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Learning a reconnecting regularization term for blood vessel variational segmentation

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Abstract—The segmentation of blood vessels in medical images is a challenging task as they are thin, connected and tortuous. The detection of a connected vascular network is of the utmost importance in clinical applications (e.g. blood flow simulations, vascular network modeling and analysis). Deep learning approaches have been developed to tackle this issue, but they require a large annotated dataset for each new application of interest, which is very challenging to build for vascular networks. In this work, rather than learning the segmentation task, we propose to learn a reconnecting regularization term that learns geometric properties of vascular networks independent of the image modality. Therefore, this term generalizes better than deep learning segmentation models, and can be easily plugged into variational segmentation frameworks to detect vascular networks in different datasets without requiring annotations. We apply this approach on retinal images by training our reconnecting term on the STARE dataset and applying it on the DRIVE dataset. We show that our approach better preserves the connectivity of vascular networks than classic regularization terms in the literature. Finally, we illustrate the generalization power of our reconnecting term by applying it to other types of data.

Index Terms—blood vessel segmentation, reconnection, deep learning, variational approach

I. INTRODUCTION

The detection of blood vessels in medical images is the key task of most vascular network analysis tools that help clinicians diagnose and treat vascular diseases. Despite being studied for the past twenty years [1], the segmentation of blood vessels is still an open problem as vascular networks are thin, connected, tortuous, sparse and low-contrasted structures.

In this work, we address the important problem of blood vessel connectivity loss upon segmentation of vascular networks. Preserving connectivity is key, as many subsequent processing tasks rely on a connected network such as blood flow simulation or vascular network modeling. Previous work have focused on this issue. Tracking strategies (see [1] Section 7) intrinsically preserve the connectivity of blood vessels by recruiting neighbors pixels recursively, but usually require to be initialized with manual information. Other methods estimate the local direction of blood vessels in order to guide the segmentation in the vessel direction [2, 3].

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Recently, deep learning strategies have been proposed to segment vascular networks. Some of them aim at enforcing the network connectivity by using topological loss [4] or by computing both the segmentation and centerline of the vessels with the same network [5]. These strategies yield interesting results, but require a large annotated dataset to provide good results. By contrast, variational segmentation strategies do not require annotated data, but are less flexible to represent complex information such as connectivity. To combine the strengths of both methods, decoupled approaches [6] were proposed. They consist in plugging an external operator to act as a regularization term in a variational framework. Early works used denoising algorithms such as BM3D or Non Local Means [7] to replace classic denoising regularization terms such as the Total Variation (TV). More recent strategies learn the external regularization term [8, 9] to take advantage of the power representation of deep learning.

In this work, we propose to learn a regularization term designed to preserve the connectivity of vascular networks, and plug it into a variational iterative segmentation framework which does not require annotated data. We show that this regularization term better preserves the connectivity of blood vessels than classic regularization terms such as TV, and more complex ones including directional information.

II. MATERIALS AND METHODS

A. Learning a reconnecting term

We propose to learn a reconnecting network by training a model to reconnect blood vessels from images with artificial disconnections. This network takes as input a binary image of disconnected blood vessels and outputs an image with reconnected vessels. We chose to work on binary images for two reasons. First, creating realistic disconnections is easier on binary images. Second, our goal is to learn a reconnecting term independent of the dataset to apply it on new data for which ground-truths are not available. Learning on binary images is then a better strategy, as the model cannot learn any intensity-based feature of the training dataset.

We first created an algorithm to simulate realistic disconnections. It creates random disconnections in blood vessels with a thickness-dependent probability (the thinner the vessel, the higher the disconnection probability). The disconnections have a random shape to emulate disconnection due to image

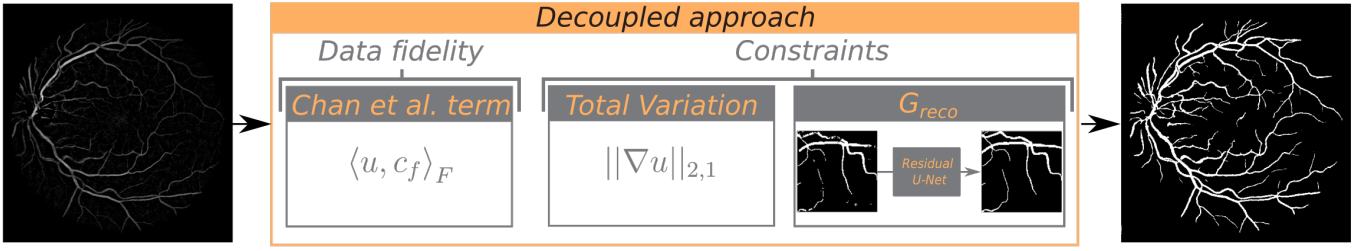


Fig. 1. Pipeline of our proposed method. A reconnecting network is first trained on a separate dataset based on a residual U-Net architecture. Then, a variational segmentation approach is used with the Chan [10] data fidelity term and either the total variation regularization term only or both the total variation and our proposed reconnecting term.

noise. Finally, some artefacts have been added to make the model distinguish parts of vessels that should be reconnected and noise that should not be connected to the vascular network. An example of disconnected image generated by this algorithm is presented in Figure 1.

To learn the reconnecting term, the residual U-Net [11] from Monai¹, a Pytorch-based framework has been used. U-Net shows great performance in medical applications and the residual units increase its robustness to image variations. 20 segmentation ground-truths from the STARE [12] dataset were used to create pairs of connected/disconnected vascular images to learn our reconnecting regularization term. We performed data augmentation by rotating each image 4 times and applying on each one different disconnections with our algorithm. This resulted in a dataset of 80 images. 2 images were kept for validation and the rest was used for training. We chose to input patches of size 96×96 instead of full images to the model to avoid learning the global architecture of the vascular network instead of the reconnecting feature. Additional random rotations and flips were applied on each patch and at each epoch to increase variability and prevent overfitting. A maximum of 1000 epochs, a batch size of 64, an Adam optimizer with a learning rate of about 10^{-3} and a Dice loss were chosen.

B. Variational approach

Segmentation may be formulated as a variational problem consisting in the optimization of a two-term energy:

$$\hat{u} = \operatorname{argmin}_{u \in [0,1]^N} E_{\text{data}}(u, f) + \lambda E_{\text{reg}}(u), \quad (1)$$

where $f \in \mathbb{R}^N$ is the initial image, E_{data} is the data fidelity, E_{reg} the regularisation term and $\lambda \in \mathbb{R}$ a regularization coefficient acting as a trade-off between both terms. The form of E_{data} depends on the choice of application. For segmentation the Chan *et al.* term [10] is classically used:

$$\begin{aligned} E_{\text{data}}(u, f) &= \langle u, c_f \rangle_F, \\ c_f &\mapsto (c_1 - f)^2 - (c_2 - f)^2, \end{aligned} \quad (2)$$

where c_1 and c_2 are constant values corresponding to the mean background and foreground intensity of the initial image, $\langle \cdot, \cdot \rangle_F$ is the Frobenius product.

¹<https://monai.io/start.html>

E_{reg} constrains the problem with regularization and/or prior knowledge. A well-known term is the Total Variation (TV), which reduces the noise in the image, and is defined as follows:

$$TV(u) = \lambda \|\nabla u\|_{2,1}, \quad (3)$$

with $\|\cdot\|_{2,1}$ the l_1 norm of l_2 norm. To solve Eq (1) with a forward-backward primal-dual algorithm (FBPD) [13], we can reformulate the model as follows:

$$\hat{u} = \operatorname{argmin}_u h(u, f) + g(Lu) + k(u), \quad (4)$$

$h(u, f) = \langle u, c_f \rangle_F$, $g(\cdot) = \lambda \|\cdot\|_{2,1}$, $L = \nabla$ and $k(u) = \iota_{u \in [0,1]^N}$ is the indicator function of the set $[0, 1]^N$. The FBPD algorithm is then :

$$\begin{aligned} u_{i+1} &= \operatorname{prox}_{\tau k}(u_i - \tau(\nabla h(x_i) + L^T v_i)) \\ v_{i+1} &= \operatorname{prox}_{\sigma g^*}(v_i + \sigma L(2u_{i+1} - u_i)), \end{aligned} \quad (5)$$

where prox_g is the proximity operator of g , g^* is the conjugate of g , $\tau \in \mathbb{R}^+$ and $\sigma \in \mathbb{R}^+$ are step-size parameters.

C. Decoupled approach

Our reconnecting term takes binary input images and returns a more connected vascular network. We thus propose to plug this reconnecting term after a few iterations of the algorithm presented in Equation 5, when the result u is close to a binary image. We thus replace, in Eq 5 $\operatorname{prox}_{\sigma k}(\cdot)$ by $\Phi(\cdot)$ defined as follows:

$$\Phi(u) = \begin{cases} \operatorname{prox}_{\sigma k}(u) & \text{if } i < \alpha \\ G_{\text{reco}}(u) & \text{otherwise} \end{cases} \quad (6)$$

where G_{reco} is the learnt reconnecting term, $\alpha \in \mathbb{N}^{+*}$ is the iteration number from which the reconnecting term is applied. Algorithm 1 present our proposed algorithm.

III. EXPERIMENTS

We applied our approach on the test set of the DRIVE dataset [14], another retinal image dataset. We compared the results with the classic Chan *et al.* model [10], and the more recent directional total variation model [2]. We optimized the parameters of both compared methods on the DRIVE training set and selected the parameters yielding the best mean results. We set the parameters of our approach experimentally to $\alpha = 350$ and $\lambda = 0.005$.

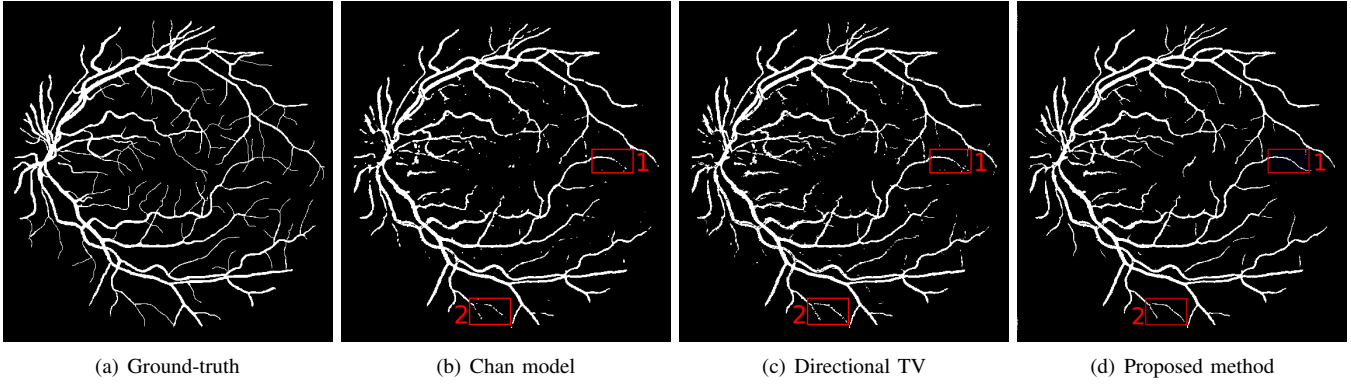


Fig. 2. Comparison of segmentation results on one DRIVE image. Zooms in the red boxes are presented in Figure 3

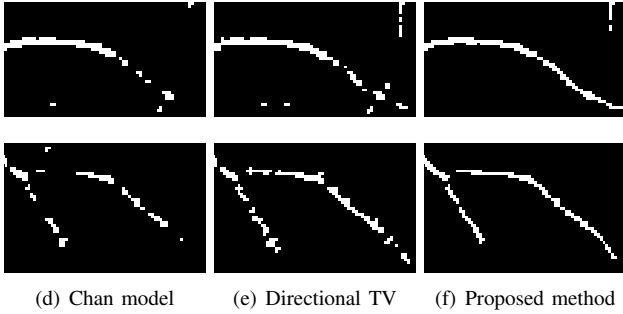


Fig. 3. Zoom of results from Figure 2. Top row corresponds to box 1 and bottom row to box 2.

Algorithm 1: Decoupled segmentation with learnt reconnecting term

Data: $\alpha \in \mathbb{N}^{+*}$, $u_0 \in \mathbb{R}^{N^2}$, $v_0 \in \mathbb{R}^{2N^2}$, $(\tau, \sigma) \in]0, +\infty[^2$, $\lambda_n \in]0, +\infty[$

for $n \geq 1$ **do**

- $p_n = (u_n - \tau(\nabla h(u_n) + L^T v_n))$
- if** $n < \alpha$ **then**
 - $p_n = \text{prox}_{\iota_{[0,1]}}(p_n)$
- else**
 - $p_n = G_{\text{reco}}(p_n)$
- $q_n = \text{prox}_{\sigma g^*}(v_n + \sigma L(2p_n - u_n))$
- $(u_{n+1}, v_{n+1}) = (u_n, v_n) + \lambda_n((p_n, q_n) - (u_n, v_n))$

We performed a quantitative evaluation of the results by computing the mean sensitivity (TPR), specificity (TNR) and accuracy (Acc) over the 20 images of the test set (see Eq. 7). All metrics were computed in the field of view provided in the dataset for a fair comparison with the literature results.

$$\begin{aligned} \text{TPR} &= \frac{\text{TP}}{\text{TP} + \text{FN}} \quad ; \quad \text{TNR} = \frac{\text{TN}}{\text{TN} + \text{FP}}, \\ \text{Acc} &= \frac{\text{TN} + \text{TP}}{\text{TP} + \text{TN} + \text{FN} + \text{FP}}, \end{aligned} \quad (7)$$

with TP (resp. TN) the number of true positives (resp. negatives), and FP (resp. FN) the number of false positives (resp. negatives).

The quantitative results are shown in Table III. We also included the quantitative results from a recent deep learning approach [15] as a comparison. Illustrations of the results are presented in Fig. 2 and Fig. 3.

Qualitatively, we observe that our reconnecting term increases the connectivity of the whole vascular network. It also yields a better connectivity than the directional total variation. Moreover, blood vessels seem smoother than the ones obtained with the other two methods, which is for instance an appealing feature for modeling and blood flow simulations. Our results also exhibit less noise, as evidenced by the higher TNR. However, the mean TPR of our approach is lower than for directional TV. We observed that directional TV tends to detect fragmented vessel parts at the extremities of the network. Such fragments are irrelevant for applications requiring connectivity. By contrast, our approach tends to only preserve connected vessels: we found that our approach detected 230% less connected components than the directional TV approach. This translates into a lower number of true positives, but a better overall quantitative performance and qualitative results for connectivity-requiring applications.

We added the quantitative results from a recent deep learning approach [15] as a baseline for comparison. These approaches are trained with images from the same dataset and thus yield better results. However, these methods do not generalize well on new data (*e.g.* blood vessels from another organ or another modality). They need to be retrained on each new dataset with their corresponding annotations. Conversely, our approach only requires annotations to train the reconnecting term. Once it is trained, it can be applied in the variational segmentation framework on new data without requiring annotations. To show the power of generalization of our method, we applied the same reconnecting term used on the DRIVE dataset to a different type of image containing blood vessel-like structures. Figure 4 shows the segmentation results on this image. We observe that even though the initial image is different from the STARE dataset, our reconnecting term generalizes well and is able to reconnect the structures of interest whereas a trained U-Net has more difficulties to generalize to this new image type. In this case it seems that

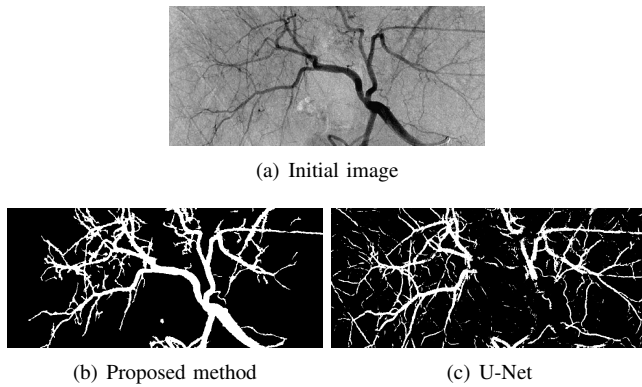


Fig. 4. Illustration of the generalization power of our reconnecting term.

TABLE I
QUANTITATIVE SEGMENTATION RESULTS ON THE DRIVE DATABASE.

	TPR	TNR	Acc
Chan model [10]	0.6615	0.9833	0.9423
Directional TV model [2]	0.6822	0.9809	0.9427
Our approach	0.6631	0.9839	0.9429
Deep learning approach (Arias <i>et al.</i> [15])	0.8597	0.9690	0.9563

the network is not able to generalize on images with a larger range of vessel diameters.

IV. CONCLUSION

We developed a reconnecting term for blood vessel segmentation. This term learns to reconnect blood vessels by focusing on their geometry instead of learning grey-level features in the image. By plugging this term into a variational segmentation framework as a decoupled approach, we are able to better preserve the connectivity of vascular networks than standard regularization terms. Our approach can be applied to unseen data and performs well, unlike deep learning segmentation approaches that have to be trained on each new dataset. Future work will extend our reconnecting term in 3D for the segmentation of various 3D vascular networks, where annotations are far more complicated to obtain. We will also modify our reconnecting network to learn a maximally monotone operator to ensure the convergence of our algorithm [16].

REFERENCES

- [1] S. Moccia *et al.* “Blood vessel segmentation algorithms—review of methods, datasets and evaluation metrics”. In: *Comp. methods and programs in biomed.* 158 (2018), pp. 71–91.
- [2] O. Merveille *et al.* “nD variational restoration of curvilinear structures with prior-based directional regularization”. In: *Trans. on Image Processing* (2019), pp. 3848–3859.
- [3] A. Wu *et al.* “Deep vessel tracking: A generalized probabilistic approach via deep learning”. In: *International Symposium on Biomedical Imaging*. IEEE, 2016, pp. 1363–1367.
- [4] M. Haft-Javaherian *et al.* “A topological encoding convolutional neural network for segmentation of 3D multiphoton images of brain vasculature using persistent homology”. In: *Computer Vision and Pattern Recognition Workshops*. 2020, pp. 990–991.
- [5] G. Tetteh *et al.* “Deepvesselnet: Vessel segmentation, centerline prediction, and bifurcation detection in 3-d angiographic volumes”. In: *Frontiers in Neuroscience* 14 (2020).
- [6] D. Gilton, G. Ongie, and R. Willett. “Neumann networks for linear inverse problems in imaging”. In: *Trans. on Comp. Imaging* 6 (2019), pp. 328–343.
- [7] F. Heide *et al.* “Flexisp: A flexible camera image processing framework”. In: *ACM Transactions on Graphics* 33.6 (2014), pp. 1–13.
- [8] A. Kofler *et al.* “Neural networks-based regularization for large-scale medical image reconstruction”. In: *Physics in Medicine & Biology* 65.13 (2020), p. 135003.
- [9] T. Meinhardt *et al.* “Learning proximal operators: Using denoising networks for regularizing inverse imaging problems”. In: *International Conference on Computer Vision*. 2017, pp. 1781–1790.
- [10] T.F. Chan, S. Esedoglu, and M. Nikolova. “Algorithms for finding global minimizers of image segmentation and denoising models”. In: *SIAM journal on applied mathematics* 66.5 (2006), pp. 1632–1648.
- [11] E. Kerfoot *et al.* “Left-ventricle quantification using residual U-Net”. In: *International Workshop on Statistical Atlases and Computational Models of the Heart*. 2018, pp. 371–380.
- [12] A.D. Hoover, V. Kouznetsova, and M. Goldbaum. “Locating blood vessels in retinal images by piecewise threshold probing of a matched filter response”. In: *Trans. on Medical imaging* 19.3 (2000), pp. 203–210.
- [13] N. Komodakis and J.C. Pesquet. “Playing with duality: An overview of recent primal-dual approaches for solving large-scale optimization problems”. In: *Signal Processing Magazine* 32.6 (2015), pp. 31–54.
- [14] J. Staal *et al.* “Ridge-based vessel segmentation in color images of the retina”. In: *Trans. on Medical Imaging* 23.4 (2004), pp. 501–509.
- [15] M.E.G. Arias, I.P. Santos D.M. Borrero, and M.J.V. Vázquez. “A new deep learning method for blood vessel segmentation in retinal images based on convolutional kernels and modified U-Net model”. In: *Comp. Methods and Programs in Biomed.* (2021), p. 106081.
- [16] J.C. Pesquet *et al.* *Learning Maximally Monotone Operators for Image Recovery*. 2021. arXiv: 2012.13247.