



École de technologie supérieure  
Department of Software Engineering  
and Information Technology  
Neuro-iX



Presentation of Ten Papers

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## **Towards Automated Neuroanatomy : Segmentation and Landmark Localization**

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1

# General Introduction



# Overview of the Core Problem Explored in the 10 Articles

- **Anatomical Segmentation:**

Precise Identification of Structures in Medical Images

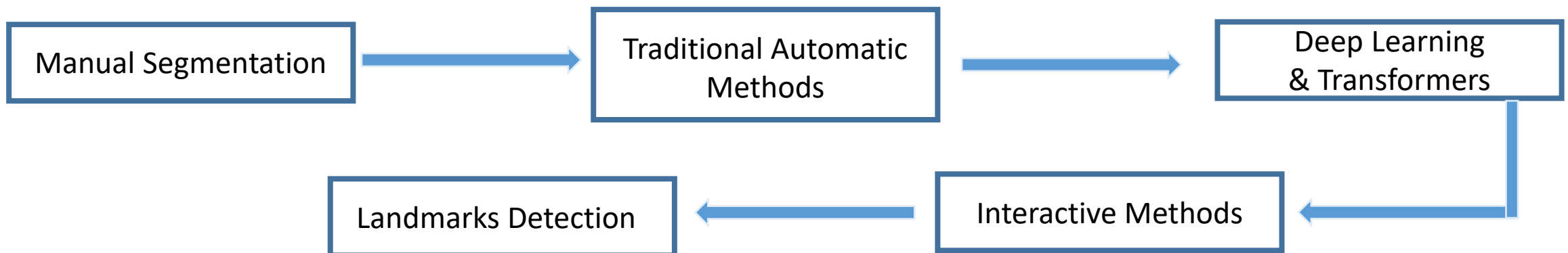
- **Importance:**

A fundamental step in many clinical analyses (e.g., volume and shape measurements), and essential for training AI models.

- **Challenges:**

Manual segmentation is time-consuming, subject to inter- and intra-rater variability, and lacks scalability — highlighting the urgent need for automation without compromising quality.

- **The Path We Follow: Segmentation Methodologies Over Time**





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# Manual Segmentation:

(Paper 1)



# Manual Segmentation – Reliability and Limitations

## Results

- High inter- and intra-rater reliability (Dice Coefficient > 0.90 for most structures).
- Detected cerebral asymmetries:  
Ex :larger nucleus accumbens on the left, hippocampus larger on the right.
- Sex-based anatomical differences: subcortical structures appear larger in males.

## Limitations

- Extremely time- and labor-intensive.
- Difficult to scale for large datasets.
- Strong dependence on expert annotators.

Region of interest (ROI)	Mean Dice	SD	Min	Max
Lateral Ventricle Left	0.95	0.02	0.92	0.98
Lateral Ventricle Right	0.95	0.02	0.93	0.98
Third Ventricle	0.84	0.05	0.75	0.90
Fourth Ventricle	0.87	0.04	0.80	0.94
Nucleus Accumbens Left	0.84	0.04	0.78	0.89
Nucleus Accumbens Right	0.84	0.05	0.76	0.93
Caudate Left	0.93	0.01	0.91	0.96
Caudate Right	0.93	0.02	0.88	0.96
Putamen Left	0.93	0.02	0.91	0.99
Putamen Right	0.93	0.02	0.91	0.99
Globus Pallidus Left	0.83	0.04	0.76	0.90
Globus Pallidus Right	0.81	0.06	0.73	0.90
Brainstem	0.95	0.01	0.94	0.98
Thalamus Left	0.88	0.04	0.78	0.92
Thalamus Right	0.88	0.03	0.82	0.93
Ventral Diencephalon Left	0.88	0.02	0.84	0.92
Ventral Diencephalon Right	0.88	0.01	0.85	0.90
Inferior Horn of Lateral Ventricle Left	0.72	0.05	0.61	0.82
Inferior Horn of Lateral Ventricle Right	0.72	0.05	0.61	0.81
Hippocampal Formation Left	0.87	0.03	0.82	0.90
Hippocampal Formation Right	0.87	0.02	0.82	0.90
Amygdala Left	0.84	0.03	0.78	0.88
Amygdala Right	0.80	0.05	0.71	0.88
Fifth Ventricle	0.76	0.07	0.65	0.84
Optic Chiasm	0.74	0.15	0.54	0.95

**Inter-rater reliability**

Region of interest (ROI)	Mean Dice	SD	Min	Max
Lateral Ventricle Left	0.95	0.02	0.93	0.97
Lateral Ventricle Right	0.96	0.03	0.93	0.98
Third Ventricle	0.89	0.04	0.84	0.91
Fourth Ventricle	0.90	0.03	0.87	0.93
Nucleus Accumbens Left	0.87	0.04	0.82	0.89
Nucleus Accumbens Right	0.89	0.02	0.87	0.91
Caudate Left	0.93	0.03	0.91	0.96
Caudate Right	0.94	0.02	0.92	0.95
Putamen Left	0.94	0.02	0.92	0.95
Putamen Right	0.94	0.02	0.91	0.95
Globus Pallidus Left	0.81	0.05	0.77	0.86
Globus Pallidus Right	0.80	0.05	0.76	0.85
Brainstem	0.96	0.01	0.95	0.97
Thalamus Left	0.91	0.01	0.91	0.92
Thalamus Right	0.91	0.03	0.88	0.93
Ventral Diencephalon Left	0.90	0.01	0.89	0.91
Ventral Diencephalon Right	0.90	0.01	0.89	0.91
Inferior Horn of Lateral Ventricle Left	0.76	0.09	0.68	0.86
Inferior Horn of Lateral Ventricle Right	0.80	0.04	0.75	0.83
Hippocampal Formation Left	0.90	0.02	0.89	0.93
Hippocampal Formation Right	0.90	0.05	0.85	0.94
Amygdala Left	0.84	0.07	0.77	0.91
Amygdala Right	0.84	0.07	0.76	0.90
Fifth Ventricle	0.75	0.08	0.66	0.83
Optic Chiasm	0.87	0.18	0.66	0.99

**Intra-rater reliability**





3

# Traditional Automatic Segmentation

(Papers 2 & 3)



# Traditional Automatic Segmentation - Probabilistic Atlas-

## Méthode de Fischl et al. (2002) :

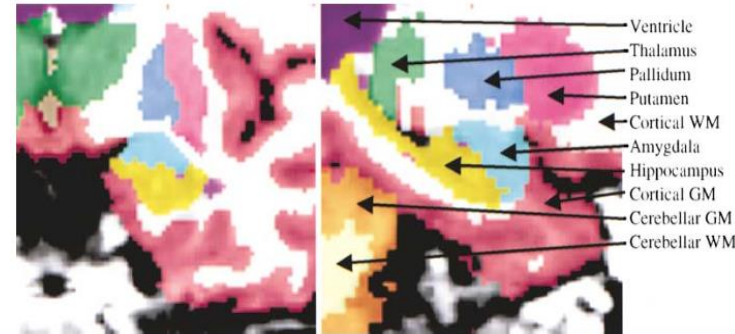
- First algorithm for whole-brain automatic labeling.
- Implemented in FreeSurfer — a reference tool in neuroimaging

## Principle of atlas-based segmentation :

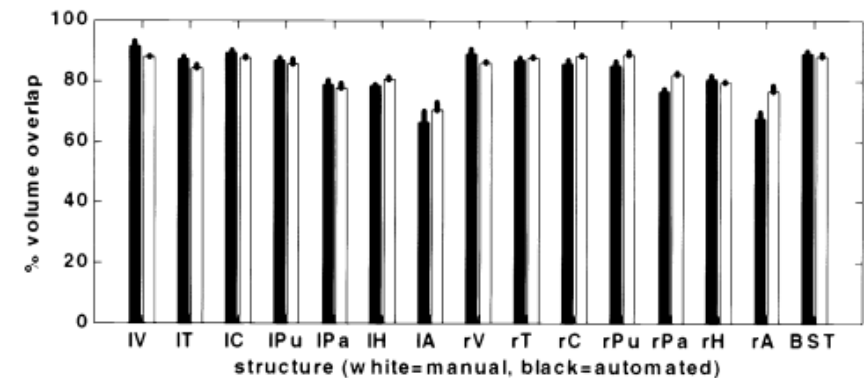
1. Statistical Atlas Construction :
  - Each voxel in the probabilistic atlas encodes intensity, location, and spatial context for anatomical labeling.
2. MRI-to-Atlas Registration :
  - Each Aligns MRI scans to the atlas reference space.
3. Bayesian Voxel-Wise Labeling :
  - Bayesian MAP estimation to assign the most probable tissue label
  - Markov Random Field modeling to enforce anatomical consistency across neighboring voxels

## Results :

- High accuracy (~90% Dice Score) for subcortical structures; detects subtle disease-related changes (e.g., in Alzheimer's)
- Limitation: Sensitive to registration errors; lacks modeling of inter-individual anatomical variability (single average atlas)

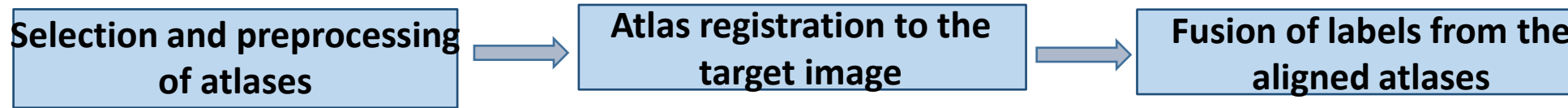


$$L = \arg \min_L \int (T(r) - I(Lr))^2 dr$$



# Traditional Automatic Segmentation -Multi-Atlas Label Fusion-

- Uses multiple manually segmented atlases to enhance segmentation accuracy.
- Each atlas is individually registered to the target MRI, and labels are fused to generate a consensus segmentation.



## JLF - Joint Label Fusion (Method proposed by the paper)

The main idea is that some atlases may have similar errors, and these dependencies must be taken into account.

## Improvement with Multi-Atlas Label Fusion

- Improved accuracy: +1.5% Dice Score for hippocampal segmentation compared to single-atlas methods.

- Key result: Multi-atlas fusion improves segmentation accuracy while reducing dependence on a single reference atlas.

$$M_x(i, j) = \mathbb{E}[\delta_i(x)\delta_j(x)] \propto \left[ \sum_{y \in \mathcal{N}(x)} |F_T(y) - F_i(y)| |F_T(y) - F_j(y)| \right]^\beta.$$

$$\mathbf{w}_x = \arg \min_{\mathbf{w}_x} \mathbf{w}_x^T M_x \mathbf{w}_x$$

$$\hat{S}_T(x) = \sum_{i=1}^n w_i(x) S_i(x)$$

Méthode	Average Dice Score (%)
Majority Voting (MV)	85.2
Joint Label Fusion (JLF - Proposé)	89.9



4

# Transformer-Based Segmentation

(Papers 4 & 5)



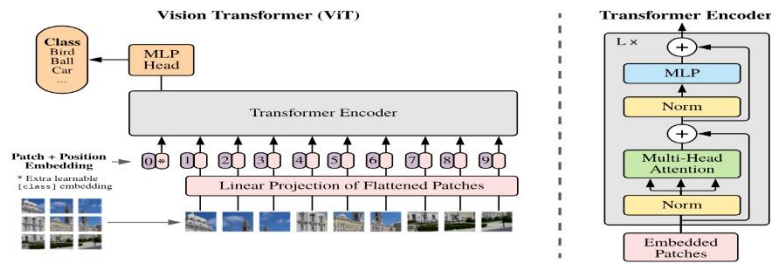
# Transformer-Based Segmentation

## Motivation in Medical Imaging

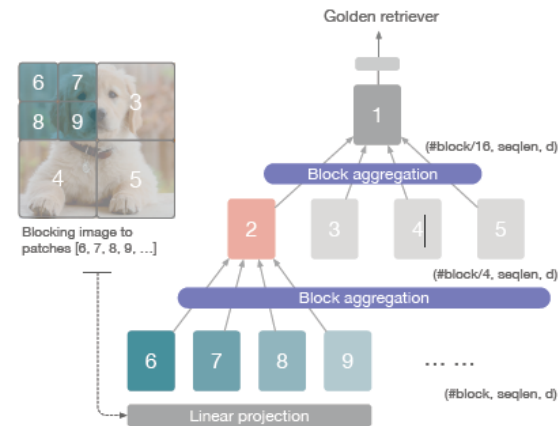
- Anatomy is contextual: Brain structures are interdependent and spatially organized
- Global context matters: A local anomaly is often meaningful only in relation to surrounding regions
- CNNs are limited by small receptive fields → Transformers provide a broader, global view

## Advances in Transformer Architectures

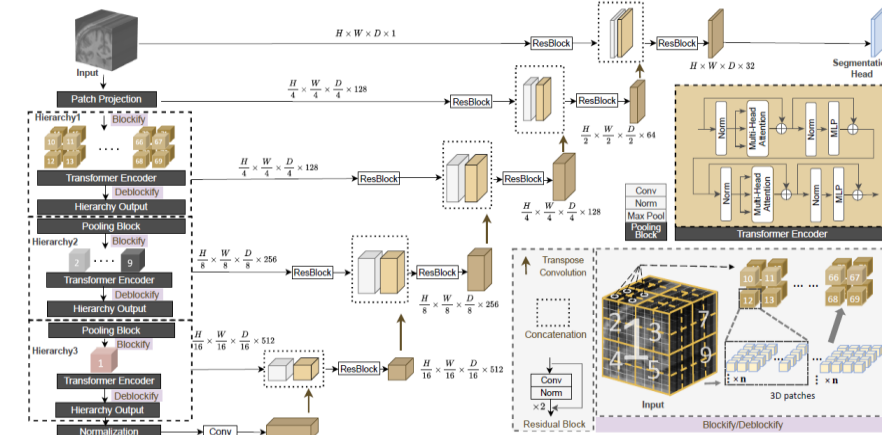
### Vision Transformer (ViT)



### Nested Transformer (NesT)



### UNesT



# Vision Transformer (ViT) - Principle and Architecture

- Inspired by Natural Language Processing (NLP) Transformers, ViT applies self-attention mechanisms to images without relying on convolutions.

1. Splitting the image into N patches of size (P×P):  $N = \frac{H \times W}{P^2}$ .

2. Patch Encoding: each patch  $x_i$  is flattened and projected into an embedding vector.

3. Addition of the classification token and positional embeddings.

$$z_0 = [x_{\text{class}}; x_p^1 E; x_p^2 E; \dots; x_p^N E]. \quad z_0 = z_0 + E_{\text{pos}}$$

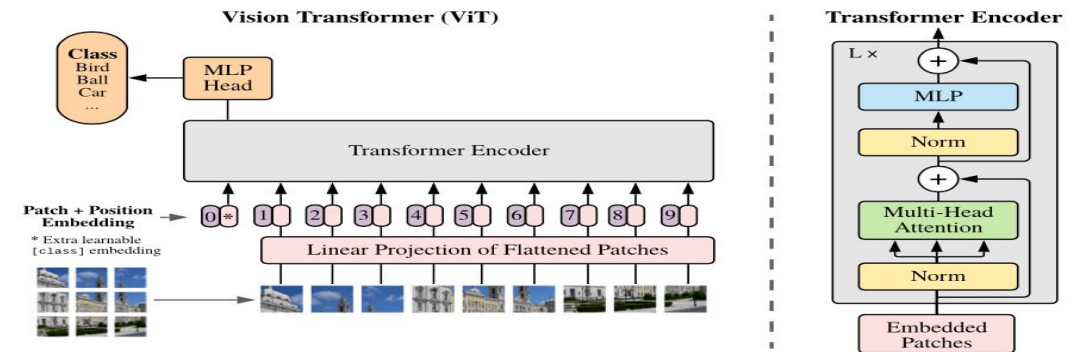
4. Transformer Encoder: Stacked  $L$  layers of Multi-Head Self-Attention (MSA) and Multi-Layer Perceptron (MLP).

$$z'_\ell = \text{MSA}(\text{LN}(z_{\ell-1})) + z_{\ell-1}, \quad z_\ell = \text{MLP}(\text{LN}(z'_\ell)) + z'_\ell.$$

5. Final representation of the classification token (first position of  $z_L$ ) is normalized to obtain the prediction.

➤ Limitation:

- Performs well on large datasets (e.g., ImageNet-21k, JFT-300M), but struggles on small datasets.



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AN IMAGE IS WORTH 16X16 WORDS:  
TRANSFORMERS FOR IMAGE RECOGNITION AT SCALE

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## ABSTRACT

While the Transformer architecture has become the de facto standard for natural language processing tasks, its application to computer vision remains limited. In vision, attention is either applied in conjunction with convolutional networks, or used to replace certain components of convolutional networks while keeping their overall structure in place. We show that this inductor on CNNs is not necessary and a pure transformer applied directly to sequences of image patches can perform very well on image classification tasks. When pre-trained on large amounts of data and maintained to maintain mid-sized or small image recognition benchmarks (ImageNet, CIFAR-100, VTAB, etc.), Vision Transformer (ViT) attains excellent results compared to state-of-the-art convolutional networks while requiring substantially fewer computational resources to train.

## 1 INTRODUCTION

Self-attention based architectures, in particular Transformers (Vaswani et al., 2017), have become the model of choice in natural language processing (NLP). The dominant approach is to pre-train on a large text corpus and then fine-tune on a smaller task-specific dataset (Devlin et al., 2019). Thanks to Transformers’ computational efficiency and scalability, it has become possible to train models of unprecedented size with over 1000 parameters (Brown et al., 2020; Lipton et al., 2020). With the models and datasets growing, there is still no sign of saturating performance.

In computer vision, however, convolutional architectures remain dominant (Lecun et al., 1998; Krizhevsky et al., 2012; He et al., 2016). Inspired by NLP successes, multiple works try combining CNN-like architectures with self-attention (Chen et al., 2018; Carion et al., 2019), some replacing the convolutional entirely (Dosovitskiy et al., 2019; Wang et al., 2020). The latter models, while theoretically efficient, have not yet been scaled effectively on modern hardware accelerators due to the use of specialized attention patterns. Therefore, in large-scale image recognition, classic ResNet-like architectures are still state of the art (Malinin et al., 2019; Xie et al., 2019; Kolesnikov et al., 2020).

Inspired by the Transformer scaling successes in NLP, we experiment with applying a standard Transformer directly to images, with the fewest possible modifications. To do so, we split an image into patches and provide the sequence of their embeddings of these patches as an input to a Transformer. Image patches are treated the same way as tokens (words) in an NLP application. We train the model on image classification in supervised fashion.

When trained on mid-sized datasets such as ImageNet without strong regularization, these models yield model accuracies of a few percentage points below ResNets of comparable size. This seemingly disappointing outcome may be expected: Transformers lack some of the inductive biases

This training code and pre-trained models are available at [https://github.com/google-research/vision\\_transformer](https://github.com/google-research/vision_transformer)

arXiv:2010.11929v2 [cs.CV] 3 Jun 2021

# Nested Transformer (NesT) – A Hierarchical Improvement

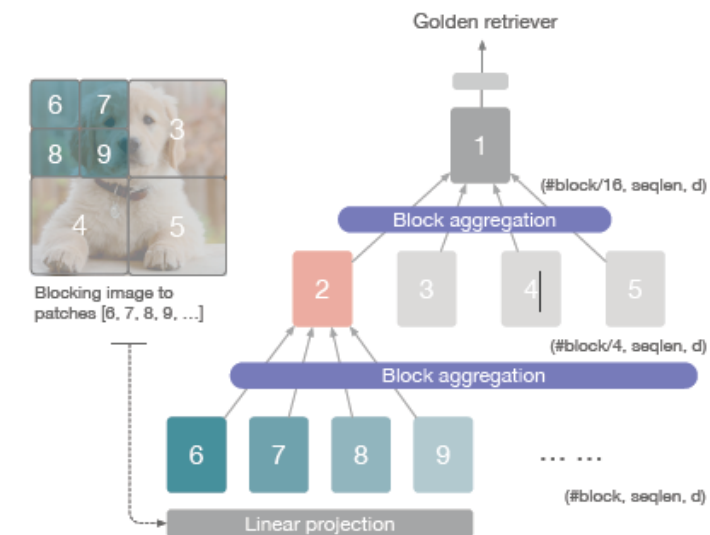
➤ NesT introduces a nested hierarchical organization to better capture spatial relationships.

1. Image partitioning into blocks (instead of individual patches).
2. Local processing: Each block is independently analyzed by a local Transformer.
3. Hierarchical block fusion: Gradual aggregation of blocks to capture global context.

- Progressive aggregation using 3×3 convolutions and max-pooling.
- Ensures better integration of local and global information.
- Each set of 4 neighboring blocks is merged into a higher-level block.
- Gradually reduces the total number of blocks.

➤ Advantages over ViT:

- Requires less data: Achieves better results on smaller datasets (CIFAR-10, ImageNet).
- Faster training: Thanks to its nested hierarchical structure.





# UNesT – A U-Net with Nested Transformers for 3D Segmentation

## Hierarchical Encoder :

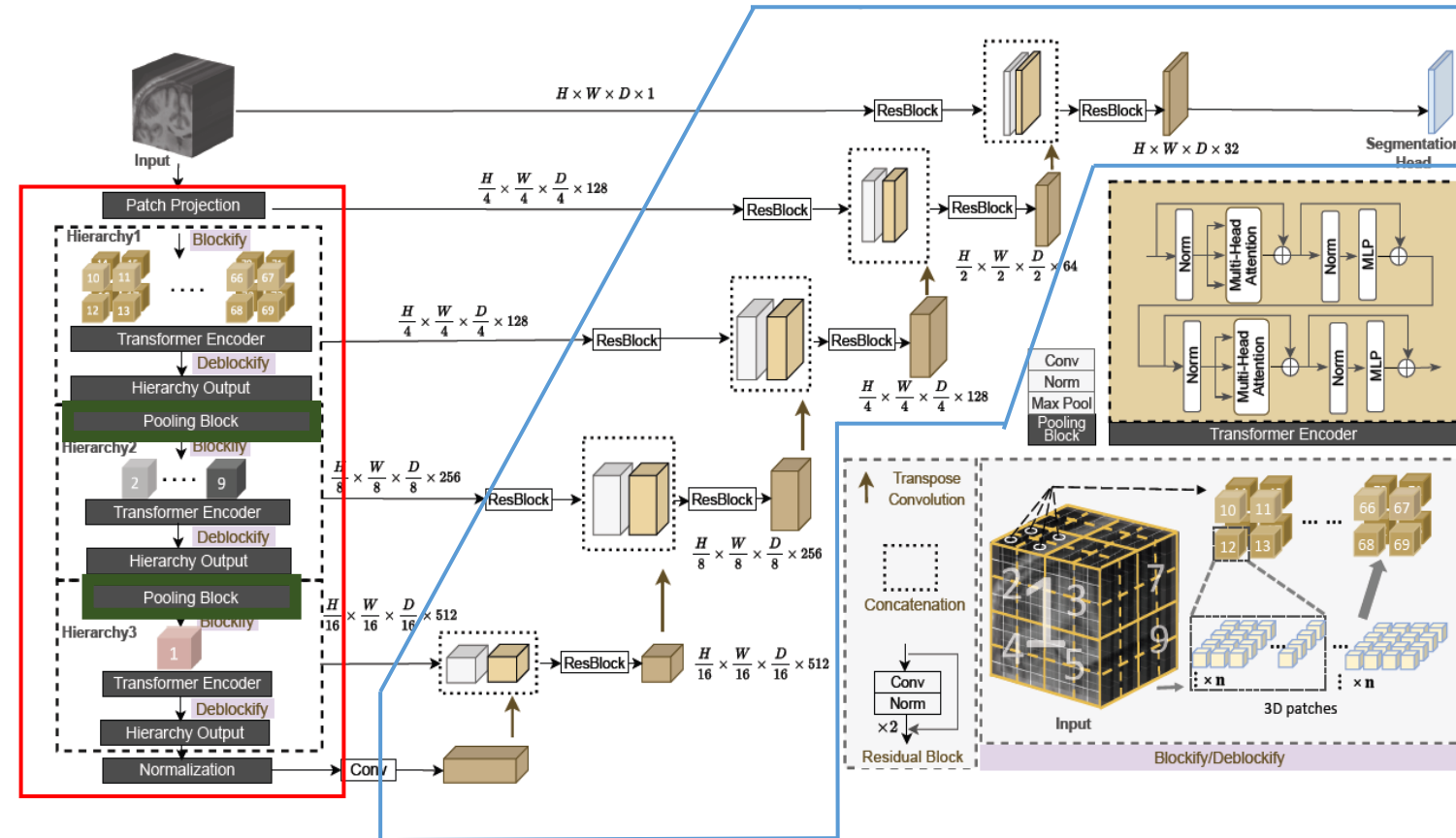
- 3D input volume is split into  $4 \times 4 \times 4$  patches, each flattened into a 128-dimensional vector.
- Patches are grouped into non-overlapping blocks for independent local processing.
- Each block is processed by a mini-transformer (MSA, MLP, LN) with skip connections between layers.

## 3D aggregation :

- Merges neighboring blocks to capture global spatial context with low memory cost.

## Hierarchical Convolutional Decoder :

- At each ascending level, features are upsampled, merged with encoder outputs, and refined toward final segmentation.
- Processes and fuses features at every level to progressively reconstruct the segmentation map.



# Comparative Results of Transformer Architectures

Criterion	ViT (2020)	NesT (2022)	UNesT (2023)
Structure	Flat, non-hierarchical Transformer	Hierarchical nested Transformers (2D)	Hierarchical 3D Transformers with convolutional decoder
Global context	Fully captured from the beginning	Progressively captured across levels	Captured at all hierarchical levels
Local detail fidelity	Low (no pyramid structure)	Moderate (nested aggregation)	Very high (fine resolution, skip connections)
Data requirements	Very high (requires massive pretraining)	Moderate (pretraining helpful but optional)	Moderate (hierarchical structure performs well with limited data)
Evaluated tasks	Classification, basic segmentation	2D/3D multi-organ segmentation	Fine volumetric segmentation (133 structures)
Performance (Dice)	Around <u>83%</u>	<u>86–88%</u> depending on task	<u>91–93%</u> on complex brain structures
Advantages	Simplicity, global view, few parameters	Good global/local balance, generalizable	High accuracy, context + detail, native 3D support
Limitations	Low spatial resolution, high data demand	Not natively 3D, nested complexity	High computational cost, long training time



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# Interactive Segmentation

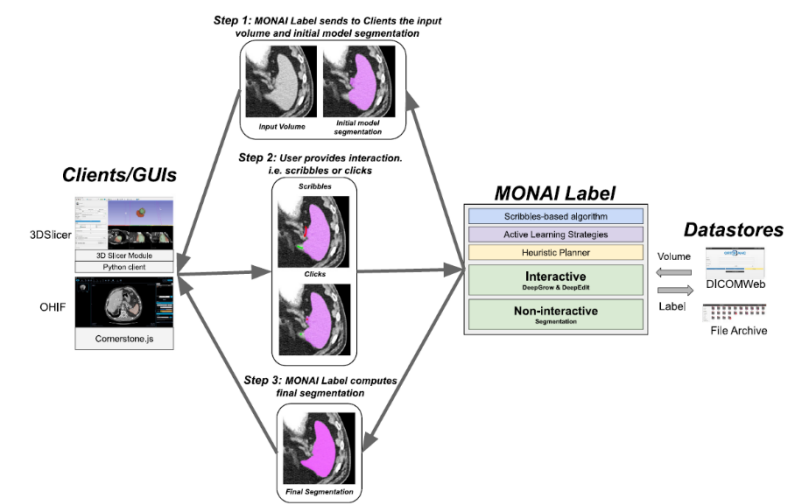
(Papers 7 & 8)



# Monai Label

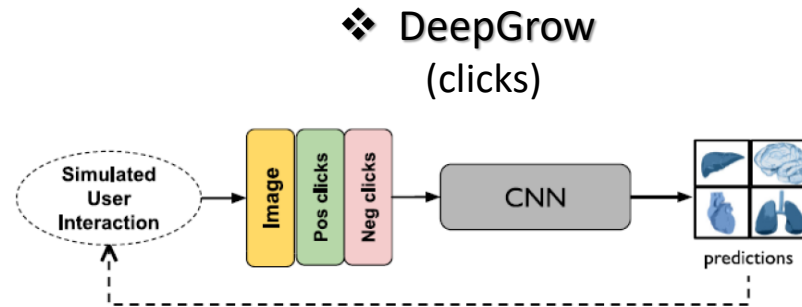
Open-source client-server platform for AI-assisted medical annotation

- Clients (3D Slicer, OHIF web): for expert interaction
- AI Server (MONAI Label): hosts pre-trained segmentation models and supports online learning strategies



## Non-interactive approach

(U-Net 2D/3D, HighResNet ou DynUNet)



- User clicks (positive/negative) are converted into binary maps added to the input image
- The model refines segmentation dynamically based on these contextual cues

## Interactive approach

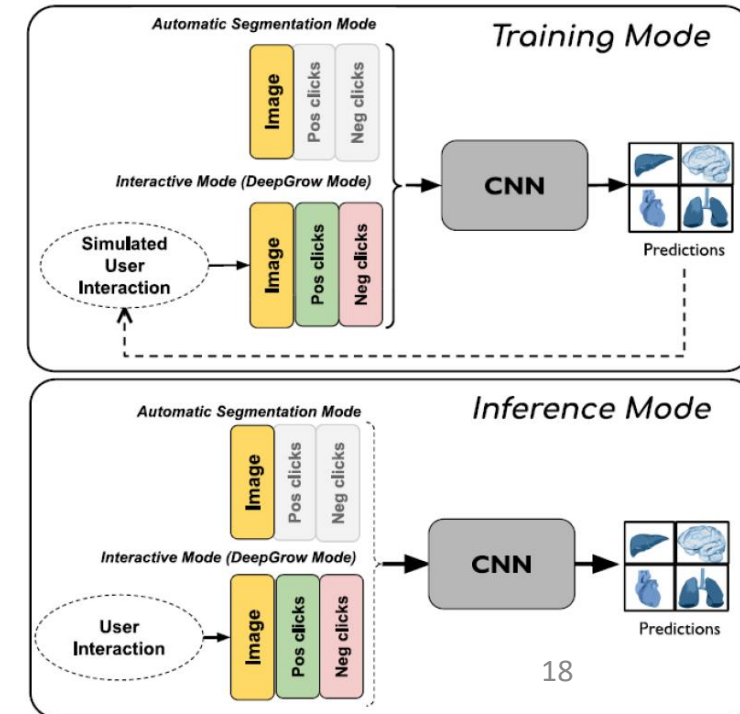
### Two-phase process:

1. Auto mask on load (no clicks)
2. User refines with corrective clicks

### Dual-mode design:

- Trained with/without clicks
- Switches between auto and interactive modes
- Enables active learning with initial masks

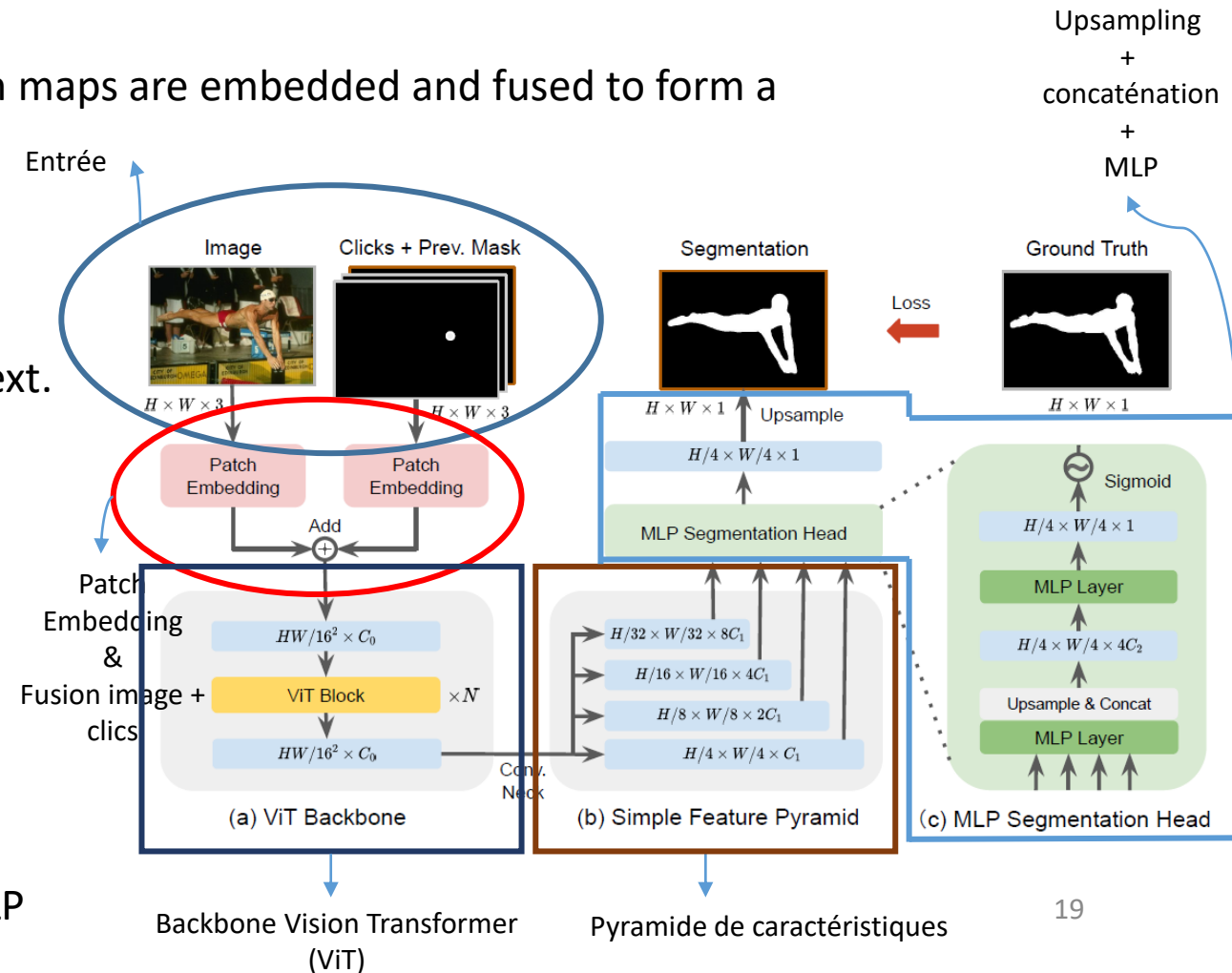
## DeepEdit (auto+clicks)



# SimpleClick

Interactive segmentation method based on Vision Transformer without hierarchical backbone

- User Click Encoding:** Generate two binary maps from clicks (positive on object, negative on background) plus a prediction map from the previous segmentation.
- Symmetric Patch Embedding:** Both image and interaction maps are embedded and fused to form a unified ViT input.
- Window-based local attention:** Attention is restricted to non-overlapping windows, reducing computational cost while preserving local context.
- Global attention blocks:** Introduced at selected layers to model long-range dependencies and ensure global context understanding.
- Multi-scale features generated via parallel convolutions with different strides, followed by upsampling and concatenation.**
- Final segmentation map predicted through a unified MLP and sigmoid activation.**



# Comparative Analysis: MONAI Label (DeepGrow, DeepEdit) vs SimpleClick

Criterion	DeepGrow (MONAI Label)	DeepEdit (MONAI Label)	SimpleClick (2023)
Core principle	Learns from a single seed point	Multiple user-guided corrections	ViT interprets positive/negative clicks
Interaction type	One central click	Multiple clicks (include/exclude)	Positive/negative clicks only
Spatial precision	High, depends on seed accuracy	Good, refined with interaction	Very high (ViT-based attention)
AI architecture	CNN encoder-decoder	CNN with correction-aware decoder	Vision Transformer pre-trained with MAE
Training data	Requires manual segmentations	Same, with local interaction examples	General pretraining + minimal tuning
Clicks vs. quality	3–6 clicks for decent quality	~5 clicks for 90% Dice	4.15 clicks for 90% IoU
2D / 3D support	Full (2D and 3D)	Full (with Slicer/MITK/OHIF)	Mainly 2D (applied per slice in 3D)
Interaction speed	Good (depends on server)	Fast and interactive	Very fast (lightweight ViT)
Deployment complexity	Medium to high (server-client)	Same as DeepGrow	Moderate (standalone, portable)





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# **Automatic Landmark Detection**

**(Papers 9 & 10)**

# Automatic Landmark Detection – Why It Matters

**Anatomical landmark** : A key point defined by the position of a structure (e.g., anterior commissure, etc)

**Use cases** :

- Image registration (align scans via shared points)
- Biometric measurements (e.g., distances)
- Extraction of standard planes in 3D imaging

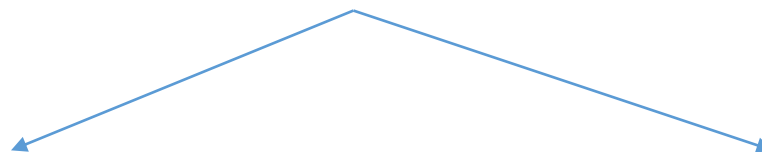
**Problem** : Manual localization is tedious and prone to high inter-operator variability.

**Challenges in automation** :

- Few annotated datasets (point labeling is as laborious as segmentation)
- Large 3D volumes
- Multiple landmarks to detect simultaneously, requiring anatomical relationship modeling.

**Solution** :

Dedicated deep learning approaches formulating landmark detection as a combined regression + classification problem for each point



PIN – Patch-based Iterative Network

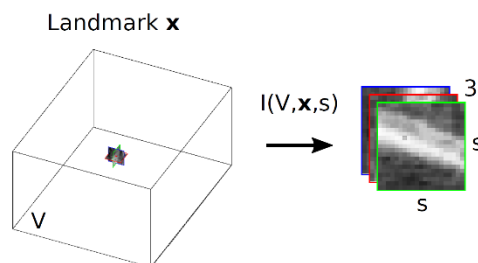
Global-to-Local Landmark Localization

# PIN – Patch-based Iterative Network

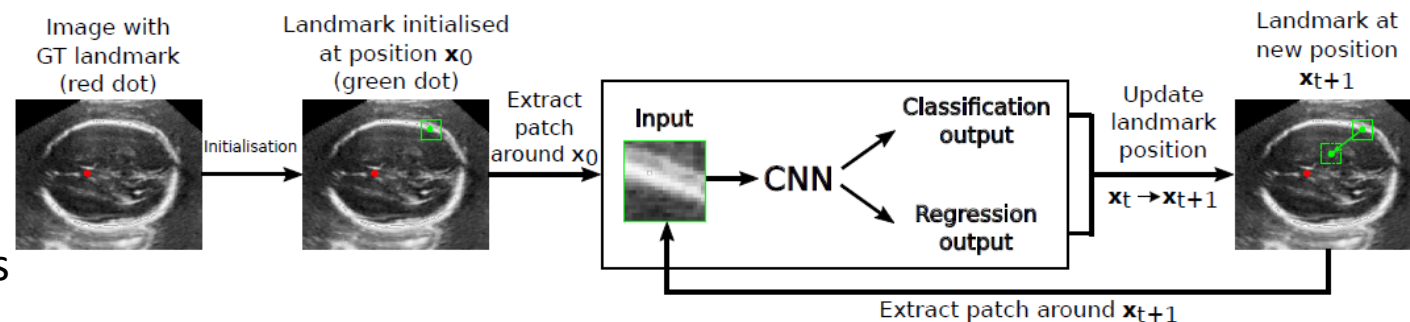
Formulates landmark localization as an iterative patch-based search.

Start from a coarse initial position, then:

- Extract a 2.5D local patch (axial, sagittal, coronal slices centered on current point)



- A CNN predicts:
  - A displacement vector  $\mathbf{d}=(\Delta x, \Delta y, \Delta z)$  toward the true point
  - A direction class  $P_{\max}$  (one of  $\pm X, \pm Y, \pm Z$ ) for the main movement axis
  - Update position :  $\mathbf{x}_{t+1} = \mathbf{x}_t + P_{\max} \cdot \mathbf{d}$ , repeat until convergence



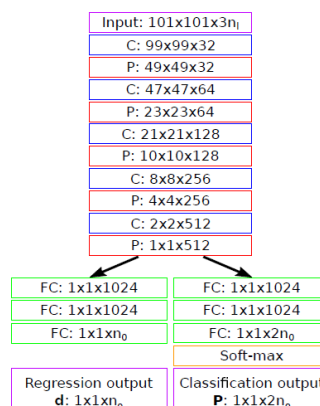
Multi-landmark extension

- All landmark coordinates are projected into a lower-dimensional space (PCA)

$$\mathbf{b} = \mathbf{W}^T (\mathbf{X} - \bar{\mathbf{X}}) \quad \mathbf{b}_{t+1} = \mathbf{b}_t + P_{\max} \cdot \mathbf{d}_b$$



Enables the CNN to predict a **compact global vector** for all points





# Global-to-Local Landmark Localization

Formulates landmark localization as an iterative patch-based search.

## Global Stage :

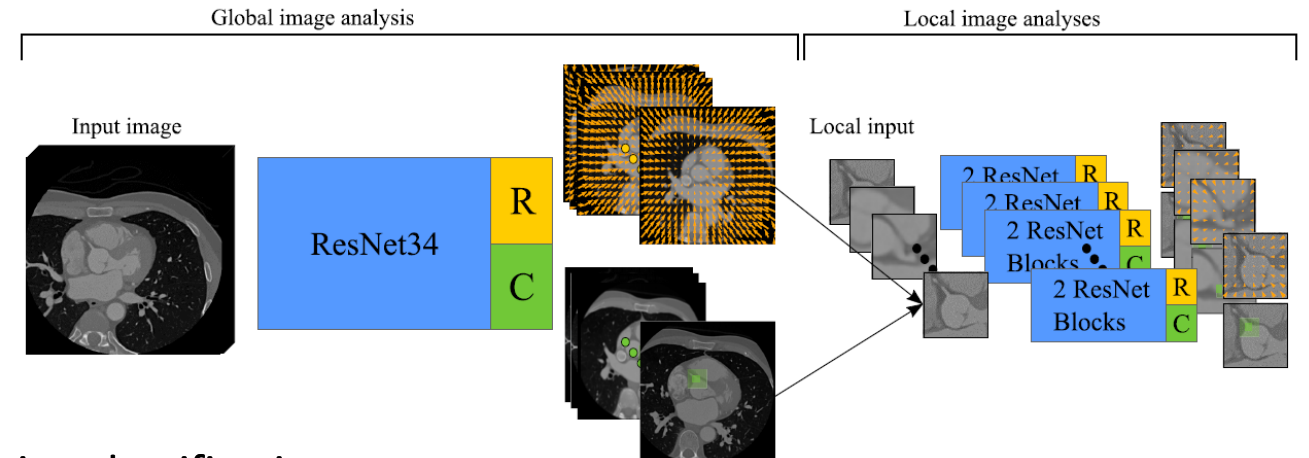
- A fully convolutional neural network (FCNN) analyzes the entire image in a patch-based manner and predicts:
  - A displacement vector from the patch center to the landmark
  - A presence probability for each landmark (classification)

Final position: weighted average of displacement vectors using classification scores

## Local Stage :

A specialized local FCNN analyzes a subvolume around each landmark to refine its position, again combining regression and classification

- For each landmark, extract a local region around the global estimate
- Apply a small local network to refine the coordinates through fine-grained search



$$\hat{p} = \frac{\sum_{i=1}^N s_i \cdot (c_i + d_i)}{\sum_{i=1}^N s_i}$$

## Performance Metrics for PIN vs Global-to-Local Approaches

Metric / Criterion	PIN (Li et al., 2018)	Global-to-Local (Noothout et al., 2020)
Mean localization error	5.59 mm (10 fetal brain landmarks)	2.0 mm (2D/3D datasets, expert-level)
Number of landmarks	10 localized jointly	Up to 19 localized jointly
Success rate ( $\geq 3$ mm)	Not reported	95% of landmarks within 3 mm
Inference time per volume	0.44 seconds (entire 3D volume)	Slower (global + local passes per point)
Initialization requirement	Requires approximate patch-centered input	No manual initialization needed
Computational cost	Low (few patches per point)	High (full volume + refinements)

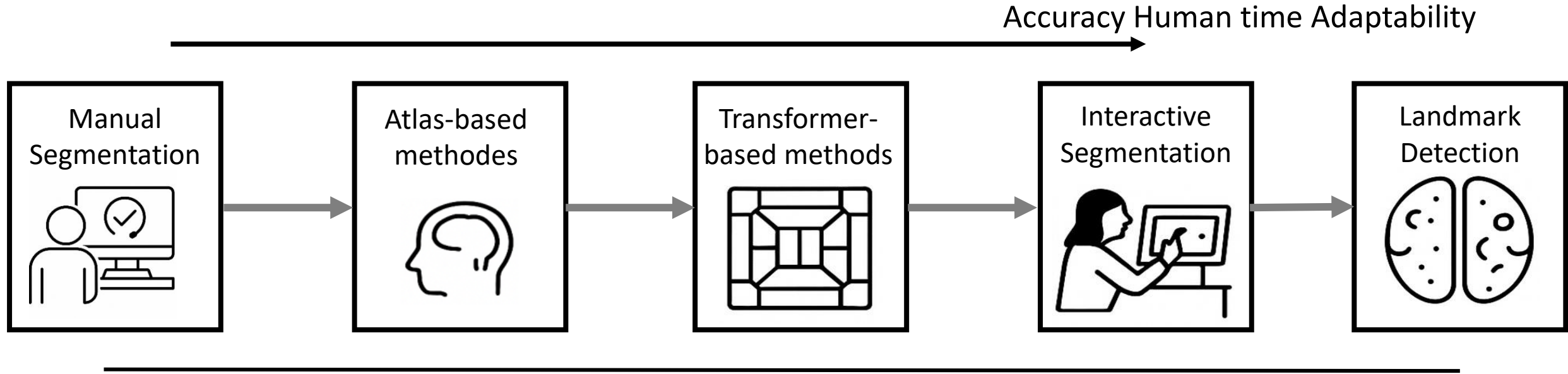


7

# Conclusion



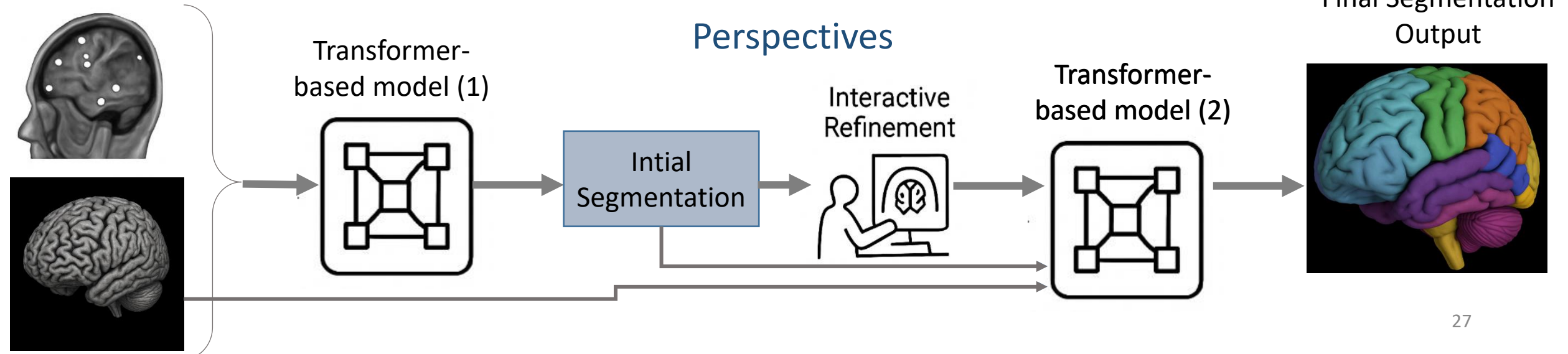
## Conclusion




Landmark Detection

Human + AI Collaboration Systems

## Perspectives



An abstract graphic featuring two large, overlapping, rounded rectangular shapes. The left shape is dark blue and contains a dense, glowing blue particle cloud. The right shape is light blue and also contains a dense, glowing blue particle cloud. The background is a light gray with a faint, white network of interconnected dots and lines, resembling a molecular or data structure. The text "Thank you for your attention!" is centered on the right side of the image.

**Thank you for your  
attention!**