BMJ Open Predicting hospitalisations related to ambulatory care sensitive conditions with machine learning for population health planning: derivation and validation cohort study

Seung Eun Yi,^{1,2} Vinyas Harish,^{3,4,5,6} Jahir Gutierrez,¹ Mathieu Ravaut,⁷ Kathy Kornas,³ Tristan Watson,^{3,8} Tomi Poutanen,¹ Marzyeh Ghassemi,^{2,6} Maksims Volkovs,¹ Laura C Rosella [©] ^{3,4,5,6,8,9}

To cite: Yi SE, Harish V, Gutierrez J, et al. Predicting hospitalisations related to ambulatory care sensitive conditions with machine learning for population health planning: derivation and validation cohort study. BMJ Open 2022;12:e051403. doi:10.1136/ bmjopen-2021-051403

Prepublication history and additional supplemental material for this paper are available online. To view these files. please visit the journal online (http://dx.doi.org/10.1136/ bmjopen-2021-051403).

Received 18 May 2021 Accepted 28 February 2022



@ Author(s) (or their employer(s)) 2022. Re-use permitted under CC BY-NC. No commercial re-use. See rights and permissions. Published by

For numbered affiliations see end of article.

Correspondence to

Dr Laura C Rosella: laura.rosella@utoronto.ca

ABSTRACT

Objective To predict older adults' risk of avoidable hospitalisation related to ambulatory care sensitive conditions (ACSC) using machine learning applied to administrative health data of Ontario, Canada. Design, setting and participants A retrospective cohort study was conducted on a large cohort of all residents covered under a single-payer system in Ontario, Canada over the period of 10 years (2008-2017). The study included 1.85 million Ontario residents between 65 and 74 years old at any time throughout the study period.

Data sources Administrative health data from Ontario, Canada obtained from the (ICES formely known as the Institute for Clinical Evaluative Sciences Data Repository. Main outcome measures Risk of hospitalisations due to ACSCs 1 year after the observation period.

Results The study used a total of 1 854 116 patients, split into train, validation and test sets. The ACSC incidence rates among the data points were 1.1% for all sets. The final XGBoost model achieved an area under the receiver operating curve of 80.5% and an area under precision-recall curve of 0.093 on the test set, and the predictions were well calibrated, including in key subgroups. When ranking the model predictions, those at the top 5% of risk as predicted by the model captured 37.4% of those presented with an ACSCrelated hospitalisation. A variety of features such as the previous number of ambulatory care visits, presence of ACSC-related hospitalisations during the observation window, age, rural residence and prescription of certain medications were contributors to the prediction. Our model was also able to capture the geospatial heterogeneity of ACSC risk in Ontario, and especially the elevated risk in rural and marginalised regions. Conclusions This study aimed to predict the 1-year

risk of hospitalisation from ambulatory-care sensitive conditions in seniors aged 65-74 years old with a single, large-scale machine learning model. The model shows the potential to inform population health planning and interventions to reduce the burden of ACSC-related hospitalisations.

Strengths and limitations of this study

- ► The study is based on a large and highly diverse cohort, with a wide range of features to predict the risk of hospitalisation for ambulatory care sensitive conditions (ACSCs).
- The model is designed in a way it can be deployed in a real-world setting at relatively low cost, using available administrative health data.
- A rigorous calibration evaluation was conducted globally and across subgroups to validate the robustness of our model.
- ► The definition of ACSCs varies across different jurisdictions, even though the set of conditions used in this study is common in most definitions.
- The model does not have access to some key indicators of patients' health and behavioural status (eq. smoking, body mass index), known to be important determinants of ACSC risks.

INTRODUCTION

Health systems globally strive to provide quality care in alignment with the Quadruple Aim.1 Ambulatory care sensitive conditions (ACSCs) are a useful indicator of health system performance often used to support Quadruple Aim efforts. Reducing the burden of ACSCs is an opportunity to promote higher value care by avoiding the usage of unnecessary hospital and health system resources.3 ACSCs refer to conditions that can be managed in primary care settings, and thus hospitalisation for these conditions is considered potentially avoidable with adequate primary care.⁴ The list of the conditions considered to be ACSCs varies by country but commonly includes major chronic diseases.^{5–7} For example, in Canada, ACSCs comprise angina, asthma, chronic



obstructive pulmonary disease (COPD), diabetes, epilepsy and heart failure (HF).8 Outcomes with respect to these ACSCs are often used to measure how a health system specifically supports individuals with chronic diseases.⁹ Although only 0.4% of Canadians under the age of 75 have an ACSC-related hospitalisation, these events use nearly 11% of hospital bed days. 10 In the UK, emergency admissions account for 67% of all hospital days, costing £12.5B annually, most of which are considered preventable. 11 ACSCs have increased by 47% over the last 15 years (as of 2017) and account for one in five unplanned admissions. 12 Furthermore, older adults disproportionately experience ACSCs. For example, adults over 65 represented 69.1% of all ACSCs in Ireland in 2016. 13 and more than half of ACSCs in Canada were from individuals over 60 years old. ¹⁰ Therefore, reducing ACSCs represent a challenge in health systems around the world and pose an increasing burden for health systems as rates of multimorbidity continue to rise. 14 15

Hospitalisations due to ACSCs are also strongly related to socioeconomic status, even in settings of universal health coverage. Studies in Canada, the UK, the USA and Ireland^{16–19} have demonstrated the association between low socioeconomic status (often measured through arealevel deprivation measures) and higher rates of ACSCs. As these ACSC disparities can be due to systems-level barriers, a number of community-based interventions such as pay for performance schemes or multidisciplinary programmes to prevent readmissions have been proposed to potentially reduce the rates of these hospitalisations.²⁰ There is a need to develop tools for predicting the admissions that are most likely to be preventable to guide the deployment of such interventions or to target resources to optimise impact.^{20 21} Risk prediction algorithms developed on big data sources such as electronic medical records and routinely collected administrative health data (AHD) offer the ability to segment populations over space and time based on their likelihood of having an outcome of interest, 22 which can be used as an input to optimise outcomes. AHD are comprehensive, widely available and automatically generated based on interactions with the healthcare system. Furthermore, in certain settings, these data can be linked with other data sources (eg, environmental exposures, area-level measures of the social determinants of health, etc). 23 24 These characteristics make them particularly appealing for developing risk prediction algorithms that can be deployed at the level of populations for health system planning. While there is an increasing number of risk prediction models intended for clinical use in individuals in an ambulatory setting, 25-28 there are few examples of a single, unified model that can be deployed on routinely collected data to regularly support population health and health system management.²⁹ Databases with analogous AHD are available in most single-payer healthcare systems such as the UK, Australia and New Zealand. However, access to extensive medical records is not limited to single-payer countries. Databases of commercial insurance claim data

are also available for large portions of the population in countries with private healthcare systems such as the USA. Consequently, we believe that with sufficient adaptation, our proposed approach has wide applicability for assessing the risk of ACSCs in the population using routinely collected data.

The specific aim of this study is to develop and internally validate a single, large-scale, machine learning model to predict hospitalisations due to ACSCs in a cohort of 1.85 million older adults in Ontario, Canada. Ontario's population is large, diverse and covered under a singlepayer system. ^{30 31} We extracted a wide variety of features from Ontario's comprehensive AHD databases including sociodemographics, prescribed medications, insurance claims from physician's visits and hospitalisations, as well as laboratory values. We evaluated model performance including discrimination, calibration and report on the top contributing features. We finally conducted a geographical analysis to demonstrate how the model captures risk distribution in the population and thus can be used for guiding population health planning and community-based interventions.

METHODS

Data sources

We used AHD from Ontario, Canada which we obtained from ICES, an independent, non-profit research institute whose legal status under Ontario's health information privacy law allows it to collect and analyse healthcare and demographic data, without consent, for health system evaluation and improvement.³² Ontario, with a population of 14.7 million people as of April 2020, 31 is Canada's most populous province with close to 250 reported ethnicities (2016 Census³⁰), making it one of the most diverse in the world. The vast majority of Ontario residents are eligible for universal health coverage, leading to highly comprehensive and representative records. Besides demographic information, the records from ICES include linked claims data that are routinely collected every time a patient interacts with the healthcare system. To link all attributes for each patient, we used a unique identification number assigned to each individual at the Registered Persons Database (RPDB). All analyses were carried out on the Health AI Data Analytics Platform, which enables high performance computing in a secure environment to protect patient privacy. All datasets used from the ICES data repository are listed in online supplemental tables 1 and 2. These datasets were linked using unique encoded identifiers and analysed at ICES.

Cohort

Our study period spanned a 10-year window from 1 January 2008 through 31 December 2017. As older adults are disproportionately affected by ACSCs, 10 we only included patients who were between 65 and 74 years of age at any point during the 10-year study period (see figure 1A) and who were covered under Ontario Health

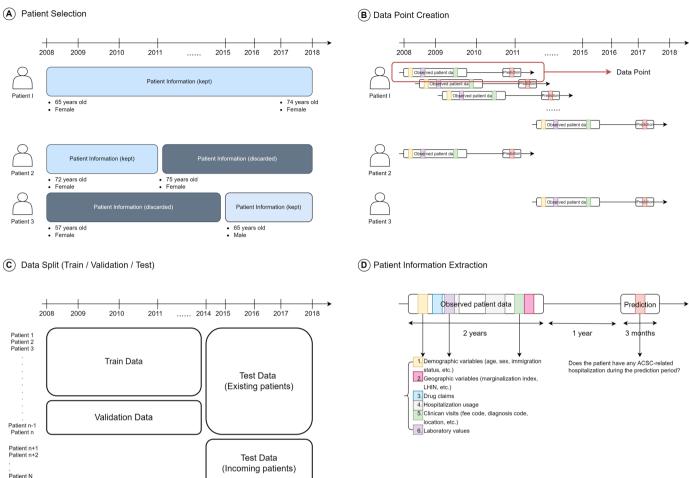


Figure 1 Overview of the data preparation. Panel A shows the patient selection process. For a given patient, only the information when they are between 65 and 74 years old was kept and the rest was discarded. This was done for the whole study period (between 2008 and 2017, included). Panel B then shows the construction of instances for the eligible patients each instance consisted of 2 years of observation window, 1 year of buffer where we did not have any information about the patient, and 3 months of target window. This instance was therefore a summary of the patient health information. Note that one patient could generate multiple instances. The first instance had an observation window from January 2008 to December 2009, a buffer from January 2010 to December 2010 and a target window from January 2011 to March 2011, all included. The last instance had an observation window from October 2014 to September 2016, a buffer from October 2016 to September 2017 and a target window from October 2017 to December 2017, all included. Instances for the same patient did not share any time period of the target window with each other, meaning each patient could have a maximum of 28 instances with non-overlapping target widows. This was often not the case due to the exclusion criteria and the individuals not fitting the age group of 65-74 years old anymore. For model development, as shown in panel C, we split our data based on patients and study period. We trained and validated our prediction model with data between 2008 and 2014, on two different sets of patients. Then, we tested the model on two types of data: data of the patients who were already used for training and validation between 2014 and 2018, and data of the patients who entered the cohort in 2014 and onwards. The latter consisted of 'young' patients between 65 and 66 years old at the beginning of the observation period. Finally, panel D shows the types of information that was extracted from each patient. ACSC, ambulatory care sensitive condition; LHIN, Local Health-Integrated Network.

Insurance Plan. Baseline characteristics of the patients included in our cohort are shown in table 1. The methodology used to extract and prepare patient data is illustrated in figure 1.

Study design

As shown in figure 1B, each patient's timeline of interactions with the healthcare system was broken down into *instances*. Each instance comprised three components: 2 years of patient history (*observation window*), 1 year during which we did not extract any information (*buffer*) and a

3-month period where we assess the risk of hospitalisation due to an ACSC (target window). We used 2 years of patient history to collect enough information about the patient and calculate predictor variables. Having a 1-year buffer period ensured that we are forecasting the risk of future preventable hospitalisations, as opposed to classifying whether or not a preventable hospitalisation has occurred within a current instance. The 3-month target window was selected due to the update frequency of administrative data at ICES, which also takes place every

BMJ Open: first published as 10.1136/bmjopen-2021-051403 on 1 April 2022. Downloaded from http://bmjopen.bmj.com/ on April 15, 2023 at Nanyang Technological University. Protected by copyright.

Table 1 Characteristics of the	Characteristics of the patients in the study					
		Train	Vali	Validation	1 1	Test
	Total	Positives	Total	Positives	Total	Positives
Cohort						
Unique patients	1 237 507	119 728	309 380	30 224	1 375 277	80 788
Data points	16 923 230	180 152	4 225 605	45 571	9 862 034	104 676
Demographics						
Sex (F)	8 845 443 (52.3%)	93 333 (51.8%)	2 213 395 (52.4%)	23 542 (51.7%)	5 153 174 (52.3%)	54 368 (51.9%)
Sex (M)	8 077 787 (47.7%)	86 819 (48.2%)	2 012 210 (47.6%)	22 029 (48.3%)	4 708 860 (47.7%)	50 308 (48.1%)
Age (mean)	69.0±2.86	69.1±2.88	69.0±2.86	69.1±2.87	69.0±2.81	69.1±2.84
Immigrant	1 619 193 (9.57%)	16 686 (9.26%)	405 211 (9.59%)	4007 (8.79%)	1 071 959 (10.9%)	10 878 (10.4%)
Non-immigrant	15 304 037 (90.4%)	163 466 (90.7%)	3 820 394 (90.4%)	41 564 (91.2%)	8 790 075 (89.1%)	93 798 (89.6%)
Geography						
Rural	2 479 688 (14.7%)	29 056 (16.1%)	616 941 (14.6%)	7069 (15.5%)	1 340 756 (13.6%)	15 499 (14.8%))
Income quintile						
1st quintile	2 982 780 (17.6%)	33 814 (18.8%)	736 716 (17.4%)	8612 (18.9%)	1 842 945 (18.7%)	20 940 (20.0%)
2nd quintile	3 335 222 (19.7%)	36 536 (20.3%)	835 276 (19.8%)	8847 (19.4%)	2 007 802 (20.4%)	21 849 (20.9%)
3rd quintile	3 351 614 (19.8%)	35 237 (19.6%)	838 012 (19.8%)	9043 (19.8%)	1 997 302 (20.3%)	21 266 (20.3%)
4th quintile	3 507 494 (20.7%)	36 756 (20.4%)	882 844 (20.9%)	9421 (20.7%)	1 913 979 (19.4%)	19 838 (19.0%)
5th quintile	3 702 063 (21.9%)	37 314 (20.7%)	921 278 (21.8%)	9521 (20.9%)	2 084 008 (21.1%)	20 613 (19.7%)
Education quintile						
1st quintile	3 075 278 (18.2%)	34 957 (19.4%)	762 687 (18.0%)	8779 (19.3%)	1 706 122 (17.3%)	19 391 (18.5%)
2nd quintile	3 385 511 (20.0%)	37 128 (20.6%)	841 692 (19.9%)	9481 (20.8%)	1 937 907 (19.7%)	21 469 (20.5%)
3rd quintile	3 493 804 (20.6%)	36 945 (20.5%)	870 402 (20.6%)	9190 (20.2%)	2 030 870 (20.6%)	21 454 (20.5%)
4th quintile	3 500 217 (20.7%)	36 902 (20.5%)	877 965 (20.8%)	9299 (20.4%)	2 084 940 (21.1%)	21 550 (20.6%)
5th quintile	3 357 516 (19.8%)	33 022 (18.3%)	844 839 (20.0%)	8485 (18.6%)	2 034 100 (20.6%)	20 045 (19.1%)
Comorbidities						
Acute Myocardial Infarction 709 309 (4.19%)	ר 709 309 (4.19%)	9667 (5.37%)	177 912 (4.21%)	2507 (5.50%)	416 924 (4.23%)	5696 (5.44%)
Arrythmia	1 530 096 (9.04%)	18 798 (10.4%)	382 872 (9.06%)	4891 (10.7%)	914 621 (9.27%)	10 990 (10.5%)
Arthritis	354 214 (2.09%)	4455 (2.47%)	89 539 (2.12%)	1113 (2.44%)	225 636 (2.29%)	2719 (2.60%)
Asthma	2 034 989 (12.0%)	26 649 (14.8%)	508 111 (12.0%)	7.093 (15.6%)	1 246 159 (12.6%)	16 445 (15.7%)
Cancer	2 198 985 (13.0%)	25 044 (13.9%)	546 554 (12.9%)	6210 (13.6%)	1 317 439 (13.4%)	14 662 (14.0%)
Chronic heart failure	838 537 (4.95%)	14 327 (7.95%)	210 965 (4.99%)	3672 (8.06%)	483 842 (4.91%)	7974 (7.62%)
Colitis	133 502 (0.79%)	1575 (0.87%)	32 232 (0.76%)	389 (0.85%)	88 715 (0.90%)	1018 (0.97%)
						Continued

lable I collillided						
	_	Train	Valid	Validation	Te	Test
	Total	Positives	Total	Positives	Total	Positives
Chronic Obstructive Pulmonary Disease	2 910 066 (17.2%)	40 751 (22.6%)	726 976 (17.2%)	10 396 (22.8%)	1 726 397 (17.5%)	23 837 (22.8%)
Coronary heart disease	3 510 361 (20.7%)	43 579 (24.2%)	879 995 (20.8%)	11 068 (24.3%)	1 899 361 (19.3%)	23 359 (22.3%)
Dementia	344 161 (2.03%)	4454 (2.47%)	86 853 (2.06%)	1158 (2.54%)	206 300 (2.09%)	2722 (2.60%)
Diabetes	4 394 754 (26.0%)	52 155 (29.0%)	1 095 489 (25.9%)	13 022 (28.6%)	2 660 592 (27.0%)	30 929 (29.5%)
Hypertension	10 494 913 (62.0%)	116 393 (64.6%)	2 622 675 (62.1%)	29 352 (64.4%)	6 033 618 (61.2%)	67 044 (64.0%)
Mental disorder*	3 410 850 (20.2%)	40 032 (22.2%)	851 249 (20.1%)	10 117 (22.2%)	2 244 412 (22.8%)	26 407 (25.2%)
Mood disorder	7 786 874 (46.0%)	85 908 (47.7%)	1 943 700 (46.0%)	21 771 (47.8%)	4 801 635 (48.7%)	52 944 (50.6%)
Osteoarthritis	10 244 732 (60.5%)	111 020 (61.6%)	2 553 893 (60.4%)	28 336 (62.2%)	6 131 185 (62.2%)	66 621 (63.6%)
Osteoporosis	1 791 824 (10.6%)	18 620 (10.3%)	450 339 (10.7%)	4656 (10.2%)	1 032 043 (10.5%)	10 525 (10.1%)
Renal failure	727 558 (4.30%)	10 545 (5.85%)	180 987 (4.28%)	2698 (5.92%)	489 046 (4.96%)	7004 (6.69%)
Stroke	965 743 (5.71%)	12 601 (6.99%)	239 406 (5.67%)	3122 (6.85%)	547 368 (5.55%)	7026 (6.71%)
Total (median)	3 (2, 7)	3 (2, 5)	3 (2, 4)	3 (2, 5)	3 (2, 5)	3 (2, 5)

*History of mental health-related visit, excluding dementia, deliberate self-harm codes and mood disorder codes. COPD, chronic obstructive pulmonary disease.

BMJ Open: first published as 10.1136/bmjopen-2021-051403 on 1 April 2022. Downloaded from http://bmjopen.bmj.com/ on April 15, 2023 at Nanyang Technological University. Protected by copyright.

3 months. This mimicked a system where we 'screen' a population quarterly for the risk of potentially avoidable hospitalisations 1 year in the future.

We split our cohort into three different sets (see figure 1C). The training and validation sets had distinct patients, which ensures that we tested the ability of the model to generalise well on unseen patients. Both sets included target windows from January 2011 to December 2015. As for the test set, it included all instances of patients that have a valid target window between January 2016 and December 2017. Therefore, two types of patient groups are included in this test set: those from the training and validation sets who still qualify at a future period of time, and patients who were under 65 years old prior to 2016 but who qualify for the test time period. The model performance on the test set thus indicated how well the model generalises to patients out of time and to new patients (ie, in-domain, in-distribution generalisation).

Inclusion and exclusion criteria

Our cohort included all residents of Ontario between 65 and 74 years old during the study period. Anyone who became 65 during any given time of the study period was included in the study cohort, for example, those who turned 65 in January 2017 were included in our test set, as shown in figure 1. We excluded instances of patients who died or ended all interaction with the healthcare system before the end of the observation window. This was to ensure that we excluded patients who were by definition known to not have an outcome. We still include patients who die during the buffer period or the prediction period when training the model, being consistent with the fact that the model cannot know any future events outside of the observation period.

Outcome of interest

The Canadian Institute for Health Information (CIHI) defines age-standardised acute care hospitalisation rate for conditions where appropriate ambulatory care is thought to prevent or reduce the need for admission to hospital.⁸ There are seven different types of avoidable hospitalisation-related ACSCs as per the CIHI definition: epilepsy, COPD, asthma, diabetes, HF, hypertension and angina. For each hospitalisation or ambulatory usage, we determined if the primary diagnosis code was in the list of CIHI ACSC diagnosis codes. We then aggregated all codes to determine whether or not an instance contained any hospitalisation related to an ACSC from the above list during the target window, in which case we labelled the instance as a positive data point (ie, our predictions are not cause-specific). The list of the ACSC diagnosis codes used as well as additional rules for defining an ACSC can be found in online supplemental table 3.

Feature preparation

The features that were extracted from the data sources included patients' demographic and geographical information, drug prescription history, chronic conditions, clinician visits, hospital usage, laboratory results, as well as past history of ACSCs (see figure 1D). In order to select the most important features for the model and reduce the likelihood of overfitting, we took a forward feature selection-based approach³³ to only select a small subset from thousands of available features that would ensure the performance of the model closely matches that of the model using all features. We ended with 140 features in total. Details of the feature preparation as well as the full list of the features are outlined in online supplemental table 4.

Model development and evaluation

Our prediction model is a gradient boosting decision tree-based model, optimised using the XGBoost library in Python. 34 35 These models have been empirically shown as computationally efficient and high-performance algorithms for tabular data through a number of machine learning competitions as well as in machine learning for health papers.³⁶ These models are able to select and combine heterogeneous features from multiple data sources that are often uncorrelated, and are thus efficient when both categorical and continuous variables with different value ranges are present in the data. Moreover, they can implicitly handle missing values without imputation. Therefore, we did not impute missing data. We undersampled the negative instances of the training data by a factor of 8, to reduce class imbalance.³⁷ This undersampling factor was chosen empirically over a grid search. Validation and the test sets were left untouched to ensure an accurate evaluation of the model performance. The details of the model's hyperparameters as well as its performance comparison against a baseline logistic regression model are presented in online supplemental methods 1 and 2, as well as in online supplemental table 5 and figure 1.

We evaluated the performance of the model on the held-out test set. We first checked the distribution of the predictions of our model after calibrating the probabilities to account for undersampling.³⁸ Model performance was then measured in terms of area under the receiver operating curve (AUC), a widely used metric for risk prediction tasks. $^{38\,39}$ Since the model was trained on highly imbalanced data, we also calculated the area under precision-recall curve (AUPRC) to focus on the probability of correctly detecting patients with the highest risk of ACSC. 40 To assess the variation of model performance on different risk groups, we compared the model predictions on multiple subgroups created from the test set. These groups comprised different age, sex, immigration and socioeconomic groups obtained using data across datasets (eg, RPDB, CENSUS, Ontario Marginalization Index (ON-MARG) in online supplemental table 1). We tested the model on patient groups split based on the number of interactions they had with the healthcare system as well.

Based on our model, we also computed Shapley Additive Explanations values, the weighted average of marginal

contributions,⁴¹ to identify features that contribute the most to model predictions. We used a randomly selected pool of 50 000 instances to generate these values.

Geospatial variation

We finally examined the performance of our model in capturing geospatial variation of ACSC incidence rates in Ontario. Ontario has 14 Local Health-Integrated Networks (LHINs), responsible for coordinating and funding local healthcare to improve resources access and patient experience. The LHINs are further divided into 74 sub-LHINs, and each of these aims at identifying healthcare needs and priorities within the region. We chose sub-LHINs to represent the subdivisions of the province. Following the predictions of our model, we computed and plotted the average risk of ACSCs for patients in each sub-LHIN.

Patient and public involvement

No patients with ACSCs or members of the public at-large were involved in the conceptualisation, analysis, or the write-up of this study. However, the public at-large is involved at ICES through the form of a Public Advisory Council which provides input into decisions made on how research is conducted using the individual-level personal health information collected. We plan to disseminate the knowledge gained through this study with the use of press releases and presentations on the value of population health planning, for both ACSCs and more broadly, tailored to general public audiences.

RESULTS

Starting from the initial cohort of 4520076 patients aged 50 years or older as of 2008, we excluded 1873139 patients who did not meet inclusion criteria due to age restrictions. From the remaining 2646937 patients, 752 505 were removed for their absence of interaction with the healthcare system, and 40 316 patients had a death

date prior to the end of the first observation window. After the exclusion criteria, a total of 1 854 116 patients were selected, resulting in 31 010 869 instances. Among them, 1237507 patients with 16 921 175 instances were used for training, 309 380 patients with 4 227 660 instances for validation and 1375277 patients yielding 9 862 034 instances for testing. The ACSC incidence rates among instances were around 1.1% for all sets. Descriptive statistics for key demographic, socioeconomic and chronic illness variables are presented in table 1.

Model performance

The model achieved an average AUC of 80.5 (range 80.4–80.5) and AUPRC of 0.093, when evaluated on all instances in the test set. Figure 2 shows the overall calibration curve of the model as well as model evaluation on important subgroups of the population. The calibration curve of figure 2A has 20 bins of equal data size, and shows the average risk of ACSC predicted by the model versus the actual ACSC-related hospitalisation rate. The inset on the right side of the graph shows that the model is slightly overpredicting the risk.

Figure 2B shows the average prediction of the model compared with the real incidence rate of ACSCs for different subgroups of the population. As in the calibration curve, the model slightly overpredicts the risk but accurately captures the variation in ACSC hospitalisation risk across different subgroups (eg, higher risk of hospitalisation in a subcohort with a higher age or a lower income index). Ranking the predictions made by the model, we found that the top 5% of test set instances predicted by the model as 'high-risk' covered 37.4% of instances who actually present an ACSC-related hospitalisation during the target window period. The top 1% and 10% of test set instances covered 15.2% and 50.8% of total positive data points, respectively. Main characteristics of these risk groups are described in online supplemental table 6.

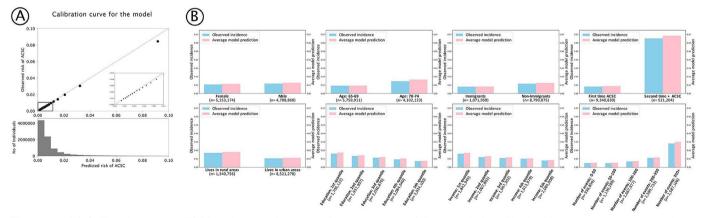


Figure 2 (A) Calibration curve. (B) Model evaluation on major subgroups of the population. The incidence rates are shown in blue and the average model prediction in pink. The subgroup sizes are displayed on the x-axis along with the subgroup types. For education and income quintiles, higher index refers to higher education level and income respectively, in the area a given patient lives in. The number of events refers to the number of any interaction a given patient had with the healthcare system—clinician visits, hospitalisation, ambulatory usage, lab tests and drug prescriptions. ACSC, ambulatory care sensitive condition.

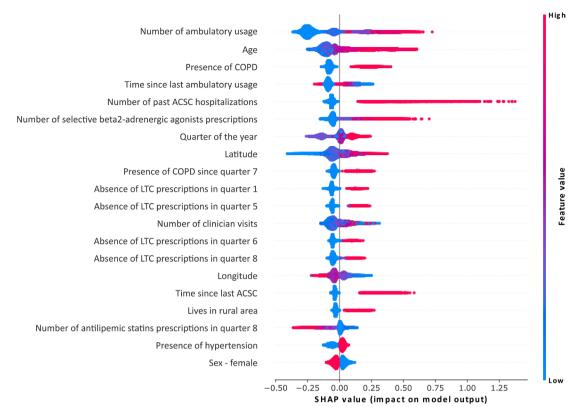


Figure 3 Global feature importance. Shapley values were generated using 50 000 random samples from the test set. Multiple runs using different samples showed the same ordering of feature contribution. ACSC, ambulatory care sensitive condition; COPD, chronic obstructive pulmonary disease.

Feature contribution

Figure 3 shows 20 features with the highest Shapley values. It is clear that the model is leveraging features across datasets, as the top features span demographics, geography, chronic conditions, prescriptions and interactions with the healthcare system. The AUC of the model when trained on individual datasets as well as different combinations of the subsets (see online supplemental figure 2) shows the impact of combining all features to achieve the best prediction performance. The previous number of ambulatory care visits, time since last ambulatory care visit, as well as presence of ACSC-related hospitalisations during the observation window were predictive of future ACSC-related hospitalisations. Several features related to medication prescription during the observation period were also found to be predictive of ACSC-related hospitalisations including the number of selective beta2-adrenergic agonists, absence of prescriptions for patients in long-term care (LTC) facilities and number of statin prescriptions. Geographical features such as latitude and rural residence were also predictive of our outcome. As shown in online supplemental figure 3, the model predictions are influenced by different features depending on whether or not a given individual already has an ACSC-related hospitalisation. Having a precedent was found to the contribution of future ACSC hospitalisations and the related features appear as the top supporting features.

Geospatial variation

Figure 4 shows the geospatial variation of ACSC incidence rates in Ontario. Figure 4A (map in red) shows the distribution of ACSC-related hospitalisations in different sub-LHINs of Ontario, weighted by the population size in each sub-LHIN. Figure 4B (map in blue) shows the distribution of ACSC risks predicted by the model. Our model is able to capture the geospatial heterogeneity of ACSC risk, in rural regions (eg, Northwestern Ontario) as well as in urban and populous areas (eg, Southern Ontario).

DISCUSSION

The goal of this work was to use AHD to develop a single, large-scale, machine learning model to predict hospitalisations due to ACSCs in a cohort of 1.85 million older adults in Ontario, Canada. This model is designed for population health planning and health resource allocation versus individual health decision-making. Our model accurately predicted the 1-year risk for ACSCs with a high AUC and the prediction values are well calibrated. The model leverages a variety of features across multiple administrative health databases to make its predictions, including outpatient visits, demographic information as well as past drug prescriptions. The presence of comorbidities such as COPD, hypertension or chronic HF as well as geography was important features in the model which is consistent with previously identified risk factors

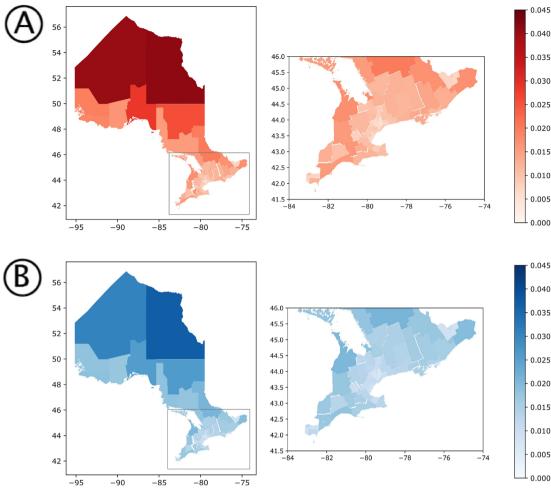


Figure 4 Distribution of ambulatory care sensitive condition (ACSC) risks predicted by the model compared with the actual variation in ACSC incidence rates in the province, in 2017. (A) The incidence rates of ACSC by sub-Local Health-Integrated Network (LHIN), normalised by the population size of the corresponding sub-LHIN. For (B), we computed the predictions of ACSC with our model for all patients and mapped them for different sub-LHINs. The model (in B) captures the normalised distribution of these patients (in A).

in Canada.¹⁰ Shapley values were provided for model transparency but should not be interpreted as causal risk factors. Our model is able to also capture geographical variations in risk across Ontario in both rural and urban settings.

Strengths and weaknesses in relation to other studies

There have been numerous studies that predict the risk of emergency admissions in community-dwelling adults, some of which focus on those that are potentially avoidable. A 2014 systematic review of 18 models that used either administrative or clinical data found that performance, as measured by the c-statistic, ranged between 0.63 and 0.83. Three of these studies focused on ACSCs. The first is an Italian study that used AHD to identify adult patients at risk of hospitalisation or death within a year who may be candidates for care through a 'patient-centred Medical Home' programme. We used similar sets of features, however, we were able to incorporate additional demographic and socioeconomic variables. Notably, we only created one single model for use in the entire population in order to facilitate the ease of

application at the population level, while achieving similar model performance in terms of discrimination and calibration. The second study involved predicting the 1-year risk of hospitalisations from ACSCs in a cohort from the Veteran's Health Administration in the USA. 44 An advantage of their work was the inclusion of individual-level socioeconomic variables. However, their cohort was much more homogenous than ours and designed for individual patient care of a very specific segment of the population. Such a model would not be applicable for population health planning or for use on routinely collected data where these individual determinants are not available. Finally, the Sussex Predictor of Key Events model leveraged a neural network architecture with 1000 features across diverse health datasets to predict admission risk due to a range of chronic diseases for every individual in Sussex. 46 While it achieved a slightly higher AUC (0.82) than our model, its complex architecture and use of almost ten times the amount of features present serious challenges for model implementation and increases the likelihood of overfitting.

Meaning of the study to clinicians and policy-makers

With our model, we are able to assess the 1-year risk of hospitalisations due to ACSCs by aggregating individual patient risks at a subregional level on a quarterly basis. It is important to see the ACSC rates not simply as a marker of access to primary care, but as an indicator for how resources could be allocated and optimised in a health system by better addressing the needs of specific regions and patient groups. In Ontario, it has been demonstrated that ambulatory care services for cardiovascular disease were actually provided more frequently in regions with lower rates of cardiovascular events (ie, myocardial infarction, stroke or cardiovascular-related death). 47 Therefore, a model such as the one developed in this work is a valuable tool for health systems planners to inform healthcare delivery models and allocate community-based interventions over space and time with a focus on health equity. 4748 Such reallocations are thought to be impactful, as it has been shown that readmission due to certain conditions, such as HF, can be reduced by the efficient and timely allocation of inpatient and outpatient counselling and monitoring efforts. 49

Strengths and weaknesses of the study

Our study has several strengths. Our cohort is large and is constructed from a highly diverse population. We developed a model that uses a wide range of features to predict the risk of hospitalisation for ACSCs, which is an indicator of the effectiveness of the healthcare system indicator. The model is designed in a way it can be deployed in a real-world setting at relatively low cost, using available AHD as well as area-level data that capture concepts related to the social determinants of health. It also allows risk assessment at different aggregation levels for geography and over time. Given that population-risk scores are known to be biased,⁵⁰ we conducted a rigorous calibration evaluation not just globally, but in subgroups as well. However, our work also has some important limitations. A challenge lies in the focus of ACSC as an outcome, which has been the focus of some debate regarding the extent of its preventable nature.⁵¹ Furthermore, the definition of ACSCs relies on diagnosis codes that are often considered to be a partial way of patient condition assessment.⁵² It varies across different jurisdictions, even though the set of conditions used in this study is common in most definitions.^{5 7 8 51} The lack of access to certain indicators (eg, smoking, body mass index, etc) that are known to be important determinants of ACSC risks^{10 44} is expected to bottleneck model performance and emphasises the difficulty of assessing an individual's risk without fully observing their health and behavioural status.⁵³ A 3-month lag period for administrative data updates also limits any potential applications in real-time; however, we have designed our system for use 1 year in the future. By excluding patients without known interactions with the health system, it is possible we are underestimating risk for individuals who have very little interaction with the health system or other barriers to access. However, we do

still capture demographic and geographical information for those individuals, which do contribute to the risk estimate and minimise any impact.

Future directions

While our study focused solely on AHD, it is theoretically possible to link this information with data captured in inpatient electronic medical records (EMRs) such that individual clinicians can use this information to direct specific care or treatments. The EMR would also facilitate the capture of individual clinical or risk behaviour features that are known to contribute ACSC-related hospitalisation risk and thus may also increase model performance. Future work could also test longer buffer periods, such as 3 or 5 years, to support resource allocation efforts at the population level with greater lead time (eg, health region-level allocations for enrollees in an intervention). Finally, there is uncertainty around what interventions are most effective at reducing rates of hospitalisations in highrisk populations. For example, trials of care coordination programmes have demonstrated mixed results for various chronic diseases. 5455 Designing and validating these interventions across the population remains an important area for future work. In cases where there are data from realworld pilots of interventions, causal machine learning approaches may be able to help improve the design and targeting of such interventions by identifying subgroups within a population that may have the largest treatment effects.29

Conclusion

In this work, we demonstrate that the development and validation of a single, large-scale machine learning model to predict the 1-year risk of hospitalisation from a series of ambulatory-care sensitive conditions is feasible in a large and diverse cohort of seniors using AHD. Such a model has the potential to reduce the burden of hospitalisations from ambulatory care conditions by supporting the allocation of community-based interventions during population health planning and health resource allocation.

Author affiliations

¹Layer6 AI, Toronto, Ontario, Canada

²Department of Computer Science, University of Toronto, Toronto, Ontario, Canada
 ³Dalla Lana School of Public Health, University of Toronto, Toronto, Ontario, Canada
 ⁴Temerty Faculty of Medicine, University of Toronto, Toronto, Ontario, Canada

⁵Temerty Centre for Artificial Intelligence Research and Education in Medicine,

University of Toronto, Toronto, Ontario, Canada

⁶Vector Institute, Toronto, Ontario, Canada

 $^{7}\mbox{School}$ of Computer Science and Engineering, Nanyang Technological University, Singapore

⁸Institute for Clinical Evaluative Sciences, Toronto, Ontario, Canada

⁹Institute for Better Health, Trillium Health Partners, Mississauga, Ontario, Canada

Twitter Laura C Rosella @LauraCRosella

Acknowledgements This study was supported by ICES, which is funded by an annual grant from the Ontario Ministry of Health (MOH) and the Ministry of Long-Term Care (MLTC). Parts of this material are based on data and information compiled and provided by the Ministry of Health (MOH) and Canadian Institute for Health Information (CIHI). However, the analyses, conclusions, opinions and statements expressed in the material are those of the author(s), and not necessarily

those of CIHI or MOH. Parts or whole of this material are based on data and/ or information compiled and provided by Immigration, Refugees and Citizenship Canada (IRCC) current to 2018. However, the analyses, conclusions, opinions and statements expressed in the material are those of the author(s), and not necessarily those of IRCC. We thank IQVIA Solutions Canada Inc. for use of their Drug Information File.

Contributors LCR conceptualised the study and secured funding for the research. SEY, TP, MV and LCR planned the first version of the study protocol. TW and KK facilitated data acquisition to the data and TW prepared the analytic cohort with support from KK. MV, MR, VH and MG critically reviewed the study protocol provided initial and ongoing input on the analysis. SEY carried out the analysis with contributions from JG, MR, MV. VH and KK took part in interpreting results and providing feedback for model refinement. SEY and VH wrote the first draft of the paper. All authors contributed to the critical revision of the manuscript for important intellectual content and approved the final version of the manuscript. LCR is the guarantor and is responsible for the overall content.

Funding This work was supported by the New Frontiers in Research Fund (NFRFE-2018-00662), a Canada Research Chair in Population Health Analytics (950-230702) (LR), Ontario Graduate Scholarship (number N/A) (VH), Canadian Institutes of Health Research Banting and Best Canada Graduate Scholarship Master's and Doctoral awards (numbers N/A) (VH), and Vector Institute Post-graduate Fellowship (number N/A) (VH).

Competing interests All authors have completed the ICMJE uniform disclosure form at www.icmje.org/coi_disclosure.pdf and declare: SY, JG, MR, MV, and TP are full-time employees of Layer 6 Al, cofounded by MV and TP, owned by Toronto-Dominion Bank. VH, KK and TW are employed at the Dalla Lana School of Public Health. The employers of the authors had no role in the design or funding of this research.

Patient and public involvement Patients and/or the public were involved in the design, or conduct, or reporting, or dissemination plans of this research. Refer to the Methods section for further details.

Patient consent for publication Not applicable.

Ethics approval This prokevct was reviewed by the University of Toronto Research Ethics Board (protocol number is 37650).ICES has obtained ethical approval (and repeats this review tri-annually) for its privacy and security policies, procedures, and practices. Each research project that is conducted at ICES is also subject to internal ethical review by the ICES Privacy and Compliance Office. ICES is a prescribed entity under section 45 of Ontario's Personal Health Information Protection Act (PHIPA). Section 45 is the provision that enables analysis and compilation of statistical information related to the management, evaluation, and monitoring of, allocation of resources to, and planning for the health system. Section 45 authorizes health information custodians to disclose personal health information to a prescribed entity, like ICES, without consent for such purposes. Projects conducted wholly under section 45, by definition, do not require review by a Research Ethics Board. As a prescribed entity, ICES must submit to trio-annual review and approval of its privacy and security policies, procedures and practices by Ontario's Information and Privacy Commissioner. These include policies, practices and procedures that require internal review and approval of every project by ICES' Privacy and Compliance Office.

Provenance and peer review Not commissioned; externally peer reviewed.

Data availability statement Data may be obtained from a third party and are not publicly available. The dataset for this study is held securely in coded form at ICES. While data sharing agreements prohibit ICES from making the dataset publicly available, access may be granted to those who meet prespecified criteria for confidential access, available at www.ices.on.ca/DAS. The full dataset creation plan is available from the authors upon request.

Supplemental material This content has been supplied by the author(s). It has not been vetted by BMJ Publishing Group Limited (BMJ) and may not have been peer-reviewed. Any opinions or recommendations discussed are solely those of the author(s) and are not endorsed by BMJ. BMJ disclaims all liability and responsibility arising from any reliance placed on the content. Where the content includes any translated material, BMJ does not warrant the accuracy and reliability of the translations (including but not limited to local regulations, clinical guidelines, terminology, drug names and drug dosages), and is not responsible for any error and/or omissions arising from translation and adaptation or otherwise.

Open access This is an open access article distributed in accordance with the Creative Commons Attribution Non Commercial (CC BY-NC 4.0) license, which permits others to distribute, remix, adapt, build upon this work non-commercially,

and license their derivative works on different terms, provided the original work is properly cited, appropriate credit is given, any changes made indicated, and the use is non-commercial. See: http://creativecommons.org/licenses/by-nc/4.0/.

ORCID ID

Laura C Rosella http://orcid.org/0000-0003-4867-869X

REFERENCES

- 1 Kruk ME, Gage AD, Arsenault C, et al. High-quality health systems in the sustainable development goals era: time for a revolution. Lancet Glob Health 2018:6:e1196–252.
- 2 Sikka R, Morath JM, Leape L. The quadruple aim: care, health, cost and meaning in work. *BMJ Qual Saf* 2015;24:608–10.
- 3 Enhancing the continuum of care report of the avoidable hospitalization advisory panel. Submitted to the Ministry of Health and Long-Term care, 2011. Available: http://www.health.gov.on.ca/en/common/ministry/publications/reports/baker_2011/baker_2011. pdf [Accessed 5 Jan 2021].
- 4 Purdy S, Griffin T, Salisbury C, et al. Ambulatory care sensitive conditions: terminology and disease coding need to be more specific to aid policy makers and clinicians. Public Health 2009;123:169–73.
- 5 Sarmento J, Rocha JVM, Santana R. Defining ambulatory care sensitive conditions for adults in Portugal. BMC Health Serv Res 2020:20:754.
- 6 Ambulatory Care Sensitive Conditions (ACSC) NHS Digital. Available: https://digital.nhs.uk/data-and-information/data-tools-and-services/data-services/innovative-uses-of-data/demand-on-healthcare/ambulatory-care-sensitive-conditions [Accessed 5 Jan 2021].
- 7 Prevention quality indicators (PQI) overview. Available: https://www.qualityindicators.ahrq.gov/Modules/pqi_resources.aspx [Accessed 5 Jan 2021].
- 8 Ambulatory care sensitive conditions. Available: http:// indicatorlibrary.cihi.ca/display/HSPIL/Ambulatory+Care+Sensitive+ Conditions [Accessed 5 Jan 2021]
- 9 Busby J, Purdy S, Hollingworth W. How do population, general practice and hospital factors influence ambulatory care sensitive admissions: a cross sectional study. BMC Fam Pract 2017;18:67.
- 10 Hospitalizations for Ambulatory Care Sensitive Conditions (ACSC). The factors that matter, 2011. Available: https://www150.statcan. gc.ca/n1/pub/82-622-x/82-622-x2011007-eng.htm [Accessed 5 Jan 2021].
- 11 Emergency admissions to hospital: managing the demand National Audit Office (NAO) Report. Available: https://www.nao.org.uk/report/emergency-admissions-hospitals-managing-demand/ [Accessed 5 Jan 2021].
- 12 QualityWatch: focus on preventable admissions. Available: https://www.health.org.uk/publications/qualitywatch-focus-on-preventable-admissions [Accessed 5 Jan 2021].
- 13 McDarby G, Smyth B. Identifying priorities for primary care investment in Ireland through a population-based analysis of avoidable hospital admissions for ambulatory care sensitive conditions (ACSC). *BMJ Open* 2019;9:e028744.
- 14 Feely A, Lix LM, Reimer K. Estimating multimorbidity prevalence with the Canadian chronic disease surveillance system. *Health Promot Chronic Dis Prev Can* 2017;37:215–22.
- 15 Rosella L, Kornas K, Huang A, et al. Accumulation of chronic conditions at the time of death increased in Ontario from 1994 to 2013. Health Aff 2018;37:464–72.
- Trachtenberg AJ, Dik N, Chateau D, et al. Inequities in ambulatory care and the relationship between socioeconomic status and respiratory hospitalizations: a population-based study of a Canadian City. Ann Fam Med 2014;12:402–7.
- 17 Grigoroglou C, Munford L, Webb R, et al. Impact of a national primary care pay-for-performance scheme on ambulatory care sensitive hospital admissions: a small-area analysis in England. BMJ Open 2020;10:e036046.
- 18 Billings J, Zeitel L, Lukomnik J, et al. Impact of socioeconomic status on hospital use in New York City. *Health Aff* 1993;12:162–73.
- 19 Conway R, O'Riordan D, Byrne D, et al. Deprivation influences the emergency admission rate of ambulatory care sensitive conditions. Clin Med 2016;16:119–23.
- 20 Wallace E, Smith SM, Fahey T, et al. Reducing emergency admissions through community based interventions. BMJ 2016;352:h6817.
- 21 Jain SH, Chandrashekar P. Implementing a targeted approach to social determinants of health interventions. Am J Manag Care 2020;26:502–4.



- 22 Bates DW, Saria S, Ohno-Machado L, et al. Big data in health care: using analytics to identify and manage high-risk and high-cost patients. Health Aff 2014;33:1123-31.
- Virnig BA, McBean M. Administrative data for public health surveillance and planning. Annu Rev Public Health 2001;22:213-30.
- Jutte DP, Roos LL, Brownell MD. Administrative record linkage as a tool for public health research. Annu Rev Public Health 2011;32:91-108.
- Srinivas S. A machine learning-based approach for predicting patient punctuality in ambulatory care centers. Int J Environ Res Public Health 2020:17:3703.
- Srinivas S, Salah H. Consultation length and no-show prediction for improving appointment scheduling efficiency at a cardiology clinic: a data analytics approach. Int J Med Inform 2021;145:104290.
- Geilleit R, Hen ZQ, Chong CY, et al. Feasibility of a real-time hand hygiene notification machine learning system in outpatient clinics. J Hosp Infect 2018;100:183-9.
- Angraal S, Mortazavi BJ, Gupta A, et al. Machine learning prediction of mortality and hospitalization in heart failure with preserved ejection fraction. JACC Heart Fail 2020;8:12-21.
- Marafino BJ, Schuler A, Liu VX, et al. Predicting preventable hospital readmissions with causal machine learning. Health Serv Res 2020;55:993-1002.
- Government of Ontario, Ministry of Finance. 2016 census highlights: Factsheet 9. Available: https://www.fin.gov.on.ca/en/economy/ demographics/census/cenhi16-9.html#:~:text=Ontario's%20 Population%20is%20Highly%20Diverse&text=About%20250%20 ethnic%20origins%20were%20reported%20by%20Ontarians%20 in%20the%202016%20Census [Accessed 5 Jan 2021].
- Ontario demographic Quarterly: highlights of first quarter, 2020. Available: https://www.ontario.ca/page/ontario-demographicquarterly-highlights-first-quarter-2020 [Accessed 5 Jan 2021].
- ICES. Available: https://www.ices.on.ca/ [Accessed 5 Jan 2021].
- Kohavi R, John GH. Wrappers for feature subset selection. Artif Intell 1997;97:273-324.
- Python API reference xgboost 1.4.0-SNAPSHOT documentation. Available: https://xgboost.readthedocs.io/en/latest/python/python api.html [Accessed 5 Jan 2021].
- Chen T, Guestrin C. XGBoost. Proceedings of the 22nd ACM SIGKDD international conference on knowledge discovery and data mining 2016.
- Bojer CS, Meldgaard JP. Kaggle forecasting competitions: an overlooked learning opportunity. *Int J Forecast* 2020.

 Akbani R, Kwek S, Japkowicz N. Applying support vector machines
- to imbalanced datasets. machine learning: ECML 2004:39-50.
- Pozzolo AD, Caelen O, Johnson RA. Calibrating probability with Undersampling for unbalanced classification. 2015 IEEE symposium series on computational intelligence 2015.
- Huang J, Ling CX. Using AUC and accuracy in evaluating learning algorithms. IEEE Transactions on Knowledge and Data Engineering 2005:17:299-310.

- 40 Ozenne B. Subtil F. Maucort-Boulch D. The precision-recall curve overcame the optimism of the receiver operating characteristic curve in rare diseases. J Clin Epidemiol 2015;68:855-9.
- 41 Lundberg SM, Erion GG, Lee S-I. Consistent individualized feature attribution for tree ensembles. arXiv [cs.LG], 2018. Available: http:// arxiv.org/abs/1802.03888
- Central West local health integration network (LHIN). Available: http:// www.centralwestlhin.on.ca/About%20Us/Frequently%20Asked% 20Questions/About%20Ontario%20LHINs.aspx [Accessed 5 Jan
- 43 Wallace E, Stuart E, Vaughan N, et al. Risk prediction models to predict emergency hospital admission in community-dwelling adults: a systematic review. Med Care 2014;52:751-65.
- Gao J, Moran E, Li Y-F, et al. Predicting potentially avoidable hospitalizations. Med Care 2014;52:164-71.
- 45 Louis DZ, Robeson M, McAna J, et al. Predicting risk of hospitalisation or death: a retrospective population-based analysis. BMJ Open 2014:4:e005223.
- Oliver-Baxter J, Bywood P, Erny-Albrecht K. PHCRIS policy issue review: predictive risk models to identify people with chronic conditions at risk of hospitalisation, 2015.
- Tu JV, Chu A, Maclagan L, et al. Regional variations in ambulatory care and incidence of cardiovascular events. CMAJ 2017;189:E494-501.
- Chen IY, Joshi S, Ghassemi M. Treating health disparities with artificial intelligence. Nat Med 2020;26:16-17.
- Amarasingham R, Patel PC, Toto K, et al. Allocating scarce resources in real-time to reduce heart failure readmissions: a prospective, controlled study. BMJ Qual Saf 2013;22:998-1005.
- Obermeyer Z, Powers B, Vogeli C, et al. Dissecting racial bias in an algorithm used to manage the health of populations. Science 2019;366:447-53.
- 51 Fleetcroft R. Hardcastle A. Steel N. et al. Does practice analysis agree with the ambulatory care sensitive conditions' list of avoidable unplanned admissions?: a cross-sectional study in the East of England. BMJ Open 2018;8:e020756.
- McKillop M, Polubriaginof F, Weng C. Exploration of temporal ICD coding bias related to acute diabetic conditions. AMIA Annu Symp Proc 2015;2015:2005-14.
- 53 Agniel D, Kohane IS, Weber GM. Biases in electronic health record data due to processes within the healthcare system: retrospective observational study. BMJ 2018;361:k1479.
- Peikes D, Chen A, Schore J, et al. Effects of care coordination on hospitalization, quality of care, and health care expenditures among Medicare beneficiaries: 15 randomized trials. JAMA 2009;301:603-18.
- McCall N. Cromwell J. Results of the Medicare health support disease-management pilot program. N Engl J Med 2011;365:1704-12.