# Prevalence of Comorbidities and their Impact on COVID-19 Outcomes: An Observational Study during Early Outbreak in Lebanon

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#### 1. Background

In December 2019, clusters of pneumonia of unknown origin broke out in Wuhan, China. The infectious agent was later identified as severe acute respiratory syndrome coronavirus 2(SARS-CoV-2), a new coronavirus that was not previously seen in humans, and belongs to the same genera of viruses that have previously caused severe acute respiratory syndrome (SARS) and Middle East Respiratory Syndrome (MERS) disease outbreaks [1-3]. The World Health Organization (WHO) first considered SARS-CoV-2 infection, or Coronavirus disease 2019 (COVID-19), as a public health emergency of international concern on 30 January 2020, and then officially declared it a pandemic on 11 March 2020 [4]. By mid-May 2020, there were more than 4 million confirmed cases of COVID-19 and almost 300 thousand deaths reported worldwide [5]. In Lebanon, the first case of COVID-19 was confirmed on February 21, 2020. By mid-May 2020, there were 891 cases and 26 deaths reported in Lebanon, but the transmission was still within "clusters of cases" and the country had not yet entered the "community transmission" phase [5,6].

The clinical presentation of COVID-19 is widely variable, ranging from asymptomatic infection, mild upper respiratory symptoms, to severe viral pneumonia leading to acute respiratory distress syndrome (ARDS), and even death. Typical initial symptoms of infection include fever, cough, and dyspnea [7, 8].

Ever since its emergence, clinical and epidemiological data on COVID-19 suggested that some comorbidities are associated with a higher risk of contracting the disease, and are correlated with an increased risk of severe disease, disease complications, and poor outcomes, including death. Older age, diabetes, hypertension, obesity, and cardiovascular disease (CVD) were considered as risk factors of particular interest in COVID-19 studies, which was supported by numerous observations since early outbreak [9-13].

In Lebanon, non-communicable diseases (NCDs) present a major health challenge. In 2018, the WHO reported that NCDs are estimated to account for 91% of all deaths in Lebanon: 47% due to CVD, 16% due to cancers, 4% due to chronic respiratory diseases, and 5% due to diabetes [14].

While different countries have reported variable case-fatality rates and disease patterns [5], epidemiological studies on COVID-19 are still lacking in Lebanon. The aim of our study was to describe epidemiological and clinical characteristics of patients presenting with COVID-19 in Lebanon during early outbreak, and to explore the association of baseline risk factors and comorbidities with COVID-19 disease course and outcomes.

#### 2. Methods

#### Study design and setting

This was a retrospective observational single-center study that included 150 patients admitted to Rafik Hariri University Hospital (RHUH) with newly diagnosed SARS-CoV-2 infections starting February 21, 2020. RHUH is the largest Lebanese public hospital located in Beirut, and until the end of data collection, it was the main central hospital in Lebanon receiving most COVID-19 cases. Data was collected between March 15 and May 15, 2020.

A database for COVID-19 patients was created once the first patient was admitted to RHUH. We randomly selected 150 patients newly diagnosed with COVID-19, regardless of age, gender, or disease severity, to be included in the study. Patients were considered as SARS-CoV-2-positive based on the result of real-time polymerase chain reaction (RT-PCR) tests performed on respiratory samples at the study center. Patients visited the center if they had any symptom (dyspnea, cough, fever), were coming from travel from a high COVID-19 prevalence country, or suspected infection based on contact with a confirmed case.

At the beginning of the national outbreak, and to limit the spread of the infection, all confirmed cases were hospitalized as a form of quarantine, regardless if patients actually required in-patient care, i.e. even if they were asymptomatic or had mild symptoms.

#### Data collected

Data collected from patients' medical records included: demographic and anthropometric characteristics (age, gender, body mass index [BMI]), comorbidities and potential risk factors (CVD, chronic lung disease, chronic kidney disease, chronic liver disease, chronic neurological disorder, malignant neoplasm, obesity, hypertension, diabetes, smoking), signs and symptoms upon admission (fever, cough, dyspnea, other), clinical characteristics and complications (intensive care unit [ICU]/regular floor admission, need for mechanical ventilation (intubation), developing viral pneumonitis, bacterial pneumonia, ARDS, congestive heart failure [CHF], cardiac arrhythmia, cardiac ischemia, cardiac arrest, acute renal injury, liver dysfunction, hyperglycemia, hypoglycemia), and disease outcomes (length of hospital stay, RT-PCR status upon discharge, disease cure [defined as having 2 consecutive negative PCRs], time to conversion [time from first positive PCR till 2 consecutive negatives], death, disease severity). Data was collected from standard forms that were completed upon admission. Comorbidities were determined based on patients' self-report. Patients were discharged if they had 2 consecutive negative PCRs or if they were completely asymptomatic with an available place to be quarantined. If discharged while positive, patients presented for a follow-up PCR to ensure that the PCR test result became negative.

The study was approved by the Institutional Review Board of RHUH and all study data were anonymized and de-identified.

#### Statistical analyses

All statistical analyses were conducted using IBM SPSS Statistics for Windows, Version 22.0. Armonk, NY: IBM Corp. No formal sample size calculation was performed since the study was done during the early outbreak period in Lebanon. At time of data collection, our sample size was considered to be sufficient, considering its representativeness of the national patient population.

All statistical tests were two-sided; p-values <0.05 were considered as statistically significant. No missing data was imputed.

Continuous variables were presented as means and standard deviations while categorical variables were presented as frequencies and percentages.

Comparison of means of continuous variables between 2 independent groups was done using Student's t-test for normally distributed data, while Mann-Whitney U test was used for data that did not meet the assumptions of a normal distribution.

Pearson chi-square or Fischer's exact tests were used for comparisons between proportions as appropriate.

Normality in distribution was tested by graphical and numerical methods. Histogram plots and Skewness/Kurtosis tests were computed for each variable. Kolmogorov-Smirnov and Shapiro-Wilk tests were used to determine whether the variable had a normal distribution.

Selected risk factors (age, obesity, CVD, hypertension, diabetes, and smoking) were evaluated for possible association with clinical characteristics and as well as disease outcomes in the sample of COVID-19 patients.

#### 3. Results

#### **Patient demographics**

A total of 150 patients were enrolled in the study. Mean age was  $45.87 \pm 20.03$  years, with 64 patients (42.67%) being 50 years or older [age range 5 – 88 years]. Most patients were male (60.67%). Thirty patients were smokers (20%) (Table 1).

 Table 1: Baseline characteristics and disease symptoms, course, and outcomes in COVID-19 patients

Baseline Characte	ristics							
Age (years)	Mean ±	45.87+20.03						
n=150	SD	10107 == 0100						
Age Category	<50 years	86 (57.33%)						
n=150	$\geq$ 50 years	64 (42.67%)						
Gender	Male	91 (60.67%)						
n=150	Female	59 (39.33%)						
BMI (kg/m <sup>2</sup> ) <sup>a</sup>	Mean ±	28 90 +7 85						
n=20	SD	20.00 ±1.05						
Smoking	No	120						
n=150		(80.00%)						
	Yes	30 (20.00%)						
COVID-19 symp	otoms							
Fever	No	67 (44.67%)						
n=150	Yes	83 (55.33%)						
Cough	No	65 (43.33%)						
n=150	Yes	85 (56.67%)						
D	N-	121						
Dyspnea	INO	(81.21%)						
n=149	Yes	28 (18.79%)						
Other	No	73 (48.67%)						
n=150	Yes	77 (51.33%)						
Disease course and o	outcomes	•						
Length of hospital stay (days)	Mean ±	14.00 . 7.50						
n=150	SD	$14.99 \pm 7.58$						
Patient was discharged while PCR-	No	88 (62.41%)						
positive, n=141	Yes	53 (37.59%)						
Discourse	No	9 (6.04%)						
Disease cure	Vaa	140						
11=149	res	(93.96%)						
Time to PCR test conversion (days)	Mean ±	21.30 ±						
n=140	SD	11.00						

<sup>a</sup>BMI was only recorded for patients admitted to the ICU

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**Prevalence of risk factors, comorbidities, and symptoms** Regarding comorbidities, hypertension (18.67%) and diabetes (16.67%) were the most commonly reported, followed by CVD (9.33%), obesity (4%), malignant neoplasms (3.33%), chronic kidney disease (2.00%), chronic neurological disorder (1.33%), and chronic lung disease (0.67%). None of the patients had chronic liver disease. Overall, 103 patients reported no comorbidities (68.67%) (Figure 1).



Figure 1: Numbers and types of comorbidities in the study population

During COVID-19 illness, the most common symptoms were cough (56.67%) and fever (55.33%), while dyspnea was less common (18.79%) (Table 1). 31 patients have not

reported having any symptoms (20.67%). The disease severity was mild in 47.33% of patients, moderate in 37.33% and severe in 15.33% (Figure 2).

DOI: 10.21275/SR201121021100



**Figure 2:** COVID-19 disease severity, complications, and outcome ICU: Intensive Care Unit; ARDS: Acute Respiratory Distress Syndrome; CHF: Congestive Heart Failure

#### COVID-19 disease course and outcomes

128 patients were admitted to regular floor (85.33%) while 22 patients (14.67%) were in ICU. The most commonly reported disease complications were viral pneumonitis (52.67%), bacterial pneumonia (14.67%), hyperglycemia (10.42%), ARDS (8.00%), and acute renal injury (7.33%). Nine patients needed mechanical ventilation (6.00%) and the same number experienced cardiac arrest. Other complications are summarized in Figure 2.

For the 150 enrolled patients, the average length of hospital stay was  $14.99 \pm 7.58$  days. Nine patients died during hospitalization (6.00%). Of those who were alive at discharge, 53 patients (37.59%) were discharged while PCR-positive. At last follow-up, one patient was lost to follow-up and 140 patients were cured (93.96%) from the disease;

among those, average time to conversion was 21.30  $\pm 11.00$  days (Table 1).

## Association of comorbidities with disease course and outcome

COVID-19 symptoms were associated with several risk factors and comorbidities. Patients with fever were more likely to be obese (100% vs. 53.47%; p=0.033), those with cough were more likely to be  $\geq$ 50 years old (67.2% vs. 48.8%; p=0.025), while dyspnea was strongly associated with age  $\geq$ 50 years (p=0.002), CVD (p=0.005), obesity (p=0.005), hypertension (p<0.001), and diabetes (p=0.001). Higher disease severity was also associated with age  $\geq$ 50 years, CVD, obesity, hypertension, and diabetes (p<0.001). Other associations are shown in table 2.

				Fever			Cough		Dyspnea			Oth	er sympton	m	Disease severity				
			No	Yes	p- value	No	Yes	p- value	No	Yes	p- value	No	Yes	p- value	Mild	Moderate	Severe	p- value	
Åio ⊂ <50 n		n	42	44	0.234	44	42	0.025	77	9	0.002	38	48	0.203	61	24	1	<0.001	
ateg	<30	%	48.84%	51.16%		51.16%	48.84%		89.53%	10.47%		44.19%	55.81%		70.93%	27.91%	1.16%		
° s s >50		n	25	39		21	43		44	19		35	29		10	32	22		
A	_50	%	39.06%	60.94%		32.81%	67.19%		69.84%	30.16%		54.69%	45.31%		15.63%	50.00%	34.38%		
scula: se	No	n	60	76	0.673	61	75	0.242	114	21	0.005	71	65	0.007	70	51	15	<0.001	
iova lisea	110	%	44.12%	55.88%		44.85%	55.15%		84.44%	15.56%		52.21%	47.79%		51.47%	37.50%	11.03%		
Card	Yes	n	7	7		4	10		7	7		2	12		1	5	8		

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		%	50.00%	50.00%		28.57%	71.43%		50.00%	50.00%		14.29%	85.71%		7.14%	35.71%	57.14%	
ant sm	No	n	65	80	1.000	64	81	0.389	119	25	0.046	72	73	0.367	70	55	20	0.044
ign pla		%	44.83%	55.17%		44.14%	55.86%		82.64%	17.36%		49.66%	50.34%		48.28%	37.93%	13.79%	
Aal	Vac	n	2	3		1	4		2	3		1	4		1	1	3	
N I	165	%	40.00%	60.00%		20.00%	80.00%		40.00%	60.00%		20.00%	80.00%		20.00%	20.00%	60.00%	
y	No	n	67	77	0.033	64	80	0.235	120	24	0.005	70	74	1.000	71	55	18	< 0.001
esit	140	%	46.53%	53.47%		44.44%	55.56%		83.33%	16.67%		48.61%	51.39%		49.31%	38.19%	12.50%	
Dbe	Vac	n	0	6		1	5		1	4		3	3		0	1	5	
•	105	%	0.00%	100.00%		16.67%	83.33%		20.00%	80.00%		50.00%	50.00%		0.00%	16.67%	83.33%	
uo	No	n	56	66	0.525	57	65	0.080	105	16	<0.001	62	60	0.271	64	48	10	< 0.001
tensi	NU	%	45.90%	54.10%		46.72%	53.28%		86.78%	13.22%		50.82%	49.18%		52.46%	39.34%	8.20%	
Dert		n	11	17		8	20		16	12		11	17		7	8	13	
HyJ	Yes	%	39.29%	60.71%		28.57%	71.43%		57.14%	42.86%		39.29%	60.71%		25.00%	28.57%	46.43%	
s	N.	n	54	71	0.419	58	67	0.090	107	17	0.001	65	60	0.068	70	46	9	< 0.001
bete	INO	%	43.20%	56.80%		46.40%	53.60%		86.29%	13.71%		52.00%	48.00%		56.00%	36.80%	7.20%	
iał	Vac	n	13	12		7	18		14	11		8	17		1	10	14	
П	165	%	52.00%	48.00%		28.00%	72.00%		56.00%	44.00%		32.00%	68.00%		4.00%	40.00%	56.00%	
50	No	n	55	65	0.565	48	72	0.099	96	23	0.739	58	62	0.870	56	46	18	0.878
kin	INO	%	45.83%	54.17%		40.00%	60.00%		80.67%	19.33%		48.33%	51.67%		46.67%	38.33%	15.00%	
mo	Vas	n	12	18		17	13		25	5		15	15		15	10	5	
S	res	%	40.00%	60.00%		56.67%	43.33%		83.33%	16.67%		50.00%	50.00%		50.00%	33.33%	16.67%	

As for COVID-19 complications, ICU admission was highly associated with age  $\geq$ 50 years, CVD, obesity, hypertension, and diabetes (p<0.001), and so were bacterial pneumonia and acute renal injury. Mechanical ventilation and cardiac arrest were associated with age  $\geq$ 50 years (p=0.005), obesity

(p<0.001), and hypertension (p=0.012). Viral pneumonitis was associated with age  $\geq$ 50 years, CVD, hypertension, and diabetes, while ARDS and hyperglycemia were associated with age  $\geq$ 50 years, obesity, hypertension, and diabetes (Table 3).

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Ι				ICU admission			Mechanical ventilation			Viral pneumonitis			ial pneu	ımonia	ARDS			CHF		
			No	Yes	p- value	No	Yes	p- value	No	Yes	p- value	No	Yes	p- value	No	Yes	p- value	No	Yes	p- value
~		n	85	1	< <u>0.001</u>	85	1	0.005	58	28	<0.001	82	4	<0.001	85	1	<0.001	86	0	0.180
tegor. urs)	<50	%	98.84 %	1.16%		98.84 %	1.16%		67.44 %	32.56 %		95.35 %	4.65%		98.84 %	1.16%		100.00 %	0.00%	
yea yea		n	43	21		56	8		13	51		46	18		53	11		62	2	
Age (	≥50	%	67.19 %	32.81 %		87.50 %	12.50 %		20.31 %	79.69 %		71.88 %	28.13 %		82.81 %	17.19 %		96.88%	3.13%	
ч		n	122	14	< 0.001	129	7	0.199	69	67	0.009	120	16	0.007	127	9	0.086	135	1	0.179
ascula ase	No	%	89.71 %	10.29 %		94.85 %	5.15%		50.74 %	49.26 %		88.24 %	11.76 %		93.38 %	6.62%		99.26%	0.74%	
iov		n	6	8		12	2		2	12		8	6		11	3		13	1	
Card	Yes	%	42.86 %	57.14 %		85.71 %	14.29 %		14.29 %	85.71 %		57.14 %	42.86 %		78.57 %	21.43 %		92.86%	7.14%	
		n	126	19	0.023	137	8	0.269	70	75	0.370	125	20	0.156	135	10	0.051	144	1	0.066
rnant lasm	No	%	86.90 %	13.10 %		94.48 %	5.52%		48.28 %	51.72 %		86.21 %	13.79 %		93.10 %	6.90%		99.31%	0.69%	
alig		n	2	3		4	1		1	4		3	2		3	2		4	1	
M	Yes	%	40.00 %	60.00 %		80.00 %	20.00 %		20.00 %	80.00 %		60.00 %	40.00 %		60.00 %	40.00 %		80.00%	20.00 %	
		n	127	17	< 0.001	139	5	< 0.001	70	74	0.213	126	18	0.004	137	7	<0.001	142	2	1.000
sity	No	%	88.19 %	11.81 %		96.53 %	3.47%		48.61 %	51.39 %		87.50 %	12.50 %		95.14 %	4.86%		98.61%	1.39%	
Dbe		n	1	5		2	4		1	5		2	4		1	5		6	0	
U	Yes	0⁄~	16.67	83.33		33.33	66.67		16.67	83.33		33.33	66.67		16.67	83.33		100.00	0.00%	
		/0	%	%		%	%		%	%		%	%		%	%		%	0.0070	
Ę		n	113	9	<0.001	118	4	0.012	63	59	0.027	112	10	<0.001	117	5	0.002	122	0	0.034
tensio	No	%	92.62 %	7.38%		96.72 %	3.28%		51.64 %	48.36 %		91.80 %	8.20%		95.90 %	4.10%		100.00 %	0.00%	
per		n	15	13		23	5		8	20		16	12		21	7		26	2	
Hyj	Yes	%	53.57 %	46.43 %		82.14 %	17.86 %		28.57 %	71.43 %		57.14 %	42.86 %		75.00 %	25.00 %		92.86%	7.14%	
		n	116	9	< <u>0.001</u>	119	6	0.173	68	57	<0.001	113	12	<0.001	119	6	0.006	124	1	0.306
oetes	No	%	92.80 %	7.20%		95.20 %	4.80%		54.40 %	45.60 %		90.40 %	9.60%		95.20 %	4.80%		99.20%	0.80%	
iat		n	12	13		22	3		3	22		15	10		19	6		24	1	
	Yes	%	48.00 %	52.00 %		88.00 %	12.00 %		12.00 %	88.00 %		60.00 %	40.00 %		76.00 %	24.00 %		96.00%	4.00%	

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		n	103	17	0.774	114	6	0.384	56	64	0.744	102	18	1.000	111	9	0.707	118	2	1.000
50	No	04	85.83	14.17		95.00	5 00%		46.67	53.33		85.00	15.00		92.50	7 50%		08 3304	1 67%	
kin		70	%	%		%	5.00%		%	%		%	%		%	7.50%		90.33%	1.0770	
mo		n	25	5		27	3		15	15		26	4		27	3		30	0	
S	Yes	0/	83.33	16.67		90.00	10.00		50.00	50.00		86.67	13.33		90.00	10.00		100.00	0.000/	
		%	0/-	0/-		0/-	0/-		0/-	0/-		0/-	0/-		0/-	0/-		0/-	0.00%	

		Cardiac arrhythmia		Cardiac ischemia			Cardiac arrest			Acute renal injur		injury	Live	r dysfu	nction	Hyp	perglyc	emia	Hyj	oglyce	emia		
			No	Yes	p- value	No	Yes	p- value	No	Yes	p- value	No	Yes	p- value	No	Yes	p- value	No	Yes	o-value	No	Yes	p- value
~		n	85	1	0.084	86	0	0.427	85	1	0.005	86	0	< 0.00	85	1	0.574	85	0	<0.001	85	0	0.010
egor. (s)	<50	%	98.84	1 1 60/		100.0	0.00		98.84	1.16		100.0	0.000/	1	98.84	1 1 60/		100.0	0.000/		100.00	0.000/	
e cate year			%	1.16%		0%	%		%	%		0%	0.00%		%	1.16%		0%	0.00%		%	0.00% -	
Age (	>50	n %	59 92.19	5		63 98.44	1 1.56		56 87.50	8 12.50		53 82.81	11		61 96.83	2		44 74.58	15 25.42		54 91.53	5	
			%	7.81%		%	%		%	%		%	%		%	3.17%		%	%		%	8.47%	
ılar		n	131	5	0.450	135	1	1.000	129	7	0.199	129	7	0.011	132	3	1.000	124	6	1.000	126	4	0.405
ascu ase	No	%	96.32 %	3.68%		99.26 %	0.74		94.85	5.15%		94.85	5.15%		97.78	2.22%		95.38 %	4.62%		96.92 %	3.08%	
liov		n	13	1		14	0		12	2		10	4		14	0		5	9		13	1	
Carc	Yes	%	92.86	7 1/1%		100.0	0.00		85.71	14.29		71.43	28.57		100.0	0.00%		35.71	64.29		92.86	7 1/1%	
<u> </u>			%	7.1470	1 000	0%	%	1 000	%	%	0.0.00	%	%	0.040	0%	0.0070	1.000	%	%	0.000	%	/.14/0	0.4.64
n t	No	n %	139	6	1.000	144	1	1.000	04.48	8	0.269	03 70	9	0.043	141	3	1.000	01.37	12	0.008	135	4	0.164
gnar lasn	110	70	%	4.14%		%	%		%	5.52%		93.19 %	6.21%		91.92 %	2.08%		%	8.63%		%	2.88%	
falig		n	5	0		5	0		4	1		3	2		5	0		2	3		4	1	
N L	Yes	%	100.0	0.00%		100.0	0.00		80.00	20.00		60.00	40.00		100.0	0.00%		40.00	60.00		80.00	20.00	
		n	0%			0%	%		%	%	<0.00	%	%		0%			%	%		%	%	
	No	п	141	3	0.001	144	0	0.040	139	5	<0.00 1	136	8	0.005	141	3	1.000	127	12	0.008	135	4	0.164
esity	110	%	97.92 %	2.08%		100.0 0%	0.00 %		96.53 %	3.47%		94.44 %	5.56%		97.92 %	2.08%		91.37 %	8.63%		97.12 %	2.88%	
0p		n	3	3		5	1		2	4		3	3		5	0		2	3		4	1	
	Yes	%	50.00	50.00		83.33	16.67		33.33	66.67		50.00	50.00		100.0	0.00%		40.00	60.00		80.00	20.00	
			%	%		%	%		%	%		%	%	<0.00	0%			%	%		%	%	<0.00
ion	No	n	120	2	0.011	122	0	0.187	118	4	0.012	119	3	<0.00	120	1	0.091	113	5	<0.001	118	0	<0.00 1
ens	110	%	98.36	1.64%		100.0	0.00		96.72	3.28%		97.54	2.46%		99.17	0.83%		95.76	4.24%		100.00	0.00%	
pert		n	% 24	4		27	% 1		23	5		20	8		26	2		% 16	10		% 21	5	
Hy	Yes	%	85.71	14.29		96.43	0.570/		82.14	17.86		71.43	28.57		92.86	-		61.54	38.46		80.77	19.23	
			%	%		%	3.57%		%	%		%	%		%	/.14%		%	%		%	%	
		n	121	4	0.262	125	0	0.167	119	6	0.173	120	5	0.003	121	3	1.000	121	1	<0.001	121	1	0.002
oetes	No	%	96.80 %	3.20%		100.0 0%	0.00%		95.20 %	4.80%		96.00 %	4.00%		97.58 %	2.42%		99.18 %	0.82%		99.18 %	0.82%	
Diał		n	23	2		24	1		22	3		19	6		25	0		8	14		18	4	
-	Yes	%	92.00	8.00%		96.00	4.00%		88.00	12.00		76.00	24.00		100.0	0.00%		36.36	63.64		81.82	18.18	
		n	% 115	5	1.000	% 119	1	1.000	% 114	%	0.384	% 113	% 7	0.231	<u>0%</u> 117	2	0.493	% 104	% 11	0 503	% 111	% 	1.000
âa	No	%	95.83		1.000	99.17	1	1.000	95.00		0.304	94.17		0.231	98.32	-	0.775	90.43		0.303	96.52	т а. кол	1.000
okin			%	4.17%		%	0.83%		%	5.00%		%	5.83%		%	1.68%		%	9.57%		%	3.48%	
Smc		n	29	1		30	0		27	3		26	4		29	1		25	4		28	1	
	I Vac	1 0/	06 67			100.0												0 6 0 1	10.70				
	res	%	90.07	3.33%		100.0	0.00%		90.00	10.00		86.67 %	13.33		96.67 %	3.33%		86.21 %	13.79		96.55 %	3.45%	

Patient mortality (death) was associated with age  $\geq$ 50 years (12.50% vs. 1.16%; p=0.005), obesity (66.67% vs. 3.47%; p<0.001), and having hypertension (17.86% vs. 3.28%; p=0.012), but not with CVD, diabetes, nor smoking (Figure

3). None of the patients who were asymptomatic or had no comorbidities died during follow-up. Younger patients were more likely to be discharged even if they were still PCR-positive (45.88% vs. 25.00%; p=0.012) (Table 4).

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DOI: 10.21275/SR201121021100



Figure 3: Mortality rate in different risk factor groups yrs: years

			Length of	hospital	Time t	o PCR		Patient w	as discharg	ged while	Б			
			sta	y	test cor	iversion		P	CR-positiv	e	Disease cuie			
			days	p-value	Days	p-value		No	Yes	p-value	No	Yes	p-value	
	<50	mean	14.79	0.926	22.26	0.561	n	46	39	0.012	1	85	0.005	
Age category	<30	SD	7.27		12.26		%	54.12%	45.88%		1.16%	98.84%		
(years)	>50	mean	15.27		19.82		n	42	14		8	55		
	≥30	SD	8.03		8.6		%	75.00%	25.00%		12.70%	87.30%		
	No	mean	15.39	0.037	21.92	0.021	n	78	51	0.211	7	128	0.201	
Cardiovascul	INU	SD	7.6		11.18		%	60.47%	39.53%		5.19%	94.81%		
ar disease	Vas	mean	11.14		14.67		n	10	2		2	12		
	168	SD	6.42		5.84		%	83.33%	16.67%		14.29%	85.71%		
	No	mean	14.96	0.850	21.39	0.608	n	84	53	0.297	8	136	0.271	
Malignant	INO	SD	7.54		11.072		%	61.31%	38.69%		5.56%	94.44%		
neoplasm	Yes	mean	16		18.25		n	4	0		1	4		
	105	SD	9.72		8.69		%	100.00%	0.00%		20.00%	80.00%		
	No	mean	15.2	0.089	21.33	0.950	n	88	51	0.140	5	138	< 0.001	
Obesity	110	SD	7.54		11.06		%	63.31%	36.69%		3.50%	96.50%		
Obesity	Vec	mean	10		19.5		n	0	2		4	2		
	105	SD	7.54		7.78		%	0.00%	100.00%		66.67%	33.33%		
	No	mean	14.83	0.639	21.72	0.503	n	70	48	0.086	4	117	0.012	
Hypertension	140	SD	7.7		11.53		%	59.32%	40.68%		3.31%	96.69%		
rrypertension	Ves	mean	15.71		19.17		n	18	5		5	23		
	103	SD	7.13		7.61		%	78.26%	21.74%		17.86%	82.14%		
	No	mean	15.3	0.155	22	0.106	n	71	48	0.117	6	119	0.160	
Diabetes	110	SD	7.28		11.41		%	59.66%	40.34%		4.80%	95.20%		
Diabetes	Ves	mean	13.44		17.33		n	17	5		3	21		
	103	SD	8.95		7.3		%	77.27%	22.73%		12.50%	87.50%		
	No	mean	15.07	0.823	21.2	0.952	n	73	41	0.413	6	113	0.386	
Smoking	140	SD	7.6		10.87		%	64.04%	35.96%		5.04%	94.96%		
Smoking	Vec	mean	14.7		21.7		n	15	12		3	27		
	Yes	SD	7.62		11.73		%	55.56%	44.44%		10.00%	90.00%		

 Table 4: Correlation between selected risk factors and disease course and outcome

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#### DOI: 10.21275/SR201121021100

#### 4. Discussion

In the present study, we report the clinical characteristics, comorbidities, disease course, and outcomes of patients affected by COVID-19 in Lebanon. Our results support previous data suggesting that comorbidities and different underlying conditions are generally associated with poorer COVID-19 prognosis. Our results are in line with those previously described in the literature, which highlighted the impact of comorbidities on COVID-19 severity [9-13].

In our sample of COVID-19 patients, age  $\geq$ 50 years was associated with higher rates of ICU admission, severe disease, need for mechanical ventilation, ARDS, and mortality. While older age is an independent risk factor for infection [15], older individuals are also more likely to have multiple comorbidities [16]. Several studies have reported that older individuals are at higher risk for COVID-19 infection and generally have a poorer prognosis when infected, having higher mortality and more severe illness [17]. Early reports from Wuhan have proposed that older age was an independent predictor of mortality in COVID-19, similar to previous observations in SARS and MERS [18].

Hypertension was the most common comorbidity in our cohort. and was highly associated with cardiac complications during hospitalization. Hypertension, but not other CVDs, was associated with need for mechanical ventilation, ARDS, CHF, arrhythmia, cardiac arrest, and higher mortality. Data from Pranata et al. showed similar results, where hypertension was associated with increased composite poor outcome, including mortality, severe COVID-19, ARDS, need for ICU care and disease progression in a pool of 6560 patients with COVID-19 [19]. One proposed explanation for this association is the upregulation of angiotensin-converting enzyme 2 (ACE2) expression in hypertensive patients; it is likely that SARS-CoV-2 uses ACE2 on the surface of epithelial cells as a receptor to attach and enter the host pneumocytes, in a similar manner to SARS-CoV. The up-regulation of ACE2 expression might facilitate viral entry and thus lead to higher disease severity and fatality [20].

Our data are in agreement with other studies that suggest that the prevalence of diabetes in COVID-19 patients is 15% (17% in our study) [21]. Our results suggest that diabetes was associated with higher risk for ICU admission, ARDS, and severe disease. Similar results have been found by Seiglie et al., who reported that compared to non-diabetics, a higher proportion of patients with diabetes was admitted to the ICU (42.1% vs. 29.8%) [22]. A meta-analyses on 6452 patients with COVID-19 has shown that diabetes was associated with poor outcome including mortality, severe disease, ARDS, and disease progression [23]. The compromised immune system of diabetic patients might put them at a higher risk of contracting respiratory infections as well as reduce their innate immune response to infection [24].

While obesity itself is a risk factor for other comorbidities including CVD, hypertension, and diabetes [25], our results have shown that it is also associated with poorer outcomes in COVID-19. One pooled analysis has shown that obese

patients are at a 46% higher risk of COVID-19 infection, a 113% higher risk of hospitalization, a 74% higher risk of ICU admission, and a 48% higher risk of mortality due to COVID-19 [26]. Evidence suggests that obesity impairs the individual's immune response through its modulatory effects on key immune cell populations [27], however, confirming this hypothesis in SARS-CoV-2 requires measuring blood immune cell counts, which was not part of our current study.

We did not find an association between smoking and COVID-19 complications nor outcomes. Conversely, a recent systematic review and meta-analysis has suggested that patients with any smoking history are at higher risk of severe disease and worse in-hospital mortality [28]. The small number of smokers in our study might have diluted this association. However, other studies have also reported no association between COVID-19 and smoking; one cross-sectional study in the UK has even found that active smoking was linked with decreased odds of a positive RT-PCR test result [29].

The mortality rate in our sample was 6%. At end of data collection (May 15, 2020), the global case-fatality rate was 6.85% based on WHO data [5]. The number of COVID-19 cases continue to increase: Lebanon has exceeded 54,000 cases by time of writing this article [6]. In this context, more studies are needed to determine which patient should be granted priority for hospitalization and to further understand who should be treated and how.

Our study had several limitations. The study was retrospective, sampled patients from one centeronly, and had a limited sample size. However, at the time of data collection, COVID-19 prevalence in Lebanon was low. In addition, the study center at that time was still the main center receiving cases in the country. Comorbidities in this study were self-reported, which leaves a place for possible under-reporting due to lack of awareness and/or the lack of diagnosis. Considering the cross-sectional nature of the study, causality could not be confirmed between different risk factors and outcome. The relative-risk of contracting COVID-19 could not be assessed in this study due to the lack of a control group.

This study is a first step in understanding disease risk factors and predictors of prognosis in the local COVID-19 population.All patients in the study were closely followed and had a known outcome, which adds value to the study's observations.

#### 5. Conclusions

In conclusion, older age and different comorbidities, especially hypertension, obesity, and diabetes should be thoroughly considered when treating patients for COVID-19. These factors should be evaluated when choosing treatment and when deciding if patients should be hospitalized for COVID-19. More studies are needed to explore the impact of comorbidities on contracting the disease, and more preventive measures should be considered in comorbid individuals.

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#### List of abbreviations:

ACE2: Angiotensin-Converting Enzyme 2 ARDS: Acute Respiratory Distress Syndrome BMI: Body Mass Index CHF: Congestive Heart Failure COVID-19: Coronavirus disease 2019 CVDs: CardioVascular Diseases ICU: Intensive Care Unit MERS: Middle East Respiratory Syndrome NCDs: Non-Communicable Diseases RHUH: Rafik Hariri University Hospital RT-PCR: Real-Time Polymerase-Chain-Reaction SARS: Severe Acute Respiratory Syndrome SARS-CoV-2: Severe Acute Respiratory Syndrome CoronaVirus 2

WHO: World Health Organization

#### 6. Declarations

#### Funding

The authors disclose receipt of the following financial support for the publication of this article: This work was supported by Novo Nordisk Pharma Sarl.

#### **Competing interests:**

The authors declare that they have no competing interests.

#### Ethics approval and consent to participate:

The study protocol was approved by the Institutional Review Board of the study site. Informed consent was not required considering the retrospective design of the study.

#### **Consent for publication**

Data records were de-identified and completely anonymous, so informed consent was waived.

#### Availability of data and materials

The datasets used and/or analyzed during the current study are available from the corresponding author on reasonable request.

#### Code availability

Not applicable

## Acknowledgements

None

#### **Authors' contributions**

Study conceptualization: SA, LO, HJ, MH, MSR, AE. Study design: SA, LO, MH, AE. Data collection: SA, LO, HJ. Formal analysis: ZEH. Data interpretation: all authors. Manuscript writing-original draft: ZEH. Manuscript writing-review and editing: SA, LO. All authors revised and approved the final manuscript.

#### References

- Zhou P, Yang XL, Wang XG, et al. A pneumonia [1] outbreak associated with a new coronavirus of probable bat origin. Nature. 2020;579(7798):270-273.
- Zhu N, Zhang D, Wang W, et al. A Novel Coronavirus [2] from Patients with Pneumonia in China, 2019. N Engl J Med. 2020;382(8):727-733.

- [3] Lu R, Zhao X, Li J, et al. Genomic characterisation and epidemiology of 2019 novel coronavirus: implications for virus origins and receptor binding. Lancet. 2020;395(10224):565-574.
- World Health Organization. Timeline of WHO's [4] response to COVID-19. https://www.who.int/newsroom/detail/29-06-2020-covidtimeline. Accessed 27 Sep 2020.
- World Health Organization. Coronavirus disease [5] (COVID-19) Situation Report 116. https://www.who.int/docs/defaultsource/coronaviruse/situation-reports/20200515-covid-19-sitrep-116.pdf?sfvrsn=8dd60956\_2. Accessed 27 Sep 2020.
- [6] Lebanese Ministry of Public Health. Coronavirus COVID-19 Lebanon Cases - Epidemiological Surveillance Program. https://www.moph.gov.lb/maps/covid19.php. Accessed 27 Sep 2020.
- Guan WJ, Ni ZY, Hu Y, et al. Clinical Characteristics [7] of Coronavirus Disease 2019 in China. N Engl J Med. 2020;382(18):1708-1720.
- Huang C, Wang Y, Li X, et al. Clinical features of [8] patients infected with 2019 novel coronavirus in Wuhan, China. Lancet. 2020;395(10223):497-506.
- [9] Singh AK, Gupta R, Ghosh A, Misra A. Diabetes in COVID-19: Prevalence, pathophysiology, prognosis and practical considerations. Diabetes Metab Syndr. 2020;14(4):303-310.
- [10] Zhao Q, Meng M, Kumar R, et al. The impact of COPD and smoking history on the severity of COVID-19: A systemic review and meta-analysis. J Med Virol. 2020.doi:10.1002/jmv.25889.
- [11] Liu K, Chen Y, Lin R, Han K. Clinical features of COVID-19 in elderly patients: A comparison with and middle-aged patients. J Infect. young 2020;80(6):e14-e18.
- [12] Wang Y, Lu X, Li Y, et al. Clinical Course and Outcomes of 344 Intensive Care Patients with COVID-19. Am J Respir Crit Care Med. 2020;201(11):1430-1434.
- [13] Zhang J, Wang X, Jia X, et al. Risk factors for disease severity, unimprovement, and mortality in COVID-19 patients in Wuhan, China. Clin Microbiol Infect. 2020;26(6):767-772.
- [14] World Health Organization. Noncommunicable Diseases (NCD) Country Profiles, Lebanon, 2018. https://www.who.int/nmh/countries/lbn\_en.pdf. Accessed 27 Sep 2020.
- [15] Shen N, Zhu Y, Wang X, et al. Characteristics and diagnosis rate of 5630 subjects receiving SARS-CoV-2 nucleic acid tests from Wuhan, China. JCI Insight. 2020;5(10):e137662.
- [16] Divo MJ, Martinez CH, Mannino DM. Ageing and the epidemiology of multimorbidity. Eur Respir J. 2014;44(4):1055-1068.
- [17] Bonanad C, García-Blas S, Tarazona-Santabalbina F, et al. The effect of age on mortality in patients with covid-19: a meta-analysis with 611,583 subjects. J Am Med Dir Assoc. 2020;21(7):915-918.
- [18] Zhou F, Yu T, Du R, et al. Clinical course and risk factors for mortality of adult inpatients with COVID-

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19 in Wuhan, China: a retrospective cohort study. The Lancet. 2020;395(10229):1054-1062.

- [19] Pranata R, Lim MA, Huang I, Raharjo SB, Lukito AA. Hypertension is associated with increased mortality and severity of disease in COVID-19 pneumonia: A systematic review, meta-analysis and meta-regression. J Renin Angiotensin Aldosterone Syst. 2020;21(2):1470320320926899.
- [20] Bosso M, Thanaraj TA, Abu-Farha M, Alanbaei M, Abubaker J, Al-Mulla F. The two faces of ace2: the role of ace2 receptor and its polymorphisms in hypertension and covid-19. Mol Ther Methods Clin Dev. 2020;18:321-327.
- [21] Hussain S, Baxi H, Chand Jamali M, Nisar N, Hussain MS. Burden of diabetes mellitus and its impact on COVID-19 patients: A meta-analysis of real-world evidence. Diabetes Metab Syndr. 2020;14(6):1595-1602.
- [22] Seiglie J, Platt J, Cromer SJ, et al. Diabetes as a risk factor for poor early outcomes in patients hospitalized with covid-19. Diabetes Care. Published online August 26, 2020.
- [23] Huang I, Lim MA, Pranata R. Diabetes mellitus is associated with increased mortality and severity of disease in COVID-19 pneumonia - A systematic review, meta-analysis, and meta-regression. Diabetes Metab Syndr. 2020;14(4):395-403.
- [24] Klekotka RB, Mizgała E, Król W. The etiology of lower respiratory tract infections in people with diabetes. Pneumonol Alergol Pol. 2015;83(5):401-408.
- [25] Jarolimova J, Tagoni J, Stern TA. Obesity: its epidemiology, comorbidities, and management. Prim Care Companion CNS Disord. 2013;15(5).
- [26] Popkin BM, Du S, Green WD, et al. Individuals with obesity and COVID-19: A global perspective on the epidemiology and biological relationships. Obesity Reviews. 2020;21(11).
- [27] van der Weerd K, Dik WA, Schrijver B, et al. Morbidly obese human subjects have increased peripheral blood CD4+ T cells with skewing toward a Treg- and Th2-dominated phenotype. Diabetes. 2012;61(2):401-408.
- [28] Reddy RK, Charles WN, Sklavounos A, Dutt A, Seed PT, Khajuria A. The effect of smoking on COVID-19 severity: A systematic review and meta-analysis. J Med Virol. Published online August 4, 2020.
- [29] de Lusignan S, Dorward J, Correa A, et al. Risk factors for SARS-CoV-2 among patients in the Oxford Royal College of General Practitioners Research and Surveillance Centre primary care network: a crosssectional study. The Lancet Infectious Diseases. 2020;20(9):1034-1042.