



Phenotyping for Cardiovascular Disease

Seng Chan You

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Standard phenotype library

- Sentinel's efforts for this

Diagnostic Accuracy Study

Medicine[®]

OPEN

Chart validation of inpatient ICD-9-CM administrative diagnosis codes for ischemic stroke among IGLV users in the Sentinel Distributed Database

Eric M. Ammann, PhD^{a,*}, Enrique C. Leira, MD MS^{a,b}, Scott K. Winiecki, MD^c, Nandakumar Nagaraja, MD^b, Sudeepta Dandapat, MD^b, Ryan M. Carnahan, PharmD, MS^a, Marin L. Schweizer, PhD^{b,d}, James C. Torner, PhD^a, Candace C. Fuller, PhD^e, Charles E. Leonard, PharmD, MSCE^f, Crystal Garcia, MPH^e, Madelyn Pimentel, MSN^e, Elizabeth A. Chrischilles, PhD^a



Standard phenotype library

- Sentinel's efforts for this

ORIGINAL REPORT

WILEY

Chart validation of inpatient ICD-9-CM administrative diagnosis codes for acute myocardial infarction (AMI) among intravenous immune globulin (IGIV) users in the Sentinel Distributed Database

Eric M. Ammann¹  | Marin L. Schweizer^{2,3} | Jennifer G. Robinson^{1,3,4} |
Jayasheel O. Eschol³ | Rami Kafa³ | Saket Girotra^{3,4} | Scott K. Winiecki⁵ |
Candace C. Fuller⁶ | Ryan M. Carnahan¹  | Charles E. Leonard⁷  | Cole Haskins^{1,4,8}  |
Crystal Garcia⁶ | Elizabeth A. Chrischilles¹ 



Standard Phenotype Library for Cardiovascular Disease

Requirements Development for the OHDSI Gold Standard Phenotype Library ✎

■ Researchers



apotvien Aaron Potvien

Aug 18

Greetings all,

Here at Georgia Tech, we've put together an initial requirements development document to help envision how an OHDSI Gold Standard Phenotype Library would function. To make the requirements more tangible, we put together a series of potential **personas** (representing different types of OHDSI stakeholders and collaborators), as well as **use cases** that capture the essence of what these users are trying to accomplish. The phenotype library is intended to be a home for validated, high-quality cohort phenotypes that can be generated using the OMOP CDM.

The full document can be [found here on Google Docs](#) ¹¹. We would greatly appreciate your input on additional personas and/or use cases that typify other characteristics you believe should be represented in order to make the OHDSI Gold Standard Phenotype Library a valuable new resource for the entire community.

Best,
Aaron

<http://forums.ohdsi.org/t/requirements-development-for-the-ohdsi-gold-standard-phenotype-library/4876>

Association of Ticagrelor vs Clopidogrel With Net Adverse Clinical Events in Patients With Acute Coronary Syndrome Undergoing Percutaneous Coronary Intervention


Seng Chan You, MD, MS; Yeunsook Rho, PhD; Behnood Bikdeli, MD, MS; Jiwoo Kim, MS; Anastasios Siapos, MSc; James Weaver, MSc; Ajit Londhe, MPH; Jaehyeong Cho, BS; Jimyung Park, BS; Martijn Schuemie, PhD; Marc A. Suchard, MD, PhD; David Madigan, PhD; George Hripcsak, MD, MS; Aakriti Gupta, MD, MS; Christian G. Reich, MD; Patrick B. Ryan, PhD; Rae Woong Park, MD, PhD; Harlan M. Krumholz, MD, SM


IMPORTANCE Current guidelines recommend ticagrelor as the preferred P2Y₁₂ platelet inhibitor for patients with acute coronary syndrome (ACS), primarily based on a single large randomized clinical trial. The benefits and risks associated with ticagrelor vs clopidogrel in routine practice merits attention.

OBJECTIVE To determine the association of ticagrelor vs clopidogrel with ischemic and hemorrhagic events in patients undergoing percutaneous coronary intervention (PCI) for ACS in clinical practice.

 [Editorial page 1](#)

 [JAMA Patient Page page 1](#)

 [Audio and Supplemental content](#)

 [CME Quiz at jamacmelookup.com and CME Questions page 0](#)

Seng Chan You¹; Yeunsook Rho²; Jiwoo Kim²; Anastasios Siapos³; Ajit Londhe⁴; Jaehyeong Cho⁵; Jimyung Park⁵; Martijn Schuemie⁴; Marc A Suchard, MD, PhD^{6,7}; David Madigan PhD⁸; George Hripcsak MD⁹; Christian G. Reich³; Patrick B. Ryan⁴; Rae Woong Park, MD, PhD^{1,5}; Harlan M. Krumholz, MD¹⁰

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Methods for phenotyping

- Rule-based phenotype
 - Human curated SNOMED-CT code
 - Leverage legacy of ICD-code system
 - ICD-9; ICD-10 (CM; K)
 - Read code
- Computational phenotype
 - APHRODITE
 - Others



Standard Phenotype Library for Cardiovascular Disease

- Why CVD? : Many OHDSI studies have been focused on CVD outcome
 - LEGEND (Lancet)
 - Ticagrelor vs Clopidogrel (JAMA)
 - Many more studies
- Targets
 - Acute myocardial infarction
 - Cerebrovascular accident (stroke)
 - Ischemic stroke
 - Hemorrhagic stroke
 - Heart failure
 - Sudden cardiac death
 - Major bleeding (hemorrhagic stroke + GI bleeding)



Why stroke?

- It is not easy to define stroke in SNOMED-CT system

Concept Set Expression

Included Concepts 51

Included Source Codes

Export

Import

Name:

Non-hemorrhagic ischemic stroke

Show

25

 entries

Showing 1 to 7 of 7 entries

	Concept Id	Concept Code	Concept Name
	4046089	230285003	Vascular dementia of acute onset
	4319146	95830009	Pituitary infarction
	379778	56267009	Multi-infarct dementia
	4046090	230287006	Mixed cortical and subcortical vascular dementia
	4046362	230706003	Hemorrhagic cerebral infarction
	4045741	230703006	Dysarthria-clumsy hand syndrome
	443454	432504007	Cerebral infarction

<http://www.ohdsi.org/web/atlas/#/cohortdefinition/1768950>



Why stroke?

Risk Prediction for Ischemic Stroke and Transient Ischemic Attack in Patients Without Atrial Fibrillation: A Retrospective Cohort Study

Zhong Yuan, MD, PhD,* Erica A. Voss, MPH,† Frank J. DeFalco, BA,†
Guohua Pan, PhD,† Patrick B. Ryan, PhD,* Daniel Yannicelli, MD,†¹ and
Christopher Neslund-Dudas, MD, PhD,†

Main Outcomes Measure

The composite of ischemic stroke or TIA (at 1-year or 3-year observable time window, respectively) was the main outcome of interest because the current study is intended to assess the performance of the existing risk schemes (i.e., CHADS₂ and CHA₂DS₂-VASc) for its prediction. The outcome postindex date was identified using the International Classification of Diseases, 9th Revision, Clinical Modification (ICD-9-CM) codes present in any diagnosis field in the database (ischemic stroke: 433.x1, 434.1, 434.x1; TIA: 435.x). We used all diagnosis fields to as-



Previous papers validating stroke in ICD code system

Validating Administrative Data in Stroke Research

PHARMACOEPIDEMIOLOGY AND DRUG SAFETY 2008; 17: 20–26

Published online 2 November 2007 in Wiley InterScience (www.interscience.wiley.com) DOI: 10.1002/pds.1518

PHARMACOEPIDEMIOLOGY AND DRUG SAFETY 2011; 20: 236–242

Published online 29 December 2010 in Wiley Online Library (wileyonlinelibrary.com) DOI: 10.1002/pds.2087



Cerebrovasc Dis Extra 2016;6:96–106

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Original Paper

Validation
value for
using
Validation
with isch

How Reliable Are Administrative Data for Capturing Stroke Patients and Their Care?

Ching-Lan Ch

search Database

Lee⁴ and Ming Liang Lai⁵

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Network, ^dInstitute for Health Policy, Management and Evaluation, University of Toronto,
and ^eOntario Stroke Network, Toronto, Ont., Canada

y, Tainan, Taiwan
l, Taiwan

Background and
data, and low
data.

Methods—Adm
algorithms fo
or the primar
data were con
discharge dia

Results—Comp
of 86%, spec
intracerebral
specificity wa
90%, 97%, 9
comorbidity &

Conclusions—S
stroke and pri
patients on
unrepresentat

Christianne
Cristina Va
and Marie

¹Veterans Adm
Center (GREC)
²HSR&D Targ
³Tennessee Val
⁴Departments
⁵Department o
⁶Pfizer Global
¹Institute of Bioph
²Institute of Clinic
³University of Illin
⁴Department of Ini
⁵Department of Ne



Systematic review for validation of stroke in ICD code system

PHARMACOEPIDEMOLOGY AND DRUG SAFETY 2012; 21(S1): 100–128
Published online in Wiley Online Library (wileyonlinelibrary.com) DOI: 10.1002/pds.2312

ORIGINAL REPORT

A systematic review of validated methods for identifying cerebrovascular accident or transient ischemic attack using administrative data

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Meyers Primary Care Institute (Reliant Medical Group, Fallon Community Health Plan, and University of Massachusetts Medical School), Worcester, MA, USA

ABSTRACT

Purpose To perform a systematic review of the validity of algorithms for identifying cerebrovascular accidents (CVAs) or transient ischemic attacks (TIAs) using administrative and claims data.

Methods PubMed and Iowa Drug Information Service searches of the English language literature were performed to identify studies published between 1990 and 2010 that evaluated the validity of algorithms for identifying CVAs (ischemic and hemorrhagic strokes, intracranial hemorrhage, and subarachnoid hemorrhage) and/or TIAs in administrative data. Two study investigators independently reviewed the abstracts and articles to determine relevant studies according to pre-specified criteria.

Results A total of 35 articles met the criteria for evaluation. Of these, 26 articles provided data to evaluate the validity of stroke, seven reported the validity of TIA, five reported the validity of intracranial bleeds (intracerebral hemorrhage and subarachnoid hemorrhage), and 10 studies reported the validity of algorithms to identify the composite endpoints of stroke/TIA or cerebrovascular disease. Positive predictive values (PPVs) varied depending on the specific outcomes and algorithms evaluated. Specific algorithms to evaluate the presence of stroke and intracranial bleeds were found to have high PPVs (80% or greater). Algorithms to evaluate TIAs in adult populations were generally found to have PPVs of 70% or greater.

Conclusions The algorithms and definitions to identify CVAs and TIAs using administrative and claims data differ greatly in the published literature. The choice of the algorithm employed should be determined by the stroke subtype of interest. Copyright © 2012 John Wiley & Sons, Ltd.

RESEARCH ARTICLE

Validity of Diagnostic Codes for Acute Stroke in Administrative Databases: A Systematic Review

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2 Arthritis Research Canada, Richmond, British Columbia, Canada, **3** Division of Rheumatology, Department of Medicine, University of British Columbia, Vancouver, British Columbia, Canada,

4 Cardiovascular Committee of the CANRAD Network, Richmond, British Columbia, Canada

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Diverse accuracy across the ICD codes

Jones[23], 2014	ICD-9-CM 433, any position	any stroke, definite or probable			10 (8-12)
	ICD-9-CM 433, primary position				15.82 (13-19)
	ICD-9-CM 434, any position				77 (74-79)
	ICD-9-CM 434, primary position				83.22 (81-86)
	ICD-9-CM 433 and 434, any position				41.56 (39.53-43.62)
	ICD-9-CM 433 and 434, primary position				54.86 (52.30-57.39)
	ICD-9-CM 436, any position				72 (66-77)
	ICD-9-CM 436, primary position				82.14 (76-87)
	ICD-9-CM 434 and 436, any position				75.60 (73.22-77.83)
	ICD-9-CM, 434 and 436, primary position				83.00 (80.61-85.15)
	ICD-9-CM 433/434/436, any position				44.94 (43.01-46.89)
	ICD-9-CM 433/434/436, primary position				58.40 (56.03-60.73)
	ICD-9-CM 433.01, 433.11, 433.21, 433.31, 433.81, 433.91, 434.01, 434.11, 434.91 (AHA/ASA code group), any position	<u>ischaemic stroke</u>			76 (74-79)



Diverse accuracy across the ICD codes

Benesch <i>et al.</i> ¹⁴	Hospitalizations at 5 academic medical centers identified using the Academic Medical Center Consortium database, 1992	Hospitalizations (stroke)	Inpatient ICD-9 codes 433 to 436	Medical record review was conducted (N = 649). Stroke was confirmed based upon the World Health Organization (WHO) definitions. Primary and secondary diagnoses: Code 433: PPV = 6.1% Code 434: PPV = 85.0% Code 435: PPV = 9.1% Code 436: PPV = 82.6% Primary diagnosis: Code 433: PPV = 9.1% Code 434: PPV = 90.3% Code 435: PPV = 6.3% Code 436: PPV = 88.9%
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Diverse accuracy across the ICD codes

Table 5. Positive predictive values of International Classification of Diseases codes to identify cerebrovascular accident and transient ischemic attacks in adult populations*

ICD-9/ICD-10 code	Number of studies reporting PPV estimate	Median PPV estimate	Range of PPV estimates (minimum, maximum)	Number of studies reporting PPV estimate	Median PPV estimate	Range of PPV estimates (minimum, maximum)
Studies evaluating acute stroke events						
Using principal or most responsible diagnosis only			Using all discharge diagnoses			
430	3	87	33, 100	4	84	74, 100
431	3	88	80, 100	4	86	83, 93
432	3	21	17, 29	4	20	0, 32
432.9	0			1	60	
433	3	17	9, 46	4	15	6, 15
433.x0	1	13		0		
433.x1	1	71		0		
434	3	90	84, 92	4	85	77, 85
434.x0	1	33		0		
434.x1	1	72		0		
434.0	1	85		1	82	
434.1	1	80		1	58	
435	4	17	3, 29	4	14	9, 26
436	4	80	48, 89	5	81	70, 86
437	3	50	45, 69	3	22	2, 31
438	2	20	8, 33	3	1	0, 7
Studies evaluating ischemic stroke events						
Using principal or most responsible diagnosis only			Using all discharge diagnoses			
433	1	4		1	14	
434	2	87	82, 92	1	77	
434.11	1	85		0		
434.91	1	82		0		
435	0			1	12	
436	1	79		1	68	
437	0			1	2	
438	0			1	0	



My conclusion for stroke

- Diagnosis code
 - ICD-9-CM: 433.x1, 434.x1
 - ICD10: I63x
- Should be considered
 - Specifiers
 - Inpatient or ED visit only?
 - Primary or secondary diagnosis only?
 - Brain CT or MRI?

Diverse accuracy across the ICD codes

Table 5. Positive predictive values of International Classification of Diseases codes to identify cerebrovascular accident and transient ischemic attacks in adult populations*

ICD-9/ICD-10 code	Number of studies reporting PPV estimate	Median PPV estimate	Range of PPV estimates (minimum, maximum)	Number of studies reporting PPV estimate	Median PPV estimate	Range of PPV estimates (minimum, maximum)
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432	3	21	17, 29	4	20	0, 32
432.9	0			1	60	
433	3	17	9, 46	4	15	6, 15
433.x0	1	13		0		
433.x1	1	71		0		
434	3	90	84, 92	4	85	77, 85
434.x0	1	33		0		
434.x1	1	72		0		
434.0	1	85		1	82	
434.1	1	80		1	58	
435	4	17	3, 29	4	14	9, 26
436	4	80	48, 89	5	81	70, 86
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Studies evaluating ischemic stroke events						
Using principal or most responsible diagnosis only				Using all discharge diagnoses		
433	1	4		1	14	
434	2	87	82, 92	1	77	
434.11	1	85		0		
434.91	1	82		0		
435	0			1	12	
436	1	79		1	68	
437	0			1	2	
438	0			1	0	



ICD-9-CM (433 & 434)

- ▶ 433 Occlusion and stenosis of precerebral arteries
 - ▶ 433.0 Occlusion and stenosis of basilar artery
 - ▶ 433.00 Occlusion and stenosis of basilar artery without mention of cerebral infarction convert 433.00 to ICD-10-CM
 - ▶ 433.01 Occlusion and stenosis of basilar artery with cerebral infarction convert 433.01 to ICD-10-CM
 - ▶ 433.1 Occlusion and stenosis of carotid artery
 - ▶ 433.10 Occlusion and stenosis of carotid artery without mention of cerebral infarction convert 433.10 to ICD-10-CM
 - ▶ 433.11 Occlusion and stenosis of carotid artery with cerebral infarction convert 433.11 to ICD-10-CM
 - ▶ 433.2 Occlusion and stenosis of vertebral artery
 - ▶ 433.20 Occlusion and stenosis of vertebral artery without mention of cerebral infarction convert 433.20 to ICD-10-CM
 - ▶ 433.21 Occlusion and stenosis of vertebral artery with cerebral infarction convert 433.21 to ICD-10-CM
 - ▶ 433.3 Occlusion and stenosis of multiple and bilateral precerebral arteries
 - ▶ 433.30 Occlusion and stenosis of multiple and bilateral precerebral arteries without mention of cerebral infarction convert 433.30 to ICD-10-CM
 - ▶ 433.31 Occlusion and stenosis of multiple and bilateral precerebral arteries with cerebral infarction convert 433.31 to ICD-10-CM
 - ▶ 433.8 Occlusion and stenosis of other specified precerebral artery
 - ▶ 433.80 Occlusion and stenosis of other specified precerebral artery without mention of cerebral infarction convert 433.80 to ICD-10-CM
 - ▶ 433.81 Occlusion and stenosis of other specified precerebral artery with cerebral infarction convert 433.81 to ICD-10-CM
 - ▶ 433.9 Occlusion and stenosis of unspecified precerebral artery
 - ▶ 433.90 Occlusion and stenosis of unspecified precerebral artery without mention of cerebral infarction convert 433.90 to ICD-10-CM
 - ▶ 433.91 Occlusion and stenosis of unspecified precerebral artery with cerebral infarction convert 433.91 to ICD-10-CM
- ▶ 434 Occlusion of cerebral arteries
 - ▶ 434.0 Cerebral thrombosis
 - ▶ 434.00 Cerebral thrombosis without mention of cerebral infarction convert 434.00 to ICD-10-CM
 - ▶ 434.01 Cerebral thrombosis with cerebral infarction convert 434.01 to ICD-10-CM
 - ▶ 434.1 Cerebral embolism
 - ▶ 434.10 Cerebral embolism without mention of cerebral infarction convert 434.10 to ICD-10-CM
 - ▶ 434.11 Cerebral embolism with cerebral infarction convert 434.11 to ICD-10-CM
 - ▶ 434.9 Cerebral artery occlusion unspecified
 - ▶ 434.90 Cerebral artery occlusion, unspecified without mention of cerebral infarction convert 434.90 to ICD-10-CM
 - ▶ 434.91 Cerebral artery occlusion, unspecified with cerebral infarction convert 434.91 to ICD-10-CM



ICD-10 (I63, I64)

I63 Cerebral infarction

Incl.: occlusion and stenosis of cerebral and precerebral arteries, resulting in cerebral infarction

Excl.: sequelae of cerebral infarction ([I69.3](#))

I63.0 Cerebral infarction due to thrombosis of precerebral arteries

I63.1 Cerebral infarction due to embolism of precerebral arteries

I63.2 Cerebral infarction due to unspecified occlusion or stenosis of precerebral arteries

I63.3 Cerebral infarction due to thrombosis of cerebral arteries

I63.4 Cerebral infarction due to embolism of cerebral arteries

I63.5 Cerebral infarction due to unspecified occlusion or stenosis of cerebral arteries

I63.6 Cerebral infarction due to cerebral venous thrombosis, nonpyogenic

I63.8 Other cerebral infarction

I63.9 Cerebral infarction, unspecified

I64 Stroke, not specified as haemorrhage or infarction

Incl.: Cerebrovascular accident NOS

Excl.: sequelae of stroke ([I69.4](#))



SNOMED-CT code

- These two ancestor concept_ids can be used to include 'maps to' SNOMED-CT OMOP concept_id for ICD-codes(433.x1, 434.x1, I63x)

concept_id	concept_name	domain_id	vocabulary_id	concept_class_id	standard_concept	concept_code
443454	Cerebral infarction	Condition	SNOMED	Clinical Finding	S	432504007
4043731	Infarction - precerebral	Condition	SNOMED	Clinical Finding	S	230692004



Unwanted 'maps from' ICD codes

	concept code	vocabulary id	source name
1	G43.6	ICD10CM	Persistent migraine aura with cerebral infarction
2	G43.60	ICD10CM	Persistent migraine aura with cerebral infarction, not intractable
3	G43.601	ICD10CM	Persistent migraine aura with cerebral infarction, not intractable, with status migrainosus
4	G43.609	ICD10CM	Persistent migraine aura with cerebral infarction, not intractable, without status migrainosus
5	G43.61	ICD10CM	Persistent migraine aura with cerebral infarction, intractable
6	G43.611	ICD10CM	Persistent migraine aura with cerebral infarction, intractable, with status migrainosus
7	G43.619	ICD10CM	Persistent migraine aura with cerebral infarction, intractable, without status migrainosus
8	G46.5	ICD10CM	Pure motor lacunar syndrome
9	G46.6	ICD10CM	Pure sensory lacunar syndrome
10	G46.7	ICD10CM	Other lacunar syndromes
11	I97.81	ICD10CM	Intraoperative cerebrovascular infarction
12	I97.810	ICD10CM	Intraoperative cerebrovascular infarction during cardiac surgery
13	I97.811	ICD10CM	Intraoperative cerebrovascular infarction during other surgery
14	I97.82	ICD10CM	Postprocedural cerebrovascular infarction
15	I97.820	ICD10CM	Postprocedural cerebrovascular infarction following cardiac surgery
16	I97.821	ICD10CM	Postprocedural cerebrovascular infarction following other surgery
17	346.6	ICD9CM	Persistent migraine aura with cerebral infarction
18	346.60	ICD9CM	Persistent migraine aura with cerebral infarction, without mention of intractable migraine without mention of status migrainosus
19	346.61	ICD9CM	Persistent migraine aura with cerebral infarction, with intractable migraine, so stated, without mention of status migrainosus
20	346.62	ICD9CM	Persistent migraine aura with cerebral infarction, without mention of intractable migraine with status migrainosus
21	346.63	ICD9CM	Persistent migraine aura with cerebral infarction, with intractable migraine, so stated, with status migrainosus
22	997.02	ICD9CM	Iatrogenic cerebrovascular infarction or hemorrhage

Unwanted 'maps from' ICD codes

	concept code	vocabulary id	source name
1	G43.6	ICD10CM	Persistent migraine aura with cerebral infarction
2	G43.60	ICD10CM	Persistent migraine aura with cerebral infarction, not intractable
3	G43.601	ICD10CM	Persistent migraine aura with cerebral infarction, not intractable, with status migrainosus
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5	G43.61	ICD10CM	Persistent migraine aura with cerebral infarction, intractable
6	G43.611	ICD10CM	Persistent migraine aura with cerebral infarction, intractable, with status migrainosus
7	G43.619	ICD10CM	Persistent migraine aura with cerebral infarction, intractable, without status migrainosus
8	G46.5	ICD10CM	Pure motor lacunar syndrome
9	G46.6	ICD10CM	Pure sensory lacunar syndrome
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22	997.02	ICD9CM	Iatrogenic cerebrovascular infarction or hemorrhage

SELECT COUNT(DISTINCT PERSON_ID) FROM
CONDITION_OCCURRENCE WHERE
condition_source_value LIKE 'G436%'
→ 0 (No one has this diagnosis code in Korea,
NHIS-NSC)



Three options

1. We can validate concept id of **443454, 4043731** in OHDSI – The best option
2. We can make a stroke cohort definition with excluding terms for same-day migraine
3. We can count how many people actually have these diagnoses in multiple databases. If no database has this condition, then I would be relieved

concept_id	concept_name	domain_id	vocabulary_id	concept_class_id	standard_concept	concept_code
443454	Cerebral infarction	Condition	SNOMED	Clinical Finding	S	432504007
4043731	Infarction - precerebral	Condition	SNOMED	Clinical Finding	S	230692004


Association of Ticagrelor vs Clopidogrel With Net Adverse Clinical Events in Patients With Acute Coronary Syndrome Undergoing Percutaneous Coronary Intervention


Seng Chan You, MD, MS; Yeunsook Rho, PhD; Behnood Bikdeli, MD, MS; Jiwoo Kim, MS; Anastasios Siapos, MSc; James Weaver, MSc; Ajit Londhe, MPH; Jaehyeong Cho, BS; Jimyung Park, BS; Martijn Schuemie, PhD; Marc A. Suchard, MD, PhD; David Madigan, PhD; George Hripcsak, MD, MS; Aakriti Gupta, MD, MS; Christian G. Reich, MD; Patrick B. Ryan, PhD; Rae Woong Park, MD, PhD; Harlan M. Krumholz, MD, SM


IMPORTANCE Current guidelines recommend ticagrelor as the preferred P2Y₁₂ platelet inhibitor for patients with acute coronary syndrome (ACS), primarily based on a single large randomized clinical trial. The benefits and risks associated with ticagrelor vs clopidogrel in routine practice merits attention.

OBJECTIVE To determine the association of ticagrelor vs clopidogrel with ischemic and hemorrhagic events in patients undergoing percutaneous coronary intervention (PCI) for ACS in clinical practice.

 [Editorial page 1](#)

 [JAMA Patient Page page 1](#)

 [Audio and Supplemental content](#)

 [CME Quiz at jamacmelookup.com and CME Questions page 0](#)

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☆ MY PROJECTS

🏠 ATLAS

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Network Overview

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🏢 DATA NETWORK

👥 EXPERT NETWORK

Service & Events

🔍 EVIX-INSIGHT™

🚀 EVIX-EXPLO™

🔧 COVID-19 STUDY

교육계정 (05 fnet) ▼



ATLAS

ATLAS는 OMOP CDM 기반의 표준화된 분석 기능을 제공하고 있는 OHDSI의 오픈 소프트웨어입니다. 에비드넷은 각 기관 서버에 ATLAS를 설치하여 원활한 CDM 연구가 진행될 수 있도록 돕고 있습니다. 기관연구자 혹은 협력연구자로 승인 받은 연구자는 해당 기관의 ATLAS에 접근하여 연구 설계를 할 수 있습니다. RFZ 참여 기관 간에는 연구자 등록 과정 없이 서로의 기관 ATLAS에 접속하셔서 자유로운 연구 설계를 진행할 수 있습니다.

ATLAS 사용법에 대한 자세한 안내는 튜토리얼을 참고하세요



[ATLAS v2.7 튜토리얼 강의자료](#)



[ATLAS v2.7 튜토리얼 교육영상](#)

Connect to ATLAS

ORGANIZATION NAME	RFZ LEVEL	API VERSION	RESEARCH RIGHT	CONNECT
Open Database	-	v2.7.6		Connect

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chandryou Merge pull request #65 from ohdsi-studies/revision ...

on 7 Oct 2020 137

 R	Extract mean and sd of age	17 months ago
 documents	fix the additional result function for revision	17 months ago
 extras	add whole code to generate figures and tables fo...	2 years ago
 inst	fix the additional result function for revision	17 months ago
 man	fix the additional result function for revision	17 months ago



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
chandryou Merge pull request #65 from ohdsi-studies/revision ...

on 7 Oct 2020 137




 R	Extract mean and sd of age	17 months ago
 documents	fix the additional result function for revision	17 months ago
 extras	add whole code to generate figures and tables fo...	2 years ago
 inst	fix the additional result function for revision	17 months ago
 man	fix the additional result function for revision	17 months ago













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 **master**  **1 branch**  **0 tags** Go to file Code

 **chandryou** undo redundant data 34a8076 16 seconds ago  **24 commits**

	2019SymposiumTutorial-Achitecture	AWS Achitecture	3 years ago
	2019SymposiumTutorial-PLE	update the materials	3 years ago
	2019SymposiumTutorial-PLP	minor change in README.md	3 years ago
	2019SymposiumTutorial-Phenotyping	phenotyping	3 years ago
	2019SymposiumTutorial-Voca	change readme file	3 years ago
	2021KOSHIS-ReplicaTutorial	revise outcome cohort defintion and add slides	9 months ago
	2022KOSHIS-Phenotyping/Cohorts	undo redundant data	17 seconds ago
	26thOct2019	update README	3 years ago

<https://github.com/ohdsi-korea/OhdsiKoreaTutorials/>

Validation using Manual chart review

eMethod 3. Individual outcome definitions

For each outcome, we developed an operational phenotype definition to determine if observational data could in fact support evaluation of the outcome. Where possible, concept sets originated with published code lists (eg ICD-9-CM and ICD-10). We developed definition of outcome cohorts and query to extract them using ATLAS, the OHDSI open-source platform (<https://github.com/OHDSI/atlas>). We executed these definitions on EHR data of Korean tertiary hospital to validate the definitions. Positive predictive values were estimated by a physician's manual chart review of discharge notes.

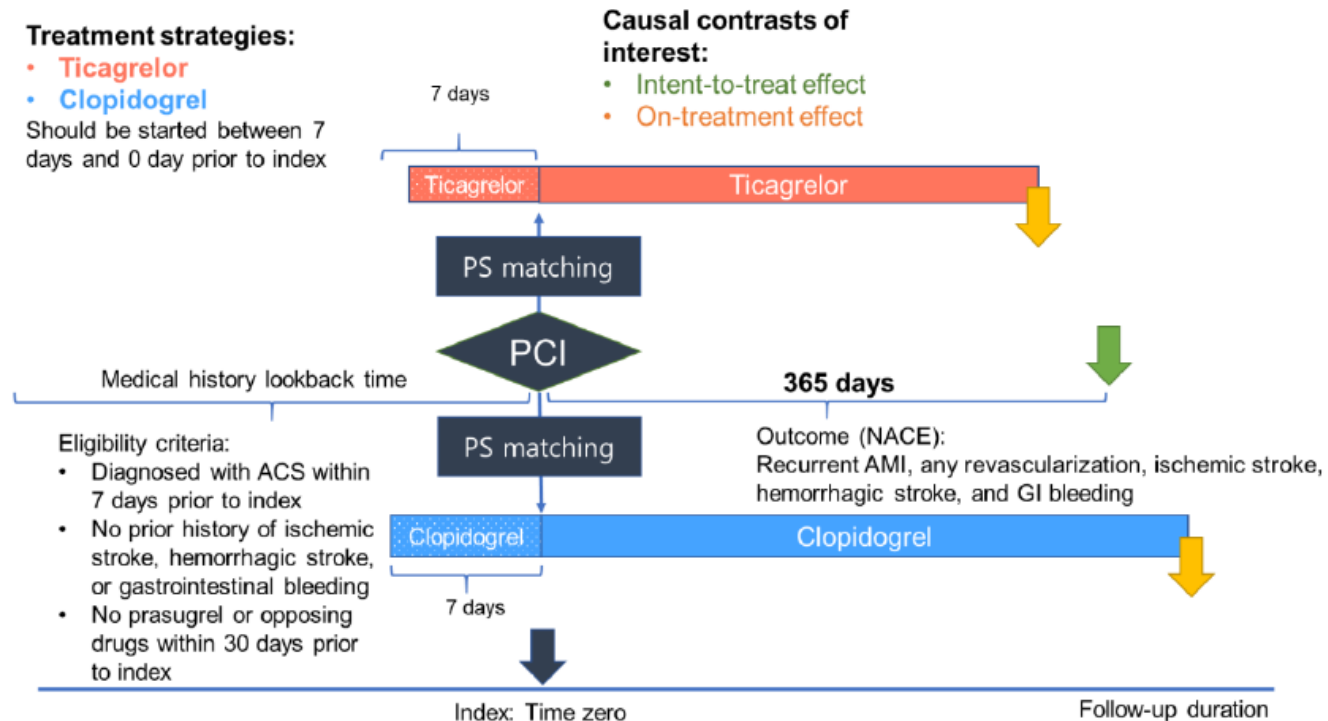
Supplementary Table. Outcome definition

Outcome	Logical description	ICD-9-CM	ICD-10	CPT4	PPV, % (n)
Acute myocardial infarction	Record of acute myocardial infarction during an inpatient or ER visit	410;410.01;410.02;410.1;410.11;410.12;410.2;410.21;410.22;410.3;410.31;410.32;410.4;410.41;410.42;410.5;410.51;410.52;410.7;410.71;410.72;410.8;410.81;410.82;410.9;410.91;410.92	I21.0;I21.1;I21.2;I21.3;I21.4;I21.9		83.8 (83/99)
Revascularization	Record of PCI or CABG during an inpatient or ER visit			566;567;33510;33511;33512;33513;33514;33516;33517;33518;33519;33521;33522;33523;33533;33534;33535;33536;33542;33545;33548;33572;33621;35506;35694;92920;92921;92924;92925;92928;92929;92933;92934;92937;92938;92941;92943;92944;1006199;1006200;1006208;1006216;1006217	100.0 (30/30)
Ischemic stroke	Earliest record of ischemic stroke during an inpatient or ER visit	346.6;346.6;346.61;346.62;346.63;433.01;433.11;433.21;433.31;433.81;433.91;434.01;434.11;434.91;997.02	I63.9;I63.8;I63.6;I63.5;I63.4;I63.3;I63.2;I63.1;I63.0;I63;G46.7;G46.6;G46.5;F01.3;F01.1;F01.0		72.9 (70/96)
Hemorrhagic stroke	Earliest record of intracranial hemorrhage without concomitant ischemic stroke during an inpatient or ER visit	430;431;432;432;432.1;432.9	I60;I60.0;I60.5;I60.6;I60.7;I60.8;I60.9;I61.0;I61.1;I61.2;I61.3;I61.4;I61.5;I62;I62.0;I62.1;I62.9		100.0 (46/46)
Gastrointestinal bleeding	Gastrointestinal hemorrhage condition record during an inpatient or ER visit	530.21;530.7;530.82;531;531.531.01;531.2;531.2;531.21;531.4;531.4;531.41;531.6;531.6;531.61;532;532;532.01;532.2;532.2;532.21;532.4;532.4;532.41;532.6;532.6;532.61;533;533;533.01;533.2;533.2;533.21;533.4;533.4;533.41;533.6;533.6;533.61;534;534.01;534.2;534.2;534.21;534.4;534.4;534.41;534.6;534.6;534.61;535.01;535.11;535.21;535.31;535.41;535.51;535.61;535.71;537.83;537.84;562.02;562.03;562.12;562.13;569.3;569.85;578;578.578.1;578.9	K22.6;K25.0;K25.2;K25.4;K25.6;K26.0;K26.2;K26.4;K26.6;K27.0;K27.2;K27.4;K27.6;K28.0;K28.2;K28.4;K28.6;K62.5;K92.0;K92.1;K92.2		95.8 (68/71)
Dyspnea	Record of dyspnea	786.02;786.05	R06.0		94.7 (18/19)



Method: Study Population

- Inclusion Criteria
 - Adults (≥ 20 yrs) who initiated ticagrelor or clopidogrel due to acute coronary syndrome (ACS) and undertook percutaneous coronary intervention (PCI)
- Exclusion Criteria
 - Prior history of stroke or gastrointestinal bleeding
 - Use of prasugrel or opposing drug within previous 30 days from index date





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Column visibility

Copy

CSV

Show 15 entries

Showing 1 to 15 of 2,175 entries

Previous

1

2

3

4

5

...

145

Next

Id	Name	Created	Updated	Author
2572	[SCYou]Ticagrelor	09/29/2021 8:46 PM	09/29/2021 8:46 PM	anonymous
2571	COPY OF: Schizo_YH (1)	09/29/2021 8:17 PM	09/29/2021 8:45 PM	anonymous
2570	COPY OF: Schizo_YH	09/29/2021 5:20 PM	09/29/2021 7:59 PM	anonymous
2500	T_Anti-TB drugs with PZA_dh	09/24/2021 7:09 PM	09/29/2021 2:15 PM	anonymous
2503	O_Toxic liver diseases_dh	09/24/2021 8:06 PM	09/29/2021 1:41 PM	anonymous
2549	Schizo_YH	09/28/2021 2:54 PM	09/29/2021 10:58 AM	anonymous
2569	epilepsy_YH	09/29/2021 10:57 AM	09/29/2021 10:57 AM	anonymous
2568	depressive disorder_YH	09/29/2021 10:55 AM	09/29/2021 10:56 AM	anonymous
2567	AD_Duloxetine_YH	09/29/2021 10:53 AM	09/29/2021 10:53 AM	anonymous
2565	AD_fluvoxamine_YH	09/29/2021 10:50 AM	09/29/2021 10:51 AM	anonymous
2566	AD mirtazapine_YH	09/29/2021 10:51 AM	09/29/2021 10:51 AM	anonymous

New Cohort

Filter:



[MyName]Ticagrelor



Definition



Concept Sets

Generation

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Text View

Graphical View

JSON

SQL

```
"STANDARD_CONCEPT_CAPTION": "Standard",  
"INVALID_REASON": "V",  
"INVALID_REASON_CAPTION": "Valid",  
"CONCEPT_CODE": "11101003",  
"DOMAIN_ID": "Procedure",  
"VOCABULARY_ID": "SNOMED",  
"CONCEPT_CLASS_ID": "Procedure",  
},  
"includeDescendants": true  
},  
{  
  "concept": {  
    "CONCEPT_ID": 4264285,  
    "CONCEPT_NAME": "Percutaneous transluminal coronary angioplasty by rotoablation",  
    "STANDARD_CONCEPT": "S",  
    "STANDARD_CONCEPT_CAPTION": "Standard",  
    "INVALID_REASON": "V",  
    "INVALID_REASON_CAPTION": "Valid"
```

Copy and paste the
cohort.json file

Copy To Clipboard

Reload



Cohort #2572

[SCYou]Ticagrelor

Definition

Concept Sets

Generation

Reporting

Export

Messages

enter a cohort definition description here

Cohort Entry Events

Events having any of the following criteria:

a procedure occurrence of PCI

+ Add attribute...

Delete Criteria

✗ for the first time in the person's history

✗ with age Greater or Equal To 20

with continuous observation of at least 365 days before and 0 days after event index date

Limit initial events to: earliest event per person.

Restrict initial events to:

having all of the following criteria:

with at least 1 using all occurrences of:

a condition occurrence of ACS

+ Add attribute...

where event starts between 7 days Before and 0 days After index start date add additional constraint

☐ restrict to the same visit occurrence

☐ allow events from outside observation period

and with at least 1 using all occurrences of:

a drug exposure of ticagrelor

+ Add attribute...

where event starts between 7 days Before and 0 days After index start date add additional constraint

☐ restrict to the same visit occurrence

☐ allow events from outside observation period

Limit initial events to: earliest event per person.

Remove initial event restriction



#1 Create Concept Set: Ticagrelor

[SCYou]Ticagrelor

Definition ? Concept Sets Generation Reporting Export Messages 8

New Concept Set Import

Show 10 entries Filter Cohort C

Id	Title
1	ACS
3	aspirin
5	clopidogrel
6	Coronary Stent Implantation
10	GI bleeding
9	ICH
8	Ischemic stroke
2	MI
0	PCI
7	Prasugrel



#1 Create Concept Set: Ticagrelor

[SCYou]Ticagrelor

Definition ? Concept Sets Generation Reporting Export Messages 8

New Concept Set Import

Show 10 entries Filter Cohort C

Id	Title
1	ACS
3	aspirin
5	clopidogrel
6	Coronary Stent Implantation
10	GI bleeding
9	ICH
8	Ischemic stroke
2	MI
0	PCI
7	Prasugrel



#1 Create Concept Set: Ticagrelor

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Q Search

Search Import

ticagrelor

Column visibility

Copy

CSV

Show 15 entries

Filter: [

Showing 1 to 15 of 232 entries

Vocabulary

RxNorm Extension (89)

NDFRT (32)

SNOMED (19)

SPL (19)

Class

Marketed Product (40)

	Id	Code	Name	Class	RC	DRC
	46240683	4f9bd52a-99a7-4606-92b4-456a917aabef	BRILINTA - ticagrelor tablet	Prescription Drug	0	5,172
	45680509	f7b3f443-e83d-4bf2-0e96-023448fed9a8	BRILINTA - ticagrelor tablet	Prescription Drug	0	5,172
	46314076	3ce9ee32-b18b-4ef3-b127-ce479d520e1a	BRILINTA - ticagrelor tablet	Prescription Drug	0	5,172
	1370827	7cee944f-e2ca-4fc8-bb97-f46ca28728c5	TICAGRELOR - ticagrelor tablet	Prescription Drug	0	5,172
	40241186	1116632	Ticagrelor	Ingredient	1,144	5,172

#1 Create Concept Set: Ticagrelor

2

← [SCYou]Ticagrelor ▶ Ticagrelor2

Search

Search

Import

ticagrelor

Column visibility

Copy

CSV

Show 15 entries

Showing 1 to 15 of 232 entries

	Id	Code	Name
	46240683	4f9bd52a-99a7-4606-92b4-456a917aabef	BRILINTA - ticagrelor tablet
	45680509	f7b3f443-e83d-4bf2-0e96-023448fed9a8	BRILINTA - ticagrelor tablet
	46314076	3ce9ee32-b18b-4ef3-b127-ce479d520e1a	BRILINTA - ticagrelor tablet
	1370827	7cee944f-e2ca-4fc8-bb97-f46ca28728c5	TICAGRELOR - ticagrelor tablet
	40241186	1116632	Ticagrelor

Vocabulary

RxNorm Extension (89)

NDFRT (32)

SNOMED (19)

SPL (19)

NDC (17)

Class

Marketed Product (40)

1



#1 Create Concept Set: Ticagrelor

Concept Set Expression

Included Concepts 104

Included Source Codes

Export

Import

Name:

ticagrelor

Show 25 entries

Search:

Showing 1 to 1 of 1 entries

Previous 1 Next

	Concept Id	Concept Code	Concept Name	Domain	Standard Concept Caption	Exclude	Descendants	Mapped
	40241186	1116632	Ticagrelor	Drug	Standard	<input type="checkbox"/>	<input checked="" type="checkbox"/>	<input type="checkbox"/>

Classification Non-Standard Standard

Delete Concept Set

Copy To Concept Set Repository

Close Concept Set



#1 Create Concept Set: ACS

Name:







ACS

Show 25 entries

Search:

Showing 1 to 5 of 5 entries

Previ

	Concept Id	Concept Code	Concept Name	Domain	Standard Concept Caption	Exclude	Descendants
	315296	4557003	Preinfarction syndrome	Condition	Standard	<input type="checkbox"/>	<input checked="" type="checkbox"/>
	444406	70422006	Acute subendocardial infarction	Condition	Standard	<input type="checkbox"/>	<input checked="" type="checkbox"/>
	438170	73795002	Acute myocardial infarction of inferior wall	Condition	Standard	<input type="checkbox"/>	<input checked="" type="checkbox"/>
	434376	54329005	Acute myocardial infarction of anterior wall	Condition	Standard	<input type="checkbox"/>	<input checked="" type="checkbox"/>
	312327	57054005	Acute myocardial infarction	Condition	Standard	<input type="checkbox"/>	<input checked="" type="checkbox"/>



#2 Create Target Cohort:Ticagrelor

Cohort Entry Events

Events having any of the following criteria:

a procedure occurrence of

PCI



✗ for the first time in the person's history

✗ with age Greater or Equal To

20

with continuous observation of at least 0 days before and 0 days after event index date

Limit initial events to: earliest event per person.



#2 Create Target Cohort:Ticagrelor

Restrict initial events to:

having of the following criteria:

with using all occurrences of:

a condition occurrence of

ACS

where between days and days

☐ restrict to the same visit occurrence

☐ allow events from outside observation period

and with using all occurrences of:

a drug exposure of

ticagrelor

where between days and days

☐ restrict to the same visit occurrence

☐ allow events from outside observation period



#2 Create Target Cohort: Ticagrelor

New inclusion criteria

1. Without clopidogrel or prasugrel on the day of PCI
2. Without previous stroke
3. Without previous GI bleeding

Without clopidogrel or prasugrel on the day of PCI

enter an inclusion rule description

having all of the following criteria:

+ Add cri

with at most 0 using all occurrences of:

a drug exposure of clopidogrel

+ Add attribute...

where event starts between 30 days Before and 0 days After

index start date [add additional constraint](#)

☐ restrict to the same visit occurrence

☐ allow events from outside observation period

and with at most 0 using all occurrences of:

a drug exposure of Prasugrel

+ Add attribute...

where event starts between 30 days Before and 0 days After

index start date [add additional constraint](#)

☐ restrict to the same visit occurrence

☐ allow events from outside observation period

#2 Create Target Cohort:Ticagrelor

Inclusion Criteria

New inclusion criteria

1. Without clopidogrel or prasugrel on the day of PCI
2. Without previous stroke
3. Without previous GI bleeding

Without previous stroke

enter an inclusion rule description

having all of the following criteria:

+ Add cr

with at most 0 using all occurrences of:

a condition occurrence of Ischemic stroke

+ Add attribute...

where event starts between All days Before and 0 days After

index start date [add additional constraint](#)

☐ restrict to the same visit occurrence

☐ allow events from outside observation period

and with at most 0 using all occurrences of:

a condition occurrence of ICH

+ Add attribute...

where event starts between All days Before and 0 days After

index start date [add additional constraint](#)

☐ restrict to the same visit occurrence

☐ allow events from outside observation period



#2 Create Target Cohort:Ticagrelor

New inclusion criteria

1. Without clopidogrel or prasugrel on the day of PCI
2. Without previous stroke
3. Without previous GI bleeding

Without previous GI bleeding

enter an inclusion rule description

having of the following criteria:

with using all occurrences of:

a condition occurrence of

GI bleeding



where between days and days

[add additional constraint](#)

☐ restrict to the same visit occurrence

☐ allow events from outside observation period



#2 Create Target Cohort:Ticagrelor

Cohort Exit

Event Persistence:

Event will persist until: end of a continuous drug exposure ▼

Continuous Exposure Persistence:


Specify a concept set that contains one or more drugs. A drug era will be added to the concept set, using the specified persistence window as a maximum, adding a specified surveillance window to the final exposure event. If the end date is inferred to be event start date + days supply in cases when day supply persistence assures that the cohort end date will be no greater than the cohort exit date.

Concept set containing the drug(s) of interest: ticagrelor ▼






- Persistence window: allow for a maximum of 7 ▼ days between events
- Surveillance window: add 0 ▼ days to the end of the era of persistence to cohort exit.




#3 Create Comparator Cohort: Clon

 Cohort #2572

[SCYou]Ticagrelor



Definition  Concept Sets Generation Reporting Export Messages 8

enter a cohort definition description here

COPY!



#3 Create Comparator Cohort: Clop



Cohort #2583

COPY OF: [SCYou]Ticagrelor

Rename -> [Name] Clopidogrel



Definition ?

Concept Sets

Generation

Reporting

Export

Messages

8

enter a cohort definition description here

Cohort Entry Events

Events having any of the following criteria:



#3 Create Comparator Cohort: Clop

with using all occurrences of:

a condition occurrence of

[+ Add attribute...](#)

where between days and days [add additional constraint](#)

☐ restrict to the same visit occurrence

☐ allow events from outside observation period

and with using all occurrences of:

a drug exposure of

[+ Add attribute...](#)

where between days and days [add additional constraint](#)

☐ restrict to the same visit occurrence

☐ allow events from outside observation period



#3 Create Comparator Cohort: Clop

New inclusion criteria

1. Without ticagrelor or prasugrel on the day of PCI
2. Without previous stroke
3. Without previous GI bleeding

Without ticagrelor or prasugrel on the day of PCI

enter an inclusion rule description

having of the following criteria:

+ Add cri

with using all occurrences of:

a drug exposure of

ticagrelor

+ Add attribute...▼

where between days and days

[add additional constraint](#)

☐ restrict to the same visit occurrence

☐ allow events from outside observation period



#3 Create Comparator Cohort: Clop

Cohort Exit

Event Persistence:

Event will persist until:

Continuous Exposure Persistence:

Specify a concept set that contains one or more drugs. A drug era will be identified based on the concept set, using the specified persistence window as a maximum length of time between events, adding a specified surveillance window to the final exposure event. If no date is inferred to be event start date + days supply in cases when days supply is not provided, persistence assures that the cohort end date will be no greater than the

Concept set containing the drug(s) of interest

- Persistence window: allow for a maximum of days between events
- Surveillance window: add days to the end of the era of persistence



#4 Create Outcome Cohort:AMI

Name:






Acute MI

Show 25 entries

Search:

Showing 1 to 4 of 4 entries

Previous 1 Next

	Concept Id	Concept Code	Concept Name	Domain	Standard Concept Caption	Exclude	Descendants	Mapped
	444406	70422006	Acute subendocardial infarction	Condition	Standard	<input type="checkbox"/>	<input checked="" type="checkbox"/>	<input type="checkbox"/>
	438170	73795002	Acute myocardial infarction of inferior wall	Condition	Standard	<input type="checkbox"/>	<input checked="" type="checkbox"/>	<input type="checkbox"/>
	434376	54329005	Acute myocardial infarction of anterior wall	Condition	Standard	<input type="checkbox"/>	<input checked="" type="checkbox"/>	<input type="checkbox"/>
	312327	57054005	Acute myocardial infarction	Condition	Standard	<input type="checkbox"/>	<input checked="" type="checkbox"/>	<input type="checkbox"/>



#4 Create Outcome Cohort:AMI

Cohort Entry Events



Events having any of the following criteria:

+ Add Initial Event ▼

a condition occurrence of

Acute MI ▼

+ Add attribute...▼

Delete Criteria

with a Visit occurrence of:

✖ Emergency Room Visit

✖ Emergency Room and Inpatient Visit



✖ Inpatient Visit

Add

Import

with continuous observation of at least 0 ▼ days before and 0 ▼ days after event index date

Limit initial events to: all events ▼ per person.

Restrict initial events



#4 Create Outcome Cohort:AMI

Show 10 entries

Filter records:

	Concept Id	Code	Concept Name	Standard Type	Vocabulary	Domain
✓	8870	23	Emergency Room - Hospital	S	CMS Place of Service	Visit
✓	38004220	261QE0002X	Ambulatory Emergency Care Clinic/Center	S	NUCC	Visit
✓	38004337	333300000X	Emergency Response System Supplier	S	NUCC	Visit
✓	38004362	343900000X	Non-emergency Medical Transport	S	NUCC	Visit
✓	9203	ER	Emergency Room Visit	S	Visit	Visit
✓	262	ERIP	Emergency Room and Inpatient Visit	S	Visit	Visit
✓	581381	OMOP4822040	Emergency Room Critical Care Facility	S	CMS Place of Service	Visit
✓	32583	7	Emergency Room	N	UB04 Point of Origin	Visit



#4 Create Outcome Cohort:AMI

Show entries

Filter records:

	Concept Id	Code	Concept Name	Standard Type	Vocabulary	Domain
✓	8717	21	Inpatient Hospital	S	CMS Place of Service	Visit
✓	38004311	315D00000X	Inpatient Hospice	S	NUCC	Visit
✓	8971	51	Inpatient Psychiatric Facility	S	CMS Place of Service	Visit
✓	8920	61	Comprehensive Inpatient Rehabilitation Facility	S	CMS Place of Service	Visit
✓	262	ERIP	Emergency Room and Inpatient Visit	S	Visit	Visit
✓	9201	IP	Inpatient Visit	S	Visit	Visit
✓	581384	OMOP4822037	Inpatient Nursery	S	CMS Place of Service	Visit
✓	581383	OMOP4822038	Inpatient Cardiac Care Facility	S	CMS Place of Service	Visit
✓	581379	OMOP4822042	Inpatient Critical Care Facility	S	CMS Place of Service	Visit
✓	32211	02	Discharged/transferred to other short term general hospital for inpatient care.	N	UB04 Pt dis status	Visit



#4 Create Outcome Cohort:AMI

Cohort Exit

Event Persistence:

Event will persist until:

Fixed Duration Persistence:

The event end date is derived from adding a number of days to the event start date. This ensures that all cohort episodes will have the same fixed duration (subject to further constraints). The cohort may have varying cohort duration times due to the varying length of stay). This event persistence assures that the cohort end date is consistent across all episodes.

- Event date to offset from:
- Number of days offset: days

Censoring Events:

Exit Cohort based on the following criteria:

No censoring events selected.