

Dr Massimo Di Pietro S.O.C Malattie Infettive I Azienda USL Toscana Centro

18-19-20 gennaio 2021

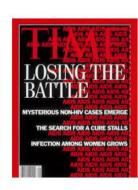
Disclosure statementPersonal fees for consultancy and lectures from Abbvie, Bristol Myers Squibb, Gilead, Janssen, Merck, ViiV.

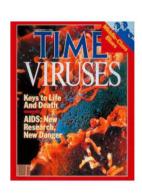










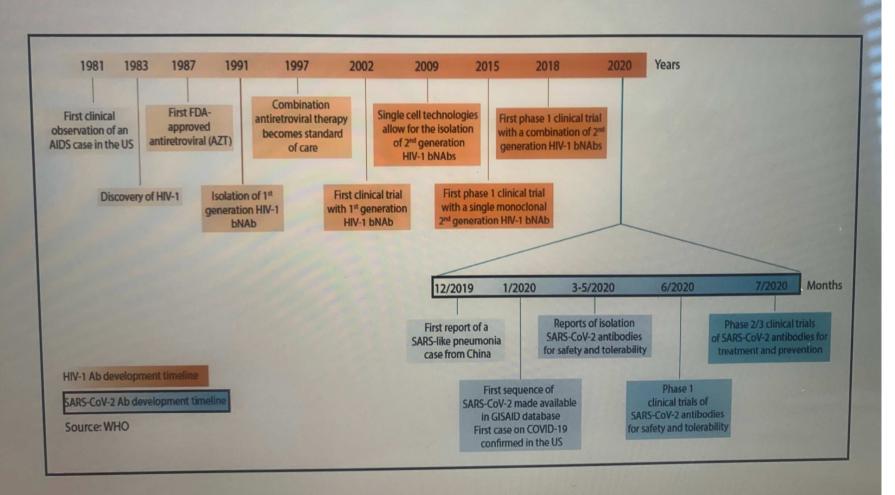






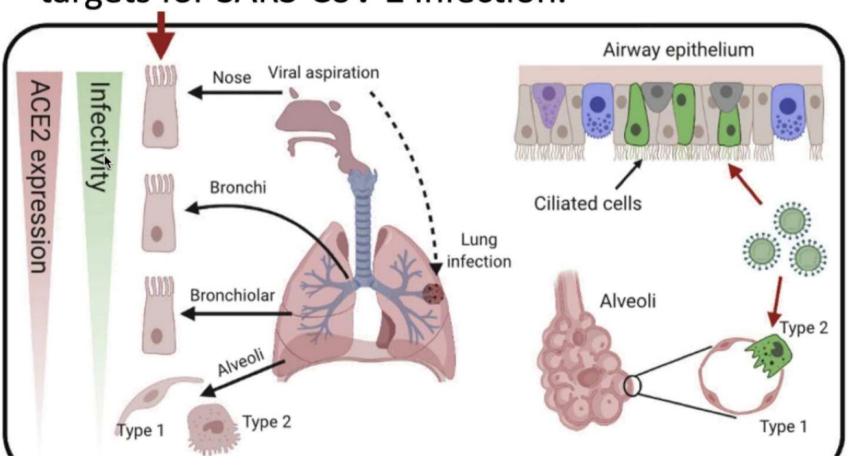
Conversazione in cors

Summary of milestones from first disease clinical manifestations to antibody discovery and development for HIV-1 and SARS-CoV-2



from the proximal to distal respiratory tract

 Ciliated airway cells and AT-2 cells are primary targets for SARS-CoV-2 infection.





Particles (purple; artificially coloured) of SARS-CoV-2 cover the surface of a human cell. Credit: NIAID/NIH/SPL

Table. Summary of SARS-CoV-2 Testing Studies

Cohort	Tested, n	SARS-CoV-2 Positive, n (%)	Positive but Asymptomatic, n (%)	Notes*	
Iceland residents (6)	13 080	100 (0.8)	43 (43.0)	R	
Vo', Italy, residents (7)	5155	102 (2.0)	43 (42.2)	R, L	
Diamond Princess cruise ship passengers and crew (8)	3711	712 (19.2)	331 (46.5)	-	
Boston homeless shelter occupants (9)	408	147 (36.0)	129 (87.8)	-	
New York City obstetric patients (11)	214	33 (15.4)	29 (87.9)	L	
U.S.S. Theodore Roosevelt aircraft carrier crew (12)	4954	856 (17.3)	~500 (58.4)	E	
Japanese citizens evacuated from Wuhan, China (2)	565	13 (2.3)	4 (30.8)	L	
Greek citizens evacuated from the United Kingdom, Spain, and Turkey (14)†	783	40 (5.1)	35 (87.5)	L	
Charles de Gaulle aircraft carrier crew (13)	1760	1046 (59.4)	~500 (47.8)	E	
Los Angeles homeless shelter occupants (10)	178	43 (24.2)	27 (62.8)	-	
King County, Washington, nursing facility residents (15)	76	48 (63.2)	3 (6.3)	L	
Arkansas, North Carolina, Ohio, and Virginia inmates (16)	4693	3277 (69.8)	3146 (96.0)	-	
New Jersey university and hospital employees (17)	829	41 (4.9)	27 (65.9)	_	
Indiana residents (18)	4611	78 (1.7)	35 (44.8)	R	
Argentine cruise ship passengers and crew (19)	217	128 (59.0)	104 (81.3)		
San Francisco residents (29)	4160	74 (1.8)	39 (52.7)	_	

E = estimated from incomplete source data; L = longitudinal data collected; R = representative sample.

* A dash indicates that the study did not have a representative sample, collected no longitudinal data, and did not require estimation of missing data.

† Clarified via e-mail communication with coauthor.

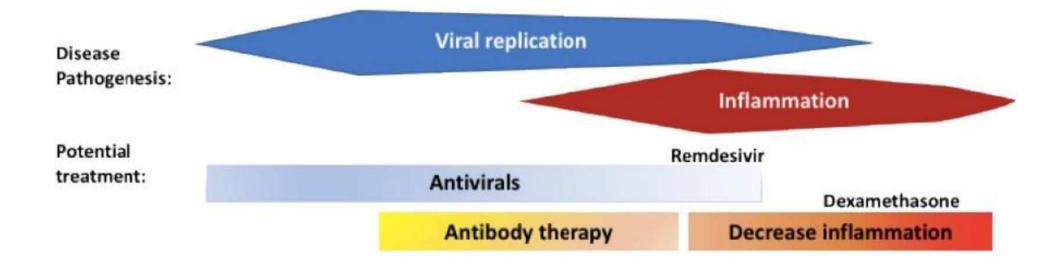
In a COVID-19 outbreak on a cruise ship where nearly all passengers and staff were screened for severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2), approximately 19 percent of the population on board tested positive; 58 percent of the 712 confirmed COVID-19 cases were asymptomatic at the time of diagnosis In studies of subsets of those asymptomatic individuals, who were hospitalized and monitored, approximately 77 to 89 percent remained asymptomatic over time (1-2)

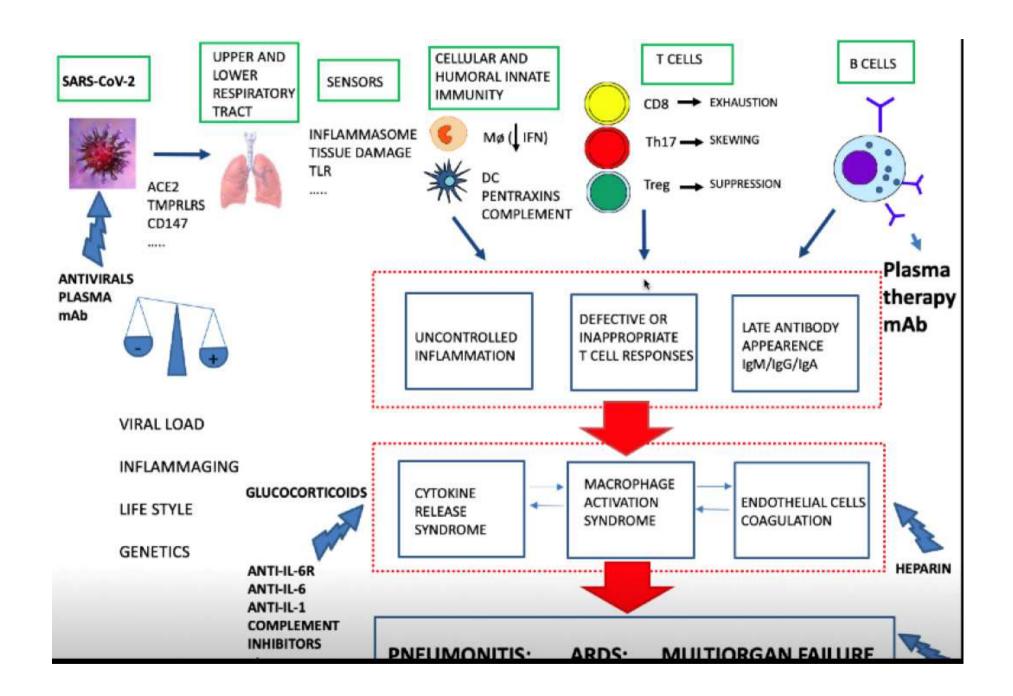
- 1. Japanese National Institute of Infectious Diseases. Field Briefing: Diamond Princess COVID-19 Cases, 20 Feb Update. https://www.niid.go.jp/niid/en/2019-ncov-e/9417-covid-dp-fe-02.html (Accessed on March 01, 2020).
- 2. Lancet Infect Dis.2020 Sep;20(9):1043-1050.doi: 10.1016/S1473-3099(20)30482-5. Epub 2020 Jun 12.

Multidimensional Challenge of Treating COVID-19

Stage/ Severity:

Asymptomatic/	Mild	Moderate	Severe	Critical illness
Presymptomatic	Illness	Illness	Illness	
+ SARS-CoV-2 test but no symptoms	Mild symptoms (eg fever, cough, taste/smell changes); no dyspnea	O ₂ saturation >=94%, lower respiratory tract disease	O₂ saturation <94%, respiratory rate >30/min; lung infiltrates >50%	Respiratory failure, shock, multi-organ dysfunction/failure





Sintomi esordio	✓	Fattori di Rischio – COVID severo				
Febbre		Malattie Cardiovascolari (no ipertensione				
		isolata)				
Tosse (secca)		Diabete (non controllato, A1c >7.5%)				
Dolori muscolari diffusi		BPCO inclusa asma				
Mal di Testa		Alterazioni ematologiche (es. anemia falciforme				
		<u>}</u>				
Mal di Gola		Alterazioni neurologiche				
Rinorrea		Trapiantati organi solidi o in terapie				
		immunosoppressive/biologici				
Congiuntivite		Chemio/immuno terapie per neoplasia				
Difficoltà respiratorie		Trapianto di midollo da < 1 anno				
Vomito		Trattamento Graft versus host				
Diarre		HIV infezione con CD4 < 200 cell				
Perdita/riduzione olfatto						
Perdita/riduzione gusto						
		Fattori di Rischio – Tromboembolici (principali)				
		BMI > 30				
		Anamnesi TVP				
		Neoplasie				
		Trombofilia				

Research

Features of 20 133 UK patients in hospital with covid-19 using the ISARIC WHO Clinical Characterisation Protocol: prospective observational cohort study

BMJ 2020; 369 doi: https://doi.org/10.1136/bmj.m1985 (Published 22 May 2020) Cite this as: BMJ 2020;369:m1985

A study among 20 133 hospitalised patients from acute care hospitals in England, Wales and Scotland identified clustering of symptoms with three common clusters:

- 1.one respiratory symptom cluster with cough, sputum, shortness of breath, and fever;
- 2.a musculoskeletal symptom cluster with myalgia, joint pain, headache, and fatigue;
- 3.a cluster of enteric symptoms with abdominal pain, vomiting, and diarrhoea

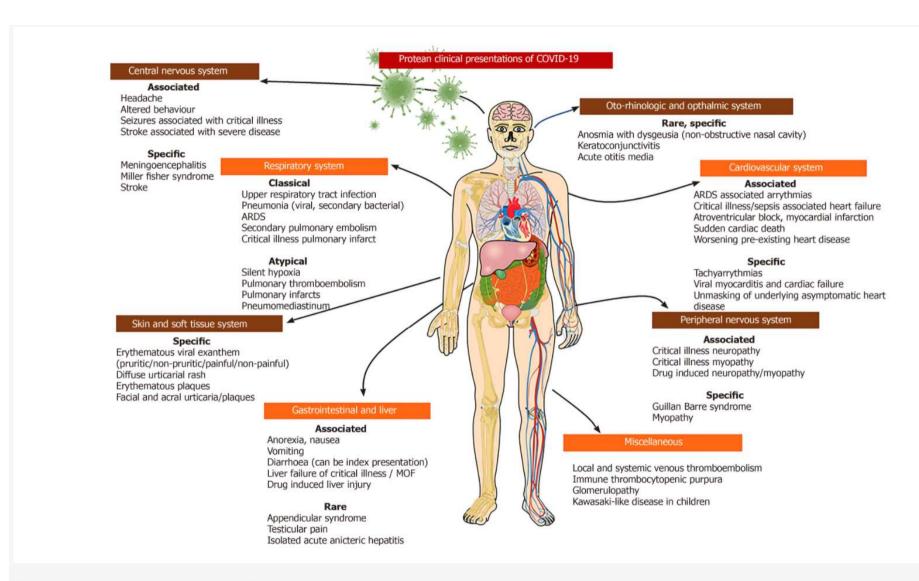
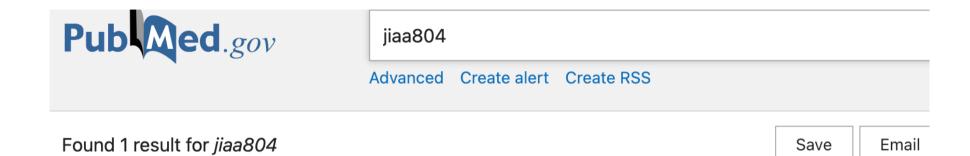


Figure 2 Summary of different clinical presentations and organ system involvement of coronavirus 2019 disease. Overall, multisystem involvement is due to the severity of the disease,



> J Infect Dis. 2021 Jan 3;jiaa804. doi: 10.1093/infdis/jiaa804. Online ahead of print.

Association between upper respiratory tract viral load, comorbidities, disease severity and outcome of patients with SARS-CoV-2 infection

```
Helena C Maltezou <sup>1</sup>, Vasilios Raftopoulos <sup>2</sup>, Rengina Vorou <sup>3</sup>, Kalliopi Papadima <sup>3</sup>, Kassiani Mellou <sup>3</sup>, Nikolaos Spanakis <sup>4</sup>, Athanasios Kossyvakis <sup>5</sup>, Georgia Gioula <sup>6</sup>, Maria Exindari <sup>6</sup>, Elisavet Froukala <sup>4</sup>, Beatriz Martinez-Gonzalez <sup>5</sup>, Georgios Panayiotakopoulos <sup>7</sup>, Anna Papa <sup>6</sup>, Andreas Mentis <sup>5</sup>, Athanasios Tsakris <sup>4</sup>
```

URT viral load status

n = 309 (%) n=316 (%) n=497 (%) Mean age±SD (years) (n=1,082) 50±22 48±21 43±21 0.001 Age group (years) <18 (n=82) 24 (7.9) 22 (7.2) 36 (7.6) 0.107 18-64 (n=767) 202 (66.9) 224 (73.2) 356 (75.1) ≥65 (n=233) 76 (25.2) 60 (19.6) 82 (17.3)	Characteristic
Age group (years) <18 (n=82) 24 (7.9) 22 (7.2) 36 (7.6) 0.107 18-64 (n=767) 202 (66.9) 224 (73.2) 356 (75.1)	
Age group (years) <18 (n=82) 24 (7.9) 22 (7.2) 36 (7.6) 0.107 18-64 (n=767) 202 (66.9) 224 (73.2) 356 (75.1)	
<18 (n=82) 24 (7.9) 22 (7.2) 36 (7.6) 0.107 18-64 (n=767) 202 (66.9) 224 (73.2) 356 (75.1)	Mean age±SD (years) (n=1,082)
18-64 (n=767) 202 (66.9) 224 (73.2) 356 (75.1)	Age group (years)
	<18 (n=82)
≥ 65 (n=233) 76 (25.2) 60 (19.6) 82 (17.3)	18-64 (n=767)
	≥65 (n=233)
Male gender (n=1,122) 156 (50.5) 189 (59.8) 274 (55.1) 0.064	Male gender (n=1,122)
Comorbidities (n=1,122) 124 (40.1) 104 (32.9) 108 (32.1) <0.001	Comorbidities (n=1,122)
Mean number of comorbidities±SD 0.62 + 0.89 0.47 + 0.79 0.32 + 0.67 < 0.001 (n=1,222)	

URT viral load status

Morbidity	High	Moderate	Low	р	-value	
	(n=309)	(n=3	16)	(n=497)		
					×	
Asymptomatic infection (n=274	42 (13	71 (2	2.5%) 16	1 (32.4%)	<0.001	
COVID-19 (n=848)	267	(86.4%)	245 (77.5	5%) 336	(67.6%)	<0.001
Hospitalization (n=518)	153 (4	9.5%) 155 (4	9.1%)	210 (42.3%)	0.064	
Complications (n=231)	88 (28.5%	69 (2:	1.8%)	74 (14.9%)	0.084	
Admission to ICU (n=99)	3	7 (12.0%)	27 (8.5%	s) 35	(7.0%)	
0.055	20)				
Mean ICU LOS ± SD (days)	6.76 <u>+</u>	12.99 5.13 <u>+</u>	13.64 3.	21 <u>+</u> 8.30	0.011*	
Intubation (n=93)	3.	5 (11.3%)	26 (8.2%	32 (6	.40%) 0.0	50
Mean intubation duration ± SD	(days) 7.5	3 <u>+</u> 13.23 5.7	9 <u>+</u> 12.54	3.29 <u>+</u> 8.2	4 0.006	*
Death (n=89)	3.	5 (11.3%)	23 (7.3%	31 (6.	.2%)	0.030



Clinical Infectious Diseases





Issues

More Content ▼

Publish ▼

Purchase

Advertise ▼

About ▼

All Clinical Infectious Disea ▼



Article Contents

Abstract

Supplementary data

Comments (0)

ACCEPTED MANUSCRIPT

Reinfection with SARS-CoV-2: Implications for Vaccines ®

Jeffrey I Cohen ™, Peter D Burbelo

Clinical Infectious Diseases, ciaa1866, https://doi.org/10.1093/cid/ciaa1866

Published: 18 December 2020 Article history ▼



PDF

■■ Split View



Permissions



Abstract

Infection with SARS-CoV-2 has become pandemic and the duration of protective immunity to the virus is unknown. Cases of persons reinfected the virus are being reported with ingreasing frequency. At present it is unconstitution.

Istantanea Schermo

47

View Metrics



Infectious Diseases Society

Table 2. Cases of reinfection with SARS-CoV-2 with different virus strains or clades based on sequence analysis^a

							Antibody	Antibody	y
							present	at onset	
				First	Second		at 1 st	of 2 nd	Virus
Patient	Age, Sex	Location	IC	infection	infection	<u>Interval</u> <u>i</u>	nfection	infection	Sequence
1	33M	Hong Kong	no	Hospitalized	Asymptomatic*	142 days	yes	no	different
clade							•		
2	25M	Nevada	no	Sx, outpatient	Hospitalized	48 days	NR	NR	5
mutatio	ns								
pneumonia									
3	51F	Belgium	no	Sx, outpatient	Sx, milder	3 month	s NR	NR	11
mutatio	ns								
4	60'sM	WA state	no	Hospitalized	Hospitalized	140 days	NR	NR	10
mutations pneumonia sx milder									
5	25M	India	no	Asymptomatic	Asymptomatic	3.5 month	ıs NR	NR	9

Table 3. Implications of reinfection with SARS-CoV-2

- 1. Precautions- masks, distancing are still important after recovery from SARS-CoV-2 in
- the
- absence of a potent vaccine or antiviral
- 2. Previously infected persons may need vaccination
- 3. Herd immunity from infection is unlikely to be sufficient to eliminate the virus if reinfection is common
- 4. Second infection is likely, but not necessarily, to be milder
- 5. Vaccination may not provide lifelong immunity; booster doses may be needed
- 6. Annual quadrivalent flu vaccine may include SARS-CoV-2 vaccine as a component