



# Fully automated guideline-compliant diameter measurements of the thoracic aorta on ECG-gated CT angiography using deep learning

Maurice Pradella<sup>1^</sup>, Thomas Weikert<sup>1^</sup>, Jonathan I. Sperl<sup>2^</sup>, Rainer Kärgel<sup>2^</sup>, Joshy Cyriac<sup>1^</sup>, Rita Achermann<sup>1</sup>, Alexander W. Sauter<sup>1^</sup>, Jens Bremerich<sup>1^</sup>, Bram Stieltjes<sup>1^</sup>, Philipp Brantner<sup>1,3^</sup>, Gregor Sommer<sup>1^</sup>

<sup>1</sup>Department of Radiology, Clinic of Radiology & Nuclear Medicine, University Hospital Basel, University of Basel, Petersgraben 4, 4031 Basel, Switzerland; <sup>2</sup>Siemens Healthineers, Siemensstraße 3, 91301 Forchheim, Germany; <sup>3</sup>Regional Hospitals Rheinfelden and Laufenburg, Riburgerstrasse 12, 4310 Rheinfelden, Switzerland

*Contributions:* (I) Conception and design: M Pradella, T Weikert, JI Sperl, AW Sauter, J Bremerich, B Stieltjes, P Brantner, G Sommer; (II) Administrative support: M Pradella, J Bremerich, B Stieltjes, P Brantner, G Sommer; (III) Provision of study material or patients: M Pradella, J Cyriac, JI Sperl, R Kärgel, J Bremerich, P Brantner, G Sommer; (IV) Collection and assembly of data: M Pradella, T Weikert, AW Sauter, J Bremerich, B Stieltjes, P Brantner, G Sommer; (V) Data analysis and interpretation: M Pradella, T Weikert, J Cyriac, R Achermann, AW Sauter, J Bremerich, B Stieltjes, P Brantner, G Sommer; (VI) Manuscript writing: All authors; (VII) Final approval of manuscript: All authors.

*Correspondence to:* Maurice Pradella, MD. Department of Radiology, Clinic of Radiology & Nuclear Medicine, University Hospital Basel, University of Basel, Petersgraben 4, 4031 Basel, Switzerland. Email: maurice.pradella@usb.ch.

**Background:** Manually performed diameter measurements on ECG-gated CT-angiography (CTA) represent the gold standard for diagnosis of thoracic aortic dilatation. However, they are time-consuming and show high inter-reader variability. Therefore, we aimed to evaluate the accuracy of measurements of a deep learning-(DL)-algorithm in comparison to those of radiologists and evaluated measurement times (MT).

**Methods:** We retrospectively analyzed 405 ECG-gated CTA exams of 371 consecutive patients with suspected aortic dilatation between May 2010 and June 2019. The DL-algorithm prototype detected aortic landmarks (deep reinforcement learning) and segmented the lumen of the thoracic aorta (multi-layer convolutional neural network). It performed measurements according to AHA-guidelines and created visual outputs. Manual measurements were performed by radiologists using centerline technique. Human performance variability (HPV), MT and DL-performance were analyzed in a research setting using a linear mixed model based on 21 randomly selected, repeatedly measured cases. DL-algorithm results were then evaluated in a clinical setting using matched differences. If the differences were within 5 mm for all locations, the cases was regarded as coherent; if there was a discrepancy >5 mm at least at one location (incl. missing values), the case was completely reviewed.

**Results:** HPV ranged up to  $\pm 3.4$  mm in repeated measurements under research conditions. In the clinical setting, 2,778/3,192 (87.0%) of DL-algorithm's measurements were coherent. Mean differences of paired measurements between DL-algorithm and radiologists at aortic sinus and ascending aorta were  $-0.45 \pm 5.52$  and  $-0.02 \pm 3.36$  mm. Detailed analysis revealed that measurements at the aortic root were over-/underestimated due to a tilted measurement plane. In total, calculated time saved by DL-algorithm was 3:10 minutes/case.

<sup>^</sup> ORCID: Maurice Pradella, 0000-0003-2449-7835; Thomas Weikert, 0000-0001-9274-053X; Jonathan I. Sperl, 0000-0003-4528-1507; Rainer Kärgel, 0000-0003-2024-5390; Joshy Cyriac, 0000-0002-4584-0623; Alexander W. Sauter, 0000-0002-6707-2258; Jens Bremerich, 0000-0002-1002-8483; Bram Stieltjes, 0000-0002-5961-802X; Philipp Brantner, 0000-0003-3996-3966; Gregor Sommer, 0000-0002-8952-0808.

**Conclusions:** The DL-algorithm provided coherent results to radiologists at almost 90% of measurement locations, while the majority of discrepant cases were located at the aortic root. In summary, the DL-algorithm assisted radiologists in performing AHA-compliant measurements by saving 50% of time per case.

**Keywords:** Deep learning; aortic aneurysm; computed tomography angiography; dimensional measurement accuracy; observer variation; time management

Submitted Feb 09, 2021. Accepted for publication May 27, 2021.

doi: 10.21037/qims-21-142

View this article at: <https://dx.doi.org/10.21037/qims-21-142>

## 1 Introduction

2 Thoracic aortic dilatation occurs with an incidence of  
3 approximately 6–16 cases per 100,000 people/year and  
4 there is an increasing prevalence and incidence of dilatation  
5 of the thoracic aorta (1-4). Regardless of the cause of  
6 dilatation, the risk of aortic dissection or rupture rises  
7 with increasing diameters (5). This leads to high mortality  
8 rates. For example, in the USA, aneurysms of the thoracic  
9 and abdominal aorta are the 14th leading cause of death in  
10 people older than 55 years (6). Main factors that cause an  
11 increase in aortic diameter are patient age, genetic disorders  
12 (such as Marfan syndrome), as well as valve pathologies such  
13 as a bicuspid aortic valve (7,8).

14 Imaging is the sole option to detect aortic dilatation,  
15 being typically an asymptomatic disease, and only cross-  
16 sectional imaging can depict the entire aortic arch; opposite  
17 to echocardiography which can only be used to visualize  
18 the aortic root. Current guidelines recommend ECG-  
19 gated CT angiography (CTA) which is considered superior  
20 to other imaging modalities (9). However, measurements  
21 differ frequently. It is well known that transverse diameter  
22 measurements are inaccurate and considered obsolete (10).  
23 Centerline-based measurements have become best practice  
24 and were established about 15 years ago (11). However, the  
25 process of evaluating the dimensions of the thoracic aorta  
26 by measurements perpendicular to the vessel centerline is  
27 still time-consuming with 5–6 minutes per case (12,13).  
28 Currently, centerline fitting is performed automatically,  
29 but measurement locations have to be chosen manually.  
30 Due to incorrectly placed centerlines or failed automatic  
31 fitting, there is often the necessity for manual adjustments/  
32 interaction (13). This increases measuring times further  
33 and is a source of variability which ranges up to 5 mm even  
34 among expert readers in a research setting (14,15).

35 There are limited studies describing tools for automatic  
36 aortic segmentation/measurements that, for example, detect  
37

abdominal aortic aneurysms, measure the descending aortic  
diameter prior to stent graft planning, segment and measure  
aortic diameters in native scans of the thoracic aorta in  
CT scans, or help to improve reading follow-up CT scans  
according to guidelines (16-19).

In this work, we analyzed the performance of a novel  
DL-algorithm that automatically detects the thoracic aorta,  
places the centerline, identifies measurement locations and  
performs measurements according to current American  
Heart Association (AHA) guidelines.

The accuracy of the DL-algorithm was analyzed in a  
patient cohort with suspected aortic dilatation. In a research  
setting, we first compared its measurements to radiologists'  
measurements who used the established semi-automatic  
procedure in order to evaluate inter- and intra-reader  
variability and the expected savings in terms of measurement  
time (MT). This was followed by an evaluation of the whole  
cohort in a clinical setting.

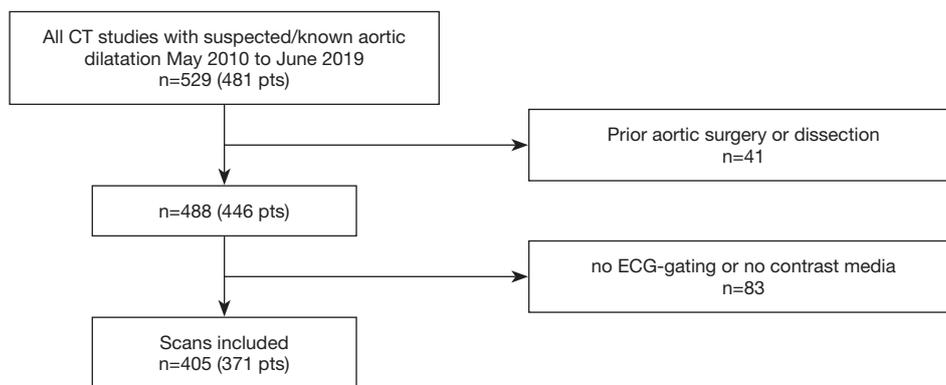
## Methods

### Ethics

The study was conducted in accordance with the  
Declaration of Helsinki (as revised in 2013). All data was  
encoded prior to any analysis to preserve patient anonymity.  
The Ethics Committee for Northwest and Central  
Switzerland approved this study (ID: 2019-01053) and  
individual consent for this retrospective analysis was waived.

### Study population

A total of 371 consecutive patients who underwent ECG-  
gated CTA at our institution between May 2010 and June  
2019 and whose radiologic reports included standardized  
diameter measurements were identified and included in this  
study (Figure 1). Those patients either were suspected to



**Figure 1** Flow chart of the dataset A. All scans with suspected or known dilatation and reported, standardized measurements were included. Pts, patients; DL, deep learning.

**Table 1** Baseline characteristics

Baseline data	Dataset A	Dataset B (inter-rater subset)	Dataset C (follow-up subset)
Number of patients	371	21	32
Age (years)	65.2±11.7	67.4±14.0	61.1±11.1
Female sex	98 (24.6%)	5 (23.8%)	5 (15.6%)
Number of CT scans	405	21	34

Baseline characteristics and study populations for both datasets (Dataset A: main cohort, Dataset B: randomly selected cases for inter-/intra-reader analysis, Dataset C: patients with more than one exam). Dataset C, age at first scan. CT, computed tomography

75 have dilatation (for example an aortic root diameter based  
76 on echocardiography of more than 40 mm) or underwent  
77 CT exams in the context of known dilatation. Exclusion  
78 criteria were aortic pathologies other than dilatation (acute  
79 or chronic dissection, rupture or intramural hematoma)  
80 or prior surgery of the thoracic aorta; the DL-algorithm  
81 was not built to evaluate those conditions. Baseline data  
82 of patients and overview of processed cases are shown  
83 in *Table 1*. While the full dataset (dataset A) was used to  
84 evaluate the overall diagnostic performance of the DL-  
85 algorithm, a subset of 21 CT studies (dataset B, inter-rater  
86 subset) was randomly selected to perform an analysis under  
87 research conditions. Thereby, the inter- and intra-observer  
88 variability associated with the common established, semi-  
89 automatic workflow was analyzed. Another subset (dataset C,  
90 follow-up subset) was created to evaluate the subcohort of  
91 patients who underwent more than one exam.

92

93

### CT scan

94

95

All scans were performed on one of four CT scanners

(Somatom Sensation 64, AS+, Edge or Definition Flash, 96  
Siemens Healthineers, Forchheim, Germany). Each exam 97  
was prospectively ECG-gated to minimize motion artifacts 98  
by cardiac movement. Image acquisition was performed 99  
during diastole, either as low pitch spiral acquisition with 100  
dose modulation over multiple heart beats (Sensation 64, 101  
AS+ or Edge scanners) or during one heart beat (Definition 102  
Flash scanner). 103

Bolus tracking was performed in the ascending aorta; 104  
trigger was  $\geq 100$  HU with a 10 s delay. 70–100 mL of 105  
contrast agent for thoracic scans were administered with a 106  
flow rate of 3–4 mL/s. No pharmacologic agent was used 107  
for heart rate control in any scan. We generally used the 108  
thinnest soft tissue kernel available (1.0 mm slice thickness, 109  
increment 0.6 mm, resolution 512×512 pixels). 110

111

### Measurement tools

112

#### Established semi-automatic workflow

113  
114  
Measurements were performed perpendicular to blood flow 115  
axis using the centerline technique in the postprocessing 116

117 software Syngo.via (Siemens Healthineers, Forchheim,  
 118 Germany) (10,14). The aortic centerline was automatically  
 119 detected; radiologists could adjust the centerline in case  
 120 it was not fitted well. If the automatic centerline wasn't  
 121 available, radiologists placed it manually. Measuring points  
 122 were used according to current AHA-guidelines (9): aortic  
 123 sinus (AS), sinotubular junction (STJ), ascending aorta (AA),  
 124 proximal aortic arch (PA), mid aortic arch (MA), and distal  
 125 aortic arch (DA) (9).

### 127 Fully-automatic DL-based workflow

128 DL-algorithm measurements were performed by an in-  
 129 house deployed prototype software (Chest AI, version  
 130 0.2.9.2, Siemens Healthineers, Forchheim, Germany). Its  
 131 development was completely independent from this study.  
 132 The thinnest soft tissue kernel series per case was sent to  
 133 the dedicated workstation which processed the cases one by  
 134 one. No further human input was necessary.

135 The DL-algorithm fully and automatically performed  
 136 three consecutive steps: detection of aortic landmarks,  
 137 segmentation of the lumen, and diameter measurements  
 138 (incl. detection of measurement locations).

139 First, landmark detection based on Deep Reinforcement  
 140 Learning was performed to detect six landmarks along  
 141 the thoracic aorta: aortic root, aortic arch center,  
 142 brachiocephalic artery bifurcation, left common carotid  
 143 artery, left subclavian artery, celiac trunk. The principles  
 144 of the underlying algorithm have been described by Ghesu  
 145 *et al.* (20). The algorithm has been trained on more than  
 146 10,000 data sets (CT data plus manual labelling of the six  
 147 landmarks).

148 The aortic root landmark was used to define a Region  
 149 of Interest (ROI) for the segmentation algorithm. The  
 150 segmentation was performed using an adversarial deep  
 151 image-to-image network (DI2IN), which is a multi-layer  
 152 convolutional neural network (CNN) taking the CT  
 153 data (cropped to the ROI) as an input and providing a  
 154 segmentation mask as an output. The technical approach  
 155 of network topology and training strategy has first been  
 156 developed in the context of liver segmentation and is  
 157 easily adapted to other organs like the aorta by providing  
 158 corresponding CT data and annotations (manually  
 159 segmented aorta masks) (21). Training was performed on  
 160 more than 1,000 CT data sets covering both native and  
 161 contrast-enhanced data with and without ECG-gating; these  
 162 data sets were completely independent from this study.

163 Given the segmented aorta mask, a centerline model  
 164 was used to generate the aortic centerline. The centerline

was used in combination with the pre-computed aortic  
 landmarks to identify the measurement planes at multiple  
 locations according to the AHA guidelines (*Figure 2*) (9).

In each of the planes, multiple diameters were measured  
 by computing intersections of rays starting from the  
 centerline with the aortic mask. Based on these diameters,  
 the maximum in-plane diameter was reported. Visual output  
 series were created in axial and sagittal orientation as well as  
 a 3D volume rendering.

### 175 *Image reading and data evaluation*

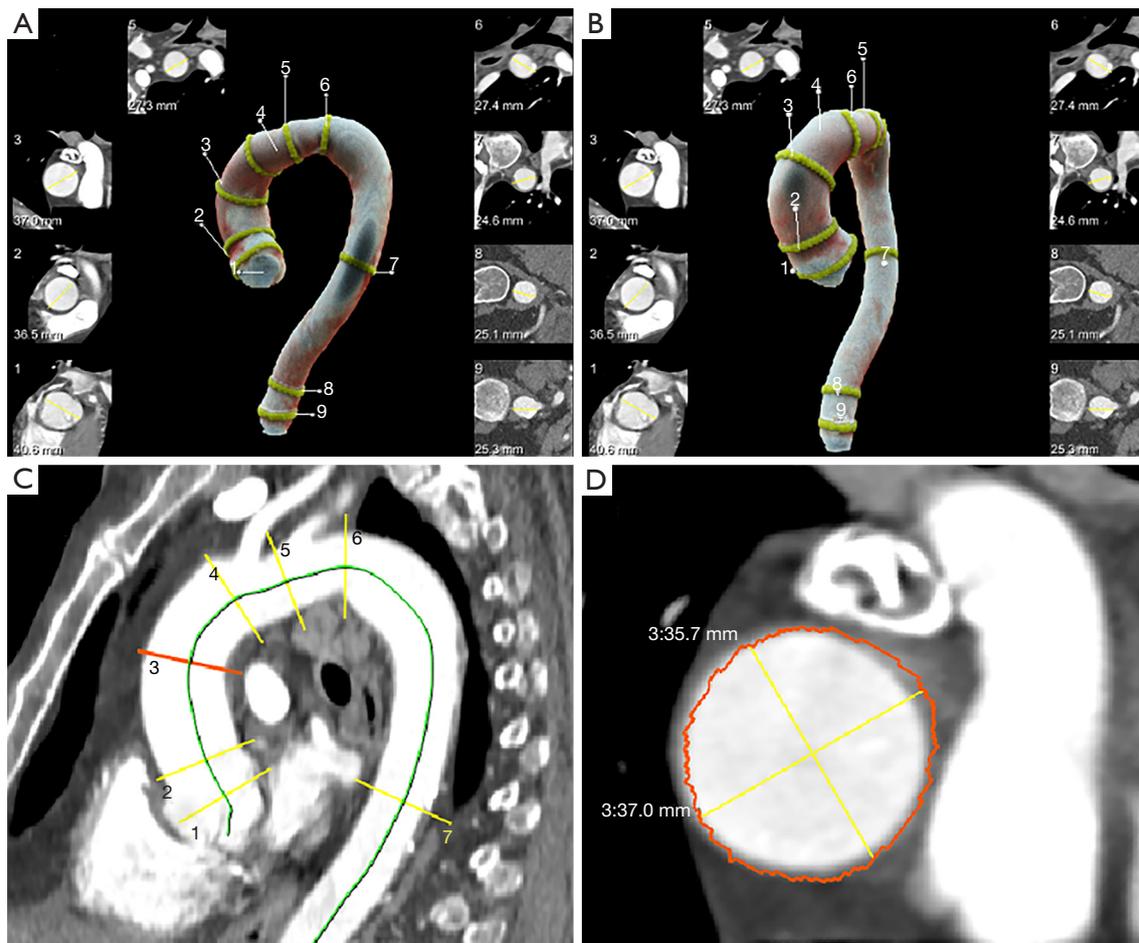
#### 176 **Research setting, inter-rater variability of semi- 177 automatic workflow compared to DL-algorithm and 178 measurement times**

179 Three readers, R1, R2, and R3 with 2, 4.5 and 8 years of  
 180 experience, respectively, performed measurements in dataset  
 181 B (inter-rater subset) twice with a blanking period of at least  
 182 7 days. Readers were blinded to reports. Reader R2 and R3  
 183 were both fellowship-trained in cardiovascular radiology.  
 184 The reading was performed in a calm environment, no  
 185 telephone or clinical duties were present to establish  
 186 optimal conditions for measurements. Each reader noted  
 187 MT of each case after it was loaded in Syngo.via software  
 188 up until all locations were measured.

189 Intraclass correlations (ICC) were calculated for both  
 190 intra- and inter-reader agreement evaluation for each  
 191 measurement location (22). To compare the the results of  
 192 the DL-algorithm with human performance and variability,  
 193 we set up a linear mixed model based on the human  
 194 measurements with reader and case as random effects and  
 195 location as a fixed effect (23). Then, a predicted value  
 196 (gold standard) and its 95% prediction interval for each  
 197 location and case was estimated in order to evaluate human  
 198 performance variability (HPV). To obtain robust prediction  
 199 intervals we applied bootstrap methods taking into account  
 200 the hierarchical structure of the data (R:library fabricatr).  
 201 Finally, the proportion of DL-measurements outside the  
 202 prediction interval was used to test (chi-squared) whether  
 203 the proportion of outliers is compatible with the expected  
 204 number of 5% (gold standard).

#### 206 **Clinical setting, performance evaluation of the DL- 207 algorithm**

208 All 405 scans included in dataset A underwent fully  
 209 automatic processing by the DL-algorithm. Each case was  
 210 processed twice to evaluate technical feasibility; to verify  
 211 reproducibility, those measurements were compared with  
 212



**Figure 2** Visual output of DL-measurements in a case with normal diameters and within human variance. (A) VRT with lateral view on thoracic aorta with thumbnails of measurements at each location; (B) VRT with anterior view on thoracic aorta; (C) Non-linear projection of the aortic centerline into a 2D plane on thoracic aorta with measurement plane at each location, measurement plane of ascending aorta highlighted in orange; (D) Measurement of AA on cross-sectional images orthogonal to the aortic centerline based on (C). AA, ascending aorta; AHA, American Heart Association; DL, deep learning; VRT, volume rendering technique.

213 each other. MT required for processing were automatically  
214 noted by the DL-algorithm.

215 The results provided by the DL-algorithm were then  
216 compared to the original diameter measurements that were  
217 retrospectively extracted from the written reports using a  
218 Python-based script. These original measurements were  
219 initially performed by residents, afterwards they were  
220 discussed with a senior, board-certified radiologist who was  
221 free to overrule measurements and who finalized the report.  
222 Mid descending aorta (MDA) and distal descending aorta  
223 (DDA) measurements were not included in the original  
224 reports as a trade-off to optimize clinical efficiency since  
225 most dilatations are found at the aortic root or ascending

aorta.

226  
227 Cases with a difference of  $>5$  mm for at least one  
228 measurement location between the two methods were  
229 regarded as discrepant (cutoff-value based on Quint  
230 *et al.* (15), this includes missing measurements). In these  
231 cases, visual outputs were used for a full review by a  
232 fellowship-trained, cardiovascular radiologist with 4.5 years  
233 of experience (R2). In addition, an analysis of classification  
234 change (dilatation versus no dilatation) between DL-  
235 measurements and original reports was performed. We  
236 defined relevant dilatation of the aorta as  $\geq 45$  mm at AS,  
237 STJ, and AA and  $\geq 40$  mm at all other locations in order to  
238 evaluate misclassification (based on current literature which

**Table 2** Technical success rates

Technical success rates	Dataset A (371 pts/405 cases)
Number of cases processed by the DL-algorithm	399/405 (98.5%)
Number of processed cases by the DL-algorithm with all measurements available	341/399 (85.5%)
Aortic sinus	399/399 (100%)
Sinotubular junction	399/399 (100%)
Ascending aorta	396/399 (99.2%)
Proximal arch	396/399 (99.2%)
Mid arch	394/399 (98.7%)
Distal arch	341/399 (85.5%)
Mid descending aorta	341/399 (85.5%)
Distal descending aorta	341/399 (85.5%)
Total technical success rate of DL-algorithm for all locations	3,007/3,192 (94.2%)

Technical success rates of the DL-algorithm divided by location. DL, deep learning.

239 also represents the standard at our institution) (2,24). For  
 240 review analysis, AS and STJ were grouped as aortic root,  
 241 PA, MA, and DA as aortic arch, and MDA and DDA as  
 242 descending aorta.

243 Dataset C which included all patients with more than  
 244 one exam was analyzed in regards if there was a difference  
 245 of >5 mm of diameters between two scans for the DL-  
 246 algorithm or the reports. The results can be found in the  
 247 supplements.

248 Data organization was performed with Excel (Microsoft  
 249 Corporation, Redmont, USA) and Python (Python  
 250 Software Foundation, Wilmington, USA). R (R Foundation  
 251 for Statistical Computing, Vienna, Austria) and SPSS (IBM,  
 252 Armonk, USA) were used for statistical analysis. We plotted  
 253 the data in scatterplots and calculated Pearson correlation  
 254 coefficients (PCC) for each location. In addition, Bland-  
 255 Altman plots were created to compare reported with DL-  
 256 measurements. To compare absolute diameters, Mann-  
 257 Whitney-U-Test was used. A P value <0.05 was defined to  
 258 indicate statistical significance. We also calculated mean  
 259 diameter measurements and standard deviations by the DL-  
 260 algorithm for each location, sorted by sex and age group  
 261 (Table S1).

262

263

## Results

264

265

### Success rates of automatic processing

266

Fully-automated diameter measurements by the DL-

algorithm were technically successful in 399 of 405 cases 268  
 (98.5%) and at 3007/3192 locations (94.2%, Table 2). 269  
 Measurements from AS until mid arch were available in 270  
 98.7%, complete measurements at all locations in 85.5% of 271  
 all cases. The algorithm's technical failure rate was highest 272  
 in the descending aorta and distal aortic arch (14.5%). 273

274

275

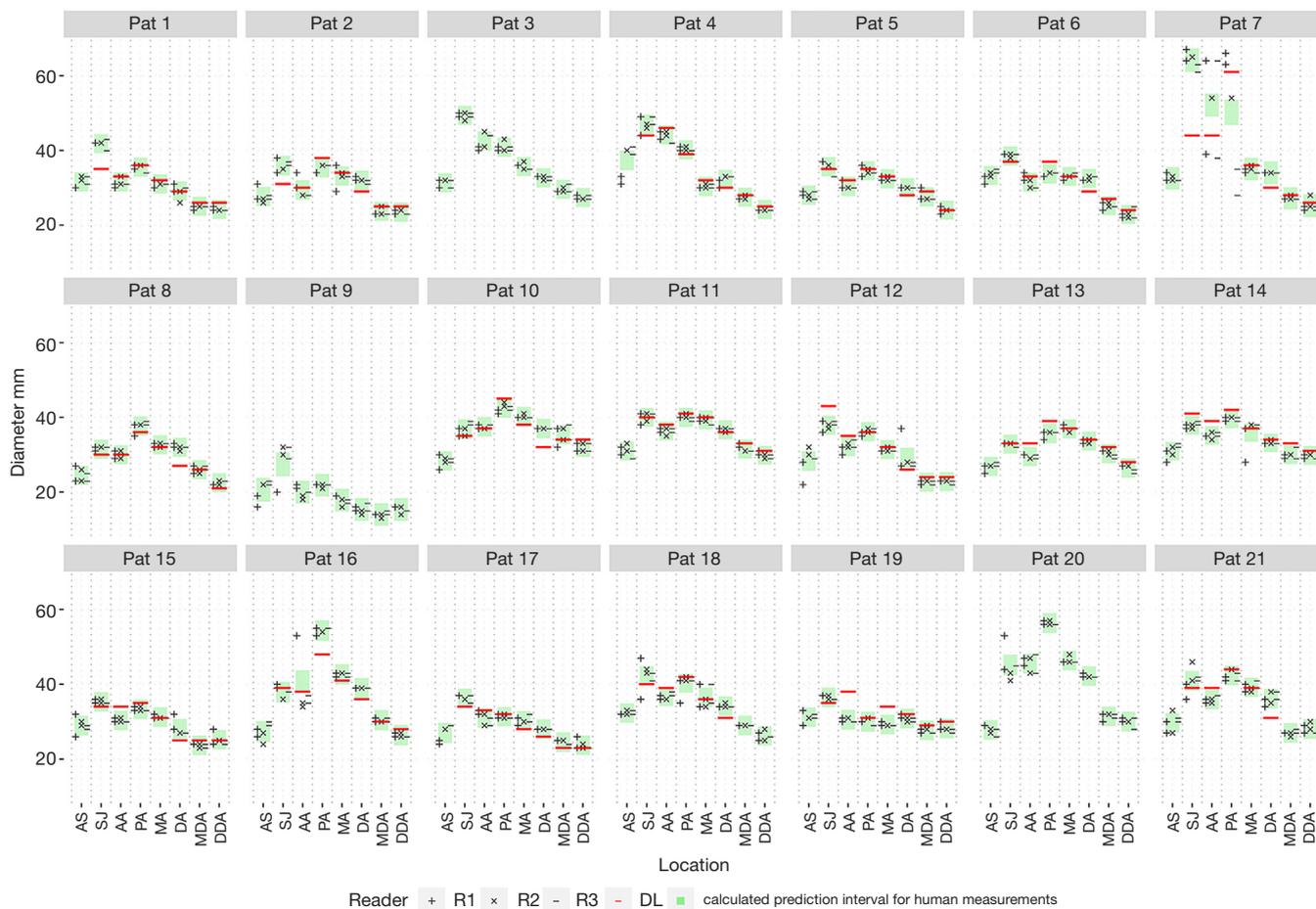
### Inter- & intra-reader comparison in research setting

The randomly selected dataset B (inter-rater subset) used 276  
 for the inter- and intra-reader analyses consisted of 12 cases 277  
 with normal diameters, 8 with aortic dilatation and one with 278  
 aortic coarctation. Figure 3 shows the manual measurement 279  
 results for all 21 patients from the subset along with the 280  
 respective DL-measurements. 281  
 282

Overall, inter-rater agreement between radiologists was 283  
 excellent: Average ICC for intra-reader agreement over 284  
 all locations was 0.94 [range: 0.81 (STJ) – 0.98 (AA)]. The 285  
 largest variation was observed for STJ measurements of R1 286  
 and R3 (ICC: 0.76 and 0.70, respectively) indicating only 287  
 moderate/good agreement. Average ICC for inter-reader 288  
 agreement over all locations was 0.94 [range: 0.85 (STJ) 289  
 – 0.98 (AA)]. An overview of all ICC can be found in the 290  
 supplements (Table S2). 291

The prediction interval for HPV that was calculated based 292  
 on the repeated measurements by the three readers varied for 293  
 each location with a median width of  $\pm 2.6$  mm, a maximum 294  
 width of  $\pm 3.4$  mm, and a minimum width of  $\pm 2.5$  mm. 295

The DL-algorithm measurements were statistically more 296



**Figure 3** Inter-reader comparison for all 21 patients. Each plot represents all measurements for one patient for the locations from AS to DDA on the x-axis. For each location, the measurements by the three readers (symbols “+”, “x” and “-”), the DL-algorithm (red line) and the predicted gold standard interval (green box) whose calculation was based on measurements by readers 1–3 (R1–3) are shown. Note: Since there was no variance in DL-measurements (which represents 100% preciseness), the red line for DL-measurements represents the two measurements performed per location. AS, aortic sinus; SJ, sinotubular junction; AA, ascending aorta; PA, proximal arch; MA, Mid arch; DA, distal arch; MDA, mid descending aorta; DDA, distal descending aorta; DL, deep learning; Pat, patient number.

297 often outside the 95% prediction interval compared to the  
 298 expected percentage of 5% as shown by the chi-squared test  
 299 (22.5%,  $P < 0.0001$ , 95% CI: 15.99–30.51).

300  
 301 **Evaluation of DL-algorithm measurement and**  
 302 **classification accuracy in clinical setting**

303 The accuracy analysis of dataset A showed that the  
 304 automated measurements in 2,540/3,192 locations (79.6%,  
 305 Table 3) differed from the human measurements by less than  
 306 a 5 mm interval and therefore, were counted as coherent.  
 307 145/399 cases (36.3%) showed a difference of >5 mm for  
 308

at least one location (this also includes cases with missing  
 measurements). After a detailed review of all measurements  
 for these 145 cases, 2,778/3,192 (87.0%) measurements were  
 identified as coherent. Aside from the aortic root (537/798,  
 67.3%), the ascending aorta (364/399, 91.2%), aortic arch  
 (1,123/1,197, 94.2%), and descending aorta (754/798,  
 94.5%) showed high rates of coherent measurements. In the  
 majority of reviewed cases, the estimation error found at the  
 aortic root was due to a tilted measurement plane (Figure 4).  
 Of all reviewed cases, classification of dilatation remained  
 unchanged in 76/145 cases (52.4%) while a change of  
 classification occurred in 69/145 cases (47.6%, Table 3).

**Table 3** Measurement and classification accuracy by location

Dataset A (399 cases/3,192 locations)	All locations	Root	Ascending aorta	Arch	Descending aorta
Cases with initially correct estimation (within 5 mm interval for all measurements)		254/399 (63.7%)			NA <sup>§</sup>
Reviewed cases <sup>†</sup>		145/399 (36.3%)			
Reviewed locations <sup>†</sup>		1,160			
Correct measurement	748/1,160 (64.5%)	24/290 (8.3%)	110/145 (75.9%)	361/435 (83.0%)	246/290 (84.8%)
Wrong measurement	377/1,160 (32.5%)	266/290 (90.3%)	32/145 (22.1%)	69/435 (15.9%)	14/290 (4.8%)
Missing measurement	35/1,160 (3.0%)	0/290	3/145 (2.0%)	5/435 (1.1%)	30/290 (10.3%)
No change of classification	76/145 (52.4%)	96/145 (66.2%)	134/145 (92.4%)	130/145 (89.7%)	NA
Change of classification	69/145 (47.6%)	49/145 (33.8%)	11/145 (7.6%)	15/145 (10.3%)	NA
Aneurysms misclassified by DL-algorithm in reviewed cases	34/145 <sup>‡</sup> (23.4%)	18/145 (12.4%)	11/145 (7.6%)	11/145 (7.6%)	NA
Finally correct measurements (incl. reviewed locations)	2,778/3,192 (87.0%)	537/798 (67.3%)	364/399 (91.2%)	1,123/1,197 (93.8%)	754/798 (94.5%)
Aneurysms misclassified by DL-algorithm (all cases)	34/399 <sup>‡</sup> (8.5%)	18/399 (4.5%)	11/399 (2.8%)	11/399 (2.8%)	NA

Measurement and classification accuracy by location. <sup>†</sup>, cases with a difference >5 mm between DL and original measurements underwent detailed review of all measurements. <sup>‡</sup>, in 6 cases the misclassified aneurysm extended to multiple locations, those cases were only counted once. <sup>§</sup>, not available in original reports. DL, deep learning; NA, not available.

321 An aneurysm was misclassified in 34/399 cases (8.5%).  
 322 An overview of measured diameters sorted by sex and age  
 323 groups can be found in the supplement (Table S1).

324 Mean differences of matched measurements by the  
 325 algorithm and the report were -0.45 mm at AS and -0.02 mm  
 326 at AA. Bland-Altman analysis revealed wider limits of  
 327 agreement ( $\pm 1.96$  SD) at AS than at AA (AS: +10.37 and  
 328 -11.28 mm, AA: +6.56 and -6.61 mm). PCC were 0.676  
 329 for AS (moderate correlation) and 0.906 for AA (high  
 330 correlation) (Figure 5). Mean differences of matched  
 331 measurements at STJ, PA, MA and DA were +3.25 mm,  
 332 +0.32 mm, -1.21 mm, and +1.53 mm, respectively. The  
 333 Bland-Altman and scatterplots for these locations can be  
 334 found in the supplement (Figures S1-S4).

335 DL-algorithm measurements were performed twice per  
 336 case, the measured diameters were the exact same in every  
 337 case and at every location, meaning perfect reproducibility  
 338 (exact to eight decimal places).

339

#### 340 *Measurement times*

341 In our research setting, the mean MT for the three human  
 342

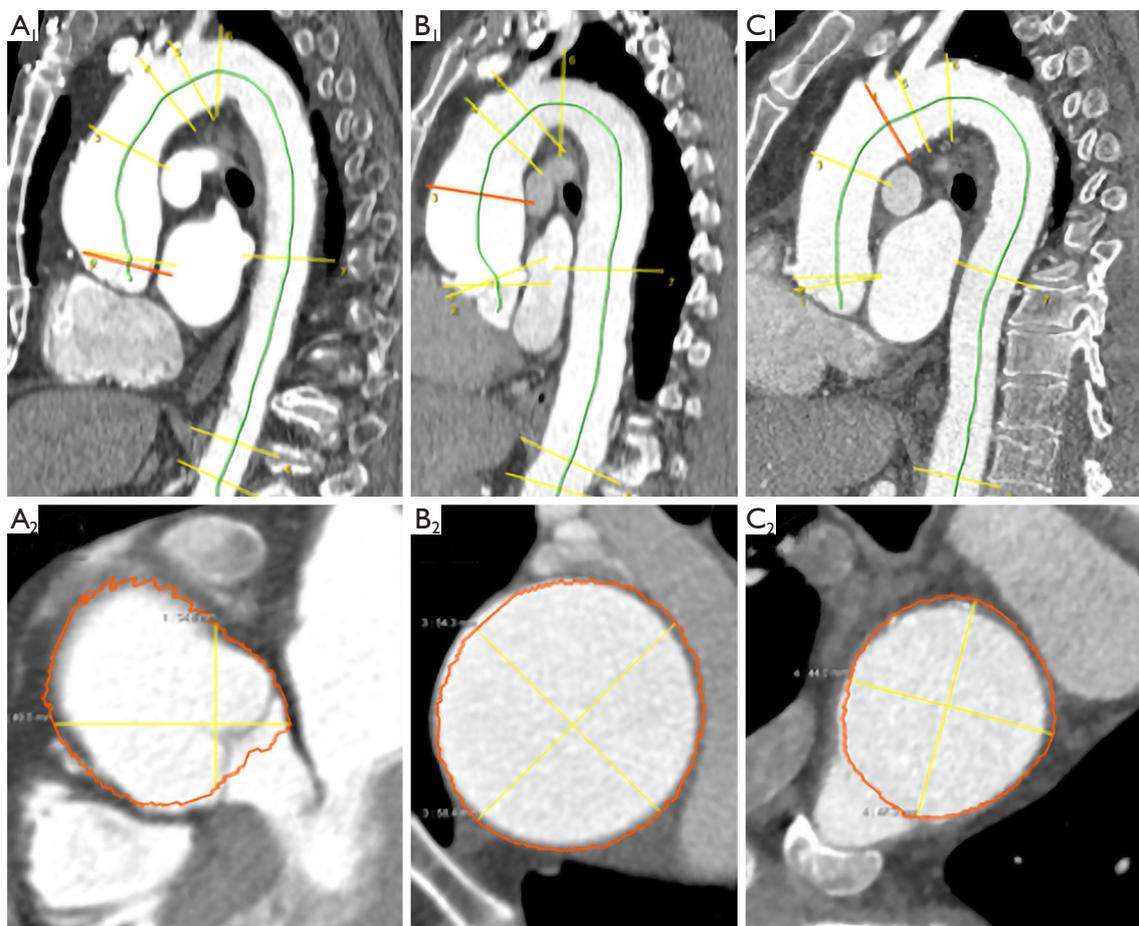
readers was 4:48 $\pm$ 1:55 min per case (range 2:00–13:00 min, 343  
 Table S2). Significant differences were seen when 344  
 comparing the less experienced reader (R1) with the two 345  
 more experienced, fellowship-trained readers (R1–R2: 346  
 P<0.001, R1–R3: P=0.001). 347

The DL-algorithm performed measurements 348  
 autonomously in 2:19 $\pm$ 0:22 min (incl. generation of visual 349  
 outputs), which was significantly faster compared to human 350  
 reader (P<0.001). 351

The calculated average time to analyze a case with 352  
 support of the DL-algorithm would be 1:38 min. We 353  
 accounted for a failure rate of 13% (success rate: 87.0%, 354  
 Table 3), a human MT for one location would be 36 seconds 355  
 (4:48 min total for 8 locations) plus one minute to check 356  
 visual outputs. This would result in an average of 3:10 min 357  
 saved for measurements per case. 358

#### 360 **Discussion**

In this study, we evaluated the accuracy of a DL-algorithm 361  
 to perform thoracic aorta diameter measurements according 362  
 to AHA-guidelines in more than 350 patients and further 363  
 364

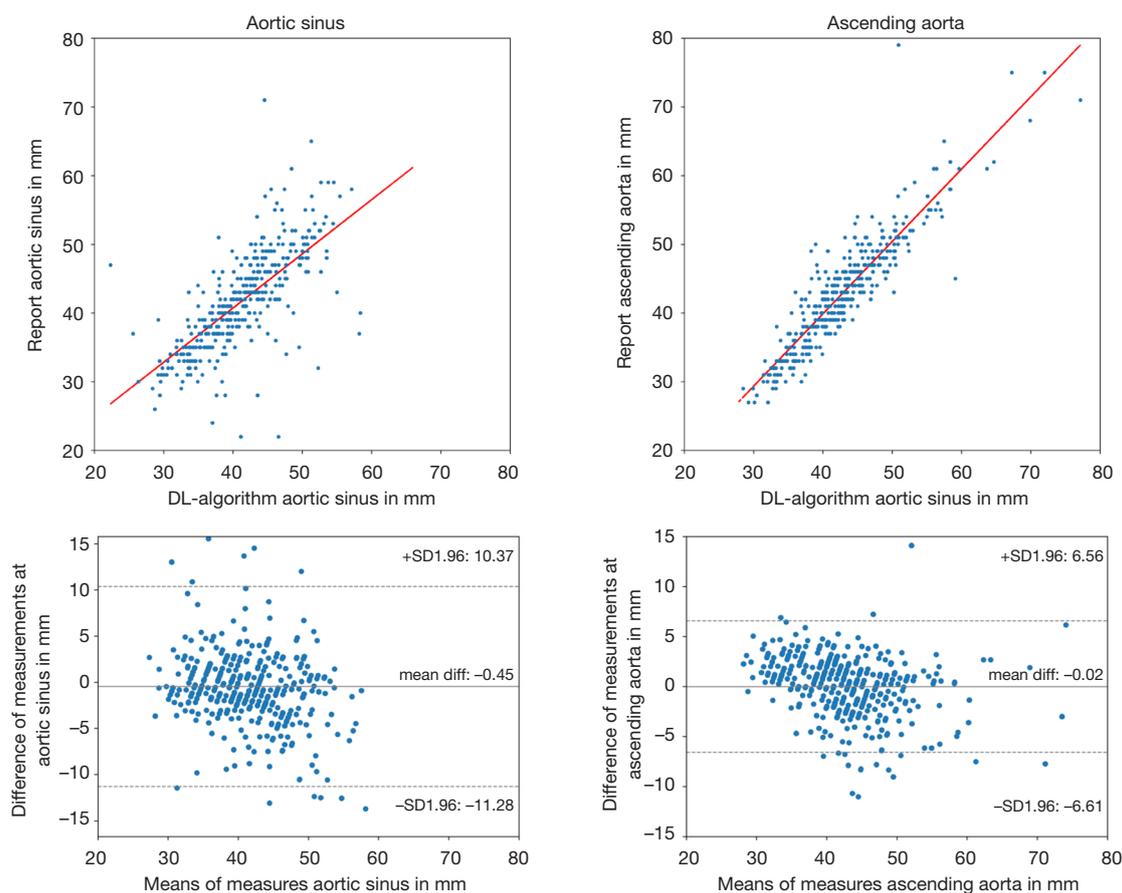


**Figure 4** Example cases. (A1) tilted measurement plane causing overestimation. AS plane (orange) showed tilt. (A2) the actual measured plane was oblique coronal, left ventricular outflow tract, aortic valve with two of three leaflets and aortic sinus were visible. (B1) correct angle of AA measurement plane. Note that AS and STJ planes are tilted. (B2) acceptable measurement of an AA aneurysm with a diameter of 5.8cm. (C1) correct angle of PA measurement plane. (C2) correct measurement of PA aneurysm with a diameter of 4.7 cm. AS, aortic sinus; AA, ascending aorta; STJ, sinotubular junction; PA, proximal arch.

365 compared them to radiologists' measurements. We  
 366 showed four major results: (I) there was a mean variance  
 367 of up to  $\pm 3.4$  mm for radiologists in a research setting  
 368 (this constitutes perfect measurement conditions) using  
 369 the established semi-automatic workflow. This observed  
 370 variance is in agreement or slightly lower than previously  
 371 described variability of up to 5 mm difference between  
 372 expert readers (15). Opposite to that, the DL-algorithm was  
 373 highly precise but less accurate in repeated measurements.  
 374 (II) We showed that time required by human readers for  
 375 guideline-compliant measurements under these perfect  
 376 conditions was about five minutes per case. (III) At 87.0%  
 377 of all measurement locations in our clinical cohort, the  
 378 DL-algorithm provided measurement results within the

379 expected margin of variance and therefore, were coherent  
 380 to results of human readers. This finding resulted in an  
 381 expected time-saving of more than 3 min per case for the  
 382 radiologist. (IV) In review of discrepant cases, errors by the  
 383 DL-algorithm were found predominantly at the aortic root  
 384 (in 139/145 cases); these cases could be easily identified  
 385 by DL-algorithms' visual outputs and therefore reduce  
 386 interaction/re-measurements to a minimum.

387 In general, there are obstacles in how to measure  
 388 diameters of tubular structures on CT data. Centerline  
 389 based measurements are today's gold standard and have  
 390 superiority over the double oblique technique based on  
 391 multiplanar reformations as previously demonstrated (14,15).  
 392 Nevertheless, the accuracy of measurements is debatable.



**Figure 5** Scatter and Bland-Altman plots of measurements (in mm) at AS and AA. AS, aortic sinus; AA, ascending aorta; STJ, sinotubular junction; DL, deep learning; Diff, difference; SD, standard deviation.

393 Eleftheriades *et al.* commented that “1–2 mm is not enough to  
 394 detect change” and “you cannot have confidence in measured  
 395 change of 3 or 4 mm” (25). These statements match our  
 396 findings: as we investigated inter-observer variability between  
 397 radiologists in the research setting, ICC analysis showed  
 398 excellent agreement overall, but the agreement at STJ was  
 399 only moderate to good between observers. Our prediction  
 400 model interval of mean values which gives an estimate of  
 401 expected variability ranged between  $\pm 2.5$ –3.4 mm depending  
 402 on case and location. The range is likely to be even  
 403 wider in a clinical setting as the current assessments were  
 404 performed under perfect conditions (quiet environment, no  
 405 telephone calls). In total, DL-measurements were outside  
 406 the prediction interval but this was mainly caused by a few  
 407 outliers. In daily clinical practice, multiple factors influence  
 408 aortic diameter measurements, for example: CT scan  
 409 technique, reader experience, measurement technique used,  
 410 stress, knowledge of previously reported diameter. Hence,

we agree that one must consider an impreciseness of up to 411  
 3–4 mm (25). Compared to a study from McComb which 412  
 used native CT datasets from a US lung cancer screening 413  
 trial to investigate normal aortic diameters, our cohort of 414  
 known and suspect dilatation had higher diameters for the 415  
 aortic root and ascending aorta and similar diameters for 416  
 arch and descending aorta (26). While the average patient 417  
 age is similar, our inclusion criteria consisted of known or 418  
 suspected dilatation which can explain this observation. 419  
 In general, absolute diameter of more than 55 mm is of 420  
 highest relevance since these patients usually require surgical 421  
 therapy (9,26). Therefore, in our cohort, we considered a 422  
 measurement difference of >5 mm as relevant which was also 423  
 justified by Quint *et al.* who found 5mm differences within 424  
 the 95% confidence interval between expert readers (15). 425

Opposite to human measurements, there was no 426  
 variance in repeated DL-measurements of the same case, 427  
 representing perfect preciseness. This was also found 428

429 in another study which showed that AI-support reduces  
430 variability of aortic measurement (19). Statistically, over all  
431 test cases, there was a difference between human and DL-  
432 measurements but this was likely caused by rather large  
433 differences in a few cases. These errors could easily be  
434 detected by inspection of DL-algorithms' visual outputs.  
435 In most cases and locations, the measurements by the DL-  
436 algorithm were quite similar to human measurements.  
437 The crucial question is how to weigh human variability or  
438 neutralization of variability in human measurements by  
439 a DL-algorithm versus a few inaccurate measurements,  
440 respectively. We believe that the variability in human  
441 measurements is bound to remain and appreciate DL's  
442 neutral measurements to achieve more objectivity.

443 In-depth analysis of undetermined cases (145/399)  
444 revealed that most differences were located at the aortic  
445 root (n=131, 90.8%). The aortic root might have to be  
446 re-measured in a third of cases (133/399, 33.3%) but the  
447 majority of all measurements (2,778/3,192, 87.0%) were  
448 approved via easily assessable DL-measurement outputs (14).  
449 Our sub-analysis of follow-up cases did not show a case with  
450 a true diameter increase of >5 mm at any location between  
451 two scans. DL produced two false-positive cases which  
452 were caused by a tilted measurement plane at the aortic  
453 root. In the rest of cases, DL-measurements were coherent;  
454 however, the full effect of the DL-algorithm on follow-up  
455 exams requires an analysis of a larger follow-up cohort.

456 In 34/399 cases, an aneurysm was misclassified by the  
457 algorithm which meant that the DL-measurement was  
458 below our cutoff. These situations could be easily identified  
459 by the visual outputs of the DL-algorithm. Opposite to that,  
460 in 35 cases radiologists overestimated diameters resulting in  
461 diagnosis of aneurysm. Deficits in understanding software  
462 applications or stress could cause those errors. In addition,  
463 centerline based measurements require a more complex  
464 understanding of both technique and anatomy.

465 Potential time savings are a major advantage of the DL-  
466 algorithm: our readers needed about 5 minutes per case for  
467 centerline measurements which is similar to in the literature  
468 reported centerline MT (13,19). The DL-algorithm would  
469 save more than 3 minutes per case, which could easily  
470 add up to multiple hours per week (27). It is important to  
471 mention that the algorithms' measurements do not involve  
472 human input so the radiologist can continue to assess the  
473 exam until the measurements are completed. In the majority  
474 of incoherent cases in our study, only the aortic root had to  
475 be re-measured. In a few cases, measurements of aortic arch  
476 and descending aorta were incoherent or missing which

could be explained by the ostia of the supraaortal arteries 477  
and lack of landmark identification by the DL-algorithm. 478

The DL-algorithm provided a high processing rate 479  
of cases (>97%), still 6 cases were not calculated which 480  
could either be a general software error (reproducible 481  
and non-reproducible) or an error in landmark detection. 482  
Furthermore, it provided additional information: DL- 483  
measurements of mid and distal descending aorta were 484  
available, and review showed that they were correct in 485  
84.8% of cases. Those measurements were not available 486  
in our radiologic reports because they were omitted in the 487  
standard workflow since most aneurysms are located at the 488  
aortic root and AA. 489

There are several limitations including some with special 490  
regard to the use of DL-software (28): first, this was a single 491  
center, retrospective analysis. Second, since all scans were 492  
ECG-gated, motion artifacts were minimized. Additionally, 493  
aortic replacement surgery or post-stenting were excluded. 494  
Third, imaging data from only one manufacturer was 495  
analyzed; performance on exams acquired on scanners from 496  
other vendors might vary. Fourth, the reference standard 497  
for dataset A were measurements extracted from the 498  
radiology reports and not re-measured in a research setting. 499  
Fifth, the algorithm is a measuring tool, it was not built 500  
to make diagnoses or detect pathologies like intramural 501  
hematoma or aortic dissection, which have to be evaluated 502  
by the radiologist. 503

In summary, the evaluated DL-algorithm performed 504  
fully automatic, guideline-compliant aortic measurements 505  
reliably in 87% of all measurements and performed repeated 506  
measurements of the same CT scan with zero variance. 507  
In about one third of cases, the aortic root had to be re- 508  
measured, however time savings in the order of 3 minutes 509  
per case were still observed. Thereby, it is a foundation for 510  
a tool supporting radiologists in guideline-compliant aortic 511  
measurements. 512

## Acknowledgments 513

We thank Mitchell A. Collins for proofreading this 514  
manuscript. 515

*Funding:* None. 516

## Footnote 517

*Conflicts of Interest:* All authors have completed the ICMJE 518  
uniform disclosure form (available at <https://dx.doi.org/10.21037/qims-21-142>). JS is an employee of Siemens 519  
520

525 Healthineers and received personal fees, RK is a consultant  
 526 for Siemens Healthineers. JS and RK both helped in  
 527 installation and maintenance of the software but were not  
 528 involved in study design, data analysis or interpretation.  
 529 They report that they have a patent US2020/0160527A1  
 530 pending to Siemens Healthineers. The other authors have  
 531 no conflict of interest to declare.

532

533 *Disclaimers:* Siemens Healthineers provided the prototype  
 534 DL-algorithm. Two co-authors are affiliated with Siemens  
 535 Healthineers (Jonathan. I. Sperl, employee and R. Kärger,  
 536 consultant). Siemens Healthineers had no influence on  
 537 study design and data analysis.

538

539 *Ethical Statement:* The authors are accountable for all  
 540 aspects of the work in ensuring that questions related  
 541 to the accuracy or integrity of any part of the work are  
 542 appropriately investigated and resolved. The study was  
 543 conducted in accordance with the Declaration of Helsinki (as  
 544 revised in 2013). All data was encoded prior to any analysis  
 545 to preserve patient anonymity. The Ethics Committee for  
 546 Northwest and Central Switzerland approved this study (ID:  
 547 2019-01053) and individual consent for this retrospective  
 548 analysis was waived.

549

550 *Open Access Statement:* This is an Open Access article  
 551 distributed in accordance with the Creative Commons  
 552 Attribution-NonCommercial-NoDerivs 4.0 International  
 553 License (CC BY-NC-ND 4.0), which permits the non-  
 554 commercial replication and distribution of the article with  
 555 the strict proviso that no changes or edits are made and the  
 556 original work is properly cited (including links to both the  
 557 formal publication through the relevant DOI and the license).  
 558 See: <https://creativecommons.org/licenses/by-nc-nd/4.0/>.

559

## 560 References

561 1. Booher AM, Eagle KA. Diagnosis and management  
 562 issues in thoracic aortic aneurysm. *Am Heart J*  
 563 2011;162:38-46.e1.  
 564 2. Goldfinger JZ, Halperin JL, Marin ML, Stewart AS, Eagle  
 565 KA, Fuster V. Thoracic aortic aneurysm and dissection. *J*  
 566 *Am Coll Cardiol* 2014;64:1725-39.  
 567 3. Mokashi SA, Svensson LG. Guidelines for the  
 568 management of thoracic aortic disease in 2017. *Gen*  
 569 *Thorac Cardiovasc Surg* 2019;67:59-65.  
 570 4. Olsson C, Thelin S, Stahle E, Ekblom A, Granath F.  
 571 Thoracic aortic aneurysm and dissection: increasing  
 572

prevalence and improved outcomes reported in a  
 nationwide population-based study of more than 14,000  
 cases from 1987 to 2002. *Circulation* 2006;114:2611-8.  
 575 5. Davies RR, Goldstein LJ, Coady MA, Tittle SL, Rizzo JA,  
 576 Kopf GS, Elefteriades JA. Yearly rupture or dissection rates  
 577 for thoracic aortic aneurysms: simple prediction based on  
 578 size. *Ann Thorac Surg* 2002;73:17-27; discussion -8.  
 579 6. Clouse WD, Hallett JW Jr, Schaff HV, Gayari MM,  
 580 Ilstrup DM, Melton LJ 3rd. Improved prognosis of  
 581 thoracic aortic aneurysms: a population-based study. *JAMA*  
 582 1998;280:1926-9.  
 583 7. Keane MG, Wiegers SE, Plappert T, Pochettino A, Bavaria  
 584 JE, Sutton MG. Bicuspid aortic valves are associated with  
 585 aortic dilatation out of proportion to coexistent valvular  
 586 lesions. *Circulation* 2000;102:III35-9.  
 587 8. Ikonomidis JS, Ivey CR, Wheeler JB, Akerman AW,  
 588 Rice A, Patel RK, Stroud RE, Shah AA, Hughes CG,  
 589 Ferrari G, Mukherjee R, Jones JA. Plasma biomarkers  
 590 for distinguishing etiologic subtypes of thoracic  
 591 aortic aneurysm disease. *J Thorac Cardiovasc Surg*  
 592 2013;145:1326-33.  
 593 9. Hiratzka LF, Bakris GL, Beckman JA, Bersin RM, Carr  
 594 VF, Casey DE Jr, et al. 2010 ACCF/AHA/AATS/ACR/  
 595 ASA/SCA/SCAI/SIR/STS/SVM guidelines for the  
 596 diagnosis and management of patients with Thoracic  
 597 Aortic Disease: a report of the American College of  
 598 Cardiology Foundation/American Heart Association  
 599 Task Force on Practice Guidelines, American Association  
 600 for Thoracic Surgery, American College of Radiology,  
 601 American Stroke Association, Society of Cardiovascular  
 602 Anesthesiologists, Society for Cardiovascular Angiography  
 603 and Interventions, Society of Interventional Radiology,  
 604 Society of Thoracic Surgeons, and Society for Vascular  
 605 Medicine. *Circulation* 2010;121:e266-369.  
 606 10. Mendoza DD, Kochar M, Devereux RB, Basson CT, Min  
 607 JK, Holmes K, Dietz HC, Milewicz DM, LeMaire SA,  
 608 Pyeritz RE, Bavaria JE, Maslen CL, Song H, Kroner BL,  
 609 Eagle KA, Weinsaft JW; GenTAC (National Registry of  
 610 Genetically Triggered Thoracic Aortic Aneurysms and  
 611 Cardiovascular Conditions) Study Investigators. Impact  
 612 of image analysis methodology on diagnostic and surgical  
 613 classification of patients with thoracic aortic aneurysms.  
 614 *Ann Thorac Surg* 2011;92:904-12.  
 615 11. Boskamp T, Rinck D, Link F, Kummerlen B, Stamm G,  
 616 Mildenerberger P. New vessel analysis tool for morphometric  
 617 quantification and visualization of vessels in CT and MR  
 618 imaging data sets. *Radiographics* 2004;24:287-97.  
 619 12. Green DB, Palumbo MC, Lau C. Imaging of  
 620

- 621 Thoracoabdominal Aortic Aneurysms. *J Thorac Imaging*  
622 2018;33:358-65.
- 623 13. Müller-Eschner M, Rengier F, Partovi S, Weber TF,  
624 Kopp-Schneider A, Geisbüsch P, Kauczor HU, von Tengg-  
625 Kobligk H. Accuracy and variability of semiautomatic  
626 centerline analysis versus manual aortic measurement  
627 techniques for TEVAR. *Eur J Vasc Endovasc Surg*  
628 2013;45:241-7.
- 629 14. Rengier F, Weber TF, Giesel FL, Böckler D, Kauczor HU,  
630 von Tengg-Kobligk H. Centerline analysis of aortic CT  
631 angiographic examinations: benefits and limitations. *AJR*  
632 *Am J Roentgenol* 2009;192:W255-63.
- 633 15. Quint LE, Liu PS, Booher AM, Watcharotone K, Myles  
634 JD. Proximal thoracic aortic diameter measurements  
635 at CT: repeatability and reproducibility according  
636 to measurement method. *Int J Cardiovasc Imaging*  
637 2013;29:479-88.
- 638 16. Lu JT, Brooks R, Hahn S, Chen J, Buch V, Kotecha  
639 G, et al. editors. *DeepAAA: Clinically Applicable and*  
640 *Generalizable Detection of Abdominal Aortic Aneurysm*  
641 *Using Deep Learning*. Cham: Springer International  
642 Publishing, 2019.
- 643 17. Biesdorf A, Rohr K, Feng D, von Tengg-Kobligk  
644 H, Rengier F, Böckler D, Kauczor HU, Wörz S.  
645 Segmentation and quantification of the aortic arch using  
646 joint 3D model-based segmentation and elastic image  
647 registration. *Med Image Anal* 2012;16:1187-201.
- 648 18. Sedghi Gamechi Z, Bons LR, Giordano M, Bos D, Budde  
649 RPJ, Kofoed KF, Pedersen JH, Roos-Hesselink JW, de  
650 Bruijne M. Automated 3D segmentation and diameter  
651 measurement of the thoracic aorta on non-contrast  
652 enhanced CT. *Eur Radiol* 2019;29:4613-23.
19. Rueckel J, Reidler P, Fink N, Sperl J, Geyer T, Fabritius  
MP, Ricke J, Ingrisch M, Sabel BO. Artificial intelligence  
assistance improves reporting efficiency of thoracic aortic  
aneurysm CT follow-up. *Eur J Radiol* 2021;134:109424.
20. Ghesu FC, Georgescu B, Zheng Y, Grbic S, Maier  
A, Hornegger J, Comaniciu D. Multi-Scale Deep  
Reinforcement Learning for Real-Time 3D-Landmark  
Detection in CT Scans. *IEEE Trans Pattern Anal Mach*  
*Intell* 2019;41:176-89.
21. Yang D, Xu D, Zhou SK, Georgescu B, Chen M, Grbic  
S, Metaxas D, Comaniciu D. editors. *Automatic Liver*  
*Segmentation Using an Adversarial Image-to-Image*  
*Network*. Cham: Springer International Publishing, 2017.
22. Popović ZB, Thomas JD. Assessing observer variability: a  
user's guide. *Cardiovasc Diagn Ther* 2017;7:317-24.
23. Koller M. *robustlmm: An R Package for Robust*  
*Estimation of Linear Mixed-Effects Models*. 2016  
2016;75:24.
24. Michelena HI, Khanna AD, Mahoney D, Margaryan  
E, Topilsky Y, Suri RM, et al. Incidence of aortic  
complications in patients with bicuspid aortic valves.  
*JAMA* 2011;306:1104-12.
25. Elefteriades JA, Farkas EA. Thoracic aortic aneurysm  
clinically pertinent controversies and uncertainties. *J Am*  
*Coll Cardiol* 2010;55:841-57.
26. McComb BL, Munden RF, Duan F, Jain AA, Tuite C,  
Chiles C. Normative reference values of thoracic aortic  
diameter in American College of Radiology Imaging  
Network (ACRIN 6654) arm of National Lung Screening  
Trial. *Clin Imaging* 2016;40:936-43.
27. McDonald RJ, Schwartz KM, Eckel LJ, Diehn FE, Hunt  
CH, Bartholmai BJ, Erickson BJ, Kallmes DF. The effects  
of changes in utilization and technological advancements  
of cross-sectional imaging on radiologist workload. *Acad*  
*Radiol* 2015;22:1191-8.
28. Bluemke DA, Moy L, Bredella MA, Ertl-Wagner BB,  
Fowler KJ, Goh VJ, Halpern EF, Hess CP, Schiebler ML,  
Weiss CR. Assessing Radiology Research on Artificial  
Intelligence: A Brief Guide for Authors, Reviewers, and  
Readers-From the Radiology Editorial Board. *Radiology*  
2020;294:487-9.

**Cite this article as:** Pradella M, Weikert T, Sperl JI, Kärger R, Cyriac J, Achermann R, Sauter AW, Bremerich J, Stieltjes B, Brantner P, Sommer G. Fully automated guideline-compliant diameter measurements of the thoracic aorta on ECG-gated CT angiography using deep learning. *Quant Imaging Med Surg* 2021;11(10):4245-4257. doi: 10.21037/qims-21-142