

APAC Community Call

Global Symposium Recap Training Session #6

November 16, 2023





- OHDSI News
- Global Symposium Recap
 - From Singapore by Mukkesh Kumar and Cindy Ho
 - From Taiwan by Septi Melisa and Dian Tri Wiyanti
- Training Session #6 by Taiwan
 - Population level estimation study: Risk of cardiovascular disease among female cancer patients using hormone therapies by Whitney Burton, Rachel Quynh Nguyen, Maz Solie
 - Patient level prediction study: Prediction of dementia among patients with chronic disease by Thanh-Phuc Phan, Daniel Chris, Sunny Lin



OHDSI News

- 2023 Our Journey Annual Report is now available online at <u>https://www.ohdsi.org/wp-content/uploads/2023/11/OHDSI-Book2023.pdf</u>
- Global Symposium's post-event page is now live at <u>https://www.ohdsi.org/ohdsi2023/</u>
- APAC made some significant contributions at the Global Symposium this year, including Titan Awards, multiple presentations at the Collaborator Showcase and a best contribution winner!
- Next month's community call has been moved to 1 week earlier,
 <u>December 14</u>, to avoid conflict with the community's year-end plans





A*STAR

DATA MANAGEMENT PLATFORM

Mukkesh Kumar, Cindy Ho

Singapore

16th November 2023

Mukkesh Kumar



Dr Mukkesh Kumar is the Head of Data Management Platform at A*STAR, leading the Multi-modal Data Management, Clinical Data Curation & Data Stewardship, Clinical Data Analytics & Reporting, Electronic Data Capture Administration and Healthcare Software Development teams. Dr Mukkesh Kumar is a PhD alumnus of the NUS Saw Swee Hock School of Public Health, he has developed a predictive care framework for diabetes & maternal health, combining coalitional game theory concepts with machine learning. The A*STAR Data Vault developed by Data Management Platform is bridging data standardization and accessibility for impactful research outcomes across the ecosystem. Working in close partnership with Singapore's Ministry of Health (MOH), Dr Mukkesh Kumar is developing the core OMOP data curation team at A*STAR to support MOH-TRUST Strategic Research Data Contributors.

Cindy Ho



Ms Cindy Ho is a Statistician at A*STAR, leading the data curation and data stewardship of population health studies. Ms Cindy Ho's software demonstration on 'GUSTO Data Vault: Laying the foundations for an open science system with OMOP Data Catalogue' has won the best community contribution award for the Open-Source Development category at the 2023 OHDSI Global Symposium. She has pioneered the OMOP data curation efforts for Growing Up in Singapore Towards healthy Outcomes (GUSTO) cohort study and is developing the integrated OMOP Data Catalogue module within the GUSTO Data Vault platform for OMOP-based research. Ms Cindy Ho graduated with a major in Statistics from the NUS Department of Statistics, with domain expertise in cardiometabolic diseases.

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Content

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Key Takeaways from the OHDSI 2023 Symposium

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Collaborator Showcase Posters & Software Demos



Common Data Model/Network Data Quality WG Meeting







Key Takeaways

- Mixture of talks, workshop and various workgroup discussions
 - (HowOften) Workshop: UI/UX
 - Workgroup: CDM/Network Data Quality and Vocabulary
- Emphasis on
 - Large scale collaboration
 - Open-source tools
 - Vocabulary building
 - Al
- Day 1 Closing
 - 'Escape Room' activity
 - OHDSI Got Talent





Collaborator Showcase: Posters & Software Demos





Full Demo Video Link: <u>https://www.ohdsi.org/wp-</u> <u>content/uploads/2023/10/407-CindyHo-</u> <u>FullVideo-Cindy-Ho.mp4</u> GUSTO OMOP Data Catalogue Link: https://gustodatavault.sg/omop/



GUSTO OMOP Data Catalogue won the Best Community Contribution Award (determined through the peer review evaluation by the OHDSI Scientific Review Committee and community voting, based on the quality and importance of contribution).

The global recognition by OHDSI helps in furthering A*STAR Data Vault's impact in the research ecosystems.

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Common Data Model/Network Data Quality WG Meeting

- OHDSI data network is an important initiative led by the OHDSI Central Coordinating Centre to mobilize coalition and forge collaborations across global research ecosystems.
- Database snapshot (querying concept IDs, display sample size of concept IDs, simple data visualisations) -> Important first step to explore the feasibility of a network study.
- The metadata of OMOP CDM databases would be hosted from AWS Cloud environment for access by OHDSI members.



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Vocabulary WG Meeting

- Community contribution to non-drug vocabularies there are templates for each use case (<u>https://github.com/OHDSI/Vocabulary-v5.0/wiki/Community-</u> <u>contribution-guidelines:-non%E2%80%90drug-vocabularies</u>)
- introducing new non-standard concept(s) to retain the granularity of research data (e.g. GUSTO Birth Cohort Study)
- promoting non-standard concepts to standard terms (e.g. standard mapping by Boston Birth Cohort Study, Shanghai Birth Cohort study)
- Community contribution to drug vocabularies (covered by Anna Ostropolets on 7th December 2023, OHDSI APAC Scientific Forum call).
- Future work by Vocabulary WG is to build a metadata which shows the uphill and downhill relationship of Concept IDs "Map to", "Map from".

#	Туре
T1	Adding new non- standard concept(s) to an existing vocabulary
T2	Adding new synonym(s) to an existing concept(s)
Т3	Adding a mapping to an existing concept
T4	Adding a new vocabulary as non- standard with mappings (full or partial) to a standard vocabulary
Т5	Modifying attributes of an existing concept(s)
Т6	Modifying mapping for an existing concept
Т7	Promoting non-standard concepts to standard









THANK YOU

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2023 OHDSI GLOBAL SYMPOSIUM

East Brunswick, New Jersey October 20-22, 2023







Schedule



OHDSI 2023 Symposium Oct. 20-22, 2023 Hilton East Brunswick Hotel & Executive Meeting Center



Agenda · Friday, Oct. 20

Time	Торіс
7:30 - 8:30 am East Brunswick Room + Grand Ballroom Foyer	Symposium Registration, Lite Breakfast Buffet, All-Day Exhibits * First-timers can meet for a quick orientation session at 7:45 am in Piscataway/Woodbridge (will conclude before the start of the first talk)
8:30 - 9:30 am Grand Ballroom	State of the Community OHDSI: Where have we been? Where are we going? George Hripcsak, Columbia Univ. Community Highlights: • OMOP CDM users and the OHDSI data network Clair Blacketer, Johnson & Johnson • OHDSI standardized vocabularies Alexander Davydov, Odysseus Data Services • OHDSI's open-source community Katy Sadowski, Boehringe Ingelheim • OHDSI Europe 2024 Peter Rijnbeek, Erasmus MC • OHDSI Asia-Pacific 2024 Mengling Feng, National Univ. of Singapore
9:30 - 10:30 am Grand Bellroom	OHDSI Community Networking Moderators: • Faaizah Arshad, Univ. of California-Los Angeles • Cynthia Sung, Duke-NUS Medical School
10:30 am - 12:00 pm Grand Bailroom	Plenary: Improving the reliability and scale of case validation Presenters: • Patrick Ryan, Johnson & Johnson, Columbia Univ. • Anna Ostropolets, Odysseus Data Services • Martiin Schuemie, Johnson & Johnson, Univ. of California-
	Los Angeles



State of the Community





Opening speech by George Hripcsak



Various leaders within OHDSI shared a presentation on the state of the community









Improving the reliability and scale of case validation



Panel Talks





Lesson learned from Network Studies



Workgroup





OHDSI Saturday Activities, 8:00am-12:00pm

Common Data Model/Network Data Quality | HADES | Health Equity | Industry Special Interest | Introduction to OHDSI Tutorial | Medical Imaging | Natural Language Processing | Oncology | Perinatal & Reproductive Health | Phenotype Evaluation

HowOften Large-Scale Characterization Workshop, Part 1, 1:00pm-5:00pm



Community and Collaboration (基) 小醫學大學







Make new friends and connections



Collaborator Showcase









Titan Awards







Interest Community Collaboration (Join The Journ



Additions Community Support Add





OHDSI Global Symposium

See you at the next year symposium







OHDSI Taiwan Study Presentation

Estimating Adverse Cardiovascular-Related Events

After Hormone Therapy Treatment

in Three Female Cancer Populations

Whitney Burton, Quynh Nguyen, Septi Melisa, Mohammad Solihuddin Muhtar, & Jason Hsu



Content

- Literature review
- Cohorts definition and Characterization
- Preliminary results



Literature review



Background for the Study

- In 2020, amongst female cancer patients:
 - **Breast cancer** ranked first for incidence and mortality rates with 24.5% of cases and 15.5% of deaths
 - Uterine cancer ranked sixth making up 4.5% of cases and 2.9% of deaths; and
 - **Ovarian cancer** ranked eight by accounting for 3.7 of cases and 4.7% of deaths.
- Hormone therapy represses the growth of cancer cells by disrupting or hinder the body's natural hormone production
 - It can be used as standalone treatment or in partnership with other strategies.
 - Aromatase Inhibitors (AIs) and Selective Estrogen Receptor Modulators (SERMs) classes are used for pre- and postmenopausal oncology patients



Background for the Study

- Als work by inhibiting the action of the enzyme aromatase, which converts androgens into estrogens (inhibiting estrogen production)
- SERMs work by binding to estrogen receptors in the tumor, but have estrogen-like activities in other tissues
- Some evidence suggests that tamoxifen (a SERMs) may reduce cholesterol level [1,2]
- Impact of AIs on cholesterol level remains unclear [3,4]
- 1. Circulation. 2005, 112, 3018–3024
- 2. Am. J. Cardiol. 1995, 76, 1072-1073
- 3. Lancet Oncol. 2006; 7:633–643.

Articles

(E) Churck for upd

Received: 21 November 2021. Review: 13 December 2021. Accepted: 23 December 2021

DOI: 10.1111/jcpt.13598 **REVIEW ARTICLE**

Detail Parmary at Temperty WILEY

Risk of cardiovascular disease in breast cancer patients receiving aromatase inhibitors vs. tamoxifen: A systematic review and meta-analysis

Qiuyan Yu Bsc Nursing | Yueping Xu Bsc Nursing | Enguang Yu Bsc Nursing | Zhufeng Zheng Bsc Nursing

etmant of Norsing, Sasing Hospital of Traditional Chinese Medicine, Japaine

Qicyan Yu. Department of numing, Jasin Hugital of Ynultiseal Chinese Medicine 1501 Zhongshan East Road, Janing. Zhajiang Province 314001, China. Email yan7082@163.com

Funding information This study was supported by Zhejang Medicine and Health Science and Technology Plan (No. 2025PV079)

Abstract What is known and objective: Breast cancer is one of the leading causes of morbidity and mortality in women worldwide. In order to reduce the risks of its recurrence, endocrine therapies, such as tamoxifen and aromatase inhibitors are commonly adninistered. Despite having a similar efficacy in preventing breast cancer recurrence, these drugs differ in terms of instigating cardiovascular morbidities. Recent randomized controlled trials and cohort studies provide inconclusive evidence of the cardiovascular risks associated with the administration of these endocrine therapies. This present review and meta-analysis evaluates the comparative cardiovascular adverse. event outcomes in breast cancer patients receiving tamoxifen and aromatase inhibitors. To evaluate the comparative cardiovascular adverse outcomes, such as venous thromboembolism, heart failure, angina, myocardial infarction and stroke in patients

with breast cancer receiving tamoxifen and aromatase inhibitors. Methods: A systematic search of the academic literature was performed according to the PRISMA guidelines across five databases, including Web of Science, EMBASE, CENTRAL, Scopus, and MEDLINE. A random-effect meta-analysis was conducted to compare the cardiovascular adverse events (i.e. venous thromboembolism, heart failure, angina, myocardial infarction, strokel in breast cancer patients treated with tamoxifen and aromatase inhibitors.

Results and discussion: From 993 studies, 20 eligible studies were identified, with 174,142 female breast cancer patients (mean age: 67.4 ± 3.8 years). A meta-analysis revealed insignificantly (p > 0.05) higher risks of venous thromboembolism (Odds ratio, 95% CI: 1.70, 0.91-3.18) in patients treated with tamoxifen as compared to aromatase inhibitors. We also observed insignificantly higher risks of stroke (0.93, 0.45-1.91), angina (0.77, 0.12-4.59), myocardial infarction (0.74, 0.30-1.79), and heart failure (0.81, 0.22-2.91) in patients receiving aromatase inhibitors as compared to tamoxifen.

What is new and conclusions: The study provides evidence regarding the comparative cantiovascular adverse outcomes between breast cancer patients consumine

Glount To and Ranging Katzentrikated equally to this work

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Research

JAMA Oncology | Original Investigation

Cardiovascular Disease After Aromatase Inhibitor Use

Reina Haque, PhD; Jiaxiao Shi, PhD; Joanne E. Schottinger, MD; Joanie Chung, MPH; Chantal Avila, MA; Britta Amundsen, MA; Xiaoqing Xu, PharmD; Ana Barac, MD; Rowan T. Chlebowski, MD

+ Supplemental content at jamaoncology.com

IMPORTANCE Cardiovascular disease (CVD) is an important cause of death in older patients with breast cancer. However, limited information exists on the long-term effect of aromatase inhibitor (AI) use on CVD risk in breast cancer survivors. To this point, no other population-based studies have been able to adjust for CVD risk factors or cardiovascular medications.

OBJECTIVE To determine the long-term influence of adjuvant endocrine therapies on CVD in a cohort of postmenopausal breast cancer survivors in analyses that accounted for major CVD risk factors, medication use, chemotherapy, and radiotherapy.

DESIGN, SETTING, AND PARTICIPANTS A retrospective cohort of postmenopausal women with breast cancer diagnosed from January 1, 1991, to December 31, 2010, and followed up through December 31, 2011 (maximum, 21 years [72 886 person-years]), was evaluated using records from a managed care organization with nearly 20 community hospitals in California. A total of 13 273 postmenopausal women with hormone receptor-positive breast cancer without prior CVD were included. Cardiovascular disease incidence was compared across endocrine therapy categories. Information on demographics, comorbidity, medication, use, and CVD risk was captured from electronic health records. Multivariate Cox proportional hazards models using time-dependent endocrine drug use variables and propensity scores were conducted. Data analysis was conducted from September 15, 2014, to February 1, 2016.

EXPOSURES Women were grouped by endocrine therapy status (tamoxifen citrate only, AI only, both, or neither).

MAIN OUTCOMES AND MEASURES Person-year rates of CVD for each therapy group.

RESULTS During 72 886 person-years in 13 273 women (mean [SD] age, 66.8 [8.1] years) with follow-up through 2011, we observed 3711 CVD events. In multivariable analyses (reported as hazard ratio [95% CI]), Al-only users had a similar risk of cardiac ischemia (myocardial infarction and angina) (adjusted, 0.97 [0.78-1.22]) and stroke (adjusted, 0.97 [0.70-1.33]) as tamoxifen-only users (reference). However, we found an increased risk of other CVD (dysrhythmia, valvular dysfunction, and pericarditis) (adjusted, 1.29 [1.11-1.50]) in women who used Als only or sequentially after tamoxifen (1.26 [1.09-1.45]) vs tamoxifen (reference) as well nonhormone users (1.18 [1.02-1.35]).

CONCLUSIONS AND RELEVANCE The risk of the most serious cardiovascular events (cardiac

European Journal of Cancer 68 (2016) 11-21



Original Research

The risk of myocardial infarction with aromatase inhibitors relative to tamoxifen in post-menopausal women with early stage breast cancer

Husam Abdel-Qadir ^{a,b,c,d}, Eitan Amir ^{a,c,e}, Hadas D. Fischer ^b, Longdi Fu^b, Peter C, Austin^{b,c}, Paula J, Harvey^{a,d,f,g}, Paula A. Rochon ^{a,b,c,d,f}, Douglas S. Lee ^{a,b,c,g,1}, Geoffrey M. Anderson b,c,f,*,1

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Received 16 March 2016; received in revised form 22 June 2016; accepted 25 August 2016 Available online 30 September 2016

KEYWORDS Hormonal therapy; Endocrine therapy; Anastrozole: Author Affiliations: Department Letrozole: Exemestane; Permanente Southern California Coronary artery Pasadena, California (Haque, Shi, disease;

Research & Evaluation, Kaiser

Schottinger, Chung, Avila, Amunds

to tamoxifen in post-menopausal women with breast cancer. This risk has not been wellquantified outside of clinical trials. Methods: Observational population-based cohort study of women aged >55 years diagnosed with stage I-III breast cancer between 2005 and 2010. Women treated with AIs or tamoxifen were followed to March 2012. The primary outcome was hospitalisation for myocardial infarction (MI). Cause-specific hazards were compared using tamoxifen as the reference group. Inverse probability of treatment weighting using the propensity score was used to

Abstract Background: Aromatase inhibitors (AIs) may increase cardiovascular risk relative

Some meta-analyses raised concerns regarding negative impact of AIs on CVD

Observational studies showed conflict results on each cardiovascular disease, for example: myocardial infarction.





Research Questions

- For cardiovascular safety, can we estimate the comparative risk of aromatase inhibitor to estrogen receptor drug classes for women with breast, urine, and ovarian cancers?
 - Sub-Question A: Does a history of only aromatase inhibitor drugs place women at greater risk for cardiovascular events?
 - Sub-Question B: Does a history of only estrogen receptor drugs place women at greater risk for cardiovascular events?
 - Sub-question C: Does exposure to both hormone classes place women at greater risk for adverse cardiovascular events compared to those with one hormone treatment history?



Cohort definition & characterization



Concept Sets

- The following were used to create the cohort definitions:
 - ICD-9,
 - ICD-10,
 - ICD-O, and
 - The World Health Organization's Collaborating Centers for Drug Statistics
 Methodology Anatomical Therapeutic Chemical Classification System



Creating Definitions

Conditions

- Diagnosis of:
 - Breast Cancer
 - Uterine Cancer
 - Ovarian Cancer
 - Female Cancer (all three)
- Age inclusion, greater than or equal to 20
- Earliest event per person

Drug Exposure

- SERM
 - Tamoxifen
- Al
 - Letrozole
 - Exemestane
 - Anastrozole
- Drug exposure inclusion criteria



Ensuring mono therapy

cardio_cancer_woman										
Cohort Entry Events		0								
Events having any of the following criter	ria:	+ Add Initial Event								
a condition occurrence of cancer_female_uterus -										
X for the first time in the person's history										
with continuous observation of at least 0 💌 days before and 0 💌 days after event index date										
Limit initial events to: earliest event V	per person.									
Inclusion Criteria		U								
New inclusion criteria	no aromatase inhibitor exposure	Copy Delete								
1. age	enter an inclusion rule description									
 use hormone therapy no aromatase inhibitor exposure 	having $\boxed{\text{all}}$ of the following criteria:	+ Add criteria to group								
	with exactly V 0 U using all occurrences of:	Delete Criteria								
	a drug exposure of aromatase inhibitors TMU 👻	+ Add attribute→								
	where event starts between									
	All v days Before v and All v days After v index start date add additional constraint									
	The index date refers to the event from the Cohort Entry criteria.									
	restrict to the same visit occurrence									
	allow events from outside observation period									
Limit qualifying events to: all events	♥ per person.									



Outcomes - Defining adverse cardiovascular events

Disease Type	ICD-9	ICD-10	Concept ID	Domain
Myocardial infarction	410, 412	121, 122, 123, 125.2	4329847	Condition
Arrhythmia	427	147, 148, 149	44784217	Condition
Cardiomyopathy	425.4, 414.8, 674.52, 674.54, 429.3	142, 142.9, 125.5, 090.3, 151.7	321319	Condition
Coronary artery disease	411, 414.[02348]	124.0, 125.1, 125.8	317576 40641917	Condition
Congestive Heart Failure	428.0, 428.2[0-3], 438.3[0-3], 428.4[0-4]	I50.2, I50.21, I50.22, I50.23 I50.3, I50.30 I50.31, I50.32, I50.33 I50.4, I50.41, I50.42, I50.43	*	Condition
Heart Valve Disease	421.1, 424.91	139 [0-8], Q23.0, Q23.4, Q24.4, Q25.3, Q22.4	4343040	Condition
Pericardial effusion	423.9, 424.91, 746.3, 746.6, 746.7,	131.3	4108814	Condition
Bradycardia	427.81, 427.89	R00.1	4169095	Condition
Constrictive Pericarditis	423.2	131.1	312334	Condition
Stroke	433, 434, 435	163, 165, 166	381591	Condition
Blood clots in legs and lung	453.[45][012], 453.[6789], 444.[29], 445.0	126, 127.82, T80.0 182. 40	440417-lung 4046884-leg 4126721-vagina 763972-stool 4091191 45768439 43531681	Condition



Preliminary results

Outcome: Female cancer Arrhythmia

Analysis	\$ D	ata source	HR	÷ 1	LB		UB		Р		Cal.HR		Cal.LB		Cal.UB		Cal.P	
сох	T	MU_CRD	1.21	(0.66		2.25		0.54		NA		NA		NA		NA	
Showing 1	Showing 1 to 1 of 1 entries								Previous	1	Next							
Power	Attrition	Population characterist	ics	Proper	nsity m	odel	P	ropen	sity sc	ores	Cov	ariate bal	ance	Systema	tic error	Kaplan-	Meier	

Table 1a. Number of subjects, follow-up time (in years), number of outcome events, and event incidence rate (IR) per 1,000 patient years (PY) in the target (*Female_cancer_Breast_Cancer_with_Aromatase_Inhibitor*) and comparator (*Female_cancer_Breast_Cancer_with_Estrogen_Receptor*) group after propensity score adjustment, as well as the minimum detectable relative risk (MDRR). Note that the IR does not account for any stratification.

Target subjects	Comparator subjects	Target years	Comparator years	Target events	Comparator events	Target IR (per 1,000 PY)	Comparator IR (per 1,000 PY)	MDRR
504	504	2,144	2,262	39	35	18.19	15.47	1.92

Table 1b. Time (days) at risk distribution expressed as minimum (min), 25th percentile (P25), median, 75th percentile (P75), and maximum (max) in the target (*Female_cancer_Breast_Cancer_with_Aromatase_Inhibitor*) and comparator (*Female_cancer_Breast_Cancer_with_Estrogen_Receptor*) cohort after propensity score adjustment.

Cohort	Min	P10	P25	Median	P75	P90	Max
Target	1	203	586	1,346	2,798	3,215	6,189
Comparator	1	164	550	1,472	3,099	3,351	6,118



Outcome: Female cancer Arrhythmia



Figure 2. Preference score distribution. The preference score is a transformation of the propensity score that adjusts for differences in the sizes of the two treatment groups. A higher overlap indicates subjects in the two groups were more similar in terms of their predicted probability of receiving one treatment over the other.

Outcome: Female cancer Arrhythmia



Figure 5. Kaplan Meier plot, showing survival as a function of time. This plot is adjusted using the propensity score: The target curve

(Female_cancer_Breast_Cancer_with_Aromatase_Inhibitor) shows the actual observed survival. The comparator curve

(*Female_cancer_Breast_Cancer_with_Estrogen_Receptor*) applies reweighting to approximate the counterfactual of what the target survival would look like had the target cohort been exposed to the comparator instead. The shaded area denotes the 95 percent confidence interval.



Outcome: Female Cancer Coronary Arteriosclerosis

Analysis	♦ D	ata source	HR	🕴 LB	\$	UB		Р		Cal.HR	$\stackrel{\mathbb{A}}{\nabla}$	Cal.LB		Cal.UB	\$	Cal.P	
сох	т	MU_CRD	1.50	0.5	54	4.47		0.45		NA		NA		NA		NA	
Showing 1 t	o 1 of 1 entr	ies													Previous	1	Next
Power	Attrition	Population characteri	stics	Propensi	ty model	I P	ropen	sity sc	ores	Cova	riate bala	ance	Systemat	ic error	Kaplan-I	Meier	

Table 1a. Number of subjects, follow-up time (in years), number of outcome events, and event incidence rate (IR) per 1,000 patient years (PY) in the target (*Female_cancer_Breast_Cancer_with_Aromatase_Inhibitor*) and comparator (*Female_cancer_Breast_Cancer_with_Estrogen_Receptor*) group after propensity score adjustment, as well as the minimum detectable relative risk (MDRR). Note that the IR does not account for any stratification.

Target	Comparator	Target	Comparator	Target	Comparator	Target IR (per	Comparator IR (per	MDRR
subjects	subjects	years	years	events	events	1,000 PY)	1,000 PY)	
513	513	2,275	2,378	13	15	5.71	6.31	2.88

Table 1b. Time (days) at risk distribution expressed as minimum (min), 25th percentile (P25), median, 75th percentile (P75), and maximum (max) in the target (*Female_cancer_Breast_Cancer_with_Aromatase_Inhibitor*) and comparator (*Female_cancer_Breast_Cancer_with_Estrogen_Receptor*) cohort after propensity score adjustment.

Cohort	Min	P10	P25	Median	P75	P90	Max
Target	15	229	669	1,439	2,869	3,229	6,203
Comparator	5	216	592	1,551	3,098	3,354	6,118



Outcome: Female cancer Coronary arteriosclerosis



Figure 2. Preference score distribution. The preference score is a transformation of the propensity score that adjusts for differences in the sizes of the two treatment groups. A higher overlap indicates subjects in the two groups were more similar in terms of their predicted probability of receiving one treatment over the other.

Outcome: Female cancer Coronary arteriosclerosis



Figure 5. Kaplan Meier plot, showing survival as a function of time. This plot is adjusted using the propensity score: The target curve (*Female_cancer_Breast_Cancer_with_Aromatase_Inhibitor*) shows the actual observed survival. The comparator curve (*Female_cancer_Breast_Cancer_with_Estrogen_Receptor*) applies reweighting to approximate the counterfactual of what the target survival would look like had the target cohort been exposed to the comparator instead. The shaded area denotes the 95 percent confidence interval.





OHDSI APAC Community training

Patient-level prediction: Dementia risk among

patients with chronic disease

PHAN THANH-PHUC, SUNNY LIN, DANIEL CHRIS, ALEX NGUYEN, JASON C. HSU



Content

- Literature review
- Cohorts definition and Characterization
- Preliminary results



Literature review





cross-sectional study

Prevalence, risk factors, and management of dementia and mild cognitive impairment in adults aged 60 years or older in China: a cross-sectional study

Lancet Public Health 2020; 5: e661–71



Meta-Analysis

Potential for primary prevention of Alzheimer's disease: an analysis of population-based data

Lancet Neurol 2014; 13: 788-94



Dementia prevention, intervention, and care: 2020 report of the Lancet Commission

Lancet 2020; 396: 413-46



THE LANCET Public Health

cross-sectional study

Prevalence, risk factors, and management of dementia and mild cognitive impairment in adults aged 60 years or older in China: a cross-sectional study

	Dementia (n=2766)		MCI (n=7125)	
	OR (95% CI)	p value	OR (95% CI)	p value
Age, years				
≥90	6.60 (5.24-8.32)	<0.0001	4.70 (3.77-5.87)	<0.0001
80-89	3.90 (3.45-4.40)	<0.0001	2.54 (2.33-2.76)	<0.0001
70-79	2.69 (2.43-2.98)	<0.0001	1.89 (1.77-2.00)	<0.0001
60-69	1 (ref)		1 (ref)	
Sex				
Female	1.43 (1.31-1.56)	<0.0001	1.51 (1.43-1.59)	<0.0001
Male	1 (ref)		1 (ref)	
Parental history of dement	ia			
Yes	7.20 (5.68-9.12)	<0.0001	1.91 (1.48-2.46)	<0.0001
No	1 (ref)		1 (ref)	
Residence location				
Rural	1.16 (1.06-1.27)	0.0010	1.45 (1.38-1.54)	<0.0001
Urban	1 (ref)		1 (ref)	
Education level, years				
<1	1.55 (1.38-1.73)	<0.0001	3.48 (3.25-3.73)	<0.0001
1-6	1.17 (1.06-1.29)	0.0021	1.48 (1.39-1.58)	<0.0001
>6	1 (ref)	**	1 (ref)	**
Marital status				
Widow	2.59 (2.30-2.90)	<0.0001	1.58 (1.44-1.73)	<0.0001
Divorced or living alone	2.66 (2.29-3.10)	<0.0001	1.74 (1.56-1.95)	<0.0001
Married	1 (ref)		1 (ref)	

Smoker				
Yes	1.85 (1.67-2.04)	<0.0001	1.27 (1.19–1.36)	<0.0001
No	1 (ref)		1 (ref)	
Hypertension				
Yes	1.86 (1.70-2.03)	<0.0001	1.62 (1.54–1.71)	<0.0001
No	1 (ref)		1 (ref)	
Hyperlipidaemia				
Yes	1.87 (1.71–2.05)	<0.0001	1·29 (1·21–1·37)	<0.0001
No	1 (ref)		1 (ref)	
Diabetes				
Yes	2.14 (1.96-2.34)	<0.0001	1.44 (1.35–1.53)	<0.0001
No	1 (ref)		1 (ref)	
Heart disease				
Yes	1.98 (1.73–2.26)	<0.0001	1.17 (1.06–1.30)	0.0023
No	1 (ref)		1 (ref)	
Cerebrovascular disease				
Yes	5.44 (4.95-5.97)	<0.0001	1.49 (1.36-1.62)	<0.0001
No	1 (ref)		1 (ref)	
ACI=mild cognitive impairme	ent. OR=odds ratio.			

Lancet Public Health 2020; 5: e661–71

THE LANCET Public Health

cross-sectional study

Prevalence, risk factors, and management of dementia and mild cognitive impairment in adults aged 60 years or older in China: a cross-sectional study

	Dementia (n=276	6)	MCI (n=7125)		Smoker				
						1.85 (1.67-2.04)	<0.0001	1.27 (1.19–1.36)	<0.0001
Age, year					No	1 (ref)		1 (ref)	
≥90					Hypertension				
80-		Hyper	tansiar	`	Yes	1.86 (1.70–2.03)	<0.0001	62 (1.54–1.71)	<0.0001
70-1		турсі	1013101	· .	No	1 (ref)		(ref)	
60-		Hyper	lipidem	nia 📕	Hyperlipidaemia				
Fer Ris	k	1990	npraom		Yes	1.87 (1.71-2.05)	<0.0001	29 (1·21-1·37)	<0.0001
Mai		Diabe	tes		No	1 (ref)		(ref)	
Parent Fac	ctor				Diabetes				
(Medicatio	n modifiable)	Heart	disease	Э 📕	Yes	2.14 (1.96-2.34)	<0.0001	44 (1·35–1·53)	<0.0001
Reside					No	1 (ref)		(ref)	
Run		CVD			Heart disease				
Urba					Yes	1.98 (1.73-2.26)	<0.0001	17 (1.06–1.30)	0.0023
	4 55 (4 20 4 72)				No	1 (ref)		(rer)	
<1	1.17 (1.06-1.29)	0.0021	3.48 (3.25-3.73)	<0.0001	Cerebrovascular disease			40 (1 26 1 62)	-0.0001
>6	1 (ref)		1 (ref)		Yes	5.44 (4.95-5.97)	<0.0001	49 (1·50-1·02)	20.0001
Marital status					No	1 (ref)	**	licit	
Widow	2.59 (2.30-2.90)	<0.0001	1.58 (1.44-1.73)	<0.0001	mei-mille cognicive impairmen	n. on-ouus latio.			
Divorced or living alon Married	1 (ref)	<0.0001	1.74 (1.56-1.95)	<0.0001	Table 3: Adjusted ORs for de	ementia and MCI			



THE LANCET Neurology

Meta-Analysis

Potential for primary prevention of Alzheimer's disease: an analysis of population-based data

	Relative risk (95% CI)*	Communality (%)†
Diabetes mellitus	1.46 (1.20–1.77)	50.9%
Midlife hypertension	1.61 (1.16–2.24)	65.0%
Midlife obesity	1.60 (1.34–1.92)	43·7%
Physical inactivity	1.82 (1.19–2.78)	49.0%
Depression	1.65 (1.42–1.92)	37.4%
Smoking	1.59 (1.15–2.20)	58·1%
Low educational attainment	1.59 (1.35-1.86)	45.6%

*Sources are provided in the appendix. †The proportion of the variance in each risk factor shared with the other risk factors, estimated using the Health Survey for England 2006.¹⁷

Table 1: Relative risks for Alzheimer's disease and shared variance between risk factors

Panel 1: Definitions used for each of the risk factors

Diabetes mellitus

Adult prevalence of diagnosed diabetes mellitus between the ages of 20 years and 79 years

Midlife hypertension

Adult midlife prevalence of hypertension between the ages of 35 years and 64 years

Midlife obesity

Adult midlife prevalence of body-mass index greater than 30 kg/m² between the ages of 35 years and 64 years

Physical inactivity

Proportion of adults who do not do either 20 min of vigorous activity on 3 or more days or 30 min of moderate activity on 5 or more days per week

Depression

Lifetime prevalence of major depressive disorder using Diagnostic and Statistical Manual of Mental Disorders or International Classification of Diseases criteria

THE LANCET Neurology

Potential for primary prevention of Alzheimer's disease: an analysis of population-based data

Meta-Analysis

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Risk Factor

(Medication modifiable)

Diabetes Hypertension Depression

*Sources are provided in the appendix. †The proportion of the variance in each risk factor shared with the other risk factors, estimated using the Health Survey for England 2006.¹⁷

Table 1: Relative risks for Alzheimer's disease and shared variance between risk factors





Review

Dementia prevention, intervention, and care: 2020 report of the Lancet Commission

	Relative risk for dementia (95% CI)	Risk factor prevalence	Communality	Unweighted PAF	Weighted PAF*
Early life (<45 years)					
Less education	1.6 (1.3-2.0)	40-0%	61.2%	19.4%	7.1%
Midlife (age 45-65 years)				
Hearing loss	1.9 (1.4-2.7)	31.7%	45.6%	22.2%	8.2%
ТВІ	1.8 (1.5-2.2)	12-1%	55.2%	9.2%	3.4%
Hypertension	1.6 (1.2-2.2)	8-9%	68.3%	5.1%	1.9%
Alcohol (>21 units/week)	1.2 (1.1-1.3)	11-8%	73.3%	2.1%	0-8%
Obesity (body-mass index ≥30)	1.6 (1.3–1.9)	3.4%	58.5%	2.0%	0.7%
Later life (age >65 years)					
Smoking	1.6 (1.2-2.2)	27.4%	62.3%	14.1%	5.2%
Depression	1.9 (1.6-2.3)	13.2%	69.8%	10.6%	3.9%
Social isolation	1.6 (1.3-1.9)	17.7%	55.2%	9.6%	3.5%
Physical inactivity	1.4 (1.2-1.7)	11-0%	28.1%	4.2%	1.6%
Diabetes	1.5 (1.3-1.8)	6-4%	71.4%	3.1%	1.1%
Air pollution	1.1 (1.1-1.1)	75-0%	13.3%	6.3%	2.3%

Data are relative risk (95% CI) or %. Overall weighted PAF=39.7%. PAF=population attributable fraction. TBI=traumatic brain injury. *Weighted PAF is the relative contribution of each risk factor to the overall PAF when adjusted for communality.

Table 1: PAF for 12 dementia risk factors

THE LANCET

Dementia prevention, intervention, and care: 2020 report of the Lancet Commission

Review

	Relative risk for dementia (95% CI)	Risk factor prevalence	Communality	Unweighted PAF	Weighted PAF*
Early life (<45 years)					
Less education	1.6 (1.3-2.0)	40.0%	61.2%	19.4%	7.1%
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ТВІ	1.8 (1.5-2.2)	12-1%	55.2%	9.2%	3-4%
Hypertension	1.6 (1.2-2.2)	8.9%	68.3%	5.1%	1.9%
Alcohol (>21 units/week)	1.2 (1.1–1.3)	11.8%	73.3%	2.1%	0-8%
Obesity (body-mass index ≥30)	1.6 (1.3-1.9)	3.4%	58.5%	2.0%	0.7%
Later life (age >65 years)					
Smoking	1.6 (1.2-2.2)	27.4%	62.3%	14.1%	5.2%
Depression	1.9 (1.6–2.3)	13.2%	69.8%	10.6%	3.9%
Social isolation	1.6 (1.3-1.9)	17.7%	55.2%	9-6%	3.5%
Physical inactivity	1.4 (1.2-1.7)	11.0%	28.1%	4.2%	1.6%
Diabetes	1.5 (1.3–1.8)	6.4%	71.4%	3.1%	1.1%
Air pollution	1.1 (1.1-1.1)	75-0%	13.3%	6.3%	2.3%

Data are relative risk (95% CI) or %. Overall weighted PAF=39.7%. PAF=population attributable fraction. TBI=traumatic brain injury. *Weighted PAF is the relative contribution of each risk factor to the overall PAF when adjusted for communality.

Risk Factor Medication modifiable

Table 1: PAF for 12 dementia risk factors







Cohort definition & characterization



Cohort Entry Events
Events having any of the following criteria:
a condition occurrence of Dementia -
with continuous observation of at least $\boxed{0 extbf{v}}$ days before and $\boxed{0 extbf{v}}$ days after event index date Limit initial events to: earliest event $ extbf{v}$ per person.
with the following event criteria: Add attribute Xwith age Greater or Equal To 18
and with at least 1 ▼ using all occurrences of: a drug exposure of Dementia Medication
where event starts between ○ ▼ days Before ▼ and All ▼ days After ▼ index start date add additional constraint The index date refers to the event from the Cohort Entry criteria. □ restrict to the same visit occurrence □ allow events from outside observation period

Concept Name	▼ Domain	Standard Concept Caption	Exclude	Descendants
Senility	Condition	Standard	\checkmark	\sim
Senile degeneration of brain	Condition	Standard	\sim	 ✓
Senile and presenile organic psychotic conditions	Condition	Standard	~	
Postconcussion syndrome	Condition	Standard	~	 ✓
General paresis - neurosyphilis	Condition	Standard	~	
Frontotemporal dementia	Condition	Standard	\sim	 ✓
Drug-induced dementia	Condition	Standard	~	
Diffuse Lewy body disease	Condition	Standard	\checkmark	\sim
Dementia following injury caused by exposure to ionizing radiation	Condition	Standard	~	 ✓
Dementia caused by volatile inhalant	Condition	Standard	~	✓
Dementia caused by toxin	Condition	Standard	~	
Dementia caused by heavy metal exposure	Condition	Standard	~	\checkmark
Dementia	Condition	Standard	\checkmark	 ✓
Age-related cognitive decline	Condition	Standard	\sim	~

Concept Name
rivastigmine
memantine
Ginko biloba leaf oil
galantamine
donepezil

Generation

□ Show only sources with results

Show 10 🗙 entries

Source Name	Generation Status	People 🝦	Records 🔶
TMU CRD	COMPLETE	6,492	6,492



Cohort Entry Events

Events having any of the following criteria:



Inclusion Criteria

New inclusion criteria

1. No Prior Type 1 DM

2. No Secondary DM

Concept Name	🔻 Domain 🛛 🔶	Standard Concept Caption	Exclude	Descendants
Type 2 diabetes mellitus without complication	Condition	Standard	\checkmark	 ✓
Type 2 diabetes mellitus	Condition	Standard	\checkmark	✓
Type 2 diabetes mellitus	Condition	Standard	\checkmark	~
Peripheral circulatory disorder due to type 2 diabetes mellitus	Condition	Standard	\checkmark	~
Ketoacidosis due to type 2 diabetes mellitus	Condition	Standard	\checkmark	~
Disorder of nervous system due to type 2 diabetes mellitus	Condition	Standard	\checkmark	~
Disorder of kidney due to diabetes mellitus	Condition	Standard	~	~
Disorder of eye due to diabetes mellitus	Condition	Standard	~	~
Disorder due to type 2 diabetes mellitus	Condition	Standard	\checkmark	~
Coma due to diabetes mellitus	Condition	Standard	~	~

Generation

Show only sources with results

Show 10 🗙 entries

Source Name	Generation Status	People 🔶	Records 🔶
TMU CRD	COMPLETE	91,394	91,394



Hypertension

Cohort Entry Events

Events having any of the following criteria:





Essential hypertension Condition Standard	Concept Name	v Domain	Standard Concept Caption	Exclude	Descendants
	Essential hypertension	Condition	Standard	~	 ✓

Concept Name	🔻 Domain	Standard Concept Caption	Exclude	Descendants
trichlormethiazide	Drug	Standard	~	\checkmark
SELECTIVE CALCIUM CHANNEL BLOCKERS WITH MAINLY VASCULAR EFFECTS	Drug	Classification	~	\checkmark
hydrochlorothiazide	Drug	Standard	~	\checkmark
Angiotensin II receptor blockers (ARBs), plain	Drug	Classification	~	\checkmark
ACE INHIBITORS, PLAIN	Drug	Classification	~	\checkmark

Generation

Show only sources with results

ihow 10 👻 entries											
Source Name	Generation Status	People 🔶	Records 🔶								
TMU CRD	COMPLETE	153,395	153,395								



Cohort Entry Events

Events having any of the following criteria:



Concept Name	Domain 🔶	Standard Concept Caption	Exclude	Descendants
Severe depression	Condition	Standard	~	 Image: A start of the start of
Schizoaffective disorder, depressive type	Condition	Standard	\checkmark	✓
Recurrent depression	Condition	Standard	\sim	✓
Reactive depressive psychosis	Condition	Standard	\checkmark	✓
Psychosis and severe depression co-occurrent and due to bipolar affective disorder	Condition	Standard	✓	\checkmark
Moderate depression	Condition	Standard	\sim	✓
Mild depression	Condition	Standard	\checkmark	✓
Major depressive disorder	Condition	Standard	\sim	✓
Depressive disorder in remission	Condition	Standard	\sim	✓
Chronic depressive personality disorder	Condition	Standard	\sim	✓
Bipolar affective disorder, currently depressed, in full remission	Condition	Standard	✓	~
Atypical depressive disorder	Condition	Standard	~	✓

Concept Name	Domain 🔶	Standard Concept Caption	÷ Exclude	Descendants
Selective serotonin reuptake inhibitors	Drug	Classification	~	✓
Other antidepressants	Drug	Classification	~	✓
Non-selective monoamine reuptake inhibitors	Drug	Classification	~	✓
Monoamine oxidase inhibitors, non-selective	Drug	Classification	~	 ✓
Monoamine oxidase A inhibitors	Drug	Classification	~	 ✓

Generation

Show only sources with results

Show 10 🗙 entries

Source Name	Generation Status	People 🔶	Records 🔶
TMU CRD	COMPLETE	31,963	31,963



Preliminary results

R packages execution – Patient level prediction

()	🚛 🔚 Source on Save 🔍 🎢 🖌 📕	1	🔹 Run	••	t	† I	Source 👻
9	library(PatientLevelPrediction)						
10							
11							
12	current_script <- rstudioapi::getSourceEditorContext()\$path						
13	<pre>setwd(dirname(current_script))</pre>						
14							
15	<pre># PatientLevelPrediction::configurePython(envname='test', envtype='python')</pre>						
16	<pre>#PatientLevelPrediction::setPythonEnvironment(envname = 'test', envtype='python')</pre>						
17	reticulate: use_virtualenv('test')						
18	# reticulate::use_condeenv('base')						
20							
20	mains - list						
22	dm2d = 1 ist(
23	analysisId = "Type 2 DM to Dementia".						
24	analysisDesc = "DM2D prediction".						
25	outcomeId = 243, # outcome cohort ID						
26	targetId = 242, # population cohort ID						
27	<pre>cohortTable = "dm2dCohort"</pre>						
28),						
29	ht2d = list(
30	<pre>analysisId = "Hyypertension to Dementia",</pre>						
31	analysisDesc = "HT2D prediction",						
32	outcomeId = 243, # outcome cohort ID						
33	targetId = 262, # population cohort ID						
34	cohortTable = "ht2dCohort"						
35							
36	aepza = List(
30	analysisia = Depression to Dementia,						
20 20	$automatic = 243 \pm automatic = cohort TD$						
40	targetId = 267, # papulation cobot ID						
41	cohortTable = "dep2dCohort"						
42							
43							
44							
45	dataOut <- "./data"						
46	saveDir <- "./out_"						
47	<pre>#DatabaseConnector::downloadJdbcDrivers('sql server','~/.config/jdbc/')</pre>						
48	conn <- DatabaseConnector::createConnectionDetails(
49	'sql server',						
50	user = 'sa',						
51	password = 'hiking@tmu2',						
52	server = 10,104,1.104',						
55	pathiouriver = ~/.config/jabc/						
54							
56 =	for (key in names(nairs)) {						
57	targetId <- pairs[[key]][['targetId']]						
58	outcomeId <- pairs[[kev]][[outcomeId']]						
59	<pre>cohortTable <- pairs[[key]][['cohortTable']]</pre>						
60	outcomeTable <- pairs[[key]][['cohortTable']]						
61							
62	<pre>cohortIds <- c(targetId, outcomeId)</pre>						



Prediction	Viewer												
Model Desig	gns Summary	,											+
Design ID	Model Type	Target Pop	Outcome	TAR	min AUROC	mean AUROC	max AUROC	Num. Diagnostic Dbs	Num. Development Dbs	Num. Validation Dbs			
1	logistic	Cohort: 242	Cohort: 243	(cohort start +	0.800	0.800	0.800	1	1	1	View	View	View
2	LightGBM	Cohort: 242	Cohort: 243	start + 1825) (cohort start + 1) - (cohort	0.737	0.737	0.737	1	1	1	View Diagnostics	View Results	View Report
3	knn	Cohort: 242	Cohort: 243	start + 1825) (cohort start + 1) - (cohort start + 1825)	0.603	0.603	0.603	1	1	1	View Diagnostics	View Results	View Report



Prediction	ı Viewer												
Model Desig	gns Summary												+
Design ID	Model Type	Target Pop	Outcome	TAR	min AUROC	mean AUROC	max AUROC	Num. Diagnostic Dbs	Num. Development Dbs	Num. Validation Dbs			
1	logistic	Cohort: 262	Cohort: 243	(cohort start + 1) - (cohort start + 1825)	0.832	0.832	0.832	1	1	1	View Diagnostics	View Results	View Report
2	LightGBM	Cohort: 262	Cohort: 243	(cohort start + 1) - (cohort start + 1825)	0.823	0.823	0.823	1	1	1	View Diagnostics	View Results	View Report
3	knn	Cohort: 262	Cohort: 243	(cohort start + 1) - (cohort start + 1825)	0.547	0.547	0.547	1	1	1	View Diagnostics	View Results	View Report



Depression to Dementia

Shiny Application	Viewer												
Model Designs Summary												+	
Design ID	Model Type	Target Pop	Outcome	TAR	min AUROC	mean AUROC	max AUROC	Num. Diagnostic Dbs	<u>Num.</u> Development Dbs	Num. Validation Dbs			
1	logistic	Cohort: 267	Cohort: 243	(cohort start + 1) - (cohort start + 1825)	0.805	0.805	0.805	1	1	1	View Diagnostics	View Results	View Report
2	LightGBM	Cohort: 267	Cohort: 243	(cohort start + 1) - (cohort start + 1825)	0.847	0.847	0.847	1	1	1	View Diagnostics	View Results	View Report
3	knn	Cohort: 267	Cohort: 243	(cohort start + 1) - (cohort start + 1825)	0.745	0.745	0.745	1	1	1	View Diagnostics	View Results	View Report



We're seeking for collaboration from our partners

• Github repository: <u>https://github.com/OHDSI-TAIWAN</u>



Thank you for your attention!



