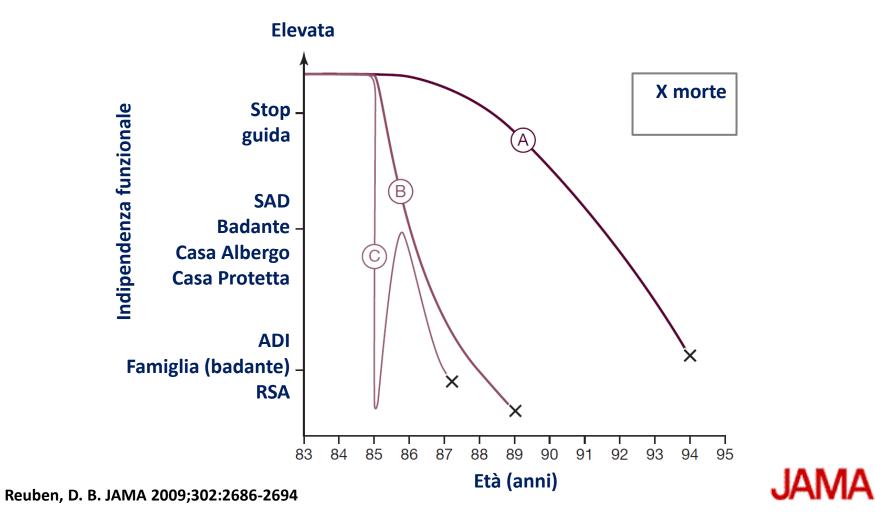
died within 30 days after hospitalisation due to CAP

~31%



died within 1 year after hospitalisation due to CAP

Traiettorie ipotetiche di declino funzionale per il signor R.



Traiettorie ipotetiche di declino funzionale per il signor R. <u>Traiettoria A</u>: stato di salute buono; buona spettanza di vita. <u>Traiettoria B</u>: il paziente ha una malattia cronica degenerativa (ad es., malattia di Alzheimer, malattia di Parkinson): declino funzionale costante con un periodo di dipendenza funzionale prolungato. <u>Traiettoria C</u>: evento catastrofico improvviso (ad es., frattura di femore, stroke, sepsi, polmonite) con qualche miglioramento funzionale, ma senza tornare allo stato di partenza, riduzione della spettanza di vita.

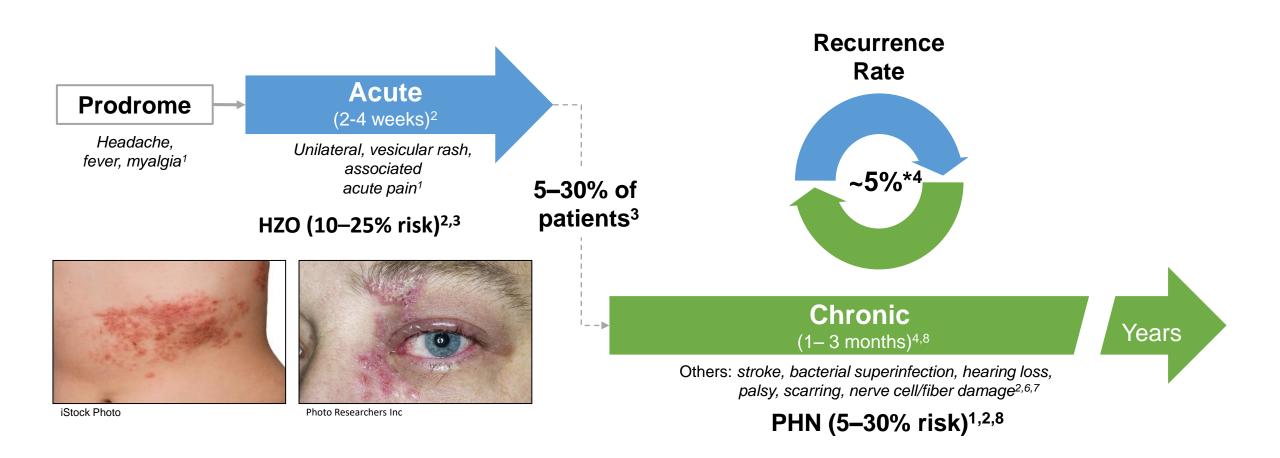
The consequence of the COVID-19 pandemic on other respiratory illnesses and potential future impact

Co-infection

- Co-infection between SARS-CoV-2 and other viruses such as influenza or RSV may result in worse outcomes compared with SARS-CoV-2 only^{1,2}
- Compared with SARS-CoV-2 alone, SARS-CoV-2/influenza co-infection was associated with:
 - ▲ Increased risk of invasive mechanical ventilation^{1,2}
 - ▲ Increased risk of in-hospital mortality^{1,2}

As social contact returned to pre-pandemic levels, we have seen a resurgence in influenza during winter 2022–2023 to levels higher than before the COVID-19 pandemic³

...Even if not associated with increased mortality, HZ acute phase can be followed by chronic complications that deeply affect quality of life¹

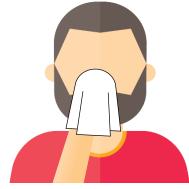


^{*}Over ~8 years follow-up
HZ, herpes zoster; HZO, herpes zoster ophthalmicus; PHN, postherpetic neuralgia
1. de Oliveira Gomes J et al. Cochrane Database Syst Rev 2023;10:CD008858; 2. Harpaz R et al. MMWR Recomm Rep 2008;57:1–30; 3. Kawai K et al. BMJ Open 2014;4:e004883; 4. Yawn BP et al. Mayo Clin Proc 2011;86:88–93; 5. Dworkin RH et al. J
Pain 2008;9:S37–44; 6. Dworkin RH et al. Clin Infect Dis 2007;44:S1–26; 7. Nagel MA and Gilden D. Curr Neurol Neurosci Rep 2015;15:16; 8. Opstelten W et al. Fam Pract 2002;19:471–475

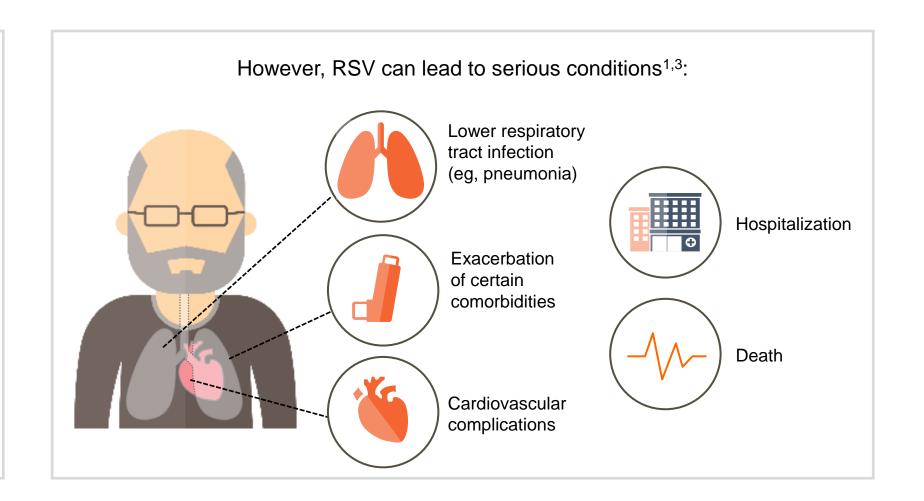
Signs and symptoms of RSV infection in adults

RSV infection is typically mild, but may lead to serious complications and poor outcomes

Typically, RSV infection in healthy adults results in mild, cold-like symptoms^{1,2}



Symptoms can be similar to those of other respiratory pathogens^{2,3}

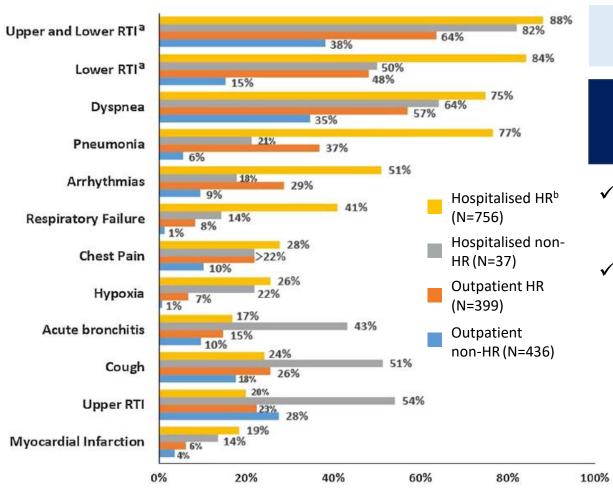


^{1.} Centers for Disease Control and Prevention (CDC), 2023. RSV in older adults and adults with chronic medical conditions. https://www.cdc.gov/rsv/high-risk/older-adults.html (accessed June 2023); 2. Nam HH and Ison MG. BMJ 2019;366:I5021;

3. Branche AR, Falsey AR. *Drugs Aging* 2015;32:261–269

Complications during follow-up period in hospitalised patients with RSV¹





Real-world observational study conducted using the US Medicare database from Jan 2011–Dec 2015

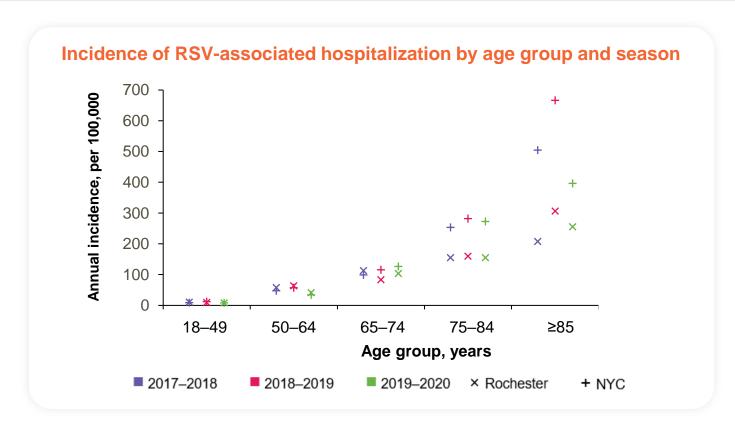
High risk patients hospitalised with RSV diagnosis (N=756)

- ✓ Significant increase in healthcare utilisation following hospitalisation
- ✓ Healthcare cost increased by \$9,210 per patient post-RSV diagnosis, mainly due to the higher rates of hospitalisation and longer LOS due to the exacerbation of exsisting comorbidities

Country-specific information may not be transferable to other countries

Risk of RSV-associated hospitalization increases with age and chronic medical conditions RSV hospitalization rate is high among older adults and those with certain chronic medical conditions

A large prospective study estimated incidence of RSV-associated hospitalization in two regions of New York State, USA, 2017–2020. N=1099 cases



Hospitalization rates for RSV were higher in adults* with underlying conditions

Comorbidity	Incidence rate ratio [†]
Asthma	2.0–3.6
CAD	3.7–7.0
Diabetes	2.4–11.4
COPD	3.2–13.4
CHF	4.0–33.2

^{*}Adults aged ≥18 years. †Ratio of rate among people with each comorbidity vs those without it, in the surveillance area population

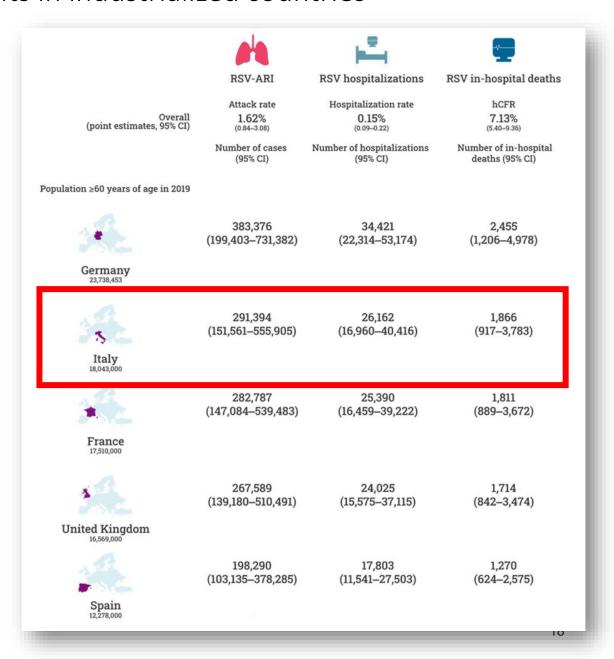
Graph and table were independently created for GSK from the original data

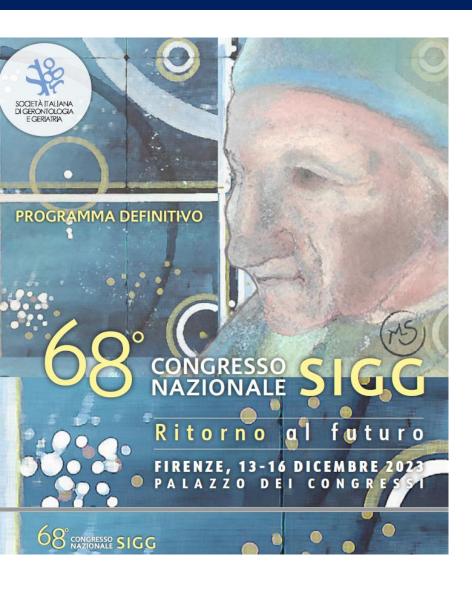
Annual burden of RSV estimates in older adults in industrialized countries

RSV is prevalent but under-recognized in older adults

Estimated cases, hospitalizations, and in-hospital deaths due to RSV-associated acute respiratory infections among adults aged 60 years and older per region, 2019 population.^a

a. Population data obtained from the United Nations Department of Economic and Social Affairs. ARI, acute respiratory infection; CI, confidence interval; hCFR, in-hospital case fatality rate; RSV, respiratory syncytial virus.





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Cause di mancata vaccinazione: cosa si può fare (prospettive)

Vaccination can provide benefits beyond acute disease prevention, contributing to better general health and healthy ageing

Infectious diseases can have health impacts **beyond the acute illness**, potentially with long-term damaging complications^{1–3}

Vaccination can help to maintain overall health by preventing not only initial infections but also the associated downstream effects^{4–7}

As global life expectancy increases, vaccination is becoming more important in supporting the health and **quality of life** of the ageing population⁸

COVID-19 vaccines* were estimated to have saved nearly 20 million lives globally in the first year of vaccination (Dec 2020–Dec 2021)

Global impact of the first year of COVID-19 vaccination:

THE LANCET

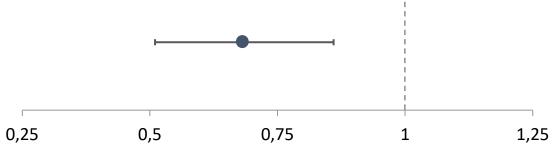
Oliver J Watson*, Gregory Barnsley*, Jaspreet Toor, Alexandra B Hogan, Peter Winskill, Azra C Ghani

Based on official reported COVID-19 deaths, we estimated that vaccinations prevented...14.4 million...deaths from COVID-19. When considering excess death, the estimate rose to 19.8 million, representing a global reduction of 63% in total deaths† during the first year of COVID-19 vaccination.

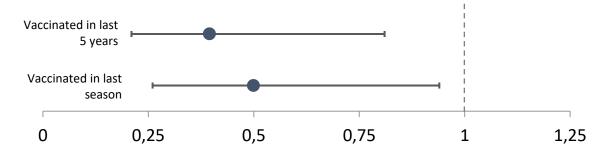
Downstream effects have been observed following vaccination

Risk of major adverse cardiovascular event in influenza-vaccinated subjects with acute coronary syndrome aged >50 years (n=221) vs unvaccinated (n=218)¹









Odds ratio (95% CI)[‡]

Patients in the influenza vaccination group had significantly lower rates of major adverse cardiovascular events than the control group (9.5% vs 19.3%)

Adjusted hazard ratio (95% CI)[†]

Adjusted HR 0.67 (95% CI 0.51–0.86); P=0.005⁺¹

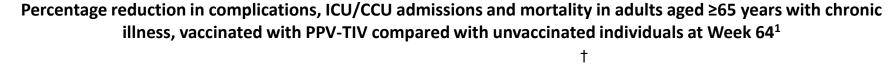
Patients with brain infarctions were less likely to be vaccinated against influenza infection than controls²

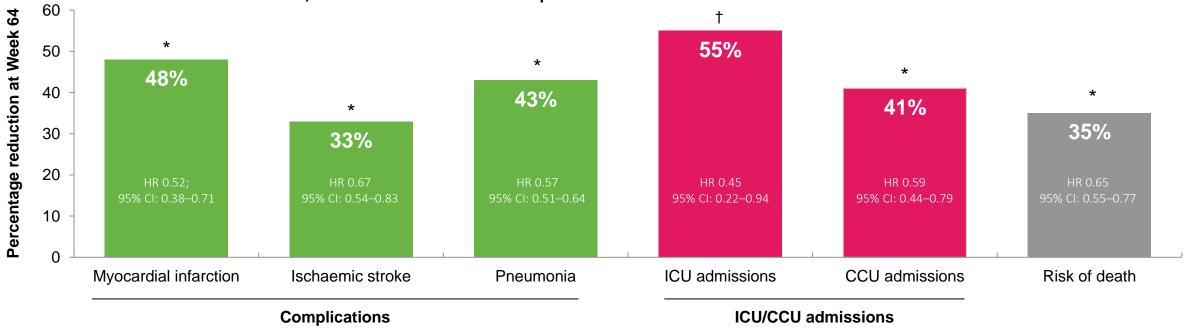
^{*}Vaccinated in last season: controls n=107/180, cases n=42/90. Vaccinated in last 5 years: controls n=101/180, cases 37/90. †Adjusted for age, sex, serum creatinine, medication and coronary revascularisation; †Adjusted for age, sex, diabetes, hypertension, body mass index, current smoking, cholesterol and use of antibiotics in the last 3 months

CI. confidence interval

^{1.} Phrommintikul A et al. Eur Heart J 2011;32:1730–1735; 2. Lavallée P et al. Stroke 2002;33:513–518

Vaccination against respiratory diseases can help protect elderly persons with chronic illness against cardiovascular and cerebrovascular events, reducing overall healthcare burden¹

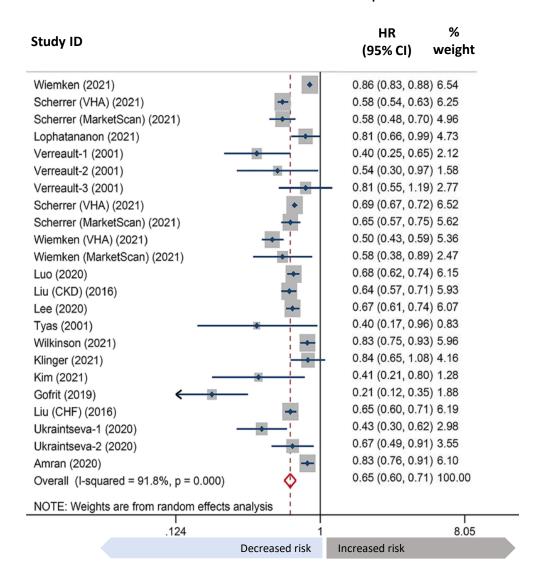




^{*}P<0.001; [†]P=0.03 CCU, coronary care unit; CI, confidence interval; HR, hazard ratio; ICU, intensive care unit; PPV, 23-valent pneumococcal vaccine; TIV, trivalent influenza vaccine 1. Hung IFN *et al. Clin Infect Dis* 2010;51:1007–1016; 2. Doherty TM *et al. Eur Geriatr Med* 2018;9:289–300

Adult vaccinations and dementia risk

Overall association between adult vaccinations and subsequent dementia risk



Vaccination can help combat AMR

- AMR is one of the top 10 global public health threats, leading to serious and life-threatening infections and death^{1,2}
- By decreasing the number of infectious disease cases, vaccines can reduce the use of antibiotics and the emergence and spread of AMR²



Vaccination against bacteria:²

- Reduces prevalence of pathogen and circulation of antimicrobial-resistant strains
- Potential to eradicate pathogens (eg smallpox)

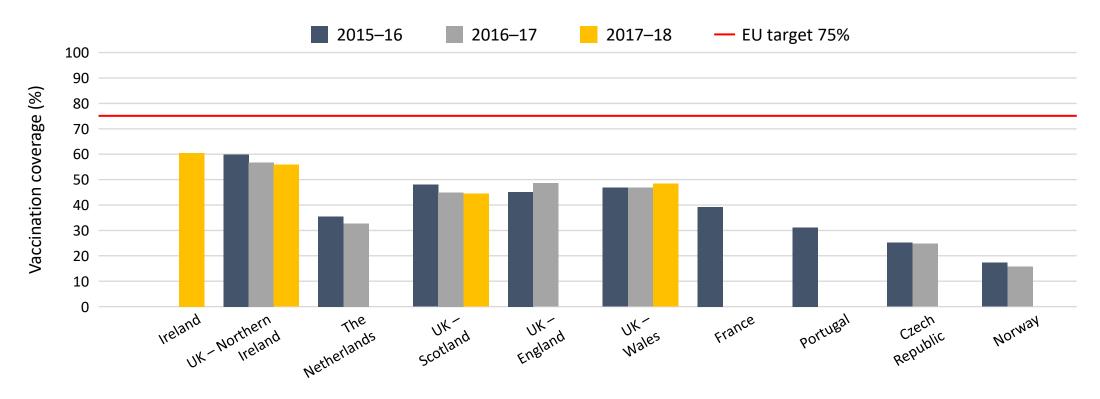


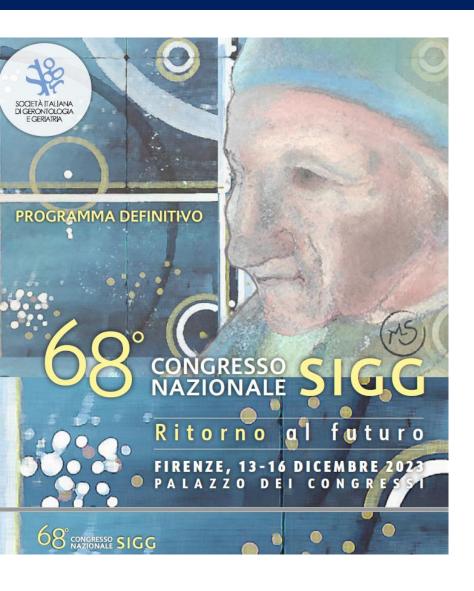
Vaccination against viral infections:²

- Can reduce inappropriate use of antibiotics
- Can prevent secondary bacterial superinfections

Patients with chronic conditions: a sub-optimally vaccinated population

Seasonal influenza vaccination coverage rates among individuals with chronic medical conditions





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Cause di mancata vaccinazione: cosa si può fare (prospettive)

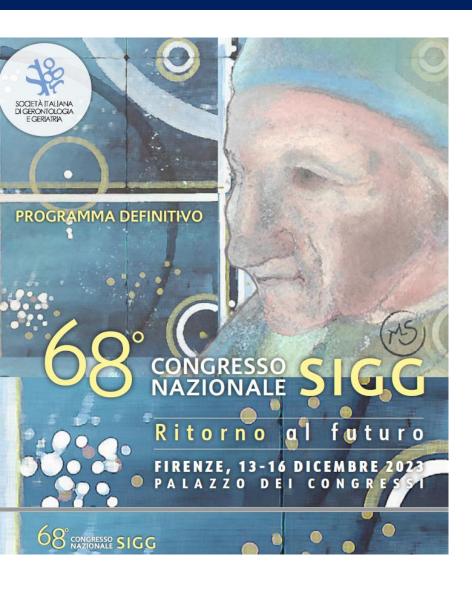
Reasons for under-vaccination in populations with chronic conditions

Lack of awareness of VPDs

Uncertainty about vaccine safety and efficacy (patients, parents and HCP)

Cost/reimbursement

Access and implementation issues



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- -Gli anziani e i pazienti con comorbilità pagano il tributo più pesante alle VPD
- -La prevenzione delle VPD può contribuire a un invecchiamento in buona salute, prolungando la vita senza disabilità e riducendo l'uso delle risorse sanitarie
- -L'adesione alla vaccinazione rimane subottimale negli adulti
- -Interventi per aumentare il tasso di vaccinazioni:
 - Migliorare le conoscenze, l'organizzazione e la formazione degli operatori sanitari
 - Utilizzare ogni forma di tecnologia comunicativa per rivolgersi a segmenti più ampi della popolazione
 - Ricerca di modalità organizzative più efficaci (e loro monitoraggio)
- -È necessaria la politica per stimolare la domanda, sostenere la ricerca e mantenere la vaccinazione come priorità nell'agenda pubblica