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# Sm: enhanced localization in Multiple Instance Learning for medical imaging classification

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# Multiple Instance Learning (MIL)

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- Bag:  $\mathbf{X} = [\mathbf{x}_1, \dots, \mathbf{x}_N]^\top \in \mathbb{R}^{N \times P}$ ,  $\mathbf{x}_n \in \mathbb{R}^P$ .
- Instance labels (**not observed**):  $\{y_1, \dots, y_N\} \subset \{0, 1\}$ .
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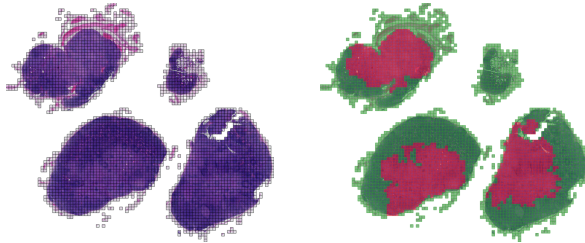
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**Why is it useful?** Minimal annotation effort.

# MIL in medical imaging

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**Figure:** Whole Slide Image (WSI, bag) and labeled patches (instances).

# MIL in medical imaging

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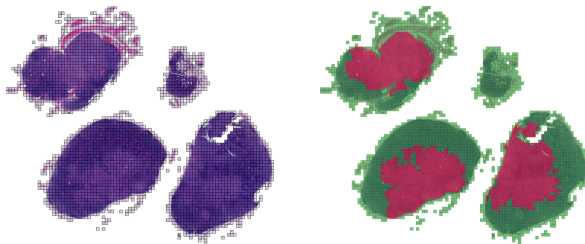


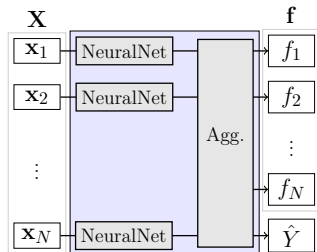
Figure: Whole Slide Image (WSI, bag) and labeled patches (instances).



Figure: Computerized Tomography (CT) scan (bag) and labeled slices (instances).

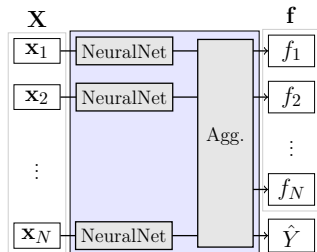
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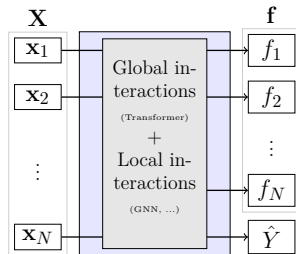
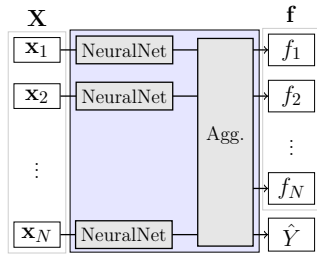


- **Attention values** ( $f_n \in \mathbb{R}$ ) are used as a proxy to estimate the instance labels.



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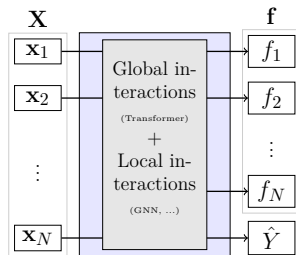
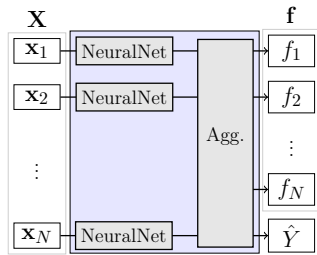
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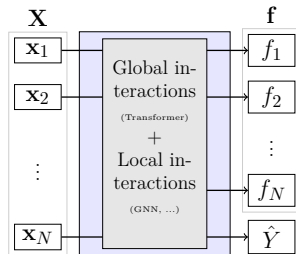
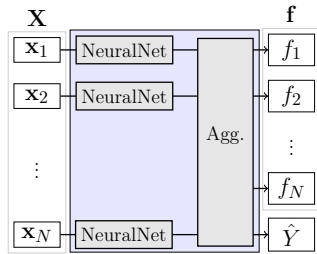
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- **Attention values** ( $f_n \in \mathbb{R}$ ) are used as a proxy to estimate the instance labels.
- Interactions have shown to improve the classification performance.
- **Problem:** previous works have been designed to target the classification task... what about **localization**?

## Method: the idea

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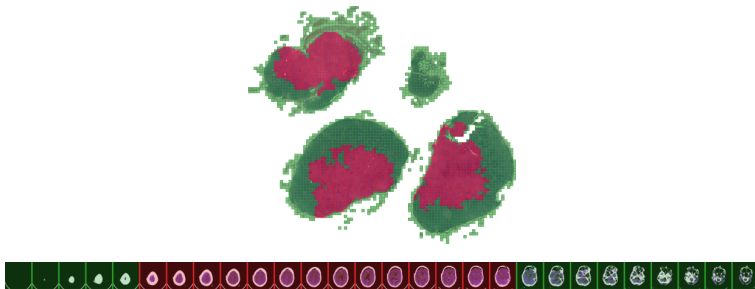


Figure: Map of labeled instances.

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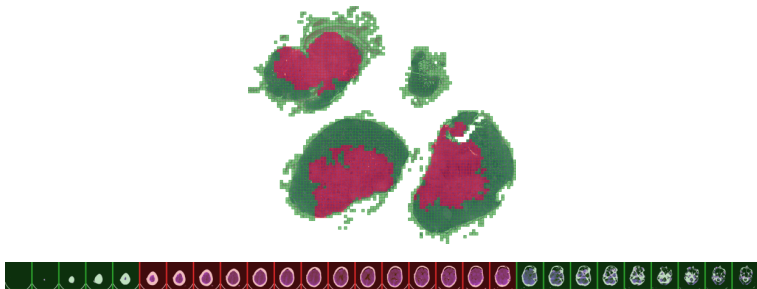


Figure: Map of labeled instances.

- Instance labels show **spatial dependencies**: an instance is likely to be surrounded by instances with the same label.
- Attention values  $f_n$  should inherit this **smoothing** property... **How?**

## Method: modelling the smoothness

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Let  $\mathbf{f} \in \mathbb{R}^N$  be attention values; interpreted as a function defined on a graph.

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**Bounding**  $\mathcal{E}_D(\mathbf{f})$ .

- $\mathcal{E}_D(\mathbf{f})$  is bounded by the Dirichlet energy of previous layers.
- **Consequence:** We can act on  $\mathbf{f}$  itself and/or on the output of previous layers.

## Method: Smooth operator ( $\mathbf{Sm}$ )

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Given  $\mathbf{U} \in \mathbb{R}^{N \times D}$ , the Smooth operator ( $\mathbf{Sm}$ ) is defined as

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**Theoretical guarantees.** If  $\mathbf{L}$  is the normalized Laplacian matrix, then

$$\mathcal{E}_D(\text{Sm}(\mathbf{U})) < \mathcal{E}_D(\mathbf{U}).$$

**Consequence:** It can be used in the different layers of a neural network to decrease  $\mathcal{E}_D$ .

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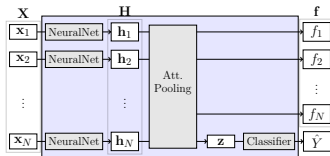
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**Avoiding matrix inversion.** It holds that

$$\begin{aligned} \text{Sm}(\mathbf{U}) &= \lim_{t \rightarrow \infty} \mathbf{G}(t), \\ \mathbf{G}(0) &= \mathbf{U}; \quad \mathbf{G}(t) = \alpha (\mathbf{I} - \mathbf{L}) \mathbf{G}(t-1) + (1 - \alpha) \mathbf{U}. \end{aligned}$$

# Method: the proposed model

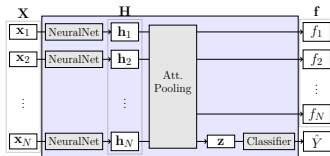
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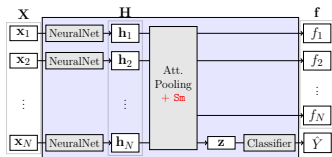
(a) ABMIL, the baseline.

# Method: the proposed model

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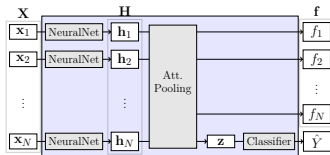
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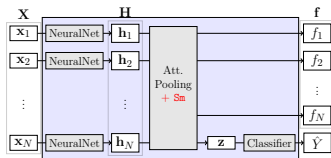
(b) SmAP.

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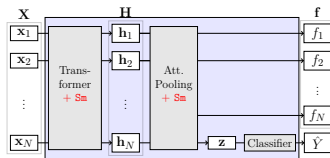
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(a) ABMIL, the baseline.



(b) SmAP.



(c) SmTAP.



# Experiments

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- 3 different medical imaging datasets:  
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- 3 different medical imaging datasets: RSNA (CT scans), PANDA (WSIs), and CAMELYON16 (WSIs).
- 4 different feature extractors, with and without self-supervised learning.
- Up to 13 different SOTA methods considered for comparison.
- **Results:** the proposed methods with **Sm** achieve the best performance in localization and remain very competitive in classification.

**Table:** Average rank (lower is better).

		Instance localization	Bag classification
Without global interactions	<b>SmAP</b>	<b>1.500</b> <sub>0.548</sub>	<b>1.833</b> <sub>0.753</sub>
	ABMIL	<u>2.500</u> <sub>1.225</sub>	2.500 <sub>1.049</sub>
	CLAM	4.167 <sub>1.329</sub>	4.500 <sub>0.837</sub>
	DSMIL	4.333 <sub>0.516</sub>	4.167 <sub>0.753</sub>
	DFTD-MIL	2.500 <sub>1.049</sub>	<u>2.000</u> <sub>1.265</sub>
With global interactions	<b>SmTAP</b>	<b>1.500</b> <sub>1.225</sub>	<b>1.833</b> <sub>0.983</sub>
	TransMIL	3.083 <sub>1.429</sub>	4.083 <sub>0.917</sub>
	SETMIL	3.667 <sub>0.816</sub>	3.583 <sub>2.010</sub>
	GTP	3.917 <sub>1.429</sub>	2.750 <sub>0.987</sub>
	CAMIL	<u>2.833</u> <sub>1.169</sub>	<u>2.750</u> <sub>1.173</sub>

## Experiments: WSI visualization.

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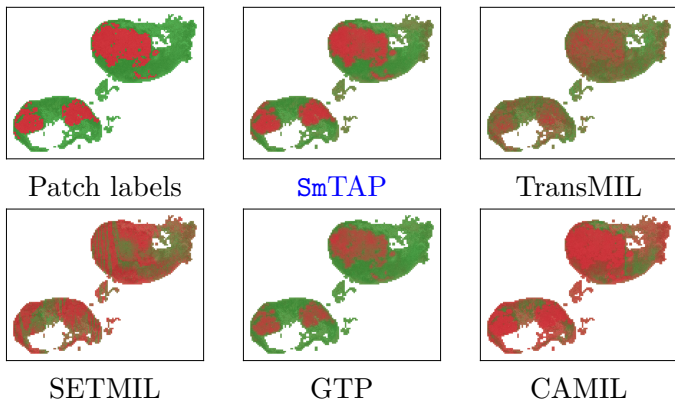


Figure: Attention maps on CAMELYON16. The novel SmTAP produces the most accurate map.

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- The proposed  $S_m$  introduces local interactions in a principled way.
- It achieves the best performance in localization while being highly competitive in classification.
- Future work: MIL methods need to quantify uncertainty so they can be deployed in clinical settings.

**Thank you!**