## Evaluating Model Performance in Medical Datasets Over Time

Helen Zhou\*

Carnegie Mellon University, United States of America

Yuwen Chen\*

Carnegie Mellon University, United States of America

Zachary Lipton

Carnegie Mellon University, United States of America

HLZHOU@ANDREW.CMU.EDU

YUWENC2@ANDREW.CMU.EDU

ZLIPTON@CMU.EDU

## Abstract

Machine learning (ML) models deployed in healthcare systems must face data drawn from continually evolving environments. However, researchers proposing such models typically evaluate them in a time-agnostic manner, splitting datasets according to patients sampled randomly throughout the entire study time period. This work proposes the Evaluation on Medical Datasets Over Time (EMDOT) framework, which evaluates the performance of a model class across time. Inspired by the concept of backtesting, EMDOT simulates possible training procedures that practitioners might have been able to execute at each point in time and evaluates the resulting models on all future time points. Evaluating both linear and more complex models on six distinct medical data sources (tabular and imaging), we show how depending on the dataset, using all historical data may be ideal in many cases, whereas using a window of the most recent data could be advantageous in others. In datasets where models suffer from sudden degradations in performance, we investigate plausible explanations for these shocks. We release the EMDOT package to help facilitate further works in deployment-oriented evaluation over time.

**Data and Code Availability** We use the following data: (1) the Surveillance, Epidemiology, and End Results (SEER) cancer dataset (National Cancer Institute, 2020), (2) the COVID-19 Case Surveillance Detailed Data provided by the CDC (Centers for Disease Control and Prevention, 2020), (3) the Southwestern Pennsylvania (SWPA) COVID-19 dataset, (4) the MIMIC-IV intensive care database (Johnson et al., 2021), (5) the Organ Procurement and Transplantation Network (OPTN) database for liver transplant candidates (Organ Procurement and Transplantation Network, 2020), and (6) the MIMIC-CXR-JPG database of chest radiographs (Johnson et al., 2019a,b). MIMIC-IV and MIMIC-CXR-JPG (referred to as MIMIC-CXR in this paper) are available on the PhysioNet repository (Goldberger et al., 2000). Except for the SWPA dataset, all are publicly accessible (after accepting a data usage agreement). Details for accessing each dataset are in Appendices C-G. The code is publicly available on GitHub.

**Institutional Review Board (IRB)** This research does not require IRB approval.

## 1. Introduction

As medical practices, healthcare systems, and community environments evolve over time, so does the distribution of collected data. Features are deprecated as new ones are introduced, data collection may fluctuate along with hospital policies, and the underlying patient and disease populations may shift.

Amidst this ever-changing environment, models that perform well on one time period cannot be assumed to perform well in perpetuity. In the MIMIC-III critical care dataset, Nestor et al. (2019) found that a change to the electronic health record (EHR) system in 2008 coincided with sudden degradations in AUROC for mortality prediction. In COVID-19 data from the Centers for Disease Control and Prevention (CDC), Cheng et al. (2021) noted that the age distribution among cases shifted continually throughout the pandemic, and that these continual shifts confounded estimates of improvements in mortality rate.

<sup>\*</sup> These authors contributed equally

We propose an evaluation framework to characterize model performance over time by simulating training procedures that practitioners could have executed up to each time point, and subsequently deployed in future time points. We argue that standard time-agnostic evaluation is insufficient for selecting deployment-ready models, showing across several datasets that it over-estimates deployment performance. Instead, we advocate for EMDOT as a worthwhile pre-deployment step to help practitioners gain confidence in the robustness of their models to shifts in the data distribution that have occurred in the past and may to some extent repeat in the future.

There is a large body of work that addresses adaptation under various structured forms of distribution shift, including covariate shift (Shimodaira, 2000; Zadrozny, 2004; Huang et al., 2006; Sugiyama et al., 2007; Gretton et al., 2009), label shift (Saerens et al., 2002; Storkey, 2009; Zhang et al., 2013; Lipton et al., 2018; Garg et al., 2020), missingness shift (Zhou et al., 2022a), and concept drift (Tsymbal, 2004; Gama et al., 2014). However, in the real-world medical datasets we analyze, none of these structural assumptions can be guaranteed, and distributional changes in covariates, labels, missingness, etc. could even occur simultaneously. This motivates our empirical work, as it is unclear across a variety of model classes and medical datasets, how existing models might degrade due to naturally occurring changes over time, and whether different training practices might impact on robustness over time.

However intuitive it might seem, evaluation of models over time remains uncommon in standard machine learning for healthcare (ML4H) research. In the proceedings of the Conference on Health, Inference, and Learning (CHIL) 2022, for example, none of the 23 papers performed evaluations which took time into account (see Appendix A for similar statistics from CHIL 2021 and the Radiology medical journal). One possible reason for this is lack of access—as noted by Nestor et al. (2019), it is common practice to remove timestamps when de-identifying medical datasets for public use. In this work, we identify six sources of medical data containing varying granularities of temporal information per-record, five of which are *pub*licly available. We profile the performance of various training strategies and model classes across time, and identify possible sources of distribution shifts within each dataset. Finally, we release the Evaluation on Medical Datasets Over Time (EMDOT) Python package (details in Appendix B) to allow researchers to apply EMDOT to their own datasets and test techniques for handling shifts over time.

### 2. Related work

The promise of ML for improving healthcare has been explored in several domains, including cancer survival prediction (Hegselmann et al., 2018), diabetic retinopathy detection (Gulshan et al., 2016), antimicrobial stewardship (Kanjilal et al., 2020; Boominathan et al., 2020), recognizing diagnoses from electronic health record data (Lipton et al., 2016), and mortality prediction in liver transplant candidates (Bertsimas et al., 2019; Byrd et al., 2021). Typically, these ML models are evaluated on randomly held out patients, and sometimes externally validated on other hospitals or newly collected data. Even with crosssite validations, we cannot be sure how models will perform in the future.

For decades, the medical community has had a history of utilizing (mostly) fixed, simple risk scores to inform patient care (Hermansson and Kahan, 2018; Kamath et al., 2001; Wilson et al., 1998; Wells et al., 1995). Risk scores often prioritize ease-of-use, are computed from few variables, verified by domain experts for clear causal connections to outcomes of interest, and validated through use over time and across hospitals. Together, these factors give clinicians confidence that the model will perform reliably for years to come. With increasingly complex models, however, trust and adoption may be hindered by a lack of confidence in robustness to changing environments.

As noted by D'Amour et al. (2022), ML models often exhibit unexpectedly poor behavior when deployed in real-world domains. A key reason for these failures, they argue, is *under-specification*, where ML pipelines yield many predictors with equivalently strong held-out performance in the training domain, but such predictors can behave very differently in deployment. By testing performance across a variety of distribution shifts that have previously occurred over time, EMDOT could serve as a stress test to help combat under-specification.

Although evaluation over time is far from standard in ML4H literature, changes in performance over time have been noted in prior work. To predict wound-healing, Jung and Shah (2015) found that when data were split by cutoff time instead of patients, benefits of model averaging and stacking disappeared. Pianykh et al. (2020) found degradation in performance of a model for wait times dependent on how much historical data was trained on. To predict severe COVID-19, Zhou et al. (2022b) found that learned clinical concept features performed more robustly over time than raw features. Closest to our work is Nestor et al. (2019), which evaluated AU-ROC in MIMIC-III critical care data from 2003–2012, comparing training on just 2001–2002; the prior year; and the full history. Using the full history and curated clinical concepts, they bridged a big drop in performance due to changing EHR systems. Whereas Nestor et al. (2019) considers three models per test year, EMDOT simulates model deployment every year and evaluates across all future years.

While we do not consider time series models in this work (instead considering those which treat data as i.i.d.), there are similarities between how training sets are defined in EMDOT and in techniques for evaluating time-series forecasts (Bergmeir and Benítez, 2012; Cerqueira et al., 2020). These techniques often roll forward in time, taking either a window of recent data or all historical data as training sets, and evaluate test performance on the next time point. Performance from each time point is then averaged to summarize performance. This type of back-testing technique is common in rapidly evolving, non-stationary applications like finance (Chauhan et al., 2020; Alberg and Lipton, 2017), where time series models are constantly updated. In the healthcare domain, however, models may not be so easily updated, with risk scores developed several years ago still being used to this day (Six et al., 2008; Kamath et al., 2001; Wilson et al., 1998; Wells et al., 1995). Thus, we track performance not only the immediate year after the training set, but all subsequent years in the dataset. Additionally, instead of collapsing performance from models trained at different time points into summary statistics, which could conceal distribution shifts over time, our framework tracks these granular fluctuations over time, and creates tools to help provide insight into the nature and potential causes of such changes.

## 3. Data

We sought medical datasets that had: (1) a timestamp for each record, (2) interesting prediction task(s), and (3) enough distinct time points to evaluate over. Six data sources satisfied these criteria: SEER cancer data, national CDC COVID-19 data, COVID-19 data from a healthcare provider in Southwestern Pennsylvania (SWPA), MIMIC-IV critical care data, OPTN data from liver transplant candidates, and MIMIC-CXR chest radiographs. All datasets are tabular except for MIMIC-CXR (medical imaging data). All but SWPA are publicly accessible.

Table 1 summarizes the dataset outcomes, time ranges, and number of samples. Figure 1 visualizes data quantity over time. Appendices C–H include cohort selection diagrams, cohort characteristics, features, heat maps of missingness, preprocessing steps, and additional details. Categorical variables are converged to dummies, and numerical variables are normalized and centered at 0. Missing values in categorical variables are treated as another category, and in numerical variables they are imputed with the mean. In all datasets except MIMIC-CXR (where each sample is a distinct radiograph), each sample corresponds to a distinct patient.

#### 3.1. SEER Cancer Data

The Surveillance, Epidemiology, and End Results (SEER) Program collects cancer incidence data from registries throughout the U.S. Each case includes demographics, primary tumor site, tumor morphology, stage, diagnosis, first course of treatment, and survival outcomes (collected with follow-up) (National Cancer Institute, 2020). We use the SEER\*Stat software (Program, 2015) to define three cohorts of interest: (1) breast cancer, (2) colon cancer, and (3) lung cancer. The outcome is 5-year survival, i.e. whether the patient was confirmed alive five years after the year of diagnosis. The amount of data has mostly increased each year (Figure 1). Performance over time is evaluated *yearly*. See Appendix C for more details.

### 3.2. National CDC COVID-19 Data

The COVID-19 Case Surveillance Detailed Data (Centers for Disease Control and Prevention, 2020) is a national dataset provided by the CDC. It has the largest number of samples among the datasets considered, and contains 33 elements, with patient-level data including symptoms, demographics, and state of residence. The cohort consists of all lab-confirmed positive COVID-19 cases that were hospitalized, so the quantity of samples over time has a seasonality reflecting surges in COVID-19 (Figure 1). The outcome of interest is mortality, defined by death\_yn = Yes in the dataset. Performance over time is evaluated on a monthly basis. See Appendix D for more details.

<sup>2.</sup> In MIMIC-CXR, all labels except "No Finding" are considered positive for the purposes of Figure 1 and Table 1.

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Dataset name	Outcome	Time Range (time point unit)	# samples	# positives
SEER (Breast)	5-year Survival	1975-2013 (year)	462,023	378,758
SEER (Colon)	5-year Survival	1975-2013 (year)	$254,\!112$	$135,\!065$
SEER (Lung)	5-year Survival	1975 - 2013 (year)	$457,\!695$	49,997
CDC COVID-19	Mortality	Mar 2020–May 2022 (month)	$941,\!140$	190,786
SWPA COVID-19	90-day Mortality	Mar 2020–Feb 2022 (month)	$35,\!293$	1,516
MIMIC-IV	In-ICU Mortality	2009-2020 (year)	$53,\!050$	$3,\!334$
OPTN (Liver)	180-day Mortality	2005-2017 (year)	143,709	$4,\!635$
MIMIC-CXR	14 diagnostic labels	2010-2018 (year)	376,204	209,088

Table 1: Summary of datasets used for analysis. For more details, see Appendices C–G.



Figure 1: Number of samples and positive<sup>2</sup>outcomes per time point.

### 3.3. SWPA COVID-19 Data

The Southwestern Pennsylvania (SWPA) COVID-19 dataset consists of EHR data from patients tested for COVID-19. It is the smallest dataset considered in this paper, and was collected by a major healthcare provider in SWPA. Features include patient demographics, labs, problem histories, medications, inpatient vs. outpatient status, and other information collected in the patient encounter. The cohort consists of COVID-19 patients testing positive for the first time, and not already in the ICU or mechanically ventilated. Similar to the CDC COVID-19 dataset, there is a seasonality to the monthly number of samples that reflects surges in COVID-19 (Figure 1). The outcome of interest is 90-day mortality, derived by comparing the death date and test date. The performance over time is evaluated on a *monthly* basis. See Appendix  $\mathbf{E}$  for more details.

### 3.4. MIMIC-IV Critical Care Data

The Medical Information Mart for Intensive Care (MIMIC)-IV (Johnson et al., 2021) database contains EHR data from patients admitted to critical care units from 2008–2019. MIMIC-IV is an update to MIMIC-III, adding time annotations placing each sample into a three-year time range, and removing elements from the old CareVue EHR system (before 2008). We approximate the year of each sample by taking the midpoint of its time range, but note that this causes certain years (2009, 2012, 2015, 2018) to have substantially more samples than others (Figure 1). The cohort is selected by taking the first encounter of all patients in the icustays table, and the outcome of interest is in-ICU mortality. Performance over time is evaluated on a *yearly* basis. See Appendix **F** for more details.

### 3.5. OPTN Liver Transplant Data

The Organ Procurement and Transplantation Network (OPTN) database tracks organ donation and transplant events in the U.S. The selected cohort consists of liver transplant candidates on the waiting list. The same pipeline as Byrd et al. (2021) is used to extract the data, except that the first record is selected for each patient. The outcome of interest is 180-day mortality from when the patient was added to the list. The performance over time is evaluated on a *yearly* basis. More details are in Appendix G.

### 3.6. MIMIC-CXR

The MIMIC Chest X-ray (MIMIC-CXR) JPG dataset (Johnson et al., 2019b) contains chest radio-

graphs in JPG format. Similar to MIMIC-IV, we approximate the year by taking the midpoint of its three-year time range. The selected cohort consists of all radiographs from 2010 to 2018. The outcomes of interest are 14 diagnostic labels: Atelectasis, Cardiomegaly, Consolidation, Edema, Enlarged Cardiomediastinum, Fracture, Lung Lesion, Lung Opacity, Pleural Effusion, Pneumonia, Pneumothorax, Pleural Other, Support Devices, and No Finding. Performance over time is evaluated on a *yearly* basis. More details are in Appendix H.

## 4. Methods

We tackle the following guiding questions:

- 1. On each dataset, what would the reported performance of a model be if it were trained using standard time-agnostic splits (**all-period**)?
- 2. **Simulating** how a practitioner might have trained and deployed models in the past, how would performance have varied **over time**?
- 3. When might it be better to train on a **recent** window of data versus all historical data?
- 4. What is the comparative performance of different **classes of models** over time?
- 5. To what extent might we be able to diagnose possible **reasons** for changes in model performance?

### 4.1. All-period Training

We mimic common practice in evaluation by using time-agnostic data splits which randomly place patients from the entire study time range into train, validation, and test sets (details in Appendix L), and reporting the test set performance. We refer to training with this type of split as *all-period* training.

#### 4.2. EMDOT Evaluation

For more realistic simulation of how practitioners train models and subsequently deploy them on future data, we define the *Evaluation on Medical Datasets Over Time* (EMDOT) framework. At each time point t (termed simulated deployment date), an inperiod subset of data from times  $\leq t$  is available for model development. After training a model on this in-period data, one might be interested in both recent



Figure 2: EMDOT training regimes, with a simulated deployment date of t = 6.

in-period performance (at time t) and future *out-of-period* performance (at times > t).

In-period data is split into train, validation, and test sets (split ratios in Appendix L). For MIMIC-CXR, where one patient could have multiple radiographs, the data is split such that there are no overlapping patients between splits. Recent in-period performance is evaluated on held-out test data from the most recent time point. Out-of-period performance is evaluated on all data from each future time point. For example, a model trained up to time 6 is tested on data from 6, 7, 8, etc. (Figure 2). At time 8, the model is considered two time points stale. Although this procedure can take O(T) times more computation than all-period training for T time points, we argue that this procedure yields a more realistic view of the type of performance that one might expect models to have over time.

Additionally, practitioners face a tradeoff between using recent data perhaps most reflective of the present and using all available historical data for a larger sample size. Intuitively, the former may be appealing in modern applications with massive datasets, whereas the latter may be necessary in data-scarce applications. We explore these two training regimes, with different definitions of in-period data (Figure 2):

- 1. Sliding window: The last W time points are considered in-period. In this paper, we use window size W = 4 for sufficient positive examples.
- 2. All-historical: Any data prior to the current time point is considered in-period.

To decouple the effect of sample size from that of shifts in the data distribution, comparisons are also performed with all-historical data that is **sub**- **sampled** to be the same size as the corresponding training set under the sliding window training regime.

To summarize more formally, let  $D_t$  refer to the set of all data points occurring at time  $t \in \{1, ..., T\}$ , where T is the number of time points that the dataset spans. Each  $D_t$  can be partitioned by splitting patients at random into disjoint train, validation, and test sets:  $D_t = D_t^{\text{train}} \cup D_t^{\text{val}} \cup D_t^{\text{test}}$ . For simulated deployment dates  $t^* \in \{W, W + 1, ..., T\}$ , training, validation, and test sets are defined for the *sliding window* training regime as follows:

- training:  $\bigcup_{k=t^*-W+1}^{t^*} D_k^{\text{train}}$
- validation:  $\bigcup_{k=t^*-W+1}^{t^*} D_k^{\text{val}}$
- in-period test:  $D_{t^*}^{\text{test}}$
- out-of-period test:  $D_k$  for  $k = t^* + 1, ..., T$

Training, validation, and test sets are defined for the *all-historical* training regime as follows:

- training:  $\bigcup_{k=1}^{t^*} D_k^{\text{train}}$
- validation:  $\bigcup_{k=1}^{t^*} D_k^{\text{val}}$
- in-period test:  $D_{t^*}^{\text{test}}$
- out-of-period test:  $D_k$  for  $k = t^* + 1, ..., T$

At each simulated deployment date  $t^*$ , models are trained using the training set, validated using the validation set, and tested on the in-period test set as well as all out-of-period test sets. If a model with simulated deployment date  $t^*$  is being evaluated on an out of period test set  $D_{t^*+j}$ , then the model is j time points *stale*.

### 4.3. Evaluation Metrics

All binary classification tasks are evaluated by AU-ROC. For multi-label prediction in MIMIC-CXR, each of the 14 diagnostic labels is treated as a separate binary classification task, and a weighted sum of AUROCs is computed, where the weight for a particular label is given by the proportional prevalence of that label among all positive labels. That is, for some class a, its weight is  $p_a / \sum_x p_x$ , where  $p_x$  is the number of positives with label x. Samples are treated in an i.i.d. manner for training.

### 4.4. Models

Logistic regression (LR), gradient boosted decision trees (GBDT) and feedforward neural networks (MLP) are trained on the tabular datasets. DenseNet-121 is trained on the MIMIC-CXR imaging dataset. Hyperparameters are selected based on in-period validation performance, and the hyperparameter grids are in Appendix M.

### 4.5. Detecting Sources of Change

To better understand possible reasons for changing performance, we create *diagnostic plots* to track model performance alongside changes in the data distribution over time.

In tabular datasets, we plot feature importances and average values of the most important features over time. Generating these plots for logistic regression, we define feature importance by the magnitudes of the coefficients, but note that other feature importance techniques could be used for more complex model classes. To avoid overcrowding the plots, we take the union of the top k most important features from each time point is taken, where kis tuned depending on the dataset. We additionally highlight (using a thicker line) categorical features with consistently high prevalence or which experience a large change in prevalence across one time point, and numerical features with high average rank (see Appendix J for thresholds for each dataset).

For the imaging dataset, where feature importance is less straightforward, we plot the distribution of pixel intensities over time, along with proportions of each of the 14 diagnostic labels.

By highlighting sudden changes in model performance and the corresponding time periods in all other plots, diagnostic plots can help bring attention to shifts in the distribution of data that coincide with changing model performance.

### 4.6. EMDOT Python Package

We release the EMDOT python package<sup>3</sup> to help practitioners move from standard model evaluation to EMDOT evaluation. See Appendix B for a schematic of the EMDOT workflow, and see the GitHub repository for a step-by-step tutorial.

<sup>3.</sup> https://github.com/acmi-lab/EvaluationOverTime

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Model	SEER (Breast)	SEER (Colon)	$\begin{array}{c} \text{SEER} \\ (\text{Lung}) \end{array}$	CDC COVID-19	SWPA COVID-19	MIMIC- IV	OPTN (Liver)	MIMIC- CXR
LR	0.888	0.863	0.894	0.837	0.928	0.935	0.846	-
GBDT	0.891	0.868	0.894	0.851	0.930	0.931	0.854	-
MLP	0.891	0.869	0.898	0.852	0.928	0.898	0.847	-
DensetNet	-	-	-	-	-	-	-	0.860

Table 2: Test AUROC from all-period training and time-agnostic evaluation



Figure 3: Average test AUROC of logistic regression vs. time. Each solid line gives the performance of a model trained up to a simulated deployment time (marked by a dot), evaluated across future time points. Error bars are  $\pm$  standard deviation computed over 5 random splits. Red dotted line gives per-timepoint test performance of a model from all-period training (infeasible in reality, as it would involve training on data after the simulated deployment date).

### 5. Results

### 5.1. All-period Training

In standard time-agnostic evaluation, GBDT and MLP achieve the highest average test AUROC on all tabular datasets except MIMIC-IV (Table 2). Note however that LR often has comparable or only slightly lower AUROC than the more complex models. The top 10 coefficients of each LR with all-period training are in Appendices C-G, and the per-label AUROC of MIMIC-CXR is in Appendix Table 11. To form a baseline for comparison across time, we also evaluate the all-period models on subsets of the allperiod test data that belong to each year (red dotted line in Figure 3), but note that this type of training (on future data) is not feasible in deployment.

## 5.2. EMDOT Evaluation

Figure 3 plots the AUROC of LR for all tabular datasets (and DenseNet-121 for MIMIC-CXR) over time when using the all-historical training regime. Plots for GBDT and MLP are in Appendix K, along with plots for AUPRC. We mainly discuss AUROC, but note that AUPRC observes similar trends as in AUROC. One difference however is that the baseline AUPRC performance is given by the label prevalence (rather than a constant 0.5, as in AUROC), and so observed trends in label prevalence over time appear to influence trends in AUPRC (Appendix Figure 44).

For both AUROC and AUPRC, the reported test performance of a model from standard all-period training (red dotted line) mostly sits above the performance of any model that could have realistically been deployed by that date. Thus, all-period training tends to provide an over-optimistic estimate of performance upon deployment.

Across the datasets, a variety of trajectories of model performance are observed over time. In the SEER datasets, the AUROC of freshly trained models increases dramatically near 1988, but several of these models experience a large drop in AUROC around 2003 (Figure 3). Additionally, in-period test AUROCs tend to increase over time. By contrast, in CDC data, in-sample test AUROCs fluctuate up and down, and model performance over time varies more smoothly, appearing to loosely follow the insample performance. Models trained after December 2020 have a slight boost in AUROC, coinciding with a surge in cases (and hence sample size, Figure 1), however by January 2022 the in-sample AUROC decreases. In SWPA COVID-19, there is more variation and uncertainty in AUROC early in the pandemic, where sample sizes are small. In December 2020, sample sizes increase, and models seem to become more robust to changes over time. Finally, in the MIMIC-IV, MIMIC-CXR, and OPTN datasets, AUROC appears relatively stable across time.

#### 5.3. Training Regime Comparison

As the staleness of training data increases (i.e. as the test date gets further from the simulated deployment date), different training regimes can fare differently depending on the dataset (Figure 4, left).

In SEER (Breast) and SEER (Lung), sliding window is initially comparable to all-historical on fresh (low-staleness) data, but significantly underperforms both all-historical and all-historical (subsampled) when data are 8 to 22 years stale. At larger stalenesses, all training regimes start to become comparable. In CDC COVID-19, sliding window outperforms all-historical regardless of how stale the data is. By contrast, in SWPA COVID-19, which has the least amount of data (Table 1), both sliding window and all-historical (subsampled) underperform all-historical. In SEER (Colon), performance is relatively stable regardless of training regime. In MIMIC-IV, OPTN (Liver), and MIMIC-CXR, sliding window is on average comparable or slightly outperforms allhistorical when staleness is 0, but at nonzero stalenesses all-historical outperforms both sliding window and all-historical subsampled.



Figure 4: AUROC – AUROC<sub>LR\* all-historical</sub> vs. staleness. i.e., AUROC difference relative to a LR\* all-historical baseline across varying stalenesses of data,<sup>5</sup> for different training regimes (left) and model classes (right). Error bars are  $\pm$  std. dev. (\*in MIMIC-CXR, DenseNet-121 is used instead of LR)

<sup>5.</sup> Note: at the largest stalenesses, there are fewer simulated deployment dates being averaged over, and they must be early in the dataset. Here, the sliding window and allhistorical can be expected to perform similarly (especially when the sliding window is not much larger than or even matches the history). Since this is an artifact of finite time ranges, we gray out stalenesses where at least half of the all-historical data is the first sliding window of data.

### 5.4. Model Comparison

In SEER (Breast) and OPTN, GBDT outperforms both LR and MLP across the entire time range (Figure 4, right). In SEER (Colon), SEER (Lung), and CDC COVID-19, both GBDT and MLP initially outperform LR when staleness of the training data is less than 4 years, 4 years, and 7 months, respectively, however both eventually underperform LR as staleness increases further. While there is an uptick in GBDT performance on CDC COVID-19 towards 21month staleness, we note this data point is derived from less data than other points on the line because the data time range is finite. In the SWPA COVID-19 dataset, LR, MLP, and GBDT appear to perform comparably over time. In the MIMIC-IV dataset, LR performed best to begin with and remained the best.

### 5.5. Detecting Possible Sources of Change

Diagnostic plots for all datasets are in Appendix J. Here, we discuss SEER (Lung) (Figure 5) in detail as it has several interesting changes in model performance over time. In 1983, as EOD 4 features from the extent of disease coding schema are introduced (Figure 5, bottom right), a sudden jump in AUROC occurs (Figure 5, top and middle left). However, models trained at this time later experience a large AU-ROC drop (Figure 5, bottom left). By 1988, EOD 4 is phased out, and EOD 10 features are introduced. This coincides with another jump in AUROC, sustained until 2003 when the EOD 10 features are removed. In this dataset, the all-historical training regime seems more robust to changes over time, as all-historical models trained after 1988 avoid the drop that sliding window models undergo once their window excludes pre-1988 data (Figure 5, bottom left).

### 6. Discussion

Reported model performance from standard allperiod training tends to be over-optimistic (Figure 3) as models are evaluated on time points already seen in their training set (unrealistic in deployment settings). Thus, AUROCs reported from all-period training do not capture degradation that would have occurred in deployment.

Comparing model classes, in all datasets except MIMIC-IV, GBDT and MLP slightly outperform LR under standard time-agnostic evaluation (Appendix Table 2). However, evaluated across time, LR is



Figure 5: SEER (Lung) diagnostic plots. AUROC vs. time for sliding window (top-left) and all-historical subsampled (mid-left), max. drop in AUROC for each simulated deployment time (low-left), absolute feature coefficients for LR models from sliding window (top-right) and all-historical subsampled (mid-right) and prevalences of important features over time (low-right).

often comparable and even outperforms more complex models once enough time passes after the simulated deployment date. For example, MLP achieves the best AUROCs in SEER Breast, Colon, and Lung in standard time-agnostic evaluation (Table 2). However, in evaluation over time, LR had superior performance once some amount of time (30, 5, 4 years respectively) had passed (Figure 4, right). In most datasets GBDT appears more robust over time than MLP, however as the training data becomes more stale it tends to become comparable to LR (in all datasets except OPTN Liver and SEER Breast, GBDT dipped below the performance of LR for several stalenesses). Thus, although complex model classes may appear to outperform simpler linear model classes in standard time-agnostic evaluation, one should consider performance over time when selecting a model class for deployment. As demonstrated by the different relative performances of model classes when evaluated over time versus in a time-agnostic manner, EMDOT can serve as a helpful stress-test to combat under-specification.

Regarding training regimes, we find that with increasing stalenesses, all-historical appears more reliable than sliding window across all datasets except for CDC COVID-19 (Figure 4, left). In SWPA COVID-19, MIMIC-IV, OPTN (Liver), and MIMIC-CXR, the benefit of all-historical data likely comes from the increased sample size, as subsampling allhistorical data to be the same size as the corresponding sliding window resulted in comparable performance to sliding window. In the SEER datasets, the effect of sample size is less pronounced, as sliding window and subsampled all-historical are frequently comparable to all-historical. There are certain stalenesses for which sliding window underperforms allhistorical, which may be due to the addition and removal of features. If the sliding window model learns to rely on recently added features which are later removed, this could result in drops in performance whereas an all-historical model which had learned to predict without the presence of such features would be more robust to such changes. On the other hand, in CDC COVID-19 (the setting with the most data and fewest features), subsampled all-historical performs comparably to all-historical, and sliding window outperforms both across all stalenesses (Figure 4, left). This suggests that the performance of LR may have been saturated even when a sub-sample of all-historical data was used, and the benefit of using more recent data outweighs the larger sample size afforded by all-historical. More broadly, in rapidly evolving environments with simple models, few features, and large quantities of data, the sliding window training regime could be advantageous.

The SEER datasets had dramatic changes in data distribution in both 1988 and 2003, when important features were added and/or removed (Figure 5). One possible reason for the robustness of all-historical models in this dataset is that after 2003, when features like EOD 10 were removed, the model could still rely on features that were introduced prior to the use of EOD 10 in 1988. More broadly, we hypothesize that if a model was trained on a mixture of distributions that occurred throughout the past, it may be better equipped to handle shifts to settings similar to those distributions in the future.

While the SEER datasets and COVID-19 datasets displayed several changes in model performance over time, the OPTN and MIMIC datasets had relatively stable behavior. One possible reason for this is that the outcomes or diseases of interest were relatively stable in nature, we did not observe any substantial changes in the distribution of data. Another is that in the MIMIC datasets, a three-year range was given for each sample rather than a specific date. This uncertainty around the date, along with the limited number of date ranges, could result in a smoothing effect on the resulting estimates of performance.

In conclusion, EMDOT not only yields insights into the suitability of different model classes or training regimes for deployment, but also helps one detect distribution shifts that occurred in the past. Understanding such shifts may help practitioners be prepared for shifts of a similar nature in the future. Although the EMDOT framework does require additional computational time than the standard timeagnostic evaluation setup, we argue that the insights that could be gained from this procedure are worthwhile, especially before deployment in high-stakes settings.

Limitations and Future Work One possible reservation that users might have about using EM-DOT is that it could involve training up to T times as many models as would normally be required (where T is number of timepoints). To help alleviate this concern, in future work we plan to implement parallelization in EMDOT. For noisier estimates of model performance in less time, one could also subsample the dataset. Another interesting extension is exploring performance over time in other data modalities (e.g. time series, natural language, etc.). Depending on the complexity of models used in these modalities. this may require additional computational resources. More broadly, we hope that others may also build upon EMDOT to shine new light on how models and methodologies fare when evaluated with an eye towards deployment.

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## Appendix A. Snapshot into the State of ML4H Model Evaluation

To get a snapshot of the current standards for model evaluation in machine learning for healthcare research, we manually reviewed all of the papers from the CHIL 2022 proceedings, the first 20 papers in the CHIL 2021 proceedings, and the first 20 papers that came up in the Radiology medical journal when searching for the keyword "machine learning" and filtering for papers from 2022 to 2023 (see README.md in https://github.com/acmi-lab/EvaluationOverTime). Out of 23 papers in the CHIL 2022 proceedings, 21 did not take time into account in their data split, and two were unclear about how they split data, but it is unlikely that they split by time. Out of the 20 papers reviewed at CHIL 2021, only one paper split by time. Out of the 20 papers reviewed at critical and the remaining models, but out of the remaining 14 papers, 13 did not take time into account in their data split.

## Appendix B. EMDOT Python Package

Figure 6 illustrates the workflow of the EMDOT Python package.



Figure 6: EMDOT Python package workflow diagram. The primary touchpoint of the EMDOT package is the EotExperiment object. Users provide a dataframe for their (mostly) preprocessed dataset (EMDOT takes care of normalization based on the relevant training set), their desired experiment configuration (e.g. sliding window), and model class (which should subclass the simple EotModel abstract class) in order to create an EotExperiment object. Running the run\_experiment() function of the EotExperiment returns a dataframe of experiment results that can then be visualized. The diagram also provides insight into some of the internals of the EotExperiment object – there is an EotDataset object that handles data splits, and an EotEvaluator object that executes the main evaluation loop.

## Appendix C. Additional SEER Data Details

The Surveillance, Epidemiology, and End Results (SEER) Program collects cancer incidence data from registries throughout the U.S. This data has been used to study survival in several forms of cancer (Choi et al., 2008; Fuller et al., 2007; Taioli et al., 2015; Hegselmann et al., 2018). Each case includes demographics, primary tumor site, tumor morphology, stage and diagnosis, first course of treatment, and survival outcomes (collected with follow-up) (National Cancer Institute, 2020). The performance over time is evaluated on a *yearly* basis. We use the November 2020 version of the SEER database with nine registries (SEER 9), which covers about 9.4% of the U.S. population. While there are SEER databases that aggregate over more registries and hence cover a greater proportion of the U.S. population, we choose SEER 9 due to the large time range it covers (1975–2018).

- Data access: After filling out a Data Use Agreement and Best Practices Agreement, individuals can easily request access to the SEER dataset.
- Cohort selection: Using the SEER\*Stat software (Program, 2015), we define three cohorts of interest: (1) breast cancer, (2) colon cancer, and (3) lung cancer. We primarily follow the cohort selection procedure from Hegselmann et al. (2018), but we use SEER 9 instead of SEER 18, and use data from all available years instead of limiting to 2004–2009. Cohort selection diagrams are given in Figures 7, 8, and 9. If there are multiple samples per patient, we filter to the first entry per patient, which corresponds to when a patient first enters the dataset. This corresponds to a particular interpretation of the prediction: when a patient is first added to a cancer registry, given what we know about that patient, what is their estimated 5-year survival probability?
- Cohort characteristics: Summaries of the SEER (Breast), SEER (Colon), and SEER (Lung) cohort characteristics are in Tables 3, 4, and 5.
- Outcome definition: 5-year survival is defined by a confirmation that the patient is alive five years after the year of diagnosis.
- Features: We list the features used in the SEER breast, colon, and lung cancer datasets in Section C.2. For all datasets, we convert all categorical variables into dummy features, and apply standard scaling to numerical variables (subtract mean and divide by standard deviation).
- Missingness heat maps: are given in Figures 10, 11, 12, 13, 14, and 15.

## C.1. Cohort Selection and Cohort Characteristics



Figure 7: Cohort selection diagram - SEER (Breast)



Figure 8: Cohort selection diagram - SEER (Colon)



Figure 9: Cohort selection diagram - SEER (Lung)

Characteristic		Missingness	Type
Sex			
Female	459.184~(99.4%)	_	categorical
Male	2,839(0.6%)	_	categorical
Age recode with single ages and $85+$	60(50-71)	0.0%	continuous
Race/ethnicity		/ •	
White	387.247(83.8%)	_	categorical
Black	40.217 (8.7%)	_	categorical
Other	34.559(7.5%)	_	categorical
Laterality			0
Right - origin of primary	224.777(48.7%)	_	categorical
Left - origin of primary	233.549(50.5%)	_	categorical
Other	3.697(0.8%)	_	categorical
Regional nodes positive $(1988+)$	0 (0-3)	21.0%	continuous
T value - based on AJCC 3rd (1988-2003)	10(10-20)	56.2%	categorical
Derived AJCC T, 7th ed (2010-2015)	13 (13-20)	85.3%	categorical
CS site-specific factor 3 (2004-2017 varying by schema)	0 (0-2)	64.8%	categorical
Regional nodes examined (1988+)	8 (2-15)	21.0%	continuous
Coding system-EOD (1973-2003)	( )		
Four-digit EOD (1983-1987)	44,066~(9.5%)	_	categorical
Ten-digit EOD (1988-2003)	202,450(43.8%)	_	categorical
Thirteen-digit (expanded) site specific EOD (1973-1982)	52,742 (11.4%)	_	categorical
Blank(s)	162,765(35.2%)	_	categorical
CS version input original (2004-2015)	10,401 (10,300-20,302)	64.8%	categorical
CS version input current (2004-2015)	20,520 (20,510-20,540)	64.8%	categorical
EOD 10 - extent (1988-2003)	10 (10-13)	56.2%	categorical
Grade (thru 2017)			
Unknown	130,713 (28.3%)	—	categorical
Moderately differentiated; Grade II	135,970~(29.4%)	_	categorical
Poorly differentiated; Grade III	$119,900 \ (26.0\%)$	_	categorical
Undifferentiated; anaplastic; Grade IV	8,081 $(1.7%)$	_	categorical
Well differentiated; Grade I	67,359~(14.6%)	-	categorical
<b>SEER historic stage A</b> $(1973-2015)$			
Regional	136,207~(29.5%)	_	categorical
Localized	286,927~(62.1%)	—	categorical
Unstaged	9,242~(2.0%)	_	categorical
Distant	29,647~(6.4%)	_	categorical
IHS Link			
Record sent for linkage, no IHS match	409,058 (88.5%)	-	categorical
Record sent for linkage, IHS match	$1,505\ (0.3\%)$	-	categorical
Blank(s)	$51,460\ (11.1\%)$	-	categorical
Histologic Type ICD-O-3	8,500 (8,500-8,500)	0.0%	categorical
EOD 10 - size (1988-2003)	18(10-30)	56.2%	categorical
Type of Reporting Source			
Hospital inpatient/outpatient or clinic	450,801 (97.6%)	—	categorical
Other CDDD	11,222 (2.4%)	—	categorical
SEER cause-specific death classification			
Anve or dead of other cause	3(8, 158 (82.0%))	—	categorical
Dead (attributable to this cancer dx)	83,203(18.0%)	-	categorical
Survival months	133 (74-220)	0.0%	categorical
o-year survival	278 758 (00 007)		antogenical
1 0	83 265 (18 0%)	_	categorical
	03,203 (10.070)	_	categorical

Table 3: SEER (Breast) cohort characteristics, with count (%) or median (Q1 – Q3).

Sex         -         categorical           Male         120,451 (47.4%)         -         categorical           Age recode with single ages and 85+         70 (61-79)         0.0%         continuous           Race recode (White, Black, Other)         -         categorical         0.0%         continuous           White         212,265 (83.5%)         -         categorical         0.0%         categorical           Other         17,806 (7.0%)         -         categorical         0.0%         categorical           Derived AJCC T, 6th ed (2004-2015)         20,510 (20,510-20,540)         72.8%         categorical           Histology ICD-0-2         8,140 (8,140-8,210)         0.0%         categorical           Record sent for linkage, no IHS match         744 (0.3%)         -         categorical           Blank(s)         44,566 (17.5%)         -         categorical           Histology trecode - broad groupings         8,797 (3.5%)         -         categorical           Blank(s)         10-10         29.8%         continuous           CS mests at (x (2004-2015)         0 (0-22)         7.8%         continuous           Regorinal nodes positive (1988+)         1 (0-10)         29.8%         continuous           CS mests at (x	Characteristic		Missingness	Type
Female       133.661 (52.6%)       -       categorical         Male       120.451 (47.4%)       -       categorical         Age recode with single ages and 85+       70 (61-79)       0.0%       continuous         Race recode (White, Black, Other)       212.265 (83.5%)       -       categorical         White       212.265 (83.5%)       -       categorical         Other       17.806 (7.0%)       -       categorical         CS version input current (2004-2015)       30 (20-40)       73.3%       categorical         Histology ICD-0-2       8,140 (8,140.8,210)       0.0%       categorical         Record sent for linkage, no IIIS match       208.802 (82.2%)       -       categorical         Black(s)       44,566 (17.5%)       -       categorical         8140-839: adenomas and adenocarinomas       213.193 (83.9%)       -       categorical         8440-8499: cystic, mucinous and serous neoplasms       8,797 (3.5%)       -       categorical         8100-8049: epithelial neoplasms, NOS       8,797 (3.5%)       -       categorical         Record sent for linkage, not sent sent set dx (2004-2015)       0 (0-22)       72.8%       continuous         CS mets at dx (2004-2015)       0 (0-22)       72.8%       continuous <tr< td=""><td>Sex</td><td></td><td></td><td></td></tr<>	Sex			
Male         120.451 (47.4%)         -         categorical           Age recode with single ages and 85+         70 (61-79)         0.0%         continuous           Race recode (White, Black, Other)         212.265 (83.5%)         -         categorical           White         212.265 (83.5%)         -         categorical           Other         17.806 (7.0%)         -         categorical           Other         17.806 (7.0%)         -         categorical           Histology ICD-O-2         8,140 (8,140-8,210)         0.0%         categorical           Record sent for linkage, no IHS match         208,802 (82.2%)         -         categorical           Blank(s)         44,566 (17.5%)         -         categorical           Blank(s)         44,566 (17.5%)         -         categorical           Blank(s)         44,566 (17.5%)         -         categorical           St40-8389: adenomas and adenocarcinomas         213,193 (83.9%)         -         categorical           Chieronecode         13,003 (5.15%)         -         categorical           Other         3,865 (1.5%)         -         categorical           Statues copic continuous         3,865 (1.5%)         -         categorical           Other	Female	133,661 (52.6%)	_	categorical
Age recode with single ages and 85+70 (61-79)0.0%continuousRace recode (White, Black, Other)212.265 (83.5%)-categoricalWhite212.265 (83.5%)-categoricalOther17.806 (7.0%)-categoricalCS version input current (2004-2015)20.510 (20.510-20.540)72.8%categoricalHistology ICD-0-28.140 (8.140-8.210)0.0%categoricalHistology ICD-0-28.140 (8.140-8.210)0.0%categoricalBlack208.802 (82.2%)-categoricalRecord sent for linkage, no IIIS match208.802 (82.2%)-categoricalBlack(s)44.6566 (17.5%)-categorical8140-8399: cystic, mucinous and serous neoplasms28.257 (11.1%)-categorical8140-8399: cystic, mucinous and serous neoplasms8.797 (3.5%)-categorical8010-8049: epithelial neoplasms, NOS8.797 (3.5%)-categoricalRegional nodes positive (1988+)1 (0-10)29.8%continuousCS mets at dx (2004-2015)30 (20-40)73.3%categoricalNot recommended13.003 (5.1%)-categoricalOther17.180 (6.8%)-categoricalDerived AJCC T, 6th ed (2004-2015)30 (20-40)73.3%categoricalCS wersion input original (2004-2015)10.401 (10.300-20.302)72.8%categoricalOther17.180 (6.8%)-categoricalcategoricalDerived AJCC T, 6th ed (2004-2015)30 (20-40)73.3% <t< td=""><td>Male</td><td>120,451 (47.4%)</td><td>_</td><td>categorical</td></t<>	Male	120,451 (47.4%)	_	categorical
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$\begin{array}{c} {\rm CS\ mets\ at\ dx\ (2004-2015)} & 0\ (0-22) & 72.8\% & {\rm continuous} \\ {\rm Reason\ no\ cancer-directed\ surgery} \\ {\rm Surgery\ performed\ } & 223,929\ (88.1\%) & - & {\rm categorical\ } \\ {\rm Not\ recommended\ } & 13,003\ (5.1\%) & - & {\rm categorical\ } \\ {\rm Other\ } & 17,180\ (6.8\%) & - & {\rm categorical\ } \\ {\rm Derived\ AJCC\ T,\ 6th\ ed\ (2004-2015)\ } & 30\ (20-40)\ 73.3\% & {\rm categorical\ } \\ {\rm Derived\ AJCC\ T,\ 6th\ ed\ (2004-2015)\ } & 10,401\ (10,300-20,302)\ 72.8\% & {\rm categorical\ } \\ {\rm Primary\ Site\ } & 184\ (182-187)\ & 0.0\% & {\rm categorical\ } \\ {\rm Diagnostic\ Confirmation\ } & \\ {\rm Positiv\ histology\ } & 244,616\ (96.3\%)\ & - & {\rm categorical\ } \\ {\rm Boy} & {\rm Confirmation\ } & \\ {\rm Positiv\ histology\ } & 244,616\ (96.3\%)\ & - & {\rm categorical\ } \\ {\rm Coller\ } & 4,674\ (1.8\%)\ & - & {\rm categorical\ } \\ {\rm Coller\ } & 4,674\ (1.8\%)\ & - & {\rm categorical\ } \\ {\rm EOD\ 10\ -\ extent\ (1988-2003)\ } & 45\ (40-85)\ 57.0\% & {\rm categorical\ } \\ {\rm Histologic\ Type\ ICD-O-3\ } & 8,140\ (8,140-8,210)\ & 0.0\% & {\rm categorical\ } \\ {\rm EOD\ 10\ -\ size\ (1988-2003)\ } & 55\ (35-999)\ 57.0\% & {\rm categorical\ } \\ {\rm Supph\ nodes\ (2004-2015)\ } & 0\ (0-210)\ & 72.8\% & {\rm categorical\ } \\ {\rm Supph\ nodes\ (2004-2015)\ } & 0\ (0-210)\ & 72.8\% & {\rm categorical\ } \\ {\rm Supph\ nodes\ (2004-2015)\ } & 0\ (0-210)\ & 72.8\% & {\rm categorical\ } \\ {\rm Supph\ nodes\ (2004-2015)\ } & 0\ (0-210)\ & 72.8\% & {\rm categorical\ } \\ {\rm Supph\ nodes\ (2004-2015)\ } & 0\ (0-210)\ & 72.8\% & {\rm categorical\ } \\ {\rm Supph\ nodes\ (2004-2015)\ } & 0\ (0-210)\ & 72.8\% & {\rm categorical\ } \\ {\rm Supph\ nodes\ (2004-2015)\ } & 0\ (0-210)\ & 72.8\% & {\rm categorical\ } \\ {\rm Supph\ nodes\ (2004-2015)\ } & 0\ (0-210)\ & 72.8\% & {\rm categorical\ } \\ {\rm Supph\ nodes\ (2004-2015)\ } & 0\ (0\ -210)\ & 72.8\% & {\rm categorical\ } \\ {\rm Supph\ nodes\ (2004-2015)\ } & 0\ (0\ -210)\ & 72.8\% & {\rm categorical\ } \\ {\rm Supph\ nodes\ (2004-2015)\ } & 0\ (0\ -210)\ & 72.8\% & {\rm categorical\ } \\ {\rm Suph\ nodes\ (2004-2015)\ } & 0\ & 0\ & 0\ & 0\ & 0\ & 0\ & 0\$	Regional nodes positive (1988+)	1 (0-10)	29.8%	continuous
Reason no cancer-directed surgery $223,929$ (88.1%) $-$ categoricalNot recommended13,003 (5.1%) $-$ categoricalOther17,180 (6.8%) $-$ categoricalDerived AJCC T, 6th ed (2004-2015)30 (20-40)73.3%categoricalCS version input original (2004-2015)10,401 (10,300-20,302)72.8%categoricalPrimary Site184 (182-187)0.0%categoricalDiagnostic Confirmation $-$ categoricalPositive histology244,616 (96.3%) $-$ categoricalRadiography without microscopic confirm $4,822$ (1.9%) $-$ categoricalOther4,674 (1.8%) $-$ categoricalEOD 10 - extent (1988-2003)45 (40-85)57.0%categoricalHistologic Type ICD-O-38,140 (8,140-8,210)0.0%categoricalCS lymph nodes (2004-2015)0 (0-210)72.8%categoricalDead (attributable to this cancer dx)119,047 (46.8%) $-$ categoricalAlive or dead of other cause135,065 (53.2%) $-$ categoricalSurvival months68 (12-151)0.0%categoricalJ135,065 (53.2%) $-$ categoricalJ135,065 (53.2%) $-$ categoricalJ135,065 (53.2%) $-$ categoricalJ135,065 (53.2%) $-$ categoricalJ119,047 (46.8%) $-$ categoricalJ135,065 (53.2%) $-$ categoricalJ10119,047 (46.8%) $-$	CS mets at dx (2004-2015)	0 (0-22)	72.8%	continuous
$\begin{array}{cccccc} Surgery performed & 223,929 (88.1\%) & - & categorical \\ Not recommended & 13,003 (5.1\%) & - & categorical \\ Other & 17,180 (6.8\%) & - & categorical \\ \hline Derived AJCC T, 6th ed (2004-2015) & 30 (20-40) & 73.3\% & categorical \\ CS version input original (2004-2015) & 10,401 (10,300-20,302) & 72.8\% & categorical \\ \hline Diagnostic Confirmation & & & \\ Positive histology & 244,616 (96.3\%) & - & categorical \\ Radiography without microscopic confirm & 4,822 (1.9\%) & - & categorical \\ Other & 4,674 (1.8\%) & - & categorical \\ \hline Diagnostic Type ICD-O-3 & 8,140 (8,140-8,210) & 0.0\% & categorical \\ \hline EOD 10 - extent (1988-2003) & 45 (40-85) & 57.0\% & categorical \\ \hline EOD 10 - size (1988-2003) & 55 (35-999) & 57.0\% & categorical \\ \hline SEER cause-specific death classification \\ Dead (attributable to this cancer dx) & 119,047 (46.8\%) & - & categorical \\ \hline Alive or dead of other cause & 135,065 (53.2\%) & - & categorical \\ \hline Survival months & 68 (12-151) & 0.0\% & categorical \\ \hline 1 & 135,065 (53.2\%) & - & categorical \\ 0 & 119,047 (46.8\%) & - & categorical \\ \hline 1 & 0.0\% &$	Reason no cancer-directed surgery	· · · ·		
Not recommended Other $13,003$ ( $5.1\%$ )-categorical categoricalDerived AJCC T, 6th ed (2004-2015) $30$ ( $20-40$ ) $73.3\%$ categoricalDerived AJCC T, 6th ed (2004-2015) $30$ ( $20-40$ ) $73.3\%$ categoricalCS version input original (2004-2015) $10,401$ ( $10,300-20,302$ ) $72.8\%$ categoricalPrimary Site $184$ ( $182-187$ ) $0.0\%$ categoricalDiagnostic Confirmation-categoricalPositive histology $244,616$ ( $96.3\%$ )-categoricalOther $4,674$ ( $1.8\%$ )-categoricalOther $4,674$ ( $1.8\%$ )-categoricalEOD 10 - extent ( $1988-2003$ ) $45$ ( $40-85$ ) $57.0\%$ categoricalHistologic Type ICD-O-3 $8,140$ ( $8,140-8,210$ ) $0.0\%$ categoricalCS lymph nodes ( $2004-2015$ ) $0$ ( $0-210$ ) $72.8\%$ categoricalSEER cause-specific death classification-categoricalDead (attributable to this cancer dx) $119,047$ ( $46.8\%$ )-categoricalAlive or dead of other cause $135,065$ ( $53.2\%$ )-categoricalSurvival months $68$ ( $12-151$ ) $0.0\%$ categorical5-year survival1 $135,065$ ( $53.2\%$ )-categorical0 $119,047$ ( $46.8\%$ )-categorical0 $119,047$ ( $46.8\%$ )-categorical	Surgery performed	223,929 (88.1%)	_	categorical
$\begin{array}{c ccccccccccccccccccccccccccccccccccc$	Not recommended	13,003(5.1%)	_	categorical
$\begin{array}{c c c c c c c c c c c c c c c c c c c $	Other	17,180(6.8%)	_	categorical
$\begin{array}{c} {\rm CS \ version \ input \ original \ (2004-2015)} \\ {\rm Primary \ Site} \\ {\rm Diagnostic \ Confirmation} \\ {\rm Positive \ histology} \\ {\rm Radiography \ without \ microscopic \ confirm} \\ {\rm Other} \\ {\rm EOD \ 10 \ - \ extent \ (1988-2003)} \\ {\rm Histologic \ Type \ ICD-O-3 \\ {\rm EOD \ 10 \ - \ size \ (1988-2003)} \\ {\rm EOD \ 10 \ - \ size \ (1988-2003)} \\ {\rm EOD \ 10 \ - \ size \ (1988-2003)} \\ {\rm SEER \ cause-specific \ death \ classification \\ {\rm Dead \ (attributable \ to \ this \ cancer \ dx) \\ {\rm Alive \ or \ dead \ of \ other \ cause} \\ {\rm SEER \ cause-specific \ death \ classification \\ {\rm Dead \ (attributable \ to \ this \ cancer \ dx) \\ {\rm Alive \ or \ dead \ of \ other \ cause} \\ \\ {\rm Survival \ months} \\ {\rm Survival \ months} \\ {\rm Survival \ months} \\ {\rm CS \ 100 \ categorical \\ {\rm CS \ 100 \ or \ categorical \ {\rm CS \ 100 \ or \ categorical \\ {\rm CS \ 100 \ or \ categorical \ {\rm CS \ 100 \ or \ 100 \ or \ categorical \\ {\rm CS \ 100 \ or \ 100 \ or \ categorical \ {\rm CS \ 100 \ or \ 100 \ $	Derived AJCC T, 6th ed (2004-2015)	30 (20-40)	73.3%	categorical
Primary Site $184 (182-187)$ $0.0\%$ categoricalDiagnostic Confirmation $244,616 (96.3\%)$ $-$ categoricalPositive histology $244,616 (96.3\%)$ $-$ categoricalRadiography without microscopic confirm $4,822 (1.9\%)$ $-$ categoricalOther $4,674 (1.8\%)$ $-$ categoricalEOD 10 - extent (1988-2003) $45 (40-85)$ $57.0\%$ categoricalHistologic Type ICD-O-3 $8,140 (8,140-8,210)$ $0.0\%$ categoricalEOD 10 - size (1988-2003) $55 (35-999)$ $57.0\%$ categoricalCS lymph nodes (2004-2015) $0 (0-210)$ $72.8\%$ categoricalSEER cause-specific death classification $119,047 (46.8\%)$ $-$ categoricalMive or dead of other cause $135,065 (53.2\%)$ $-$ categoricalSurvival months $68 (12-151)$ $0.0\%$ categorical $0$ $119,047 (46.8\%)$ $-$ categorical $0$ $119,047 (46.8\%)$ $-$ categorical $0$ $0.0\%$ $0.0\%$ $0$ $0.0\%$ $0.0\%$ $0$ $0.0\%$ $0.0\%$ $0$ $0.0\%$ $0.0\%$ $0$ $0.0\%$ $0.0\%$ $0$ $0.0\%$ $0.0\%$ $0$ $0.0\%$ $0.0\%$ $0$ $0.0\%$ $0.0\%$ $0$ $0.0\%$ $0.0\%$ <t< td=""><td>CS version input original (2004-2015)</td><td>10,401 (10,300-20,302)</td><td>72.8%</td><td>categorical</td></t<>	CS version input original (2004-2015)	10,401 (10,300-20,302)	72.8%	categorical
$\begin{array}{c c c c c c c c c c c c c c c c c c c $	Primary Site	184 (182-187)	0.0%	categorical
$\begin{array}{cccccc} \mbox{Positive histology} & 244,616 (96.3\%) & - & categorical \\ \mbox{Radiography without microscopic confirm} & 4,822 (1.9\%) & - & categorical \\ \mbox{Other} & 4,674 (1.8\%) & - & categorical \\ \mbox{EOD 10 - extent (1988-2003)} & 45 (40-85) & 57.0\% & categorical \\ \mbox{Histologic Type ICD-O-3} & 8,140 (8,140-8,210) & 0.0\% & categorical \\ \mbox{EOD 10 - size (1988-2003)} & 55 (35-999) & 57.0\% & categorical \\ \mbox{CS lymph nodes (2004-2015)} & 0 (0-210) & 72.8\% & categorical \\ \mbox{SEER cause-specific death classification} & & & \\ \mbox{Dead (attributable to this cancer dx)} & 119,047 (46.8\%) & - & categorical \\ \mbox{Survival months} & 68 (12-151) & 0.0\% & categorical \\ \mbox{Survival months} & 68 (12-151) & 0.0\% & categorical \\ \mbox{5-year survival} & & \\ \mbox{119,047 (46.8\%)} & - & categorical \\ 119,047 (4$	Diagnostic Confirmation			-
Radiography without microscopic confirm Other $4,822 (1.9\%)$ $-$ categoricalConterner $4,674 (1.8\%)$ $-$ categoricalEOD 10 - extent (1988-2003) $45 (40-85)$ $57.0\%$ categoricalHistologic Type ICD-O-3 $8,140 (8,140-8,210)$ $0.0\%$ categoricalEOD 10 - size (1988-2003) $55 (35-999)$ $57.0\%$ categoricalCS lymph nodes (2004-2015) $0 (0-210)$ $72.8\%$ categoricalSEER cause-specific death classification $0 (0-210)$ $72.8\%$ categoricalMive or dead of other cause $119,047 (46.8\%)$ $-$ categoricalSurvival months $68 (12-151)$ $0.0\%$ categorical5-year survival $1135,065 (53.2\%)$ $-$ categorical1 $135,065 (53.2\%)$ $-$ categorical0 $119,047 (46.8\%)$ $-$ categorical	Positive histology	244,616 (96.3%)	_	categorical
$\begin{array}{cccccccc} {\rm Other} & 4,674 (1.8\%) & - & {\rm categorical} \\ {\rm EOD \ 10 - extent \ (1988-2003)} & 45 \ (40\text{-}85) & 57.0\% & {\rm categorical} \\ {\rm Histologic \ Type \ ICD-O-3} & 8,140 \ (8,140\text{-}8,210) & 0.0\% & {\rm categorical} \\ {\rm EOD \ 10 - size \ (1988-2003)} & 55 \ (35\text{-}999) & 57.0\% & {\rm categorical} \\ {\rm CS \ lymph \ nodes \ (2004-2015)} & 0 \ (0\text{-}210) & 72.8\% & {\rm categorical} \\ {\rm SEER \ cause-specific \ death \ classification} & & \\ {\rm Dead \ (attributable \ to \ this \ cancer \ dx)} & 119,047 \ (46.8\%) & - & {\rm categorical} \\ {\rm Survival \ months} & 68 \ (12\text{-}151) & 0.0\% & {\rm categorical} \\ {\rm Survival \ months} & 68 \ (12\text{-}151) & 0.0\% & {\rm categorical} \\ {\rm J} & 135,065 \ (53.2\%) & - & {\rm categorical} \\ {\rm J} & 135,065 \ (53.2\%) & - & {\rm categorical} \\ {\rm J} & 135,065 \ (53.2\%) & - & {\rm categorical} \\ {\rm J} & 10,047 \ (46.8\%) & - & {\rm categorical} \\ {\rm J} & 10,047 \ (46.8\%) & - & {\rm categorical} \\ {\rm J} & 10,047 \ (46.8\%) & - & {\rm categorical} \\ {\rm J} & 10,047 \ (46.8\%) & - & {\rm categorical} \\ {\rm J} & 10,047 \ (46.8\%) & - & {\rm categorical} \\ {\rm J} & {\rm J} & {\rm J} & {\rm J} & {\rm J} \\ {\rm J} & {\rm J} \\ {\rm J} & {\rm J} & {\rm J} & {\rm J} & {\rm J} \\ {\rm J} & {\rm J} & {\rm J} & {\rm J} & {\rm J} \\ {\rm J} & {\rm J} & {\rm J} & {\rm J} & {\rm J} \\ {\rm J} & {\rm J} & {\rm J} & {\rm J} & {\rm J} \\ {\rm J} & {\rm J} & {\rm J} & {\rm J} \\ {\rm J} & {\rm J} & {\rm J} & {\rm J} \\ {\rm J} & {\rm J} & {\rm J} & {\rm J} \\ {\rm J} & {\rm J} & {\rm J} \\ {\rm J} & {\rm J} & {\rm J} \\ {\rm J} & {\rm J} & {\rm J} \\ {\rm J} & {\rm J} & {\rm J} \\ {\rm J} & {\rm J} & {\rm J} \\ {\rm J} & {\rm J} & {\rm J} \\ {\rm J} & {\rm J} \\ {\rm J} & {\rm J} & {\rm J} \\ {\rm J} \\ {\rm J} & {\rm J} \\ {\rm J} & {\rm J} \\ {\rm J} & {\rm J} \\ $	Radiography without microscopic confirm	4,822 (1.9%)	_	categorical
$\begin{array}{c ccccccccccccccccccccccccccccccccccc$	Other	4,674 $(1.8%)$	_	categorical
Histologic Type ICD-O-3 $8,140 (8,140-8,210)$ $0.0\%$ categoricalEOD 10 - size (1988-2003) $55 (35-999)$ $57.0\%$ categoricalCS lymph nodes (2004-2015) $0 (0-210)$ $72.8\%$ categoricalSEER cause-specific death classification $0 (0-210)$ $72.8\%$ categoricalDead (attributable to this cancer dx) $119,047 (46.8\%)$ $-$ categoricalAlive or dead of other cause $135,065 (53.2\%)$ $-$ categoricalSurvival months $68 (12-151)$ $0.0\%$ categorical5-year survival $135,065 (53.2\%)$ $-$ categorical1 $135,065 (53.2\%)$ $-$ categorical0 $119,047 (46.8\%)$ $-$ categorical	EOD 10 - extent (1988-2003)	45 (40-85)	57.0%	categorical
$\begin{array}{c c c c c c c c c c c c c c c c c c c $	Histologic Type ICD-O-3	8,140 (8,140-8,210)	0.0%	categorical
$\begin{array}{c c c c c c c c c c c c c c c c c c c $	EOD 10 - size (1988-2003)	55 (35-999)	57.0%	categorical
SEER cause-specific death classification $-$ categoricalDead (attributable to this cancer dx) $119,047$ (46.8%) $-$ categoricalAlive or dead of other cause $135,065$ (53.2%) $-$ categoricalSurvival months $68$ (12-151) $0.0\%$ 5-year survival $-$ categorical1 $135,065$ (53.2%) $-$ categorical0 $119,047$ (46.8%) $-$ categorical	CS lymph nodes (2004-2015)	0 (0-210)	72.8%	categorical
$\begin{array}{cccc} \mbox{Dead} \mbox{(attributable to this cancer dx)} & 119,047 \ (46.8\%) & - & categorical \\ \mbox{Alive or dead of other cause} & 135,065 \ (53.2\%) & - & categorical \\ \mbox{Survival months} & 68 \ (12-151) & 0.0\% & categorical \\ \mbox{5-year survival} & & & & \\ \mbox{1} & 135,065 \ (53.2\%) & - & categorical \\ \mbox{0} & 119,047 \ (46.8\%) & - & categorical \\ \mbox{0} $	SEER cause-specific death classification			-
Alive or dead of other cause       135,065 (53.2%)       -       categorical         Survival months       68 (12-151)       0.0%       categorical         5-year survival       1       135,065 (53.2%)       -       categorical         0       119,047 (46.8%)       -       categorical	Dead (attributable to this cancer $dx$ )	119,047~(46.8%)	_	categorical
Survival months         68 (12-151)         0.0%         categorical           5-year survival         1         135,065 (53.2%)         -         categorical           0         119.047 (46.8%)         -         categorical	Alive or dead of other cause	135,065(53.2%)	_	categorical
	Survival months	68 (12-151)	0.0%	categorical
$\begin{array}{cccccccccccccccccccccccccccccccccccc$	5-year survival	. ,		
0 119,047 (46.8%) – categorical	1	135,065~(53.2%)	_	categorical
	0	119,047~(46.8%)	_	categorical

Table 4: SEER (Colon) cohort characteristics, with count (%) or median (Q1–Q3).

Characteristic		Missingness	Type
Sex			
Female	187,967 (41.1%)	_	categorical
Male	269,728(58.9%)	_	categorical
Age recode with single ages and $85+$	68 (60-76)	0.0%	continuous
Race recode (White, Black, Other)			
White	384,184 (83.9%)	_	categorical
Black	47,237 (10.3%)	_	categorical
Other	26,274(5.7%)	_	categorical
Histologic Type ICD-O-3	8,070 (8,041-8,140)	0.0%	categorical
Laterality	, , , , ,		0
Left - origin of primary	178,661 (39.0%)	_	categorical
Right - origin of primary	245,321(53.6%)	_	categorical
Paired site, but no information concerning laterality	23,196(5.1%)	_	categorical
Other	10,517 $(2.3%)$	_	categorical
EOD 10 - nodes (1988-2003)	2 (1-9)	56.3%	categorical
EOD 4 - nodes (1983-1987)	3 (0-9)	88.4%	categorical
Type of Reporting Source			0
Hospital inpatient/outpatient or clinic	445,606 (97.4%)	_	categorical
Other	12.089(2.6%)	_	categorical
SEER historic stage A (1973-2015)			0
Regional	79,409 (17.3%)	_	categorical
Distant	182,467(39.9%)	_	categorical
Blank(s)	123,161(26.9%)	_	categorical
Localized	50,375 (11.0%)	_	categorical
Unstaged	22,283 (4.9%)	_	categorical
CS version input current (2004-2015)	20,520 (20,510-20,540)	70.6%	categorical
CS mets at dx (2004-2015)	23 (0-40)	70.6%	continuous
CS version input original (2004-2015)	10,401 (10,300-20,302)	70.6%	categorical
CS tumor size (2004-2015)	50 (29-999)	70.6%	categorical
EOD 10 - size (1988-2003)	80 (35-999)	56.3%	categorical
CS lymph nodes (2004-2015)	200 (0-200)	70.6%	categorical
Histology recode - broad groupings			C
8140-8389: adenomas and adenocarcinomas	147,127(32.1%)	_	categorical
8010-8049: epithelial neoplasms, NOS	179,848 (39.3%)	_	categorical
8440-8499: cystic, mucinous and serous neoplasms	6,266 (1.4%)	_	categorical
Other	124,454(27.2%)	_	categorical
EOD 10 - extent (1988-2003)	78 (40-85)	56.3%	categorical
SEER cause-specific death classification			0
Alive or dead of other cause	49,997 (10.9%)	_	categorical
Dead (attributable to this cancer dx)	407,698 (89.1%)	_	categorical
Survival months	7 (2-19)	0.0%	categorical
5-year survival			C
1	49,997~(10.9%)	_	categorical
0	407,698 (89.1%)	—	categorical

Table 5: SEER (	(Lung) cohort	characteristics,	with count	(%)	or median	(Q1 -	- Q3).
Table 0. DEBIC	(Lang) conord	, chiaracteristics,	with count	(10)	or mountain	(~~+	

## C.2. Features

### **SEER** (Breast):

AJCC stage 3rd edition (1988-2003) AYA site recode/WHO 2008 Age recode with single ages and 85+ Behavior code ICD-0-2 Behavior code ICD-0-3 Behavior recode for analysis Breast - Adjusted AJCC 6th M (1988-2015) Breast - Adjusted AJCC 6th N (1988-2015) Breast - Adjusted AJCC 6th Stage (1988-2015) Breast - Adjusted AJCC 6th T (1988-2015) Breast Subtype (2010+) CS Schema - AJCC 6th Edition CS extension (2004-2015) CS extension (2004-2015) CS lymph nodes (2004-2015) CS mets at dx (2004-2015) CS site=specific factor 1 (2004-2017 varying by schema) CS site=specific factor 15 (2004-2017 varying by schema) CS site=specific factor 2 (2004-2017 varying by schema) CS site=specific factor 3 (2004-2017 varying by schema) CS site=specific factor 3 (2004-2017 varying by schema) CS site=specific factor 5 (2004-2017 varying by schema) CS site=specific factor 5 (2004-2017 varying by schema) CS site=specific factor 6 (2004-2017 varying by schema) CS site=specific factor 7 (2004-2017 varying by schema) CS site=specific factor 7 (2004-2017 varying by schema) CS version derived (2004-2015) CS version derived (2004-2015) CS version input current (2004-2015) CS version input original (2004-2015) CS version input original (2004-2015) Coding system-EDD (1973-2003) Derived AJCC M, 6th ed (2004-2015) Derived AJCC M, 7th ed (2010-2015) Derived AJCC N, 7th ed (2010-2015) Derived AJCC Stage Group, 6th ed (2004-2015) Derived AJCC Stage Group, 7th ed (2010-2015) Derived AJCC T, 7th ed (2010-2015) Derived AJCC T, 7th ed (2010-2015) Derived AJCC T, 7th ed (2010-2015) Derived HER2 Recode (2010+) EDD 10 - extent (1988-2003) EDD 10 - size (1988-2003) EDD 10 - size (1988-2003) EDS 10 - size (19 ER Status Recode Breast Cancer (1990+) First malignant primary indicator Grade (thru 2017) Histologic Type ICD-0-3 Histology recode - Brain groupings Histology recode - broad groupings ICCC site rec extended ICD-0-3/WHD 2008 IHS Link Laterality Lymphoma subtype recode/WHO 2008 (thru 2017) M value - based on AJCC 3rd (1988-2003) Origin recode NHIA (Hispanic, Non-Hisp) PR Status Recode Breast Cancer (1990+) Primary Site Primary by international rules Race recode (W, B, AI, API) Race recode (White, Black, Other) Mace recode (White, Black, Uther) Race/ethnicity Regional nodes examined (1988+) SEER historic stage A (1973-2015) SEER modified AJCC stage 3rd (1988-2003) Cra Site recode ICD-0-3/WHO 2008 T value - based on AJCC 3rd (1988-2003) Tumor marker 1 (1990-2003) Tumor marker 2 (1990-2003) Tumor marker 3 (1998-2003) Type of Reporting Source

#### SEER (Colon):

Age recode with <1 year olds Age recode with single ages and 85+ Behavior code ICD-O-2 Behavior code ICD-O-3 CS extension (2004-2015) CS lymph nodes (2004-2015) CS site-specific factor 1 (2004-2017 varying by schema) CS tumor size (2004-2015) CS version input current (2004-2015) Derived AJCC M, 7th ed (2010-2015) Derived AJCC M, 7th ed (2010-2015) Derived AJCC N, 7th ed (2010-2015) Derived AJCC Stage Group, 7th ed (2010-2015) Derived AJCC Stage Group, 7th ed (2010-2015) Derived AJCC Stage Group, 7th ed (2010-2015) Derived AJCC T, 7th ed (2010-2015) Diagnostic Confirmation EDD 10 - extent (1988-2003) EOD 10 - size (1988-2003) Histologic Type ICD-0-3 Histology ICD-0-2 Histology recode - broad groupings IHS Link Origin recode NHIA (Hispanic, Non-Hisp) Primary Site Primary by international rules RX Summ--Surg Prim Site (1998+) Race recode (White, Black, Other) Reason no cancer-directed surgery Regional nodes positive (1988+) SEER modified AJCC stage 3rd (1988-2003) Sex

### SEER (Lung):

AYA site recode/WHO 2008 Age recode with single ages and 85+ Behavior code ICD-O-2 Behavior code ICD-O-3 CS extension (2004-2015) CS symph nodes (2004-2015) CS site-specific factor 1 (2004-2017 varying by schema) CS tumor size (2004-2016) CS version input current (2004-2015) Derived AJCC M, 6th ed (2004-2015) Derived AJCC M, 6th ed (2004-2015) Derived AJCC N, 6th ed (2004-2015) Derived AJCC N, 7th ed (2010-2015) Derived AJCC T, 7th ed (2010-2015) Derived AJCC T, 7th ed (2004-2015) Derived AJCC T, 6th ed (2004-2015) Derived AJCC T, 7th ed (2004-2015) Derived AJCC T, 7th ed (2010-2015) Derived AJCC T, 7th ed (2010-2015) Derived AJCC T, 7th ed (2004-2015) Derived AJCC T, 7th

Sex Type of Reporting Source

### C.3. Missingness heatmaps

This section plots missingness heatmaps of categorical and numerical features in each SEER dataset over time. Darker color means larger proportion of missing data.



Figure 11: Missingness of numerical features in SEER (Breast).

Time (year)



Figure 15: Missingness of numerical features in SEER (Lung).

## Appendix D. Additional CDC COVID-19 Data Details

The COVID-19 Case Surveillance Detailed Data (Centers for Disease Control and Prevention, 2020) is a national, publicly available dataset provided by the CDC. It contains 33 elements, with patient-level data including symptoms, demographics, and state of residence. The performance over time is evaluated on a *monthly* basis. We use the version the released on June 6th, 2022. Disclaimer: "The CDC does not take responsibility for the scientific validity or accuracy of methodology, results, statistical analyses, or conclusions presented."

- Data access: To access the data, users must complete a registration information and data use restrictions agreement (RIDURA).
- Cohort selection: The cohort consists of all patients who were lab-confirmed positive for COVID-19, had a non-null positive specimen date, and were hospitalized (hosp\_yn = Yes). Cohort selection diagrams are given in Figures 16
- Cohort characteristics: Cohort characteristics are given in Table 6.
- Outcome definition: mortality, defined by death\_yn = Yes
- Features: We list the features used in the CDC COVID-19 datasets in Section D.2. We convert all categorical variables into dummy features, and apply standard scaling to numerical variables (subtract mean and divide by standard deviation).
- Missingness heat map: is given in Figure 17.
- Additionally, we provide stacked area plots showing how the distribution of ages (Figure 18(a) and states 18(b) shifts over time.





Figure 16: Cohort selection diagram - CDC COVID-19

Characteristic		Missingness	Type
Sex			
Female	455,376~(48.4%)	_	categorical
Male	475,223~(50.5%)	—	categorical
Unknown/Missing	10,541~(1.1%)	—	categorical
Age Group			
0 - 9	16,373~(1.7%)	—	categorical
10 - 19	$17,252\ (1.8\%)$	—	categorical
20 - 29	48,505~(5.2%)	—	categorical
30 - 39	71,776 $(7.6%)$	_	categorical
40 - 49	88,531 $(9.4%)$	_	categorical
50 - 59	141,805~(15.1%)	_	categorical
60 - 69	$189,354\ (20.1\%)$	_	categorical
70 - 79	189,018 (20.1%)	_	categorical
80+	177,765 (18.9%)	_	categorical
Missing	$761 \ (0.1\%)$	_	categorical
Race			
White	544,199~(57.8%)	_	categorical
Black	173,847 (18.5%)	_	categorical
Other	205,547 ( $21.8%$ )	_	categorical
State of Residence			
NY	189,684~(20.2%)	_	categorical
OH	70,097 (7.4%)	_	categorical
$\operatorname{FL}$	35,679 $(3.8%)$	_	categorical
WA	58,854 $(6.3%)$	_	categorical
MA	31,441 ( $3.3%$ )	_	categorical
Other	555,353 (59.0%)	_	categorical
Mechanical Ventilation	L		
Yes	38,009~(4.0%)	_	categorical
No	138,331 (14.7%)	_	categorical
Unknown/Missing	764,800 (81.2%)	_	categorical
Mortality			_
1	190,786~(20.3%)	_	categorical
0	750,354 (79.7%)	_	categorical

 $\label{eq:covid-19} \mbox{Table 6: CDC COVID-19 cohort characteristics, with count (\%) or median (Q1-Q3).}$ 

### D.2. Features

abdom\_yn, abxchest\_yn, acuterespdistress\_yn, age\_group, chills\_yn, cough\_yn, diarrhea\_yn, ethnicity, fever\_yn, hc\_work\_yn, headache\_yn, hosp\_yn, icu\_yn, mechvent\_yn, medcond\_yn, month, myalgia\_yn, nauseavomit\_yn, pna\_yn, race, relative\_month, res\_county, res\_state, runnose\_yn, sex, sfever\_yn, sob\_yn, sthroat\_yn,

#### D.3. Missingness heatmaps



Figure 17: Missingness over time for features in CDC COVID-19 dataset after cohort selection. The darker the color, the larger the proportion of missing data.



### **D.4.** Additional Figures

Figure 18: Proportion of deaths over time for each age group and state of residence.

## Appendix E. Additional SWPA COVID-19 Data Details

The Southwestern Pennsylvania (SWPA) COVID-19 dataset consists of EHR data from patients tested for COVID-19. It was collected by a major healthcare provider in SWPA, and includes patient demographics, labs, problem histories, medications, inpatient vs. outpatient status, and other information collected in the patient encounter. The performance over time is evaluated on a *monthly* basis.

- Data access: This is a private dataset.
- Cohort selection: The cohort consists of COVID-19 patients who tested positive for COVID-19 and were not already in the ICU or mechanically ventilated. We filter for the first positive test, and define features and outcomes relative to that time. Cohort selection diagrams are given in Figures 19. If there are multiple samples per patient, we filter to the first entry per patient, which corresponds to when a patient first enters the dataset. This corresponds to a particular interpretation of the prediction: when a patient is first tests positive, given what we know about that patient, what is their estimated risk of 90-day mortality?
- Cohort characteristics: Cohort characteristics are given in Table 7.
- Outcome definition: 90-day mortality by comparing the death date and test date
- Features: We list the features used in the SWPA COVID-19 datasets in Section E.2. We convert all categorical variables into dummy features, and apply standard scaling to numerical variables (subtract mean and divide by standard deviation). To create a fixed length feature vector, where applicable we take the most recent value of each feature (e.g. most recent lab values).
- Missingness heat maps: are given in Figures 20, 21, 22, and 23,

### E.1. Cohort Selection and Cohort Characteristics



Figure 19: Cohort selection diagram - SWPA COVID-19

Characteristic		Missingness	Type
Gender			
Female	20,283~(57.5%)	_	categorical
Male	15,003~(42.5%)	_	categorical
Unknown	7~(0.0%)	_	categorical
Age			
Under 20	3,210~(9.1%)	—	categorical
20 - 30	4,349~(12.3%)	—	categorical
30 - 40	4,667~(13.2%)	—	categorical
40 - 50	4,653~(13.2%)	—	categorical
50 - 60	6,111~(17.3%)	—	categorical
60 - 70	5,700~(16.2%)	_	categorical
70 +	6,603~(18.7%)	_	categorical
Location of test			
Inpatient	$14,911 \ (42.2\%)$	_	categorical
Outpatient	$17,661\ (50.0\%)$	_	categorical
Unknown	2,721~(7.7%)	—	categorical
90-day mortality			
True	1,516~(4.3%)	—	categorical
False	33,777~(95.7%)	—	categorical

Table 7: SWPA COVID-19 cohort characteristics, with count (%) or median (Q1–Q3).

#### E.2. Features

Asthm CAD CKD COPD CVtest\_ICD\_Acute pharyngitis, unspecified CVtest\_ICD\_Acute upper respiratory infection, unspecified CVtest\_ICD\_Acute upper respiratory infection, unspecified CVtest\_ICD\_Contact with and (suspected) exposure to other viral communicable diseases CVtest\_ICD\_Encounter for general adult medical examination without abnormal findings CVtest\_ICD\_Encounter for screening for respiratory disorder NEC CVtest\_ICD\_Encounter for screening for respiratory disorder NEC CVtest\_ICD\_Nasal congestion CVtest\_ICD\_Nasal congestion CVtest\_ICD\_Other general symptoms and signs CVtest\_ICD\_Other specified symptoms and signs involving the circulatory and respiratory systems CVtest\_ICD\_Parageusia communicable diseases CVtest\_ICD\_Parageusia CVtest ICD R05.9 CVtest\_ICD\_R51.9 CVtest\_ICD\_U07.1 CVtest\_ICD\_Viral infection, unspecified CVtest\_ICD\_Z20.822 ESLD Hypertension IP\_ICD\_z20.828 Inglocations Immunocompromised Interstitial Lung disease OP\_ICD\_Abdominal Pain OP ICD Chest Pain OP\_ICD\_Chills OP\_ICD\_Coronavirus Concerns OP\_ICD\_Covid Infection DP\_ICD\_Exposure To Covid-19 DP\_ICD\_Generalized Body Aches DP\_ICD\_Headache OP\_ICD\_Labs Only OP ICD Medication Refill OP\_ICD\_Nasal Congestion OP\_ICD\_Nausea OP ICD Other OP TCD Results OP\_ICD\_Results OP\_ICD\_Shortness of Breath OP\_ICD\_Sore Throat OP\_ICD\_URI UP\_ICD\_UR1 age\_bin\_(20, 30] age\_bin\_(30, 40] age\_bin\_(40, 50] age\_bin\_(50, 60] age\_bin\_(60, 70] age\_bin\_(70, 200] bmi bmi cancer cough covid\_vaccination\_given diabetes fatigue fever gender hyperglycemia lab ANION GAP lab\_ATRIAL RATE lab\_BASOPHILS ABSOLUTE COUNT lab\_BASOPHILS RELATIVE PERCENT lab\_BLOOD UREA NITROGEN lab\_CALCIUM lab\_CALCUALTED T AXIS lab CALCULATED R AXIS lab\_CHLORIDE lab\_CO2 lab\_CREATININE lab EOSINOPHILS ABSOLUTE COUNT lab\_EOSINOPHILS ABSOLVE COUNT lab\_EOSINOPHILS RELATIVE PERCENT lab\_GFR MDRD AF AMER lab\_GFR MDRD NON AF AMER lab\_GLUCOSE lab\_GLUCUSE lab\_IMMATURE GRANULOCYTES RELATIVE PERCENT lab\_LYMPHOCYTES ABSOLUTE COUNT lab\_LYMPHOCYTES RELATIVE PERCENT lab\_MEAN CORFUSCULAR HEMOGLOBIN lab\_MEAN CORFUSCULAR HEMOGLOBIN CONC lab\_MEAN PLATELET VOLUME lab\_MONOCYTES ABSOLUTE COUNT lab\_MONOCYTES ABSOLUTE COUNT lab\_MONOCYTES RELATIVE PERCENT lab\_NEUTROPHILS RELATIVE PERCENT lab\_NUCLEATED RED BLOOD CELLS lab POTASSTUM lab\_PROTEIN TOTAL lab\_Q-T INTERVAL lab\_QRS DURATION lab\_QRC CALCULATION lab\_RED CELL DISTRIBUTION WIDTH lab\_SODIUM lab\_VENTRICULAR RATE lab\_merged\_CRP

lab\_merged\_albumin lab\_merged\_alkalinePhosphatase lab\_merged\_alt lab\_merged\_ast lab\_merged\_bnp lab\_merged\_ddimer lab\_merged\_directBilirubin lab merged ggt lab\_merged\_bgt lab\_merged\_hct lab\_merged\_hgb lab\_merged\_indirectBilirubin lab\_merged\_lactate lab\_merged\_ldh lab\_merged\_mcv lab\_merged\_neutrophil lab\_merged\_platelets lab\_merged\_pt lab\_merged\_rbc lab\_merged\_sao2 lab\_merged\_totalBilirubin lab\_merged\_totalProtein lab\_merged\_troponin lab\_merged\_troponin lab\_merged\_troponin labs\_ICD\_Acute pharyngitis, unspecified labs\_ICD\_Chete phan, unspecified labs\_ICD\_Chest pain, unspecified labs\_ICD\_Contact with and (suspected) exposure to other viral communicable diseases labs\_ICD\_Escounter for other preprocedural examination labs\_ICD\_Escontar for other preprocedural examination labs\_ICD\_Escontar for other preprocedural examination labs\_ICD\_Escontar for other preprocedural examination labs\_ICD\_Fever, unspecified labs\_ICD\_Heart failure, unspecified labs\_ICD\_Other general symptoms and signs labs\_ICD\_Other pulsonary embolism without acute cor pulmonale labs\_ICD\_Other specified abnormalities of plasma proteins labs\_ICD\_Shortness of breath lab\_merged\_troponin labs\_ICD\_Shortness of breath labs\_ICD\_Syncope and collapse labs\_ICD\_U07.1 labs\_ICD\_Unspecified atrial fibrillation labs\_ICD\_Viral infection, unspecified labs ICD Z20.822 liver disease location\_covidtest\_ordered\_Inpatient location covidtest ordered Outpatient lung disease med\_dx\_Acquired hypothyroidism med\_dx\_Anxiety med\_dx\_COVID-19 med\_dx\_Encounter for antineoplastic chemotherapy
med\_dx\_Encounter for antineoplastic chemotherapy and immunotherapy
med\_dx\_Encounter for antineoplastic immunotherapy med dx Encounter for immunization med\_dx\_Gastroesophageal reflux disease without esophagitis med\_dx\_Gastroesophageal reflux disease, esophagitis presence med\_dx\_Gastroesophageal reflux disease, esophagitis presence not specified med\_dx\_Generalized anxiety disorder med\_dx\_Hyperlipidemia, unspecified hyperlipidemia type med\_dx\_Hypothyroidism, unspecified type med\_dx\_Hron deficiency anemia, unspecified iron deficiency anemia type med\_dx\_Mixed hyperlipidemia med\_dx\_Himary osteoarthritis of right knee medication\_ACETAMINOPHEN 325 %G TABLET medication\_ALEDITENDI SULFATE 2.5 %G/3 ML (0.083 % FOR NEULIZATION medication\_ALBUTENDI SULFATE HFA 90 MCG/ACTUATION AEROSOL INHALER medication\_ALBUTENUL SULFATE HFA 90 MCG/ACTUATION AERUSUL INHALEM medication\_SPIRIN 81 NG TABLET, DELAYED RELEASE medication\_DEXAMETHASONE SODIUM PHOSPHATE 4 MC/ML INJECTION SOLUTION medication\_EPINEPHRINE 0 MC/ML INJECTION (WRAPPER) medication\_EPINEPHRINE 0.3 MG/0.3 ML INJECTION, AUTO-INJECTOR medication\_FENTANYL (PF) 50 MCG/ML INJECTION SOLUTION medication\_HVDROCODNE 5 MC-ACETHAINOPHEN 325 MG TABLET medication\_HVDROCONFISONE SOD SUCCINATE (PF) 100 MG/2 ML SOLUTION FDR INJECTION FOR INJECTION medication\_IOPAMIDOL 76 % medication\_IACTATED RINGERS INTRAVENOUS SOLUTION medication\_MIDAZOLAM 1 MG/ML INJECTION SOLUTION medication\_MALDXONE 0.4 MG/ML INJECTION SOLUTION medication\_NALDADAE OF WAYNE INSECTION SOLUTION medication\_ONDANSETRON HEL (FP) 4 MG/2 ML INJECTION SOLUTION medication\_OXYCODONE 5 MG TABLET medication\_PANTOPRAZOLE 40 MG TABLET,DELAYED RELEASE medication PROPOFOL 10 MG/ML INTRAVENOUS BOLUS (20 ML) medication\_SODIUM CHLORIDE 0.9 % medication\_SODIUM CHLORIDE 0.9 % myalgia obesitv past7Dprobhx\_ICD\_Acute kidney failure, unspecified past7Dprobhx\_ICD\_Anemia, unspecified past7Dprobhx\_ICD\_Anxiety disorder, unspecified past/Dprobhx\_ICD\_Lnext pain, uspecified past/Dprobhx\_ICD\_Lnext pain, unspecified past/Dprobhx\_ICD\_Encounter for general adult medical examination without abnormal findings past7Dprobhx\_ICD\_Encounter for immunization past7Dprobhx\_ICD\_Encounter for screening for malignant neoplasm of colon past7Dprobhx\_ICD\_F32.A past7Dprobhx\_ICD\_Gastro-esophageal reflux disease without esophagitis

past7Dprobk\_ICD\_Hyperlipidemia, unspecified
past7Dprobk\_ICD\_Hypotalemia
past7Dprobk\_ICD\_Hypothyroidism, unspecified
past7Dprobk\_ICD\_Unspecified atrial fibrillation
past7Dprobk\_ICD\_Unspecified atrial fibrillation
probk\_ICD\_Acute kidney failure, unspecified
probk\_ICD\_Chest pain, unspecified
probk\_ICD\_Dizziness and giddiness
probk\_ICD\_Encounter for general adult medical examination without
abnormal findings
probk\_ICD\_Encounter for screening for malignant neoplasm of colon
probk\_ICD\_Factore-acophageal reflux disease without esophagitis
probk.ICD\_Hypetlipidemia, unspecified
probk\_ICD\_Hypothyroidism, unspecified
probk\_ICD\_Hypothyroidism, unspecified
probk\_ICD\_Hypothyroidism, unspecified
probk\_ICD\_Mypokalemia
probk\_ICD\_Unspecified atrial fibrillation
transplant
troponin
vaccine\_COVID-19 RNA (PF) Vaccine (Jansen)
vaccine\_INFLUENZA, CCIV4
vaccine Influenza Kigh PF
vaccine\_Influenza Kigh PF
vaccine\_Influenza PF
vaccine\_Influenza Recombinant (RIV4)
vaccine\_Influenza, Piceuse
vaccine\_INFLUENZA, CCIV4
vaccine\_Influenza, Recombinant (RIV4)
vaccine\_Influenza, Pr
vaccine\_Influenza, Pr
vaccine\_Influenza, Pr
vaccine\_Influenza, Recombinant (RIV4)
vacci

## E.3. Missingness heatmaps

This section plots missingness heatmaps of categorical and numerical features over time. Darker color means larger proportion of missing data.



Figure 20: Missingness of categorical features in SWPA COVID-19 dataset (part 1).



Figure 21: Missingness of categorical features in SWPA COVID-19 dataset (part 2).



Figure 22: Missingness of categorical features in SWPA COVID-19 dataset (part 3).



Figure 23: Missingness of numerical features in SWPA COVID-19.

## Appendix F. Additional MIMIC-IV Data Details

The Medical Information Mart for Intensive Care (MIMIC)-IV (Johnson et al., 2021) database contains EHR data from patients admitted to critical care units from 2008–2019. MIMIC-IV is an update to MIMIC-III, adding time annotations placing each sample into a three-year time range, and removing elements from the old CareVue EHR system (before 2008). Each patient has an anchor\_year\_group, anchor\_year and intime. For each patient, we first calculated an offset as the difference between intime and anchor\_year. Then, we approximated the admit time as the midpoint of anchor\_year\_group after applying the computed offset.

The performance over time is evaluated on a *yearly* basis. Our study uses MIMIC-IV-1.0.

- Data access: Users must create a Physionet account, become credentialed, and sign a data use agreement (DUA).
- Cohort selection: We select all patients in the icustays table, filtering for their first encounter (minimum intime), and defining a feature vector only using information available by the first 24 hrs of their first encounter. (Selection diagram in Figure 24). If there are multiple samples per patient, we filter to the first entry per patient, which corresponds to when a patient first enters the dataset. This corresponds to a particular interpretation of the prediction: when a patient first visits the ICU, given what we know about that patient, what is their estimated risk of in-ICU mortality?
- Outcome definition: The outcome of interest is in-ICU mortality, defined by comparing the outtime of the patient's ICU visit with the patient's dod (date of death, in the patients table). As noted in the documentation, out-of-hospital mortality is not recorded.
- Cohort characteristics: Cohort characteristics are given in Table 8.
- Features: We list the features used in the MIMIC-IV datasets in Section F.2. We convert all categorical variables into dummy features, and apply standard scaling to numerical variables (subtract mean and divide by standard deviation). To create a fixed length feature vector, we take the most recent value of any patient history data available (e.g. most recent lab values).
- Missingness heat maps: are given in Figures 25, 26, 27, 28.

F.1. Cohort Selection and Cohort Characteristics



Figure 24: Cohort selection diagram - MIMIC-IV

Characteristic		Missingness	Type
Gender			
Female	23,313(43.9%)	_	categorical
Male	29,737~(56.1%)	_	categorical
Age at Admission	66 (54-78)	0.0%	continuous
O2 Delivery Device(s)	· · · ·		
Use device	33,359~(62.9%)	_	categorical
None	18,549(35.0%)	_	categorical
Missing	1,142(2.2%)	_	categorical
Pupil Response R	, , ,		0
Brisk	39,708 (74.9%)	_	categorical
Sluggish	4,603 (8.7%)	_	categorical
Non-reactive	1,812(3.4%)	_	categorical
Missing	6,927 (13.1%)	_	categorical
first_careunit	, , ,		0
Medical Intensive Care Unit (MICU)	10,213~(19.3%)	_	categorical
Surgical Intensive Care Unit (SICU)	8,241 (15.5%)	_	categorical
Medical/Surgical Intensive Care Unit (MICU/S	8,808 (16.6%)	_	categorical
Cardiac Vascular Intensive Care Unit (CVICU)	9,437 (17.8%)	_	categorical
Coronary Care Unit (CCU)	6.098(11.5%)	_	categorical
Trauma SICU (TSICU)	6.947(13.1%)	_	categorical
Other	3,306(6.2%)	_	categorical
Anion Gap	13 (11-16)	0.5%	continuous
Heart Rhythm	- ( -)		
SR (Sinus Rhythm)	34.004~(64.1%)	_	categorical
Abnormal heart rhythm	18.657 (35.2%)	_	categorical
Missing	389(0.7%)	_	categorical
Glucose FS (range 70 -100)	131 (110-164)	32.7%	continuous
Eve Opening			
Spontaneously	39,216 (73.9%)	_	categorical
To Speech	7.387 (13.9%)	_	categorical
None	4.538(8.6%)	_	categorical
To Pain	1.702(3.2%)	_	categorical
Missing	207(0.4%)	_	categorical
Lactate	2(1-2)	22.0%	continuous
Motor Response			
Obeys Commands	44,409 (83.7%)	_	categorical
Localizes Pain	3.419(6.4%)	_	categorical
Flex-withdraws	1.673(3.2%)	_	categorical
No response	2.930(5.5%)	_	categorical
Abnormal extension	157(0.3%)	_	categorical
Abnormal Flexion	238(0.4%)	_	categorical
Missing	224(0.4%)	_	categorical
Respiratory Pattern	( )		0
Regular	29,373 (55.4%)	_	categorical
Not regular	1,739(3.3%)	_	categorical
Missing	21,938 (41.4%)	_	categorical
Richmond-RAS Scale	0 (-1-0)	15.4%	categorical
in-icu mortality	- /		0
0	49,716 (93.7%)	_	categorical
1	3,334(6.3%)	_	categorical

Table 8 MIMIC-IV	cohort characteristics	with count (	%)	or median	(01 - 0)	3)
1abic 0. $101010-10$	conore characteristics,	with count (	/0/	or moutan	ાજા જા	<b>J</b> ),

### F.2. Features

18 Gauge Dressing Occlusive 18 Gauge placed in outside facility 20 Gauge Dressing Occlusive 20 Gauge placed in outside facility 20 Gauge placed in the field Abdominal Assessment Activity Activity Tolerance Admission Weight (Kg) Admission Weight (lbs.) Alanine Aminotransferase (ALT) Alarms On Albumin Alkaline Phosphatase All Medications Tolerated Ambulatory aid Anion Gap Anion gap Anti Embolic Device Anti Embolic Device Status Asparate Aminotransferase (AST) Assistance BUN Balance Base Excess Basophils Bath Bicarbonate Bilirubin, Total Bowel Sounds Braden Activity Braden Friction/Shear Braden Mobility Braden Moisture Braden Nutrition Braden Sensory Perception CAM-ICU MS Change Calcium non-ionized Calcium, Total Calculated Total CO2 Capillary Refill L Capillary Refill R Chloride Chloride (serum) Commands Commands Response Cough Effort Cough Type Creatinine Creatinine (serum) Currently experiencing pain Daily Wake Up Delirium assessment Dialysis patient

Diet Type Difficulty swallowing Dorsal PedPulse L Dorsal PedPulse R ETOH Ectopy Type 1 Edema Amount Edema Location Education Barrier Education Existing Knowledge Education Learner Education Method Education Readiness/Motivation Education Response Education Topic Eosinophils Epithelial Cells Eye Opening Family Communication Flatus GU Catheter Size Gait/Transferring Glucose (serum) Glucose FS (range 70 -100) Goal Richmond-RAS Scale HCO3 (serum) HOB HR HR Alarm - High HR Alarm - Low Heart Rhythm Height Height (cm) Hematocrit Hematocrit (serum) Hemoglobin History of falling (within 3 mnths)\* History of slips / falls Home TF INR. INR(PT) IV/Saline lock Insulin pump Intravenous / IV access prior to admission Judgement LLE Color LLE Temp LLL Lung Sounds LUE Color LUE Temp LUL Lung Sounds Lactate Lactic Acid Living situation Lymphocytes

MCH MCHC MCV Magnesium Mental status Monocytes Motor Response NBP Alarm - High NBP Alarm - Low NBP Alarm Source NBPd NBPm NBPs Nares L Nares R Neutrophils O2 Delivery Device(s) Oral Care Oral Cavity Orientation PT PTT Pain Assessment Method Pain Cause Pain Level Pain Level Acceptable Pain Level Response Pain Location Pain Management Pain Present Pain Type Parameters Checked Phosphate Phosphorous Platelet Count Position PostTib Pulses L PostTib Pulses R Potassium Potassium (serum) Potassium, Whole Blood Pressure Reducing Device Pressure Ulcer Present Pupil Response L Pupil Response R Pupil Size Left Pupil Size Right RBC RDW RLE Color RLE Temp RLL Lung Sounds RR RUE Color RUE Temp

RUL Lung Sounds Radial Pulse L Radial Pulse R Red Blood Cells Resp Alarm - High Resp Alarm - Low Respiratory Effort Respiratory Pattern Richmond-RAS Scale ST Segment Monitoring On Safety Measures Secondary diagnosis Self ADL Side Rails Skin Color Skin Condition Skin Integrity Skin Temp Sodium Sodium (serum) SpO2 SpO2 Alarm - High SpO2 Alarm - Low SpO2 Desat Limit Specific Gravity Specimen Type Speech Strength L Arm Strength L Leg Strength R Arm Strength R Leg Support Systems Temp Site Temperature F Therapeutic Bed Tobacco Use History Turn Untoward Effect Urea Nitrogen Urine Source Verbal Response Visual / hearing deficit WBC White Blood Cells Yeast admit\_age gender pC02 pН p02

## F.3. Missingness heatmaps



Figure 25: Missingness over time for labevents features in MIMIC-IV dataset after cohort selection. The darker the color, the larger the proportion of missing data.



Figure 26: Missingness over time for chartevents features in MIMIC-IV dataset after cohort selection. The darker the color, the larger the proportion of missing data. (part 1)



Figure 27: Missingness over time for chartevents features in MIMIC-IV dataset after cohort selection. The darker the color, the larger the proportion of missing data. (part 2)



Figure 28: Missingness over time for chartevents features in MIMIC-IV dataset after cohort selection. The darker the color, the larger the proportion of missing data. (part 3)

## Appendix G. Additional OPTN (Liver) Data Details

The Organ Procurement and Transplantation Network (OPTN) database Organ Procurement and Transplantation Network (2020) tracks organ donation and transplant events in the U.S. Our study uses data from candidates on the liver transplant wait list. The performance over time is evaluated on a *yearly* basis.

- First, we provide the disclaimer: "The data reported here have been supplied by the United Network for Organ Sharing as the contractor for the Organ Procurement and transplantation Network. The interpretation and reporting of these data are the responsibility of the author(s) and in no way should be seen as an official policy of or interpretation by the OPTN or the U.S. Government".
- Data access: After signing the Data Use Agreement I from Organ Procedurement And Transplantation network, users can access the OPTN (Liver) dataset.
- Cohort selection: The cohort consists of liver transplant candidates on the waiting list (2005-2017). We follow the same pipeline as Byrd et al. (2021) to extract the data, except that we select the first record for each patient. Cohort selection diagrams are given in Figures 29. This corresponds to a particular interpretation of the prediction: when a patient is first added to the transplant list, given what we know about that patient, what is their estimated risk of 180-day mortality?
- Outcome definition: 180-day mortality from when the patient was first added to the list
- Cohort characteristics: Cohort characteristics are given in Table 9.
- Features: We list the features used in the OPTN liver dataset in Section G.2. We convert all categorical variables into dummy features, and apply standard scaling to numerical variables (subtract mean and divide by standard deviation).
- Missingness heat maps: are given in Figures 30 and 31.

## G.1. Cohort Selection and Cohort Characteristics



Figure 29: Cohort selection diagram - OPTN (Liver)

Table 9: OPTN	(Liver) c	ohort charact	eristics, with	n count (%)	) or median	(Q1 –	Q3).
---------------	-----------	---------------	----------------	-------------	-------------	-------	------

Feature name (value)		Empty (ratio)	Type
Gender			
Male	92,560~(64.4%)	_	categorical
Female	51,149 ( $35.6%$ )	_	categorical
INIT_AGE	56(49-62)	0.0%	continuous
FUNC_STAT_TCR	2,070 (2,050-2,080)	0.0%	categorical
INIT_OPO_CTR_CODE	11,036 (3,782-19,282)	0.0%	categorical
ALBUMIN	3(3-4)	0.0%	$\operatorname{continuous}$
HCC_DIAGNOSIS_TCR			
No	31,390~(21.8%)	—	categorical
Yes	11,312~(7.9%)	—	categorical
Missing	101,007~(70.3%)	—	categorical
$\mathbf{PERM}_{-}\mathbf{STATE}$			
CA	$19,\!645~(13.7\%)$	_	categorical
TX	14,692~(10.2%)	_	categorical
NY	9,976~(6.9%)	_	categorical
$\operatorname{GA}$	4,052~(2.8%)	_	categorical
MD	4,050~(2.8%)	_	categorical
$\operatorname{FL}$	$7,\!602~(5.3\%)$	_	categorical
PA	8,013~(5.6%)	_	categorical
MI	3,989~(2.8%)	_	categorical
Other	71,007~(49.4%)	_	categorical
EDUCATION	4(3-5)	0.0%	categorical
ASCITES	2(1-2)	0.0%	categorical
MORTALITY_180D			
1	4,635~(3.2%)	_	categorical
0	$139{,}074~(96.8\%)$	_	categorical

### G.2. Features

ABO BACT\_PERIT\_TCR CITIZENSHIP DGN\_TCR DGN2\_TCR DIAB EDUCATION FUNC\_STAT\_TCR GENDER LIFE\_SUP\_TCR MALIG\_TCR OTH\_LIFE\_SUP\_TCR PERM\_STATE PORTAL\_VEIN\_TCR PREV\_AB\_SURG\_TCR PRI\_PAYMENT\_TCR REGION TIPSS\_TCR VENTILATOR\_TCR WORK\_INCOME\_TCR ETHCAT HCC\_DIAGNOSIS\_TCR MUSCLE\_WAST\_TCR INIT\_OPO\_CTR\_CODE WLHR WLIN WLKI WLLU WLPA INACTIVE ASCITES ENCEPH DIALYSIS\_PRIOR\_WEEK  $\texttt{INIT}_\texttt{HGT}_\texttt{CM}$ INIT\_WGT\_KG INIT\_BMI\_CALC INIT\_AGE UNOS\_CAND\_STAT\_CD BILIRUBIN SERUM\_CREAT INR SERUM\_SODIUM ALBUMIN BILIRUBIN\_DELTA SERUM\_CREAT\_DELTA INR\_DELTA SERUM\_SODIUM\_DELTA ALBUMIN\_DELTA





Figure 30: Missingness over time for categorical features in OPTN (Liver) dataset after cohort selection. The darker the color, the larger the proportion of missing data.



Figure 31: Missingness over time for numerical features in OPTN (Liver) dataset after cohort selection. The darker the color, the larger the proportion of missing data. (Near-zero missingness here.)

## Appendix H. Additional MIMIC-CXR Data Details

The MIMIC Chest X-ray (MIMIC-CXR-JPG) (Johnson et al., 2019b) is a publicly available dataset containing chest radiographs in JPG format from 2009–2018. Similar to MIMIC-IV, MIMIC-CXR add time annotations placing each sample into a three-year time range. We approximate the year of each sample by taking the midpoint of its time range. Each patient has an anchor\_year\_group, anchor\_year and StudyDate. For each patient, we first calculated an offset as the difference between StudyDate and anchor\_year. Then, we approximated the admit time as the midpoint of anchor\_year\_group after applying the computed offset. The performance over time is evaluated on a *yearly* basis. Our study uses MIMIC-IV-JPG-2.0. A similar training setup to that in Seyyed-Kalantari et al. (2020) was used (learning rate, architecture, data augmentation, stopping criteria, etc.).

- Data access: Users must create a Physionet account, become credentialed, and sign a data use agreement (DUA).
- Cohort selection: We removed the records from 2009 due to the tiny sample size. (Selection diagram in Figure 32). We keep all records for each patients and split the data based on patient subject id.
- Outcome definition: The outcome is the probabilities of all labels given the input images. The labels includes 13 abnormal outcomes and 1 normal outcome. (Atelectasis, Cardiomegaly, Consolidation, Edema, Enlarged Cardiomediastinum, Fracture, Lung Lesion, Lung Opacity, Pleural Effusion, Pneumonia, Pneumothorax, Pleural Other, Support Devices, No Finding)
- Cohort characteristics: Cohort characteristics are given in Table 10.

## H.1. Cohort Selection and Cohort Characteristics



Figure 32: Cohort selection diagram - MIMIC-CXR

	~		
Feature name (value)	Summary statistic	Empty (ratio)	Status
Gender			
$\mathbf{F}$	179,765~(47.8%)	_	categorical
М	196,439~(52.2%)	—	categorical
Age	64(51-76)	0.0%	continuous
Diseases			
Atelectasis	65,390~(17.4%)	—	categorical
Cardiomegaly	56,404~(15.0%)	—	categorical
Consolidation	14,394~(3.8%)	—	categorical
Edema	36,026~(9.6%)	—	categorical
Enlarged Cardiomediastinum	9,821~(2.6%)	_	categorical
Fracture	6,314~(1.7%)	_	categorical
Lung Lesion	10,574~(2.8%)	_	categorical
Lung Opacity	76,074~(20.2%)	_	categorical
Pleural Effusion	75,526~(20.1%)	—	categorical
Pleural Other	$3,\!432~(0.9\%)$	—	categorical
Pneumonia	25,065~(6.7%)	—	categorical
Pneumothorax	12,828~(3.4%)	—	categorical
Support Devices	69,148~(18.4%)	_	categorical
No Finding	167,116~(44.4%)	_	categorical

Table 10: MIMIC-CXR cohort characteristics, with count (%) or median (Q1–Q3).



## H.2. Label level AUROC over time for MIMIC-CXR

Figure 33: Absolute AUROC over time of each label in MIMIC-CXR



Figure 34: Weighted test AUROC vs. year for the DenseNet architecture on MIMIC-CXR.

Table 11: MIMIC-CXR label-level AUROC from time-agnostic evaluation of all-period training. The format is mean (±std. dev. across splits)

Label	AUROC	Label	AUROC
Atelectasis	$0.826~(\pm 0.003)$	Cardiomegaly	$0.837 (\pm 0.002)$
Consolidation	$0.841 \ (\pm 0.003)$	Edema	$0.904~(\pm 0.002)$
Enlarged Cardiomediastinum	$0.759~(\pm 0.005)$	Fracture	$0.745~(\pm 0.006)$
Lung Lesion	$0.784~(\pm 0.003)$	Lung Opacity	$0.770 \ (\pm 0.002)$
Pleural Effusion	$0.929 \ (\pm 0.001)$	Pleural Other	$0.844 \ (\pm 0.009)$
Pneumonia	$0.755 (\pm 0.004)$	Pneumothorax	$0.918 \ (\pm 0.006)$
Support Devices	$0.928~(\pm 0.001)$	No Finding	$0.876~(\pm 0.002)$

## Appendix I. Logistic Regression Coefficients from Splitting by Patient

To help with intuition in important features for the predictive task on each dataset, here we have the coefficients of logistic regression models trained from splitting by patient.

Table 12: SEER (Breast) top 10 important features for LR models, all-period training.

Feature	Coefficient
SEER historic stage A (1973-2015)_Distant	-2.113944
SEER historic stage A (1973-2015)_Localized	1.676493
Regional nodes examined $(1988+)_95.0$	-1.167844
CS lymph nodes $(2004-2015)_{-750}$	1.100824
CS lymph nodes $(2004-2015)_{-755}$	1.023753
Histologic Type ICD-O-3_8530	-0.913494
Histologic Type ICD-O-3_8543	0.902798
Breast - Adjusted AJCC 6th T (1988-2015)_T4d	0.899491
Histologic Type ICD-O-3_8211	0.877848
EOD 10 - extent (1988-2003)_85	-0.791136

Table 13: SEER (Colon) top 10 important features for LR models, all-period training.

Feature	Coefficient
Reason no cancer-directed surgery_Surgery performed	2.360161
Regional nodes positive $(1988+)_00$	1.897706
Regional nodes positive $(1988+)_01$	1.872008
modified AJCC stage 3rd (1988-2003)_40	-1.787481
EOD 10 - extent (1988-2003)_13	1.766066
Reason no cancer-directed surgery_Not recommended,	-1.752474
contraindicated due to other cond; autopsy only (1973-2002)	
EOD 10 - extent (1988-2003)_85	-1.732619
EOD 10 - extent (1988-2003)_70	-1.704333
CS mets at dx (2004-2015)_99	1.619905
CS mets at dx (2004-2015)_00	1.609454

Feature	Coefficient
Histologic Type ICD-O-3_8240	2.514539
EOD 4 - nodes $(1983-1987)_{-0}$	2.074730
EOD 4 - nodes (1983-1987)_7	-1.777530
EOD 10 - size (1988-2003)_140	-1.587893
Histologic Type ICD-O-3_8141	-1.546566
CS tumor size (2004-2015)_998.0	-1.515856
EOD 4 - nodes $(1983-1987)_{-6}$	-1.497022
Type of Reporting Source_Nursing/convalescent home/hospice	-1.338998
CS mets at dx $(2004-2015)_{-51}$	-1.326595
EOD 10 - size (1988-2003)_150	-1.326196

Table 14: SEER (Lung) top 10 important features for LR models, all-period training.

Table 15: CDC COVID-19 top 10 important features for LR models, all-period training.

Feature	Coefficient
res_state_DE	2.202055
$age_group_0 - 9$ Years	-2.114818
$age\_group\_80+$ Years	1.965279
$age\_group_10 - 19$ Years	-1.681099
res_state_GA	1.391469
$age_group_70 - 79$ Years	1.379589
res_county_WICHITA	1.290644
$age\_group_20 - 29$ Years	-1.189734
$res\_county\_SUMNER$	-1.135073
$mechvent_yn_Yes$	1.117372

Table 16: SWPA COVID-19 top 10 important features for LR models according to experiments splitting by patient.

Feature	Coefficient
age_bin_(70, 200]_0	-0.781337
age_bin_(70, 200]_1	0.780673
medication_FENTANYL (PF) 50 MCG/ML INJECTION SOLUTION_0.0	0.651419
medication_EPINEPHRINE $0.3 \text{ MG}/0.3 \text{ ML}$ INJECTION, AUTO-INJECTOR_nan	-0.627565
medication_HYDROCORTISONE SOD SUCCINATE (PF) 100 MG/2 ML SOLUTION FOR INJECTION_0.0	0.544222
medication_HYDROCODONE 5 MG-ACETAMINOPHEN 325 MG TABLET_nan	-0.520368
medication_DEXAMETHASONE SODIUM PHOSPHATE 4 MG/ML INJECTION SOLUTION_0.0	0.502954
medication_ASPIRIN 81 MG TABLET,DELAYED RELEASE_nan	-0.479100
bmi_nan	-0.427569
age_bin_(60, 70]_0	-0.380688

Table 17: MIMIC-IV top 10 important features for LR models, all-period training.

Feature	Coefficient
O2 Delivery Device(s)_None	-0.307334
Eye Opening_None	0.301737
admit_age	0.299712
O2 Delivery Device(s)_Nasal cannula	-0.248463
Motor Response_Obeys Commands	-0.230931
Pupil Response L_Non-reactive	0.223776
Richmond-RAS Scale_ 0 Alert and calm	-0.205476
Temp Site_Blood	-0.204514
HR_0.0	0.197299
Diet Type_NPO	0.195156

Table 18: OPTN (Liver) top 10 important features for LR models, all-period training.

Feature	Coefficient
SERUM_CREAT_DELTA	0.660589
FUNC_STAT_TCR_2020.0	0.241507
FUNC_STAT_TCR_2080.0	-0.236288
DGNC_4110.0	-0.234680
REGION_5.0	0.223940
EDUCATION_998.0	0.218549
ASCITES_3.0	0.218329
ASCITES_1.0	-0.214076
INIT_OPO_CTR_CODE_1054	-0.209265
INIT_OPO_CTR_CODE_4743	-0.207778

## Appendix J. Diagnostic plots

We took the union of the top k most important features from each time point to be included in the diagnostic plots, where k was tuned depending on the dataset so that the resulting plots would not be overcrowded. For categorical features, we additionally highlighted (using a thicker line) features that had consistently high prevalence ( $\geq p$ ) or experienced a large change in prevalence across one time point ( $\geq \Delta$ ). The specific parameters of each dataset are defined in each subsection. For numerical features, we highlighted features whose average ranking across all time points was  $\leq 3$  (also chosen to avoid overcrowding).

## J.1. SEER (Breast)

For SEER (Breast) diagnostic plots, important features were selected using  $k = 5, p = 0.4, \Delta = 0.2$ .



Figure 35: Diagnostic plot of SEER (Breast) dataset. The important features are selected as the union of the top 5 features that have the highest absolute value model coefficients. The left column includes AUROC versus time for both sliding window and all-historical subsampled, and the maximum AUROC drop for each trained model. The right column provides the absolute coefficients of each trained model from both regimes, and positive proportion of the significant features over time. As shown in the gray highlighted region, there are jumps in performance around 1988 and 2003, which coincides with the introducing and removal of several features (e.g. T value - based on AJCC 3rd (1988-2003)\_T1). The latency of jumps in coefficients are caused by length of sliding window.





For SEER (Colon) diagnostic plots, important features were selected using  $k = 3, p = 0.4, \Delta = 0.2$ .

Figure 36: Diagnostic plot of SEER (Colon) dataset. The important features are selected as the union of the top 3 features that have the highest absolute model coefficients. The left column includes AUROC versus time for both sliding window and all-historical subsampled, and the maximum AUROC drop for each trained model. The right column provides the absolute coefficients of each trained model from both regimes, and positive proportion of the significant features over time. As shown in the gray highlighted region, there are jumps in performance around 1988 and 2003, which coincides with the introducing and removal of several features (e.g. SEER modified AJCC stage 3rd (1988-2003)\_40). The latency of jumps in coefficients are caused by length of sliding window.







Figure 37: Diagnostic plot of SEER (Lung) dataset. The important features are selected as the union of the top 5 features that have the highest absolute model coefficients. The left column includes AUROC versus time for both sliding window and all-historical subsampled, and the maximum AUROC drop for each trained model. The right column provides the absolute coefficients of each trained model from both regimes, and positive proportion of the significant features over time. As shown in the gray highlighted region, there are jumps in performance around 1988 and 2003, which coincides with the introducing and removal of several features (e.g. EOD 10 - nodes (1988-2013)\_0 & EOD 10 - extent (1988-2003)\_85). The latency of jumps in coefficients are caused by length of sliding window.

### J.4. CDC COVID-19



For CDC COVID-19 diagnostic plots, important features were selected using  $k = 5, p = 0.15, \Delta = 0.15$ .

Figure 38: Diagnostic plot of CDC COVID-19. The important features are selected as the union of the top 5 features that have the highest absolute model coefficients. The left column includes AUROC versus time for both sliding window and all-historical subsampled, and the maximum AUROC drop for each trained model. The right column provides the absolute coefficients of each trained model from both regimes, and positive proportion of the significant features over time. As shown in the gray highlighted region, the models trained around June 2021 suffer the largest maximum AUROC drop, coinciding with a shift in distribution of ages (Figure 18(a)) and states (Figure 18(b)). The latency of jumps in coefficients are caused by length of sliding window.

### J.5. SWPA COVID-19



For SWPA COVID-19 diagnostic plots, important features were selected using  $k = 3, p = 0.4, \Delta = 0.2$ .

Figure 39: Diagnostic plot of SWPA COVID-19. The important features are selected as the union of the top 3 features that have the highest absolute model coefficients. The left column includes AUROC versus time for both sliding window and all-historical subsampled, and the maximum AUROC drop for each trained model. The right column provides the absolute coefficients of each trained model from both regimes, and positive proportion of the significant features over time. One of the hypotheses for relatively large uncertainty is smaller sample size.

## J.6. MIMIC-IV





Figure 40: Diagnostic plot of MIMIC-IV. The important features are selected as the union of the top 3 features that have the highest absolute model coefficients. The left column includes AUROC versus time for both sliding window and all-historical subsampled, and the maximum AUROC drop for each trained model. The right column provides the absolute coefficients of each trained model from both regimes, and positive proportion of the significant features over time. The model performance is relatively stable, coinciding with relatively stable distributions of a majority of important features.

### J.7. OPTN (Liver)



For OPTN (Liver) diagnostic plots, important features were selected using  $k = 3, p = 0.4, \Delta = 0.2$ .

Figure 41: Diagnostic plot of OPTN (Liver). The important features are selected as the union of the top 3 features that have the highest absolute model coefficients. The left column includes AUROC versus time for both sliding window and all-historical subsampled, and the maximum AUROC drop for each trained model. The right column provides the absolute coefficients of each trained model from both regimes, and positive proportion of the significant features over time. Although the HCC DIAGNOSIS TCR binary features change in positive proportion over time, these features were not always important, and the other important features (faded) maintain relatively stable proportions across time. Overall, model performance is quite stable over time.

### J.8. MIMIC-CXR



Figure 42: Diagnostic plot of MIMIC-CXR. The top and mid left includes AUROC versus time for both sliding window and all-historical subsampled. The top right is the maximum AUROC drop for each trained model. The mid-right provides the label proportions over time. The bottom shows pixel intensities for images in each year. The histogram of pixel intensity is stable over time, which is consistent with the small variation in model performance over time

# Appendix K. Model performance over time from three models K.1. AUROC

All plots in this section are for the all-historical training regime.





Figure 43: AUROC versus test timepoints from three model classes on all datasets.

## K.2. AUPRC

All plots in this section are for the all-historical training regime.



Test AUPRC vs. Timepoint (year or month)

Figure 44: AUPRC versus test timepoints from three model classes on all datasets. Label prevalance refers to the ratio of accumulated positive labels over time.

## Appendix L. Data Split Details

Table 19: Split ratio for each dataset for training, validation and testing (both for time-agnostic splits and in-period splits).

Dataset	Split ratio
SEER (Breast)	0.8-0.1-0.1
SEER (Colon)	0.8 - 0.1 - 0.1
SEER (Lung)	0.8 - 0.1 - 0.1
CDC COVID-19	0.8 - 0.1 - 0.1
SWPA COVID-19	0.5 - 0.25 - 0.25
MIMIC-IV	0.5 - 0.25 - 0.25
OPTN (Liver)	0.5 - 0.25 - 0.25
MIMIC-CXR	0.5 - 0.25 - 0.25

## Appendix M. Hyperparameter Grids

Parameter	Values Considered		
LR			
С	$0.01, 0.1, 1, 10, 10^2, 10^3, 10^4, 10^5$		
GBDT			
$n_{-}$ estimators	50,100		
$\max\_depth$	3, 5		
learning_rate	0.01,  0.1		
MLP			
hidden_layer_sizes	3, 5		
learning_rate_init	$10^{-4},  10^{-3},  0.01$		

Table 20: Hyperparameter grids for model training.

# Appendix N. AUROC from full-period training

Dataset	Model	Full-period AUROC
SEER (Breast)	LR GBDT MLP	$\begin{array}{c} 0.888 \ (\pm 0.002) \\ 0.891 \ (\pm 0.002) \\ 0.891 \ (\pm 0.002) \end{array}$
SEER (Colon)	LR GBDT MLP	$\begin{array}{c} 0.863 \ (\pm 0.003) \\ 0.868 \ (\pm 0.002) \\ 0.869 \ (\pm 0.003) \end{array}$
SEER (Lung)	LR GBDT MLP	$\begin{array}{c} 0.894 \ (\pm 0.002) \\ 0.894 \ (\pm 0.002) \\ 0.898 \ (\pm 0.002) \end{array}$
CDC COVID-19	LR GBDT MLP	$\begin{array}{c} 0.837 \ (\pm 0.001) \\ 0.851 \ (\pm 0.001) \\ 0.852 \ (\pm 0.002) \end{array}$
SWPA COVID-19	LR GBDT MLP	$\begin{array}{c} 0.928 \ (\pm 0.005) \\ 0.930 \ (\pm 0.004) \\ 0.928 \ (\pm 0.006) \end{array}$
MIMIC-IV	LR GBDT MLP	$\begin{array}{c} 0.935 \ (\pm 0.003) \\ 0.931 \ (\pm 0.002) \\ 0.898 \ (\pm 0.008) \end{array}$
OPTN (Liver)	LR GBDT MLP	$\begin{array}{c} 0.846 \ (\pm 0.005) \\ 0.854 \ (\pm 0.005) \\ 0.847 \ (\pm 0.006) \end{array}$
MIMIC-CXR	DenseNet	$0.860 \ (\pm 0.001)$

Table 21: AUROC report from full-period training, the results are in format mean ( $\pm$ std. dev. across splits)