

## Motivation & Challenges

### Motivation

- ✓ Improving discriminative in survival models often compromises calibration.
- ✓ A new approach is need to enhance discrimination without sacrificing calibration, using the embedding space through contrastive learning.

### Challenges

- ✓ Combining NLL with ranking loss improves discrimination but misaligns model outputs with the actual risk distribution, negatively affecting calibration and clinical applicability.

## Preliminaries

### Discrete-Time Survival Analysis

- ✓ Survival function  $S$  represents the probability that the event occurs after time  $t$  for a patient with features  $\mathbf{x}$ .
- ✓ Hazard function  $\lambda$  is the instantaneous risk of the event at time  $t$  given feature  $\mathbf{x}$

$$S(t|\mathbf{x}) = \mathbb{P}(T > t|\mathbf{x}) = \prod_{t' \leq t} (1 - \lambda(t'|\mathbf{x}))$$

$$\mathcal{L}_{NLL} = - \sum_{i=1}^N [\delta_i \log \hat{p}(\tau_i|\mathbf{x}_i) + (1 - \delta_i) \log \hat{S}(\tau_i|\mathbf{x}_i)]$$

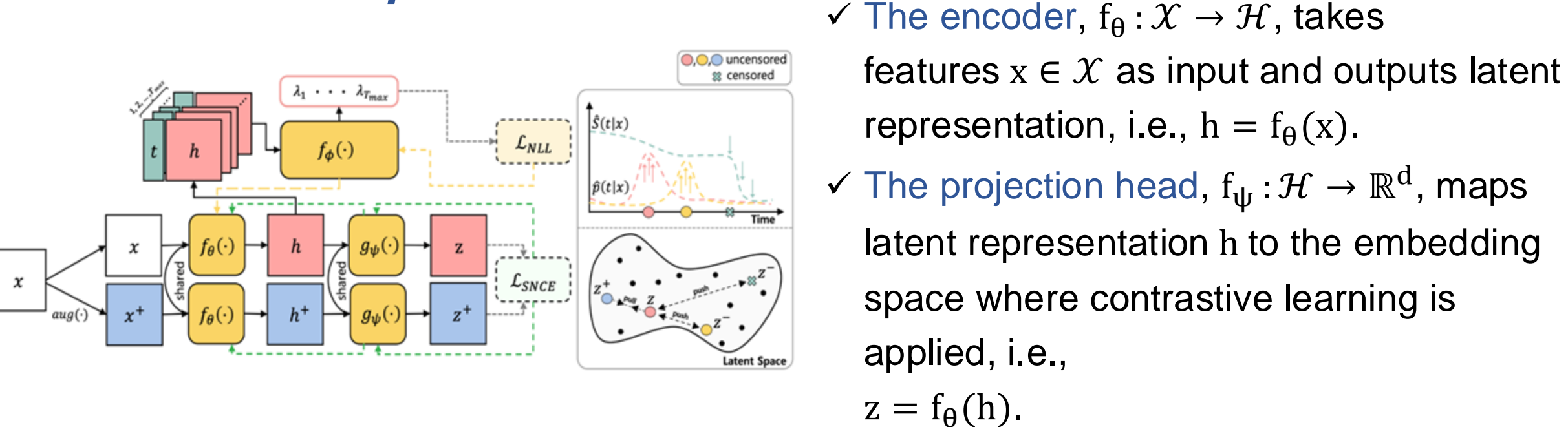
where  $\hat{p}$  represents the estimate for the probability of an event occurring at time  $t$

### Ranking Loss for Survival Analysis

Aim to maximize a relaxed proxy of the concordance index.

$$\mathcal{L}_{Rank} = \sum_{i \neq j} A_{i,j} \cdot \eta \left( \hat{R}(\tau_i|\mathbf{x}_i), \hat{R}(\tau_j|\mathbf{x}_j) \right)$$

## Network Description



- ✓ The hazard network,  $f_\phi: \mathcal{H} \times \mathcal{T} \rightarrow [0,1]$ , predicts the hazard rate at each time point  $t \in \mathcal{T}$  given input latent representation  $\mathbf{h}$ , i.e.,  $\hat{\lambda}(t|\mathbf{x}) = f_\phi(f_\theta(\mathbf{x}), t)$ .

## Method

- ✓ Propose a novel contrastive learning approach for deep survival model
- ✓ Deviate from directly ranking the model outcome in the form of risk/survival functions
- ✓ Goal : Aligns with our inductive bias that **patients with similar survival outcomes should share similar clinical status**, which manifests through similar representations

### Noise Contrastive Estimation (NCE)

To learn mapping  $f = g_\psi \circ f_\theta$  utilizing a positive sample  $\mathbf{x}^+ \sim p_{X^+}$ , and negative samples  $\mathbf{x}^- \sim q$

$$\mathbb{E}_{\mathbf{x}^+ \sim p_{X^+}, \mathbf{x}^- \sim q} \left[ -\log \frac{e^{s(\mathbf{x}, \mathbf{x}^+)}}{M \cdot \mathbb{E}_{\mathbf{x}^- \sim q} [e^{s(\mathbf{x}, \mathbf{x}^-)}]} \right]$$

### Weighted Distribution $q$ for Time-to-Event Differences

To reflect the difference in the time-to-events in the embedding space, we design a novel distribution  $q$  by utilizing the available information from survival outcomes.

$$w(\tau^-; \tau) = 1 - e^{-|\tau - \tau^-|/\sigma}$$

- ✓ Hence, given an anchor  $(\mathbf{x}, \tau)$  and a negative  $(\mathbf{x}^-, \tau^-)$ , we define the weight function,  $\sigma > 0$  is a temperature coefficient.
- ✓ This function assigns larger weights to samples with large differences in time-to-event outcomes, and smaller weights to samples with small differences.

$$q(\mathbf{x}^-; \mathbf{x}) = \frac{1}{Z} w(\mathbf{x}^-; \mathbf{x}) p(\mathbf{x}^-)$$

- ✓ Designing  $q$  based on the following inductive bias similar patients are more likely to experience the event at similar time points than the ones who are not.

### Importance Sampling Using Survival Outcomes

$$\begin{aligned} E_{\mathbf{x}^- \sim q} [e^{s(\mathbf{x}, \mathbf{x}^-)}] &= E_{\mathbf{x}^- \sim p} \left[ \left( \frac{q(\mathbf{x}^-; \mathbf{x})}{p(\mathbf{x}^-)} \right) \cdot e^{s(\mathbf{x}, \mathbf{x}^-)} \right] \\ &= E_{\mathbf{x}^- \sim p} \left[ \left( \frac{w(\mathbf{x}^-; \mathbf{x})}{Z} \right) \cdot e^{s(\mathbf{x}, \mathbf{x}^-)} \right] \\ &\approx \frac{1}{Z \cdot M} \sum_{j=1}^M w(\mathbf{x}_j^-; \mathbf{x}) \cdot e^{s(\mathbf{x}, \mathbf{x}_j^-)} \end{aligned}$$

## Experiments

### Effect of Contrastive Learning

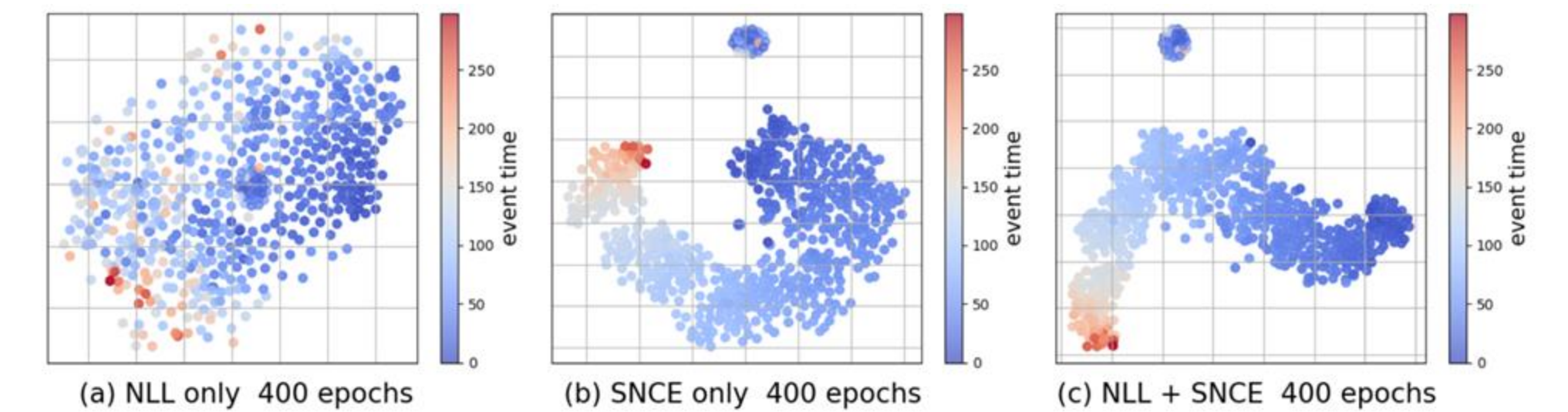


Figure 2: t-SNE visualization for latent representations learned with  $\mathcal{L}_{NLL}$  only,  $\mathcal{L}_{SNCE}$  only, and ConSurv for the METABRIC dataset, colored by event times (for uncensored samples).

### Calibration Plot

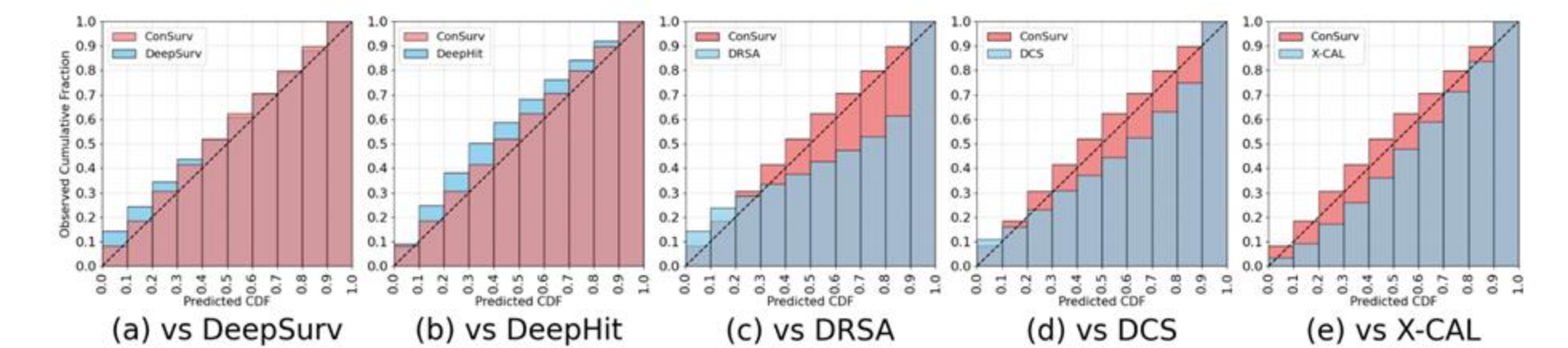


Figure 3: Calibration plots for ConSurv in comparison with benchmarks for the METABRIC dataset.

### Subgroup Analysis

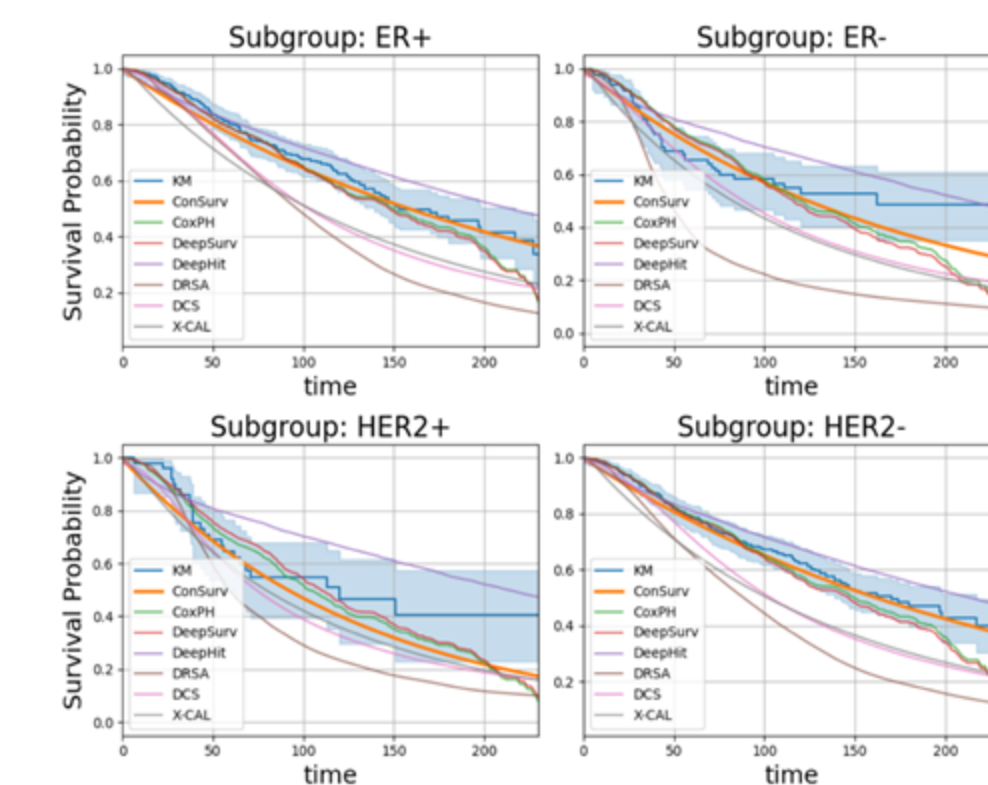


Figure 4: Comparison of the survival curves across various patient subgroups for the METABRIC dataset.

### Quantitative Analysis

METHOD	CI $\uparrow$	METABRIC		
		IBS $\downarrow$	DDC $\downarrow$	D-CAL
COXPH	0.645 $\pm$ 0.019	<b>0.175<math>\pm</math>0.028</b>	0.111 $\pm$ 0.024	<b>25</b>
DEEPSURV	0.625 $\pm$ 0.025	0.183 $\pm$ 0.029	0.103 $\pm$ 0.026	<b>25</b>
DEEPHIT	0.604 $\pm$ 0.019	0.204 $\pm$ 0.018	0.292 $\pm$ 0.017	0
DRSA	0.604 $\pm$ 0.032	0.249 $\pm$ 0.038	0.178 $\pm$ 0.060	0
DCS	0.612 $\pm$ 0.029	0.206 $\pm$ 0.043	<b>0.054<math>\pm</math>0.039</b>	2
X-CAL	0.632 $\pm$ 0.027	0.182 $\pm$ 0.023	0.065 $\pm$ 0.037	2
<b>CONSURV</b>	<b>0.665<math>\pm</math>0.023</b>	0.186 $\pm$ 0.021	0.110 $\pm$ 0.024	<b>23</b>

METHOD	CI $\uparrow$	GBSG		
		IBS $\downarrow$	DDC $\downarrow$	D-CAL
COXPH	0.662 $\pm$ 0.179	0.181 $\pm$ 0.007	0.183 $\pm$ 0.037	<b>25</b>
DEEPSURV	0.653 $\pm$ 0.042	0.182 $\pm$ 0.009	0.153 $\pm$ 0.066	24
DEEPHIT	0.633 $\pm$ 0.032	0.205 $\pm$ 0.006	0.342 $\pm$ 0.023	3
DRSA	0.668 $\pm$ 0.016	0.278 $\pm$ 0.018	0.402 $\pm$ 0.055	0
DCS	<b>0.677<math>\pm</math>0.017</b>	0.181 $\pm$ 0.008	<b>0.124<math>\pm</math>0.025</b>	10
X-CAL	0.675 $\pm$ 0.017	0.181 $\pm$ 0.010	0.166 $\pm$ 0.020	8
<b>CONSURV</b>	<b>0.677<math>\pm</math>0.020</b>	<b>0.179<math>\pm</math>0.007</b>	0.160 $\pm$ 0.026	<b>18</b>

- ✓ Significantly improves the alignment of representations with event time information.
- ✓ Compares survival plots of the models with the Kaplan-Meier curve to confirm calibration.
- ✓ ConSurv outperforms all benchmarks in discrimination and maintains comparable calibration.