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Robust extraction of tumorous vascular networks in high resolution 3D images

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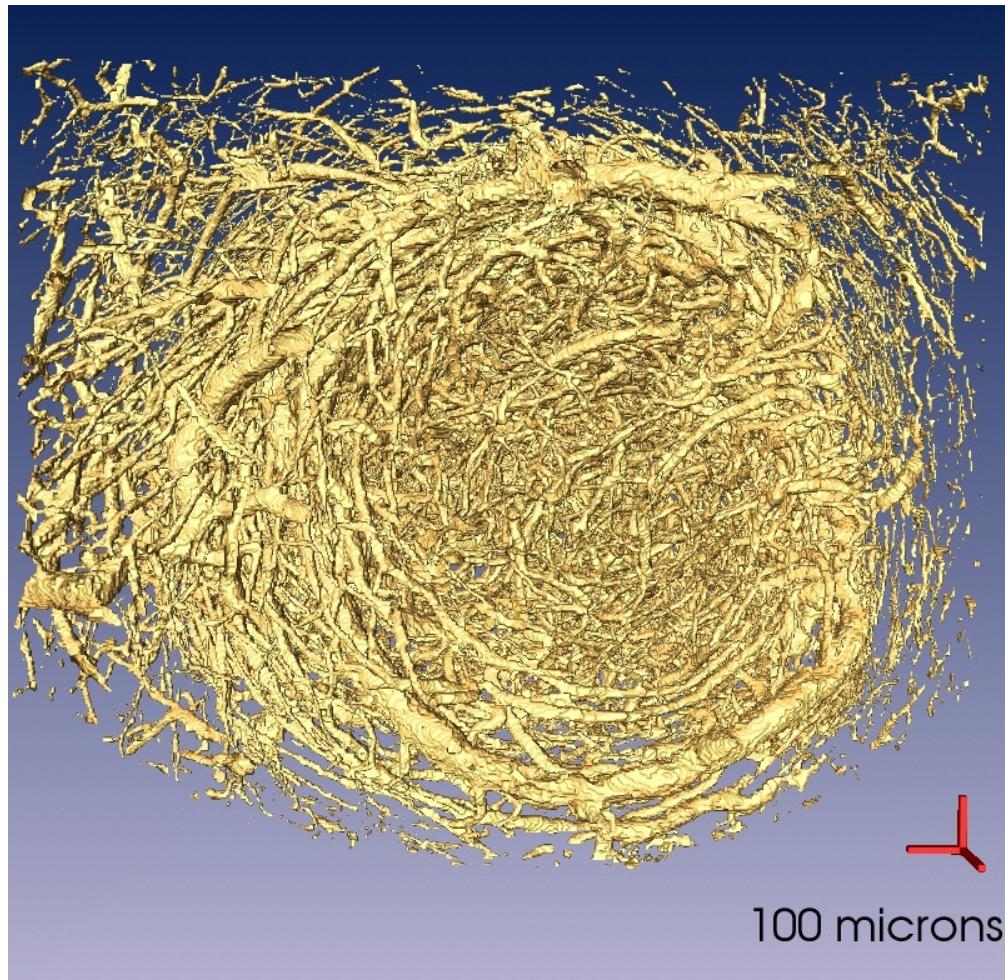
Joint work with:

- 2006 – 2008: F. Plouraboué (CNRS/IMFT), X. Descombes (INRIA Sophia Antipolis),
- 2015 - ..: R. Bates (Univ. Oxford), J.A. Schnabel (Univ. Oxford / King's college London)

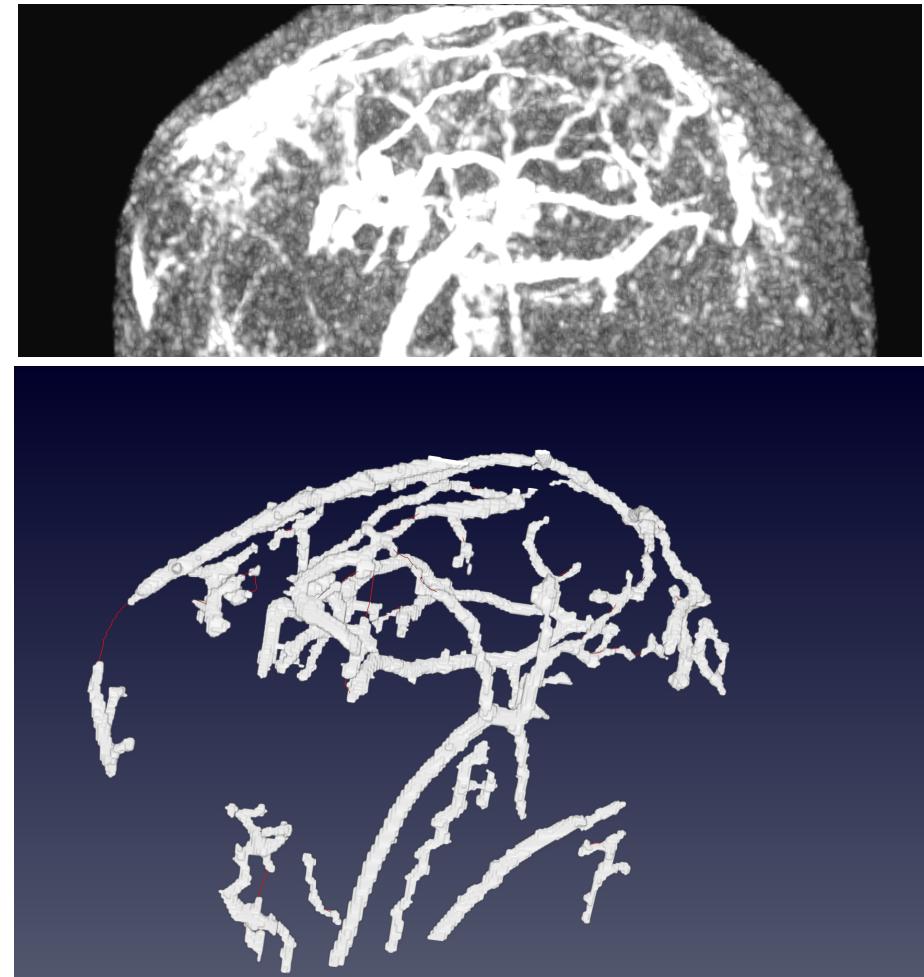
1: Context of this work

Driving motivations:

- Analysing the blood flow around tumours in pre-clinical models
- Extracting the vasculature topology around tumours out of high resolution 3D images



**Synchrotron tomography images
ESRF Grenoble → 8mm³ / 1.4μm**
[Risser et al., IEEE TMI 2007]



**Contrast enhanced micro-CT
Siemens Inveon CT → 250mm³ / 32.7μm**
[Bates et al., MICCAI 2015]

1: Context of this work

Standard pipeline:

1. Segmentation of the vessels enhanced by a contrast agent
2. Skeletonization of the extracted vessels
3. **If required: Gap filling**



Example of skeletonized network



Evidence for discontinuities (or gaps)

1: Context of this work

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Example of skeletonized network



Evidence for discontinuities (or gaps)

Existing strategies to tackle the *gap issue* in vascular networks:

1. Improving the robustness of the segmentation [Quek et al., TMI 2001] [Szymczak et al., SPIE 2005] [Pock et al., CVW 2005]
2. Learning heuristics to guide manual interactions [Kaufhold et al., MedIA 2012]
3. Simulating angiogenesis [Schneider et al. MICCAI 2014]
4. Tensor models to fill small gaps due to the noise [Cetin et al., TMI 2013]

No method adapted to fill relatively large gaps with in irregular vascular networks

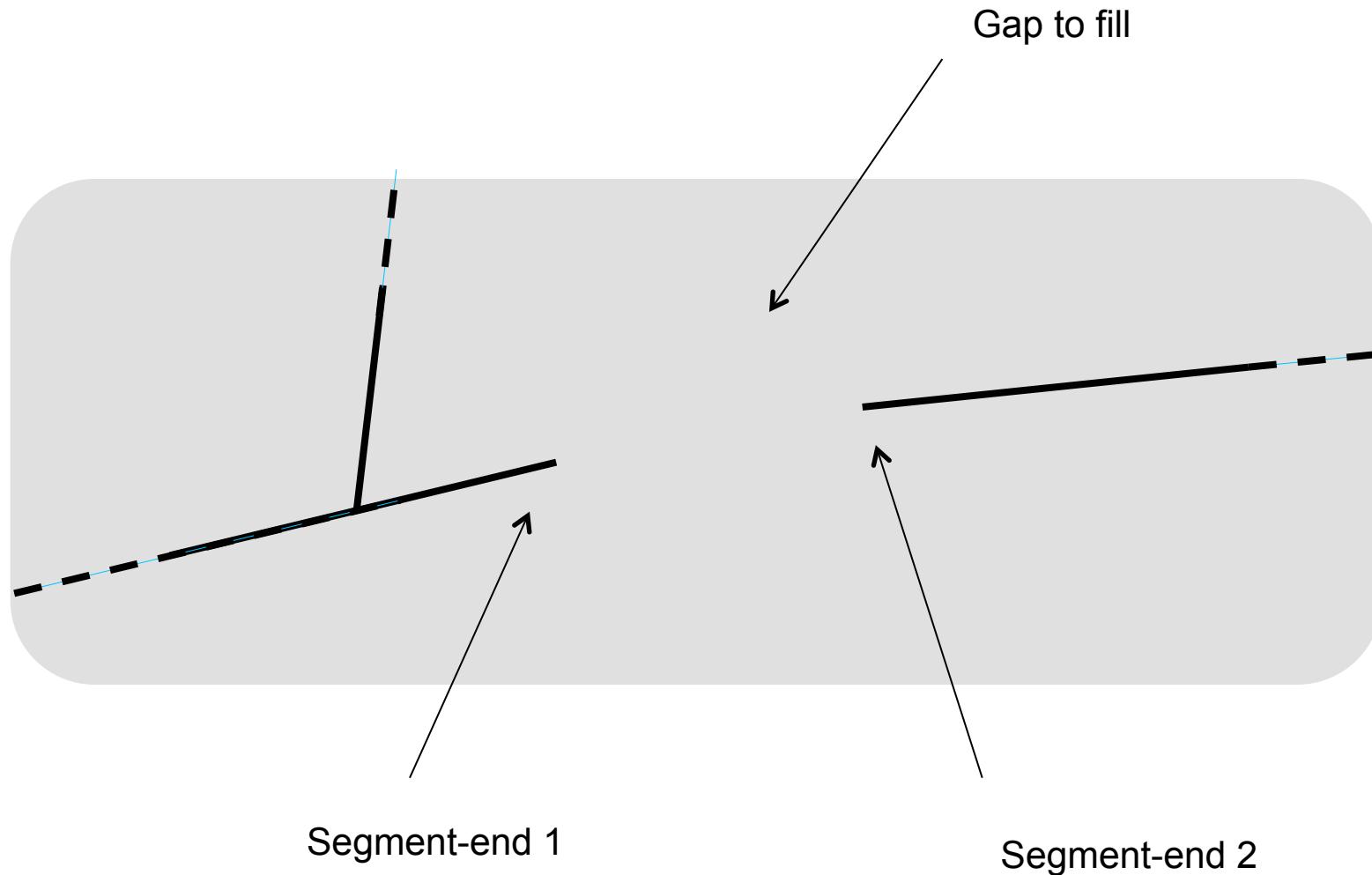
1: Context of this work

Overview of this talk:

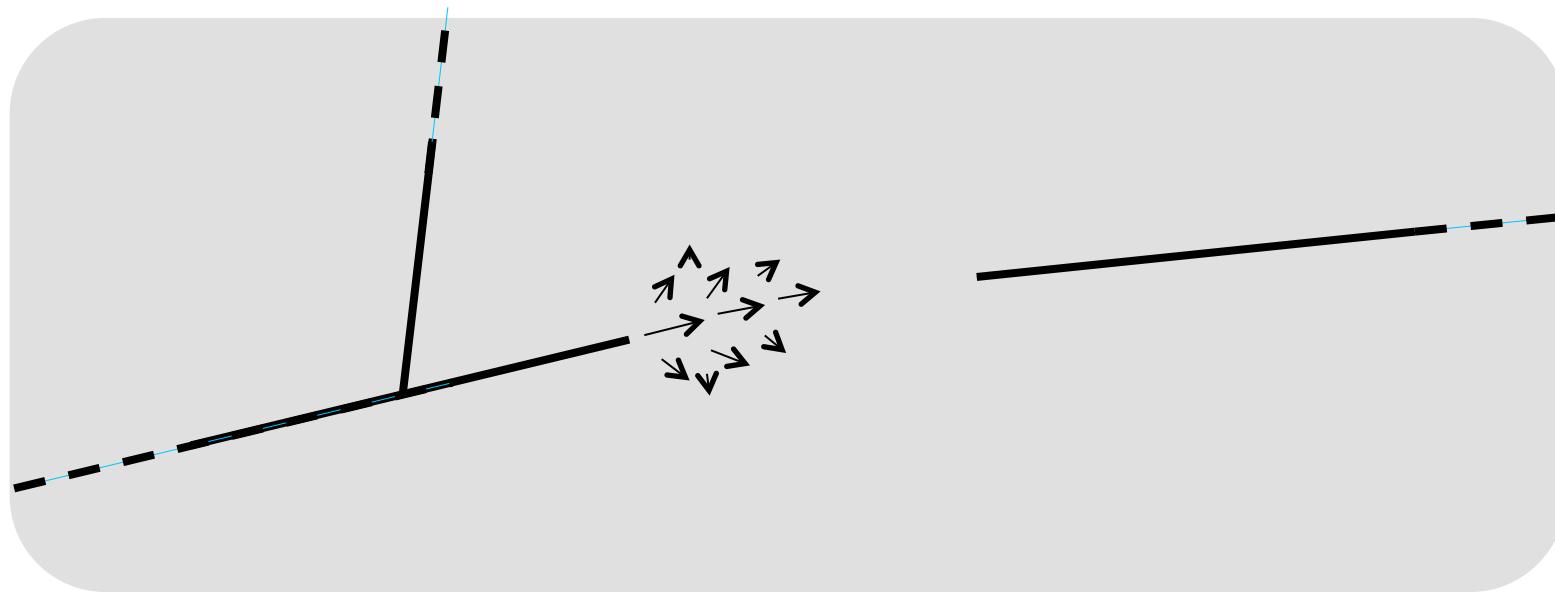
- Using tensor fields to fill discontinuities
- Gap filling using skeleton-based information
- Gap filling using skeleton- and intensity-based information

2: Using tensor fields to fill discontinuities

Idea adapted from [Guy & Medioni, IEEE PAMI 1997] in [Risser et al., IEEE TMI 2007]



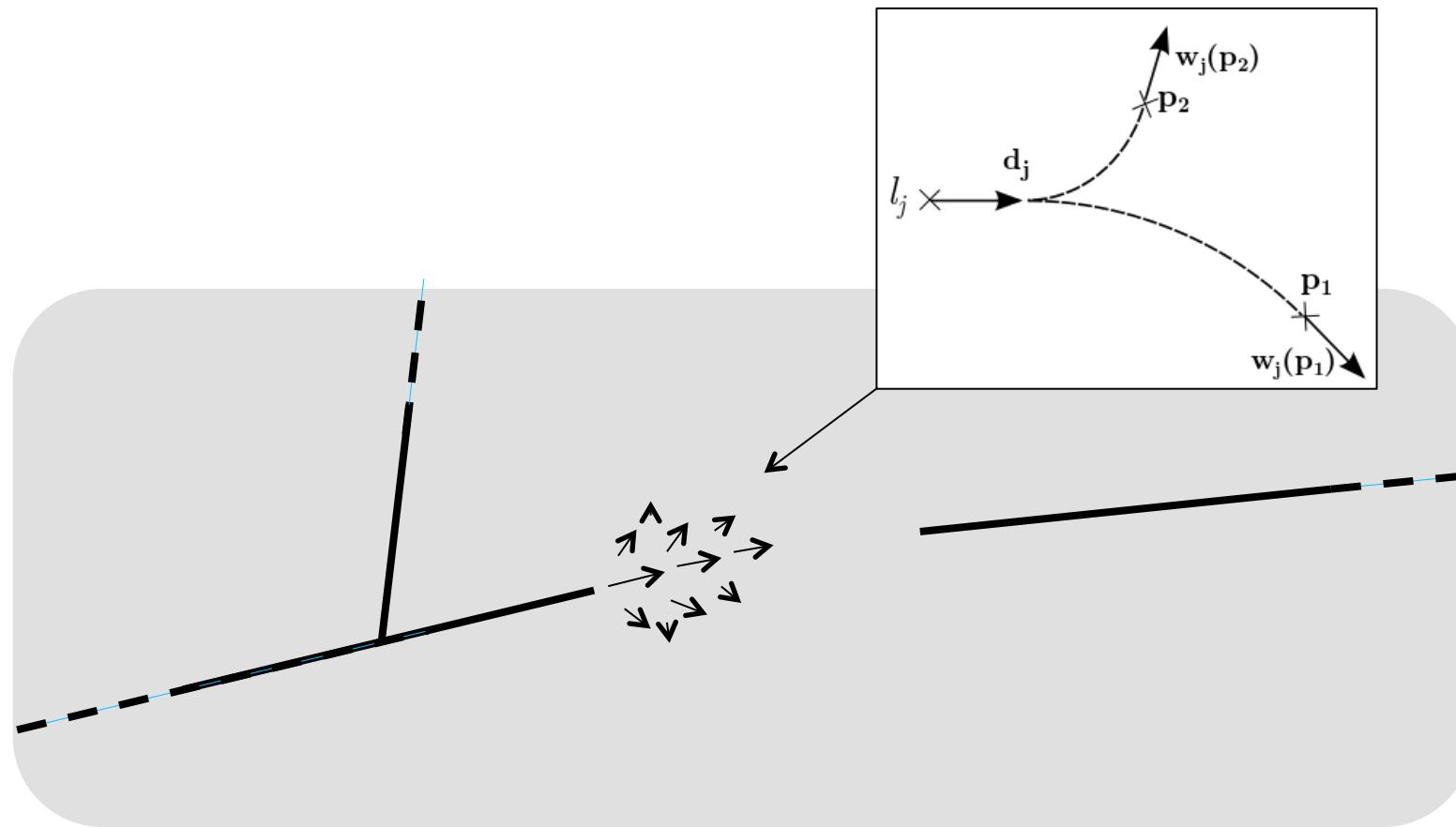
2: Using tensor fields to fill discontinuities



For segment-end 1: generate a vector field P_1

- Represents how the vessel could be extended
- Here: $P_1(i,j,k) = (x_1, y_1, z_1)$

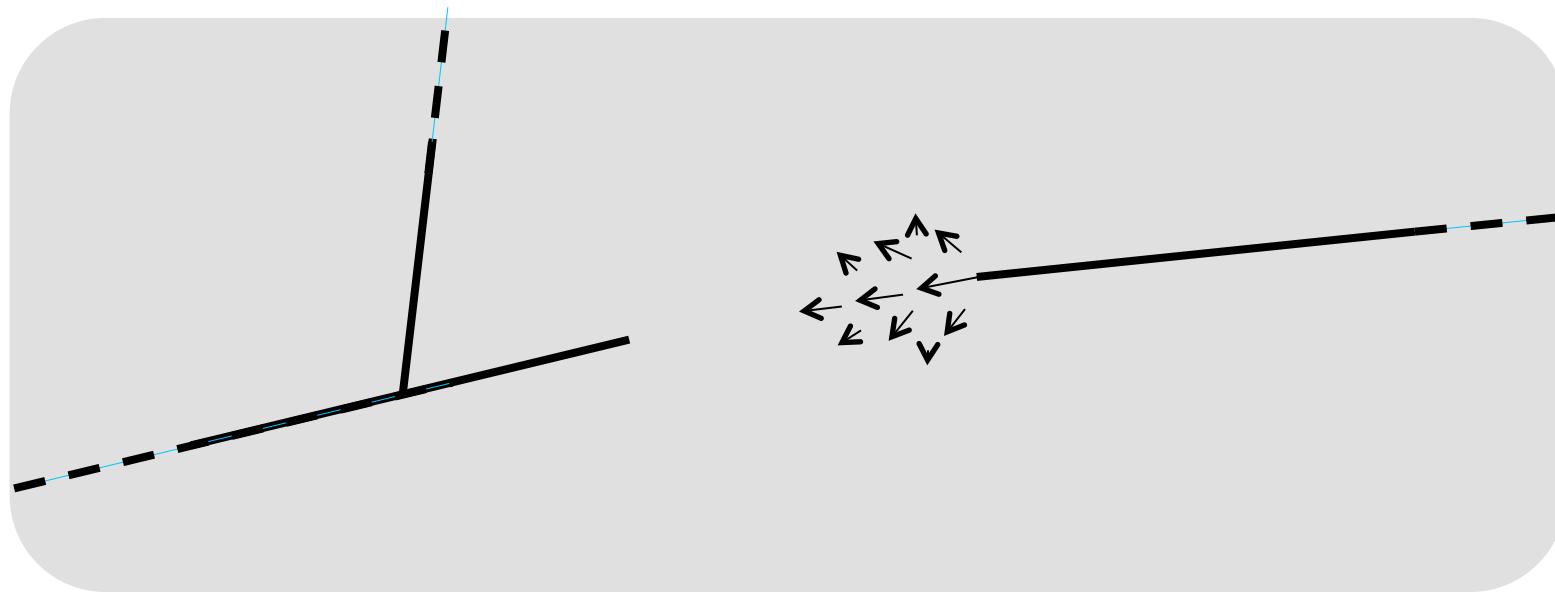
2: Using tensor fields to fill discontinuities



For segment-end 1: generate a vector field \mathbf{P}_1

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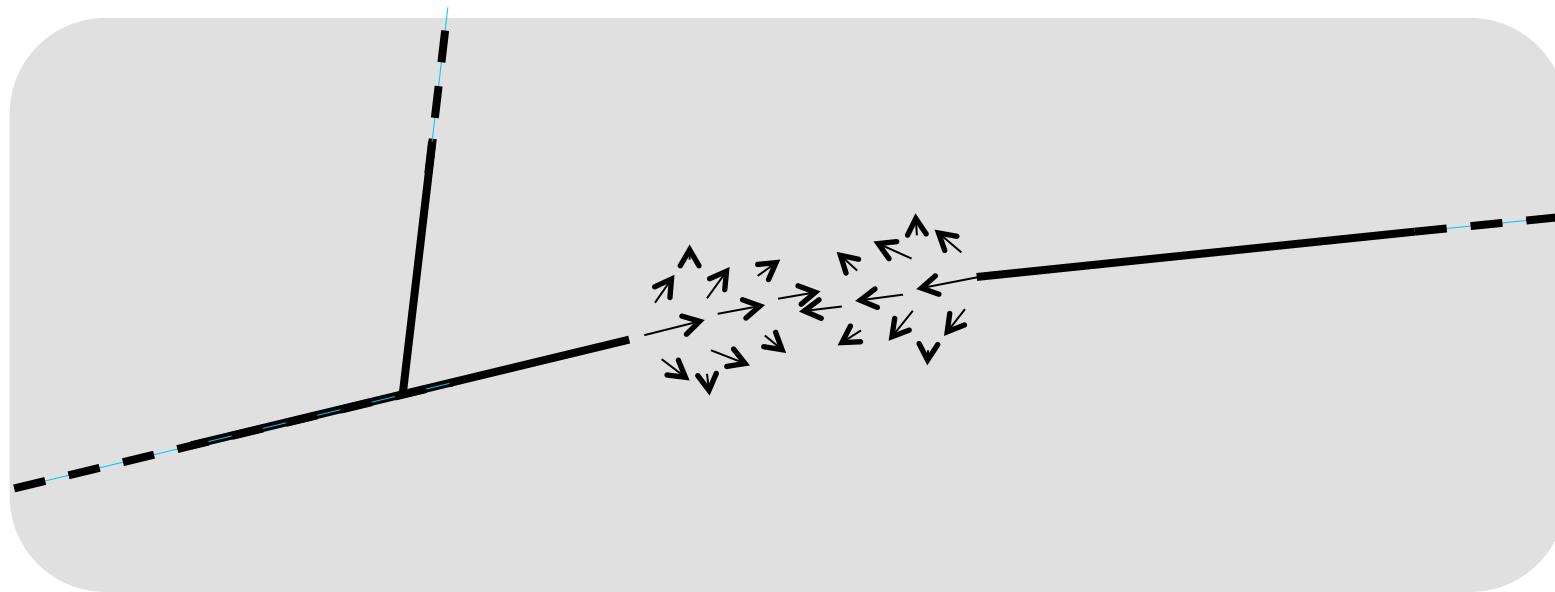
2: Using tensor fields to fill discontinuities



For segment-end 2: generate a vector field \mathbf{P}_2

- Represents how the vessel could be extended
- Here: $\mathbf{P}_2(i,j,k) = (x_2, y_2, z_2)$

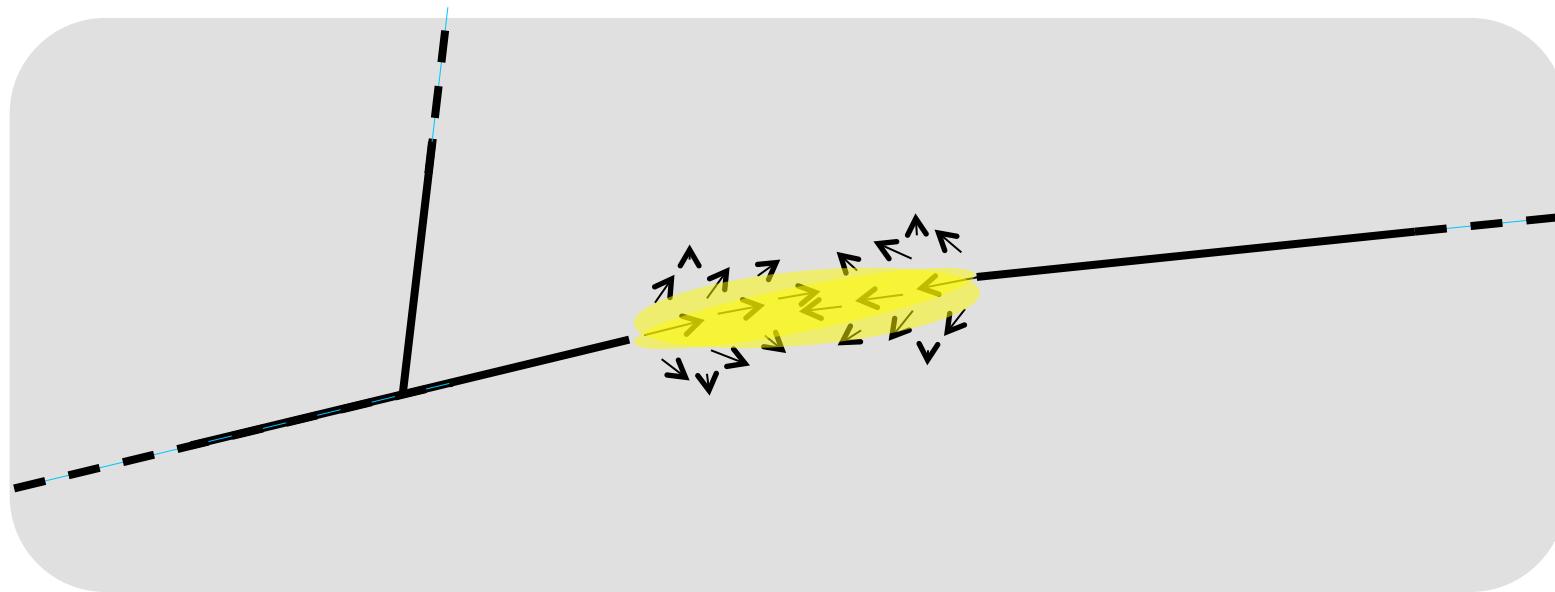
2: Using tensor fields to fill discontinuities



Communication at point (i,j,k) between segment-end 1 and segment-end 2 through a tensor field:

$$\rightarrow \quad T(i,j,k) = P_1(i,j,k) \otimes P_1(i,j,k) + P_2(i,j,k) \otimes P_2(i,j,k)$$

2: Using tensor fields to fill discontinuities



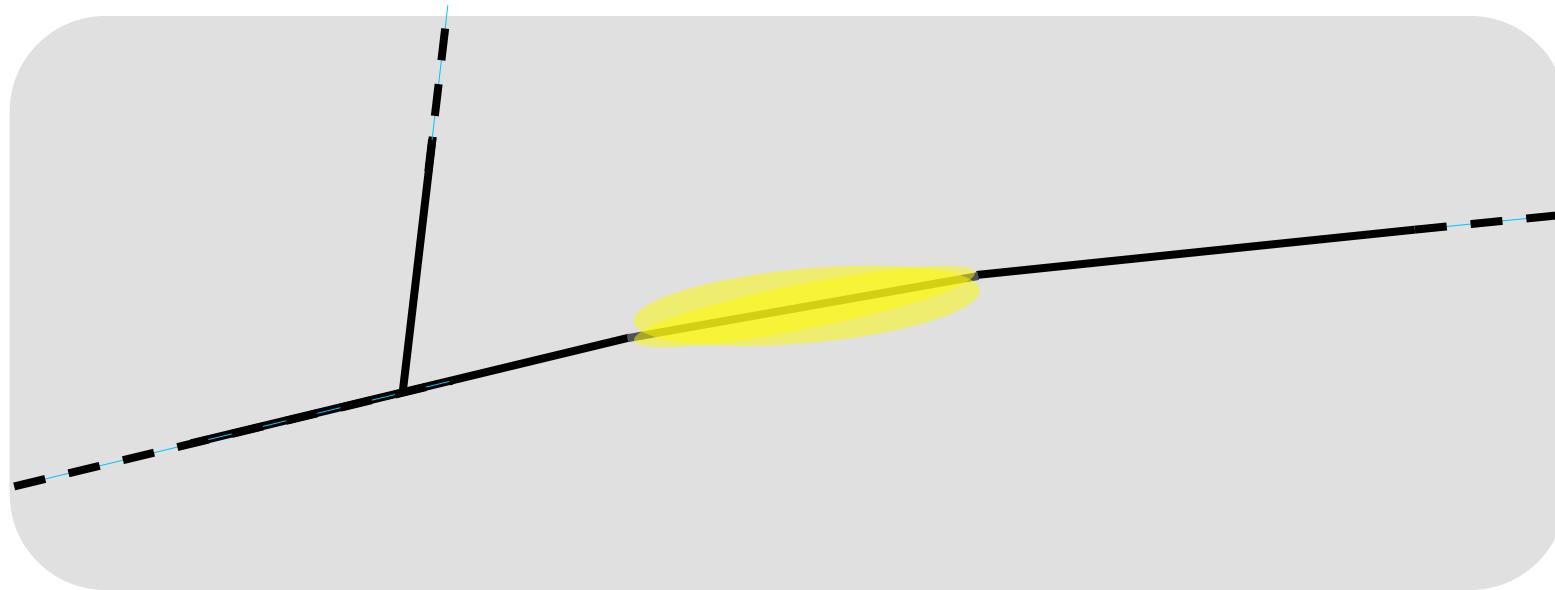
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Saliency to a curve computed as the difference between the two first eigenvalues of $T(i,j,k)$:

$$\rightarrow \quad S(i,j,k) = \lambda_1(i,j,k) - \lambda_2(i,j,k)$$

2: Using tensor fields to fill discontinuities



Path search:

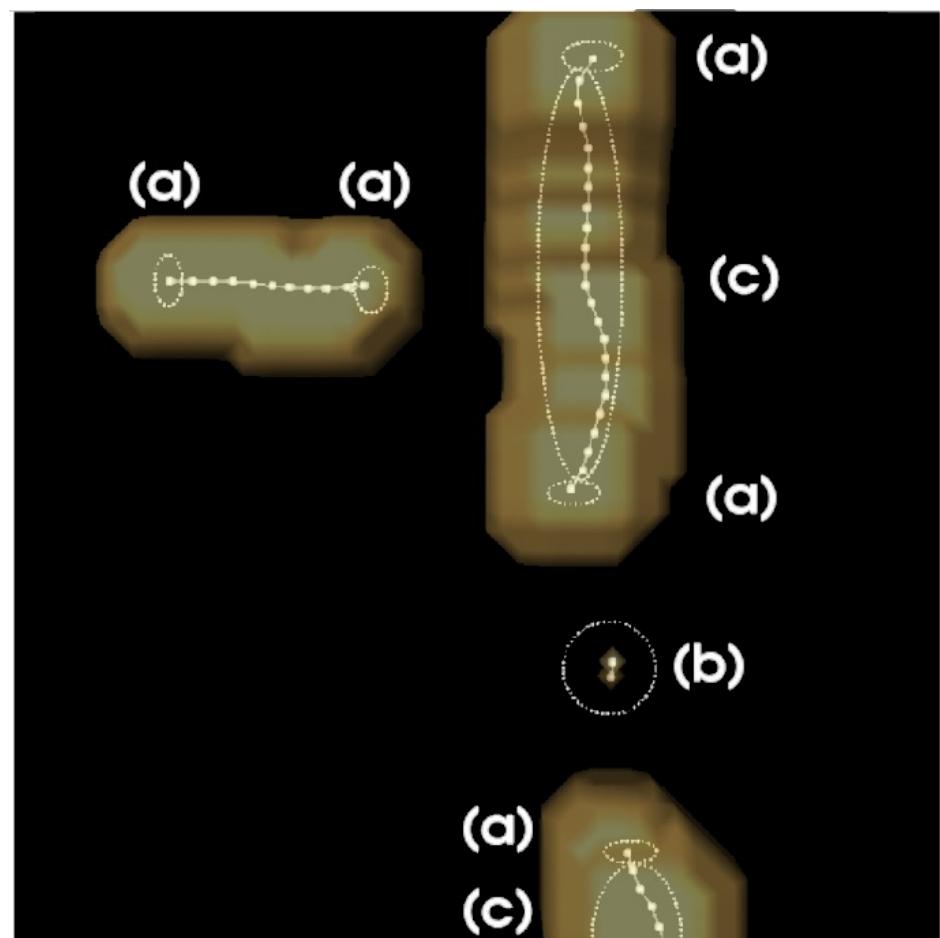
- Start from a segment end
- Follow the local maxima of the saliency to a curve

3: Gap filling using skeleton based information

Three types of tokens

[Risser et al., IEEE TMI 2007]

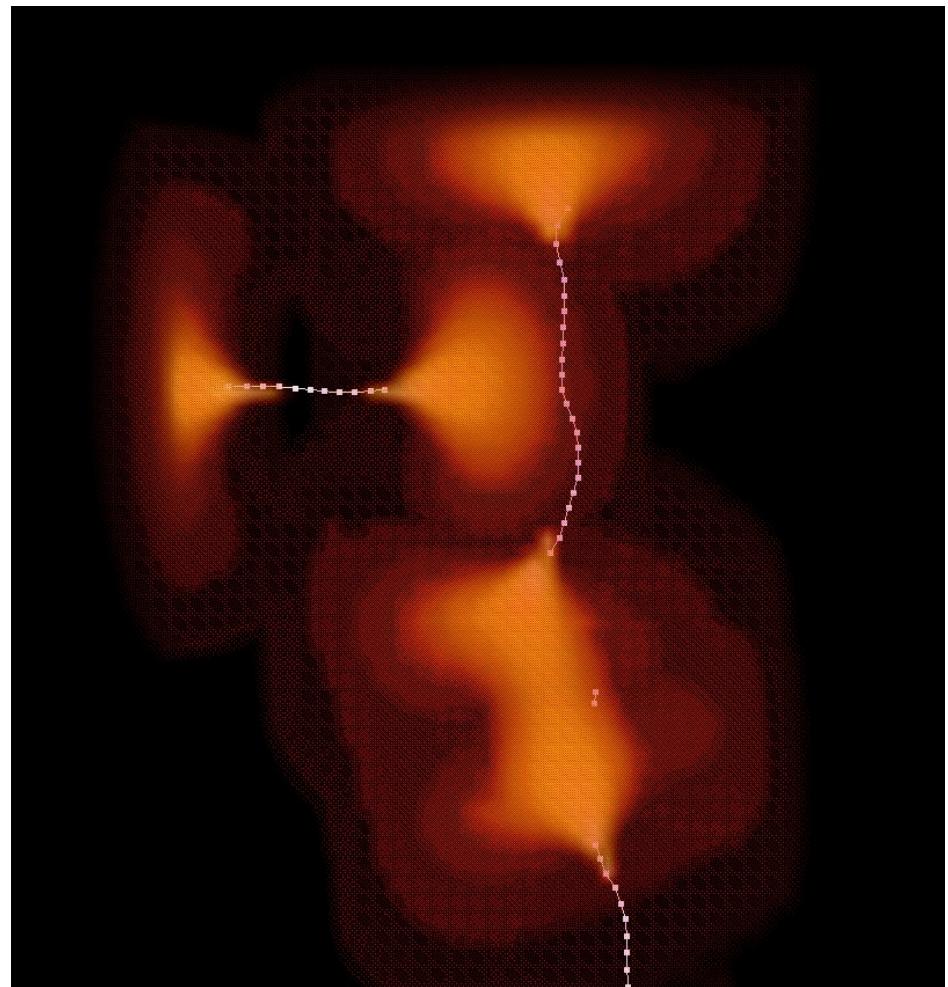
- Segment-end tokens (a)
- Island tokens (b)
- Segment tokens (c)



3: Gap filling using skeleton based information

Communication

- Segment-end tokens
- Island tokens
- Segment tokens

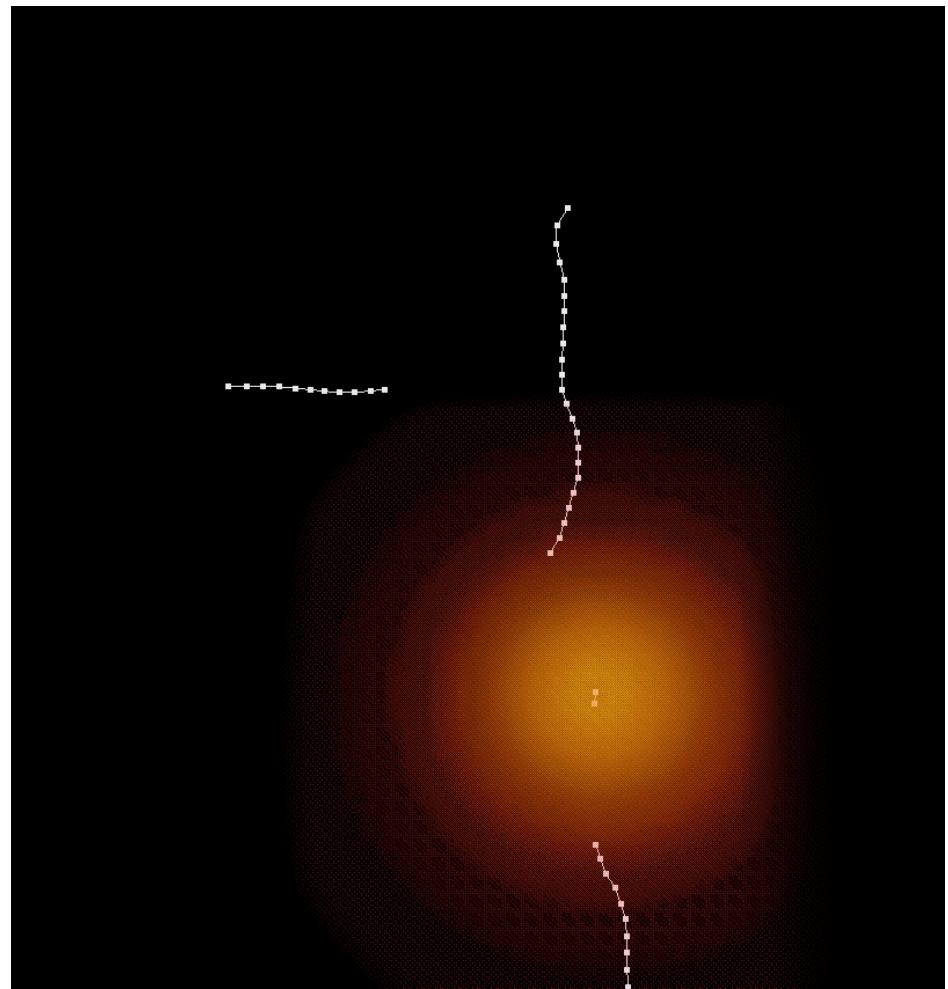


$$\mathbf{T}(i, j, k) = \sum_{n=1}^N \mathbf{TE}_n(i, j, k) + \sum_{m=1}^M \mathbf{TI}_m(i, j, k) + \sum_{q=1}^Q \mathbf{TS}_q(i, j, k)$$

3: Gap filling using skeleton based information

Communication

- Segment-end tokens
- Island tokens
- Segment tokens

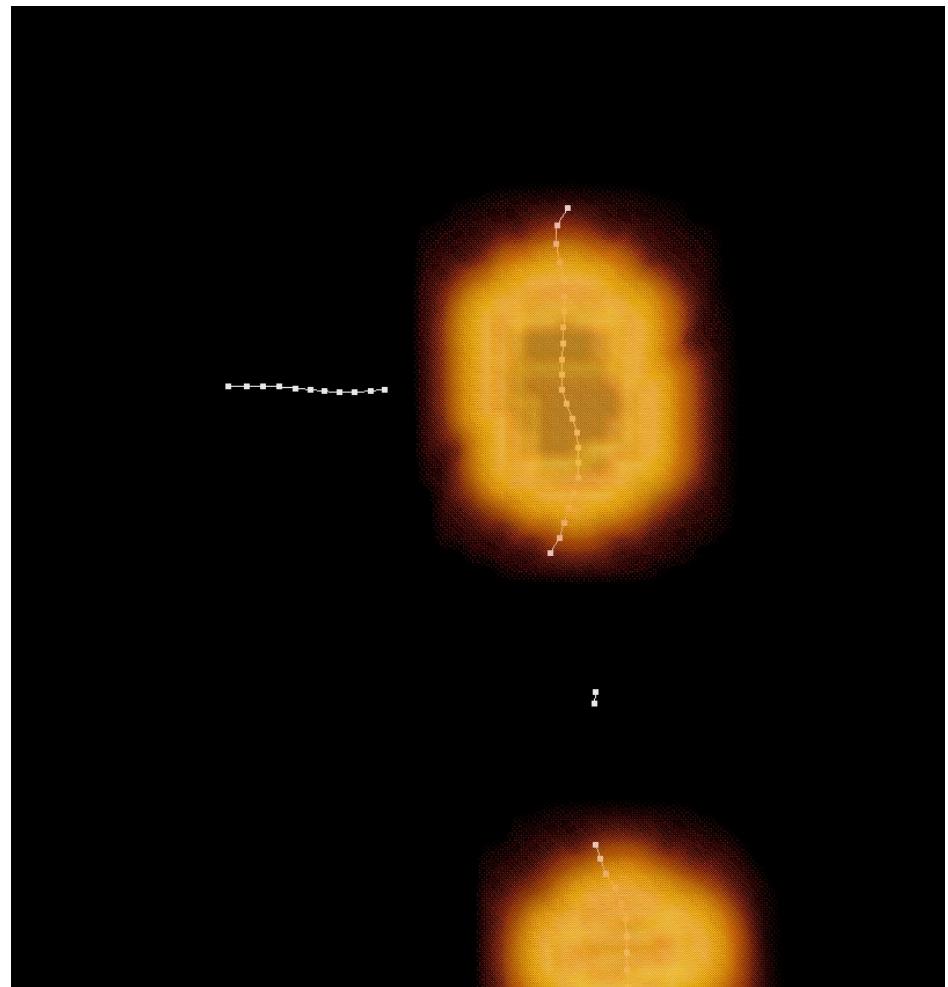


$$\mathbf{T}(i, j, k) = \sum_{n=1}^N \mathbf{TE}_n(i, j, k) + \sum_{m=1}^M \mathbf{TI}_m(i, j, k) + \sum_{q=1}^Q \mathbf{TS}_q(i, j, k)$$

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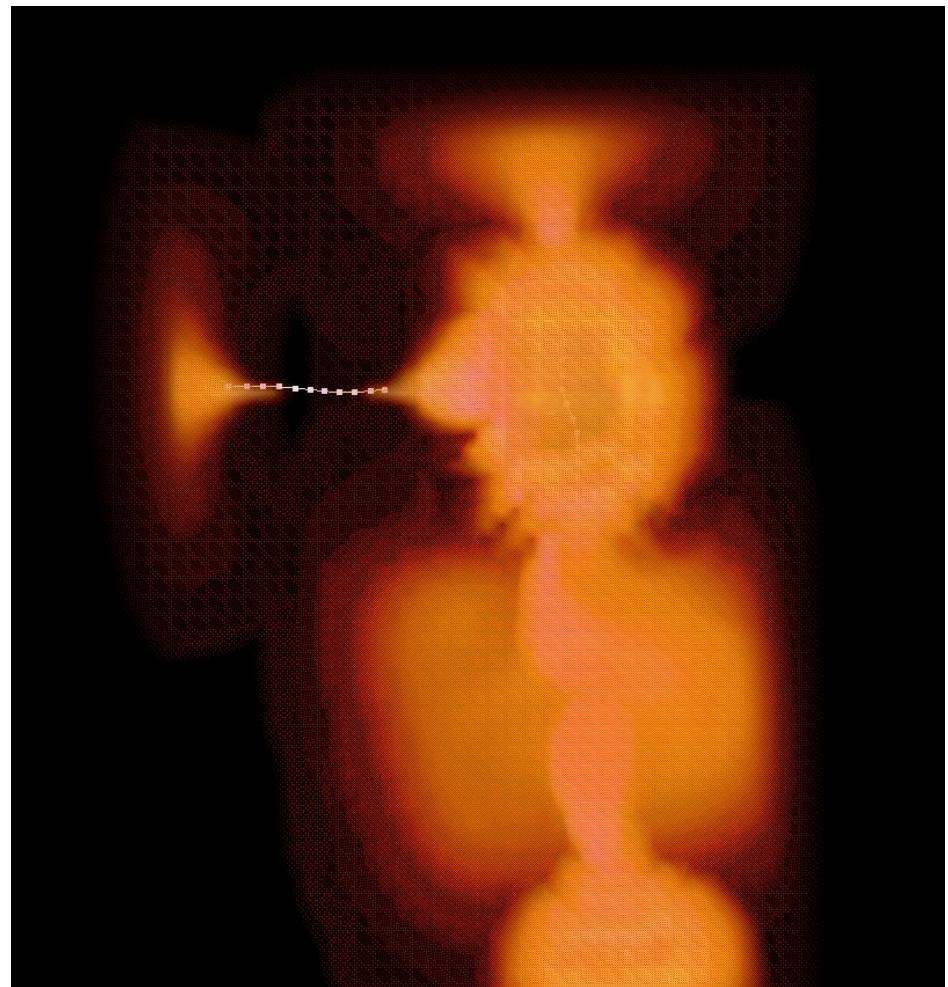


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Communication

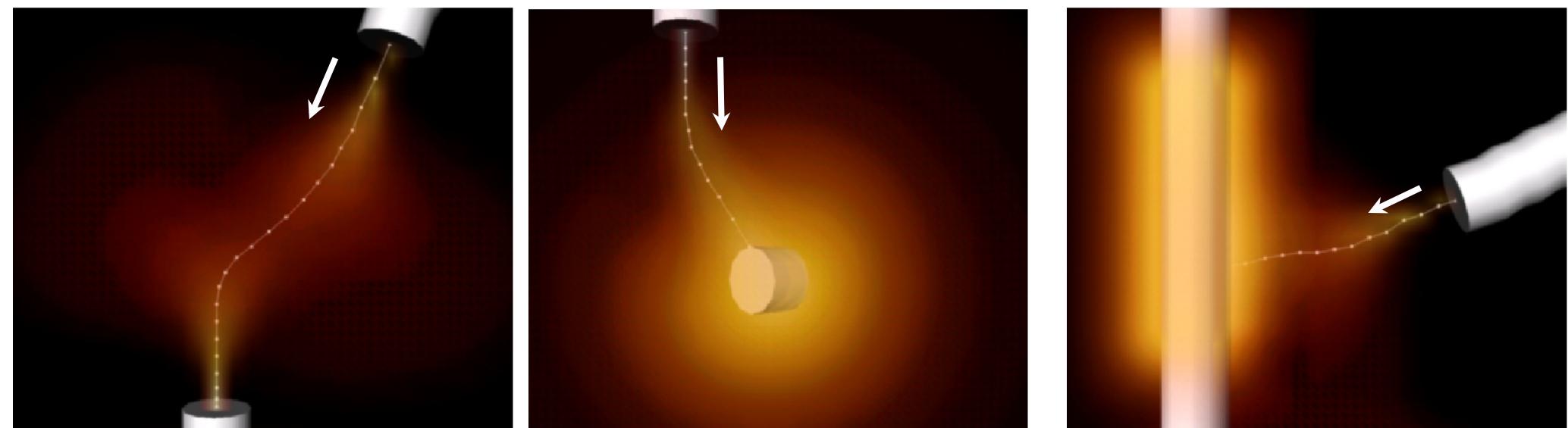
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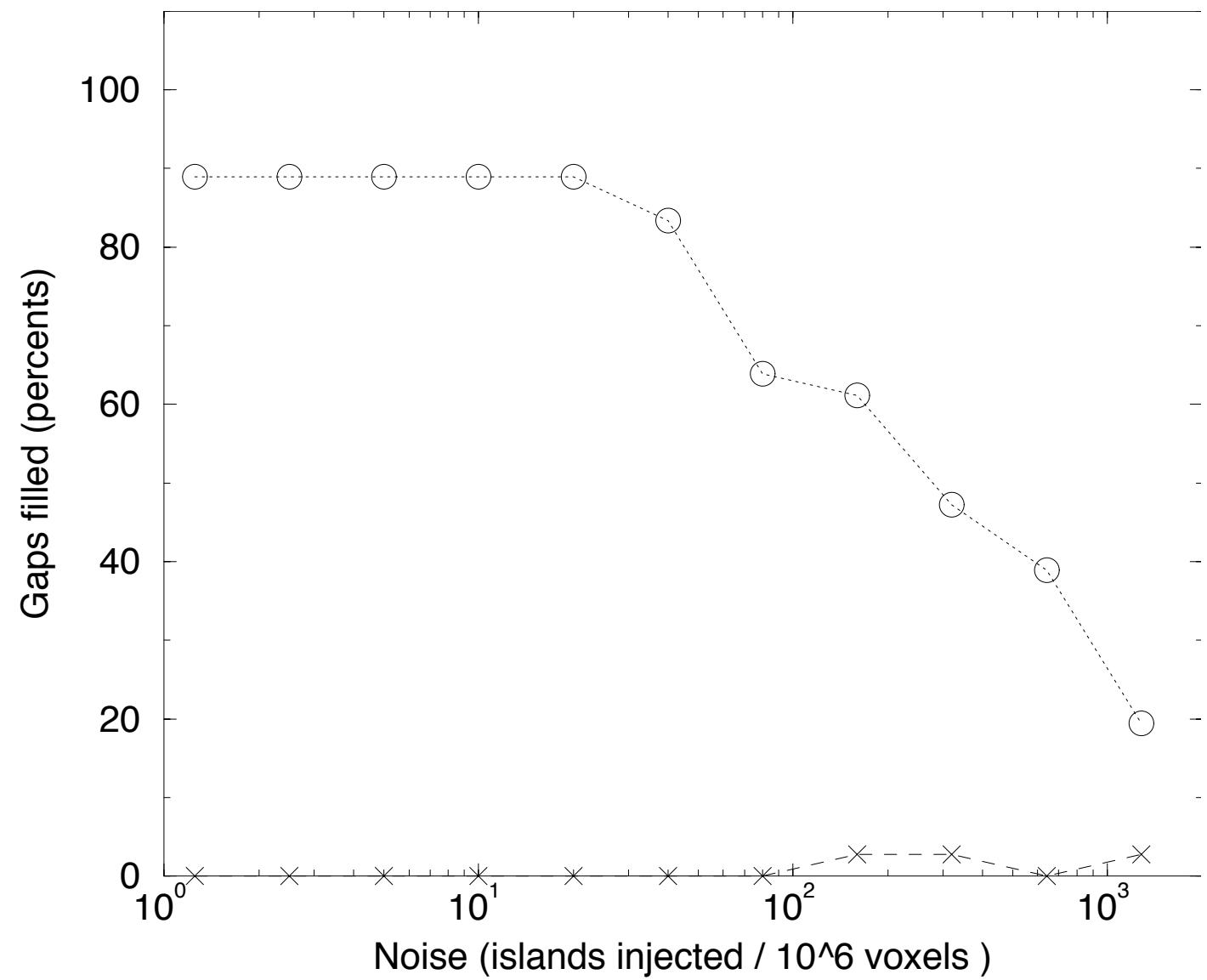
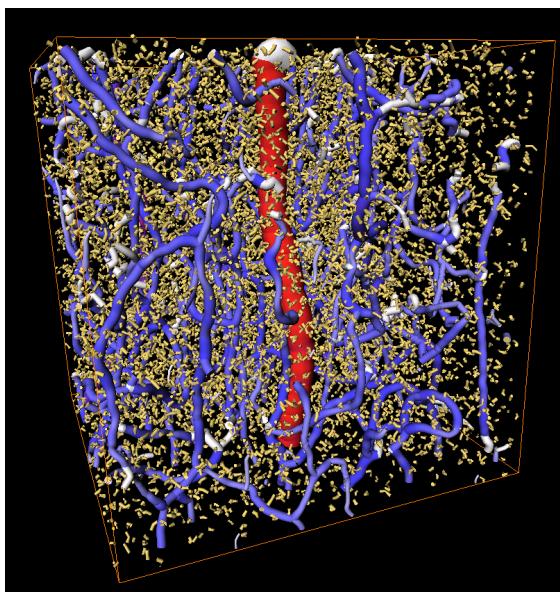
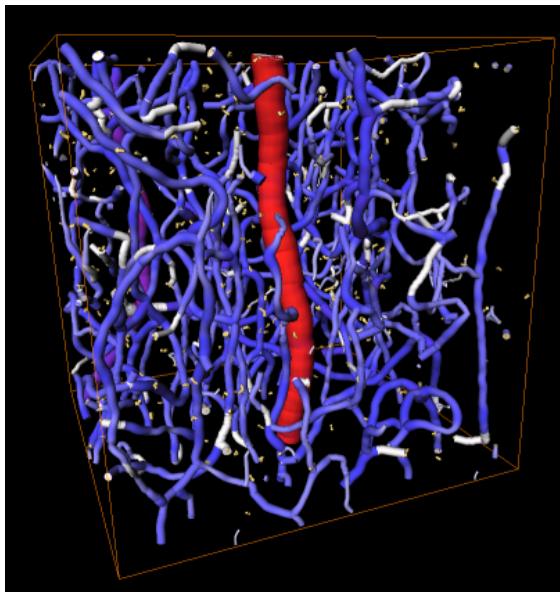
3: Gap filling using skeleton based information

Path search: from segment-ends to other tokens



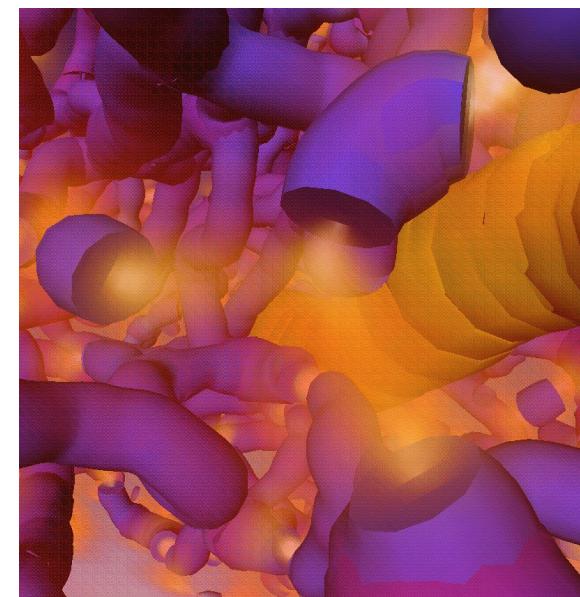
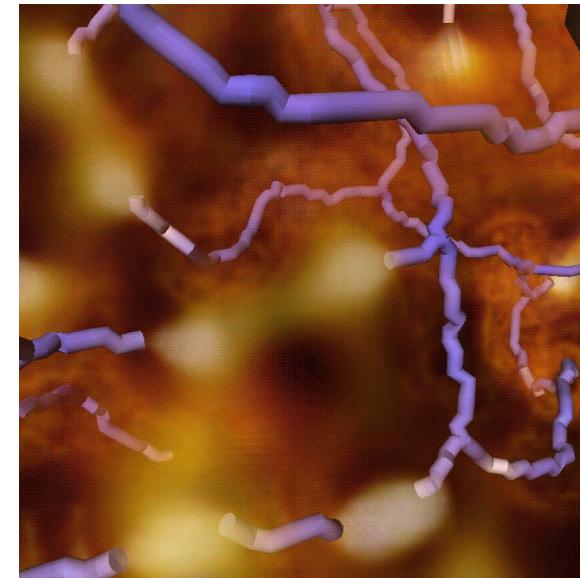
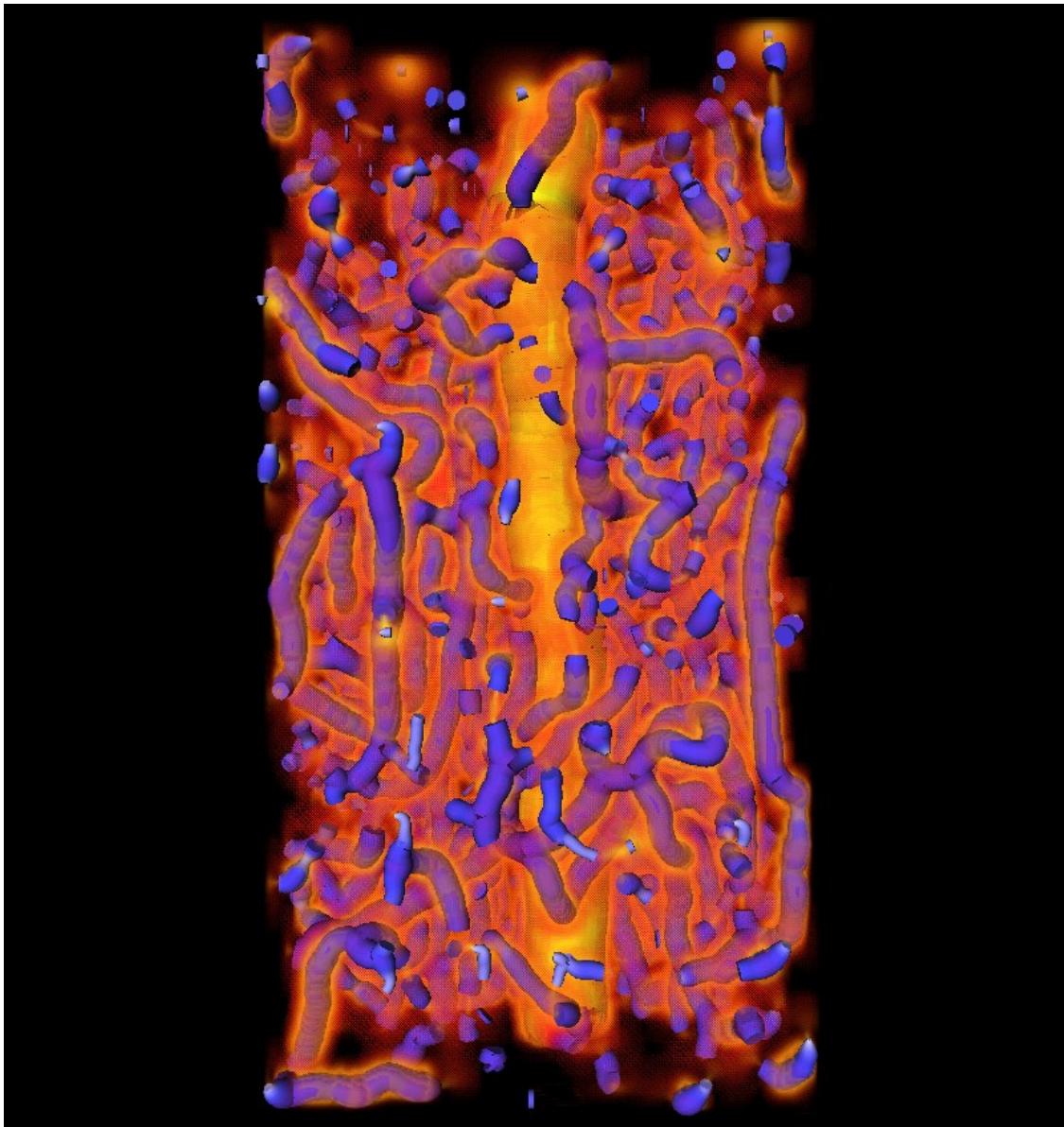
3: Gap filling using skeleton based information

Results on phantom data



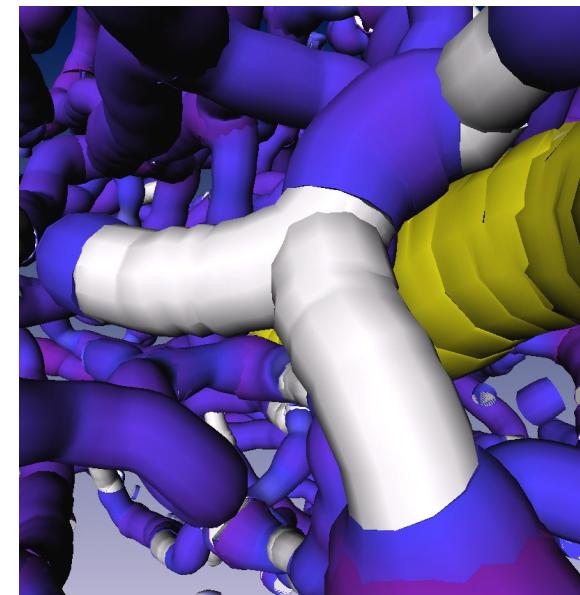
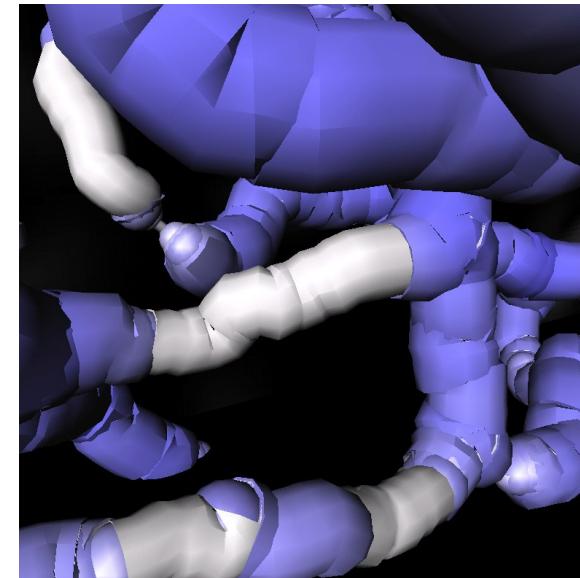
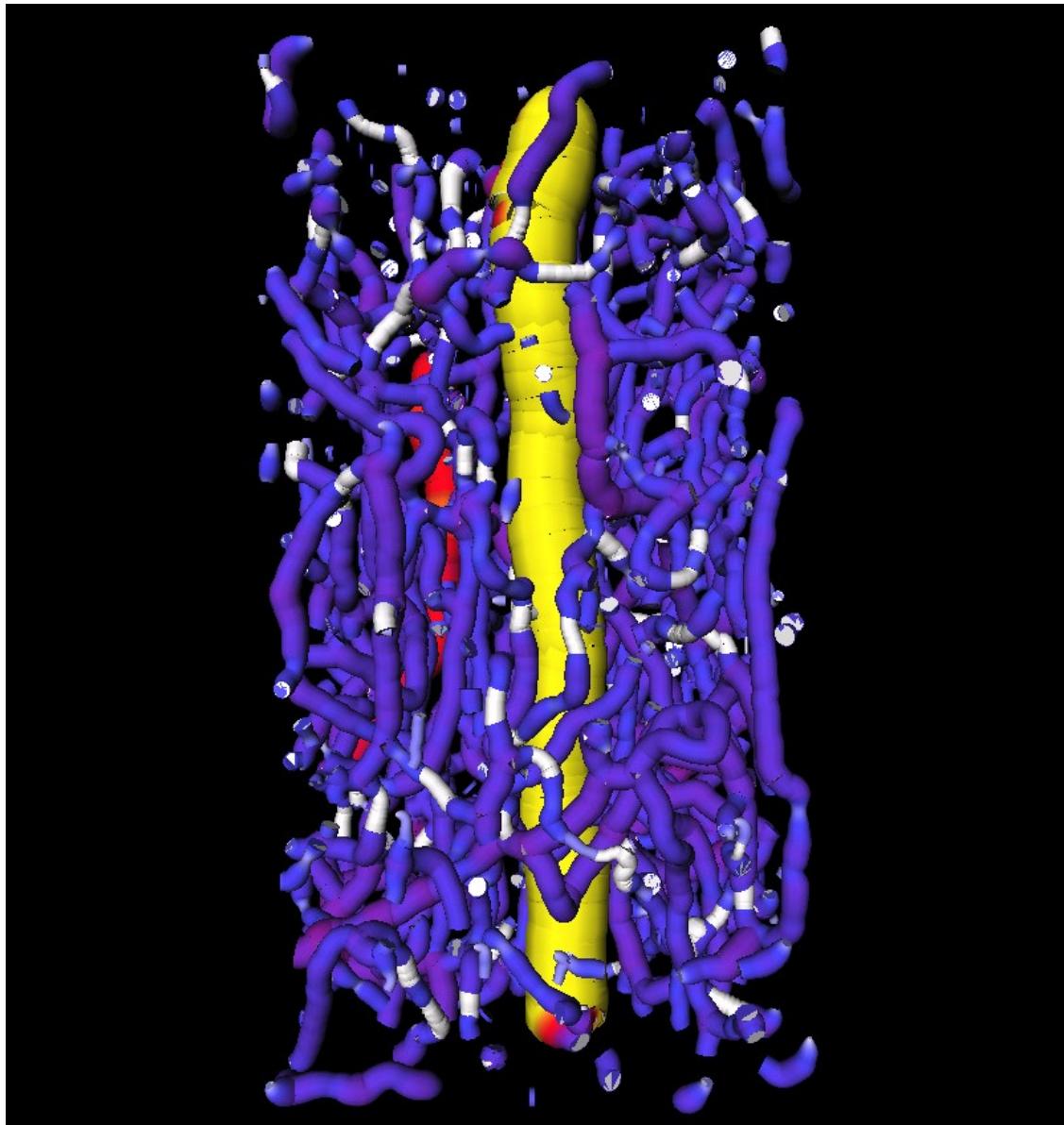
3: Gap filling using skeleton based information

Results on real data



3: Gap filling using skeleton based information

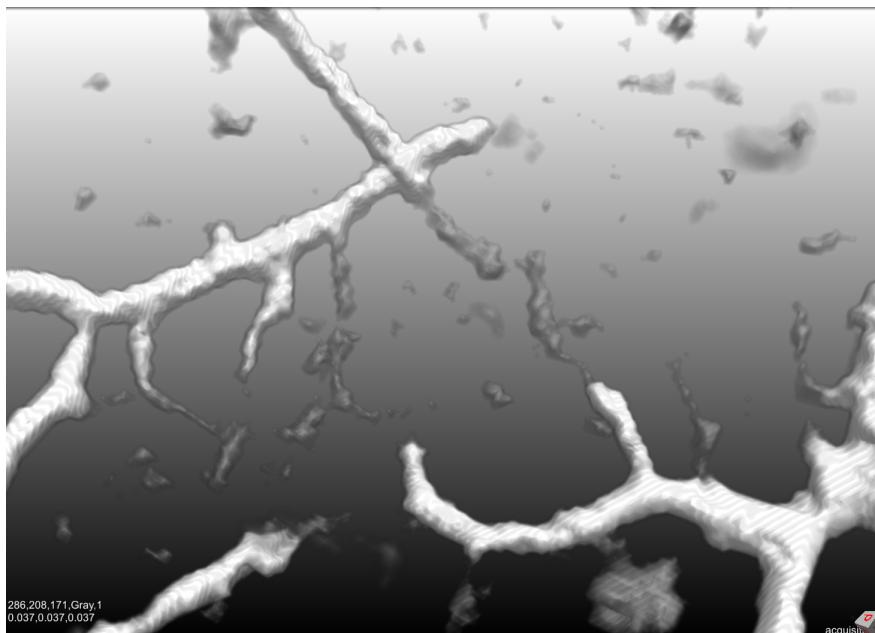
Results on real data → About 90% of gaps filled



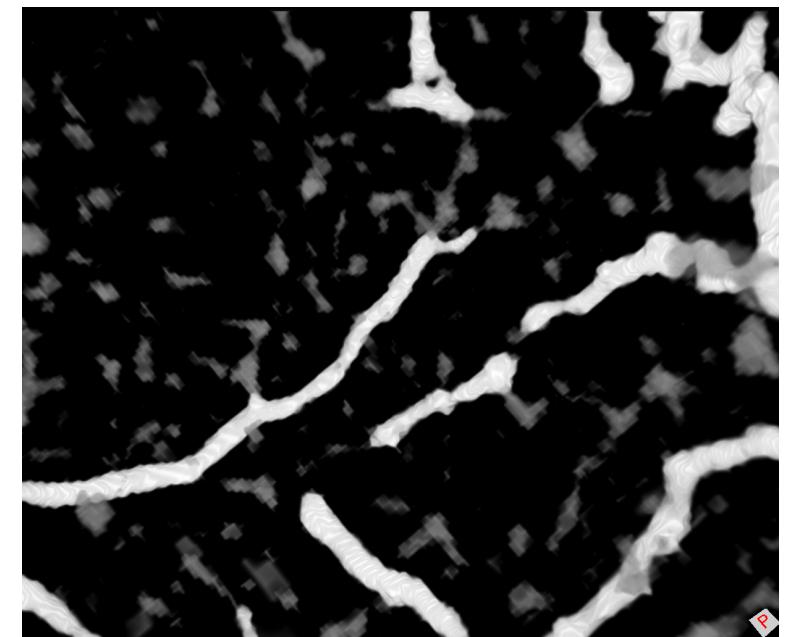
4: Gap filling using skeleton- and intensity-based information

Good results obtained using skeleton-based information, but...

... intensity-based information would be helpful for robust long distance gap filling



Example 1



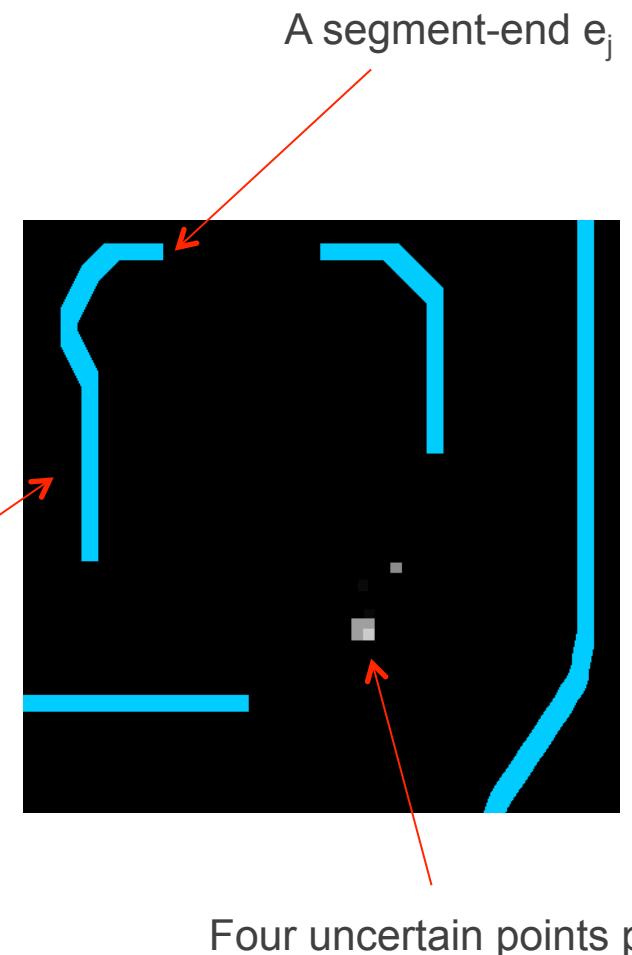
Example 2

4: Gap filling using skeleton- and intensity-based information

Proposed strategy

Consider:

- A set of segments $s_i, i \in \{1, \dots, I\}$
- A set of segment-ends $e_j, j \in \{1, \dots, J\}$
- A set of uncertain points p_k with weights $\omega_k, k \in \{1, \dots, K\}$



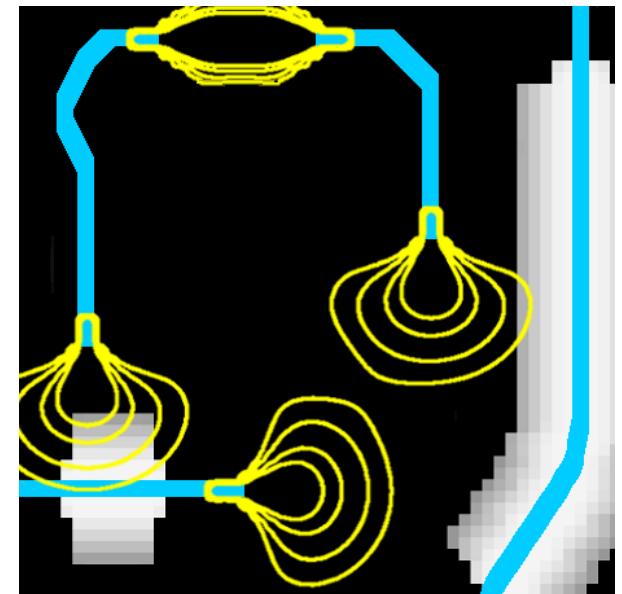
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Generate a tensor field T using the e_j and s_i



4: Gap filling using skeleton- and intensity-based information

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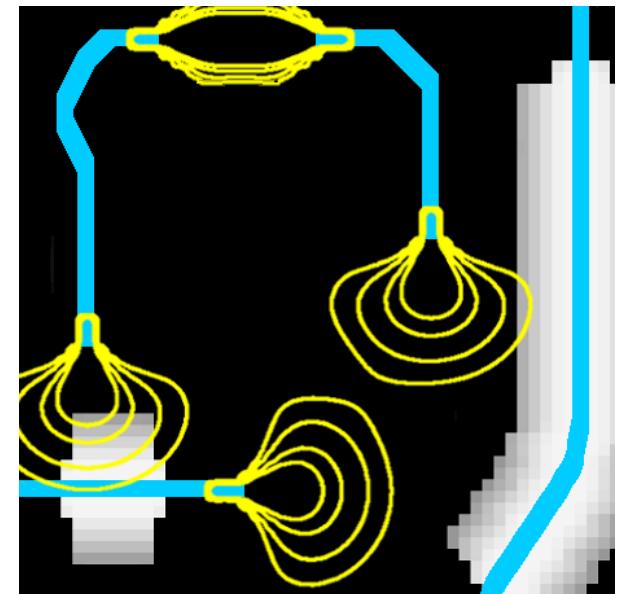
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Generate a tensor field T using the e_j and s_i

After performing a SVD at each point, derivate from T :

- Saliency map to a curve $S = \lambda_1 - \lambda_2$
- Preferential directions $D = v_1$



4: Gap filling using skeleton- and intensity-based information

Proposed strategy

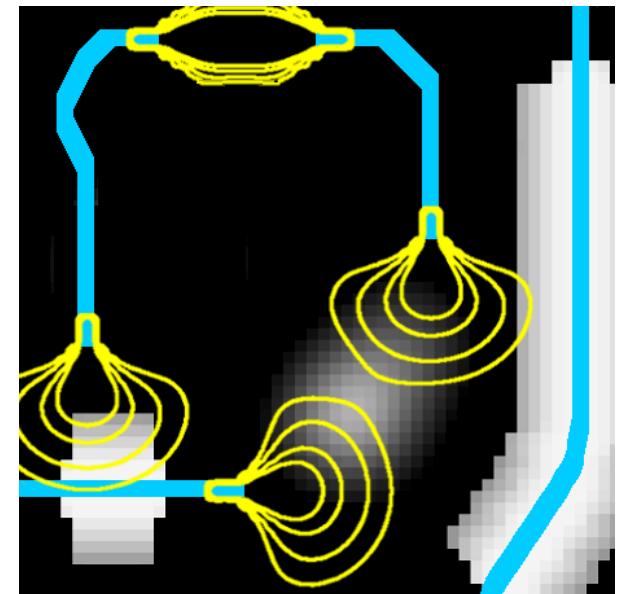
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After constructing and image P using the (p_k, ω_k) compute an enhancement map E:

$$E = \frac{K_\sigma \otimes P}{K_{2\sigma} \otimes I(P) + \varepsilon}$$

Inspired from the normalized convolution strategy of [Knutsson and Westin CVPR 1993]

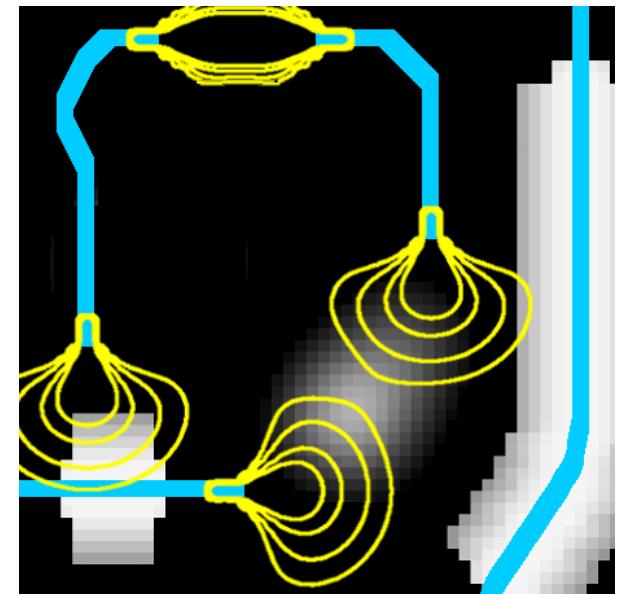
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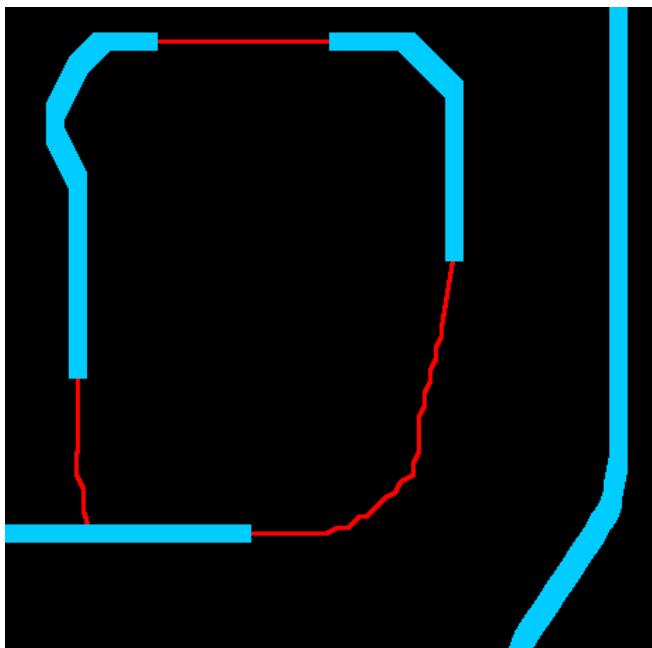
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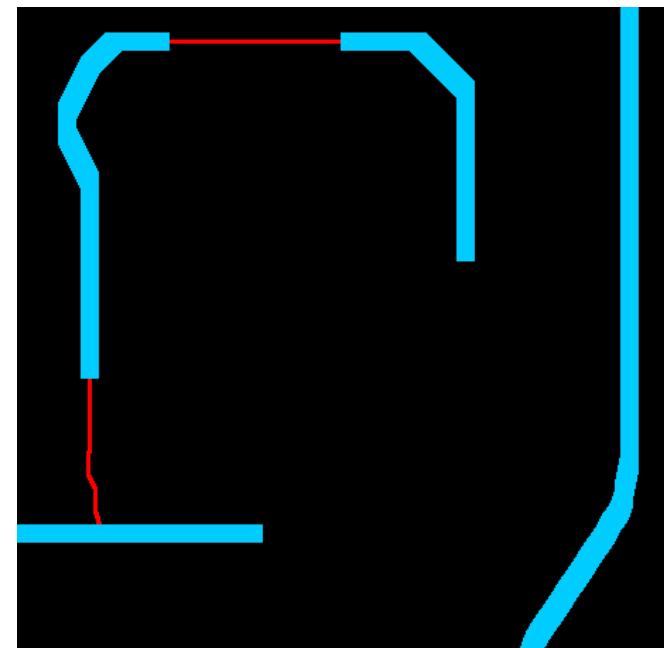
... then path search

4: Gap filling using skeleton- and intensity-based information

Results in the 2D synthetic example:



Using intensity information

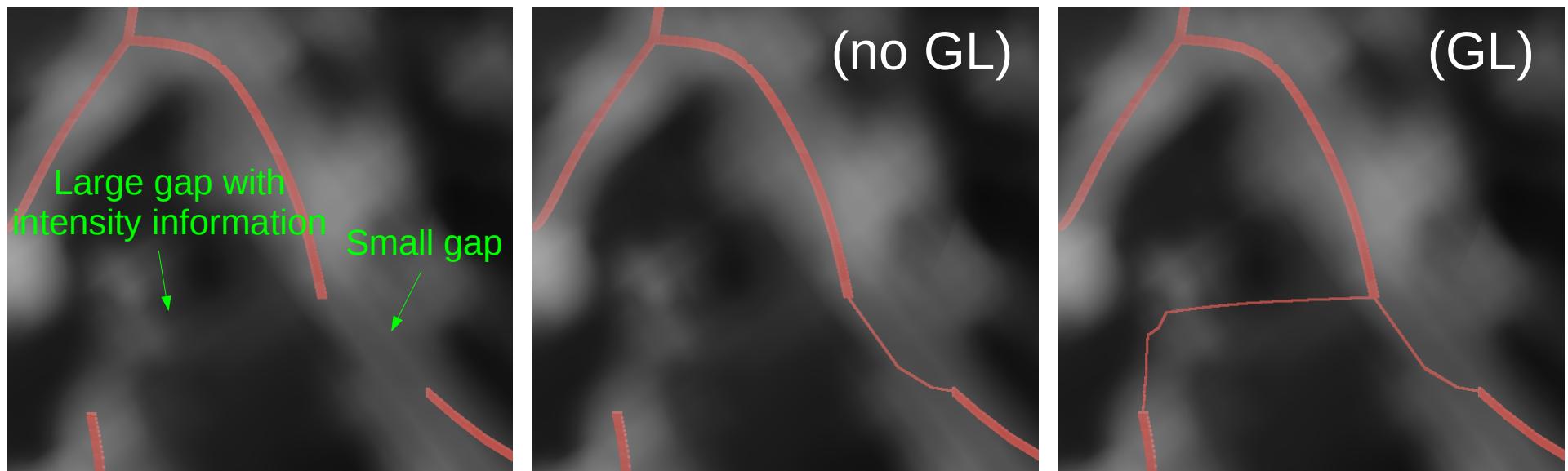


No intensity information

4: Gap filling using skeleton- and intensity-based information

Results on 3D real data:

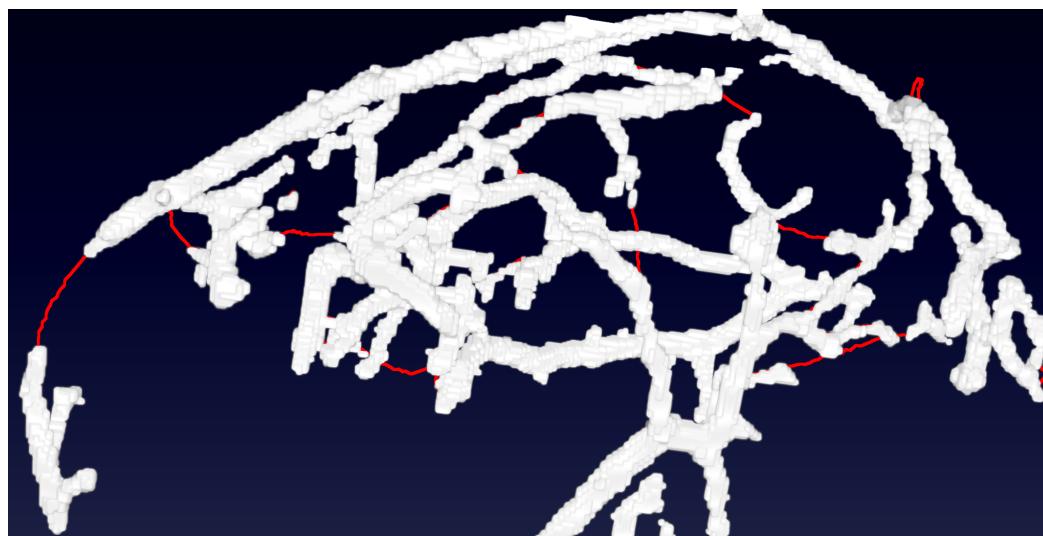
- Ten volumes of preclinical tumour model
- Female CBA mouse with murine adenocarcinoma NT (CaNT) implanted subcutaneously on the right flank.
- Contrast enhanced micro-CT scanning using an Inveon system (Siemens Healthcare)
- Isotropic voxel size of $32.7\mu\text{m}$ / $300\times 200\times 170$ voxels



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- 75 gaps filled using no intensity information
- 95 gaps filled using intensity information
- No obvious false positive

4: Gap filling using skeleton- and intensity-based information

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- No obvious false positive

Conclusion:

- Model adapted to the vasculature in tumours
- Encouraging results

Current work:

- Stochastic path search strategy

References

- [Risser et al., TMI 2007] L. Risser, F. Plouraboué, and X. Descombes. Gap Filling in Vessel Networks by Skeletonization and Tensor Voting. *IEEE Trans. Med. Imaging*, 2008.
- [Bates et al., MICCAI 2015] R. Bates, L. Risser, B. Irving, B. Papiez, P. Kannan, V. Kersemans, J.A Schnabel. Filling Large Discontinuities in 3D Vascular Networks using Skeleton- and Intensity-based information. In Proc. MICCAI 2015
- [Guy & Medioni, PAMI, 1997] G. Guy and G. Medioni. Inference of surfaces, 3-D curves, and junctions from sparse, noisy, 3-D data. *IEEE Trans. Pat. Anal. Mach. Int.*, 1997
- [Quek and Kirbas, TMI 2001] F. K. H. Quek and C. Kirbas. Vessel extraction in medical images by wave-propagation and traceback. *IEEE Trans. Med. Imaging*, 2001.
- [Szymczak et al., SPIE MI 2005] A. Szymczak, A. Tannenbaum, and K. Mischaikow. Coronary vessel cores from 3-D imagery: A topological approach. In Proc. SPIE Med. Imag., 2005.
- [Pock et al., CVW 2005] T. Pock, C. Janko, R. Beichel, and H. Bischof. Multiscale medialness for robust segmentation of 3-d tubular structures. In Proc. CVW workshop, 2005.
- [Kaufhold et al., Media 2012] P. Kaufhold, P. S. Tsai, P. Blinder, and D. Kleinfeld. Vectorization of optically sectioned brain microvasculature: Learning aids completion of vascular graphs by connecting gaps and deleting open-ended segments. *Medical Image Analysis*, 2012.
- [Schneider et al. MICCAI 2014] M. Schneider, S. Hirsch, B. Weber, G. Szekely, and B.H. Menze. TGIF: Topological Gap In-Fill for Vascular Networks. In Proc. MICCAI, 2014.
- [Cetin et al., TMI 2013] S. Cetin, A. Demir, A. J. Yezzi, M. Degertekin, and G. B. Unal. Vessel Tractography using an Intensity Based Tensor Model With Branch Detection. *IEEE Trans. Med. Imaging*, 2013.