# UNIVERSITE DE LAUSANNE – FACULTE DE BIOLOGIE ET DE MEDECINE

Département de radiologie médicale Service de radiodiagnostic et de radiologie interventionnelle

# Ascending aorta measurements as assessed by ECG-gated multi-detector computed tomography: a pilot study to establish normative values for transcatheter therapies

THESE

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# DOCTEUR EN MEDECINE

par

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Ascending aorta measurements as assessed by ECG-gated multi-detector computed tomography : a pilot study to establish normative values for transcatheter therapies

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Cl adles

Madame le Professeur Stephanie Clarke Directrice de l'Ecole doctorale

# Rapport de synthèse

# Mesures de l'aorte ascendante par scanner synchronisé au rythme cardiaque: une étude pilote pour établir des valeurs normatives dans le cadre des futures thérapies par transcathéter

# Objectif

L'objectif de cette étude est d'établir les valeurs morphométriques normatives de l'aorte ascendante à l'aide de l'angiographie par scanner synchronisé au rythme cardiaque, afin d'aider au développement des futurs traitements par transcathéter.

## Matériels et méthodes

Chez soixante-dix-sept patients (âgé de 22 à 83 ans, âge moyen: 54,7 ans), une angiographie par scanner synchronisé au rythme cardiaque a été réalisée pour évaluation des vaisseaux coronaires. Les examens ont été revus afin d'étudier l'anatomie de la chambre de chasse du ventricule gauche jusqu'au tronc brachio-céphalique droit. A l'aide de programmes de reconstructions multiplanaires et de segmentation automatique, différents diamètres et longueurs considérés comme importants pour les futurs traitements par transcathéter ont été mesurés. Les valeurs sont exprimées en moyennes, médianes, maximums, minimums, écart-types et en coefficients de variation. Les variations de diamètre de l'aorte ascendante durant le cycle cardiaque ont été aussi considérées.

## Résultats

Le diamètre moyen de la chambre de chasse du ventricule gauche était de 20.3+/-3.4 mm. Au niveau du sinus coronaire de l'aorte, il était de 34.2+/-4.1 mm et au niveau de la jonction sinotubulaire il était de 29.7+/-3.4 mm. Le diamètre moyen de l'aorte ascendante était de 32.7+/-3.8 mm. Le coefficient de variation de ces mesures variait de 12 à 17%. La distance moyenne entre l'insertion proximale des valvules aortiques et le départ du tronc brachio-céphalique droit était de 92.6+/-11.8 mm. La distance moyenne entre l'insertion proximale des valvules aortiques et le départ du tronc brachio-céphalique droit était de 31%. La distance moyenne entre les deux ostia coronaires était de 7.2+/-3.1 mm avec un coefficient de variation de 43%. La longueur moyenne du petit arc de l'aorte ascendante entre l'artère coronaire gauche et le tronc brachio-céphalique droit était de 52.9+/-9.5 mm. La longueur moyenne de la continuité fibreuse entre la valve aortique et la valvule mitrale antérieure était de 14.6+/-3.3 mm avec un coefficient de variation de 23%. L'aire moyen de la valve aortique était de 582.0+/-131.9 mm<sup>2</sup>. La variation du diamètre antéro-postérieur et transverse de l'aorte ascendante était respectivement de 8.4% et de 7.3%.

## Conclusion

Il existe d'importantes variations inter-individuelles dans les mesures de l'aorte ascendante avec cependant des variations intra-individuelles faibles durant le cycle cardiaque. De ce fait, une approche personnalisée pour chaque patient est recommandée dans la confection des futures endoprothèses de l'aorte ascendante. Le scanner synchronisé au rythme cardiaque jouera un rôle prépondérant dans le bilan préthérapeutique.

## VASCULAR-INTERVENTIONAL

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#### Introduction

# Ascending aorta measurements as assessed by ECG-gated multi-detector computed tomography: a pilot study to establish normative values for transcatheter therapies

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Abstract The aim of this study was to provide an insight into normative values of the ascending aorta in regards to novel endovascular procedures using ECG-gated multi-detector CT angiography. Seventy-seven adult patients without ascending aortic abnormalities were evaluated. Measurements at relevant levels of the aortic root and ascending aorta were obtained. Diameter variations of the ascending aorta during cardiac cycle were also considered. Mean diameters (mm) were as follows: LV outflow tract  $20.3 \pm 3.4$ , coronary sinus 34.2±4.1, sino-tubular junction  $29.7 \pm 3.4$  and mid ascending aorta  $32.7\pm3.8$  with coefficients of variation (CV) ranging from 12 to 17%. Mean distances (mm) were: from the plane passing through the proximal

insertions of the aortic valve cusps to the right brachio-cephalic artery (BCA)  $92.6 \pm 11.8$ , from the plane passing through the proximal insertions of the aortic valve cusps to the proximal coronary ostium 12.1±3.7, and between both coronary ostia  $7.2\pm3.1$ . minimal arc of the ascending aorta from left coronary ostium to right BCA 52.9  $\pm 9.5$ , and the fibrous continuity between the aortic valve and the anterior leaflet of the mitral value  $14.6\pm3.3$ , CV 13–43%. Mean aortic valve area was  $582.0\pm131.9$  mm<sup>2</sup>. The variation of the antero-posterior and transverse diameters of the ascending aorta during the cardiac cycle were 8.4% and 7.3%, respectively. Results showed large inter-individual variations in diameters and distances but with limited intra-individual variations during the cardiac cycle. A personalized approach for planning endovascular devices must be considered.

Keywords ECG-gated computed tomography · Normative values · Ascending aorta · Endovascular repair · Endograft

Historically treatment of abdominal aortic aneurysms (AAA) was based on surgical repair until the 1990s [1–3]. Since then endovascular graft systems have been widely recognized and used for AAA repair (EVAR) [4]. The lack

of off-the-shelf endografts warrants precise prepoperative imaging to define adequate device selection. Therefore increasing importance has been placed on preoperative CT imaging protocols for the planning of endovascular abdominal aortic aneurysms [5]. With the advent of endovascular aneurysm repair and valve replacement, more focus is now turned to ascending aorta and aortic arch problems [6, 7]. Experience with endografts for ascending aortic disease is scarce because few groups are using such devices due to technological restrictions. On the one hand the lower caseload for ascending aorta endoluminal procedures limits the device evolution. On the other hand the large diameter and the curvature of ascending aorta and aortic arch render access to these vascular structures difficult and hinder the introduction, placement, tracking and deployment of the graft, thus challenging device engineering. Blood flow in the coronary arteries and the supra-aortic arteries should also be maintained, by means of different fenestrated technologies and surgical bypasses depending on the graft landing zone [8, 9].

All these anatomical and technical constraints require precise knowledge of normal ascending aorta. Although normative thoracic aortic dimensions have already been determined by means of CT angiography [10–13], to our knowledge, no exhaustive measurements have been established by means of ECG-gated CT angiography.

Despite improvement in spatial and temporal resolution (below 150 ms) with fast-rotating multi-detector CT technology (MDCT), the thoracic aorta remains a vascular segment that is difficult to analyse due to axial and transverse motion during the cardiac cycle, which is particularly true for the ascending aorta [14–16]. ECG assistance for aortic studies allows the synchronisation of data acquisition to the heart cycle in order to obtain CT data during the diastolic phase when heart movement is minimal thus reducing motion artefact [17–19].

The purpose of this study was to provide an insight into normative values of the ascending aorta and its branches in adults, regarding endovascular procedures and devices by using ECG-gated MDCT.

#### Methods

#### Study population

Seventy-seven patients were included in this study. There were 59 men and 18 women. Age ranged from 22 to 83 years. Mean age and median were 54.7 and 54 years, respectively. Inclusion criteria were: patients older than 18 years old evaluated for coronary artery disease with data available from the outflow tract up to the mid ascending aorta or up to the right brachio-cephalic artery. Exclusion criteria were: outflow tract and/or aortic valve diseases, atherosclerosis of the ascending aorta, dissection or aneurysm of the ascending aorta, surgical procedures of the ascending aorta, important artefacts. Patients who were not under beta-blocker therapy received oral beta-blocker before examination and mean heart rate during acquisition was 68.9 bpm.

#### CT protocol

Data were acquired on a 64-detector VCT system (General Electric Medical Systems, Milwaukee, USA). Acquisitions were obtained after an antecubital intravenous injection of 100-120 ml nonionic contrast medium (Accupaque 350, GE Healthcare, Giles, UK) with 40 ml saline solution as bolus chaser at 4-5 ml s<sup>-1</sup>. Helical CT acquisition was started during a single breath hold when sufficient contrast enhancement (at least 200 HU) was detected in the ascending aorta (bolus tracking technique). All CT data were obtained with a collimation of 0.625 × 40 mm, a FOV of 350 mm, rotation time of 0.35 s and 120 kV with mA modulation during the cardiac cycle (typically 250-750 mA). High resolution (0.625-mm slice thickness, 512×512 matrix) retrospective ECG-gated axial transverse reconstructions were obtained for all patients at 75% of the R-R cycle. Multi-phase reconstructions were also done from 0 to 99% with increments of 6 or 10%.

Data analysis and aortic measurements

In 49 patients, data were available from the outflow tract up to the mid ascending aorta. Data were available up to the right brachio-cephalic artery in the remaining patients (n=28).

Analysis of ascending aorta and aortic arch morphology was performed on a Advantage Window workstation with version 4.3 software (GE Medical Systems). Twodimensional multiplanar reformations and automatic, interactive, centerline-based segmentation of the aorta were used and reviewed in consensus by two radiologists. The reported precision of measurements of the software is less than 0.7 mm.

In close collaboration with cardiovascular surgeons, relevant measurements (Fig. 1) were identified as key parameters for development of future transcatheter cardiovascular therapies (TCT): aortic diameters perpendicular to the aortic centerline axis (Fig. 2) at different levels (outflow tract measured at the level of the insertion of the anterior leaflet of the mitral valve, coronary sinus, sino-tubular junction and mid ascending aorta level); distances from the plane passing through the proximal insertions of the aortic valve cusps in the sinus wall to the right brachio-cephalic artery, from the plane passing through the proximal insertions of the aortic valve cusps to the proximal coronary ostium, between both coronary ostia; and the minimal arc of the ascending aorta from the left coronary ostium to the level of the right brachio-cephalic artery. Aortic valve area was also measured.

Dynamic analysis was done for measurement of the length of the aorto-mitral fibrous continuity in every patient. Anatomically it refers to the septum extending from below the non-coronary and the left coronary cusps to the insertion of the anterior leaflet of the mitral valve [20].

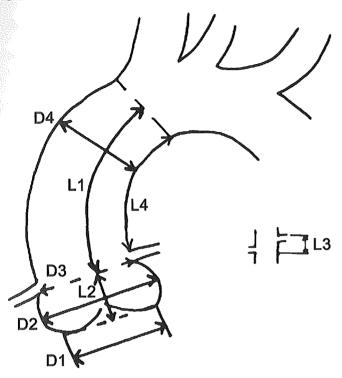


Fig. 1 D1 outflow tract, D2 coronary sinus, D3 sino-tubular junction, D4 mid ascending aorta, L1-L2 distance from the plane passing through the proximal insertions of the aortic cusps to the right brachio-cephalic artery, L2 distance from the plane passing through the proximal insertions of the aortic cusps to the proximal coronary ostia, L3 distance between both coronary ostia, L4 minimal arc of the ascending aorta from the left coronary ostium to the right brachio-cephalic artery

This septum was deemed to be a potential site for graft anchoring. It was identified on a three-cavities view (Fig. 3) during the cardiac cycle and its length was measured from the proximal insertions of the left coronary and the non-coronary cusps in the sinus wall



Fig. 3 Three-cavities view demonstrates the fibrous continuity (*black asterisk*) extending from the left coronary cusp (*white asterisk*) to the insertion of the anterior leaflet (*white circle*) of the mitral valve

to the insertion of the anterior leaflet of the mitral valve. Dynamic data were also used for measurement of the ascending aorta pulsatility. Pulsatility was defined as the percentage difference of minimal and maximal anteroposterior and latero-medial diameters of the mid ascending aorta.

Mean, median, maximum and minimum values were considered with standard deviations. Coefficients of variation were also computed.

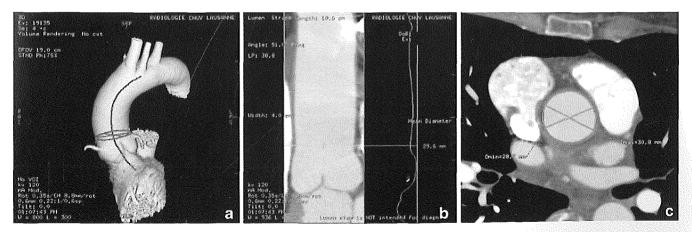


Fig. 2 Example of measurement at the level of the sino-tubular junction in a 54-year-old woman. Volume rendering reconstruction of the ascending aorta and of the aortic arch (a). Lumen view of the ascending aorta (b). Automatic segmentation of the ascending aorta (c)

#### Results

Mean diameters were respectively  $20.3\pm3.4$  mm for outflow tract,  $34.2\pm4.1$  mm for coronary sinus,  $29.7\pm$ 3.4 mm for sino-tubular junction and  $32.7\pm3.8$  mm for mid ascending aorta. Coefficients of variation ranged from 12 to 17%. See Table 1 for details. The stratification of diameters by age is presented in Fig. 4.

Mean distance from the plane passing through the proximal insertions of the aortic valve cusps to the right brachio-cephalic artery was  $92.6\pm11.8$  mm. Mean distance from the plane passing through the proximal insertions of the aortic valve cusps to the proximal coronary artery was  $12.1\pm3.7$  mm with a coefficient of variation of 31%. Mean distance between both coronary ostia was  $7.2\pm3.1$  mm with a coefficient of the minimal arc of the ascending aorta from the left coronary ostium to the level of the right brachio-cephalic artery was  $52.9\pm9.5$  mm. Mean value for the aorto-mitral fibrous continuity was  $14.6\pm3.3$  mm with a coefficient of variation of 23%.

Antero-posterior and transverse diameters of the mid ascending aorta during the cardiac cycle varied from a maximum of 34.2 mm to a minimum of 31.3 mm and from a maximum of 33.6 mm to a minimum of 31.2 mm, respectively. Computed variations are respectively 8.4% and 7.3%.

Mean area of the aortic valve was  $582.0\pm131.9 \text{ mm}^2$  with a coefficient of variation of 23%. The median value was 579.1. The maximum and minimum values were 971.0 and 286.5 mm<sup>2</sup>, respectively.

#### Discussion

Transcatheter therapies of the aortic valve and of the ascending aorta will bring an innovative approach to the treatment of thoracic aorta abnormalities. Nevertheless indications for such types of minimal invasive procedure are still limited. The main restrictions are due to the size limitation of the vascular access and the availability of

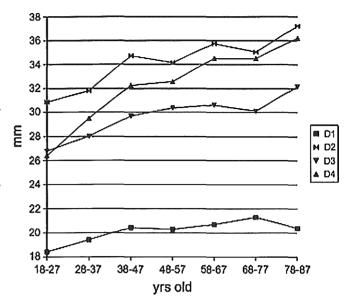


Fig. 4 Mean aortic diameters by age. D1 outflow tract, D2 coronary sinus, D3 sino-tubular junction, D4 mid ascending aorta

dedicated devices. The latter are now being developed, but in the pursuit of the ideal ascending aorta endograft precise morphometric measurements of this vascular segment are needed. This study presents an insight into measurements of the ascending aorta by means of ECG-gated tomography.

ECG-gated MDCT provides high resolution images in a near isotropic condition that allows precise 3D measurements. Although MDCT imparts a high dose of ionizing radiation [21], it remains the gold standard for quantitative imaging owing to its high spatial resolution. MDCT allows fast and readily accessible acquisition of volume data that are prone to easy post-processing. Besides, as already mentioned, motion artefacts—particularly in the ascending aorta—can affect precision of measurements when examination is not ECG-gated.

To our knowledge, no previous studies have reported complete assessment of the ascending aorta by quantifying various parameters. However our results agreed with those

Table 1 Results obtained from ECG-gated multi-detector CT angiography (mm)

	Mean	SD	Median	Minimum	Maximum	CV (%)
D1	20.3	3.4	20.1	12.1	28.6	17
D2	34.2	4.1	34	25.1	45.4	12
D3	29.7	3.4	29.8	21.5	37.8	12
D4	32.7	3.8	32.3	23.3	42.5	12
L1–L2	92.6	11.8	93.7	72.3	119.9	13
L2	12.1	3.7	12.1	2.9	25.6	31
L3	7.2	3.1	6.6	2.4	17.5	43
L4	52.9	9.5	52	32.7	78	18
Fibrous continuity	14.6	3.3	14.7	9.2	26	23

reported in previous studies. In two studies with non-gated  $\hat{\mathbf{CT}}$  in adult patients, the diameter of the aortic sinus was measured between 29.8 and 36.2 mm and the diameter of the ascending aorta between 30.9 and 35.1 mm [10, 13]. A Dutch group used gated CT to measure the distance between the aortic valve and the right brachio-cephalic in 14 patients [22]. Their results ranged from 72 to 99 mm. This group also showed that motion and stress forces in the ascending aorta are higher than in the abdominal aorta with a maximum difference of diameter during the cardiac cycle of up to 27.5%. Our results differ with this reported variation. Such a difference may be due to the studied populations. Our sample was larger and younger (77 patients versus 15, mean age 54.7 years versus 72.2) and heart rates during examination were perhaps different. Nevertheless based on the age-related reduction of distensibility of the aortic wall [23] we think that pulsatility in the ascending aorta especially in the elder patients is probably not that high.

Several other imaging techniques have already been used for evaluating the ascending aorta. In the pediatric population, morphometric measurements have been done with transthoracic echocardiography [24]. Although transthoracic echocardiography or even transesophageal echocardiography are readily available techniques and although gated-ultrasound technologies offer even better temporal resolution compared with computed tomography [25], image acquisition and interpretation are still operator dependent [26]. Interposition between the ascending aorta and the trachea make visualization difficult and sometimes incomplete [27]. Artefacts can also appear in the ascending portion of the aorta due to reflection of ultrasound on the atrio–aortic interface and can cause false-positive findings for example in diagnosis of aortic dissection [28].

Normative data were also obtained by means of angiographic acquisition in the pediatric population [29, 30]. The technique is based on projectional views and summation images. Thus angle of incidence of X-rays and overlapping structures can induce geometrical distortion [31]. Considering that angiography also requires an invasive procedure, it is not the most suitable technique to get aortic measurements.

Magnetic resonance angiography experience for normative measurements was also described. However few data are available [32]. Cardiac magnetic resonance imaging is a promising technology. Nevertheless reduced availability, cost and patient discomfort are disadvantages compared

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 Dubost C, Allary M, Oeconomos N (1951) Aneurysm of the abdominal aorta treated by resection and graft. Arch Mal Coeur Vaiss 44:848–851 with tomography. From a technical point of view fast acquisition sequences like echo-planar imaging (EPI) introduce geometric distortion and ghosting in the cardiac examination post-processing and reconstruction [33]. Strong magnetic fields can also interfere with the ECG signal recording during MRI scans, the so-called magnetohydrodynamic effect [34].

Ideally one standard endovascular device should fit any patient's anatomy. Experience with EVAR showed that this is practically impossible and careful selection of size, geometrical configuration and anchoring options is needed, depending on anatomical morphology and the nature of the lesion to be treated [35]. Our results confirmed that large inter-individual variations exist. Distances between coronary ostia and the level of insertions of the aortic valve cusps and mean distances between both ostia showed coefficients of variation up to 43%. Variation of coronary anatomy is common [36] and becomes problematic when a fenestrated endograft is in scope [37]. When results are stratified by age, differences exist with a tendency to have larger aortic diameters with older patients [11, 13]. Difference may also exist between both sexes. Thus a personalized approach is required for every endograft procedure for ascending aorta lesions.

Our aim was to obtain an insight into normative values for the ascending aorta. The limitation of our study is the relatively low number of the studied population. In order to establish more solid normative tables, a larger population is needed. Besides we do not stratify our results by height and weight. Such normalization will be useful particularly in the pediatric population. Nevertheless, this study is a first step in establishing normative values as well as potential preoperative aortic characteristics in the perspective of future TCT of the ascending aorta. We think that ECGgated computed tomography is the most suitable and convenient approach for such a purpose.

#### Conclusion

Large inter-individual variations exist in measurements of the ascending aorta but with limited intra-individual variations during the cardiac cycle. Therefore a personalized design of transcatheter aortic devices for each patient might become necessary. ECG-gated MDCT could have a major role in the decision-making process and the treatment planning.

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