

Characteristics of SARS-CoV-2 patients dying in Italy Report based on available data on December 2nd, 2020

1. Sample

The present report describes characteristics of 55,824 SARS-CoV-2 patients dying in Italy^{*}. Data are update to December 2nd, 2020. Geographic distribution across the 19 regions and 2 autonomous provinces of Trento and Bozen is presented in *Table 1*. Absolute number and percentage of deaths are reported in descending order of the total, according to the 3 periods since the beginning of the pandemic in 2020: the initial period (March-May), the second period (June-September), and the third (October-December), although the latter is still ongoing.

	March	ch-May June-September		ptember	October-D	December	Total	
REGION	N	%	N	%	N	%	N	%
Lombardia	16,360	47.6	606	32.3	5,286	27.0	22,252	39.9
Emilia Romagna	4,313	12.6	173	9.2	1,318	6.7	5,804	10.4
Piemonte	4,077	11.9	168	9.0	1,306	6.7	5,551	9.9
Veneto	1,950	5.7	248	13.2	1,701	8.7	3,899	7.0
Lazio	840	2.4	141	7.5	1,539	7.9	2,520	4.5
Liguria	1,516	4.4	138	7.4	764	3.9	2,418	4.3
Toscana	1,045	3.0	96	5.1	1,253	6.4	2,394	4.3
Campania	477	1.4	56	3.0	1,621	8.3	2,154	3.9
Sicilia	300	0.9	56	3.0	1,206	6.2	1,562	2.8
Puglia	524	1.5	75	4.0	947	4.8	1,546	2.8
Marche	981	2.9	7	0.4	169	0.9	1,157	2.1
Abruzzo	446	1.3	37	2.0	443	2.3	926	1.7
Friuli Venezia Giulia	343	1.0	20	1.1	373	1.9	736	1.3
Trento	402	1.2	4	0.2	256	1.3	662	1.2
Bolzano	290	0.8	2	0.1	270	1.4	562	1.0
Sardegna	131	0.4	24	1.3	269	1.4	424	0.8
Umbria	75	0.2	9	0.5	323	1.7	407	0.7
Valle d'Aosta	143	0.4	6	0.3	168	0.9	317	0.6
Calabria	96	0.3	4	0.2	152	0.8	252	0.5
Basilicata	29	0.1	2	0.1	105	0.5	136	0.2
Molise	22	0.1	2	0.1	104	0.5	128	0.2
Total	34,360	100.0	1,874	100.0	19,573	100.0	55,807	100.0

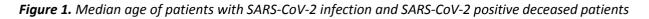
Table 1. Geographic distribution of deceased patients SARS-CoV-2 positive, by period

Note: For 17 deceased persons, period was not possible to be evaluated

* 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 23,596 (42.3%). *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).



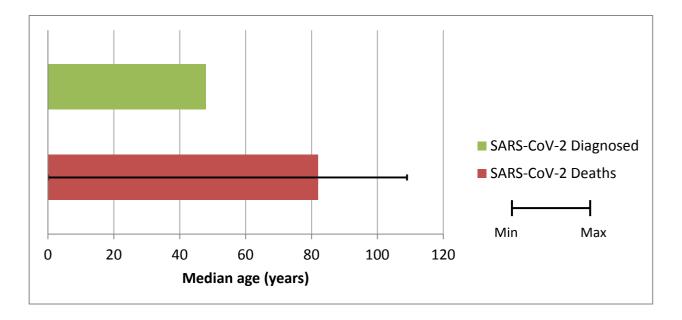


Figure 2. Absolute number of deaths by age group

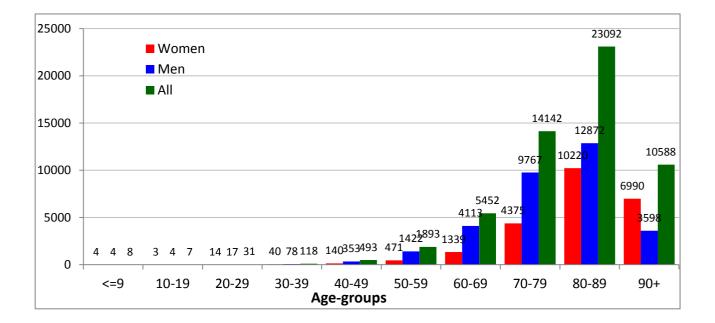


Figure 3 shows the trend in the average age of SARS-CoV-2 positive deceased patients per calendar week, starting from the 3rd week of February 2020 (the date of the first death dates back to 21st February 2020). The average age of weekly deceased persons has substantially increased up to 85 years (1st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

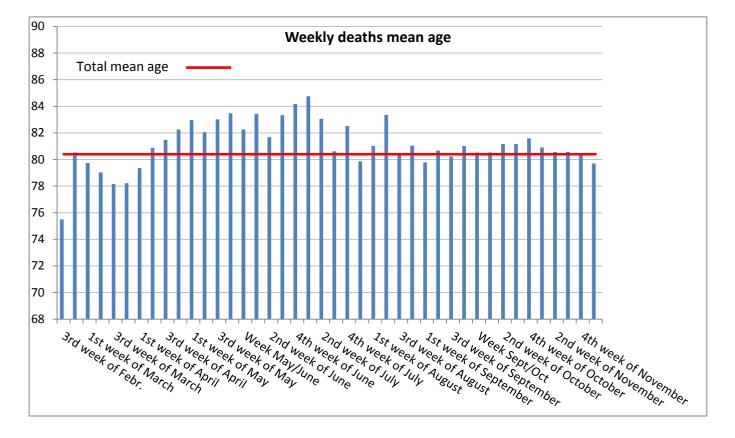


Figure 3. Mean age of SARS-CoV-2 positive deceased patients by week of death

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 5,726 patients dying in-hospital for whom it was possible to analyse clinic charts. Mean number of diseases was 3.6 (median 3, SD 2.1). Overall, 3.1% of the sample presented with a no comorbidities, 12.4% with a single comorbidity, 18.5% with 2, and 65.9% 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.

Diseases	N	%
Ischemic heart disease	1591	27.8
Atrial Fibrillation	1388	24.2
Heart failure	936	16.3
Stroke	662	11.6
Hypertension	3779	66.0
Type 2-Diabetes	1671	29.2
Dementia	1323	23.1
COPD (Chronic Obstructive Pulmonary Disease)	992	17.3
Active cancer in the past 5 years	975	17.0
Chronic liver disease	268	4.7
Chronic renal failure	1207	21.1
Dialysis	121	2.1
Respiratory failure	390	6.8
HIV Infection	14	0.2
Autoimmune diseases	252	4.4
Obesity	591	10.3
Number of comorbidities		
0 comorbidities	180	3.1
1 comorbidity	712	12.4
2 comorbidities	1060	18.5
3 comorbidities and over	3774	65.9

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

Table 3 presents the most common pre-existing chronic pathologies in patients who died, separately in men (n = 3,441) and women (n = 2,285). The average number of pathologies observed in women is 3.8 (median 4, Standard Deviation 2.0). In men the average number of pathologies observed is 3.5 (median 3, Standard Deviation 2.1).

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

Diseases	N	%	N
Ischemic heart disease	534	23.4	1057
Atrial Fibrillation	588	25.7	800
Heart Failure	434	18.5	502
Stroke	286	12.5	376
Hypertension	1570	68.7	2209
Type 2-Diabetes	614	26.9	1057
Dementia	733	32.1	590
COPD (Chronic Obstructive Pulmonary Disease)	318	13.9	674
Active cancer in the past 5 years	360	15.8	615
Chronic liver disease	93	4.1	175
Chronic renal failure	438	19.2	769
Dialysis	36	1.6	85
Respiratory failure	161	7.0	229
HIV Infection	2	0.1	12
Autoimmune diseases	142	6.2	110
Obesity	232	10.2	359
Number of comorbidities			
0 comorbidities	47	2.1	133
1 comorbidity	246	10.8	466
2 comorbidities	390	17.1	670
3 comorbidities and over	1602	70.1	2172

Women

Men

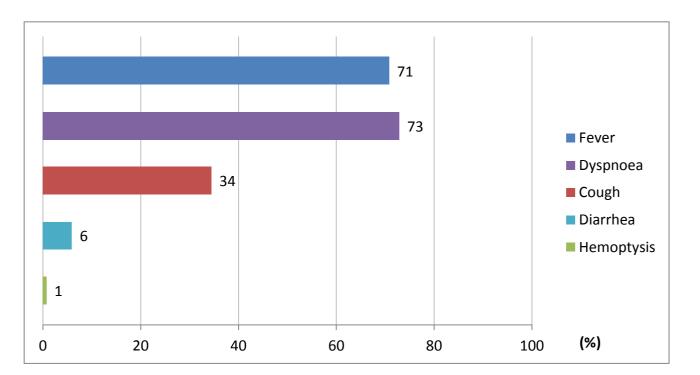
N	%		
1057	30.7		
800	23.2		
502	14.4		
376	10.9		
2209	64.2		
1057	30.7		
590	17.1		
674	19.6		
615	17.9		
175	5.1		
769	22.3		
85	2.5		
229	6.7		
12	0.3		
110	3.2		
359	10.4		
133	3.9		
466	13.5		
670	19.5		
2172	63.1		

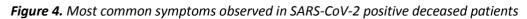
4. Diagnosis of hospitalization

In 90.8% of hospitalizations, conditions (e.g. pneumonia, respiratory failure) or symptoms (e.g. fever, dyspnoea, cough) compatible with SARS-CoV-2 were mentioned. In 491 cases (9.2% of cases) the diagnosis of hospitalization was not related to the infection. In 72 cases the diagnosis of hospitalization concerned exclusively neoplastic pathologies, in 165 cases cardiovascular pathologies (for example Acute Myocardial Infarction-AMI, heart failure, stroke), in 70 cases gastrointestinal pathologies (for example cholecystitis, perforation of the intestine, intestinal obstruction, cirrhosis), in 184 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, 8.1% of patients did not present any symptoms at hospital admission.





6. Acute conditions

Acute Respiratory Distress syndrome was observed in the majority of patients (94.1% of cases), followed by acute renal failure (23.6%). Superinfection was observed in 19.3% and acute cardiac injury in 10.8% of cases.

7. Treatments

Antibiotics were used by 85.9% of patients during hospital stay, while less used were antivirals (50.2%) and corticosteroids (49.8%). Concomitant use of these 3 treatments was observed in 24.5% of cases.

Out of SARS-CoV-2 positive deceased patients, 4.1% 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).

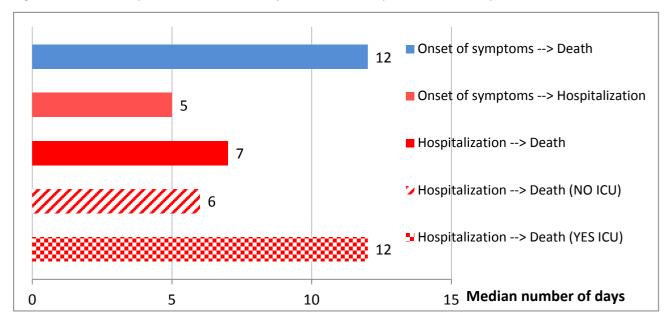


Figure 5. Median hospitalization times (in days) in SARS-CoV-2 positive deceased patients

9. Provenience

Most of the SARS-CoV-2 positive deceased patients, at the time of the hospital admission, came from their home (53.3%); 22.9% came from social-health or social-welfare residential structures (nursing homes, hospices, wards or long-term care facilities); 18.6% had been transferred from another hospital and 5.2% from other unspecified facilities.

10. Deaths under the age of 50 years

As of December 2nd 2020, 657 out of the 55,824 (1.2%) positive SARS-CoV-2 patients under the age of 50 died. In particular, 163 of these were less than 40 years (102 men and 61 women), age range between 0 and 39 years. For 29 patients under the age of 40 years no clinical information is available; out of the remaining ones, 119 had serious pre-existing pathologies (cardiovascular, renal, psychiatric pathologies, diabetes, obesity) and 15 had no major pathologies.

11. Comparison of death characteristics in the 3 quarters March-May, June- September, and October-December 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-September, and the third period October-December. Overall, the sample represents 10.5% of all deaths from the beginning of the pandemic; in particular, the 13.3% of those who died between March and May, 26.9% of those who died between June and September, and 4.0% of those who died between October and December.

Both the mean age of deaths and the proportion of women increase only in the second period; 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 periods 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 treatmentsin deaths with COVID-19 in the 3 periods March-May, June- September, and October-December 2020

All deaths	All (n=55,807) (Missing=17)	March-May 2020 (n=34,360)	June- Sept 2020 (n=1,874)	Oct-Dec 2020 (n=19,573)	p- value*	
Mean age (years)	80.3	80.1	82.3	80.7	<0.001	
	n (%)	n (%)	n (%)			
Women	23,587 (42.3)	14,421 (42.0)	1087 (58.0)	8079 (41.3)	<0.001	
Sample of the evaluated clinical charts	All (n=5,726)	March-May 2020 (n=4,466)	June- Sept 2020 (n=499)	Oct-Dec 2020 (n=761)	p- value*	
N of comorbidities						
0	180 (3.1)	169 (3.8)	5 (1.0)	6 (0.8)		
1	712 (12.4)	605 (13.5)	44 (8.8)	63 (8.3)		
2	1060 (18.5)	894 (20.0)	66 (13.2)	100 (13.1)	<0.001	
3 or more	3774 (65.9)	2798 (62.7)	384 (77.0)	592 (77.8)	1	
Provenience						
Nursing homes, hospices, wards or long-term care facilities	285 (22.9)	93 (25.1)	45 (26.0)	147 (21.0)	0.181	
Home	664 (53.3)	200 (53.9)	78 (45.1)	386 (55.1)	0.060	
Other Hospital	231 (18.6)	54 (1.2)	43 (24.9)	134 (19.1)	0.013	
Other	65 (5.2)	24 (0.5)	7 (4.0)	34 (4.9)	0.398	
Complications during hospitalization						
Acute Respiratory Distress Syndrome	5274 (94.1)	4138 (95.1)	410 (83.2)	726 (95.0)	<0.001	
Acute renal failure	1323 (23.6)	1001 (23.0)	133 (27.0)	189 (24.7)	0.106	
Acute cardiac injury	608 (10.8)	471 (10.8)	49 (9.9)	88 (11.5)	0.678	
Superinfection	1081 (19.3)	721 (16.6)	201 (40.8)	159 (20.8)	<0.001	
Treatments						
Antibiotics	4854 (85.9)	3810 (86.7)	422 (85.9)	622 (81.5)	0.001	
Antivirals	2835 (50.2)	2597 (59.1)	156 (31.8)	82 (10.7)	<0.001	
Steroids	2816 (49.8)	1901 (43.3)	319 (65.0)	596 (78.1)	<0.001	
Tocilizumab	210 (4.1)	170 (4.4)	28 (5.9)	12 (1.6)	<0.001	

* *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, stroke, dementia, COPD, 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*).

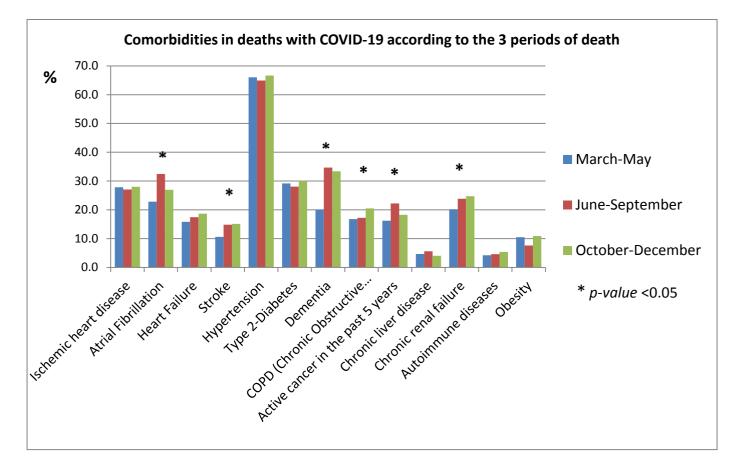


Figure 6. Pre-existing pathologies in deaths with COVID-19 in the 3 periods

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 doubles 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 doubles; it decreases again in the third 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=5,726)		March-May 2020 (n=4,466)		June- Sept 2020 (n=499)		Oct-Dec 2020 (n=761)		p- value*
Times(in days)	Median	IQR	Median	IQR	Median	IQR	Median	IQR	
From symptoms onset to death	12	7-20	12	7-19	22	9-55	11	6-18	<0.001
From symptoms onset to SARS-CoV-2 testing	5	2-8	5	2-9	3	1-7	3	1-6	<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	15	5-43	7	3-12	<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

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