

Leveraging the OHDSI ecosystem

as a developer

OMOP Team, IQVIA March 2021



Training series plan

- + Session 1 : Course Introduction
 - OMOP CDM and vocabulary overview, OMOP conversion, data quality, examples of previous research and use cases, introducing ATLAS and OHDSI tools
- + Session 2: OMOP CDM/Vocabulary Tutorial
 - Concept, Concept mapping, Hierarchy, Ancestors, and OMOP CDM
- + Session 3: Cohort and Cohort Characterization
 - Concept sets, cohort definition, and cohort characterization
- + Session 4: Treatment Pathways and Incident Rates
 - Treatment pathways, Incidence rates, and Characterization using R





Table of contents

- + Exercise review from previous session
- + Introduction of influenza study in ATLAS

+ ATLAS concept sets and cohorts

- Data sources
- Concept sets
- Cohort building
- Build cohorts for example case study exercise

+ ATLAS characterizations

- Cohort characterization (default and customized features)
- Build characterization for example case study exercise

+ Homework





Ground Rules

- +This session will be recorded
- +Please make sure your microphones are muted
- +Type your questions in the chat or bring them to the Q& A session
- +Turn off your camera





Exercise Review from Session #1

Session #1 Homework – Proposed concepts

		CONCEPT ID	CONCEPT NAME	OHDSI SOURCE
Influenza		4266367		
Type 2 diabetes				https://atlas.ohdsi.org/#/conceptset/27/expression
Lung disease			chronic obstructive pulmonary disease (COPD)	https://atlas.ohdsi.org/#/conceptset/59/expression
Cancer		443392	Malignant neoplastic disease	
		4144289	H/O: malignant neoplasm	
Immunodeficiency		433740	Immunodeficiency disorder	
Heart Disease	Cardiomyopathy	321319	Cardiomyopathy	
	Myocardial infarction			https://atlas.ohdsi.org/#/conceptset/44/expression
	Heart Failure			https://atlas.ohdsi.org/#/conceptset/57/expression
Hypertension				https://atlas.ohdsi.org/#/conceptset/21/expression
Asthma				https://atlas.ohdsi.org/#/conceptset/60/expression
Renal Disease		45768812	Anemia in chronic kidney disease	https://atlas.ohdsi.org/#/cohortdefinition/228/conceptsets
		194385	Aneurysm of renal artery	
		46271022	Chronic kidney disease	
		192279	Disorder of kidney due to diabetes mellitus	
		4263367	Glomerulonephritis	
		261071	Glomerulosclerosis	
		201313	Hypertensive renal disease	
		193253	Nephritis	
		195314	Nephrotic syndrome	
		192359	Renal failure syndrome	





Exercise Review from Session #2

Session #2 Homework – Proposed solutions

https://drive.google.com/drive/folders/1-zqBUMOKFC2zk EvPsGPg-c36uWSjNTa





Introduction of Influenza Study in ATLAS

Influenza Study

Treatments and outcomes of influenza patients during hospital stay

Study Topic

Baseline demographics and clinical characteristics, treatment patterns and outcomes of patients diagnosed with influenza initiating treatment in the US hospital setting: a retrospective cohort study using administrative data.

Objectives

Primary Objectives

- Describe the treatment patterns of hospitalized influenza patients:
 - > Drugs (a) antivirals (b) antibiotics (c) corticosteroids
 - > Procedures (a) mechanical ventilation (b) tracheostomy (c) extracorporeal membrane oxygenation (d) oxygen therapy
- Describe the length of the hospital stay by conditions of interest:
 - > (a) diabetes (b) lung disease (c) cancer (d) immunodeficiency (e) heart disease (f) hypertension (g) asthma (h) kidney disease

Secondary Objectives

- Describe the baseline demographics and clinical characteristics of hospitalized influenza patients.



Influenza Study — Cohort Definition

Study Population

• Persons hospitalized during the **2008-2009 influenza season** with a diagnosis of influenza **21 days prior** or during the **hospital stay**, with **no prior continuous enrollment** required and with **no influenza hospitalization in the 6 months prior** to hospital admission.

Inclusion Criteria

- Patients with claims for a hospital stay between 1st September 2008 and 1st April 2009 (index date). All hospital stays during
 the study period are of interest.
- Patient is ≥ 18 years of age at index date.
- Patient has at least 1 diagnosis/measurement of influenza 21 days prior to index start date (hospital admission) or up to index end date (hospital discharge date).
- Patient has 0 months of prior continuous enrollment prior to hospital admission.
- EXCLUDE patients with evidence of hospitalization for influenza in the 6 months prior to index date.



ATLAS

- ★ Home
- Data Sources
- **Q** Search
- Concept Sets
- Cohort Definitions
- Characterizations
- A Cohort Pathways
- Incidence Rates
- Profiles
- △ Estimation
- Prediction
- **Jobs**
- **Configuration**
- Feedback

Apache 2.0 open source software

provided by



Analysis plan for the example study in ATLAS

- Part 1
 - Build concepts sets
 - Build cohort definitions

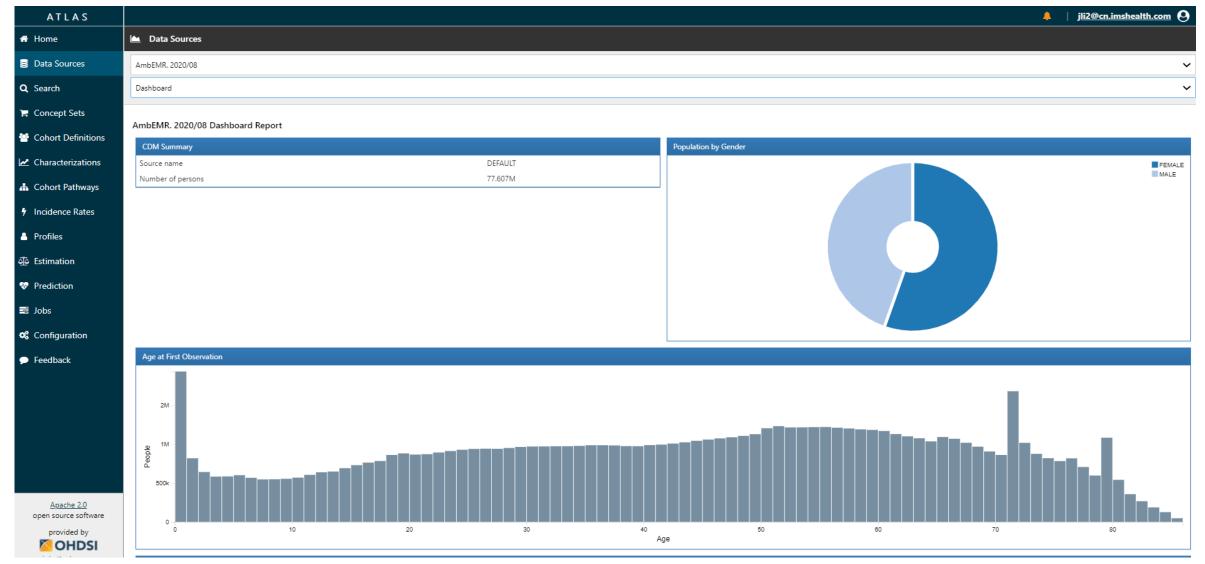
- Part 2
 - Characterization of the study cohort



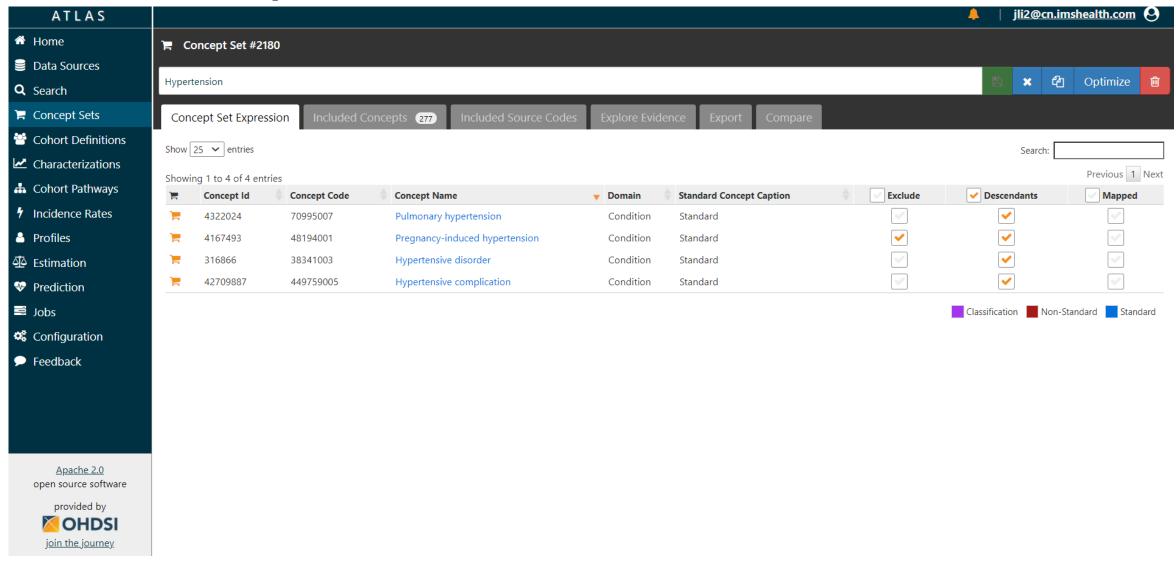


ATLAS Concept Sets Lecture, Demo & Exercise

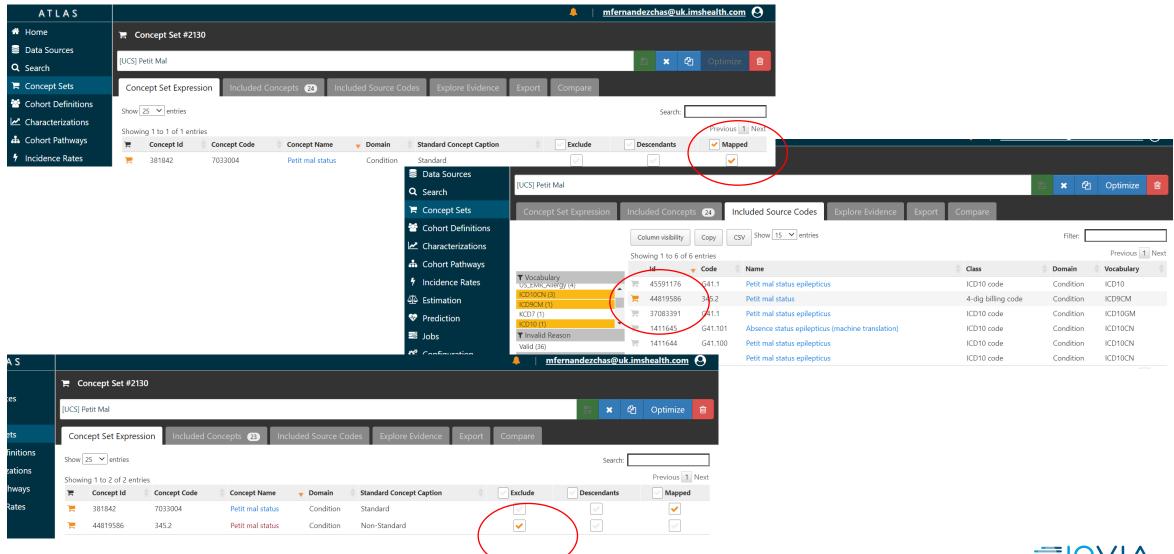
ATLAS – Data Sources



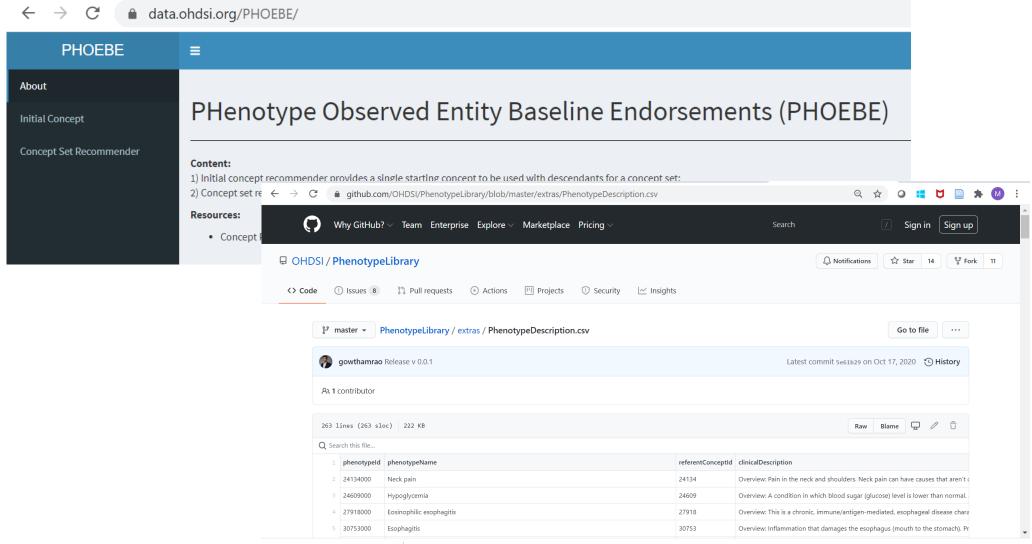
ATLAS – Concept Sets



ATLAS – Concept Sets



ATLAS – Concept Sets (Resources: phenotypes, Phoebe)



ATLAS – Concept Sets – Demo

- influenza
- type 2 diabetes
- lung disease
- cancer
- immunodeficiency
- heart disease
- hypertension
- asthma
- kidney disease

ATLAS – Concept Sets – Exercise

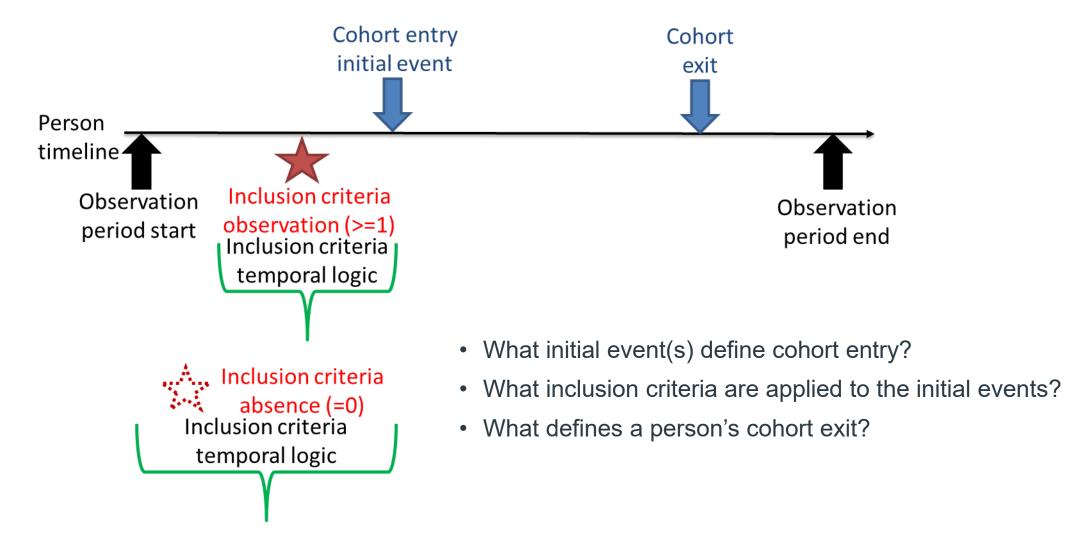
- influenza
- type 2 diabetes
- lung disease
- cancer
- immunodeficiency
- heart disease
- hypertension
- asthma
- kidney disease



ATLAS Cohort Definition Lecture, Demo & Exercise

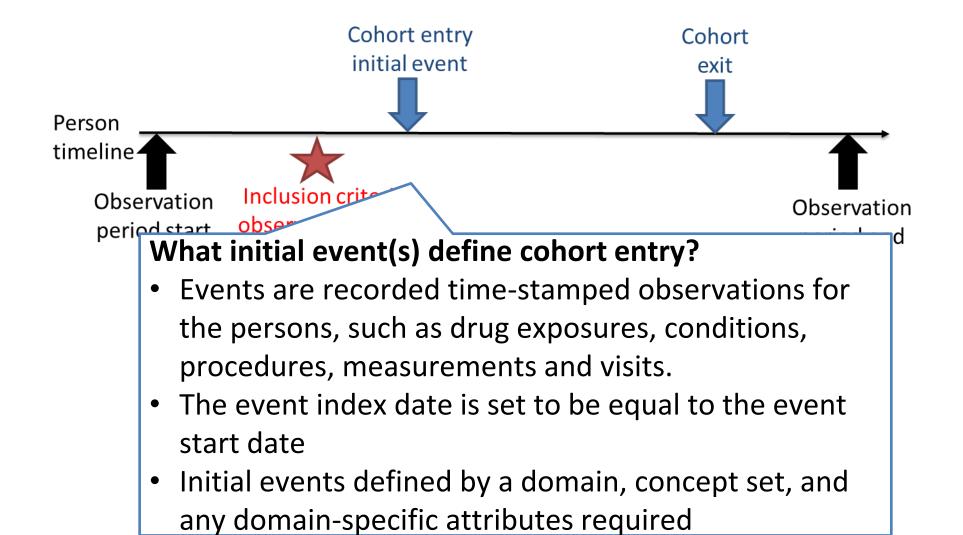
ATLAS – Cohort definitions

Dissecting the anatomy of a cohort definition



ATLAS – Cohort entry event

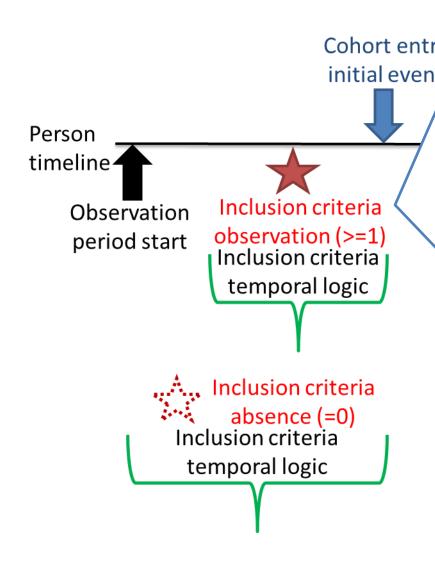
Dissecting the anatomy of a cohort definition





ATLAS – Cohort inclusion criteria

Dissecting the anatomy of a cohort definition



Cohort entr initial even initial events? What inclusion criteria are applied to the

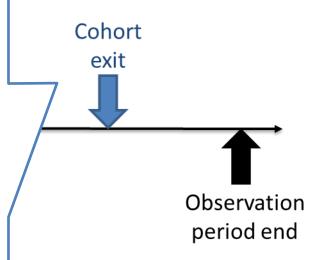
- The qualifying cohort will be defined as all persons who have an initial event and satisfy all qualifying inclusion criteria.
- Each inclusion criteria is defined by domain(s), concept set(s), domain-specific attributes, and the temporal logic relative to initial events
- Each qualifying inclusion criteria can be evaluated to determine the impact of the criteria on the attrition of persons from the initial cohort (example use case: clinical trial feasibility)

ATLAS – Cohort exit

Dissecting the anatomy of a cohort definition

What defines a person's cohort exit?

- Cohort exit signifies when a person no longer qualifies for cohort membership
- Cohort exit can be defined in multiple ways:
 - End of observation period
 - Fixed time interval relative to initial event
 - Last event in a sequence of related observations (ex: persistent drug exposure)
 - Censoring observations
- Cohort exit strategy will impact whether a person can belong to the cohort multiple times during different time intervals





BREAK

10 mins



ATLAS – Cohort Definition – Demo

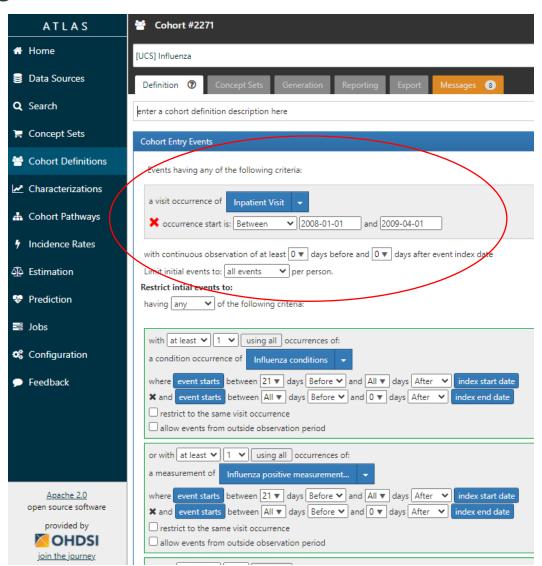
Persons hospitalized with influenza, no prior observation required, 2008-2009 season

https://atlas.ohdsi.org/#/cohortdefinition/117

Initial Event Cohort

People having any of the following:

- A visit occurrence of inpatient visit
 - Occurrence start is between 2008-09-01 and 2009-04-01 (inclusive)
- With continuous observation of at least 0 days prior and 0 days after event index date
- Limit initial events to: all events per person.



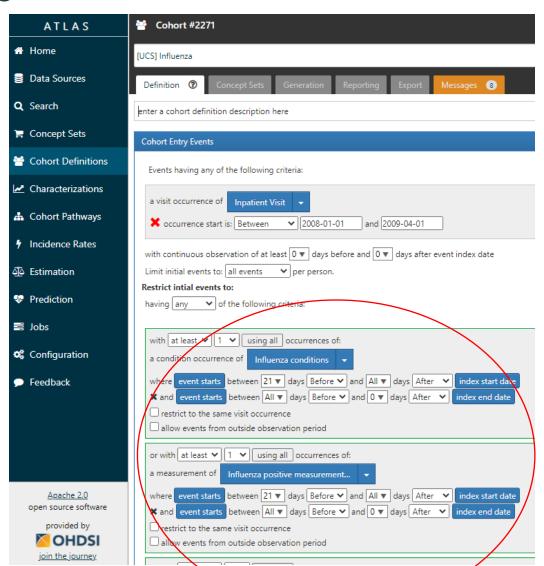


ATLAS – Cohort Definition – Demo

For people matching the **Primary Events**, include:

Having any of the following criteria:

- at least 1 occurrence of a condition occurrence of influenza
 where event
 starts between 21 days before and all days after index start
 date and event
 starts between all days before and 0 days after index end
 date
- or at least 1 occurrence of a measurement of influenza (precoordinated positive measurements) where event starts between 21 days before and event all days after index start date starts between all days before and 0 days after index end date





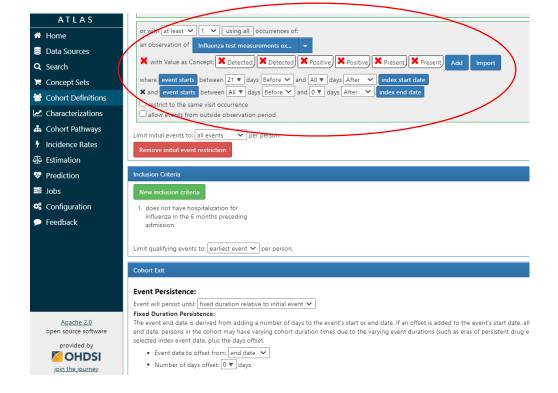
ATLAS – Cohort Definition – Demo

For people matching the **Primary Events**, include:

Having any of the following criteria:

- at least 1 occurrences of an observation of influenza testing (excluding Ab testing)
 - value as concept is any of: detected, positive, present
 where event
 starts between 21 days before and all days after index start
 date and event
 - starts between all days before and 0 days after index end date
- or at least 1 occurrences of a measurement of Influenza testing (excluding Ab testing)
 - value as concept is any of: detected, positive, present
 where event starts between 21 days before and all days after index start date and event
 - starts between all days before and 0 days after index end date*

Limit cohort of initial events to: all events per person.





^{*} Exercise

ATLAS – Cohort Definition – Exercise

1.- Download the concept sets from OHDSI Atlas (4 in total – 1xcondition diagnosis, 2xcondition measurements and 1xinpatient visit):

https://atlas.ohdsi.org/#/cohortdefinition/117/conceptsets

- 2.- Build the influenza cohort as shown in the demo and save it as '[UCS] Influenza cohort 2008 to 2009'.
- 3.- Now, add the following criteria:

For people matching the **Primary Events**, include:

Having any of the following criteria:

- or at least 1 occurrences of a measurement of influenza testing (excluding Ab testing)
 - value as concept is any of: detected, positive, present (SNOMED and LOINC)

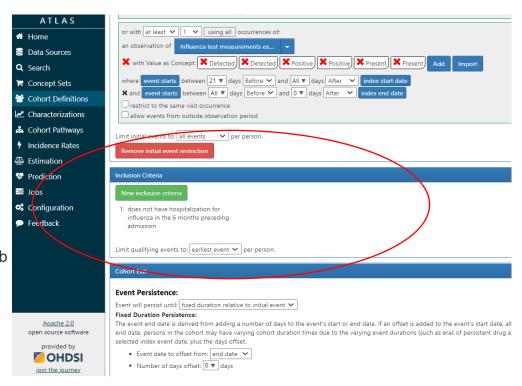
where event starts between 21 days Before and all days After index start date and event starts between all days Before and 0 days After index end date

4.- Add the following inclusion rules:

Inclusion Criteria #1: age >=18

<u>Inclusion Criteria #2</u>: does not have hospitalization for influenza in the 6 months preceding admission

Limit qualifying cohort to: earliest event per person.





BREAK

30 mins

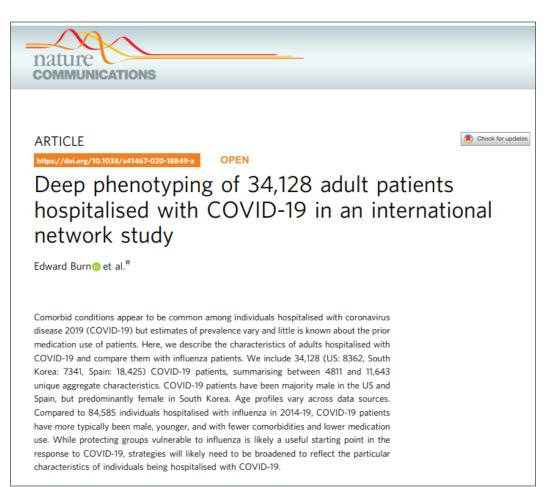




ATLAS Characterization Lecture, Demo & Exercise

Characterization Study Example

- Describe baseline characteristics for those hospitalized for COVID-19 as compared to those hospitalized for influenza
- Findings:
 - Patients hospitalized with COVID are systematically different from those hospitalized with flu
 - COVID hospitalized patients, when compared those hospitalized for influenza:
 - Greater proportion are male and slightly younger
 - Fewer comorbidities and lower medication use
- Utilized claims and electronic medical records from 10 databases across 3 different countries



https://www.nature.com/articles/s41467-020-18849-z



Characterization Results – Comparison of Sources/Locations

Fig. 1: Age of patients hospitalised with COVID-19 and of patients hospitalised with influenza.

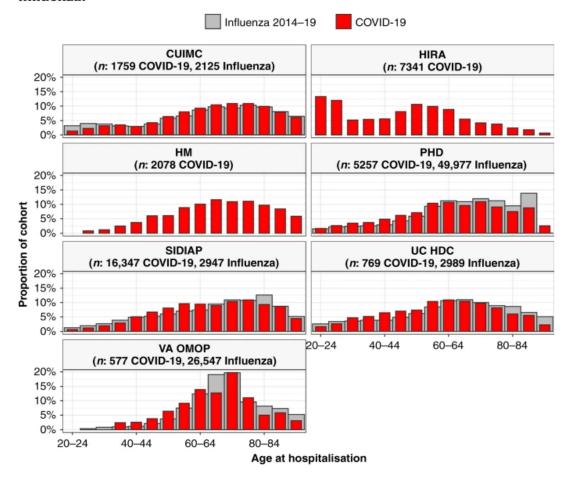
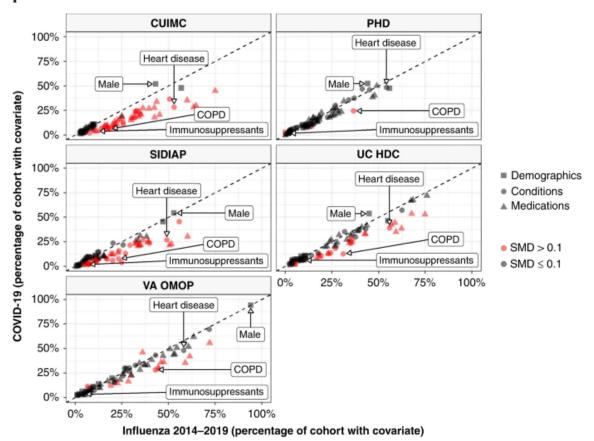
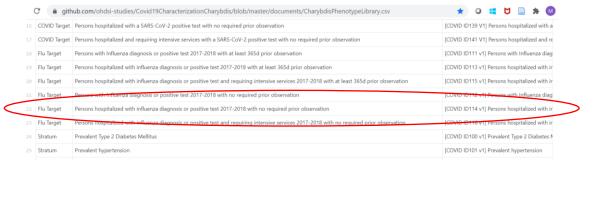


Fig. 4: Characteristics of COVID-19 patients compared to 2014–2019 Influenza patients.

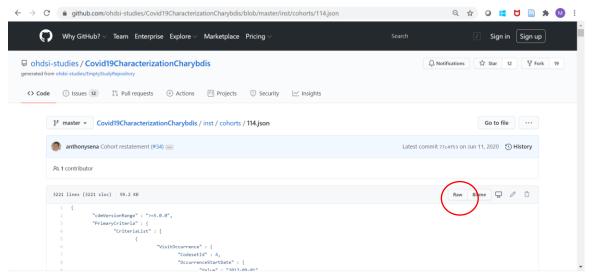


Characterizations from GitHub to ATLAS – part 1

https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis







To recreate one of the cohorts from the paper:

Locate phenotype library in the documents folder

https://github.com/ohdsistudies/Covid19CharacterizationCharybdis/bl ob/master/documents/CharybdisPhenotypeLi brary.csv

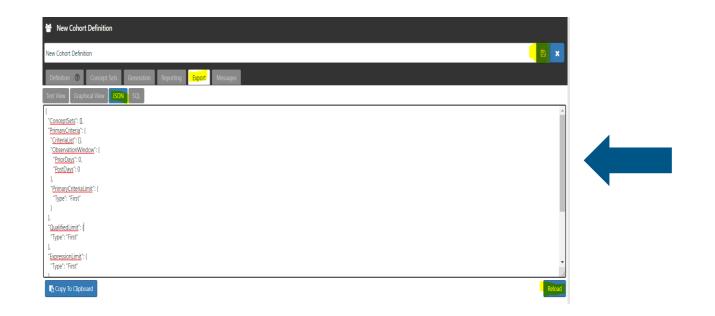
2. Choose the ones of interest from the list (114 and 135)

- 3. Go to the inst folder
- 4. Click on the JSON file you chose (114 and 135)
- 5. Click on the raw button
- 6. Copy all lines of code or 'Save as...' a JSON file (114.json and 135.json)



Characterizations from GitHub to ATLAS – part 1

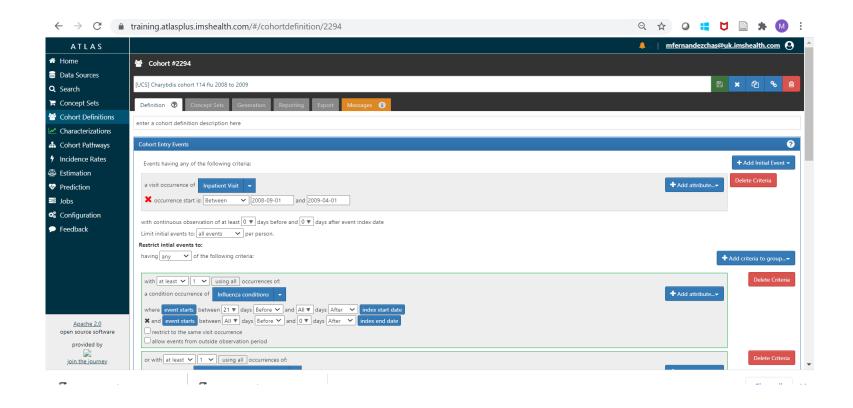
https://github.com/ohdsi-studies/Covid19CharacterizationCharybdis



In ATLAS:

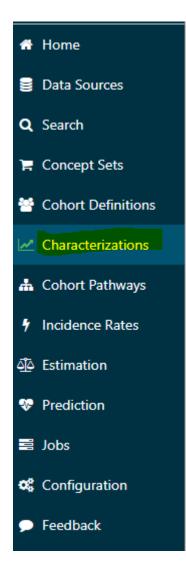
- 1. Create a brand new Cohort Definition
- 2. Under the Export JSON tabcpaste your copied code into this section
- 3. Press the **Reload** green button
- 4. Press the **SAVE** green button

ATLAS characterization stage 1 - definition



On the 'Definition' tab, the contents of the JSON file are automatically converted into a cohort definition

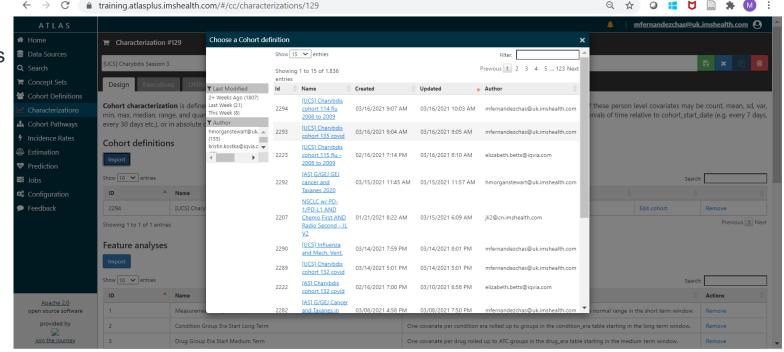
You can now update the cohort with the dates of the influenza season under study (2008-2009 in our case)

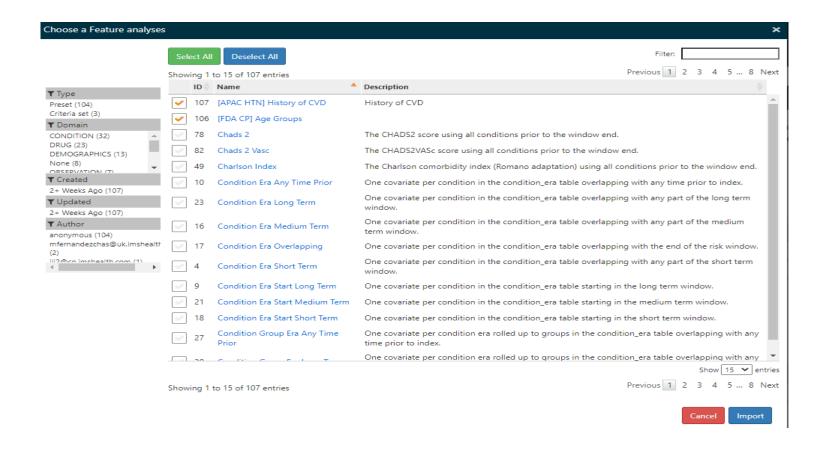


Go to Characterizations in ATLAS;

Create a New Characterization

Import your cohort(s)



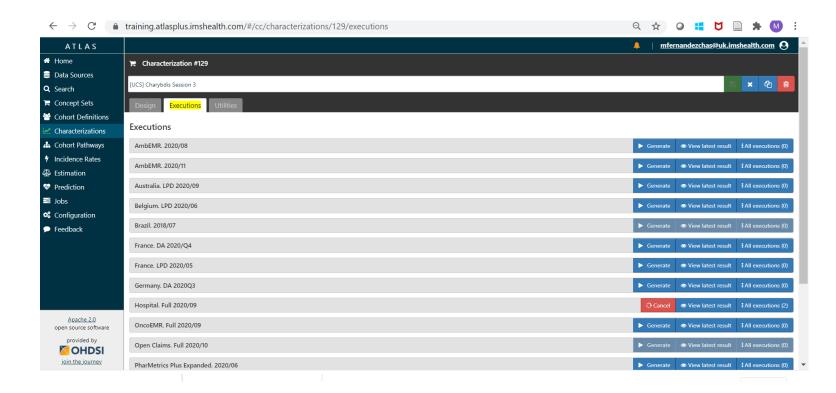


Select preset feature analysis, import

Save the characterization (green button by title)

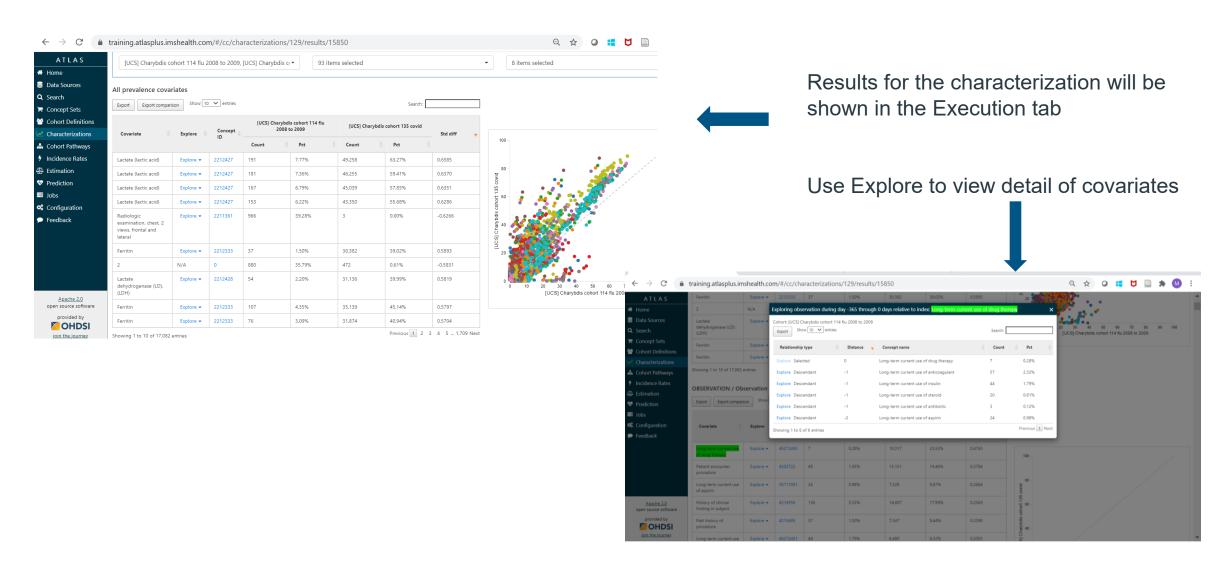
Go to the Executions tab, click Generate to create results from data sources

View latest results to see data for your chosen cohort(s) on that data source – next slide



Go to the Executions tab, click 'generate' for desired data source(s)

View latest results to see data for your chosen cohort(s) on that data source



ATLAS Characterization – Exercise 1

From the list of the RD2D consortium questions already answered :

• Among adults hospitalized with influenza, how many were put on a mechanical ventilator per subgroup (age, gender)?

Influenza Cohort

- Use your influenza cohort or download the one from the OHDSI ATLAS website:

https://atlas.ohdsi.org/#/cohortdefinition/117

And save it as '[UCS] Influenza cohort 2008 to 2009' or any other name of your preference

- Update the dates to reflect the 2008-2009 influenza season (1st September 2008 and 1st April 2009)

Mechanical Ventilation Inclusion

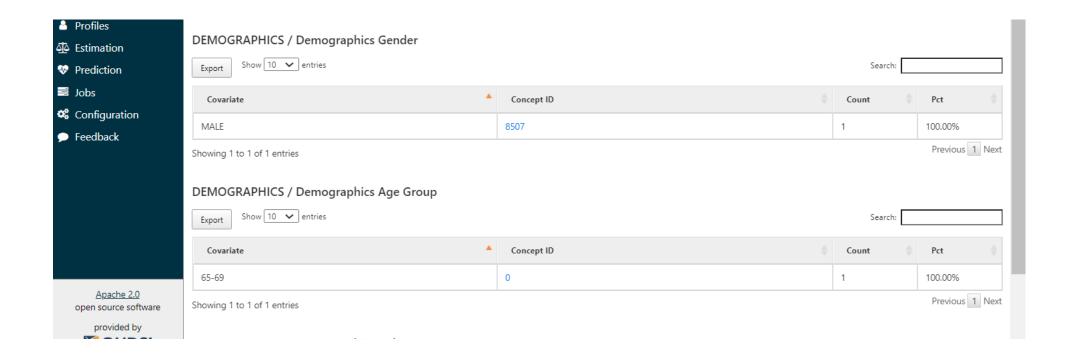
- Add mechanical ventilation to the inclusion criteria during the hospital stay (between index date and end date). Note that this is a multidomain concept (condition, observation and procedure). You will need to add one criteria per domain, and then select 'having ANY of the following criteria', otherwise, if the default 'ALL' is left, you will get 0 counts:
 - https://atlas.ohdsi.org/#/conceptset/49/expression
- Save the new cohort as '[UCS] Influenza cohort 2008 to 2009 mech. vent.' or any other name of your preference.

Characterization

- Create a new characterization and save it as '[UCS] Influenza cohort 2008 to 2009 mech. vent. analysis'
- Import your new influenza and ventilation cohort
- Include the age group and gender features
- Execute in the OHDSI CDM V5 database and check the results



ATLAS Characterization – Exercise 1 Answer



ATLAS Characterization – Exercise 2

Among adults hospitalized with influenza and pneumonia between 1st September 2008 and 1st April 2009, which group has the highest Charlson comorbidity index?

Influenza Cohort

- It has already been defined in the previous exercise

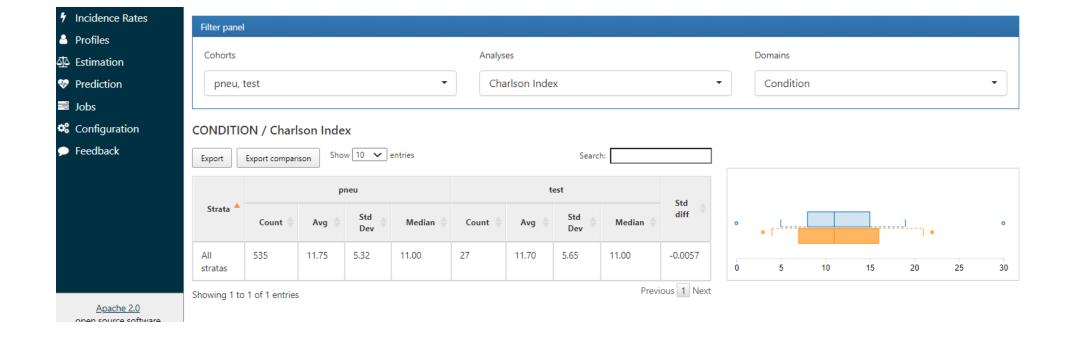
Pneumonia Cohort

- Create a copy of this influenza cohort and rename as '[UCS] Pneumonia 2008 to 2009'. Replace the condition 'influenza' by the condition 'pneumonia' and delete the measurement criteria. You will need a new concept for 'pneumonia' (concept id = 255848 and descendants).

Characterization

- Create a new characterization and save it as '[UCS] Influenza and Pneumonia cohort 2008 to 2009 Analysis'
- Import your new influenza and pneumonia cohorts
- Include the Charlson Index as the feature of the analysis (Preset feature ID = 49 calculates the Charlson comorbidity index using all conditions prior to the window end)
- Execute in the OHDSI CDM V5 database and check the results

ATLAS Characterization – Exercise 2 Answer



ATLAS Characterization – Homework

From the list of the RD2D consortium questions already answered :

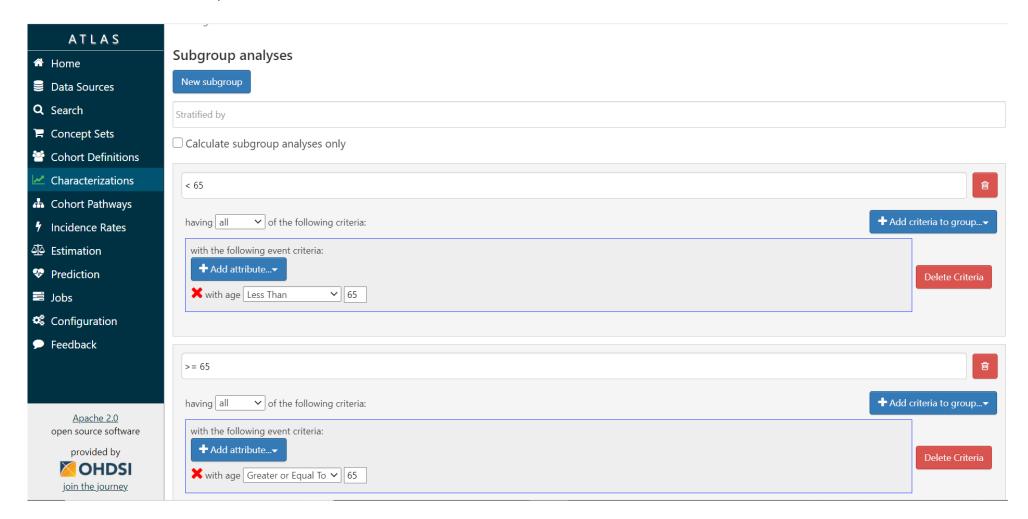
- Among adults hospitalized with COVID-19, how many were put on a mechanical ventilator per subgroup (age, gender)?
 - Download the COVID-19 cohort from the Charybdis website:
 - https://atlas.ohdsi.org/#/cohortdefinition/198
 - Add mechanical ventilation as described in Exercise 1
 - Save the new cohort as '[UCS] Covid-19 cohort mech. vent.' or any other name of your preference.
 - Create a new characterization and save it as '[UCS] Covid-19 cohort mech. vent. analysis'
 - Import your new COVID-19 and mechanical ventilation cohort
 - Include the age group and gender features
 - Execute in your own database and check the results
- Compare your results with the RD2D consortium results for COVID-19:

https://covid19questions.org/component/content/article/32-q-a/70-among-adults-hospitalized-with-covid-19-how-many-were-put-on-a-mechanical-ventilator-per-subgroup-age-ethnicity-gender-and-race?ltemid=279



Subgroup Analyses

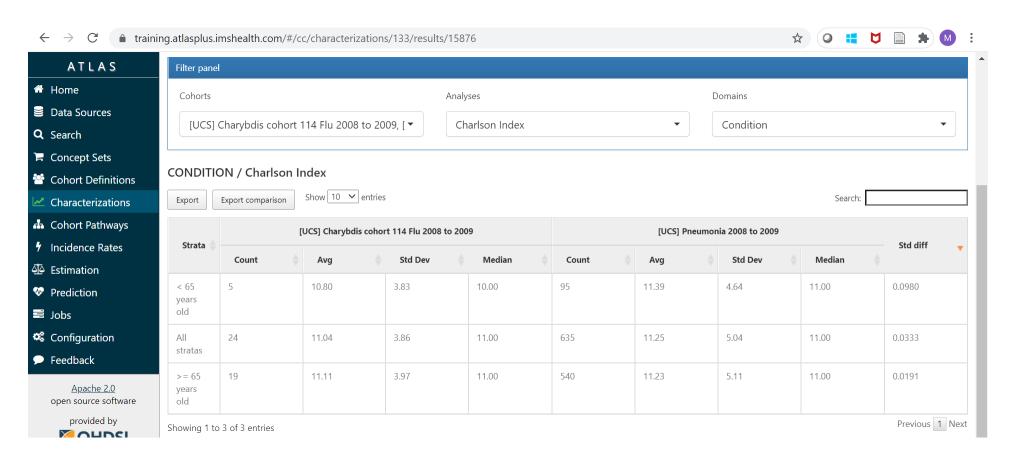
• Patients under 65, over 65





Subgroup Analyses

• Patients under 65, over 65

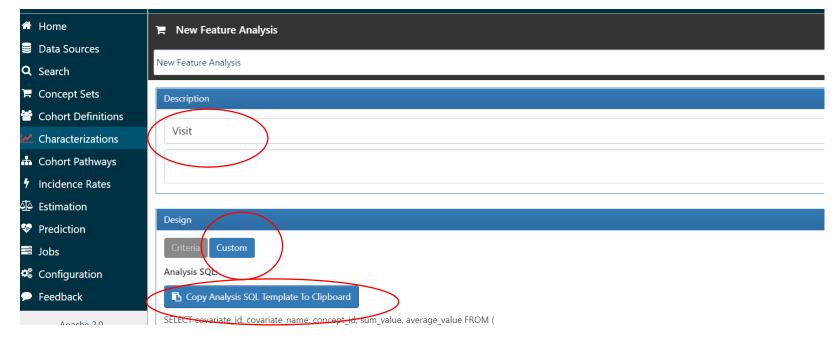


Customized features using SQL



- 1.- Select Feature Analysis on the left
- 2.- Select New Feature Analysis on the right

- 3.- Choose the Visit Domain
- 4.- Click on the Custom button
- 5.- Click on 'CopyAnalysis SQLTemplate to Clipboard'



Customized features using SQL

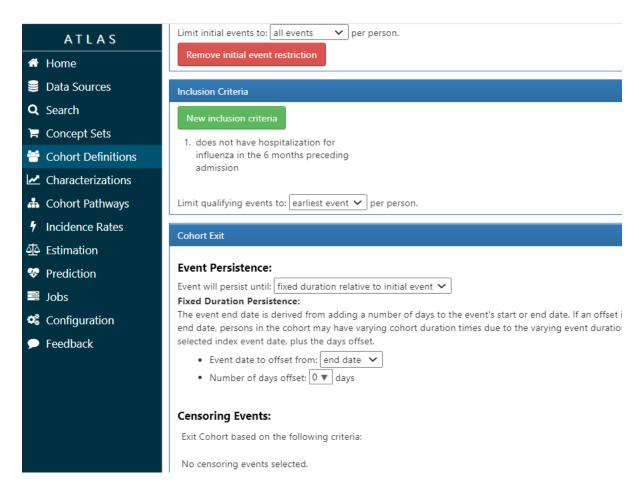
SQL Template

```
Feature SQL Template sql
     SELECT covariate id, covariate name, concept id, sum value, average value FROM (
 2
              -- Custom analysis producing same results as Feature Extraction's "One covariate per drug in
               -- the drug era table overlapping with any time prior to index."
               -- Available variables:
  6
                   -- @cdm database schema
                   -- @cohort table
 8
                   -- @cohort id
 9
                   -- @analysis id
 10
                   -- all variables specified in Cohort Characterization parameters
 11
               SELECT
                                                                                                AS covariate id,
 12
                CAST (drug concept id AS BIGINT) * 1000 + @analysis id
 13
                                                                                                AS covariate name,
                c.concept name
                drug concept id
 14
                                                                                                AS concept id,
                                                                                                AS sum value,
 15
                COUNT (*)
 16
                COUNT(*) * 1.0 / stat.total cnt * 1.0
                                                                                                AS average value
 17
               FROM (
 18
                      SELECT DISTINCT
 19
                        drug concept id,
 20
                        cohort.subject id,
                       cohort.cohort start date
 21
 22
                      FROM @cohort table cohort
                        INNER JOIN @cdm database schema.drug era ON cohort.subject id = drug era.person id
 23
 24
                      WHERE drug era start date <= cohort.cohort start date</pre>
 25
                            AND drug concept id !=
 26
                            AND cohort.cohort definition id = @cohort id
 27
                JOIN @cdm database schema.concept c ON drug entries.drug concept id = c.concept id
 28
 29
                CROSS JOIN (SELECT COUNT(*) total cnt
                             FROM @cohort table
                            WHERE cohort definition id = @cohort id) stat
 31
 32
               GROUP BY drug concept id, c.concept name, stat.total cnt
```

Paste the SQL code in your editor of preference and update the code highlighted in gray accordingly.

Customized features using SQL

Length of Visit



- Make sure the index event end date is set to 0 days from end date.
- This way the patients will exit the cohort at the end of the hospital visit.

Customized features using SQL

SELECT covariate_id, covariate_name, concept_id, sum_value, average_value FROM

avg(datediff(day, cohort.cohort start date, cohort.cohort end date))

1000 + @analysis id

0 AS average value

FROM @cohort table cohort

'Visit Length'

Length of Visit

Feedback

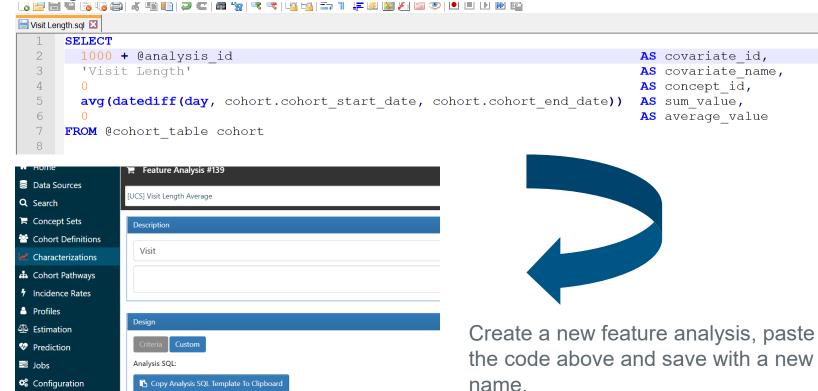
Apache 2.0

open source software

provided by

OHDSI

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AS covariate id,

AS covariate name

- Simple code to extract the length of the visit from our influenza cohort
- Since our index event is the hospital stay, the start and end date of the cohort will match the start and end date of the hospital stay.
- Hence the length of the stay is just the difference between the two dates.



BREAK

10 mins



Customized features using SQL

Length of Visit

From the list of the RD2D consortium pending questions :

• Among adults hospitalized with influenza what is the length of stay in hospital for patients with (a) Diabetes, (b) Lung Disease, (c) Cancer, (d) Immunodeficiency, (e) Heart Disease, (f) Hypertension, (g) Asthma, (h) Kidney Disease (1) Overall and (2) per subgroup >= 65 and < 65?

Influenza Cohort

- It has already been defined in the previous exercises

Inclusion Criteria

- Create a copy of this influenza cohort and rename as '[UCS] Covid-19 cohort and diabetes'. Add the 'diabetes' condition as a new inclusion criteria any time prior to the index date. Use the answers from the Session #1 Exercise to build the concept set for 'diabetes'
- Repeat this process for all conditions in the list above (8 in total)

Characterization 1

- Create a new characterization
- Import all your new influenza cohorts (8)
- Create a customised new feature analysis for the length of the visit and import this new feature into the characterization
- Execute and check the results

Characterization 2

- Make a copy of Characterization 1 and rename it
- Create the subgroups for >= 65 and < 65 years old
- Execute and check the results



Customized features using SQL

Top Drugs

 Among adults hospitalized with influenza what are the top drugs taken during the hospital stay?

Influenza Cohort

- As already defined during this session

Characterization

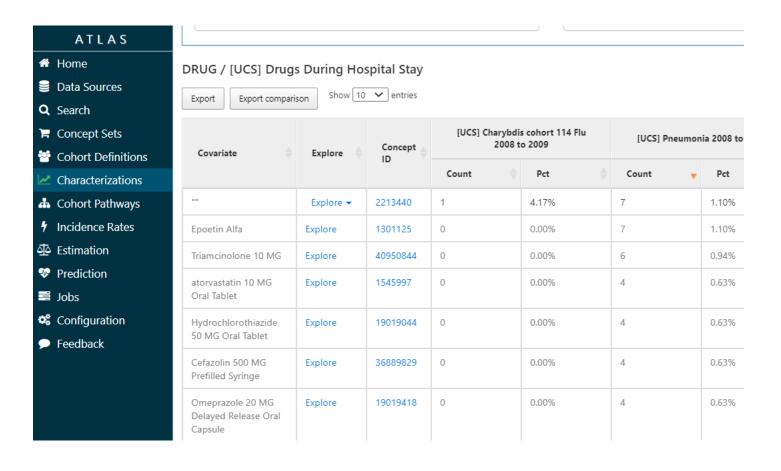
- Create a new characterization
- Import your influenza cohort
- Create a customised new feature analysis for the drugs below taken during the hospital stay as shown in the slide and import this new feature into the characterization
- Execute and check the results

```
🔚 [UCS] Drugs During Hospital Stay.sql 🗵
       SELECT
         CAST (drug concept id AS BIGINT) * 1000 + @analysis id
                                                                                             AS covariate id,
                                                                                             AS covariate name,
         c.concept name
         drug concept id
                                                                                             AS concept id,
         COUNT (*)
                                                                                             AS sum value,
         COUNT(*) * 1.0 / stat.total cnt * 1.0
                                                                                             AS average value
     FROM (
              SELECT DISTINCT
  9
                drug concept id,
 10
                cohort.subject id,
 11
                cohort.cohort start date
 12
              FROM @cohort table cohort
 13
                INNER JOIN @cdm database schema.drug exposure ON cohort.subject id = drug exposure.person id
              WHERE drug exposure start date >= cohort.cohort start date
 14
                    AND drug exposure start date <= cohort.cohort end date
 15
                    AND drug concept id != 0
 16
 17
                    AND cohort.cohort definition id = @cohort id
 18
            ) drug entries
 19
         JOIN @cdm database schema.concept c ON drug entries.drug concept id = c.concept id
 20
         CROSS JOIN (SELECT COUNT (*) total cnt
 21
                     FROM @cohort table
 22
                     WHERE cohort definition id = @cohort id) stat
      GROUP BY drug concept id, c.concept name, stat.total cnt
```

ATLAS Characterization – Customized Features – Homework Answer

Customized features using SQL

Top Drugs





Discussion & Homework

Homework





Training series plan

- + Session 1 : Course Introduction
 - OMOP CDM and vocabulary overview, examples of previous research and use cases, introducing ATLAS and OHDSI tools
- + Session 2: Common Data Model
 - Data source profiling, building concept sets, building cohorts, exercises and discussion
- + Session 3: Leveraging the OHDSI ecosystem as a developer
 - Concept sets, cohort definitions and their application to analytical use cases
 - Study design and development in ATLAS
- + Session 4: Leveraging the OHDSI ecosystem to run Network Studies
 - Characterization and treatment pathways, in ATLAS, SQL and R





Thank you