Brief Summary of Findings on the Association Between Interstitial Lung Diseases and Severe COVID-19 Outcomes

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Overall, 7 cohort studies¹⁻⁷ and 1 modeling study⁸ were retrieved that reported data on any interstitial lung disease and severe COVID-19 outcomes including mortality, intensive care unit (ICU), hospital admission, and ventilation. All studies were rated as having a moderate threat to internal validity.

- Evidence from six studies^{2-4, 6-8} (N=230,262) indicates the presence of underlying interstitial lung disease (ILD) is associated with an increase in mortality. Limited evidence suggests an increase in the risk of ICU admission² (N=89,530) and ventilation³ (N=483) with underlying ILD, however the evidence is insufficient to assure confidence in this assessment.
- Evidence from two cohort studies^{1,3} (N=8,256,644) conducted in Europe indicates the presence of underlying idiopathic pulmonary fibrosis is associated with an increase in mortality. Limited evidence from one study¹ (N=8,256,161) suggests increases in ICU admission and hospitalization with underlying idiopathic pulmonary fibrosis, however the evidence is insufficient to assure confidence in this assessment.
- Evidence from two cohorts^{1, 2} (N=8,345,691) conducted in Europe indicates the presence of underlying sarcoidosis is associated with increases in mortality and ICU admission. Evidence from two cohort studies^{1, 5} (N=8,256,982) conducted in the US and Europe suggests an increase in hospitalization with underlying sarcoidosis.
- Limited evidence from one cohort study¹ (N=8,256,161) conducted in Europe suggests underlying hypersensitivity pneumonitis, also called extrinsic allergic alveolitis, is associated with increases in mortality and hospitalization, however the evidence is insufficient to assure confidence in this assessment.
- Evidence from two cohorts^{1,3} (N=8,256,644) conducted in Europe indicates the presence of underlying interstitial lung diseases other than idiopathic pulmonary fibrosis or sarcoidosis is associated with an increase in mortality. Limited evidence from one study¹ (N=8,256,161) suggests an increase in hospitalization with underlying interstitial lung diseases other than idiopathic pulmonary fibrosis or sarcoidosis, however the evidence is insufficient to assure confidence in this assessment.
- Limited evidence from one cohort study³ (N=483) suggests that mortality is greater among patients with moderate or severe ILD when compared to patients with mild ILD. However, the evidence is insufficient to assure confidence in this assessment.

References

- 1. Aveyard P, Gao M, Lindson N, Hartmann-Boyce J, Watkinson P, Young D, et al. Association between pre-existing respiratory disease and its treatment, and severe COVID-19: a population cohort study. The Lancet Respiratory Medicine 2021;9(8):909-923.
- 2. Beltramo G, Cottenet J, Mariet A-S, Georges M, Piroth L, Tubert-Bitter P, et al. Chronic respiratory diseases are predictors of severe outcome in COVID-19 hospitalised patients: a nationwide study. European Respiratory Journal 2021:2004474.
- 3. Drake TM, Docherty AB, Harrison EM, Quint JK, Adamali H, Agnew S, et al. Outcome of hospitalization for COVID-19 in patients with interstitial lung disease an international multicenter study. American Journal of Respiratory and Critical Care Medicine 2020;202(12):1656-1665.
- 4. Estiri H, Strasser ZH, Klann JG, Naseri P, Wagholikar KB, Murphy SN. Predicting COVID-19 mortality with electronic medical records. npj Digital Medicine 2021;4(1).

- 5. Halalau A, Odish F, Imam Z, Sharrak A, Brickner E, Lee PB, et al. Epidemiology, Clinical Characteristics, and Outcomes of a Large Cohort of COVID-19 Outpatients in Michigan. Int J Gen Med 2021;14:1555-1563.
- 6. Kokturk N, Babayigit C, Kul S, Duru Cetinkaya P, Atis Nayci S, Argun Baris S, et al. The predictors of COVID-19 mortality in a nationwide cohort of Turkish patients. Respir Med 2021;183:106433.
- 7. Oh TK, Song IA. Impact of coronavirus disease-2019 on chronic respiratory disease in South Korea: an NHIS COVID-19 database cohort study. BMC Pulmonary Medicine 2021;21(1).
- 8. Mollalo A, Rivera KM, Vahabi N. Spatial statistical analysis of pre-existing mortalities of 20 diseases with COVID-19 mortalities in the continental United States. Sustainable Cities and Society 2021;67:102738.

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A. Methods

The aim of this review is to identify and synthesize the best available evidence on the association between interstitial lung disease (ILD) and severe COVID-19 in order to update the Centers for Disease Control and Prevention (CDC) website on underlying conditions and enable the creation of a provider-specific website with more rigorous information.

A.1. Literature Search

A list of search terms was developed to identify the literature most relevant to the population, exposure, comparator, and outcomes (PECO) question. Clinical experts and library scientists were consulted to develop a robust list of search terms. These terms were then incorporated into search strategies, and these searches were performed in OVID using the COVID-19 filter from the end of the previous literature search (December 2020). The detailed search strategies for identifying primary literature and the search results are provided in <u>Section B.1</u>. Subject matter experts supplemented the literature search results by recommending relevant references published before December 2020. References were included if retrieved by the chronic lung disease literature search and reported exposures and outcomes relevant to this review.

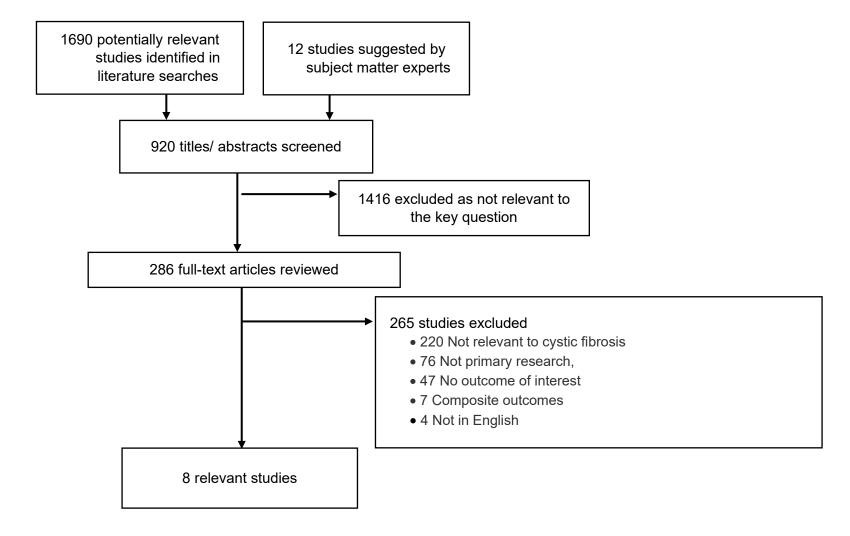
A.2. Study Selection

Titles and abstracts from references were screened by dual review (M.C., J.K.K., C.O., D.O.S., T.R., C.S., E.C.S., or M.W.). Full-text articles were retrieved if they were:

- 1. relevant to the PECO question;
- 2. primary research; and
- written in English.

Section B.2 presents the full list of exclusion criteria. The full texts of selected articles were then screened by two independent reviewers, and disagreements were resolved by discussion (J.K.K., C.O., D.O.S., K.T.R., C.S., E.C.S., or M.W.). After the full-text screening was complete, a bibliography of the articles selected for inclusion was vetted with subject matter experts. Additional studies suggested by the subject matter experts were screened for inclusion as described above. The results of the study selection process are depicted in Figure 1.

Figure 1. Results of the Study Selection Process



A.3. Data Extraction and Synthesis

Methodologic data and results of relevant outcomes from the studies meeting inclusion criteria were extracted into standardized evidence tables. Data and analyses were extracted as presented in the studies. For the purposes of this review, statistical significance was defined as $p \le 0.05$.

A.4. Aggregation of the Evidence

The internal validity associated with each study was assessed using scales developed by the Division of Healthcare Quality Promotion and scores were recorded in the evidence tables. Table 4 in <u>Section B.3.c.</u> includes the signaling questions used to assess the quality of each study design. The strength, magnitude, precision, consistency, and applicability of results were assessed for all comparators. The overall confidence in the evidence base is reported in the aggregation tables in <u>Section B.3.a.</u>

A.5 Reviewing and Finalizing the Systematic Review

Draft findings, aggregation tables, and evidence tables, are presented to CDC subject matter experts for review and input. Following further revisions, the summary will be published on the CDC website.

B. Systematic Literature Review Results

B.1. Search Strategies and Results

Table 1 Chronic Lung Disease search conducted March 17, 2021

#	Search History		
1	chronic lung disease		
2	respiratory system disease*		
3	reactive airway disease*		
4	emphysema		
5	chronic bronchitis		
6	COPD		
7	Chronic obstructive pulmonary disease		
8	Asthma *		
9	allergic asthma		
10	irritant asthma		
11	Interstitial lung disease		
12	Pulmonary fibrosis		
13	idiopathic pulmonary fibrosis		
14	nonspecific interstitial pneumonitis		
15	hypersensitivity pneumonitis		
16	sarcoidosis		
17	pneumoconiosis		
18	asbestosis		
19	coal workers pneumoconiosis		
20	silicosis		
21	bronchiectasis		
22	cystic fibrosis		
23	pulmonary vascular disease		
24	pulmonary hypertension		
25	bronchopulmonary dysplasia		
26	bronchiolitis obliterans		
27	asthma*		
28	reactive airway disease*		
29	CF		

30	1 or 2 or 3 or 4 or 5 or 6 or 7 or 8 or 9 or 10 or 11 or 12 or 13 or 14 or 15 or 16 or 17 or 18 or 19 or 20 or 21 or 22 or 23 or 24 or 25 or
	26 or 27 or 28 or 29
31	Limit 30 to covid-19
32	(202012* or 2021*).dt
33	(202012* or 2021*).dc
34	32 or 33
35	31 and 34
36	Deduplicate

B.2. Study Inclusion and Exclusion Criteria

Inclusion Criteria: Studies were included at the title and abstract screen if they:

- were relevant to the key question "what is the association between interstitial lung disease and severe COVID-19?";
- were primary research;
- were written in English (can be seen as [language] in title); and
- examined humans only.

Exclusion Criteria: Studies were excluded at full text review if they:

- were not available as full-text;
- were a conference abstract, poster, letter to the editor, or reply letter;
- examined lung transplant, cancer, or immunocompromised populations;
- reported autopsy results; and
- reported only composite outcome measures for "severe COVID-19".

B.3. Evidence Review: Interstitial Lung Disease and Severe COVID-19

B.3.a. Strength & Direction of Evidence

Table 2. Evidence examined for associations with interstitial lung disease and severe COVID-19

Outcome	Results
Mortality	Overall, the evidence suggests the presence of underlying interstitial lung disease (ILD) is associated with an
	increase in mortality. All six studies were found to have a moderate threat to internal validity.
	• Strength of Association: Six studies reported adjusted measures of association ranging from 1.027 to 5.27.
	Precision of Association: Three studies reported wide confidence intervals, one of which crossed the null.
	Three studies reported narrow confidence intervals.
	 Consistency of Association: All six studies reported measures of association in the same direction.

Outcome	Results
	 Applicability of Association: The populations and settings were directly applicable to the question. Five studies were conducted in high-income countries, four of which were in the US or Europe,
	Six studies suggested an increased risk of mortality in patients with ILD.
	 Five cohort studies^{2-4, 6, 7} (N=230,262) reported an effect measure suggesting that underlying ILD is associated with an increase in mortality in patients with COVID-19.
	 Of these studies, three^{3, 6, 7} (N=124,023) reported wide confidence intervals, decreasing confidence in the measure of effect. One study³ (N=483) had a small sample size. Another study¹ (N=122,040) which reported a confidence interval that spanned the null, was conducted in South Korea and reported a low rate of ILD among the study population (421/122,040 (0.3%)). The other study⁶ (N=1,500) was the only study reporting on ILD that was conducted in a middle-income country and reported few events. One study⁴ (N=16,709) reported an increase in the odds and risk of mortality among patients with interstitial pulmonary disease after adjusting for age, gender, and history of underlying conditions. One modeling study³ (N=NR) evaluating the geospatial distributions of COVID-19 mortality in the US reported an association between county-level COVID-19 case fatality rates and county-level, age-adjusted mortality due to ILD among counties with high COVID-19 mortality that were surrounded by counties with high COVID-19 mortalities. There was a protective association among counties with low COVID-19 mortality that were surround by counties with low COVID-19 mortalities.
ICU admission	The evidence is limited but suggests the presence of underlying ILD is associated with an increase in ICU admission. Aggregation indices cannot be measured for one study, which was found to have a moderate threat to internal validity.
	 One large retrospective cohort² (N=89,530) conducted in France reported an increase in odds of ICU admission among patients with ILD compared to patients with no underlying chronic lung disease. After adjusting for obesity, diabetes, hypertension, heart failure, atherosclerotic heart disease, sex, and age as a continuous variable, the increased odds of ICU admission remained.

Outcome	Results
Ventilation	The evidence is limited on the association between underlying ILD and the risk of ventilation but suggests a protective effect for underlying ILD. Aggregation indices cannot be measured for one study, which was found to have a moderate threat to internal validity. One prospective cohort study³ (N=483) evaluating outcomes among ILD patients hospitalized in Denmark, Germany, Italy, the Republic of Ireland, Spain, and the UK suggested a decrease in the prevalence of ventilation among patients with ILD compared to those without (3.7% vs. 9.0%, p=NR), however the study population size was small.

Table 3. Evidence examined for associations between idiopathic pulmonary fibrosis and severe COVID-19

	Describe
Outcome	Results
Mortality	The evidence suggests the presence of underlying idiopathic pulmonary fibrosis is associated with an increase in
	mortality. Both studies were found to have a moderate threat to interval validity (Aveyard 2021 and Drake 2020)
	 Strength of Association: Two studies reported adjusted measures of association ranging from 1.47 to 1.74.
	 Precision of Association: One study reported a wide confidence interval that did not cross the null. One
	study reported a narrow confidence interval.
	 Consistency of Association: Both studies reported measures of association in same direction.
	Applicability of Association: Both studies were implemented in Europe.
	Two cohort studies ^{1, 3} (N=8,256,644) reported that underlying idiopathic pulmonary fibrosis is associated with an
	increase in mortality in patients with COVID-19. In the larger study ¹ (N=8,256,161), this increase remained after
	adjusting for age and sex, other demographic factors, or other comorbidities. The smaller study ³ (N=483) reported
	results adjusting for age and comorbidity.
	 One study³ (N=483) reported a wide confidence interval and a small sample size, decreasing confidence in
	the measure of effect.
ICU admission	The evidence is limited but suggests the presence of underlying idiopathic pulmonary fibrosis associated with an
	increase in ICU admission. Aggregation indices cannot be measured for one study, which was found to have a
	moderate threat to internal validity.
	 One large retrospective cohort¹ (N=8,256,644) conducted in England suggested an increase in the hazard
	of ICU admission among patients with idiopathic pulmonary fibrosis. This increase remained after adjusting
	for age and sex, other demographic factors, or other comorbidities; however, confidence intervals were

Outcome	Results
	wide and crossed the null, decreasing confidence in the measure of effect. The study reported a small
	number of events and the hazard of ICU admission was calculated among the entire study population, not
	limited to those hospitalized with COVID-19.
Hospitalization	The evidence is limited but suggests the presence of underlying idiopathic pulmonary fibrosis associated with an
	increase in hospitalization. Aggregation indices cannot be measured for one study, which was found to have a
	moderate threat to internal validity.
	 One large retrospective cohort¹ (N=8,256,161) conducted in England reported an increase in the hazard of
	hospitalization among patients with idiopathic pulmonary fibrosis. This increase remained after adjusting
	for age and sex, other demographic factors, or other comorbidities.

Table 4. Evidence examined for associations between sarcoidosis and severe COVID-19

Outcome	Results
Mortality	 The evidence suggests the presence of underlying sarcoidosis is associated with an increase in mortality. Both studies were found to have a moderate threat to internal validity. Strength of Association: Two studies reported adjusted measures of association ranging from 1.41 to 2.11. Precision of Association: Both studies reported wide confidence intervals, one of which crossed the null. Consistency of Association: Both studies reported measures of association in same direction. Applicability of Association: Both studies were implemented in Europe. Two large cohort studies^{1, 2} (N=8,345,691) reported an increase in mortality among patients with sarcoidosis and both were found to have a moderate threat to internal validity. In one study¹, this increase in sarcoidosis remained after adjusting for age and sex, other demographic factors, or other comorbidities. In the other², the increase in pulmonary sarcoidosis remained after adjusting for age, sex, and other underlying medical conditions. Both studies reported wide confidence intervals and one¹ crossed the null, decreasing confidence in the measure of effect. Both studies reported a small number of events.
ICU admission	The evidence suggests the presence of underlying sarcoidosis is associated with an increase in ICU admission. Both studies were found to have a moderate threat to interval validity. • Strength of Association: Two studies reported adjusted measures of association ranging from 1.51 to 2.65. • Precision of Association: Both studies reported wide confidence intervals, one of which crossed the null. • Consistency of Association: Both studies reported measures of association in same direction. • Applicability of Association: Both studies were implemented in Europe.

Outcome	Results
	 Two cohorts^{1, 2} (N=8,345,691) suggested an increase in ICU admission among patients with sarcoidosis. In one study¹, this increase remained after adjusting for age and sex, other demographic factors, or other comorbidities. In the other², the increase remained after adjusting for age, sex, and other underlying medical conditions. Both studies reported wide confidence intervals and one¹ crossed the null, decreasing confidence in the measure of effect. Both studies reported a small number of events, and one¹ calculated the hazard of ICU admission among the entire study population, not limited to only those hospitalized with COVID-19.
Hospitalization	 The evidence is limited but suggests the presence of underlying sarcoidosis is associated with an increase in hospitalization. Both studies were found to have a moderate threat to internal validity. Strength of Association: One study reported an adjusted measure of association of 1.36 and one study did not report a measure of association. Precision of Association: One study reported a narrow confidence interval, and one study did not report a confidence interval. Consistency of Association: One study reported an association, and one found no association. Applicability of Association: Both studies were implemented in US or Europe.
	 Two studies suggested an increased risk of hospitalization in patients with sarcoidosis. One large retrospective cohort study¹ (N=8,256,161) conducted in England reported an increase in the hazard of hospitalization among patients with sarcoidosis. This increase remained after adjusting for age and sex, other demographic factors, or other comorbidities. One retrospective cohort study⁵ (N=821) in Michigan, US suggested an increase in the rate of hospitalization for people with sarcoidosis, however this difference was not statistically significant (2.3% vs. 0.5%, p=0.123). Of note, the prevalence of sarcoidosis was low in this study (6/821 (0.7%)).

Table 5. Evidence examined for associations between hypersensitivity pneumonitis (extrinsic allergic alveolitis) and severe COVID-19

Outcome	Results
Mortality	The evidence is limited but suggests the presence of underlying hypersensitivity pneumonitis, also called extrinsic
	allergic alveolitis, is associated with an increase in mortality. Aggregation indices cannot be measured for one
	study, which was found to have a moderate threat to internal validity. One retrospective cohort ¹ (N=8,256,161)
	conducted in England suggested an increase in the hazard of mortality among patients with extrinsic allergic

Outcome	Results
	alveolitis. This increase remained after adjusting for age and sex, other demographic factors, or other
	comorbidities.
	 This study reported a wide confidence interval that spanned the null, decreasing confidence in the
	measure of effect. This study reported a small number of events.
Hospitalization	The evidence is limited but suggests the presence of underlying hypersensitivity pneumonitis or extrinsic allergic alveolitis is associated with an increase in hospitalization. Aggregation indices cannot be measured for one study, and the study was found to have a moderate threat to internal validity. One retrospective cohort ¹ (N=8,256,161) conducted in England suggested an increase in the hazard of hospitalization among patients with extrinsic allergic alveolitis. This increase remained after adjusting for age and sex, other demographic factors, or other comorbidities.
	 This study reported a wide confidence interval that spanned the null, decreasing confidence in the measure of effect. This study reported a small number of events.

Table 6. Evidence examined for associations between other interstitial lung diseases and severe COVID-19

Outcome	Results
Mortality	The evidence suggests the presence of underlying other ILD excluding either idiopathic pulmonary fibrosis or sarcoidosis are associated with an increase in mortality. Both studies were found to have a moderate threat to internal validity. • Strength of Association: Two studies reported adjusted measures of association ranging from 1.50 to 2.05. • Precision of Association: Both studies reported wide confidence intervals that did not span the null. • Consistency of Association: Both studies reported measures of association in same direction. Applicability of Association: Both studies were implemented in Europe.
	 Two cohort studies^{1, 3} (N=8,256,644) reported an increase in mortality among patients with other ILD excluding either idiopathic pulmonary fibrosis or both idiopathic pulmonary fibrosis and sarcoidosis. One study³ (N=483) had a small study size and the other¹ (N=8,256,161) reported a small number of events.
Hospitalization	The evidence is limited but suggests the presence of underlying other ILD excluding either idiopathic pulmonary fibrosis or sarcoidosis are associated with an increase in hospitalization. Aggregation indices cannot be measured for one study, which was found to have a moderate threat to internal validity. One large retrospective cohort¹ (N=8,256,161) conducted in England reported an increase in the hazard of hospitalization among patients with other ILD, excluding idiopathic pulmonary fibrosis or sarcoidosis. This increase remained after adjusting for age and sex, other demographic factors, or other comorbidities. This

Outcome	Results
	study reported a wide confidence interval decreasing confidence in the measure of effect; however, it does
	not span the null. This study reported a small number of events.

Table 7. Evidence examined for associations between severity of interstitial lung disease and severe COVID-19

Outcome	Results
Mortality	The evidence is limited regarding the association between the severity of underlying ILD and the risk of mortality.
	Aggregation indices cannot be measured for one study, which was found to have a moderate threat to internal
	validity.
	 One prospective cohort study³ (N=483) conducted in Denmark, Germany, Italy, the Republic of Ireland,
	Spain, and the UK reported that mortality was greater among patients with moderate or severe ILD when
	compared to patients with mild ILD.
	 The analysis comparing severity levels was restricted to only those patients with ILD, limiting the
	confidence in the results as the sample size decreased.

B.3.b. Extracted Evidence

Table 8. Extracted Studies Reporting the Association between Interstitial Lung Disease and Severe COVID-19 Outcomes

Study	Population and Setting	Intervention	Definitions	Results
Author: Aveyard ¹	Population: N= 8,256,161	Health Condition Category:	Medical	Severe COVID-19:
		Chronic lung disease, Risk	Condition(s):	aHR: Adjusted Hazard Ratio for all
Year: 2021	Setting: 1,205 general practices	factors, Multiple comorbid	COPD: ND (Not	other respiratory diseases,
		conditions, Cancer	defined)	ethnicity, socioeconomic status,
Data Extractor:	Location: England, UK		Asthma: ND	region of England, body-mass
TR		Medical Condition, n/N	Bronchiectasis: ND	index, smoking status, non-
	Study dates: January 24, 2020-April 30, 2020	(%):	Cystic fibrosis: ND	smoking-related illness
Reviewer: DOS		COPD: 193,520/8,256,161	Sarcoidosis: ND	(hypertension, type 1 diabetes,
	Inclusion criteria:	(2.3%)	Extrinsic allergic	chronic liver disease, chronic
Study design:	All patients aged 20 years and older registered with one of	Asthma:	alveolitis: ND	neurological disease) and smoking-
Retrospective	the 1,205 general practices in England contributing to the	1,090,028/8,256,161	Idiopathic pulmonary	related illness (coronary heart
cohort study	QResearch database (version 44, uploaded March 23,	(13.2%)	fibrosis: ND	disease, stroke, atrial fibrillation,
	2020) were included in this population cohort study. Data	Bronchiectasis:	Other interstitial lung	type 2 diabetes, chronic kidney
Study Objective:	were linked to Public Health England's database of SARS-	41271/8,256,161 (0.5%)	diseases: ND	disease)
To assess	CoV-2 testing and English hospital admissions, ICU	Cystic fibrosis:	Lung cancer: ND	HR: Hazard Ratio
whether chronic	admissions, and deaths for COVID-19.	2081/8,256,161 (<1%)		
lung disease or		Sarcoidosis:	Severity Measure(s):	Mortality, n/N (%):
use of inhaled	Exclusion criteria:	17624/8,256,161 (0.2%)	Active asthma:	COPD:
corticosteroids	NR (Not reported)	Extrinsic allergic alveolitis:	having at least one	• aHR: 1.54 (95% CI: 1.42-1.67)
(ICS) affects the		2331/8,256,161 (<1%)		

Study	Population and Setting	Intervention	Definitions	Results
risk of		Idiopathic pulmonary	prescription for	• HR: 6.66 (95% CI: 6.19-7.18)
contracting		fibrosis: 7454/8,256,161	asthma medication	• COPD: 811/193,520 (0.4%)
severe COVID-		(0.1%)	Severe asthma: being	Asthma:
19.		Other interstitial lung	prescribed at least	• aHR: 0.99 (95% CI: 0.91-1.07)
		diseases: 5677/8,256,161	three different	• HR: 0.96 (95% CI: 0.89-1.04)
Internal validity		(0.1%)	classes of medication	• Asthma: 762/1,090,028 (0.1%)
assessment		Lung cancer:	for asthma in the	Cystic fibrosis:
(IVA) Score: 24		10792/8,256,161 (0.1%)	year before cohort	• Cystic fibrosis: 0/2081 (0%)
(moderate)			entry	Bronchiectasis:
		Control/Comparison group,		• aHR: 1.12 (95% CI: 0.94-1.33)
		n/N (%):	Clinical marker: NR	• HR: 4.77 (95% CI: 4.03-5.65)
		COPD: 8,062,641/8,256,161		 Bronchiectasis: 138/41,271
		(97.7%)	Treatment/	(0.3%)
		Asthma:	Associated Therapy:	Sarcoidosis:
		7,166,133/8,256,161	NR	• aHR: 1.41 (95% CI: 0.99-1.99)
		(86.6%)	Inhaled	
		Bronchiectasis:	corticosteroids (ICS):	• HR: 2.53 (95% CI: 1.79-3.58)
		8,214,890/8,256,161	commonly used	Sarcoidosis: 32/17,624 (0.2%) Futringia allorgia alloquitis:
		(99.5%)	treatments for	Extrinsic allergic alveolitis:
		Cystic fibrosis:	airways disease	• aHR: 1.56 (95% CI: 0.78-3.13)
		8,254,080/8,256,161		• HR: 4.82 (95% CI: 2.41-9.65)
		(99.9%)	Outcome Definitions:	Extrinsic allergic alveolitis:
		Sarcoidosis:	Mortality: confirmed	8/2,331 (0.3%)
		8,238,537/8,256,161	or suspected COVID-	Letter and the second second second second
		(99.8%) Extrinsic allergic	19 (ICD-10 codes	Idiopathic pulmonary fibrosis:
		alveolitis:	U07.1 and U07.2) on	• aHR: 1.47 (95% CI: 1.12-1.92)
		8,253,830/8,256,161	the death certificate,	• HR: 12.09 (95% CI: 9.42-15.53)
		(99.9%)	including deaths in	Idiopathic pulmonary fibrosis:
		Idiopathic pulmonary	and out of hospital	62/7,454 (0.8%)
		fibrosis:	ICU admission:	Other interstitial lung diseases:
		8,248,707/8,256,161	admission to an ICU	• aHR: 2.05 (95% CI: 1.49-2.81)
		(99.9%)	with severe COVID-	• HR: 11.37 (95% CI: 8.48-15.25)
		Other interstitial lung	19 (ICD-10 code	Other interstitial lung
		diseases:	U07.1 or U07.2) in	diseases: 45/5,677 (0.8%)
		8,250,484/8,256,161	Intensive Care	Lung cancer:
		(99.9%)	National Audit and	• aHR: 1.77 (95% CI: 1.37-2.29)
		Lung cancer:	Research Centre	• HR: 8.33 (95% CI: 6.46-10.74)
		8,245,369/8,256,161	(ICNARC) records	 Lung cancer: 60/10,792 (0.6%)
		(99.9%)	Intubation: NR	
			Ventilation: NR	ICU admission, n/N (%):
			Hospitalization:	COPD:
			positive test for	• aHR: 0.89 (95% CI: 0.68-1.17)
			SARS-CoV-2 and	• HR: 1.68 (95% CI: 1.29-2.18)
			appearing in the	• COPD: 59/193,520 (<0.1%)

Study	Population and Setting	Intervention	Definitions	Results
			Hospital Episode	Asthma:
			Statistics dataset as	 aHR: 1.08 (95% CI: 0.93-1.25)
			an in-patient within	• HR: 1.05 (95% CI: 0.91-1.22)
			30 days of that test	• 213/1,090,028 (<0.1%)
			or having an	Bronchiectasis:
			International	• aHR: 1.47 (95% CI: 0.91-2.36)
			Classification of	• HR: 2.37 (95% CI: 1.49-3.78)
			Diseases (ICD)-10	Bronchiectasis: 18/41,271
			code U07.1 for	(<0.1%)
			confirmed COVID-19	Sarcoidosis:
			or U07.2 for suspected COVID-19	• aHR: 1.51 (95% CI: 0.81-2.81)
			Non-elective	• HR: 3.06 (95% CI: 1.64-5.70)
			readmissions: NR	• Sarcoidosis: 10/17,624 (0.1%)
			readiniosions. Wit	Idiopathic pulmonary fibrosis:
			Comments: None	• aHR: 1.97 (95% CI: 0.85-4.55)
				• HR: 4.48 (95% CI: 2.01-9.99)
				Idiopathic pulmonary fibrosis: (7, 45,4 (9, 48))
				6/7,454 (0.1%)
				Hospitalization, n/N (%):
				COPD:
				• aHR: 1.54 (95% CI: 1.45-1.63)
				• HR: 5.09 (95% CI: 4.83-5.36)
				• COPD: 1,555/193,520 (0.8%)
				Asthma:
				• aHR: 1.18 (95% CI: 1.13-1.24)
				• HR: 1.22 (95% CI: 1.17-1.28)
				 Asthma: 2,266/1,090,028
				(0.2%)
				Bronchiectasis:
				• aHR: 1.34 (95% CI: 1.20-1.50)
				• HR: 4.53 (95% CI: 4.06-5.07)
				Bronchiectasis: 319/41,271
				(0.8%)
				Cystic fibrosis:
				• aHR: 1.55 (95% CI: 0.65-3.73)
				• HR: 1.37 (95% CI: 0.57-3.30)
				• Cystic fibrosis: 5/2,081 (0.2%)
				Sarcoidosis:
				• aHR: 1.36 (95% CI: 1.10-1.68)
				• HR: 2.74 (95% CI: 2.21-3.39)
				• Sarcoidosis: 84/17,624 (0.5%)
				Extrinsic allergic alveolitis:

Study	Population and Setting	Intervention	Definitions	Results
				• aHR: 1.35 (95% CI: 0.82-2.21)
				• HR: 3.97 (95% CI: 2.43-6.48)
				 Extrinsic allergic alveolitis:
				16/2,331 (0.7%)
				Idiopathic pulmonary fibrosis:
				• aHR: 1.59 (95% CI: 1.30-1.95)
				• HR: 8.80 (95% CI: 7.29-10.62)
				 Idiopathic pulmonary fibrosis: 110/7,454 (1.5%)
				Other interstitial lung diseases:
				• aHR: 1.66 (95% CI: 1.30-2.12)
				• HR: 7.57 (95% CI: 6.02-9.53)
				 Other interstitial lung
				diseases: 73/5,677 (1.3%)
				Lung cancer:
				• aHR: 2.24 (95% CI: 1.89-2.65)
				• HR: 7.92 (95% CI: 6.70-9.36)
				 Lung cancer: 139/10,792
				(1.3%)
				Severity of Condition:
				Mortality, n/N (%):
				Active asthma:
				• aHR: 1.05 (95% CI: 0.96-1.15)
				• HR: 1.62 (95% CI: 1.49-1.77)
				 Active asthma: 602/535,126
				(0.1%)
				Severe asthma:
				• aHR: 1.08 (95% CI: 0.98-1.19)
				• HR: 1.78 (95% CI: 1.62-1.95)
				• Severe asthma: 476/385,702
				(0.1%)
				ICU admission, n/N (%):
				Active asthma:
				• aHR: 1.34 (95% CI: 1.14-1.58)
				• HR: 1.73 (95% CI: 1.47-2.03)
				 Active asthma: 165/535,126
				(<0.1%)
				Severe asthma:
				• aHR: 1.30 (95% CI: 1.08-1.58)
				 HR: 1.79 (95% CI: 1.49-2.15)

Study	Population and Setting	Intervention	Definitions	Results
				• Severe asthma: 124/385,702
				(<0.1%)
				Hospitalization, n/N (%):
				Active asthma:
				• aHR: 1.26 (95% CI: 1.20-1.33)
				• HR: 1.95 (95% CI: 1.85-2.05)
				 Active asthma: 1,720/535,126
				(0.3%)
				Severe asthma:
				• aHR: 1.29 (95% CI: 1.22-1.37)
				• HR: 2.14 (95% CI: 2.02-2.26)
				• Severe asthma: 1,369/385,702 (0.4%)
				Duration of Condition: NR
				Treatment/ Associated Therapy:
				Mortality:
				ICS:
				• aHR: 1.15 (95% CI: 1.01-1.31)
				• HR: 2.63 (95% CI: 2.44-2.84)
				ICU admission:
				ICS:
				• aHR: 1.63 (95% CI: 1.18-2.24)
				• HR: 2.10 (95% CI: 1.78-2.46)
				Hospitalization:
				ICS:
				• aHR: 1.13 (95% CI: 1.03-1.23)
				• HR: 2.72 (95% CI: 2.60-2.85)
				Comorbid Conditions: NR
				Risk Markers:
				Mortality among COPD patients,
				n/N (%):
				Age: p<0.001
				40-59:
				• HR: 4.61 (95% CI: 2.93-7.26)
				• Died: 20/31,175 (0.06%)
				60-79:

Study	Population and Setting	Intervention	Definitions	Results
				• HR: 2.26 (95% CI: 1.99-2.57)
				• Died: 310/115,046 (0.30%)
				≥ 80:
				• HR: 1.28 (95% CI: 1.16-1.42)
				• Died: 481/46,194 (1.04%)
				Sex: p=0.005
				Women:
				• HR: 1.77 (95% CI: 1.56-2.00)
				• Died: 321/92,676 (0.35%)
				Men:
				• HR: 1.42 (95% CI: 1.28-1.57)
				• Died: 490/100,844 (0.49%)
				Ethnic group: p=0.009
				White:
				• HR: 1.55 (95% CI: 1.41-1.69)
				• Died: 635/161,376 (0.39%)
				Asian:
				• HR: 1.01 (95% CI: 0.70-1.44)
				• Died: 33/4,463 (0.74%)
				Black:
				• HR: 1.10 (95% CI: 0.70-1.73)
				• Died: 20/1,900 (1.05%)
				Chinese:
				• HR: 0.68 (95% CI: 0.09-5.05)
				• Died: <5/178 (2.81%)
				Other or not recorded:
				• HR: 1.89 (95% CI: 1.56-2.29)
				• Died: 122/25,603 (0.48%)
				Smoking status:
				Non-smoker: p=0.360
				• HR: 1.51 (95% CI: 1.27-1.79)
				• Died: 145/23,935 (0.61%)
				Ex-smoker:
				• HR: 1.52 (95% CI: 1.37-1.67)
				• Died: 547/104,638 (0.52%)
				Current smoker:
				• HR: 1.72 (95% CI: 1.37-2.14)
				• Died: 145/64,775 (0.22%)
				ICU admission among COPD
				patients, n/N (%):
				Age: p=0.466
				40-59:

Study	Population and Setting	Intervention	Definitions	Results
				• HR: 1.40 (95% CI: 0.69-2.83)
				• ICU admission: 8/31,175
				(0.03%)
				60-79:
				• HR: 0.90 (95% CI: 0.66-1.22)
				 ICU admission: 45/115,046
				(0.04%)
				≥ 80:
				• HR: 1.21 (95% CI: 0.51-2.85)
				 ICU admission: 6/46,194
				(0.01%)
				Sex: p=0.025
				Women:
				• HR: 1.43 (95% CI: 0.91-2.27)
				• ICU admission: 20/92,676
				(0.02%)
				Men:
				• HR: 0.74 (95% CI: 0.53-1.04)
				• ICU admission: 39/100,844
				(0.04%)
				Ethnic group: p=0.826
				White:
				• HR: 0.91 (95% CI: 0.66-1.26)
				• ICU admission: 42/161,376
				(0.03%)
				Asian:
				• HR: 0.74 (95% CI: 0.30-1.79)
				• ICU admission: 5/4,463 (0.11%)
				Black:
				• HR: 1.18 (95% CI: 0.44-3.20)
				• ICU admission: <5/1,900
				(0.26%)
				Other or not recorded:
				• HR: 0.79 (95% CI: 0.38-1.54)
				• ICU admission: 8/25,603
				(0.03%)
				Smoking status: p=0.732 Non-smoker:
				• HR: 0.76 (95% CI: 0.38-1.54)
				• ICU admission: 8/23,935
				(0.03%)
				Ex-smoker:
				• HR: 0.89 (95% CI: 0.65-1.21)

Study	Population and Setting	Intervention	Definitions	Results
				• ICU admission: 45/104,638
				(0.04%)
				Current smoker:
				• HR: 1.18 (95% CI: 0.51-2.72)
				• ICU admission: 6/64,775
				(0.01%)
				Hospitalization among COPD
				patients, n/N (%):
				Age: p<0.0001
				40-59:
				• HR: 2.57 (95% CI: 2.08-3.17)
				• Hospitalized: 91/31,175 (0.29%)
				60-79:
				• HR: 1.93 (95% CI: 1.78-2.09)
				 Hospitalized: 725/115,046
				(0.63%)
				≥ 80:
				• HR: 1.31 (95% CI: 1.21-1.42)
				• Hospitalized: 739/46,194 (1.60%)
				Sex: p=0.090
				Women:
				• HR: 1.63 (95% CI: 1.50-1.78)
				Hospitalized: 635/92,676
				(0.69%)
				Men:
				HR: 1.49 (95% CI: 1.38-1.60)Hospitalized: 920/100,844
				(0.91%)
				Ethnic group: p=0.0002 White:
				• HR: 1.55 (95% CI: 1.46-1.66)
				 Hospitalized: 1,223/161,376
				(0.76%)
				Asian:
				• HR: 0.98 (95% CI: 0.76-1.27)
				• Hospitalized: 61/4,463 (1.4%)
				Black:
				• HR: 1.17 (95% CI: 0.85-1.61)
				• Hospitalized: 39/1,900 (2.10%)
				Chinese:

Study	Population and Setting	Intervention	Definitions	Results
				• HR: 1.33 (95% CI: 0.33-5.45)
				 Hospitalized: <5/178 (2.81%)
				Other or not recorded:
				• HR: 1.83 (95% CI: 1.59-2.10)
				 Hospitalized: 230/25,603
				(0.90%)
				Smoking status: p=0.0002
				Non-smoker:
				• HR: 1.37 (95% CI: 1.21-1.56)
				 Hospitalized: 253/23,935
				(1.06%)
				Ex-smoker:
				• HR: 1.51 (95% CI: 1.41-1.62)
				 Hospitalized:1,031/104,638
				(0.99%)
				Current smoker:
				• HR: 1.94 (95% CI: 1.69-2.23)
				 Hospitalized: 265/64,775
				(0.41%)
				Mortality among asthma patients,
				n/N (%):
				Age: p=0.001
				20-39:
				• HR: 2.11 (95% CI: 1.00-4.42)
				• Died: 9/459,751 (<0.01%)
				40-59:
				• HR: 1.27 (95% CI: 0.95-1.69)
				• Died: 54/352,853 (0.02%)
				60-79:
				• HR: 1.09 (95% CI: 0.96-1.24)
				• Died: 275/218,881 (0.13%)
				≥ 80:
				• HR: 0.85 (95% CI: 0.77-0.95)
				• Died: 424/58,543 (0.72%)
				Sex: p=0.628
				Women:
				• HR: 0.97 (95% CI: 0.86-1.08)
				• Died: 362/571,497 (0.06%)
				Men:
				• HR: 1.01 (95% CI: 0.90-1.12)
				• Died: 400/518,531 (0.08%)
				Ethnic group: p=0.448

Study	Population and Setting	Intervention	Definitions	Results
				White:
				• HR: 0.96 (95% CI: 0.87-1.05)
				• Died: 514/84.083 (0.61%)
				Asian:
				• HR: 1.00 (95% CI: 0.78-1.27)
				• Died: 80/68,014 (0.12%)
				Black:
				• HR: 0.97 (95% CI: 0.72-1.32)
				• Died: 48/2,835 (1.69%)
				Chinese:
				• HR: 0.95 (95% CI: 0.22-4.03)
				• Died: <5/3,503 (0.14%)
				Other or not recorded:
				• HR: 1.14 (95% CI: 0.94-1.38)
				• Died: 118/206,076 (0.06%)
				Smoking status: p=0.396
				Non-smoker:
				• HR: 0.99 (95% CI: 0.89-1.10)
				• Died: 374/624,797 (0.06%)
				Ex-smoker:
				• HR: 0.99 (95% CI: 0.88-1.11)
				• Died: 341/257,566 (0.13%)
				Current smoker:
				• HR: 0.91 (95% CI: 0.65-1.26)
				• Died: 40/193,373 (0.02%)
				ICU admission among asthma
				patients, n/N (%):
				Age: p=0.015
				20-39:
				• HR: 2.16 (95% CI: 1.40-3.33)
				 ICU admission: 28/459,751
				(0.01%)
				40-59:
				• HR: 1.03 (95% CI: 0.81-1.30)
				• ICU admission: 78/352,853
				(0.02%)
				60-79:
				• HR: 1.03 (95% CI: 0.83-1.27)
				• ICU admission: 103/218,881
				(0.05%)
				≥ 80:
				• HR: 0.61 (95% CI: 0.22-1.69)

Study	Population and Setting	Intervention	Definitions	Results
				• ICU admission: <5/58,543
				(0.01%)
				Sex: p=0.021
				Women:
				• HR: 1.36 (95% CI: 1.07-1.74)
				• ICU admission: 84/571,497
				(0.01%)
				Men:
				• HR: 0.95 (95% CI: 0.79-1.15)
				• ICU admission: 129/518,531 (0.02%)
				Ethnic group: p=0.230
				White:
				• HR: 1.18 (95% CI: 0.97-1.43)
				• ICU admission: 124/784,083 (0.02%)
				Asian:
				• HR: 0.94 (95% CI: 0.65-1.34)
				• ICU admission: 34/68,014
				(0.05%)
				Black:
				• HR: 1.33 (95% CI: 0.88-2.02)
				• ICU admission: 26/28,352
				(0.09%)
				Chinese:
				• HR: 0.99 (95% CI: 0.13-7.56)
				• ICU admission: <5/3,503 (0.14%)
				Other or not recorded:
				• HR: 0.77 (95% CI: 0.52-1.13)
				• ICU admission: 28/206,076 (0.01%)
				Smoking status: p=0.725
				Non-smoker:
				• HR: 1.06 (95% CI: 0.88-1.28)
				• ICU admission: 124/624,797
				(0.02%)
				Ex-smoker:
				• HR: 1.14 (95% CI: 0.90-1.45)
				• ICU admission: 81/257,566
				(0.03%)
				Current smoker:
				• HR: 0.79 (95% CI: 0.36-1.73)

Study	Population and Setting	Intervention	Definitions	Results
				• ICU admission: 7/193,373
				(<0.01%)
				Hospitalization among asthma
				patients, n/N (%):
				Age: p<0.0001
				20-39:
				• HR: 1.59 (95% CI: 1.37-1.86)
				• Hospitalized: 206/459,751 (0.04%)
				40-59:
				• HR: 1.43 (95% CI: 1.29-1.57)
				• Hospitalized: 507/352,853
				(0.14%)
				60-79:
				• HR: 1.19 (95% CI: 1.10-1.28)
				• Hospitalized: 847/218,881
				(0.39%)
				≥ 80:
				• HR: 0.93 (95% CI: 0.86-1.00)
				• Hospitalized: 706/58,543 (1.21%)
				Sex: p=0.0001
				Women:
				• HR: 1.29 (95% CI: 1.21-1.37)
				 Hospitalized: 1,238/571,497
				(0.22%)
				Men:
				• HR: 1.08 (95% CI: 1.01-1.15)
				 Hospitalized: 1,028/518,531
				(0.20%)
				Ethnic group: p=0.868
				White:
				• HR: 1.20 (95% CI: 1.14-1.27)
				 Hospitalized: 1,539/748,083
				(0.21%)
				Asian:
				• HR: 1.16 (95% CI: 1.01-1.33)
				 Hospitalized: 252/68,014
				(0.37%)
				Black:
				• HR: 1.10 (95% CI: 0.93-1.31)
				 Hospitalized: 149/28,352
				(0.53%)

Study	Population and Setting	Intervention	Definitions	Results
				Chinese:
				• HR: 1.07 (95% CI: 0.43-2.67)
				 Hospitalized: 5/3,503 (0.14%)
				Other or not recorded:
				• HR: 1.15 (95% CI: 1.02-1.29)
				 Hospitalized: 321/206,076
				(0.16%)
				Smoking status: p=0.286
				Non-smoker:
				• HR: 1.18 (95% CI: 1.11-1.25)
				• Hospitalized: 1,205/624,797 (0.19%)
				Ex-smoker:
				• HR: 1.16 (95% CI: 1.07-1.25)
				 Hospitalized: 868/257,566
				(0.34%)
				Current smoker:
				• HR: 1.32 (95% CI: 1.12-1.55)
				• Hospitalized: 182/193,373
				(0.09%)
				Long-term Sequelae: NR
Author: Beltram	Population: N= 89,530 COVID-19 patients	Health Condition Category:	Medical Condition(s):	Severe COVID-19:
o ²		Chronic heart	Pulmonary	aOR: Adjusted odds ratio; adjusted
	Setting: Public and private hospitals	disease, Chronic lung	hypertension: ICD-10	for obesity, diabetes, hypertension,
Year: 2021		disease, Cancer	1270	heart failure, atherosclerotic
	Location: France		Any CRD: includes	heart disease, sex, and age as a
Data		Medical Condition, n/N	chronic respiratory	continuous variable
Extractor: MC	Study dates: COVID-19 cohort: March 1 - April 30, 2020	(%):	failure, asthma,	OR: Odds ratio
		Pulmonary hypertension:	COPD, ILD,	Mortality, n/N (%):
Reviewer: DOS	Inclusion criteria: For the COVID-19 cohort, all patients	341/89,530 (0.38%)	pulmonary	Pulmonary hypertension:
	hospitalized for COVID-19 during the study dates were	Any CRD: 14351/89530	hypertension,	
Study design:	included and identified by the primary, related, or	(16.0%)	sarcoidosis, CF, and	• aOR: 1.24 (95% CI: 0.91- 1.67)
Cohort	associated diagnoses by the ICD-10 codes U0710, U0711,	Chronic respiratory	lung cancer	,
	U0712, U0714 or U0715, regardless of their	failure: 1433/89,530	Chronic respiratory	• OR: 2.01 (95% CI: 1.50- 2.68)
Study	age. Data obtained from the	(1.60%)	failure: ICD-10 J961	·
Objective: To	national Programme de Medicalisation des Systemes d'Info	Sleep apnea: 3581/89,530	Sleep apnea: ICD-10	Pulmonary hypertension: 96/341 (28.2%)
describe and	rmation (PMSI) database.	(4.00%)	G473	
compare chronic		Chronic obstructive	COPD: ICD-10 J40,	• No CRD: 11222/75179 (14.93%)
respiratory	Exclusion criteria:	pulmonary disease	J41, J42, J44	• p<0.05
diseases (CRD) in	NR	(COPD): 4866/89,530	Emphysema: ICD-10	Any CRD:
hospitalized		(5.44%)	J43, J982	7.11, CRD.

respiratory complications for COVID-19 patients with CRD to COVID-19 patients without CRD and to influenza patients. (ILD): 1611/89,530 (1.80%) Lung cancer: ICD-10 1.59) Pulmonary C34, C45 ● OR: 2.10 (95 Severity ● Chronic responsion group, n/N (%): Treatment/ Sleep apnea:	
suffering from COVID-19 or influenza (2018- 2019 season), and to describe and compare respiratory complications for COVID-19 patients with CRD to COVID-19 patients without CRD and to influenza patients. Control/Comparison group, influenza patients. Control/Comparison group, patients. Control/Comparison group, n/N (%): NR Control (23.43%) CF: ICD-10 E840 (23.43%) No CRD: 112 (95.112 (14.93%) Control/Comparison group, n/N (%): NR No CRD: 121 (95.112 (95.112 (14.93%) No CRD: 1579/89530 (1.06) No CRD: 1579/89530 (1.06) OR: 1.12 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.112 (95.	363/14351
influenza (2018- 2019 season), and to describe and compare respiratory complications for COVID-19 patients with CRD to COVID-19 patients without CRD and to influenza patients. ILD: ICD-10 J84 Pulmonary 20/89,530 (0.02%) sarcoidosis: ICD-10 Interstitial lung disease (ILD): 1611/89,530 (1.80%) Pulmonary Sarcoidosis: 159/89,530 (0.18%) COVID-19 patients without CRD and to influenza patients. IVA Score: 24 (moderate) (14.93%) (14.93%) ILD: ICD-10 J84 Pulmonary Sarcoidosis: ICD-10 Chronic respiratory fa ■ QR: 1.30 (9 1.59) ■ OR: 2.10 (95 2.54) ■ OR: 2.10 (95 2.54) ■ Chronic respiratory fa ■ QR: 0.18%) Lung cancer: ICD-10 1.59) ■ OR: 2.10 (95 2.54) ■ Chronic respiratory fa ■ QR: 1.30 (9 1.59) ■ OR: 2.10 (95 2.54) ■ OR: 2.10 (95 2	•
2019 season), and to describe and compare respiratory complications for COVID-19 patients with CRD to COVID-19 patients without CRD and to influenza patients. CYSTIC fibrosis (CF): 20/89,530 (0.02%) sarcoidosis: ICD-10 Chronic respiratory fallows (1.09%) Severity Sarcoidosis: ISD/89,530 (0.02%) Lung cancer: ICD-10 (1.59) (0.18%) Severity Sarcoidosis: 159/89,530 (0.18%) Severity Severity Sarcoidosis: 159/89,530 (0.18%) Severity Sarcoidosis: ISD/89,530 (0.18%) Severity Sarcoidosis: ISD-10 (1.59) (0.18%) Severity Sarcoidosis: ISD-10 (1.59) (0.18%) Severity Sarcoidosis: ISD/89,530 (0.18%) Severity Sarcoido	222/75179
and to describe and compare respiratory complications for COVID-19 patients with CRD and to influenza patients. IVA Score: 24 (moderate) and to describe and compare respiratory (and compare respiratory (ILD): 1611/89,530 (0.02%) Interstitial lung disease (ILD): 1611/89,530 (1.80%) Pulmonary (ILD): 1611/89,530 (1.80%) Pulmonary (ILD): 1611/89,530 (1.80%) Pulmonary (C34, C45	
and compare respiratory (ILD): 1611/89,530 (1.80%)	
respiratory complications for COVID-19 patients with CRD to COVID-19 patients without CRD and to influenza patients. IVA Score: 24 (moderate) COVID-19 COVID-1	ilure:
complications for COVID-19 patients with CRD to COVID-19 patients without CRD and to influenza patients. Pulmonary sarcoidosis: 159/89,530 (0.18%) CSeverity (0.18%) Everity (0.18%) Chronic resp. failure: 413/1433 (28%) Measure(s): NR failure: 413/1433 (28%) No CRD: 112 (14.93%) No CRD: 112 (14.93%) Treatment/ No CRD: 75179/89530 Associated Therapy, n/N (%): NR aOR: 0.95 (98%) 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00	95% CI: 1.06-
COVID-19 patients with CRD to COVID-19 patients without CRD and to influenza patients. (0.18%) Severity (0.18%) Measure(s): NR € Chronic resp. failure: 413/1433 (28 € failure: 413/1433 (28 € failure: 413/1433) € No CRD: 112 € failure: 413/1433 (28 € failure: 413/1433) € No CRD: 112 € failure: 413/1433 (28 € failure: 413/1433) € No CRD: 112 € failure: 413/1433 (28 € failure: 413/1433) € No CRD: 112 € failure: 413/1433 (28 € failure: 413/1433) € No CRD: 112 € failure: 413/1433 (28 € failure: 413/1433) € No CRD: 112 € failure: 413/1433 (28 € failure: 413/1433) € No CRD: 112 € failure: 413/1433 (28 € failure: 413/1433) € No CRD: 112 € failure: 413/1433 (28 € failure: 413/1433) € No CRD: 112 € failure: 413/1433 (28 € failure: 413/1433) € No CRD: 112 € failure: 413/1433 (28 € failure: 413/1433) € No CRD: 112 € failure: 413/1433 (28 € failure: 413/1433) € No CRD: 112 € failure: 413/1433 (28 € failure: 413/1433) € No CRD: 112 € failure: 413/1433 (28 € failure: 413/1433) € No CRD: 112 € failure: 413/1433 (28 € failure: 413/1433) € No CRD: 112 € failure: 413/1433 (28 € failure: 413/1433) € No CRD: 112 € failure: 413/1433 (28 € failure: 413/1433) € No CRD: 112 € failure: 413/1433 (28 € f	
patients with CRD to COVID-19 patients without CRD and to influenza patients. (0.18%) Lung cancer: 977/89,530 (1.09%) Severity Measure(s): NR failure: 413/1433 (28 failure: 413/1433 (28 failure: 413/1433) (28 fail	5% CI: 1.74-
CRD to COVID-19 patients without CRD and to influenza patients. Lung cancer: 977/89,530 (1.09%) Measure(s): NR failure: 413/1433 (28	
patients without CRD and to influenza patients. (1.09%) Clinical marker: NR	piratory
CRD and to influenza patients. Control/Comparison group, n/N (%): NR Clinical marker: NR (14.93%) IVA Score: 24 (moderate) No CRD: 75179/89530 (84.0%) Treatment/ Associated Therapy, n/N (%): NR ■ OR: 0.95 (91.12 (95.12))	8.8%)
influenza patients. Control/Comparison group, n/N (%): Treatment/ Sleep apnea: IVA Score: 24 (moderate) No CRD: 75179/89530 (84.0%) Associated Therapy, n/N (%): NR ■ OR: 0.95 (90.05)	222/75179
patients. n/N (%): Treatment/ Sleep apnea: IVA Score: 24 (moderate) (84.0%) Associated Therapy, n/N (%): NR ■ aOR: 0.95 (9 1.06) ■ OR: 1.12 (95	
No CRD: 75179/89530 IVA Score: 24 (moderate) No CRD: 75179/89530 (84.0%) Associated Therapy, n/N (%): NR 1.06) OR: 1.12 (95)	
IVA Score: 24 (moderate) (84.0%) n/N (%): NR 1.06) • OR: 1.12 (95)	
(moderate) • OR: 1.12 (95	95% CI: 0.85-
4.25)	
Outcome 1.25)	5% CI: 1.02-
Definitions: ● Sleep apnea	a: 672/3581
Mortality: in- (18.8%)	
hospital mortality • No CRD: 112	222/75179
during (14.93%)	
hospitalization • p<0.05	
ICU admission: ND COPD:	
	95% CI: 1.06-
Ventilation: NR 1.22)	
Hospitalization: NR OR: 1.72 (95	5% CI: 1.61-
Non-elective 1.84)	
readmissions: NR	9/4886 (25.3
Comments: none • No CRD: 112	222/75179
(14.93%)	
1.22)	DE% CI+ 0 92
● OR: 1.18 (95	95% CI: 0.83-
1.42)	

Study	Population and Setting	Intervention	Definitions	Results
				• Emphysema: 312/1426
				(21.8%)
				• No CRD: 11222/75179
				(14.93%)
				Asthma:
				• aOR: 0.82 (95% CI: 0.71-
				0.94)
				• OR: 0.51 (95% CI: 0.45-
				0.58)
				 Asthma: 310/3273
				(9.5%)
				• No CRD: 11222/75179
				(14.93%)
				• p<0.05
				Cystic fibrosis:
				• 0/20 (0.0%)
				ILD:
				• aOR: 1.20 (95% CI: 1.05-
				1.28)
				• OR: 1.41 (95% CI: 1.24-
				1.61)
				• ILD: 363/1611 (22.5%)
				• No CRD: 11222/75179
				(14.93%)
				• p<0.05
				Pulmonary sarcoidosis:
				• aOR: 2.11 (95% CI: 1.36-
				3.26)
				• OR: 1.38 (95% CI: 0.92-
				2.09)
				• Pulmonary
				sarcoidosis: 32/159 (20.1%)
				• No CRD: 11222/75179
				(14.93%)
				Lung cancer:
				• aOR: 3.67 (95% CI: 3.20-
				4.21)
				• OR: 3.64 (95% CI: 3.20-
				4.14)
				• Lung cancer: 402/977
				(41.2%)

Study	Population and Setting	Intervention	Definitions	Results
				No CRD: 11222/75179
				(14.93%)
				• p<0.05
				ICU admission, n/N (%):
				Pulmonary hypertension:
				• aOR: 1.73 (95% CI: 1.27-
				2.37)
				• OR: 1.97 (95% CI: 1.46-
				2.65)
				Pulmonary
				hypertension: 97/341 (28.5%)
				• No CRD: 12119/75179
				(16.12%)
				• p<0.05
				Any CRD:
				• Any CRD: 2985/14351 (20.80%)
				• No CRD: 12119/75179 (16.12%)
				• p<0.0001
				Chronic respiratory failure:
				• aOR: 1.03 (95% CI: 0.81-
				1.30)
				• OR: 1.18 (95% CI: 0.94-
				1.49)
				• Chronic
				respiratory failure: 320/1433
				(22.3%)
				• No CRD: 12119/75179
				(16.12%)
				Sleep apnea:
				• aOR: 1.39 (95% CI: 1.27-
				1.53)
				• OR: 2.74 (95% CI: 2.52-
				2.98)
				• Sleep apnea: 1172/3581
				(32.7%)
				• No CRD: 12119/75179
				(16.12%)
				• p<0.05
				COPD:

Study	Population and Setting	Intervention	Definitions	Results
				• aOR: 1.16 (95% CI: 1.07-
				1.26)
				• OR: 1.47 (95% CI: 1.37-
				1.58)
				• COPD: 986/4866
				(20.6%)
				• No CRD: 12119/75179
				(16.12%)
				• p<0.05
				Emphysema:
				• aOR: 1.83 (95% CI: 1.56-
				2.16)
				• OR: 2.09 (95% CI: 1.78-
				2.45)
				• Emphysema: 405/1426
				(28.4%)
				• No CRD: 12119/75179
				(16.12%)
				• p<0.05
				Asthma:
				• aOR: 1.23 (95% CI: 1.12-
				1.36)
				• OR: 1.35 (95% CI: 1.23-
				1.48)
				• Asthma: 640/3273
				(19.6%)
				• No CRD: 12119/75179
				(16.12%)
				p<0.05Cystic fibrosis:
				• aOR: 0.60 (95% CI: 0.1- 2.60)
				• OR: 0.63 (95% CI: 0.15- 2.73)
				• Cystic fibrosis: 2/20
				(10.0%)
				• No CRD: 12119/75179
				(16.12%)
				(10.12%) ILD:
				• aOR: 2.42 (95% CI: 2.14-
				2.72)

				 OR: 2.77 (95% CI: 2.47-3.11) ILD: 527/1611 (32.7%) No CRD: 12119/75179 (16.12%) p<0.05 Pulmonary sarcoidosis: aOR: 2.65 (95% CI: 1.83-3.84) OR: 2.94 (95% CI: 2.07-4.19) Pulmonary sarcoidosis: 53/159 (33.3%) No CRD: 12119/75179 (16.12%)
				 p<0.05 Lung cancer: aOR: 0.77 (95% CI: 0.63-0.94) OR: 0.78 (95% CI: 0.64-0.94) Lung cancer: 117/977 (12.0%) No CRD: 12119/75179 (16.12%) p<0.05 Severity of Condition: NR Duration of Condition: NR Treatment/ Associated Therapy: NR Comorbid Conditions: NR Risk Markers: NR Long-term Sequelae: NR
Author: Drake ³ Pop	opulation: N=483 patients	Health Condition Category: Chronic heart	Medical Condition(s):	Severe COVID-19:

Study	Population and Setting	Intervention	Definitions	Results
Year: 2020	Setting: tertiary ILD centers and secondary care hospitals	disease, Diabetes, Chronic Lung Disease, Risk	Chronic Heart Disease: ND	aHR: Adjusted hazard ratio (95% CI); matched adjusted analysis
Data Extractor: JKK	Location: Denmark, Germany, Italy, Republic of Ireland, Spain, and the UK	Factors, Multiple Comorbid Conditions	Hypertension: ND Diabetes: type 1 or	Mortality, n/N (%):
			Diabetes: type 1 or type 2 ILD: any of the following conditions: chronic hypersensitivity pneumonitis, connective tissue disease-related ILD, IPF, rheumatoid-related ILD, sarcoidosis, Age-related ILD/presbyotic lung, ANCA vasculitis, asbestosis, autoimmune pneumonitis, chronic eosinophilic pneumonia, combined emphysema, and pulmonary fibrosis, desquamative interstitial pneumonia, non-specific interstitial pneumonia, organizing pneumonia, PPFE, smoking related ILD, or unclassifiable ILD IPF: ND	Mortality, n/N (%): Chronic Heart Disease:
			Moderate/Severe ILD: FVC<80% Mild ILD: FVC>80%	• No ILD: 29/322 (9.0%)

Study	Population and Setting	Intervention	Definitions	Results
				Severity of Condition:
			Clinical marker: NR	Mortality, n/N (%):
				Moderate/Severe ILD:
			Treatment/	• aHR: 1.72 (95%CI: 1.05 –
			Associated	2.83); p=0.032
			Therapy: NR	 Moderate/severe ILD:
			,	42/69 (60.9%)
			Outcome	• Mild ILD: 27/69 (39.1%)
			Definitions:	• p=0.039
			Mortality: in-hospital	,
			mortality	Duration of Condition: NR
			ICU admission: NR	
			Intubation: NR	Treatment/ Associated
			Ventilation: ND	Therapy: NR
			Hospitalization: NR	
			Non-elective	Comorbid Conditions:
			readmissions: NR	Mortality, n/N (%) <u>:</u>
			redumissions. NR	Obesity:
			Commonte None	Obese &ILD:
			Comments: None	• aHR: 2.27 (95%CI: 1.39–
				3.71); p=0.001
				5.1.44
				Risk Markers:
				Mortality, n/N (%):
				Sex, male:
				• aHR: 1.98 (1.14-
				3.43); p=0.015
				Female <60 years:
				• ILD: 3/11 (27%)
				• No ILD: 8/25 (32%)
				Female 60-75 years:
				• ILD: 10/21 (48%)
				• No ILD: 14/35 (40%)
				Female >75 years:
				• ILD: 4/19 (21%)
				• No ILD: 18/42 (43%)
				Male <60 years:
				• ILD: 2/8 (25%)
				 No ILD: 5/31 (16%)
				Male 60-75 years:
				• ILD: 27/49 (55%)
				• No ILD: 23/86 (27%)

Study	Population and Setting	Intervention	Definitions	Results
				Male >75 years:
				• ILD: 33/53 (62%)
				• No ILD: 46/103 (45%)
				Long-term Sequelae: NR
Author: Estiri ⁴	Population: N=16709	Health Condition	Medical	Severe COVID-19:
		Category: Cerebrovascular	Condition(s):	aOR: Adjusted odds ratio from
Year: 2021	Setting: Medical system consisting of 10 hospital	Disease, Chronic Heart	Cerebrovascular	Generalized Linear Model
		Disease, Neurocognitive	accident: ICD9 434.x,	(GLM) boosting model; median over 10 model iterations; model
Data	Location: MA, US	Disorders,	436, 437.x, 438.x;	included age, history of
Extractor: DOS		Diabetes, Chronic Kidney	ICD10 I63.x, I69.x,	pneumonia, type 2 diabetes
	Study dates: March 3 - November 10, 2020	Disease, Chronic Lung	G46.x	mellitus with complications, heart
Reviewer: MW		Disease, Risk	Thoracic aortic	failure, chronic kidney disease,
	Inclusion criteria: Electronic health record (EHR) data from	Factors, Immunocompromis	aneurysm: ICD9	interstitial pulmonary disease,
Study	patients with a confirmed case for COVID-19 (confirmed	ed Status, Blood Disorders,	441.1, 441.2; ICD10	chronic obstructive pulmonary
design: Cohort	PCR test) who had at least 1 year of medical history (i.e., a	Cancer	171.1, 171.2	disease, pulmonary embolism,
	1-year time difference between the first and last medical		Abdominal aortic	benign prostate hypertrophy, atrial
Study	record before the COVID-19 positive PCR test) with medical	Medical Condition, n/N	aneurysm: ICD9	fibrillation and flutter,
Objective: To	system. Included data from beginning of electronic record	(%):	441.3, 441.4; ICD10	hypertensive urgency or
predict risk of	(as far back as January 1, 2020) up to 14 days prior to the	Cerebrovascular accident:	171.3, 171.4	emergency, coronary artery
mortality and	positive COVID-19 PCR test date.	1337/16709 (8.0%)	Atrial fibrillation and	disease, gout, lung neoplasm,
study risk factors		Thoracic aortic aneurysm:	flutter: ICD9 427.3x;	history of a cerebrovascular
for death across	Exclusion criteria: NR	212/16709 (0.1%)	ICD10 I48.x	accident, abdominal aortic
different age		Abdominal aortic	Aortic valve	aneurysm, cardiomegaly, and
groups.		aneurysm: 151/16709	disorder: IDC9 424.1;	female RR: Univariate relative risk
		(0.9%)	ICD10 I35.x	OR: Univariate odds ratio
IVA Score: 25		Atrial fibrillation and flutter:	Cardiomegaly: ICD9	OK. Onivariate dads ratio
(moderate)		1262/16709 (7.6%)	429.3; ICD10 I51.7	Mortality, n/N (%):
		Aortic valve disorder:	Coronary artery	Cerebrovascular accident:
		727/16709 (4.4%)	disease: ICD9 414.x;	• aOR: 1.009 (IQR:
		Cardiomegaly: 893/16709	ICD10 I25.x, R93.1;	0.003)
		(5.3%)	ERX 705113; OMA	• RR: 5.10 (95%CI: 4.45-
		Coronary artery disease:	NSPA3	5.84)
		1924/16709 (11.5%)	Heart failure: ICD9	• OR: 6.07 (95%CI: 5.16-
		Heart failure: 1382/16709	428.3x, 428.4x; ICD10	7.11)
		(8.3%)	150.3x,	• Non-
		Hyperlipidemia:	I50.4x, ICD9 398.91,	survivors: 255/830 (30.7%)
		5733/16709 (34.3%)	402.11, 402.91,	• Survivors: 1082/15,879
		Hypertension: 6539/16709	404.01, 404.11,	(6.9%)
		(39.1%)	404.13, 404.91,	• p<0.001

Study	Population and Setting	Intervention	Definitions	Results
		Hypertensive emergency:	404.93, 428.0, 428.1,	Thoracic aortic aneurysm:
		353/16709 (2.1%)	428.2x, 428.4x; ODA	• RR: 3.43 (95%CI: 2.51-
		Mitral valve disorder:	MLFT5; ICD10 I09.81,	4.67)
		1131/16709 (6.8%)	113.0, 111.0, 150.2x,	• OR: 3.92 (95%CI: 2.67-
		Occlusion of the carotid	150.82, 150.84, 150.89,	5.60)
		artery: 573/16709 (3.4%)	150.4x, 150.9	Non-survivors: 35/830
		Pulmonary hypertension:	Hyperlipidemia: ICD9	(4.2%)
		322/16709 (1.9%)	272.x; ICD10 E78.x	 Survivors: 177/15,879
		Tricuspid valve disorder:	Hypertension: ICD9	(1.1%)
		695/16709 (4.2%)	401.x, 405.x, 642.13,	• p<0.001
		Ventricular tachycardia:	997.91; ICD10 I10.x,	Abdominal aortic aneurysm:
		331/16709 (2.0%)	O10.x, O11.x; ERX	• aOR: 1.006 (IQR:
		Parkinson's disease:	100141; LMA 953;	0.000)
		165/16709 (1.0%)	OMA MTDH2	• RR: 5.70 (95%CI: 4.35-
		Epilepsy: 623/16709 (3.7%)	Hypertensive	7.46)
		Diabetes mellitus, type 1:	emergency: ICD9	• OR: 7.47 (95%CI: 5.12-
		590/16709 (3.5%)	401.0, 405.01,	10.68)
		Diabetes mellitus, type 2	405.09; ICD10 I16.x	• Non-survivors: 41/830
		with complications:	Mitral valve	(4.9%)
		2082/16709 (12.5%)	disorder: ICD9 424.0;	• Survivors: 110/15,879
		Diabetes mellitus, type 2	ICD10 I34.x	(0.7%)
		without complications:	Occlusion of the	• p<0.001
		3018/16709 (18.1%)	carotid artery: ICD9	Atrial fibrillation and flutter:
		End-stage renal disease:	433.1; ICD10 I65.2x	• aOR: 1.016 (IQR:
		383/16709 (2.3%)	Pulmonary	0.003)
		Chronic kidney disease:	hypertension: ICD9	• RR: 5.77 (95%CI: 5.05-
		1554/16709 (9.3%)	416.0; ICD10 I27.x	6.60)
		Chronic obstructive	Tricuspid valve	• OR: 7.05 (95%CI: 6.00-
		pulmonary	disorder: IDC9 387.0;	8.26)
		disease (COPD): 910/16709	ICD10 I36.x	• Non-
		(5.4%)	Ventricular	survivors: 266/830 (32.0%)
		Interstitial pulmonary	tachycardia: IDC9	• Survivors: 996/15,879
		disease: 260/16709 (1.6%)	427.1; ICD10 I47.2	(6.3%)
		Obstructive sleep apnea:	Parkinson's	• p<0.001
		1657/16709 (9.9%)	disease: ICD9 332.x;	Aortic valve disorder:
		Pulmonary embolism:	ICD10 G21.x	• RR: 5.66 (95%CI: 4.87-
		940/16709 (5.6%)	Epilepsy: ICD9 345.x,	6.59)
		Rheumatoid arthritis:	649.4x; ICD10 G40.x,	• OR: 7.09 (95%CI: 5.86-
		396/16709 (2.4%)	Z82.0	8.54)
		Anemia: 4974/16709		• Non-
		(29.8%)		survivors: 170/830 (20.5%)

Study	Population and Setting	Intervention	Definitions	Results
		Peripheral vascular disease:	Diabetes mellitus,	• Survivors: 557/15,879
		674/16709 (4.0%)	type 1: ICD9 25.x1,	(3.5%)
		Breast neoplasm:	250.x3; ICD10 E10.x	• p<0.001
		407/16709 (2.4%)	Diabetes mellitus,	Cardiomegaly:
		Lung neoplasm: 190/16709	type 2 with	• aOR: 1.006 (IQR:
		(1.1%)	complications: ICD9	0.000)
		Prostate neoplasm:	249.x, 250.x0, 250.x2;	• RR: 4.03 (95%CI: 3.43-
		314/16709 (1.9%)	ICD10 E08.x, E09.x,	4.74)
			E11.x, E13.x	• OR: 4.67 (95%CI: 3.85-
		Control/Comparison group,	Diabetes mellitus,	5.63)
		n/N (%):	type 2 without	• Non-
		No cerebrovascular	complications: ICD9	survivors: 154/830 (18.6%)
		accident: 15372/16709	240.0x, 250.0x; ICD10	Survivors: 739/15,879
		(92.0%)	E08.9, E11.9, E13.9	(4.7%)
		No thoracic aortic	End-stage renal	• p<0.001
		aneurysm: 16497/16709	disease: ICD9 458.21,	Coronary artery disease:
		(98.7%)	585.5, 585.6, 996.68,	• aOR: 1.014 (IQR:
		No abdominal aortic	996.68,	0.003)
		aneurysm: 16558/16709	996.73, V45.1,	• RR: 5.52 (95%CI: 4.85-
		(99.1%)	V45.11, V45.12,	6.28)
		No atrial fibrillation and	V56.x; ICD10 N18.5,	• OR: 6.28 (95%CI: 5.62-
		flutter: 15447/16709	N18.6, I13.11, I13.2,	7.55)
		(92.4%)	112.0, 195.3,	• Non-
		No aortic valve	T85.611x, T85.621A,	survivors: 347/830 (41.8%)
		disorder: 15982/16709	T85.691A, T85.71XD,	Survivors: 1577/15,879
		(95.6%)	T82.41Xx, T82.43Xx,	(10.0%)
		No	T82.49Xx, Y62.2, Y84.	• p<0.001
		cardiomegaly: 15816/16709	1, Z49.01, Z49.02,	Heart failure:
		(94.7%)	Z49.3, Z91.15, Z99.2;	• aOR: 1.047 (IQR:
		No coronary artery disease:	ERX122830, LMA	0.011)
		14785/16709 (88.5%)	3772, OMA BASX6	• RR: 6.31 (95%CI: 5.54-
		No heart failure:	Chronic kidney	7.19)
		15327/16709 (91.7%)	disease: ICD9 585.x;	• OR: 7.79 (95%CI: 6.67-
		No	ICD10 D63.1, E08.22,	9.08)
		hyperlipidemia: 109761670	E09.22, E10.22,	• Non-
		9 (65.7%)	E11.22, E13.22, I12.x,	survivors: 301/830 (36.3%)
		No	I13.x, N18.x	 Survivors: 1081/15,879
				(6.8%)
				• p<0.001
		(Hyperlipidemia:
		hypertension: 10170/16709 (60.9%)	COPD: ICD9 434.x, 436, 437.x, 438.x; ICD10 I63.x, I69.x, G46.x	• p<0.001

Study	Population and Setting	Intervention	Definitions	Results
		No hypertensive	Interstitial pulmonary	• RR: 4.04 (95%CI: 3.50-
		emergency: 16356/16709	disease: ICD9 516.x;	4.65)
		(97.9%)	ICD10 J84.x	• OR: 4.37 (95%CI: 3.76-
		No mitral valve disorder:	Obstructive sleep	5.08)
		15578/16709 (93.2%)	apnea: ICD9 780.57,	• Non-
		No occlusion of the carotid	780.53, 327.23;	survivors: 563/830 (67.8%)
		artery: 16136/16709	ICD10 G47.33	• Survivors: 5170/15,879
		(96.6%)	Pulmonary	(32.7%)
		No pulmonary	embolism: ICD9	• p<0.001
		hypertension: 16387/16709	415.x, 453.x, 445.x,	Hypertension:
		(98.1%)	673.x; ICD10 I26.x,	• RR: 6.61 (95%CI: 5.58-
		No tricuspid valve	174.x, 175.x, 182.x,	7.84)
		disorder: 16014/16709	O88.x, Z86.711,	• OR: 7.25 (95%CI: 6.10-
		(95.8%)	Z86.718	8.68)
		No ventricular	Rheumatoid	• Non-
		tachycardia: 16378/16709	arthritis: ICD9 714.x;	survivors: 672/830 (81.0%)
		(98.0%)	ICD10 M05.x, M06.x,	• Survivors: 5867/15,879
		No Parkinson's disease:	M08.x	(37.2%)
		16544/16709 (99.0%)	Anemia: ICD9 280.x,	• p<0.001
		No epilepsy: 16086/16709	281.x, 282.x, 283.x,	Hypertensive emergency:
		(96.3%)	285.x; ICD10 D50.x,	• aOR: 1.015 (IQR:
		No diabetes mellitus, type	D50.x-53.x, D55.x-	0.021)
		1: 16119/16709 (96.5%)	D59.x, D60x-D64.x,	• RR: 3.87 (95%CI: 3.07-
		No diabetes mellitus, type 2	O90.81, O99.01x	4.88)
		with	Peripheral vascular	• OR: 4.51 (95%CI: 3.38-
		complications: 14627/1670	disease: ICD9 443.9;	5.94)
		9 (87.5%)	ICD10 I73.9	• Non-survivors: 64/830
		No diabetes mellitus, type 2	Breast	(7.7%)
		without	neoplasm: ICD9	• Survivors: 289/15,879
		complications: 13691/1670	174.x; ICD10 C50.x	(1.8%)
		9 (81.9%)	Lung neoplasm: ICD9	• p<0.001
		No end-stage renal	162.x (excluding x=0);	Mitral valve disorder:
		disease: 16326/16709 (97.7	ICD10 C34.x	• RR: 3.98 (95%CI: 3.42-
		%)	Prostate	4.63)
		No chronic kidney disease:	neoplasm: ICD9	• OR: 4.57 (95%CI: 3.82-
		15155/16709 (90.7%)	185.x; ICD10 C61.x	5.43)
		No COPD: 15799/16709		• Non-
		(94.6%)	Severity	survivors: 186/830 (22.4%)
		No interstitial pulmonary	Measure(s): NR	• Survivors: 945/15,879
		disease: 16449/16709		(6.0%)
		(98.4%)	Clinical marker: NR	• p<0.001

Study	Population and Setting	Intervention	Definitions	Results
		No obstructive sleep apnea:		Occlusion of the carotid artery:
		15052/16709 (90.1%)	Treatment/	• RR: 5.23 (95%CI: 4.42-
		No pulmonary	Associated	6.18)
		embolism: 15769/16709	Therapy: NR	• OR: 6.47 (95%CI: 5.23-
		(94.4%)		7.96)
		No rheumatoid	Outcome	• Non-
		arthritis: 16313/16709	Definitions:	survivors: 130/830 (15.7%)
		(97.6%)	Mortality: from	Survivors: 443/15,879
		No anemia: 11735/16709	various data sources	(2.8%)
		(70.2%)	and included	• p<0.001
		No peripheral vascular	mortality unrelated	Pulmonary hypertension:
		disease: 16035/16709	to visit	• RR: 4.91 (95%C
		(96.0%)	ICU admission: NR	I: 3.96-6.07)
		No breast	Intubation: NR	• OR: 6.06 (95%
		neoplasm: 16302/16709	Ventilation: NR	CI: 4.59-7.91)
		(97.6%)	Hospitalization: NR	• Non-
		No lung neoplasm:	Non-elective	survivors: 73/830 (8.8%)
		16519/16709 (98.9%)	readmissions: NR	Survivors: 249/
		No prostate		15,879 (1.6%)
		neoplasm: 16395/16709	Comments: None	• p<0.001
		(98.1%)		Tricuspid valve disorder:
				• RR: 4.12 (95%C
				I: 3.47-4.91)
				• OR: 4.82 (95%
				CI: 3.90-5.91)
				• Non-
				survivors: 126/830 (15.2%)
				• Survivors: 569/
				15,879 (3.6%)
				• p<0.001
				Ventricular tachycardia:
				• RR: 5.42 (95%C
				I: 4.44-6.63)
				• OR: 6.89 (95%
				CI: 5.28-8.90)
				• Non-
				survivors: 82/830 (9.9%)
				• Survivors: 249/
				15,879 (1.6%)
				• p<0.001
				Parkinson's disease:

Study	Population and Setting	Intervention	Definitions	Results	
				•	RR: 4.02 (95%C
				I: 2.92-5.53)	
				•	OR: 4.76 (95%
				CI: 3.16-6.96)	
				•	Non-
				survivors: 32/8	30 (3.9%)
				•	Survivors: 133/
				15,879 (0.8%)	
				•	p<0.001
				Epilepsy:	
				•	RR: 2.45 (95%C
				I: 1.95-3.08)	
				•	OR: 2.65 (95%
				CI: 2.03-3.40)	
				•	Non-
				survivors: 72/8	30 (8.7%)
				•	Survivors: 551/
				15,879 (3.5%)	
				•	p<0.001
				Diabetes mellitus	s, type 1:
				•	RR: 3.20 (95%C
				I: 2.60-3.93)	
				•	OR: 3.58 (95%
				CI: 2.80-4.53)	
				•	Non-
				survivors: 87/8	30 (10.5%)
				•	Survivors: 503/
				15,879 (3.2%)	
				•	p<0.001
				Diabetes mellitus	s, type 2 with
				complications:	
				•	aOR: 1.047
				(Interquartile R	Range (IQR):
				0.015)	
				•	RR: 3.82 (95%C
				I: 3.35-4.39)	
				•	OR: 4.30 (95%
				CI: 3.69-4.99)	
				•	Non-
				survivors: 293/	830 (35.3%)
				•	Survivors: 178
				9/15,879 (11.3	%)

Study	Population and Setting	Intervention	Definitions	Results
				• p<0.001
				Diabetes mellitus, type 2 without
				complications:
				• RR: 3.47 (95%
				I: 3.05-3.96)
				• OR: 3.81 (95%
				CI: 3.30-4.40)
				• Non-
				survivors: 360/830 (43.4%)
				Survivors: 265
				8/15,879 (16.8%)
				• p<0.001
				End-stage renal disease:
				• RR: 4.74 (95%
				I: 3.87-5.80)
				• OR: 5.78 (95%
				CI: 4.46-7.41)
				• Non-
				survivors: 83/830 (10.0%)
				Survivors: 300
				15,879 (1.9%)
				• p<0.001
				Chronic kidney disease:
				• aOR: 1.042
				(IQR: 0.014)
				• RR: 6.83 (95%
				I: 6.01-7.77)
				• OR: 8.48 (95%
				CI: 7.29-9.85)
				• Non-
				survivors: 342/830 (41.2%)
				Survivors: 121
				2/15,879 (7.7%)
				• p<0.001
				COPD:
				• aOR: 1.024
				(IQR: 0.021)
				• RR: 4.77 (95%
				I: 4.10-5.55)
				• OR: 5.70 (95%
				CI: 4.74-6.82)

Study	Population and Setting	Intervention	Definitions	Results	
				•	Non-
				survivors: 179/8	830 (21.6%)
				•	Survivors: 731/
				15,879 (4.6%)	
				•	p<0.001
				Interstitial pulmo	nary disease:
				•	aOR: 1.027
				(IQR: 0.025)	
				•	RR: 4.32 (95%C
				I: 3.36-5.54)	
				•	OR: 5.17 (95%
				CI: 3.76-7.00)	
				•	Non-
				survivors: 53/83	30 (6.4%)
				•	Survivors: 207/
				15,879 (1.3%)	
				•	p<0.001
				Obstructive sleep	apnea:
				•	RR: 2.84 (95%C
				I: 2.49-3.24)	
				•	OR: 3.04 (95%
				CI: 2.64-3.51)	
				•	Non-
				survivors: 128/8	830 (15.4%)
				•	Survivors: 152
				9/15,879 (9.7%	
				•	p<0.001
				Pulmonary embo	
				•	aOR: 1.018
				(IQR: 0.019)	
				•	RR: 3.55 (95%C
				I: 3.01-4.19)	
				•	OR: 4.02 (95%
				CI: 3.30-4.86)	
				•	Non-
				survivors: 145/8	
				•	Survivors: 795/
				15,879 (5.0%)	
				•	p<0.001
				Rheumatoid arth	ritis:

Study	Population and Setting	Intervention	Definitions	Results	
				•	RR: 2.36 (95%C
				I: 1.78-3.14)	
				•	OR: 2.54 (95%
				CI: 1.83-3.46)	
				•	Non-
				survivors: 45/83	0 (5.4%)
				•	Survivors: 351/
				15,879 (2.2%)	
				•	p<0.001
				Anemia:	
				•	RR: 3.38 (95%C
				I: 2.96-3.87)	
				•	OR: 3.64 (95%
				CI: 3.16-4.20)	
				•	Non-
				survivors: 489/8	
				•	Survivors: 448
				5/15,879 (28.4%	
				•	p<0.001
				Peripheral vascula	
				•	RR: 5.04 (95%C
				I: 4.28-5.92)	
				0, 500 7.40)	OR: 6.15 (95%
				CI: 5.02-7.48)	
				• 145/0	Non-
				survivors: 145/8	
				• 15,879 (3.4%)	Survivors: 529/
				13,879 (3.4%)	n<0.001
				Breast neoplasm:	p<0.001
				•	RR: 2.62 (95%C
				I: 2.01-3.42)	KK. 2.02 (93/0C
				1. 2.01-3.42)	OR: 2.86 (95%
				CI: 2.09-3.84)	ON. 2.00 (33%
				•	Non-
				survivors: 51/83	
				•	Survivors: 356/
				15,879 (2.3%)	Jul vivois. 330/
				13,073 (2.370)	p<0.001
				Lung neoplasm:	p 10.001

Study	Population and Setting	Intervention	Definitions	Results
				• aOR: 1.012
				(IQR: 0.000)
				• RR: 4.75 (95%C
				I: 3.62-6.23)
				• OR: 5.86 (95%
				CI: 4.09-8.22)
				• Non-
				survivors: 43/830 (5.2%)
				• Survivors: 147/
				15,879 (0.9%)
				• p<0.001 Prostate neoplasm:
				• RR: 3.63 (95%C
				l: 2.82-4.68)
				• OR: 4.19 (95%
				CI: 3.07-5.62)
				• Non-
				survivors: 54/830 (6.5%)
				Survivors: 260/
				15,879 (1.6%)
				• p<0.001
				Severity of Condition: NR
				Duration of Condition: NR
				Treatment/ Associated
				Therapy: NR
				Comorbid Conditions: NR
				Risk Markers:
				Mortality, n/N (%):
				Female:
				• aOR: 0.972
				(IQR: 0.017)
				• RR: 0.67
				(95%CI: 0.58-0.76)
				• OR: 0.65
				(95%CI: 0.57-0.75)
				• Non-
				survivors: 391/830 (47.1%)

Study	Population and Setting	Intervention	Definitions	Results
				• Survivors: 9,16 8/15,879 (58.1%) • p<0.001
				Ethnicity, Hispanic: RR: 0.32 (95%CI: 0.22-0.47) OR: 0.31 (95%CI: 0.20-0.45) Non-
				survivors: 27/830 (3.3%) Survivors: 1,56 7/15,879 (9.9%) p<0.001 Race, White: RR: 2.36
				(95%CI: 2.03-2.74) • OR: 2.45 (95%CI: 2.10-2.88) • Non- survivors: 608/830 (73.3%)
				● Survivors: 8,37 4/15,879 (53%) ● p<0.001 Race, Black or African American: ● RR: 0.84 (95%CI: 0.69-1.03)
				• OR: 0.84 (95%CI: 0.67-1.03) • Non- survivors: 104/830 (12.5%) • Survivors: 2,32 6/15,879 (14.7%)
				• p=0.102 Age: • aOR: 2.816 (IQ R: 0.052)
				Age, under 45 years: ■ RR: 0.04 (95%CI: 0.02-0.06)

Study	Population and Setting	Intervention	Definitions	Results
				• OR: 0.03
				(95%CI: 0.02-0.05)
				• Non-
				survivors: 21/830 (2.5%)
				• Survivors: 6,90
				3/15,879 (43.7%)
				• p<0.001
				Age, 45-65 years:
				• RR: 0.39
				(95%CI: 0.33-0.47)
				• OR: 0.38
				(95%CI: 0.31-0.45)
				Non-
				survivors: 142/830 (17.1%)
				Survivors: 5,62
				9/15,879 (35.7%)
				• p<0.001
				Age, 65-85 years:
				• RR: 3.88
				(95%CI: 3.41-4.43)
				• OR: 4.30 (3.73-
				4.96)
				• Non-
				survivors: 385/830 (46.4%)
				Survivors: 2,65
				9/15,879 (16.8%)
				• p<0.001
				Age, over 85 years:
				• RR: 8.35
				(95%CI: 7.35-9.49)
				• OR: 11.36
				(95%CI: 9.65-13.36)
				• Non-
				survivors: 282/830 (34.0%)
				• Survivors: 688/
				15,879 (4.4%)
				• p<0.001
				· '
				Smoking history:
				• RR: 2.09
				(95%CI: 1.76-2.50)

Study	Population and Setting	Intervention	Definitions	Results
				• OR: 2.21 (95%CI: 1.82-2.67) • Non- survivors: 137/830 (16.5%) • Survivors: 130 4/15,879 (8.3%) • p<0.001 Long-term Sequelae: NR
Author: Halalau ⁵	Population: N=821	Health Condition	Medical	Severe COVID-19:
Year: 2021	Setting: Large healthcare system including 8 hospitals	Category: Chronic Heart Disease, Chronic Liver Disease,	Condition(s): Cerebrovascular Disease: ND	aOR: Adjusted odds ratio; multivariate regression model included age, BMI, diabetes
Data Extractor: MW	Location: Michigan, USA Study dates: Up to April 12, 2020	Neurocognitive Disorders, Diabetes, Chronic Kidney Disease, Chronic Lung	Hypertension: ND Hyperlipidemia: ND Coronary Artery	mellitus, obstructive sleep apnea, hypertension, COPD, dementia, and chronic kidney disease
Reviewer: DOS	Inclusion criteria: Patients who tested positive for SARS-	Disease, Risk Factors, Immunocompromis	Disease: ND Heart failure: ND	Hospitalization, n/N (%): Cerebrovascular disease:
Study Design: Cohort	CoV-2 at any date up to April 1, 2020, after evaluation at any of the emergency departments across the 8	ed Status, Blood Disorders, Cancer	Cardiac arrhythmia: ND	Admitted patients: 10/86 (11.6%)
Study Objective: To describe the	study hospitals, and subsequently discharged home. Laboratory confirmation for COVID-19 was defined as a positive result of real-time RT-PCR assay of nasopharyngeal swabs. Testing was offered if patients	Medical Condition, n/N (%): Cerebrovascular Disease: 71/821 (8.6%)	Chronic liver disease: ND Chronic Hepatitis B: ND	 Outpatients: 61/735 (8.3%) p=0.299 Transient Ischemic Attack:
demographics, initial clinical presentation, and outcomes of	experienced moderate cough or fever over 100.4°F, and if they had chronic kidney disease, heart disease, diabetes, chronic lung disease, were receiving immunosuppression medication, or were	Transient Ischemic Attack: 48/821 (5.8%) Hypertension: 261/821 (31.8%)	C: ND Cognitive impairment or dementia: ND	 Admitted patients: 8/86 (9.3%) Outpatients: 40/735 (5.4%)
a large cohort of outpatients with COVID-19.	immunocompromised due to cancer treatment, recent surgeries, or other conditions.	Hyperlipidemia: 168/821 (20.5%) Coronary Artery Disease:	Seizure disorder: ND Transient Ischemic Attack: ND	p=0.149 Hypertension: Admitted patients:
IVA Score: 23 (moder	Exclusion criteria: All patients with a negative test for SARS-CoV-2.	125/821 (15.2%) Heart failure: 18/821 (2.2%)	Psychiatric Disorder: ND Diabetes	45/86 (52.3%) • Outpatients: 216/735 (29.4%)
ate)		Cardiac arrhythmia: 116/821 (14.1%) Chronic liver disease: 11/821 (1.3%) Chronic Hepatitis B: 1/821	Mellitus: ND Prediabetes: ND Chronic kidney disease: ND COPD: ND	 p<0.001 Hyperlipidemia: Admitted patients: 29/86 (33.7%)
		(0.1%)	33, 2, 112	Outpatients: 139/735 (18.9%)

Study	Population and Setting	Intervention	Definitions	Results
		Chronic Hepatitis C: 1/821	Bronchial	• p=0.001
		(0.1%)	Asthma: ND	Coronary artery disease:
		Cognitive impairment or	Obstructive Sleep	 Admitted patients:
		dementia: 153/821 (18.6%)	Apnea: ND	24/86 (27.9%)
		Seizure disorder: 150/821	Interstitial Lung	 Outpatients: 101/735
		(18.3%)	Disease: ND	(13.7%)
		Psychiatric Disorder:	Pulmonary	• p=0.001
		103/821 (12.5%)	hypertension: ND	Heart failure:
		Diabetes Mellitus: 279/821	Sarcoidosis: ND	 Admitted patients: 2/86
		(34%)	Venous	(2.3%)
		Prediabetes: 82/821	thromboembolic	Outpatients: 16/735
		(10.2%)	disease: ND	(2.2%)
		Chronic kidney disease:	Rheumatologic	• p=1.0
		86/821 (10.5%)	disorders: ND	Cardiac arrhythmia:
		Chronic obstructive	Immunosuppression:	 Admitted patients:
		pulmonary disease (COPD):	ND	15/86 (17.4%)
		74/821 (9%)	Venous	 Outpatients: 101/735
		Bronchial Asthma: 92/821	thromboembolic	(13.7%)
		(11.2%)	disease: ND	• p=0.351
		Obstructive Sleep Apnea:	Cancer: ND	Chronic liver disease:
		171/821 (20.8%)		 Admitted patients: 0/86
		Interstitial Lung Disease:	Severity	(0%)
		0/821 (0%)	Measure(s): NR	Outpatients: 11/735
		Pulmonary hypertension:		(1.5%)
		8/821 (1%)	Clinical marker: NR	• p=0.617
		Sarcoidosis: 6/821 (0.7%)		Chronic hepatitis B:
		Venous thromboembolic	Treatment/	 Admitted patients: 0/86
		disease: 103/821 (12.5%)	Associated	(0%)
		Rheumatologic disorders:	Therapy: NR	Outpatients: 1/735
		146/821 (17.8%)		(0.1%)
		Immunosuppression:	Outcome	• p=1.0
		11/821 (1.3%)	Definitions:	Chronic hepatitis C:
		Cancer: 75/821 (9.1%)	Mortality: NR	 Admitted patients: 0/86
			ICU admission: NR	(0%)
		Control/Comparison group,	Intubation: NR	• Outpatients: 1/735
		n/N (%):	Ventilation: NR	(0.1%)
		None of the above: 295/821	Hospitalization: Emer	• p=1.0
		(35.9%)	gency department	Cognitive impairment or
			visits for the patients	dementia:
			that resulted in	Admitted patients:
			admission to hospital	30/86 (34.9%)

Study	Population and Setting	Intervention	Definitions	Results
			Non-elective	Outpatients: 123/735
			readmissions: NR	(16.7%)
				• p < 0.001
			Comments: None	Seizure disorder:
				Admitted patients:
				27/86 (31.4%)
				 Outpatients: 123/735
				(16.7%)
				• p=0.001
				Psychiatric Disorder:
				 Admitted patients: 9/86
				(10.5%)
				Outpatients: 94/735
				(12.8%)
				• p=0.538
				Diabetes Mellitus:
				• aOR: 3.27 (95%CI: 1.49-
				7.15), p=0.003
				Admitted patients:
				64/86 (74.4%)
				Outpatients: 215/735
				(29.3%)
				• p<0.001
				Prediabetes:
				Admitted patients:
				22/86 (25.6%)
				• Outpatients: 62/735
				(8.4%)
				• p<0.001
				Chronic kidney disease:
				• Admitted patients:
				15/86 (17.4%)
				 Outpatients: 71/735 (9.7%)
				• p=0.026 COPD:
				Admitted patients:
				34/86 (39.5%)
				• Outpatients: 40/735
				(5.4%)
<u> </u>				• p<0.001

Study	Population and Setting	Intervention	Definitions	Results
				Bronchial Asthma:
				 Admitted patients:
				12/86 (14.0%)
				Outpatients: 80/735
				(10.9%)
				• p=0.393
				Obstructive Sleep Apnea:
				• aOR: 3.44 (95%CI: 1.11- 10.65), p=0.032
				Admitted patients:
				31/86 (36.0%)
				 Outpatients: 140/735 (19.0%)
				• p<0.001
				Interstitial Lung Disease:
				Admitted patients: 0/86 (0%)
				 Outpatients: 0/735 (0%)
				• p=N/A
				Pulmonary hypertension:
				• Admitted patients: 0/86 (0%)
				• Outpatients: 8/735
				(1.1%)
				• p=1.0
				Sarcoidosis:
				Admitted patients: 2/86 (2.3%)
				• Outpatients: 4/735 (0.5%)
				• p=0.123
				Venous thromboembolic disease:
				Admitted patients:
				11/86 (12.8%)
				Outpatients: 92/735
				(12.5%)
				• p=0.942
				Rheumatologic disorders:
				 Admitted patients: 24/86 (28.9%)

Study	Population and Setting	Intervention	Definitions	Results
				 Outpatients: 122/735
				(16.6%)
				• p=0.010
				Immunosuppression:
				 Admitted patients: 1/86
				(1.2%)
				Outpatients: 10/735
				(1.4%)
				• p=1.0
				Cancer:
				 Admitted patients:
				14/86 (16.3%)
				Outpatients: 61/735
				(8.3%)
				• p=0.015
				Severity of Condition: NR
				Duration of Condition: NR
				Treatment/ Associated
				Therapy: NR
				Comorbid Conditions: NR
				Risk Markers:
				Hospitalization, n/N (%) or mean
				±SD:
				Age, years:
				 Admitted
				patients: 54.4±15.6
				Outpatients: 48.7±15.7
				• p=0.002
				Sex, male:
				 Admitted patients:
				50/86 (58.1%)
				Outpatients: 334/735
				(45.4%)
				• p=0.026
				Ethnicity; p=0.679
				Caucasian:

Study	Population and Setting	Intervention	Definitions	Results
				 Admitted patients: 34/86 (39.5%) Outpatients: 269/735 (36.6%) African American: Admitted patients: 47/86 (54.6%) Outpatients: 405/735 (55.1%) Other: Admitted patients: 5/86 (5.8%) Outpatients: 61/735 (8.3%) Long-term Sequelae: NR
Author: Kokturk ⁶	Population: N=1500	Health Condition Category:	Medical	Severe COVID-19:
W 2024	Setting: 26 Centers (17 university hospitals, 2 large tertiary	Cerebrovascular disease,	Condition(s):	aOR: Adjusted odds ratio;
Year: 2021	hospitals, 2 secondary care hospitals and 5 private hospitals)	Chronic heart disease,	Cerebrovascular disease: ND	multivariable logistic regression with 1228 cases including clinical
Data Extractor:	Location: Turkey	Chronic liver disease, Diabetes, Chronic kidney	Atherosclerosis: ND	parameters, disease spectrum and
MW	Location. Turkey	disease, Obesity, Chronic	Hypertension: ND	comorbidities
IVIVV	Study dates: March 11 – July 18, 2020	lung disease, Risk factors,	Heart failure: ND	OR: Odds ratio; univariable logistic
Reviewer: DOS	Study dates. March 11 – July 16, 2020	Multiple comorbid	Chronic hepatic	regression
Reviewer. DO3	Inclusion criteria: Patients admitted to the hospital during	conditions,	disease: ND	regression
Study Design:	study dates with a proven presence of a positive nucleic	Immunocompromised	Diabetes: ND	Mortality, n/N (%):
Cohort	acid amplification test or a positive rapid antigen detection	status, Blood disorders,	Chronic kidney	Cerebrovascular disease:
Conort	test together with clinical and radiographic findings that	Cancer	disease: ND	
Study Objective:	were strongly suggestive of COVID-19, and Highly probable	Cancer	Obesity: BMI ≥30	• OR: 1.97 (95%CI: 0.69–5.65); p=0.209
To evaluate the	cases presented with similar clinical and radiographic	Medical Condition, n/N	Asthma: ND	'
clinical outcomes	findings but could not be confirmed with an RT-PCR test.	(%):	COPD: ND	• Non-survivors: 4/67 (6.3%)
of hospitalized	mangs sat coald not be committed with all Ki T cit test.	Cerebrovascular disease:	Bronchiectasis: ND	• Survivors: 45/1433 (3.3%)
patients and to	Exclusion criteria: NR	49/1500 (3.4%)	Interstitial lung	Atherosclerosis:
predict COVID-19		Atherosclerosis: 145/1500	disease: ND	• OR: 4.12 (95%CI: 2.34–7.26);
mortality among		(10%)	Immunosuppressive	p=0.001
highly suspected		Hypertension: 402/1500	conditions: ND	• Non-survivors: 19/67 (29.2%)
patients.		(27.4%)	Connective tissue	• Survivors: 126/1433 (9.1%)
		Heart failure: 64/1500	disorder: ND	Hypertension:
IVA Score: 24		(4.4%)	Malignancy: ND	• OR: 3.08 (95%CI: 1.88–5.05);
(moderate)		Chronic hepatic disease:	Others: ND	p=0.001
		11/1500 (0.8%)		• Non-survivors: 35/67 (52.2%)
		Diabetes: 236/1500 (16.3%)		• Survivors: 367/1433 (26.2%)

Study	Population and Setting	Intervention	Definitions	Results
•		Chronic kidney disease:	Severity Measure(s):	Heart failure:
		51/1500 (3.5%)	NR	• OR: 5.08 (95%CI: 2.51–10.27);
		Obesity: 175/1500 (21.4%)		p=0.001
		Asthma: 111/1500 (7.7%)	Clinical marker: NR	• Non-survivors: 11/67 (16.9%)
		Chronic obstructive		• Survivors: 53/1433 (3.9%)
		pulmonary disease (COPD):	Treatment/	Chronic hepatic disease:
		90/1500 (6.2%)	Associated Therapy:	• OR: 2.16 (95%CI: 0.27–17.15);
		Bronchiectasis: 12/1500	NR	p=0.466
		(0.8%)		• Non-survivors: 1/67 (1.6%)
		Interstitial lung disease:	Outcome Definitions:	• Survivors: 10/1433 (0.7%)
		22/1500 (1.5%)	Mortality: ND	Diabetes:
		Immunosuppressive conditions: 25/1500 (1.7%)	ICU admission: NR Intubation: NR	• OR: 1.28 (95%CI: 0.69–2.39);
		Connective tissue disorders:	Ventilation: NR	p=0.439
		25/1500 (1.7%)	Hospitalization: NR	'
		Malignancy: 76/1500 (5.3%)	Non-elective	• Non-survivors: 13/67 (19.7%)
			readmissions: NR	• Survivors: 223/1433 (16.1%)
		Control/Comparison group,	redamissions. W	Chronic kidney disease:
		n/N (%):	Comments: None	• OR: 4.35 (95%CI: 1.96–9.68);
		No cerebrovascular disease:		p=0.001
		1451/1500 (96.7%)		• Non-survivors: 8/67 (12.3%)
		No atherosclerosis:		• Survivors: 43/1433 (3.1%)
		1355/1500 (90.3%)		Obesity:
		No hypertension:		• OR: 5.01 (95%CI: 0.66–37.80);
		1098/1500 (73.2%)		p=0.118
		No heart failure: 1436/1500		• Not obesity: 18/643 (2.8%)
		(95.7%)		• Obesity: 1/175 (0.5%)
		No chronic hepatic disease:		Asthma:
		1489/1500 (99.3%)		• OR: 0.57 (95%CI: 0.17–1.84);
		No diabetes: 1264/1500		p=0.345
		(84.3%) No chronic kidney disease:		 Non-survivors: 3/67 (4.6%)
		1449/1500 (96.6%)		• Survivors: 108/1433 (7.9%)
		Not obese (BMI<30):		COPD:
		643/1500 (78.6%)		• OR: 5.24 (95%CI: 2.81–9.76);
		No asthma: 1389/1500		p=0.001
		(92.6%)		• Non-survivors: 15/67 (23.1%)
		No COPD: 1410/1500 (94%)		• Survivors: 75/1433 (5.4%)
		No bronchiectasis:		Bronchiectasis:
		1488/1500 (99.2%)		• OR: 1.93 (95%CI: 0.25–15.21);
		No interstitial lung disease:		p=0.531
		1478/1500 (98.5%)		• Non-survivors: 1/67 (1.5%)
				• Survivors: 11/1433 (0.8%)
				Interstitial lung disease:

Study	Population and Setting	Intervention	Definitions	Results
		No immunosuppressive		• aOR: 5.27 (95%CI: 1.17–23.8);
		conditions: 1475/1500		p=0.031
		(98.3%)		• OR: 4.99 (95%CI: 1.64–15.21);
		No connective tissue		p=0.005
		disorders: 1475/1500 (98.3%)		 Non-survivors: 4/67 (6.3%)
		(98.3%) No malignancy: 1424/1500		• Survivors: 18/1433 (1.3%)
		(94.9%)		Immunosuppressive conditions:
		(3 1.370)		• OR: 5.73 (95%CI: 2.08–15.78);
				p=0.001
				Non-survivors: 5/67 (7.8%)
				• Survivors: 20/1433 (1.5%)
				Connective tissue disorder:
				• OR: 4.21 (95%CI: 1.4–12.66);
				p=0.001
				• Non-survivors: 4/67 (6.0%)
				• Survivors: 21/1433 (1.4%)
				Malignancy:
				• aOR: 19.99 (95%CI: 8.14-49.1);
				p=0.001
				• OR: 10.49 (95%CI: 5.81–18.94);
				p=0.001
				 Non-survivors: 20/67 (30.8%)
				• Survivors: 56/1433 (4.1%)
				Severity of Condition: NR
				Duration of Condition: NR
				Treatment/ Associated Therapy: NR
				Comorbid Conditions: NR
				Risk Markers:
				Mortality, n/N (%) or mean \pm SD:
				Age (years):
				• aOR: 1.09 (95%CI: 1.06–1.12);
				p=0.001
				• OR: 1.08 (95%CI: 1.06–1.10);
				p=0.001
				 Non-survivors: 71.3 ± 15.06

Study	Population and Setting	Intervention	Definitions	Results
				• Survivors: 50.98 ± 17.24
				Age group:
				• OR: 6.71 (95%CI: 3.97–11.33);
				p=0.001
				 Non-survivors ≥65: 45/379
				(11.9%)
				 Non-survivors <65: 22/1117
				(2.0%)
				Sex:
				• aOR: 2.47 (95%CI: 1.05-5.82);
				p=0.038
				• OR: 1.96 (95%CI: 1.14-3.36);
				p=0.015
				Male non-survivors: 48/850
				(5.6%)
				• Female non-survivors: 19/640
				(3.0%)
				Smoking:
				Active smokers:
				• OR: 0.76 (95%CI: 0.26-2.23);
				p=0.622
				Active smoker non-survivors:
				4/204 (2.0%)
				Never smoker non-survivors:
				23/901 (2.5%)
				Ex-smokers:
				• OR: 3.77 (95%CI: 2.08–6.84);
				p=0.001
				• Ex-smoker non-survivors:
				23/256 (9.0%)
				Never smoker non-survivors:
				23/901 (2.5%)
				Ever smokers:
				• OR: 2.38 (95%CI: 1.35-4.20);
				p=0.003
				Ever smoker non-survivors:
				27/460 (5.9%)
				 Never smoker non-survivors:
				23/901 (2.5%)
				Long-term Sequelae: NR

Study	Population and Setting	Intervention	Definitions	Results
Author: Mollalo ⁸	Population: N=NR	Health Condition Category:	Medical	Severe COVID-19:
		Cerebrovascular disease,	Condition(s):	Mixed-effects multinomial logistic
Year: 2021	Setting: nationwide	Chronic heart disease,	Cerebrovascular	regression model odds ratio [OR]
		Chronic liver disease,	disease: ND	(95% CI) for association between
Data Extractor:	Location: US	Respiratory disease, Risk	Cardiovascular	COVID-19 CFR classification (HH or
DOS		factors, Cancer, Blood	disease: ND	LL) and mortalities of other
I	Study dates: January 22 – November 22, 2020	diseases	Cardiomyopathy &	diseases:
Reviewer: CS			myocarditis: ND	
	Inclusion criteria:	Medical Condition:	Peripheral vascular	Cerebrovascular disease:
Study design:	cumulative COVID-19 cases and deaths collected from	Cerebrovascular disease:	disease: ND	• HH: 1.267 (95% CI: 1.169-
Predictive	USAFacts; age-adjusted mortality rates of 20 covariates	NR	Hypertensive heart:	1.374), p<0.001
modeling	collected from <i>University of Washington Global Health</i>	Cardiovascular disease: NR	ND	• LL: 1.053 (95% CI: 0.977-1.135),
Charles Obtantions	Data Exchange	Cardiomyopathy &	Atrial fibrillation: ND	p=0.180
Study Objective:	Exclusion criteria: counties with less than 16 reported	myocarditis: NR	Hepatitis: ND	Cardiovascular disease:
To apply spatial and statistical	deaths were excluded from subsequent analyses	Peripheral vascular disease:	Asthma: ND	• HH: 0.817 (95% CI: 0.759-
analysis to better	deaths were excluded from subsequent analyses	NR	Pulmonary	0.880), p<0.001
understand the		Hypertensive heart: NR	,	• LL: 0.949 (95% CI: 0.884-1.018),
geospatial		Atrial fibrillation: NR	sarcoidosis &	p=0.142
distributions of		Hepatitis: NR	interstitial lung	Cardiomyopathy & myocarditis:
the COVID-19		Asthma: NR	disease: ND	• HH: 1.233 (95% CI: 1.113-
mortality rate		Pulmonary sarcoidosis &	COPD: ND	1.366), p<0.001
(MR) and case		interstitial lung disease: NR	Tracheal, bronchus,	1 - 1
fatality rate		COPD: NR	and lung cancer: ND	• LL: 1.130 (95% CI: 1.024-1.246),
(CFR) in US		Tracheal, bronchus, and	Pancreatic cancer:	p=0.014
, ,		lung cancer: NR	ND	Peripheral vascular disease:
IVA Score: 22		Pancreatic cancer: NR	Leukemia: ND	• HH: 1.034 (95% CI: 0.729-
(moderate)		Leukemia: NR	Hodgkin lymphoma:	1.468), p=0.851
		Hodgkin lymphoma: NR	ND	• LL: 0.953 (95% CI: 0.711-1.276),
		Mesothelioma: NR	Mesothelioma: ND	p=0.745
		HIV/AIDS: NR	HIV/AIDS: ND	Hypertensive heart:
		HIV/AIDS. NK		• HH: 1.214 (95% CI: 1.126-
		High high (IIII), counties	Severity Measure(s):	1.309), p<0.001
		High-high (HH): counties	NR	• LL: 1.034 (95% CI: 0.963-1.109),
		with high COVID-19		p=0.361
		mortality surrounded by	Clinical marker: NR	Atrial fibrillation:
		counties with high COVID-		• HH: 1.324 (95% CI: 1.130-
		19 mortalities	Treatment/	1.551), p=0.001
			Associated Therapy:	• LL: 0.992 (95% CI: 0.872-1.128),
		Low-low (LL): counties with	NR	p=0.902
		low COVID-19 mortality		Hepatitis:
		surrounded by counties	Outcome Definitions:	· '
		with low COVID-19	COVID-19 case	• HH: 5.602 (95% CI: 1.265-
		mortalities	fatality ratio (CFR):	24.814), p=0.023

Study	Population and Setting	Intervention	Definitions	Results
			proportion of	• LL: 0.808 (95% CI: 0.187-3.483),
		Control/Comparison group:	recorded death over	p=0.774
		Non-significant (NS): non-	the confirmed cases	Asthma:
		significant counties	001/12 1011 111	• HH: 4.584 (95% CI: 2.583-
			COVID-19 Mortality	8.137), p<0.001
			rate (MR): mean COVID-19 mortality	• LL: 0.818 (95% CI: 0.461-1.452),
			rate per 100,000	p=0.492
			individuals	Interstitial lung disease:
				• HH: 1.218 (95% CI: 1.012-
			Comments: none	1.466), p=0.037
				• LL: 0.826 (95% CI: 0.702-0.972),
				p=0.021
				COPD:
				• HH: 0.996 (95% CI: 0.976-
				1.016), p=0.705
				• LL: 1.028 (95% CI: 1.010-1.046),
				p=0.002
				Tracheal, bronchus, and lung
				cancer:
				• HH: 1.043 (95% CI: 1.016-
				1.070), p=0.002
				• LL: 0.967 (95% CI: 0.947-0.986),
				p=0.001
				Pancreatic cancer:
				• HH: 0.474 (95% CI: 0.377-
				0.596), p<0.001
				• LL: 1.343 (95% CI: 1.082-1.666),
				p=0.007
				Leukemia:
				• HH: 2.172 (95% CI: 1.518-
				3.106), p<0.001
				• LL: 0.432 (95% CI: 0.325-0.573),
				p<0.001
				Hodgkin lymphoma:
				• HH: 0.008 (95% CI: 0.000-
				1.196), p=0.059
				• LL: 48.361 (95% CI: 0.644-
				3633.023), p=0.078
				Mesothelioma:

Study	Population and Setting	Intervention	Definitions	Results
				• HH: 0.847 (95% CI: 0.522-
				1.375), p=0.502
				• LL: 1.954 (95% CI: 1.130-3.380),
				p=0.017
				HIV/AIDS:
				• HH: 0.850 (95% CI: 0.753-
				0.960), p=0.009
				• LL: 2.061 (95% CI: 1.641-2.588),
				p<0.001
				p<0.001
				Severity of Condition: NR
				Duration of Condition: NR
				Treatment/ Associated Therapy: NR
				Comorbid Conditions: NR
				Risk Markers:
				Alcohol use disorder:
				• HH: 1.088 (95% CI: 0.965-
				1.227), p=0.168
				• LL: 1.149 (95% CI: 1.044-1.266),
				p=0.005
				Drug use disorder:
				• HH: 1.016 (95% CI: 0.972-
				1.061), p=0.491
				• LL: 0.960 (95% CI: 0.928-0.992),
				p=0.016
				Long-term Sequelae: NR
Author : Oh ⁷	Population: N=122,040 n=7,780 COVID-19 +	Health Condition Category:	Medical	Severe COVID-19:
V 2024	Continue National Haalth Inc.	Cerebrovascular disease,	Condition(s):	aOR: Multivariable Logistic
Year: 2021	Setting: National Health Insurance Service database	Chronic heart disease,	ICD-10 codes were	Regression: Multivariable Logistic
Data Extractor	Location: South Koroa	Chronic liver disease,	used to evaluate CRDs and other	Regression
Data Extractor: MW	Location: South Korea	Neurocognitive disorders, Diabetes, Chronic kidney	comorbid conditions	Mortality
IVIVV	Study dates: January 1-June 26, 2020	disease, Chronic lung	in the study	Mortality: Cerebrovascular disease:
Reviewer: CS	Study dates. January 1-June 20, 2020	disease, Risk factors,	population:	• aOR: 0.57 (95% CI: 0.38-0.87);
vicwei. co	Inclusion criteria: Individuals ≥20 years old, had a	Immunocompromised	population.	p=0.009
Study design:	respiratory disease diagnosis by the International	status, Blood disorders,		1
Cohort	Classification of Diseases codes, and prescription	Cancer		Hemiplegia/paraplegia:

Study	Population and Setting	Intervention	Definitions	Results
	information concerning drugs and/or procedures from		Cerebrovascular	• aOR: 1.92 (95% CI: 1.03-3.59);
Study Objective:	2015-2020 were included. COVID-19 negative individuals	Medical Condition, n/N	disease: G45.x, G46.x,	p=0.040
To investigate	were extracted from the national database using	(%):	H34.0, I60.x - I69.x	Hypertension:
various chronic	stratification methods with regard to age, sex, and	Cerebrovascular disease:	Hemiplegia/paraplegi	• aOR: 1.36 (95% CI: 0.89-2.06);
respiratory	residence in February 2020.	5763/122,040 (4.6%)	a: G04.1, G11.4,	p=0.153
diseases (CRDs)		Hemiplegia/paraplegia:	G80.1, G80.2, G81.x,	Congestive heart failure:
that affect the	Exclusion criteria: NR	568/122,040 (0.5%)	G82.x, G83.0 - G83.4,	• aOR: 1.91 (95% CI: 1.38-2.66);
risk of COVID-19		Hypertension:	G83.9	p<0.001
among the		32,727/122,040 (26.3%)	Hypertension: ND	Myocardial infarction:
general		Congestive heart failure:	Congestive heart	'
population in		3683/122,040 (3.0%)	failure: 109.9, 111.0,	• aOR: 0.79 (95% CI: 0.47-1.33);
South Korea, and		Myocardial infarction:	113.0, 113.2, 125.5,	p=0.374
to examine the		1187/122,040 (1.0%)	142.0, 142.5 - 142.9,	Dementia:
effect of		Mild liver disease:	143.x, 150.x, P29.0	• aOR: 1.61 (95% CI: 1.11-2.32);
different CRDs		13,612/122,040 (10.9%)	Myocardial	p=0.011
on hospital		Moderate or severe liver	infarction: I21.x,	Diabetes without chronic
mortality among		disease: 146/122,040	122.x, 125.2	complication:
patients with		(0.1%)	Dementia: F00.x -	• aOR: 1.87 (95% CI: 1.35-2.59);
COVID-19 in		Dementia: 3926/122,040	F03.x, F05.1, G30.x,	p<0.001
South Korea.		(3.2%)	G31.1	Diabetes with chronic
IVA Score: 25		Diabetes without chronic complication:	Diabetes without chronic complication:	complication:
(Moderate)		13,781/122,040 (11.1%)	E10.0, E10.1, E10.6,	• aOR: 1.61 (95% CI: 1.06-2.45);
(iviouerate)		Diabetes: with chronic	E10.8, E10.9, E11.0,	p=0.027
		complication: 4255/122,040	E11.1, E11.6, E11.8,	Renal disease:
		(3.4%)	E11.9, E12.0, E12.1,	• aOR: 1.47 (95% CI: 0.87-2.47);
		Renal disease:	E12.6, E12.8, E12.9,	p=0.148
		1392/122,040 (1.1%)	E13.0, E13.1, E13.6,	Any chronic respiratory disease:
		Any chronic respiratory	E13.8, E13.9, E14.0,	• aOR: 1.19 (95% CI: 0.86-1.64);
		disease: 36,365/122,040	E14.1, E14.6, E14.8,	p=0.299
		(29.2%)	E14.9	COPD:
		Chronic obstructive	Diabetes with chronic	• aOR: 1.56 (95% CI: 1.06-2.2);
		pulmonary disease (COPD):	complication: E10.2 -	p=0.024
		4488/122,040 (3.6%)	E10.5, E10.7, E11.2 -	Asthma:
		Asthma: 33,858/122,040	E11.5, E11.7, E12.2 -	• aOR: 1.03 (95% CI: 0.76-1.41);
		(27.2%)	E12.5, E12.7, E13.2 -	p=0.834
		Interstitial lung disease:	E13.5, E13.7, E14.2 -	Interstitial lung disease:
		421/122,040 (0.3%)	E14.5, E14.7	• aOR: 1.83 (95% CI: 0.74-4.55);
		Lung disease d/t external	Renal disease: I12.0,	p=0.193
		agent: 437/122,040 (0.4%)	I13.1, N03.2 - N03.7,	Lung disease d/t external agent:
		Obstructive sleep apnea:	N05.2 - N05.7, N18.x,	• aOR: 3.54 (95% CI: 1.70-7.38);
		550/122,040 (0.4%)	N19.x, N25.0, Z49.0 -	p<0.001
		Tuberculosis of lung:	Z49.2, Z94.0, Z99.2	Obstructive sleep apnea:
		608/122,040 (0.5%)		obstructive sieep aprilea.

Study	Population and Setting	Intervention	Definitions	Results
		AIDS/HIV: 32/122,040	Any chronic	• aOR: 0.47 (95% CI: 0.06-3.94);
		(0.0%)	respiratory disease:	p=0.486
		Peripheral vascular disease:	ND	Tuberculosis of lung:
		7198/122,040 (5.8%)	COPD: 127.8, 127.9,	aOR: 1.65 (95% CI: 0.48-5.64);
		Malignancy:	J40.x - J47.x, J60.x -	p=0.423AIDS/HIV:
		22,013/122,040 (17.7%)	J67.x, J68.4, J70.1,	• aOR: 1.43 (95% CI: 0.11-19.37);
		Metastatic solid tumor:	J70.3	p=0.788
		4072/122,040 (3.3%)	Asthma: J45	Peripheral vascular disease:
		Lung cancer: 769/122,040	Interstitial lung	• aOR: 1.19 (95% CI: 0.81-1.76);
		(0.6%)	disease: J84.9	p=0.76
			Lung disease d/t	Malignancy:
		Control/Comparison group,	external agent: J60-	• aOR: 1.07 (95% CI: 0.78-1.46);
		n/N (%):	J70	,
		No cerebrovascular disease:	Obstructive sleep	p=0.694
		116,277/122,040 (95.3%)	apnea: G47.33	Metastatic solid tumor:
		No hemiplegia/paraplegia:	Tuberculosis of lung:	• aOR: 1.37 (95% CI: 0.85-2.19);
		121,472/122,040 (99.5%)	A15	p=0.192
		No hypertension:	AIDS/HIV: B20.x -	Lung cancer:
		89,313/122,040 (73.2%)	B22.x, B24.x	• aOR: 1.82 (95% CI: 0.80-4.14);
		No congestive heart failure:	Peripheral vascular	p=0.154
		118,357/122,040 (97.0%)	disease: I70.x, I71.x,	
		No myocardial infarction:	173.1, 173.8, 173.9,	Severity of Condition:
		120,853/122,040 (99.0%)	177.1, 179.0, 179.2,	Mortality:
		No mild liver disease:	K55.1,	Charlson comorbidity index:
		108,428/122,040 (88.8%) No moderate or severe liver	K55.8, K55.9, Z95.8, Z95.9	comorbidity index:
		disease: 121,894/122,040	Malignancy: except	• aOR: 1.80 (95% CI: 1.32-2.44);
		(99.9%)	malignant neoplasm	p<0.001
		No dementia:	of skin; C00.x - C26.x,	
		118,114/122,040 (96.8%)	C30.x - C34.x, C37.x -	Duration of Condition: NR
		No diabetes without	C41.x, C43.x, C45.x -	_
		chronic complication:	C58.x, C60.x - C76.x,	Treatment/ Associated Therapy:
		108,259/122,040 (88.7%)	C81.x - C85.x, C88.x,	NR
		No diabetes with chronic	C90.x - C97.x	
		complication:	Metastatic solid	Comorbid Conditions: NR
		117,785/122,040 (96.5%)	tumor: C77.x - C80.x	
		No renal disease:	Lung cancer: C34	Risk Markers:
		120,648/122,040 (98.9%)		Mortality:
		No chronic respiratory	Severity Measure(s):	Age, 10-year increase:
		disease: 85,675/122,040	Mild liver disease:	• aOR: 2.85 (95% CI: 2.40-3.38);
		(70.2%)	B18.x, K70.0 - K70.3,	p<0.001
		No COPD: 117,552/122,040	K70.9, K71.3 - K71.5,	Sex, male:
		(96.3%)	K71.7, K73.x, K74.x,	• aOR: 2.12 (95% CI: 1.55-2.88);
			, ,	p<0.001

Study	Population and Setting	Intervention	Definitions	Results
		No asthma: 88,182/122,040 (72.3%) No interstitial lung disease: 121,619/122,040 (99.7%) No lung disease d/t external agent: 121,603/122,040 (99.6%) No obstructive sleep apnea: 121,490/122,040 (99.5%) No tuberculosis of lung: 121,432/122,040 (99.5%) No AIDS/HIV: 122,008/122,040 (99.9%) No peripheral vascular disease: 114,842/122,040 (94.1%) No malignancy: 100,027/122,040 (82.0%) No metastatic solid tumor: 117,968/122,040 (96.7%) No lung cancer: 121,271/122,040 (99.4%)	K76.0, K76.2 - K76.4, K76.8, K76.9, Z94.4 Moderate or severe liver disease: 185.0, 185.9, 186.4, 198.2, K70.4, K71.1, K72.1, K72.9, K76.5, K76.6, K76.7 Charlson comorbidity index: NDND Clinical marker: NR Treatment/ Associated Therapy: NR Outcome Definitions: Mortality: ND ICU admission: NR Intubation: NR Ventilation: NR Hospitalization: NR Non-elective readmissions: NR Comments: The paper reports different study sizes in the text (N=122,040; n=7,669 COVID-19+) and Figure 1 (N=124,330; n=7,780 COVID-19+).	Income level by quartile grouping: Q1 (lowest): 1 Q2:

B.3.c. Internal Validity Assessments of Extracted Studies

Table 9. Internal Validity Assessments of Extracted Studies reporting the Association between Interstitial Lung Diseases and Severe COVID-19 Outcomes

	Author Year	Aveyard 2021 ¹	Beltramo 2021 ²	Drake 2020³	Estiri 2021 ⁴	Halalau 2021 ⁵	Kokturk 2021 ⁶	Mollalo 2021 ⁸	Oh 2021 ⁷
	Outcome	Mortality, ICU, Hospitalization	Mortality, ICU admission	Mortality, ventilation	Hospitalization	Hospitalization	Mortality	Mortality for COVID- 19 and other diseases	Mortality
Domain	Signaling question	data extracted from medical records	Data extracted from hospital records	data extracted from medical records	medical records	Data extracted from electronic medical records	Data extracted from medical records	Data from USAFacts and UW Global Health Data Exchange	Data extracted from database
	Design appropriate to			1	1	1	1	1	1
	research question	1	1	1	1	1	1	1	1
	Well described population	1	1	1	1	1	1	1	1
	Well described setting	1	1	1	1	1	1	1	1
Study Elements	Well described intervention/ exposure	1	1	1	1	1	1	1	1
	Well described control/ comparator	1	1	1	1	1	1	1	1
	Well described outcome	1	1	1	1	1	1	1	1
	Clear timeline of exposures/ interventions and outcomes	1	0	1	1	1	1	0	1
	Randomization appropriately performed	0	0	0	0	0	0	0	0
Selection Bias: Sampling	Allocation adequately concealed	0	0	0	0	0	0	0	0
	Population sampling appropriate to study design	1	1	1	1	1	1	1	1
	Attrition not significantly different between groups	1	1	1	1	1	1	1	1
Selection Bias: Attrition	Attrition <10-15% of population	1	1	1	1	1	1	1	1
	Attrition appropriately analyzed	1	1	1	1	1	1	1	1
Information Bias:	Measure of intervention/ exposure is valid	1	1	1	1	1	1	1	1
Measurement	Measure of outcome is valid	1	1	1	1	1	1	0	1

	Author Year	Aveyard 2021 ¹	Beltramo 2021 ²	Drake 2020 ³	Estiri 2021 ⁴	Halalau 2021 ⁵	Kokturk 2021 ⁶	Mollalo 2021 ⁸	Oh 2021 ⁷
	Outcome	Mortality, ICU, Hospitalization	Mortality, ICU admission	Mortality, ventilation	Hospitalization	Hospitalization	Mortality	Mortality for COVID- 19 and other diseases	Mortality
Domain	Signaling question	data extracted from medical records	Data extracted from hospital records	data extracted from medical records	medical records	Data extracted from electronic medical records	Data extracted from medical records	Data from USAFacts and UW Global Health Data Exchange	Data extracted from database
and	Fidelity to intervention is			0	0	0	0	0	0
Misclassification	measured	0	0	U	U	U	U	U	U
	Fidelity to intervention is valid	0	0	0	0	0	0	0	0
	Prospective study	1	1	1	1	1	1	0	1
	Adequately powered to detect result	0	0	0	1	0	0	0	1
	Outcome assessor blinded	0	0	0	0	0	0	0	0
	Study participant blinded	0	0	0	0	0	0	0	0
Information	Investigator/ data analyst blinded	0	0	0	0	0	0	0	0
Bias: Performance &	Data collection methods described in sufficient detail	1	1	1	1	1	1	1	1
Detection	Data collection methods appropriate	1	1	1	1	1	1	1	1
	Sufficient follow up to detect outcome	1	1	1	1	1	1	1	1
	Appropriate statistical analyses for collected data	1	1	1	1	1	1	1	1
Information Bias: Analytic	Appropriate statistical analyses are conducted correctly	1	1	1	1	1	1	1	1
	Confidence interval is narrow	0	1	0	0	0	0	1	0
	Potential confounders identified	1	1	1	1	1	1	1	1
Confounding	Adjustment for confounders in study design phase	0	0	0	0	0	0	0	0

	Author Year	Aveyard 2021 ¹	Beltramo 2021 ²	Drake 2020³	Estiri 2021 ⁴	Halalau 2021 ⁵	Kokturk 2021 ⁶	Mollalo 2021 ⁸	Oh 2021 ⁷
	Outcome	Mortality, ICU, Hospitalization	Mortality, ICU admission	Mortality, ventilation	Hospitalization	Hospitalization	Mortality	Mortality for COVID- 19 and other diseases	Mortality
Domain	Signaling question	data extracted from medical records	Data extracted from hospital records	data extracted from medical records	medical records	Data extracted from electronic medical records	Data extracted from medical records	Data from USAFacts and UW Global Health Data Exchange	Data extracted from database
	Adjustment for confounders in data analysis phase	1	1	1	1	0	1	1	1
Reporting Bias	All pre-specified outcomes are adequately reported	1	1	1	1	1	1	1	1
Other Bias	No other sources of bias	1	1	1	1	1	1	1	1
COI	Funding sources disclosed and no obvious conflict of interest	1	1	1	1	1	1	1	1
SCORE	Threat to internal validity Low, Moderate, High	24 Moderate	24 Moderate	24 Moderate	25 Moderate	23 Moderate	24 Moderate	22 Moderate	25 Moderate

C. References

- 1. Aveyard P, Gao M, Lindson N, Hartmann-Boyce J, Watkinson P, Young D, et al. Association between pre-existing respiratory disease and its treatment, and severe COVID-19: a population cohort study. The Lancet Respiratory Medicine 2021;9(8):909-923.
- 2. Beltramo G, Cottenet J, Mariet A-S, Georges M, Piroth L, Tubert-Bitter P, et al. Chronic respiratory diseases are predictors of severe outcome in COVID-19 hospitalised patients: a nationwide study. European Respiratory Journal 2021:2004474.
- 3. Drake TM, Docherty AB, Harrison EM, Quint JK, Adamali H, Agnew S, et al. Outcome of hospitalization for COVID-19 in patients with interstitial lung disease an international multicenter study. American Journal of Respiratory and Critical Care Medicine 2020;202(12):1656-1665.
- 4. Estiri H, Strasser ZH, Klann JG, Naseri P, Wagholikar KB, Murphy SN. Predicting COVID-19 mortality with electronic medical records. npj Digital Medicine 2021;4(1).
- 5. Halalau A, Odish F, Imam Z, Sharrak A, Brickner E, Lee PB, et al. Epidemiology, Clinical Characteristics, and Outcomes of a Large Cohort of COVID-19 Outpatients in Michigan. Int J Gen Med 2021;14:1555-1563.
- 6. Kokturk N, Babayigit C, Kul S, Duru Cetinkaya P, Atis Nayci S, Argun Baris S, et al. The predictors of COVID-19 mortality in a nationwide cohort of Turkish patients. Respir Med 2021;183:106433.

- 7. Oh TK, Song IA. Impact of coronavirus disease-2019 on chronic respiratory disease in South Korea: an NHIS COVID-19 database cohort study. BMC Pulmonary Medicine 2021;21(1).
- 8. Mollalo A, Rivera KM, Vahabi N. Spatial statistical analysis of pre-existing mortalities of 20 diseases with COVID-19 mortalities in the continental United States. Sustainable Cities and Society 2021;67:102738.

D. Abbreviations

Acronym	Full
95% CI	95% confidence interval
aHR	adjusted hazard ratio
aOR	adjusted odds ratio
BMI	body mass index
BPD	bronchopulmonary dysplasia
CF	cystic fibrosis
CFR	case fatality ratio
COI	conflict of interest
COPD	chronic obstructive pulmonary disease
CRD	chronic respiratory disease
ECMO	extracorporeal membrane oxygenation
EHR	electronic health record
EMR	electronic medical record
IQR	Interquartile range
GLM	generalized linear model
HH	high-high counties
HR	hazard ratio
ICD10	International Classification of Diseases 10
ICNARC	Intensive Care National Audit and Research Centre
ICS	inhaled corticosteroids
ICU	intensive care unit
ILD	interstitial lung disease
IPF	idiopathic pulmonary fibrosis
IVA	Internal validity assessments
LL	low-low counties
MR	mortality rate
ND	not defined

NR	not reviewed
OR	odds ratio
PCR	polymerase chain reaction
PECO	population, exposure, comparator, and outcomes
PMSI	Programme de Medicalisation des Systemes d'Information
RR	Rate ratio
RT-PCR	real time polymerase chain reaction