

Characteristics of SARS-CoV-2 patients dying in Italy Report based on available data on November 11th, 2020

1. Sample

The present report describes characteristics of 41,737 SARS-CoV-2 patients dying in Italy.* Geographic distribution across the 19 regions and 2 autonomous provinces of Trento and Bozen is presented in the table below. Data are update to November 11th, 2020.

Table 1. Geographic distribution of deceased patients SARS-CoV-2 positive

REGION	N	%
Lombardia	18,689	44.8
Emilia Romagna	4,845	11.6
Piemonte	4,121	9.9
Veneto	2,689	6.4
Liguria	2,000	4.8
Lazio	1,534	3.7
Toscana	1,403	3.4
Marche	1,031	2.5
Campania	928	2.2
Puglia	894	2.1
Sicilia	776	1.9
Abruzzo	636	1.5
Trento	477	1.1
Friuli Venezia Giulia	444	1.1
Bolzano	360	0.9
Sardegna	256	0.6
Valle d'Aosta	207	0.5
Umbria	201	0.5
Calabria	128	0.3
Basilicata	61	0.1
Molise	57	0.1
Total	41,737	100.0

^{*} SARS-CoV-2 related deaths presented in this report are those occurring in patients who test positive for SARS-CoV-2RT by PCR, independently from pre-existing diseases.

2. Demographics

Mean age of patients dying for SARS-CoV-2 infection was 80 years (median 82, range 0-109, IQR 74-88). Women were 17,736 (42,5%). Figure 1 shows that median age of patients dying for SARS-CoV-2 infection was more than 30 years higher as compared with the national sample diagnosed with SARS-CoV-2 infection (median age 48 years). Figure 2 shows the absolute number of deaths by age group. Women dying for SARS-CoV-2 infection had an older age than men (median age women 85 - median age men 80).

Figure 1. Median age of patients with SARS-CoV-2 infection and SARS-CoV-2 positive deceased patients

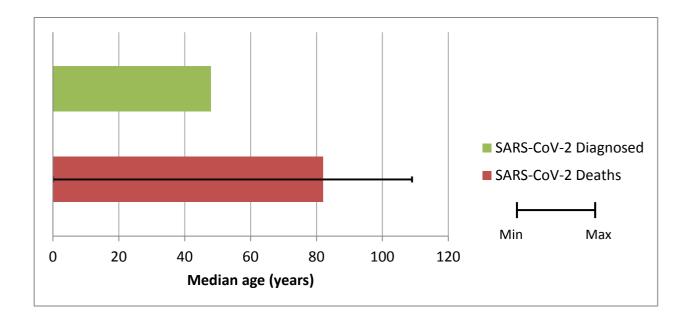
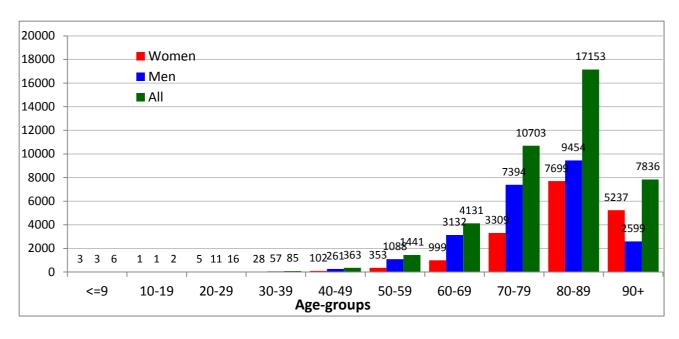


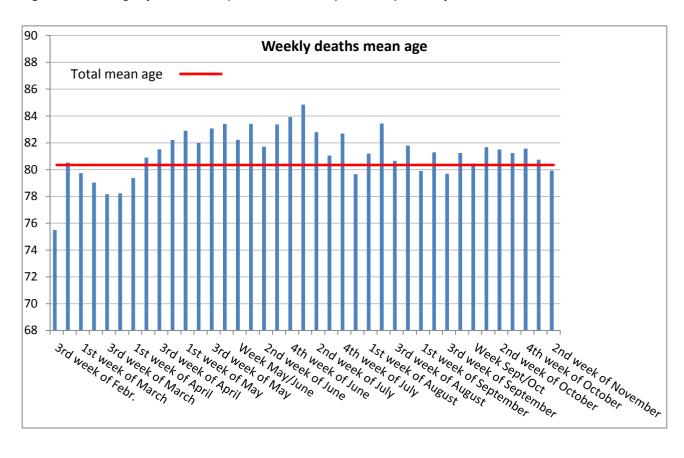
Figure 2. Absolute number of deaths by age group



Note: For 1 deceased person, age was not possible to be evaluated

Figure 3 shows the trend in the average age of SARS-CoV-2 positive deceased patients per calendar week, starting from the 3rd week of February 2020 (the date of the first death dates back to 21st February 2020). The average age of weekly deceased persons has substantially increased up to 85 years (1st week of July) and then dropped slightly.

Figure 3. Mean age of SARS-CoV-2 positive deceased patients by week of death



3. Pre-existing conditions

Table 2 presents most common comorbidities diagnosed before SARS-CoV-2 infection. Data on diseases were based on chart review and was available on 5,234 patients dying in-hospital for whom it was possible to analyse clinic charts. Mean number of diseases was 3.5 (median 3, SD 2.0). Overall, 3.3% of the sample presented with a no comorbidities, 13.0% with a single comorbidity, 18.9% with 2, and 64.8% with 3 or more.

Before hospitalization, 21% of SARS-CoV-2 positive deceased patients followed ACE-inhibitor therapy and 14% angiotensin receptor blockers-ARBs therapy. This information can be underestimated because data on drug treatment before admission were not always described in the chart.

Table 2. Most common comorbidities observed in SARS-CoV-2 positive deceased patients

Diseases	N	%
Ischemic heart disease	1449	27.7
Atrial Fibrillation	1266	24.2
Heart failure	858	16.4
Stroke	595	11.4
Hypertension	3441	65.7
Type 2-Diabetes	1531	29.3
Dementia	1148	21.9
COPD (Chronic Obstructive Pulmonary Disease)	895	17.1
Active cancer in the past 5 years	896	17.1
Chronic liver disease	244	4.7
Chronic renal failure	1099	21.0
Dialysis	109	2.1
Respiratory failure	345	6.6
HIV Infection	11	0.2
Autoimmune diseases	223	4.3
Obesity	541	10.3
Number of comorbidities		
0 comorbidities	174	3.3
1 comorbidity	682	13.0
2 comorbidities	988	18.9
3 comorbidities and over	3390	64.8

Table 3 presents the most common pre-existing chronic pathologies in patients who died, separately in men (n = 3,162) and women (n = 2,072). The average number of pathologies observed in women is 3.7 (median 3, Standard Deviation 2.0). In men the average number of pathologies observed is 3.4 (median 3, Standard Deviation 2.1).

Table 3. Most common comorbidities observed in SARS-CoV-2 positive deceased patients by gender

Women Men

Diseases	N	%
Ischemic heart disease	474	22.9
Atrial Fibrillation	531	25.6
Heart Failure	391	18.4
Stroke	254	12.3
Hypertension	1415	68.3
Type 2-Diabetes	556	26.8
Dementia	636	30.7
COPD (Chronic Obstructive Pulmonary Disease)	282	13.6
Active cancer in the past 5 years	335	16.2
Chronic liver disease	88	4.2
Chronic renal failure	394	19.0
Dialysis	31	1.5
Respiratory failure	143	6.9
HIV Infection	0	0.0
Autoimmune diseases	125	6.0
Obesity	215	10.4
Number of comorbidities		
0 comorbidities	44	2.1
1 comorbidity	235	11.3
2 comorbidities	369	17.8
3 comorbidities and over	1424	68.7

N	%
975	30.8
735	23.2
467	14.5
341	10.8
2026	64.1
975	30.8
512	16.2
613	19.4
561	17.7
156	4.9
705	22.3
78	2.5
202	6.4
11	0.3
98	3.1
326	10.3
130	4.1
447	14.1
619	19.6
1966	62.2

4. Diagnosis of hospitalization

In 90.6% of hospitalizations, conditions (e.g. pneumonia, respiratory failure) or symptoms (e.g. fever, dyspnoea, cough) compatible with SARS-CoV-2 were mentioned. In 455 cases (9.4% of cases) the diagnosis of hospitalization was not related to the infection. In 67 cases the diagnosis of hospitalization concerned exclusively neoplastic pathologies, in 163 cases cardiovascular pathologies (for example Acute Myocardial Infarction-AMI, heart failure, stroke), in 61 cases gastrointestinal pathologies (for example cholecystitis, perforation of the intestine, intestinal obstruction, cirrhosis), in 164 cases other pathologies.

5. Symptoms

Figure 4 shows symptoms most commonly observed at hospital admission. Fever, dyspnoea and cough were the most commonly observed symptoms, while diarrhoea and haemoptysis were less commonly observed. Overall, 7.8% of patients did not present any symptoms at hospital admission.

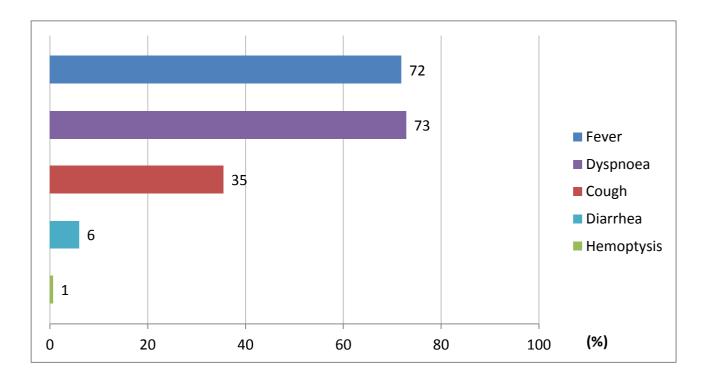


Figure 4. Most common symptoms observed in SARS-CoV-2 positive deceased patients

6. Acute conditions

Acute Respiratory Distress syndrome was observed in the majority of patients (93.9% of cases), followed by acute renal failure (23.5%). Superinfection was observed in 19.2% and acute cardiac injury in 11.2% of cases.

7. Treatments

Antibiotics were used by 86.0% of patients during hospital stay, while less used were antivirals (53.8%) and corticosteroids (46.9%). Concomitant use of these 3 treatments was observed in 25.9% of cases.

Out of SARS-CoV-2 positive deceased patients, 4.4% were treated with Tocilizumab during hospitalization.

8. Time-line

Figure 5 shows, for SARS-CoV-2 positive deceased patients, the median times, in days, from the onset of symptoms to death (12 days), from the onset of symptoms to hospitalization (5 days) and from hospitalization to death (7 days). The time from hospitalization to death was 6 days longer in those who were transferred to intensive care than those who were not transferred (12 days vs. 6 days).

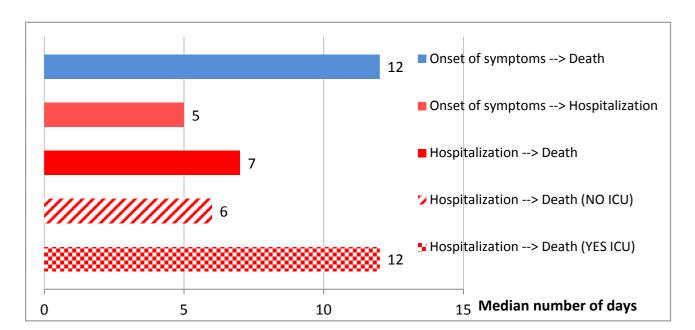


Figure 5. Median hospitalization times (in days) in SARS-CoV-2 positive deceased patients

9. Deaths under the age of 50 years

As of November 11th 2020, 472 out of the 41,737 (1.1%) positive SARS-CoV-2 patients under the age of 50 died. In particular, 109 of these were less than 40 years (72 men and 37 women), age range between 0 and 39 years. For 31 patients under the age of 40 years no clinical information is available; out of the remaining ones, 64 had serious pre-existing pathologies (cardiovascular, renal, psychiatric pathologies, diabetes, obesity) and 14 had no major pathologies.

10. Comparison of death characteristics in the 3 quarters March-May, June-August, and September-November 2020

Table 4 summarizes the main characteristics of deaths with COVID-19 that occurred in 3 periods of time from the beginning of the pandemic in 2020: the initial quarter, March-May, the second quarter, June-August, and the third period September-November. Overall, the sample represents 12.8% of all deaths from the beginning of the pandemic; in particular, the 13.2% of those who died between March and May, 26.9% of those who died between June and August, and 7.1% of those who died between September and November.

In the second and third periods, the average age of deaths slightly increased (particularly in the second period) in comparison to the first period; the proportion of women increases in the second period only; deaths of people with 3 or more pre-existing pathologies increase and those of persons with fewer pathologies or none decrease: this seems to indicate that in the second and third period deaths concern older people and persons with a pre-existing health condition worse than those dying in the first quarter (table 4).

The use of drugs is also extremely different in the three periods, with a clear reduction in the use of antivirals and an increase in the use of steroids in the second and third periods.

Table 4. Mean age, prevalence of women, number of pre-existing diseases, complications and treatments in deaths with COVID-19 in the 3 periods March-May, June-August, and September-November 2020

All deaths	All (n=41,725) (Missing=11)	March-May 2020 (n=34,194)	June-August 2020 (n=1,399)	Sept-Nov 2020 (n=6,132)	p- value*
Age (years)	80.3	80.1	82.8	81.0	<0.001
	n (%)	n (%)	n (%)		
Women	17,732 (42.5)	14,321 (41.9)	865 (61.8)	2,546 (41.5)	<0.001
Sample of the evaluated clinical charts	All (n=5,234)	March-May 2020 (n=4,431)	June-August 2020 (n=371)	Sept-Nov 2020 (n=432)	p- value*
N of comorbidities					
0	174 (3,3)	168 (3,8)	3 (0,8)	3 (0,7)	
1	682 (13,0)	604 (13,6)	34 (9,2)	44 (10,2)	<0.001
2	988 (18,9)	882 (19,9)	49 (13,2)	57 (13,2)	0.001
3 or more	3,390 (64,8)	2,777 (62,7)	285 (76,8)	328 (75,9)	=
Complications during hospitalization					
Acute Respiratory Distress Syndrome	4,799 (93.9)	4,108 (95.2)	293 (79.4)	398 (93.0)	<0.001
Acute renal failure	1,199 (23.5)	992 (23.0)	100 (27.1)	107 (25.0)	0.148
Acute cardiac injury	574 (11.2)	468 (10.8)	38 (10.3)	68 (15.9)	0.006
Superinfection	984 (19.2)	715 (16.6)	173 (46.9)	96 (22.4)	<0.001
Treatments					
Antibiotics	4,435 (86.0)	3,779 (86.7)	311 (85.0)	345 (80.4)	0.001
Antivirals	2,773 (53.8)	2,578 (59.1)	129 (35.2)	66 (15.4)	<0.001
Steroids	2,420 (46.9)	1,883 (43.2)	220 (60.1)	317 (73.9)	<0.001
Tocilizumab	203 (4.4)	169 (4.4)	25 (7.1)	9 (2.2)	0.004

^{*} *p-value* for difference between the 3 quarters

The distribution of the main pre-existing diseases in the different periods is presented in *figure 6*. The prevalence of atrial fibrillation, heart failure, stroke, dementia, cancer and renal insufficiency varies significantly in the three periods. These pathologies are more frequently diagnosed in the deceased in the second and third period than in the first (*Figure 6*).

Figure 6. Pre-existing pathologies in deaths with COVID-19 in the 3 periods

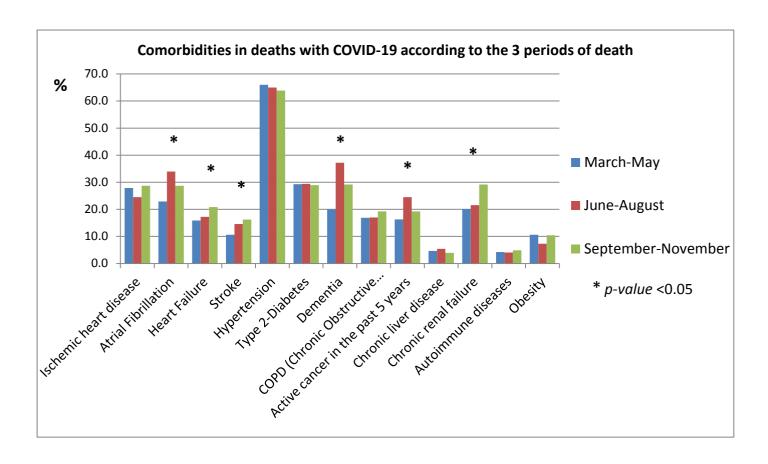


Table 5 shows the durations, as median times (in days), from the symptoms onset to death, SARS-CoV-2 testing, and hospitalization, and from the hospitalization to death, in the 3 periods considered. Between the first and second period triples the time that passes from the onset of symptoms to death, while it returns to the initial levels in the third period; decreases the time from the onset of symptoms to the swab for the detection of SARS-CoV-2 infection in both the second and third period, as well as the time between the onset of symptoms and hospitalization; the median duration in days from hospitalization to death between the first and second period triples; it decreases again in the third period, even if it remains above the levels of the first period. These results seem to suggest a greater reactivity of the Health System evidenced by the greater speed in carrying out diagnostic tests and hospitalization.

Table 5. Median times (in days) between symptoms onset, PCR test, hospitalization and death in the 3 periods

	All (n=5,2		March-May 2020 (n=4,431)		June-August 2020 (n=371)		September- November 2020 (n=432)		p- value*
Times(in days)	Median	IQR	Median	IQR	Median	IQR	Median	IQR	
From symptoms onset to death	12	7-20	12	7-19	37	10-65	12	6-19	<0.001
From symptoms onset to SARS-CoV-2 testing	5	2-8	5	2-9	3	1-10	2	0-6	<0.001
From symptoms onset to hospitalization	4	2-7	4	2-7	3	1-7	3	1-6	<0.001
From hospitalization to death	7	3-14	7	3-13	21	5-51	8	3-13	<0.001

^{*} *p-value* for difference between the 3 quarters

IQR=Inter-Quartile Range

The data here presented can be explained by a greater knowledge about the infection and a greater ability and timeliness of treatment in the period June-August in comparison to the previous quarter. In addition, it is likely that in the months of March and April SARS-CoV-2 infection was under-diagnosed in many frail elderly who died (such as those living in RSA). This may have led to an underestimation of the burden of the diseases in persons dying in that period

This report was produced by SARS-CoV-2 Surveillance Group

Members of the SARS-CoV-2 Surveillance Group

Luigi Palmieri, Elvira Agazio, Xanthi Andrianou, Pierfrancesco Barbariol, Antonino Bella, Stefania Bellino, Eva Benelli, Luigi Bertinato, Matilde Bocci, Stefano Boros, Gianfranco Brambilla, Giovanni Calcagnini, Marco Canevelli, Maria Rita Castrucci, Federica Censi, Alessandra Ciervo, Elisa Colaizzo, Fortunato D'Ancona, Martina Del Manso, Corrado Di Benedetto, Chiara Donfrancesco, Massimo Fabiani, Francesco Facchiano, Antonietta Filia, Marco Floridia, Fabio Galati, Marina Giuliano, Tiziana Grisetti, Cecilia Guastadisegni, Yllka Kodra; Martin Langer, Ilaria Lega, Cinzia Lo Noce, Pietro Maiozzi, Fiorella Malchiodi Albedi, Valerio Manno, Margherita Martini, Alberto Mateo Urdiales, Eugenio Mattei, Claudia Meduri, Paola Meli, Giada Minelli, Manuela Nebuloni, Lorenza Nisticò, Marino Nonis, Graziano Onder, Lucia Palmisano, Nicola Petrosillo, Patrizio Pezzotti, Flavia Pricci, Ornella Punzo, Vincenzo Puro, Federica Quarata, Valeria Raparelli, Giovanni Rezza, Flavia Riccardo, Simone Rocchetto, Maria Cristina Rota, Paolo Salerno, Giulia Sarti, Debora Serra, Andrea Siddu, Paola Stefanelli, Manuela Tamburo De Bella, Dorina Tiple, Marco Toccaceli Balzi, Federica Trentin, Brigid Unim, Luana Vaianella, Nicola Vanacore, Maria Fenicia Vescio, Monica Vichi, Emanuele Rocco Villani, Amerigo Zona, Silvio Brusaferro.