

# Leveraging Tumor Heterogeneity: Heterogeneous Graph Representation Learning for Cancer Survival Prediction in Whole Slide Images

Junxian Wu, Xinyi Ke, Xiaoming Jiang, Huanwen Wu, Youyong Kong, Lizhi Shao



# Background

**Task:** Survival prediction based on Whole Slide Images (WSIs).

## The Ultra-High Resolution of WSIs

- Contain much redundant information.
- Typically employ the **Multiple Instance Learning (MIL)** method to partition and process WSIs, may result in the loss of important information.

## Abundant Domain Priors

- Pathological research is sufficiently extensive.
- The domain priors regarding **intratumoral tissues** that may affect prognosis are abundantly available<sup>[1,2,3]</sup>.

[1] Oliver, Amanda J., et al. "Tissue-dependent tumor microenvironments and their impact on immunotherapy responses." *Frontiers in immunology* 9 (2018): 70.

[2] Mountain, Clifton F. "New prognostic factors in lung cancer: biologic prophets of cancer cell aggression." *Chest* 108.1 (1995): 246-254.

[3] Bremnes, Roy M., et al. "The role of tumor stroma in cancer progression and prognosis: emphasis on carcinoma-associated fibroblasts and non-small cell lung cancer." *Journal of thoracic oncology* 6.1 (2011): 209-217.

# Motivation

Can we **retain** as much information as possible within the MIL pipeline, and **guide** the model to focus on intratumoral tissues that are highly relevant to prognosis based on pathological priors?

## Re-modeling the lost information

- Spatial relationship information between patches<sup>[1]</sup>.
- Hierarchical perspective information<sup>[1,2]</sup>.

## Incorporation of histopathological priors

- Intratumoral tissues commonly recognized as relevant to prognosis (tumor, stroma, necrosis, infiltration, etc.)
- Guiding the model to focus primarily on tissues relevant to prognosis while not entirely neglecting other tissues.

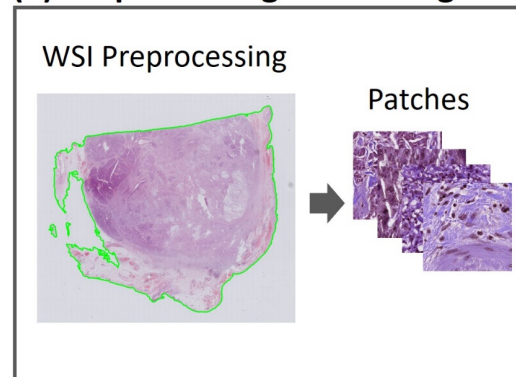
[1] Chen, Richard J., et al. "Whole slide images are 2d point clouds: Context-aware survival prediction using patch-based graph convolutional networks." *Medical Image Computing and Computer Assisted Intervention–MICCAI 2021: 24th International Conference, Strasbourg, France, September 27–October 1, 2021, Proceedings, Part VIII 24*. Springer International Publishing, 2021.

[2] Chen, Richard J., et al. "Scaling vision transformers to gigapixel images via hierarchical self-supervised learning." *Proceedings of the IEEE/CVF Conference on Computer Vision and Pattern Recognition*. 2022.

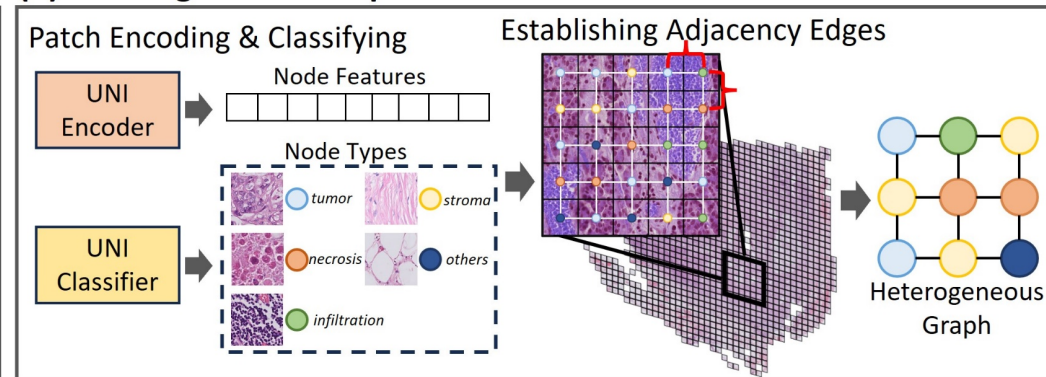
# Our Method: ProtoSurv

- A dual-stream heterogeneous graph model that is decoupled into the **Structure View (SV)** and **Histology View (HV)**.
- **SV** employs GNN to model the spatial relationship between patches and simulate hierarchical receptive fields.
- **HV** constructs a prototype network, extracting multi-prototypes globally based on pre-obtained node categories, with a focus on intratumoral tissues that are highly relevant to prognosis based on histopathological priors.

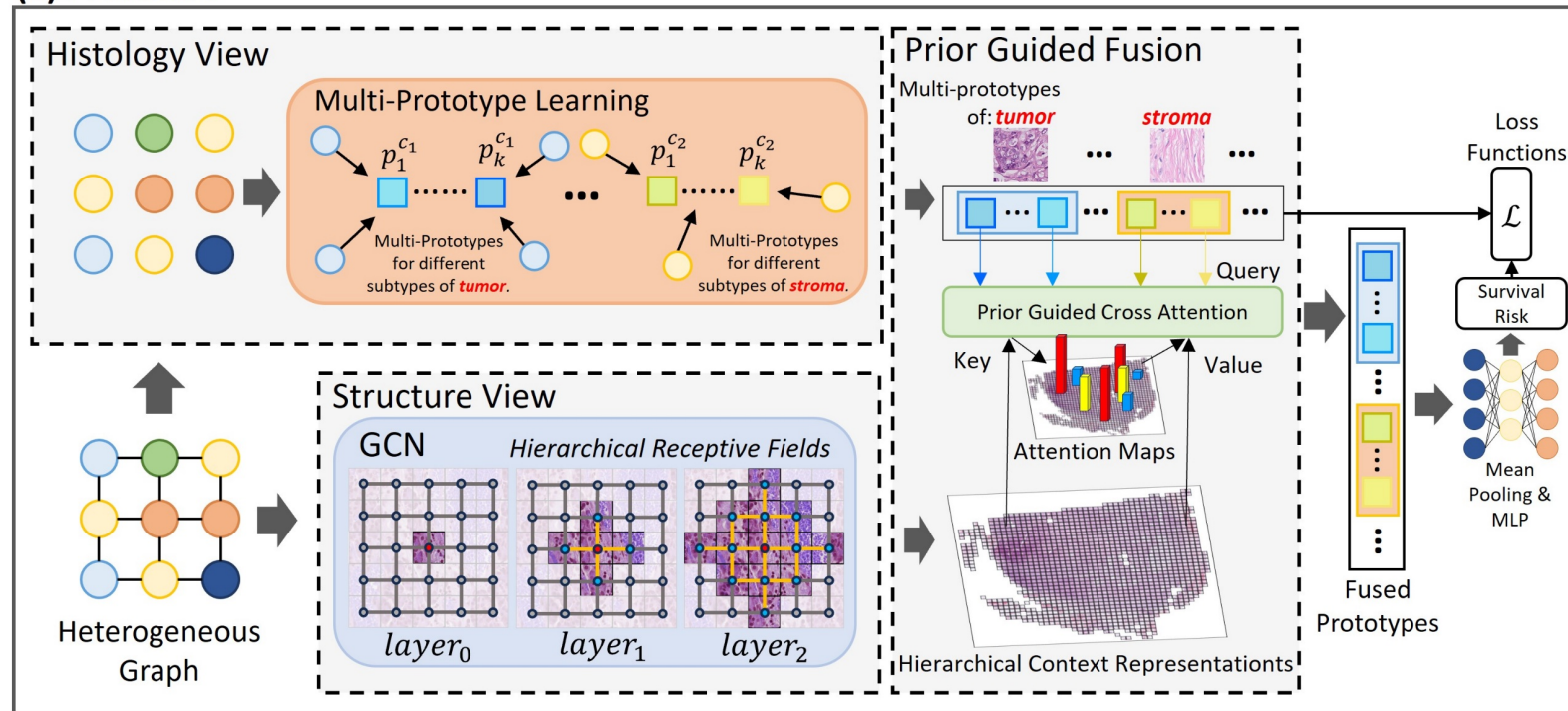
## (1) Preprocessing & Patching



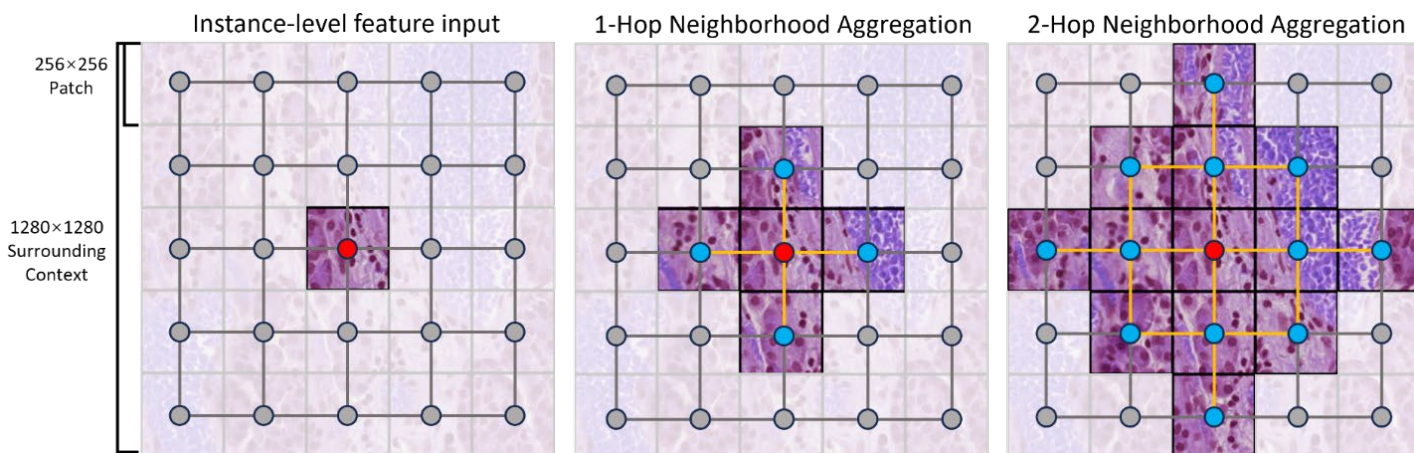
## (2) Heterogeneous Graph Construction



## (3) ProtoSurv

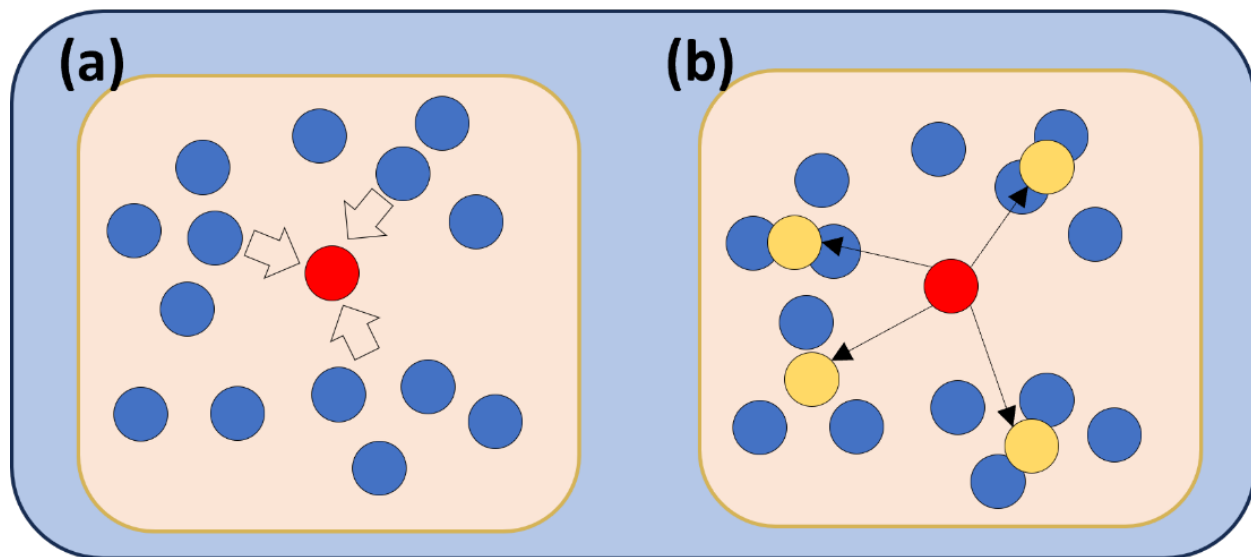


# Our Method: Component Details



**Illustration of the hierarchical receptive fields of GNN within the **Structure View**.**

We leverage the characteristic of GNNs of aggregating k-hop neighbor information in k-th layer to model the hierarchical receptive fields.



**Illustration of the multi-prototypes within the **Histology View**.** We extract multiple prototypes for each tissue category to obtain more diverse information.



# Summary

- To incorporate **pathological priors** within the MIL pipeline, we propose **ProtoSurv**, which decouples the heterogeneous graph into Structure View (SV) and Histology View (HV),
- **SV** re-models the lost spatial information between patches
- **HV** uses a prototype network to guide the model to focus on intratumoral tissues relevant to prognosis based on priors.

**THANK YOU!**