RESEARCH PROTOCOL:

Characterizing Health Associated Risks, and Your Baseline Disease In SARS-COV-2 (CHARYBDIS): protocol for an OHDSI network study

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# 1. List of Abbreviations

|  |  |
| --- | --- |
| AKI | Acute Kidney Injury |
| ARDS | Acute Respiratory Distress Syndrome |
| CHARYBDIS | Characterizing Health Associated Risks, and Your Baseline Disease In SARS-COV-2 (CHARYBDIS) |
| CDM | Common Data Model |
| COVID-19 | 2019 novel coronavirus / coronavirus disease 2029 |
| DVT | Deep Venous Thrombosis |
| EHDEN | European Health Data and Evidence Network |
| HIV | Human Immunodeficiency Virus |
| OMOP | Observational Medical Outcomes Partnership |
| OHDSI | Observational Health Data Science and Informatics |
| PE | Pulmonary Embolism |
| RxNorm | US-specific terminology in medicine that contains all medications available on the US market |
| SARS-CoV-2 | Severe Acute Respiratory Syndrome-Coronavirus-2 |
| SNOMED | Systematized Nomenclature of Medicine |
| WHO | World Health Organization |

# 

# 2. Responsible Parties

## 2.1. Investigators and Authors

|  |  |
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## 

Authorship will vary from paper to paper. The Responsible Parties involved in this protocol take accountability for the overarching protocol, package development, providing assistance to sites running the analysis and ensuring site-specific governance is adhered to in all publications generated from this protocol.

For subgroup analysis, the following table represents individuals who will be contributing to follow-on work to write manuscripts on these specific topics. This list is not exhaustive of all possible authors for future papers:

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| --- | --- | --- |
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| Asieh Golozar | Regeneron Pharmaceuticals, NY USA; Johns Hopkins Bloomberg School of Public Health, Baltimore, MD, USA | COVID-19 testing |
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| Julianna Kohler | United States Agency for International Development, Washington, DC, USA | COVID-19 in HIV/Immunocompromised (inclusive of Hepatitis C) and tuberculosis populations |
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| Seng Chan You | Department of Biomedical Informatics, Ajou University | COVID-19 uncomplicated hypertension |
| Martina Recalde | Fundació Institut Universitari per a la recerca a l'Atenció Primària de Salut Jordi Gol i Gurina (IDIAPJGol), Barcelona, Spain; Universitat Autònoma de Barcelona, Spain | COVID-19 in hypertension, type 2 diabetes patients, obesity |
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| Daniel Prieto-Alhambra | 1) Fundació Institut Universitari per a la recerca a l'Atenció Primària de Salut Jordi Gol i Gurina (IDIAPJGol), 2) Centre for Statistics in Medicine, NDORMS, University of Oxford | COVID-19 drug utilization during treatment (descriptive study) |

## 2.2 Sponsor

This study was undertaken by Observational Health Data Science and Informatics (OHDSI), an open collaboration. **DPA** receives partial support from the National Institute for Health Research Oxford Biomedical Research Centre and Senior Research Fellowship. **SCY** receives grant funding from the Korean Ministry of Health & Welfare and from the Korean Ministry of Trade, Industry & Energy. **GH** receives grant funding from the US National Institutes of Health and contracts from Janssen Research and Development. **PBR**, **AGS, GR** and **AS** are employees of Janssen Research and Development and shareholders in Johnson & Johnson. **KK** is an employee of IQVIA. **APU** receives grants from MRC and Fundacion Alfonso Martin Escudero. Participating data owners will be responsible for self-reporting any grants funding the conversion and maintenance of their OMOP CDM. Disclosures will be reported in accordance with publication policies of journals papers are submitted to. No other disclosures reported.

# 3. Abstract

In this study we will describe the baseline demographic and clinical characteristics, as well as the occurrence of treatments and outcomes of individuals tested or with COVID-19 in 2019-2020. We will compare these with the characteristics of individuals with Influenza in 2017-2018.

# 4. Amendments and Updates

|  |  |  |  |  |
| --- | --- | --- | --- | --- |
| **Number** | **Date** | **Section of study protocol** | **Amendment or update** | **Reason** |
| V2 | 27APR2020 | All | Incorporating community feedback | Collaboration |
| V3 | 04MAY2020 | All | Incorporating final changes from core team | Consistency |
| V4 | 10JUN2020 | All | Updated name to CHARYBDIS and added final cohort definitions | Consistency |
| V5 | 11JUN2020 | Stratifications  Features of Interest | Adding additional requirements from DPA/PBR | Collaboration |
| V5.1 | 16JUN2020 | Study leads  Appendix | Adding additional study leads  Fixing ATLAS link error | Consistency |

# 5. Rationale and Background

Since December 31st 2019, when the first case was reported to WHO, infections by SARS-CoV-2 have spread worldwide. Our current understanding of the disease's clinical outcomes is limited to case series in hospitalised populations. The baseline characteristics of 138 patients with Coronavirus disease 2019 (COVID-19) hospitalized in Wuhan due to pneumonia, for example, have been reported.1 In another study, a series of 21 cases admitted to an ICU in Washington state reported clinical and demographic characteristics at hospital admission.2 A more recent study has described the baseline characteristics and complications in 393 confirmed COVID-19 cases admitted to two hospitals in New York City. 3

Government and healthcare strategies for testing and diagnosing suspected COVID-19 patients vary extensively between countries and by time. Some countries have been urged to implement widespread testing following WHO's recommendation, and other countries are testing fewer cases often due to lack of availability of laboratory tests.4 Often these strategies focus on testing more severe and high-risk patients. Selective testing might lead to overestimation of poor outcomes and mortality. An understanding of the characteristics of tested populations by country government policies would better inform epidemiology studies and understanding of disease burden, magnitude and outcomes.

As more routinely-collected data emerges on individuals with COVID-19, a federated network study could provide a fuller picture of these individuals, particularly among specific population groups that might be more susceptible to COVID-19 and for which limited information is currently available, such as the pediatric or pregnant populations, or persons with comorbidities such as diabetes, hypertension or cancer. Moreover, the use of such data to inform comparisons with previous viral outbreaks, like regular seasonal influenza epidemics, could also help us to better understand the profiles of individuals with COVID-19. Data assets mapped to the Observational Medical Outcomes Partnership (OMOP) common data model (CDM)5 provide a unique opportunity to make a difference in the current crisis, allowing for robust analyses to be performed in a timely across a network of sites.

# 6. Objective

The primary objective of the Characterizing Health Associated Risks, and Your Baseline Disease In SARS-COV-2 (CHARYBDIS) study is to describe the baseline demographic and clinical characteristics, as well as treatments and occurrence of outcomes of interest among individuals tested for SARS-CoV-2 or diagnosed with COVID-19 after December 1st 2019, overall and by sex, age and comorbidities. We will also describe the characteristics and outcomes of the population with seasonal influenza infection between September 1st 2017 and April 1st 2018 as a benchmark.

(***Note:*** *For computational efficiency, this analysis will deviate from the prior OHDSI Hospitalized in COVID-19 analysis which included a larger sample of years (2014-2019) of the seasonal influenza population and a separate cohort for H1N1 (2009-2010). The computational run time to pull these cohorts required a significant investment by data owners and, at the time of authoring this analysis, it was decided a smaller window of years – particularly years in which flu severity was highest – would be the most efficient use of compute.)*

# 7. Methods

### 7.1 Data Sources

This study is a multinational cohort study describing the baseline characteristics, treatments and outcomes of 12 cohorts, namely:

1. SARS-CoV-2 tested
2. SARS-CoV-2 tested positive
3. SARS-CoV-2 tested with a COVID-19 diagnosis record or a positive test
4. COVID-19 diagnosed or with a positive test
5. COVID-19 diagnosed/positive tested and hospitalized
6. COVID-19 diagnosed/positive tested, hospitalized, and requiring intensive services
7. Influenza between 2017-2018 diagnosed or with a positive test
8. Influenza between 2017-2018 diagnosed/positive tested and hospitalized
9. Influenza between 2017-2018 diagnosed/positive tested, hospitalized, and requiring intensive services

A South Korean national claims database, multiple U.S. (New York City, San Francisco, Boston, Denver, US Veterans’ Affairs) health systems databases, and one Spanish EHR database are accumulating COVID-19 patients and have tested the operability of our analysis package at their sites (**Table 1**). As more data becomes available, we will include additional databases that are formatted to the OMOP-CDM. These will reflect the clinical experience of patients from seven European countries (Belgium [general practice EHR], Netherlands [general practice EHR], Germany [general practice EHR, hospital EHR], France [general practice EHR, outpatient specialist EHR], Spain [general practice EHR, outpatient specialist EHR and linked inpatient administrative data], Estonia [EHR, claims, and registry data], and the United Kingdom [general practice EHR, linked to hospital EHR]), the United States of America (general practice EHR, outpatient specialist EHR, hospital EHR, insurance claims], South Korea [EHR, claims, and registry data], Japan [insurance claims], and any other country with mapped data willing to participate.

The study will be conducted using data from real world data sources that have been mapped to the OMOP Common Data Model in collaboration with the Observational Health Data Sciences and Informatics (OHDSI) and European Health Data and Evidence Network (EHDEN) initiatives. The OMOP Common Data Model (<https://github.com/OHDSI/CommonDataModel/wiki>) includes a standard representation of health care experiences (such as information related to drug utilization and condition occurrence), as well as common vocabularies for coding clinical concepts, and enables consistent application of analyses across multiple disparate data sources.6

**Table 1**. Data sources formatted to the OMOP CDM that as of 10 JUN 2020 include COVID-19 patients.

|  |  |  |  |  |
| --- | --- | --- | --- | --- |
| **Data source** | **Source population** | **Sample size** | **Data type** | **Longitudinal history** |
| South Korea: Health Insurance and Review Assessment (HIRA) | All citizens in South Korea | ≈ 50 million | Administrative fee-for-service claims data collected for healthcare reimbursement, including healthcare services such as treatments, pharmaceuticals, procedures, and diagnoses. | 5-years of available look-back (data older than 5-years is deleted from the database) |
| Columbia University Irving Medical Center | Patients of the Columbia University Irving Medical Center (New York City, USA) | ≈ 6 million | General practice electronic health records, outpatient specialist electronic health records, inpatient hospital electronic health records, hospital billing/summary | 1989 (1978 for diagnoses) |
| Stanford Medicine Research Data Repository (STARR) | Patients of Stanford University (San Francisco Bay Area, USA) | ≈ 3 million | General practice electronic health records, outpatient specialist electronic health records, inpatient hospital electronic health records, hospital billing/summary | 2008 |
| Tufts Clinical Academic Research Enterprise Trust (CLARET) | Patients of Tufts Medical Center (Boston, USA) | ≈ 1 million | Electronic medical record data on approximately 1 million patients who received care beginning in 2006 at Tufts Medical Center (TMC). TMC is an academic medical center that includes Tuft Medical Center's main downtown Boston hospital for adult patients, the Floating Hospital for Children, and associated primary and specialty care clinics. CLARET contains TMC's EHR data fused with data on the same patients from TMC's CoC accredited tumor registry, its oncology EHR, and death data from the Massachusetts State Registry of Vital Statistics. EHR data streams ingested into CLARET include controlled vocabulary data on all domains except cost, and select free text sources and devices. | 2006 |
| Veteran’s Affairs | Patients of the Veterans Affairs in the United States | ≈ 9 million | VA OMOP data reflects the national Department of Veterans Affairs health care system, which is the largest integrated provider of medical and mental health services in the United States. Care is provided at 170 VA Medical Centers and 1,063 outpatient sites serving more than 9 million enrolled Veterans each year. | 2000 |
| UC Health Data Compass | Patients of UC Denver and Children’s Hospital of Colorado (Denver, USA) | To be confirmed in analysis | Health Data Compass (HDC) is a multi-institutional data warehouse. HDC contains inpatient and outpatient electronic medical data including patient, encounter, diagnosis, procedures, medications, laboratory results from two electronic medical record systems (UCHealth and Children's Hospital of Colorado), state-level all-payers claims data, and the Colorado death registry. | To be confirmed in analysis |
| University of Washington | Patients of UW Medical Center (Seattle, USA) | ≈ 5 million | The clinical data warehouse of University of Washington Medical Center - Montlake, Harborview Medical Center, UW Medical Center – Northwest, Seattle WA, based on its current electronic health record systems, with data spanning over 10 years and including roughly 5 million patients. We also have de-identified and annotated xrays for COVID-19 patients who were imaged in a specific COVID-19 PACS. | 2009 |
| The Information System for Research in Primary Care (SIDIAP) | General population in Catalonia, Spain | ≈ 8 million | Electronic health records from primary care partially linked to inpatient data. SIDIAP is also linked to pharmacy dispensations and primary care laboratories. Healthcare is universal and tax-payer funded in the region, and primary care physicians are gatekeepers for all care and responsible for repeat prescriptions. | 2006 |

### 7.2 Study design

The study will be an observational cohort study based on routinely-collected health care data which has been mapped to the Observational Medical Outcomes Partnership (OMOP) Common Data Model (CDM). Cohorts of individuals with COVID-19 or influenza will be identified. Characteristics of these individuals at their index date will be described. Treatments and outcomes of these individuals after their index date will be described.

### 7.3 Target cohorts

COVID-19 cohorts

**Target Cohort #1:** persons in the *tested for SARS-CoV-2 cohort* will:

* have a record of a first test for SARS-CoV-2 (index event) after December 1st 2019.

**Target Cohort #2:** persons in the *tested positive for SARS-CoV-2 cohort* will:

* have a record of a positive test for SARS-CoV-2 (index event will be the earliest test date that occurs within 7 days of positive test result) after December 1st 2019.

***Note: For computational efficiency, there will not be a separate tested negative SARS-CoV-2 cohort.***

**Target Cohort #3:** persons in the *tested for SARS-CoV-2 with a COVID-19 diagnosis record or a SARS-CoV-2 positive test cohort* will:

* have a record of a test for SARS-CoV-2 (index event first test) after December 1st 2019, and either,
  + have a record of COVID-19 diagnosis, or
  + have a record of a positive test for SARS-CoV-2.

**Target Cohort #4:** persons in the *COVID-19 diagnosis or a SARS-CoV-2 positive test cohort* will:

* have a record of a positive test for SARS-CoV-2 after December 1st 2019, or,
* have a record of a COVID-19 diagnosis (index date is date of the first of these 2 events).

**Target Cohort #5:** persons in the *hospitalized with a COVID-19 diagnosis record or a SARS-CoV-2 positive test cohort* will:

● have a hospitalization (index event) after December 1st 2019, and,

● a record of a COVID-19 diagnosis or a SARS-CoV-2 positive test result in the 3 weeks prior and up to end of hospitalization, and,

● have no COVID-19 associated hospitalization in the six months prior to the index event.

**Target Cohort #6:** persons in the *hospitalized and requiring intensive services with a COVID-19 diagnosis record or a SARS-CoV-2 positive test cohort* will:

● have a hospitalization (index event) after December 1st 2019, and,

● a record of Mechanical Ventilation or Tracheostomy or ECMO during hospitalization, and,

● a record of a COVID-19 diagnosis or a SARS-CoV-2 positive test result in the 3 weeks prior and up to end of hospitalization, and,

* have no COVID-19 associated hospitalization in the six months prior to the index event.

**Target Cohort #7:** persons in the *hospitalized with a SARS-CoV-2 positive test cohort* will:

have a hospitalization (index event) after December 1st 2019, and,

● a record of a SARS-CoV-2 positive test result in the 3 weeks prior and up to end of hospitalization, and,

* have no COVID-19 associated hospitalization in the six months prior to the index event.

**Target Cohort #8:** persons in the *hospitalized and requiring intensive services with a SARS-CoV-2 positive test cohort* will:

have a hospitalization (index event) after December 1st 2019, and,

● a record of Mechanical Ventilation or Tracheostomy or ECMO during hospitalization, and,

● a record of a SARS-CoV-2 positive test result in the 3 weeks prior and up to end of hospitalization, and,

* have no COVID-19 associated hospitalization in the six months prior to the index event.

Influenza cohorts

**Target Cohort #9:** persons in the *influenza diagnosed or with a positive test cohort* will:

* have a record of a positive test for influenza between September 1st 2017 and April 1st 2018, or,
* have a record of influenza diagnosis (index on first event).

**Target Cohort #10:** persons in the *hospitalized with influenza diagnosis or positive test cohort* will:

● have a hospitalization (index event) between September 1st 2017 and April 1st 2018,

● with a record of a diagnosis/positive test of influenza in the 3 weeks prior and up to end of hospitalization,

● have no influenza associated hospitalization in the six months prior to the index event.

**Target Cohort #11:** persons in the *hospitalized with influenza diagnosis or positive test and require intensive services cohort* will:

● have a hospitalization (index event) between September 1st 2017 and April 1st 2018, and

● a record of Mechanical Ventilation or Tracheostomy or ECMO during hospitalization, and,

● have a record of a diagnosis/positive test of influenza in the 3 weeks prior and up to end of hospitalization, and,

● have no influenza associated hospitalization in the six months prior to the index event.

These cohorts will all be identified without any requirement for prior observation time, and also with the added restriction of having a minimum of 365 days of prior observation time available in order to assess how large proportion of subjects fall ill without previous healthcare encounters, and if their inclusion in analyses would change results or conclusions.

### 7.4 Follow-up

Different index dates will be used for each of the cohorts above:

**Target Cohort #1:**

Index event: First test for COVID-19/influenza.

Follow-up: 30 days after the index event.

**Target Cohort #2:**

Index event: Earliest test date that occurs within 7 days of positive test result.

Follow-up: 30 days after the index event.

**Target Cohort #3:**

Index event: Earliest test date that occurs within 7 days of positive test result or COVID-19/influenza diagnosis.

Follow-up: 30 days after the index event.

**Target Cohort #4,9:**

Index event: First positive test result for COVID-19/Flu or first COVID-19/Flu Diagnosis (whichever appears first).

Follow-up: 30 days after the index event.

**Target Cohort #5,7,10:**

Index event: First hospitalization, with a record of COVID-19/Flu 3w prior.

Follow-up: 30 days after the index event.

**Target Cohort #6,8,11:**

Index event: First hospitalization, with a record of COVID-19/Flu 3w prior, and Ventilation or Tracheostomy or ECMO.

Follow-up: 30 days after the index event.

### 7.5 Stratifications

Each target cohort will be analysed in full and stratified on factors based on the following pre-index characteristics, all stratum are pending meeting minimum reportable cell counts (as specified by data owners):

* Follow-up time: overall, with full 30 days follow-up, without full 30 days follow-up
* Sex (Male vs. Female)
* All reportable age groups as well as specifically:
  + Pediatrics (Age <18). If sample size allows, results will be reported stratified in the following age categories: overall (0 up to 17 years); neonates (0-28d), infants (29d- less than 1 year), toddler (1-less than 2y), early childhood (2-5y), middle childhood (6-11y), and adolescents (12-17y). In case of limited sample size for granular classification, the following categories will be used: neonates, infants, and children (0-12y); and adolescents (12-17y)
  + Elderly (Age >= 65). If sample size allows, results will be reported stratified in the following age categories: 65-84 years, and >=85 years, or in finer age strata (65-69, 70-74, 75-79, 80-84, >=85 years)
* Type 2 Diabetes Mellitus
* Hypertension
* Chronic obstructive pulmonary disease (COPD) without asthma
* Asthma without COPD
* Chronic kidney disease
* End-stage renal disease
* Heart disease
* Malignant neoplasm excluding non-melanoma skin cancer
* Pregnant women
* Human immunodeficiency virus infection
* Tuberculosis
* Hepatitis C
* Obesity
* Dementia
* Pre-existing condition of COVID risk factor
* Autoimmune condition
* Flu-like symptom episodes

### 7.6 Features of interest

These features span across the full set of target cohorts and research questions of interest in subgroups (incl. pediatrics, pregnant women, etc) so some features will only be relevant in some target cohorts or some subgroups, but the full list is given here.

Pre-index characteristics

These features will be described as assessed in two different time windows: the last 30 days (-1 to -30 days) and the year (-1 to -365 days) pre-index:

**Demographics**:

- Age: calculated as year of cohort start date – year of birth and with 5 year groupings

- Sex

* Race

**Concept-based:**

- Condition groups (SNOMED + descendants), >=1 occurrence during the interval

- Drug era groups (ATC/RxNorm + descendants), >=1 day during the interval which overlaps with at least 1 drug era

**Cohort-based: (to be added if relevant from lists)**

* Gestational diabetes
* Preeclampsia or eclampsia

Post-index characteristics

These features will be described in two different time windows: at index date (day 0) and in the 30 days from index date (0 to 30 days). As time elapses, additional windows of time will be investigated (eg. in the 60 days from index date [0 to 60 days]). The characteristics will include:

**Concept-based:**

* Condition groups (SNOMED + descendants), >=1 occurrence during the interval
* Drug era start groups (ATC/RxNorm + descendants), >=1 drug era start during the interval

**Cohort-based:**

* Abortion
* Acute kidney injury (AKI) diagnosis during hospitalization
* Acute kidney injury (AKI) using diagnosis codes and change in measurements during hospitalization
* Acute myocardial infarction events
* Acute pancreatitis events
* Acute Respiratory Distress syndrome (ARDS) during hospitalization
* Angina during hospitalization
* Anosmia OR Hyposmia OR Dysgeusia
* Asthma/COPD Step 1
* Asthma/COPD Step 2
* Asthma/COPD Step 3
* Bleeding during hospitalization
* Bradycardia or heart block during hospitalization
* Cardiac arrhythmia during hospitalization
* Cardiovascular-related mortality
* Cesarean section
* Cough
* Death
* Deep vein thrombosis events
* Dialysis during hospitalization
* Discharge from hospitalization
* Dyspnea
* Eclampsia and pre-eclampsia
* ECMO during hospitalization
* Fetal growth restriction
* Fever
* Gastrointestinal bleeding events
* Gestational diabetes
* Heart failure during hospitalization
* Hemorrhagic stroke (intracerebral bleeding) events
* Hospitalization episodes
* Hospitalization for Asthma
* Hospitalization for COPD
* Hospitalization for psychosis
* Incident depression with no prior treatment and no mania/psychoses
* Intensive services during hospitalization
* Ischemic stroke events
* Livebirth Delivery
* Livebirth excluding preterm and post term delivery
* Livebirth Post term Delivery
* Livebirth Preterm Delivery
* Malaise or fatigue
* Mechanical ventilation during hospitalization
* Multi-system inflammatory syndrome (Kawasaki disease or toxic shock syndrome)
* Myalgia
* Persons with additional testing for SARS-Cov-2 (prior test >=1d before test)
* Persons with additional testing for SARS-Cov-2 (prior test >=5d before test)
* Persons with chest pain or angina
* Persons with hepatic failure
* Pneumonia during hospitalization
* Pneumonia episodes
* Premature Rupture of Membranes
* Prevalent Asthma or Chronic obstructive pulmonary disease (COPD)
* Pulmonary Embolism events
* Sepsis during hospitalization
* Stillbirth
* Stroke (ischemic or hemorrhagic) events
* Suicide and suicidal ideation
* Supraventricular arrythymia during hospitalization
* Total cardiovascular disease events
* Tracheostomy during hospitalization
* Transient ischemic attack events
* Venous thromboembolic (pulmonary embolism and deep vein thrombosis) events
* Ventricular arrhythmia or cardiac arrest during hospitalization

In addition, DRUG\_ERA features will be generated using the same drug list for the Project Sc(y)lla: SARS-Cov-2 Large-scale Longitudinal Analyses on the comparative safety and effectiveness of treatments under evaluation for COVID-19 across an international observational data network. The full protocol for this investigation can be found at <https://www.ohdsi.org/wp-content/uploads/2020/05/OHDSI-SCYLLA-PLE-COVID-effectiveness-protocol-v1.0-final-with-appendix.pdf>.

### 7.7 Analysis: Characterizing cohorts

All analyses will be performed using code developed for the OHDSI Methods library. The code for this study can be found at <https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis>. A diagnostic package, built off the OHDSI Cohort Diagnostics ( <https://ohdsi.github.io/CohortDiagnostics/>) library, is included in the base package as a preliminary step to assess the fitness of use of phenotypes on your database. If a database passes cohort diagnostics, the full study package will be executed. Baseline covariates will be extracted using an optimized SQL extraction script based on principles of the FeatureExtraction package (<http://ohdsi.github.io/FeatureExtraction/>) to quantify Demographics (Gender, Prior Observation Time, Age Group), Condition Group Eras and Drug Group Eras (at and within 30 days after index date, at index date, within 30 days before index date, and within 365 days before index date). Additional cohort-specific covariates will be constructed using OMOP standard vocabulary concepts.

At the time of executing Feature Extraction, the package will create a data frame in which individuals’ age and sex will be extracted. Individuals’ medical conditions, procedures, measurements and medications will be summarized 1) over the year prior to their index date (-365d to -1d), 2) over the 30 days before index date (-30d to -1d), 3) at index date (0d), and 4) at and over the 30 days after index date (0d to 30d). Number and proportion of persons with feature variables during time-at-risk windows will be reported by target cohort and specific stratifications. Standardized mean differences (SMD) will be calculated when comparing characteristics of study cohorts, with plots comparing the mean values of characteristics for each of the characteristics (with the color indicating the absolute value of the standardized difference of the mean).

### 7.8 Logistics of Executing a Federated Analysis

Sites will run the study analysis package locally on their data coded according to OMOP CDM. Only aggregate results will be shared with the study coordinator. Result files will be automatically staged into a ZIP file that can be transmitted using the OhdsiSharing R Library (<http://ohdsi.github.io/OhdsiSharing/>) or through a site’s preferred SFTP client using a site-specific key provisioned by the OHDSI Study Coordinator. Local data stewards are encouraged to review study parameters to ensure minCellCount function follows local governance. At a minimum, it is encouraged to keep this value to >5 to avoid any potential issues with re-identification of patients. An example of tables and figures can be seen in Section 13. (Note: covariates are constructed using controlled ontologies from the OMOP standard vocabularies though some labels may be replaced with publication-friendly labels due to space restrictions of the submitting journal.)

# 8. Sample Size and Study Power

The study package is designed to suppress any analyses which have less than 140 unique persons. This cut point was informed by a power calculation performed by the OHDSI COVID Consortia to assess the computational cut point of when a cell count would be too small to merit additional subdivision within the target-stratum-feature combination. This means that each data owner will only generate results for target-stratum-feature pairs that meet this minimum threshold.

# 9. Strengths and Limitations

### 9.1 Strengths

To our knowledge, this is one of the world’s largest observational sets of analyses of secondary health data for SARS-Cov-2 tested population and COVID-19 cases. We are running a multi-country, multi-center characterization study to understand baseline covariates, treatments and outcomes observed in COVID-19 and influenza cases. The use of a common data model and standard vocabularies ensures interoperability and portability of phenotypes utilized in this analysis. This will be the first time any analysis compares patient samples with these infectious conditions within the same population. The use of a federated study model will ensure no movement of patient-level data from institutions participating in this analysis. This is critically important to ensure the protection of patient privacy in the secondary use of routinely collected patient data. Data custodians will remain in control of the analysis run on these data and will conduct their own site-based validation processes to evaluate case reports against public health reporting.

### 9.2 Limitations

The potential for case misclassification of Influenza and COVID-19 is a concern, especially for COVID-19 for which testing plays a large part. COVID-19 cases may be underreported due to shortages in testing resources and the fact that many infected patients may remain asymptomatic or do not require observed healthcare utilization. There is temporal variability in the extent to which underdiagnosis/underreporting may impact case rates as countries develop different national testing strategies, diagnosis guidance and the sensitivity of utilized testing instruments (e.g. nasopharyngeal versus saliva swabs, antibody testing). The case definition will inherently vary over calendar time and may require, as time continues, adjustment for known issues in case classification. The use of seasonal vaccinations for influenza strains may contribute to disruption in the natural history of influenza in susceptible subgroups. In the absence of vaccination, the adoption and implementation of social distancing measures to isolate and contain SARS-COV-2 spread may also disrupt the natural history of COVID-19. In both cases the success of preventative measures may lead to lower case rates than unmitigated populations though the impact is largely negligible as large populations still remain at-risk for potential infection.

Furthermore, classification of the cases could also vary with respect to calendar time, since underdiagnosis could become more frequent over the course of the pandemic, with positive cases tending to be identified mostly in severe cases. To address this inherent limitation, the study also includes a hospitalization-based COVID-19 case definition which will likely be more consistent across space and time and better classified for the more serious cases of disease in these data, to provide additional context.

Medical conditions may be underestimated as they will be based on the presence of condition codes, with the absence of such a record taken to indicate the absence of a disease. Meanwhile, medication records indicate that an individual was prescribed or dispensed a particular drug, but this does not necessarily mean that an individual took the drug as originally prescribed or dispensed. Our study could be subject to exposure misclassification with false positives if a patient had a dispensing but did not ingest the drug, but may also be subject to false negatives for non-adherent patients who continued their medication beyond the days supply due to stockpiling. Medication use estimates on the date of hospitalization is particularly sensitive to misclassification, and may conflate baseline concomitant drug history with immediate treatment upon admission. Comparisons of individuals with COVID-19 with individuals previously with influenza has limitations. In particular, observed differences may be explained by changes in clinical practice or data capture procedures over time, rather than by differences in the individuals themselves. This is likely a particular relevant drawback for any comparison of medication use. There may be less time-to-event data in the more emergent population at-risks this will inevitably become a non-issue as the pandemic continues on and more follow-up time is captured. In the immediate analysis, we will investigate the potential variation in complete versus incomplete follow-up time.

# 10. Protection of Human Subjects

The study uses only de-identified data. Confidentiality of patient records will be maintained at all times. Data custodians will remain in full control of executing the analysis and packaging results. There will be no transmission of patient-level data at any time during these analyses. Only aggregate statistics will be captured. Study packages will contain minimum cell count parameters to obscure any cells which fall below allowable reportable limits. All study reports will contain aggregate data only and will not identify individual patients or physicians.

# 11. Management and Reporting of Adverse Events and Adverse Reactions

This study will provide a descriptive summary of individuals at time of diagnosis/test for COVID-19. Adverse events after the first 30 days of a COVID-19 record are outside the scope of the study.

# 12. Plans for Disseminating and Communicating Study Results

The results will be used across multiple papers by the target cohorts, by stratification features, and/or by baseline characteristics, treatments, or outcomes. All results will be posted on the OHDSI website (evidence.ohdsi.org) after completion of the study. At least one paper per topic presented in Section 2.1 describing the study and its results will be written and submitted for publication to a peer-reviewed scientific journal. The results will also be presented at the OHDSI in-person events.

# 

# References

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4. Jon Cohen, Kai Kupferschmidt, Countries test tactics in ‘war’ against COVID-19. Science. 2020. Vol. 367, Issue 6484, pp. 1287-1288. 10.1126/science.367.6484.1287
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6. Voss EA, Makadia R, Matcho A, et al. Feasibility and utility of applications of the common data model to multiple, disparate observational health databases. J Am Med Inf Assoc 2015;22:553–64. doi:10.1093/jamia/ocu023

# ***Appendix 1: Cohort Definitions***

COVID Target Definitions –

|  |  |  |
| --- | --- | --- |
| Name | Atlas Link | JSON Link |
| Persons tested for SARS-CoV-2 with at least 365d prior observation | https://atlas.ohdsi.org/#/cohortdefinition/206 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/126.json |
| Persons tested positive for SARS-CoV-2 with at least 365d prior observation | https://atlas.ohdsi.org/#/cohortdefinition/204 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/128.json |
| Persons tested with a COVID-19 diagnosis record or a SARS-CoV-2 positive test with at least 365d prior observation | https://atlas.ohdsi.org/#/cohortdefinition/202 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/130.json |
| Persons with a COVID-19 diagnosis or a SARS-CoV-2 positive test with at least 365d prior observation | https://atlas.ohdsi.org/#/cohortdefinition/200 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/132.json |
| Persons hospitalized with a COVID-19 diagnosis record or a SARS-CoV-2 positive test with at least 365d prior observation | https://atlas.ohdsi.org/#/cohortdefinition/197 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/134.json |
| Persons hospitalized and requiring intensive services with a COVID-19 diagnosis record or a SARS-CoV-2 positive test with at least 365d prior observation | https://atlas.ohdsi.org/#/cohortdefinition/305 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/136.json |
| Persons hospitalized with a SARS-CoV-2 positive test with at least 365d prior observation | https://atlas.ohdsi.org/#/cohortdefinition/191 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/138.json |
| Persons hospitalized and requiring intensive services with a SARS-CoV-2 positive test with at least 365d prior observation | https://atlas.ohdsi.org/#/cohortdefinition/306 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/140.json |
| Persons tested for SARS-CoV-2 with no required prior observation | https://atlas.ohdsi.org/#/cohortdefinition/205 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/127.json |
| Persons tested positive for SARS-CoV-2 with no required prior observation | https://atlas.ohdsi.org/#/cohortdefinition/203 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/129.json |
| Persons tested with a COVID-19 diagnosis record or a SARS-CoV-2 positive test with no required prior observation | https://atlas.ohdsi.org/#/cohortdefinition/201 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/131.json |
| Persons with a COVID-19 diagnosis or a SARS-CoV-2 positive test with no required prior observation | https://atlas.ohdsi.org/#/cohortdefinition/199 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/133.json |
| Persons hospitalized with a COVID-19 diagnosis record or a SARS-CoV-2 positive test with no required prior observation | https://atlas.ohdsi.org/#/cohortdefinition/198 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/135.json |
| Persons hospitalized and requiring intensive services with a COVID-19 diagnosis record or a SARS-CoV-2 positive test with no required prior observation | https://atlas.ohdsi.org/#/cohortdefinition/308 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/137.json |
| Persons hospitalized with a SARS-CoV-2 positive test with no required prior observation | https://atlas.ohdsi.org/#/cohortdefinition/194 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/139.json |
| Persons hospitalized and requiring intensive services with a SARS-CoV-2 positive test with no required prior observation | https://atlas.ohdsi.org/#/cohortdefinition/309 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/141.json |

Influenza Target Definitions –

|  |  |  |
| --- | --- | --- |
| Name | Atlas Link | JSON Link |
| Persons with Influenza diagnosis or positive test 2017-2018 with at least 365d prior observation | https://atlas.ohdsi.org/#/cohortdefinition/211 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/111.json |
| Persons hospitalized with influenza diagnosis or positive test 2017-2018 with at least 365d prior observation | https://atlas.ohdsi.org/#/cohortdefinition/212 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/113.json |
| Persons hospitalized with influenza diagnosis or positive test and requiring intensive services 2017-2018 with at least 365d prior observation | https://atlas.ohdsi.org/#/cohortdefinition/307 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/115.json |
| Persons with Influenza diagnosis or positive test 2017-2018 with no required prior observation | https://atlas.ohdsi.org/#/cohortdefinition/210 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/112.json |
| Persons hospitalized with influenza diagnosis or positive test 2017-2018 with no required prior observation | https://atlas.ohdsi.org/#/cohortdefinition/209 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/114.json |
| Persons hospitalized with influenza diagnosis or positive test and requiring intensive services 2017-2018 with no required prior observation | https://atlas.ohdsi.org/#/cohortdefinition/310 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/116.json |

Stratum Definitions –

|  |  |  |
| --- | --- | --- |
| Name | Atlas Link | JSON Link |
| Prevalent Type 2 Diabetes Mellitus | https://atlas.ohdsi.org/#/cohortdefinition/311 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/100.json |
| Prevalent hypertension | https://atlas.ohdsi.org/#/cohortdefinition/227 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/101.json |
| Prevalent chronic obstructive pulmonary disease (COPD) without asthma | https://atlas.ohdsi.org/#/cohortdefinition/219 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/119.json |
| Prevalent Asthma without COPD | https://atlas.ohdsi.org/#/cohortdefinition/218 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/120.json |
| Prevalent chronic kidney disease | https://atlas.ohdsi.org/#/cohortdefinition/228 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/102.json |
| Prevalent end stage renal disease | https://atlas.ohdsi.org/#/cohortdefinition/232 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/103.json |
| Prevalent heart disease | https://atlas.ohdsi.org/#/cohortdefinition/231 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/104.json |
| Prevalent malignant neoplasm excluding non-melanoma skin cancer | https://atlas.ohdsi.org/#/cohortdefinition/222 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/105.json |
| Pregnant women | https://atlas.ohdsi.org/#/cohortdefinition/233 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/199.json |
| Prevalent Human immunodeficiency virus infection | https://atlas.ohdsi.org/#/cohortdefinition/229 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/106.json |
| Prevalent tuberculosis | https://atlas.ohdsi.org/#/cohortdefinition/315 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/117.json |
| Prevalent Hepatitis C | https://atlas.ohdsi.org/#/cohortdefinition/230 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/107.json |
| Prevalent obesity | https://atlas.ohdsi.org/#/cohortdefinition/224 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/108.json |
| Prevalent Dementia | https://atlas.ohdsi.org/#/cohortdefinition/226 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/109.json |
| Prevalent pre-existing condition of COVID risk factor | https://atlas.ohdsi.org/#/cohortdefinition/213 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/125.json |
| Prevalent Autoimmune condition | https://atlas.ohdsi.org/#/cohortdefinition/220 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/118.json |
| Flu-like symptom episodes | https://atlas.ohdsi.org/#/cohortdefinition/290 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/200.json |
| Prevalent chronic kidney disease broad | https://atlas.ohdsi.org/#/cohortdefinition/312 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/203.json |
| Prevalent end stage renal disease broad | https://atlas.ohdsi.org/#/cohortdefinition/313 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/204.json |
| Prevalent Human immunodeficiency virus infection broad | https://atlas.ohdsi.org/#/cohortdefinition/314 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/205.json |
| Prevalent tuberculosis broad | https://atlas.ohdsi.org/#/cohortdefinition/315 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/206.json |

Feature Definitions –

|  |  |  |
| --- | --- | --- |
| Name | Atlas Link | JSON Link |
| Prevalent Asthma or Chronic obstructive pulmonary disease (COPD) | https://atlas.ohdsi.org/#/cohortdefinition/217 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/121.json |
| Asthma/COPD Step 1 | https://atlas.ohdsi.org/#/cohortdefinition/216 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/122.json |
| Asthma/COPD Step 2 | https://atlas.ohdsi.org/#/cohortdefinition/215 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/123.json |
| Asthma/COPD Step 3 | https://atlas.ohdsi.org/#/cohortdefinition/214 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/124.json |
| Gestational diabetes | https://atlas.ohdsi.org/#/cohortdefinition/225 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/110.json |
| Eclampsia and pre-eclampsia | https://atlas.ohdsi.org/#/cohortdefinition/289 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/142.json |
| Fever | https://atlas.ohdsi.org/#/cohortdefinition/288 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/143.json |
| Cough | https://atlas.ohdsi.org/#/cohortdefinition/287 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/144.json |
| Myalgia | https://atlas.ohdsi.org/#/cohortdefinition/286 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/145.json |
| Malaise or fatigue | https://atlas.ohdsi.org/#/cohortdefinition/285 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/146.json |
| Dyspnea | https://atlas.ohdsi.org/#/cohortdefinition/284 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/147.json |
| Anosmia OR Hyposmia OR Dysgeusia | https://atlas.ohdsi.org/#/cohortdefinition/283 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/148.json |
| Persons with additional testing for SARS-Cov-2 (prior test >=1d before test) | https://atlas.ohdsi.org/#/cohortdefinition/282 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/149.json |
| Persons with additional testing for SARS-Cov-2 (prior test >=5d before test) | https://atlas.ohdsi.org/#/cohortdefinition/281 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/150.json |
| Hospitalization episodes | https://atlas.ohdsi.org/#/cohortdefinition/280 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/151.json |
| Pneumonia during hospitalization | https://atlas.ohdsi.org/#/cohortdefinition/279 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/152.json |
| Acute Respiratory Distress syndrome (ARDS) during hospitalization | https://atlas.ohdsi.org/#/cohortdefinition/278 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/153.json |
| Acute kidney injury (AKI) diagnosis during hospitalization | https://atlas.ohdsi.org/#/cohortdefinition/276 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/154.json |
| Sepsis during hospitalization | https://atlas.ohdsi.org/#/cohortdefinition/277 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/155.json |
| Venous thromboembolic (pulmonary embolism and deep vein thrombosis) events | https://atlas.ohdsi.org/#/cohortdefinition/275 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/156.json |
| Pulmonary Embolism events | https://atlas.ohdsi.org/#/cohortdefinition/274 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/157.json |
| Deep vein thrombosis events | https://atlas.ohdsi.org/#/cohortdefinition/273 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/158.json |
| Heart failure during hospitalization | https://atlas.ohdsi.org/#/cohortdefinition/272 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/159.json |
| Cardiac arrhythmia during hospitalization | https://atlas.ohdsi.org/#/cohortdefinition/248 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/160.json |
| Bradycardia or heart block during hospitalization | https://atlas.ohdsi.org/#/cohortdefinition/271 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/161.json |
| Supraventricular arrythymia during hospitalization | https://atlas.ohdsi.org/#/cohortdefinition/270 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/162.json |
| ventricular arrhythmia or cardiac arrest during hospitalization | https://atlas.ohdsi.org/#/cohortdefinition/269 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/163.json |
| death | Not in Atlas | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/164.json |
| Stillbirth | https://atlas.ohdsi.org/#/cohortdefinition/268 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/165.json |
| Livebirth Delivery | https://atlas.ohdsi.org/#/cohortdefinition/267 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/166.json |
| Livebirth Preterm Delivery | https://atlas.ohdsi.org/#/cohortdefinition/264 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/167.json |
| Livebirth Post term Delivery | https://atlas.ohdsi.org/#/cohortdefinition/265 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/168.json |
| Livebirth excluding preterm and post term delivery | https://atlas.ohdsi.org/#/cohortdefinition/266 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/169.json |
| Abortion | https://atlas.ohdsi.org/#/cohortdefinition/263 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/170.json |
| Premature Rupture of Membranes | https://atlas.ohdsi.org/#/cohortdefinition/262 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/171.json |
| Fetal growth restriction | https://atlas.ohdsi.org/#/cohortdefinition/261 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/172.json |
| Cesarean section | https://atlas.ohdsi.org/#/cohortdefinition/260 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/173.json |
| Hospitalization for Asthma | https://atlas.ohdsi.org/#/cohortdefinition/259 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/174.json |
| Hospitalization for COPD | https://atlas.ohdsi.org/#/cohortdefinition/258 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/175.json |
| Pneumonia episodes | https://atlas.ohdsi.org/#/cohortdefinition/257 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/176.json |
| intensive services during hospitalization | https://atlas.ohdsi.org/#/cohortdefinition/316 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/177.json |
| mechanical ventilation during hospitalization | https://atlas.ohdsi.org/#/cohortdefinition/318 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/178.json |
| tracheostomy during hospitalization | https://atlas.ohdsi.org/#/cohortdefinition/317 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/179.json |
| ECMO during hospitalization | https://atlas.ohdsi.org/#/cohortdefinition/253 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/180.json |
| dialysis during hospitalization | https://atlas.ohdsi.org/#/cohortdefinition/252 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/181.json |
| Persons with chest pain or angina | https://atlas.ohdsi.org/#/cohortdefinition/251 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/182.json |
| Angina during hospitalization | https://atlas.ohdsi.org/#/cohortdefinition/250 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/183.json |
| Persons with hepatic failure | https://atlas.ohdsi.org/#/cohortdefinition/249 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/184.json |
| Acute pancreatitis events | https://atlas.ohdsi.org/#/cohortdefinition/247 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/185.json |
| Total cardiovascular disease events | https://atlas.ohdsi.org/#/cohortdefinition/246 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/186.json |
| Gastrointestinal bleeding events | https://atlas.ohdsi.org/#/cohortdefinition/245 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/187.json |
| Cardiovascular-related mortality | https://atlas.ohdsi.org/#/cohortdefinition/244 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/188.json |
| Transient ischemic attack events | https://atlas.ohdsi.org/#/cohortdefinition/243 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/189.json |
| Stroke (ischemic or hemorrhagic) events | https://atlas.ohdsi.org/#/cohortdefinition/242 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/190.json |
| Ischemic stroke events | https://atlas.ohdsi.org/#/cohortdefinition/241 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/191.json |
| Hemorrhagic stroke (intracerebral bleeding) events | https://atlas.ohdsi.org/#/cohortdefinition/240 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/192.json |
| Acute myocardial infarction events | https://atlas.ohdsi.org/#/cohortdefinition/239 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/193.json |
| Bleeding during hospitalization | https://atlas.ohdsi.org/#/cohortdefinition/238 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/194.json |
| Incident depression with no prior treatment and no mania/psychoses | https://atlas.ohdsi.org/#/cohortdefinition/237 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/195.json |
| Hospitalization for psychosis | https://atlas.ohdsi.org/#/cohortdefinition/236 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/196.json |
| Suicide and suicidal ideation | https://atlas.ohdsi.org/#/cohortdefinition/235 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/197.json |
| Multi-system inflammatory syndrome (Kawasaki disease or toxic shock syndrome) | https://atlas.ohdsi.org/#/cohortdefinition/234 | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/198.json |
| Acute kidney injury (AKI) using diagnosis codes and change in measurements during hospitalization | Not in Atlas | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/201.json |
| Discharge from hospitalization | Not in Atlas | https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis/blob/master/inst/cohorts/202.json |