



Segmentation of Retinal Arteries in Adaptive Optics Images

Nicolas Lermé, Florence Rossant, Isabelle Bloch, Michel Paques, Edouard Koch

► To cite this version:

Nicolas Lermé, Florence Rossant, Isabelle Bloch, Michel Paques, Edouard Koch. Segmentation of Retinal Arteries in Adaptive Optics Images. 2013. hal-00868639v4

HAL Id: hal-00868639

<https://hal.science/hal-00868639v4>

Preprint submitted on 21 Apr 2014 (v4), last revised 23 Apr 2014 (v5)

HAL is a multi-disciplinary open access archive for the deposit and dissemination of scientific research documents, whether they are published or not. The documents may come from teaching and research institutions in France or abroad, or from public or private research centers.

L'archive ouverte pluridisciplinaire **HAL**, est destinée au dépôt et à la diffusion de documents scientifiques de niveau recherche, publiés ou non, émanant des établissements d'enseignement et de recherche français ou étrangers, des laboratoires publics ou privés.

Segmentation of Retinal Arteries in Adaptive Optics Images

Nicolas Lermé^{*†}, Florence Rossant^{*}, Isabelle Bloch[†], Michel Paques[‡] and Edouard Koch[‡]

^{*} LISITE, Institut Supérieur d'Électronique de Paris, Paris, France

[†] LTCI, CNRS UMR5141, Institut Mines-Télécom, Télécom ParisTech, Paris, France

[‡] CIC 503, Centre Hospitalier National des Quinze-Vingts, Paris, France

Abstract—In this paper, we present a method for automatically segmenting the walls of retinal arteries in adaptive optics images. To our best knowledge, this is the first method addressing this problem in such images. To achieve this goal, we propose to model these walls as four curves approximately parallel to a common reference line located near to the center of vessels. Once this line detected, the curves are simultaneously positioned as near as possible to the borders of walls using an original tracking procedure to cope with deformations along vessels. Then, their positioning is refined using a deformable model embedding a parallelism constraint. Such an approach enables us to control the distance of the curves to their reference line and improve the robustness to image noise. This model was evaluated on healthy subjects by comparing the results against segmentations from physicians. Noticeably, the error introduced by this model is smaller or very near to the inter-physicians error.

Keywords: Active contours model, adaptive optics, approximate parallelism, retina imaging.

I. INTRODUCTION

Arterial hypertension (AH) and diabetic retinopathy (DR) mainly and precociously affect the physiology and structure of retinal blood vessels of small diameter (i.e. less than $150\mu\text{m}$). According to the Public Health Agency of Canada, AH and DR affected 15 to 20% of the world's adult population in 2009. These modifications can result in a thickening or a narrowing of walls and are predictive of end-organ damage such as stroke or visual loss [1], [2]. For instance, the authors of [3] estimate that 98% of visual damages could be avoided if DR was treated in time. Accurate measurements such as WLR (Wall-to-Lumen Ratio) are therefore necessary to better prevent the DR and the complications of AH. However, classical fundus photographs and Doppler-based measurements cannot capture such level of details due to their limited spatial resolution [4].

Adaptive-Optics (AO) based cameras improve the lateral resolution of fundus photographs, thus enabling the visualization of microstructures such as photoreceptors [6], capillaries [7] and vascular walls [8], noninvasively. In the present study, the camera rtx1 [9] is used to acquire 2D images by flood illumination at 10Hz using a 850nm LED light source with a pixel-resolution of $0.8\mu\text{m}$. Flood-illumination systems usually produce noisy images making walls hardly visible. A commonly used solution is to register these images and average them to increase the signal-to-noise ratio [5]. In the latter, blood vessels appear as dark elongated structures with a bright linear axial reflection, over a textured background. Outer

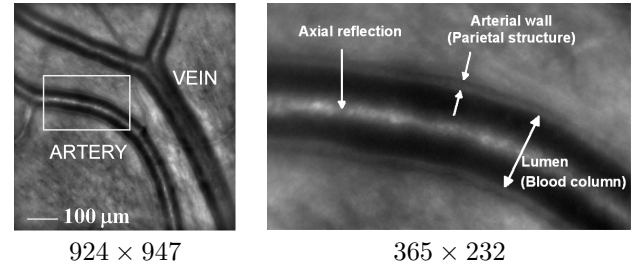


Figure 1: A mean image from [5] (left) and a detailed view of an artery (right). The size of each image is indicated below.

borders of walls are however only visible along arteries and the present study will therefore focus on them. Parietal structures (arterial walls) appear as a gray line along both sides of the lumen (blood column), with a typical thickness of about 15% of the latter [10] (see Figure 1).

In this paper, we propose an automatic procedure for segmenting arterial walls in a selected region of interest ¹ in the mean images produced by [5]. To our best knowledge, this is the first method addressing this problem in such images. This task is challenging for multiple reasons: (i) background is highly textured, (ii) lumens are globally dark but with significant intensity variation along them, (iii) axial reflections may locally show discontinuities or poor contrast, (iv) outer borders of walls are low-contrasted, (v) segments can be locally blurred due to the geometry of the retina, and (vi) deformations can occur along vessels in case of pathologies.

To overcome these difficulties, we propose a strategy exploiting geometric, radiometric and topological *a priori* information of vessels. More precisely, we model arterial walls as four curves approximately parallel to a common reference line located near to the axial reflection. Once this line detected, the curves are simultaneously positioned as near as possible to the borders of walls using a tracking procedure to cope with deformations along vessels. Then, their positioning is refined using an active contours approach [11] where curves evolve towards large image gradients under a parallelism constraint. Such an approach enables us to control the distance of the curves to their reference line and improve the robustness to image noise. Finally, we also want to mention that this work has recently permitted us to establish relationships between morphometric measurements and clinical parameters [12].

¹In particular, we do not aim at segmenting the whole vascular tree. This problem will be the subject of a forthcoming paper.

The rest of this paper is organized as follows. In Section II, we first summarize the steps for detecting axial reflections. Then, we detail in Section III the procedure for segmenting the arterial walls. Finally, we evaluate the relevance of the approach against manual segmentations in Section IV.

II. AXIAL REFLECTION DETECTION

In this section, we consider images as functions mapping points from $\Omega \subset \mathbb{Z}^2$ into the interval $[0, 1]$.

A. Pre-processing

The original image (see Figure 2(a)) is first enhanced by applying a median filter followed by a non-linear diffusion filter [13]. This combination of filters allows us to denoise the image and smooth the blood vessels while preserving the contrast along their edges. We denote by I_P the resulting image (see Figure 2(b)).

B. Detection of bright elongated structures

To enhance the bright elongated structures, a white top-hat transform is applied on the pre-processed image I_P with a binary disk whose radius is about $1/3$ of the axial reflection diameter. We denote by I_T the resulting image (see Figure 2(c)). Then, we binarize the image I_T by hysteresis thresholding and denote it by I_{ES} (see Figure 2(d)). Parts of the axial reflection of vessels are thus extracted, but also other bright areas of the textured background. Further processing steps are therefore necessary to discard these undesired areas.

C. Detection of the darkest areas

K-means classification ($k = 3$) is performed on the pre-processed image I_P (see Figure 2(e)) and the cluster of lowest mean intensity value provides a first binary image of the darkest regions. It is then post-processed with morphological operations to get the main connected components corresponding to the dark areas of lumens. We denote by I_{DA} the resulting image (see Figure 2(f)).

D. Extraction of vascular segments by information fusion

A first selection of vascular segments is performed based on a simple measure of the tortuosity. Let us denote by I_{ES}^L the binary image of a tested connected component of the image I_{ES} . This component is retained if

$$\frac{\#I_{ES}^L}{\#(I_{ES}^L \bullet S)} > 0.8, \quad (1)$$

where $\#$ denotes the cardinality of a set, \bullet denotes the closing operator and S is a binary disk whose radius is 15 pixels. Moreover, a segment of axial reflection must lie inside a dark area, and conversely, a dark region of the lumen must contain at least one axial reflection segment. We denote by $I_{ES'}$ the binary image made of the components satisfying (1) and $I_{ES'}^L$ a tested connected component of it. Such a component is kept as part of an axial reflection segment if

$$\#(I_{DA} \cap (I_{ES'}^L \oplus S')) > \frac{\#I_{ES'}^L}{5}, \quad (2)$$

where \oplus denotes the dilation operator and S' is a binary disk whose radius is 15 pixels. Notice that the radius of S and S' are in accordance to the minimum size of the vessels that are studied. We denote by $I_{ES''}$ the binary image made of the components satisfying (2) (see Figure 2(g)). Morphological operations are applied to I_{DA} , including reconstruction by dilation with the marker $I_{ES''}$, in order to get the final lumen mask I_{LM} (see Figure 2(h)).

E. Segment labeling and reconnection

We first compute the skeleton of the image $I_{ES''}$ to get the end-points of the retained segments (see Section II-D). These end-points are then reconnected using minimal path techniques [14], [15]. These techniques aim at extracting curves of minimal length, in a Riemannian metric computed from the image and depending on the targeted application. A minimal path C connecting two end-points p and q is obtained by minimizing the following functional:

$$L[C] = \int_p^q \mathcal{P}(C(s)) ds,$$

where s denotes the curvilinear abscissa and \mathcal{P} is a potential inducing the metric defined as

$$\mathcal{P}(x) = w_1(1 - I_T(x))^2 + w_2(1 - I_{SM}(x))^2 + w_3, \quad \forall x \in \Omega,$$

where I_{SM} is the spurred skeleton of the lumen mask I_{LM} filtered by a Gaussian of standard deviation σ and $w_1, w_2, w_3 \in \mathbb{R}^+$ are free parameters. These parameters are empirically set with $\sigma = 10$, $w_1 = 0.5$, $w_2 = 0.45$ and $w_3 = 0.05$. In the latter expression, the first term is derived from the top-hat image I_T (considering that the values should ideally be close to one along the axial reflection) while the second one encourages the path C to pass near the middle of the lumen mask I_{LM} . The last term is a regularization constant. The combination of the above criteria allow for a good robustness against the variety of the encountered images.

Two end-points form a candidate pair for reconnection if they belong to the same connected component in the lumen mask I_{LM} and if they do not belong to the same connected component in the image $I_{ES''}$. The candidate pairs are then processed by decreasing order of the Euclidian distance (to start with points that are close to each other) and reconnected using the above procedure. A new skeleton is then calculated, providing the axial reflection of the vessels, and the vessel branches are then labeled (see Figure 2(i)). The vessel branches are individually regularized using a classical parametric active contour [16] with Gradient Vector Flow [17]. The lumen mask I_{LM} is also labeled such that every non-null pixel receives the label of the closest branch (see Figure 2(j)).

Although the above described steps rely on a number of parameters, we empirically found that they are stable for the images studied (including those presented in Section IV).

III. SEGMENTATION OF ARTERIAL WALLS

For convenience, we detail the procedure for segmenting arterial walls on a single regularized vessel branch obtained at the end of the axial reflection detection step (see Section II). We denote this regularized branch as the reference line $V(s) = (x(s), y(s))^T$ of the vessel, parameterized by s . Once obtained,

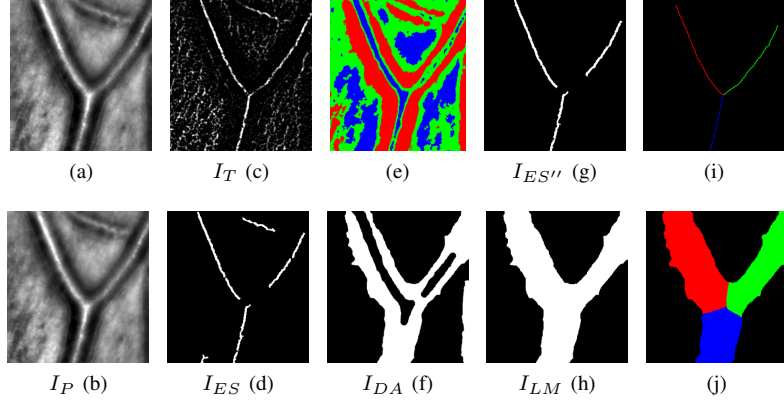


Figure 2: Steps for axial reflection detection: The source image (a) is first pre-processed (b). Dark elongated structures are detected from (b) by top-hat filtering (c) followed by hysteresis thresholding (d). Darkest areas are detected from (b) by k-means (e) followed by post-processing (f). Axial segments (g) and lumens (h) are extracted from (d) and (f). Axial segments (i) and lumens (j) are labeled from (c), (g) and the spurred skeleton of (h).

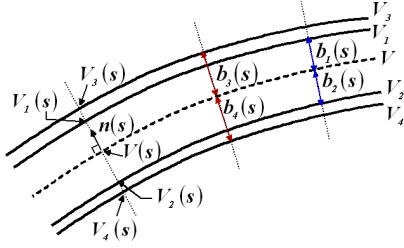


Figure 3: Parametric representation of the proposed model.

this line is considered to be fixed and will therefore no longer evolve in the subsequent steps. Additionally, we choose to model the walls as four curves approximately parallel to this line. We respectively denote by V_1 , V_2 and V_3 , V_4 the inner and outer borders of walls, and define them by

$$\begin{cases} V_1(s) = V(s) + b_1(s)\vec{n}(s) \\ V_2(s) = V(s) - b_2(s)\vec{n}(s) \\ V_3(s) = V(s) + b_3(s)\vec{n}(s) \\ V_4(s) = V(s) - b_4(s)\vec{n}(s), \end{cases}$$

where $\vec{n}(s)$ is the normal vector to the curve V and $b_k(s)$ is the local distance between the reference line V and the curve V_k , $\forall k \in \{1, \dots, 4\}$ (see Figure 3). This model allows a direct correspondence between each curve V_k and the reference line V . In what follows, we first detail how these curves are roughly positioned near to the borders of walls. Next, we describe the model used to refine their positioning.

A. Pre-segmentation

First, we discretize the reference line V in m equally spaced points and denote by $V_i = (x(s = ih), y(s = ih))^T$ the discrete coordinates at the point V_i and \vec{n}_i the associated normal vector (h is the discretization step). We also discretize the curves representing the walls with

$$\begin{cases} V_i^1(b_i^{int}) = V_i + b_i^{int}\vec{n}_i \\ V_i^2(b_i^{int}) = V_i - b_i^{int}\vec{n}_i \\ V_i^3(b_i^{ext}) = V_i + b_i^{ext}\vec{n}_i \\ V_i^4(b_i^{ext}) = V_i - b_i^{ext}\vec{n}_i, \end{cases} \quad (3)$$

where b^{int} and b^{ext} respectively denote the half-diameter of the inner and outer borders. The model (3) assumes that the inner and outer borders lie at the same distance from the reference line V . Although this assumption could appear to be strong, it is verified for a large number of the images presented in Section I. Also, the presegmentation described below leads to a preliminary result which will be refined in Section III-B.

Let us now denote by I a grayscale image with values in $[0, 1]$. We denote by $D_{\vec{u}}I(p)$ the derivative of I in the direction \vec{u} at the point p . Moreover, for an half-diameter $b \in \mathbb{R}^+$, a point V_i and a window of size $(2r + 1)$, we define the mean intensity along the curves V^1 and V^2 by

$$\bar{I}^{int}(b, i, r) = \frac{1}{2(2r + 1)} \sum_{j=-r}^{+r} \left(I(V_{i+j}^1(b)) + I(V_{i+j}^2(b)) \right), \quad (4)$$

the mean local gradient along the curves V^1 and V^2 by

$$\bar{D}^{int}(b, i, r) = \frac{1}{2(2r + 1)} \sum_{j=-r}^{+r} \left(D_{\vec{n}_{i+j}} I(V_{i+j}^1(b)) + D_{-\vec{n}_{i+j}} I(V_{i+j}^2(b)) \right), \quad (5)$$

and the mean local gradient along the curves V^3 and V^4 by

$$\bar{D}^{ext}(b, i, r) = \frac{1}{2(2r + 1)} \sum_{j=-r}^{+r} \left(|D_{\vec{n}_{i+j}} I(V_{i+j}^3(b))| + |D_{-\vec{n}_{i+j}} I(V_{i+j}^4(b))| \right). \quad (6)$$

Increasing the window radius r makes gradient measures more robust to noise but less reliable where strong deformations occur along vessels. This parameter therefore requires a trade-off. Due to the profile of lumens and since we do not use absolute values in (5), we expect this quantity to be larger near to the inner borders of walls than near to the axial reflection. We now summarize the steps necessary to estimate the half-diameters b^{int} and b^{ext} in (3).

Step 1

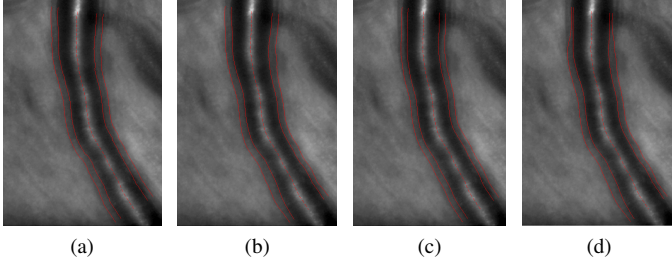


Figure 4: Benefit of using a presegmentation with varying half-diameters (a),(b) against constant ones (c),(d) for a pathological case. Here, we set $r = 10$ and $\alpha = 0.95$. In (a) and (c), the image is superimposed with the presegmentations while it is superimposed in (b) and (d) with the segmentations obtained using the model described in Section III-B.

We jointly search for a constant half-diameter of inner and outer borders maximizing the mean gradient divided by the mean intensity along the curves V^1, V^2, V^3 and V^4 . Using Equations (4), (5) and (6), such half-diameters are given by

$$(b^{int*}, b^{ext*}) = \underset{\substack{b^{int}, b^{ext} \\ b^{int} < b^{ext}}}{\operatorname{argmax}} \left[\frac{\sum_i \bar{D}^{int}(b^{int}, i, r)}{\sum_i \bar{I}^{int}(b^{int}, i, r)} + \frac{1}{m} \sum_i \bar{D}^{ext}(b^{ext}, i, r) \right], \quad (7)$$

where we remind that m is the number of discrete points on V . Dividing by the intensity encourages gradients in dark areas. To speed up this step, the search interval for b^{ext} is restricted to typical WLR values [10]. Such a simplistic approach can however fail to accurately segment the walls when deformations occur along vessels. Estimating varying half-diameters for inner and outer borders is therefore crucial to take into account these deformations (see Figure 4).

Step 2

We search for the position along the vessels having the largest contrast along the curves defined by the half-diameters b^{int*} and b^{ext*} found at Step 1. This position is given by

$$i^* = \underset{i}{\operatorname{argmax}} \frac{1}{2} \left(\bar{D}^{int}(b^{int*}, i, r) + \bar{D}^{ext}(b^{ext*}, i, r) \right).$$

Step 3

Finally, we jointly search for a variable half-diameter of inner and outer borders whose difference (the wall thickness) is constant and having the largest contrast along the curves defined by the half-diameters b^{int*} and b^{ext*} found at Step 1. These half-diameters are constructed iteratively under a regularity constraint from each side of the position i^* found at Step 2. For a fixed gap e , the half-diameter of inner borders b^{int*} is constructed as follows:

$$b_i^{int*} = \begin{cases} b^{int*} & \text{if } i = i^* \\ \operatorname{argmax}_{b_i^{int}} E(b_i^{int}, b_{i-1}^{int*}, i, e, r) & \text{if } i > i^* \\ \operatorname{argmax}_{b_i^{int}} E(b_i^{int}, b_{i+1}^{int*}, i, e, r) & \text{if } i < i^*, \end{cases} \quad (8)$$

with

$$E(b_i^{int}, b_j^{int*}, i, e, r) = \alpha \left(\frac{\bar{D}^{int}(b_i^{int}, i, r)}{\bar{I}^{int}(b_i^{int}, i, r)} + \bar{D}^{ext}(b_i^{int} + \bar{e} + e, i, r) \right) + (1 - \alpha)(b_i^{int} - b_j^{int*})^2, \quad (9)$$

where $\bar{e} = (b^{ext*} - b^{int*})$ and $\alpha \in [0, 1]$ a parameter controlling the amount of regularity. In (9), notice that the term on the left has the same form as (7), except that it only concerns a single point. Finally, we search the best gap with

$$e^* = \underset{e}{\operatorname{argmax}} \sum_{i>0} E(b_i^{int*}, b_{i-1}^{int*}, i, e, r), \quad (10)$$

and where b^{int*} is constructed using (8) and (9). Notice that the discretization step of the search intervals of (8) and (10) is divided by two. The half-diameters estimates of inner and outer borders are finally respectively given by $b^{int} = b^{int*}$ and $b^{ext} = b^{int*} + (b^{ext*} - b^{int*}) + e^*$. Rough estimates of inner and outer borders are now fully defined using (3).

B. Refined segmentation

The model proposed in [11] simultaneously evolves two curves under a parallelism constraint. In what follows, we describe an extension of this model for extracting four curves V_1, V_2, V_3 and V_4 almost parallel to a reference line V . Since this line is fixed in our situation, the energy becomes

$$E(V_1, \dots, V_4, b_1, \dots, b_4) = \sum_{k=1}^4 \left(E_{Image}(V_k) + R(V_k, b_k) \right), \quad (11)$$

where the term

$$E_{Image}(V_k) = \int_0^1 P(V_k(s)) ds$$

is designed to attract the curve V_k towards large intensity gradients (see [16]). In this context, the term E_{Image} is based on the Gradient Vector Flow [17]. The role of the term R in (11) is to control the variation of the distance b_k , thus imposing a local parallelism. The authors of [11] proposed a function of the derivative of b_k with

$$R(V_k, b_k) = \int_0^1 Q(s, b'_k) ds = \int_0^1 \varphi_k(s) (b'_k(s))^2 ds,$$

where $\varphi_k(s) \in \mathbb{R}^+$ are application-dependent parameters that locally control the strength of the parallelism of the curve V_k with respect to the reference line V . More precisely, the larger the parameter $\varphi_k(s)$ is, the more strict is the parallelism to the reference line V . It is worth underlying that the distance between both curves has not to be known. It is adjusted during the evolution process and can vary along boundaries. Notice also that the energy (11) does not ensure that $b_1(s) < b_3(s)$ and $b_2(s) < b_4(s)$. However, we never encountered such behavior in our experiments (and in particular, for those detailed in Section IV).

Since the energy (11) does not have crossing terms involving different curves, the minimization can be independently

done for each curve V_k . For any $k \in \{1, \dots, 4\}$, the Euler-Lagrange equation expresses the minimization of (11) with respect to $b_k(s)$

$$\frac{\partial P(V_k(s))}{\partial b_k} - \frac{d}{ds} \frac{\partial Q(s, b'_k)}{\partial b'_k} = 0,$$

and the evolution of the distance b_k to the line V is driven by

$$\langle \vec{n}, -\nabla P(V_k(s)) \rangle - 2[\varphi_k(s)b''_k(s) + \varphi'_k(s)b'_k(s)] = 0. \quad (12)$$

The latter equation is solved by discretizing it and introducing the time variable using standard numerical approximations of derivatives (central difference in space, backward difference in time). The resolution of the above equations stops when

$$\max_{k \in \{1, \dots, 4\}} \left\{ \max_s |b_k^n(s) - b_k^{n-1}(s)| \right\} \leq \varepsilon.$$

In the latter expression, $b_k^n(s)$ is the estimate of the distance of the curve V_k to the reference line V at iteration n and $\varepsilon \simeq 0$ is an accuracy parameter. An example of resolution using this model is illustrated in Figure 4(c).

IV. EVALUATION

Thirteen images from healthy subjects were manually delineated by three physicians. These physicians have several years of experience in the field of AO image interpretation. The images were selected to ensure the representativeness of the quality and the noise levels encountered by physicians during routine clinical. Let us denote resp. by V^M and V^A a manual segmentation and an automatic segmentation. For each image, using the same axial reflection, we measure the absolute relative difference on the inner diameter, the outer diameter and the total wall thickness (i.e. the difference between outer and inner diameters), resp. defined for each point by

$$\delta_{int}(V^M, V^A) = \frac{|d_{int}(V^M) - d_{int}(V^A)|}{d_{int}(V^M)} \times 100, \quad (13)$$

$$\delta_{ext}(V^M, V^A) = \frac{|d_{ext}(V^M) - d_{ext}(V^A)|}{d_{ext}(V^M)} \times 100, \quad (14)$$

$$\delta_{wt}(V^M, V^A) = \frac{|d_{wt}(V^M) - d_{wt}(V^A)|}{d_{wt}(V^M)} \times 100, \quad (15)$$

where d_{int} , d_{ext} and d_{wt} denote resp. the inner diameter, the outer diameter and the total wall thickness. The measure δ_{wt} is of great importance for us due to its high sensitivity. Also, measurements were only taken into account where no vessel bifurcation occurs (see Figure 5). For the parameters, we use $\varepsilon = 0.1$, $\alpha = 0.9$, $r = 10$ and $\varphi_k = 100$, $\forall k \in \{1, \dots, 4\}$. Also, we choose manual segmentations from the most experienced physician as a reference (denoted by $Phys_{Ref}$). For each image, we then compute the mean and standard deviation of (13), (14) and (15) between the segmentations obtained by the automatic procedure and the those from the physician $Phys_{Ref}$. The same statistics are also computed between the segmentations from physicians (different of $Phys_{Ref}$) and those from the physician $Phys_{Ref}$. The results of these experiments are detailed in Table I and illustrated in Figure 5. To put in perspective these results, we provide between parentheses in Table I, the above statistics but for a displacement of one pixel all along a curve. In that case, the numerators of (13), (14) and (15) become equal to one.

Due to the size of arterial walls, we first remark that the error (i.e. mean and standard deviation) is much more larger on total wall thickness compared to inner and outer diameters. Noticeably, the error introduced by the automatic procedure is globally smaller than the inter-physicians error on total wall thickness and outer diameter. Although this is not the case concerning the inner diameter, the errors remain globally very near from it. For inner / outer diameters and total wall thickness measurements between the automatic procedure and the physician $Phys_{Ref}$, the standard deviation is globally smaller than the mean. This means a good robustness of the proposed approach. For some subjects, the error introduced by the automatic procedure is larger than the inter-physicians error. For some of them, we do believe that this is due to a lack of accuracy in manual segmentations. Due to the poor contrast along walls and the size of these structures, it is indeed difficult for physicians to delineate them all along a vessel with high accuracy (see Subject 5). For other subjects, the outer border of walls computed by the automatic procedure slightly differ from manual segmentations. This is for instance the case for the Subject 3 where a kind of double contour occur all along the vessel. However, there is currently no consensus among physicians about the ideal location of the outer border of walls in such situation. Finally, we are currently evaluating the proposed approach on pathological subjects. First experiments show a good robustness with respect to morphological deformations.

REFERENCES

- [1] N.H. Buus et al. Small artery structure during antihypertensive therapy is an independent predictor of cardiovascular events in essential hypertension. *Journal of Hypertension*, 31(4):791–797, 2013.
- [2] A.M. Heagerty et al. Small artery structure in hypertension: Dual processes of remodeling and growth. *Journal of Hypertension*, 21:391–397, 1993.
- [3] H. Taylor and J. Keeffe. World blindness: A 21st century perspective. *British Journal of Ophthalmology*, 85(3):261–266, 2001.
- [4] F. Rossant, M. Badellino, A. Chavillon, I. Bloch, and M. Paques. A morphological approach for vessel segmentation in eye fundus images, with quantitative evaluation. *Journal of Medical Imaging and Health Informatics*, 1(1):42–49, 2011.
- [5] C. Kulcsár, G. Le Besnerais, E. Ödlund, and X. Levecq. Robust processing of images sequences produced by an adaptive optics retinal camera. In *Optical Society of America, Adaptive Optics: Methods, Analysis and Applications*, page OW3A.3, 2013.
- [6] J. Liang, D.R. Williams, and D.T. Miller. Supernormal vision and high-resolution retinal imaging through adaptive optics. *Journal of Optical Society America A*, 14(11):2884–2892, 1997.
- [7] J.A. Martin and A. Roorda. Direct and noninvasive assessment of parafoveal capillary leukocyte velocity. *Ophthalmology*, 112(12):2219–2224, 2005.
- [8] T.Y. Chan, D.A. Vannasdale, and S.A. Burns. The use of forward scatter to improve retinal vascular imaging with an adaptive optics scanning laser ophthalmoscope. *Biomedical Optics Express*, 3(10):2537–2549, 2012.
- [9] C. Viard, K. Nakashima, B. Lamory, X. Paques, M. ad Levecq, and N. Château. Imaging microscopic structures in pathological retinas using a flood-illumination adaptive optics retinal camera. In *Photonics West: Biomedical Optics (BiOS)*, volume 7885, pages 788 509+, 2011.
- [10] E. Koch. Morphometric study of human retina arteriolar in high resolution imaging. Master's thesis, Université Pierre et Marie Curie, 2012.
- [11] I. Ghorbel, F. Rossant, I. Bloch, and M. Paques. Modeling a parallelism constraint in active contours. Application to the segmentation of eye vessels and retinal layers. In *International Conference on Image Processing (ICIP)*, pages 445–448, 2011.

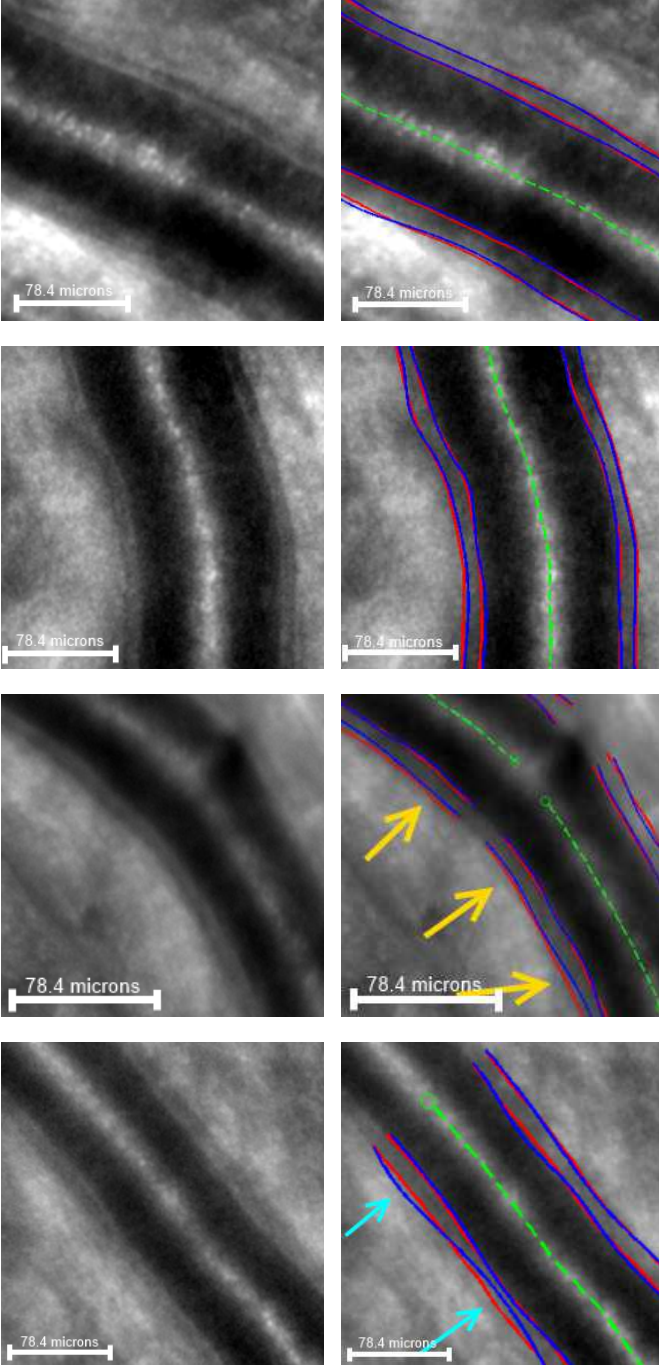


Figure 5: From top to bottom: positive (half-up) and negative (half-down) results against the reference physician $Phys_{Ref}$ for Subjects 7, 11, 3 and 5 (see Table I). The manual and the automatic segmentation are drawn in blue and red, respectively. The green dashed line is the reference line. The original images are provided in the left column while they are superimposed to the segmentations in the right column. Yellow and cyan arrows point misplacement of borders in automatic and manual segmentations, respectively.

Subject	Inter-physicians	Our method / $Phys_{Ref}$
1	2.29 ± 1.54 (1.13 ± 0.04)	2.07 ± 1.06 (1.13 ± 0.04)
2	3.50 ± 2.37 (2.47 ± 0.16)	4.17 ± 3.29 (2.47 ± 0.16)
3	2.86 ± 2.57 (1.37 ± 0.04)	1.63 ± 1.31 (1.37 ± 0.04)
4	2.29 ± 1.7 (0.75 ± 0.03)	3.16 ± 1.65 (0.75 ± 0.03)
5	2.79 ± 1.91 (0.89 ± 0.05)	3.21 ± 2.09 (0.89 ± 0.05)
6	3.75 ± 2.8 (1.13 ± 0.05)	2.14 ± 2.34 (1.13 ± 0.05)
7	2.82 ± 2.38 (0.87 ± 0.04)	2.51 ± 2.64 (0.87 ± 0.04)
8	2.84 ± 2.41 (0.97 ± 0.03)	3.78 ± 2.92 (0.97 ± 0.03)
9	3.26 ± 2.65 (0.87 ± 0.03)	4.58 ± 3.19 (0.87 ± 0.03)
10	3 ± 2.08 (0.8 ± 0.02)	1.87 ± 1.96 (0.8 ± 0.02)
11	2.75 ± 1.8 (0.84 ± 0.02)	2.36 ± 1.88 (0.84 ± 0.02)
12	2.7 ± 2.26 (0.9 ± 0.04)	2.51 ± 2.05 (0.9 ± 0.04)
13	3.38 ± 2.27 (0.84 ± 0.05)	6.07 ± 4.25 (0.84 ± 0.05)
Overall	2.92 ± 2.24 (0.99 ± 0.35)	3.06 ± 2.73 (0.99 ± 0.35)

Subject	Inter-physicians	Our method / $Phys_{Ref}$
1	2.08 ± 1.69 (0.91 ± 0.02)	2.06 ± 1.83 (0.91 ± 0.02)
2	2.2 ± 1.76 (1.7 ± 0.07)	3.86 ± 1.92 (1.7 ± 0.07)
3	2.16 ± 1.53 (1.05 ± 0.03)	5.64 ± 2.82 (1.05 ± 0.03)
4	2.8 ± 1.94 (0.6 ± 0.02)	1.57 ± 1.04 (0.6 ± 0.02)
5	2.94 ± 2.19 (0.67 ± 0.03)	2.28 ± 1.66 (0.67 ± 0.03)
6	5.11 ± 3.81 (0.74 ± 0.03)	3.05 ± 2.1 (0.74 ± 0.03)
7	4.05 ± 2.59 (0.65 ± 0.01)	1.96 ± 1.38 (0.65 ± 0.01)
8	1.93 ± 1.36 (0.69 ± 0.02)	2.23 ± 1.34 (0.69 ± 0.02)
9	2.93 ± 1.84 (0.62 ± 0.01)	2.33 ± 1.01 (0.62 ± 0.01)
10	3.03 ± 1.97 (0.6 ± 0.01)	1.67 ± 1.45 (0.6 ± 0.01)
11	2.18 ± 1.95 (0.68 ± 0.01)	2.19 ± 1.33 (0.68 ± 0.01)
12	2.48 ± 1.54 (0.69 ± 0.03)	5.04 ± 2.27 (0.69 ± 0.03)
13	1.7 ± 1.28 (0.68 ± 0.02)	2.95 ± 2.68 (0.68 ± 0.02)
Overall	2.83 ± 2.31 (0.74 ± 0.24)	2.7 ± 2.16 (0.74 ± 0.24)

Subject	Inter-physicians	Our method / $Phys_{Ref}$
1	10.57 ± 8.1 (4.68 ± 0.59)	9.12 ± 6.33 (4.68 ± 0.59)
2	8.78 ± 6.59 (5.53 ± 0.58)	9.79 ± 6.03 (5.53 ± 0.58)
3	11.63 ± 9.54 (4.63 ± 0.45)	23.11 ± 13.1 (4.63 ± 0.45)
4	14.52 ± 11.95 (3.04 ± 0.43)	13.52 ± 8.35 (3.04 ± 0.43)
5	17.46 ± 14.5 (2.8 ± 0.57)	15.62 ± 11 (2.8 ± 0.57)
6	14.32 ± 10.93 (2.2 ± 0.29)	10.9 ± 7.87 (2.2 ± 0.29)
7	16.12 ± 10.57 (2.54 ± 0.33)	9.82 ± 8.59 (2.54 ± 0.33)
8	9.85 ± 7.64 (2.46 ± 0.22)	12.22 ± 6.22 (2.46 ± 0.22)
19	14.77 ± 12.36 (2.17 ± 0.32)	11.42 ± 5.6 (2.17 ± 0.32)
10	13.17 ± 8.52 (2.36 ± 0.23)	7.4 ± 5.87 (2.36 ± 0.23)
11	15.98 ± 13.26 (3.67 ± 0.36)	15.15 ± 13.8 (3.67 ± 0.36)
12	12.58 ± 10.11 (2.88 ± 0.22)	25.52 ± 8.41 (2.88 ± 0.22)
13	15.69 ± 10.53 (3.5 ± 0.4)	14.68 ± 11.8 (3.49 ± 0.4)
Overall	14.09 ± 11.34 (3.1 ± 0.98)	13.84 ± 10.52 (3.1 ± 0.98)

Table I: From top to bottom: inter-physicians and automatic / $Phys_{Ref}$ absolute relative error for inner diameter outer diameter and total wall thickness. The numbers between parentheses denote a unit displacement all along the curve (see text).

- [12] E. Koch, D. Rosenbaum, A. Brolly, J.-A. Sahel, P. Chaumet-Riffaud, X. Girard, F. Rossant, and M. Paques. Morphometric analysis of small arteries in the human retina using adaptive optics imaging: Relationship with blood pressure and focal vascular changes. *Journal of Hypertension*, 2013.
- [13] J. Weickert, B.H. Romeny, and M. Viergever. Efficient and reliable schemes for nonlinear diffusion filtering. *IEEE Transactions on Image Processing*, 7(3):398–410, 1998.
- [14] L.D. Cohen and R. Kimmel. Global minimum for active contour models: A minimal path approach. *International Journal of Computer Vision*, 24(1):57–78, 1997.
- [15] J.A. Sethian. *Level Set Methods and Fast Marching Methods: Evolving Interfaces in Computational Geometry, Fluid Mechanics, Computer Vision, and Materials Science*. Cambridge University Press, 1999.
- [16] M. Kass, A. Witkin, and D. Terzopoulos. Snakes: Active contour models. *International Journal of Computer Vision*, 1(4):321–331, 1988.
- [17] C. Xu and J.L. Prince. Snakes, shapes and gradient vector flow. *IEEE Transactions on Image Processing*, 7(3):359–369, 1998.