1	New morphological factor for predicting late proximal type I endoleak after
2	endovascular aneurysm repair.
3	
4	Hiroshi Banno, Masayuki Sugimoto, Tomohiro Sato, Shuta Ikeda, Yohei Kawai, Takuya
5	Tsuruoka, Akio Kodama, Kimihiro Komori
6	
7	Division of Vascular and Endovascular Surgery, Department of Surgery, Nagoya
8	University Graduate School of Medicine, Nagoya, Japan
9	
10	Corresponding author: Hiroshi Banno; Division of Vascular and Endovascular Surgery,
11	Department of Surgery, Nagoya University Graduate School of Medicine, 65 Tsurumai-
12	cho, Showa-ku, 466-8550, Nagoya, Japan (e-mail: hbanno@med.nagoya-u.ac.jp)
13	
14	Short title: New morphological factor predictive of late T1AEL
15	
16	Word count: 4820

#### 18 ABSTRACT

**Background:** Although we have witnessed several cases of late proximal type I endoleak (T1AEL) after endovascular aneurysm repair (EVAR), most patients did not have "hostile neck" preoperatively. We hypothesized that the distance between the lowest renal artery and the neck angulation point and neck length are the two most important factors for maintaining long-term proximal sealing. This study evaluated "neck hostility", which is the product of the distance to the angulation point and the neck length, as a preoperative morphological risk factor for the development of late T1AEL after EVAR.

Methods: A retrospective review of a prospectively assembled database was performed for all patients who had undergone EVAR at a single institution from June 2007 to May 2017. Patient demographics and preoperative imaging data were collected, and Cox regression analysis was performed to identify the risk factors for late T1AEL.

30 Results: Of the 655 patients who underwent EVAR during the study period, 115 were 31 excluded due to complex EVAR (n=14), primary indications for iliac aneurysms (n=86), 32 primary T1AEL (n = 3), or other reasons (n=15). Of the remaining 537 patients, twelve 33 patients (2.2%) developed late T1AEL a median of 3.2 (interquartile range [IQR]; 3.0, 34 5.4) years after EVAR. Receiver operating characteristic (ROC) curve analysis revealed a neck hostility cutoff value of 8. Cox regression analysis revealed that a neck hostility 35 36 value  $\leq 8$  and conical neck anatomy were risk factors for the development of late T1AEL 37 after EVAR. Well-known hostile neck factors such as short neck, severe angulated neck, 38 and severe calcification/thrombus in the proximal neck were not significantly different. 39 Conclusions: The present study demonstrated a correlation between late T1AEL and the

- 40 product of the angulation distance and the neck length. This factor may be useful for
- 41 predicting poor late proximal outcomes after EVAR.
- 42
- 43 Keywords: EVAR, AAA, Hostile neck, Endoleak, type Ia endoleak

44

#### 46 **INTRODUCTION**

47 Endovascular aneurysm repair (EVAR) has become a standard treatment for 48 abdominal aortic aneurysm (AAA) and has significantly improved short-term outcomes 49 compared with open surgical repair (OSR).<sup>1-3</sup> However, this procedure can fail in the long 50 term, partly due to the interdependence of endograft sealing and the morphological 51 aspects of the proximal aortic neck. Accordingly, manufacturers traditionally have 52 defined a short proximal neck length (< 10 mm or < 15 mm), severe neck angulation (>53 60°), or significant thrombus and/or calcification as outside their instructions for use 54 (IFUs). These IFUs, however, have been based on benchtop research and aimed to achieve 55 immediate optimal outcomes after EVAR.

The term "hostile neck" was first advocated by the Pittsburgh group in 2003 to characterize outcomes after EVAR in patients with unfavourable neck anatomy.<sup>4</sup> The term is currently used when the aortic anatomy is beyond the eligibility criteria for manufacturers' regulatory clinical trials. However, nearly half (47.6%) of cases violate the IFUs in clinical practice.<sup>5</sup> The impact of each factor on late proximal outcomes after EVAR is not clear because various outcomes have been reported in regards to the association between hostile neck anatomy and late proximal type I endoleak (T1AEL).<sup>6–</sup>

63

9

Although we have experienced several cases of late proximal type I endoleak (T1AEL) after EVAR, most were without "hostile neck" anatomy preoperatively. In clinical settings, a short proximal neck, including conical neck anatomy, is of course challenging for primary proximal sealing, but a severely angulated neck, which is

68	included in the hostile neck criteria, is not so difficult if the angulation point is far from
69	the lowest renal artery. Accordingly, we hypothesized that the distance between the lowest
70	renal artery and the neck angulation point and neck length are the two most important
71	factors for maintaining proximal sealing in the long term. The purpose of this study was
72	therefore to evaluate a factor multiplied by these two factors as a preoperative
73	morphological factor associated with late T1AEL after EVAR.
74	
75	METHODS
76	The Nagoya University Institutional Review Board approved this study (2019-
77	0096), and the need for individual patient consent was waived since all data were obtained
78	from routine clinical care.
79	Study population
80	Patients who underwent elective EVAR for infrarenal AAA between June 2007
81	and May 2017 at our institution were reviewed. The indication for repair was $AAA \ge 5$
82	cm in diameter, a rapidly growing an eurysm ( $\geq$ 5 mm per 6 months), or a saccular
83	aneurysm. In principle, EVAR was applied to patients older than 75 years of age or
84	patients at high risk with OSR even if they were younger than 75 years of age. <sup>10</sup> Patients
85	with targeted aneurysms located in the abdominal aorta (not the iliac artery) and treated
86	with a commercially available bifurcated endograft were included in this study. Patients
07	
87	treated in the emergent setting and patients who required concomitant procedures to the
87 88	treated in the emergent setting and patients who required concomitant procedures to the renal artery, including chimney/snorkel or fenestrated/branched techniques, were

90	excluded from this study (in contrast, patients who were treated immediately after the
91	detection of primary T1AEL on postoperative computed tomography (CT) angiography
92	were included).

93

### Study/follow-up protocol and procedures

Patient baseline demographics, comorbidities, medications, operative details, and outcomes were collected. All patients underwent CT angiography with threedimensional reconstruction before surgery. All procedures were performed in a fully equipped operating room with the patients under regional or general anaesthesia and with fluoroscopic guidance. Patients underwent CT angiographies (if renal function permitted) or plain CT with Doppler ultrasound tests at 3, 6, and 12 months and annually thereafter.

#### 101 Image analysis

A 3D imaging workstation (TeraRecon, Inc., Durham, NC) was used to generate multiple 3D reconstructions of volumetric data sets from preoperative CT angiography scans. A greater curve line was extracted automatically after generating a 3D aortic lumen centerline (a greater curve line is indicated as a yellow line in **Figure 1A**, and a centerline is indicated as a yellow line in **Figure 1B**).

#### 107 **Definitions**

Preoperative coronary artery disease (CAD) was defined as an abnormal result on coronary angiography and a history of myocardial infarction or open or percutaneous coronary artery revascularization. Chronic obstructive pulmonary disease (COPD) was identified by pulmonary function studies or active treatment with medication.

Hypertension (HT), dyslipidaemia (DL), and diabetes were identified in patients undergoing active medical treatment or diet modification. Cerebrovascular disease (CVD) was defined as a history of stroke, transient ischaemic attack, or carotid intervention.

116 In this study, the angulation point was defined as the point perpendicular to the 117 greater curve line at 45 degrees from just below the lowest renal artery. Then, the 118 angulation distance (mm) was measured to be the greater curve line length from the 119 lowest renal artery to the angulation point (Figure 1A). An angulation distance over 50 120 mm and aortic angulation less than 45 degrees were set to 50 mm as the angulation 121 distance. Neck length was defined as the centerline length between the level of the lowest 122 renal artery and the level where the diameter of the infrarenal aortic neck exceeded that 123 at the lowest renal artery by 10%. "Neck hostility" was defined as the neck length x the 124 angulation distance/100. A short neck was defined as an infrarenal neck length less than 125 15 mm. Other hostile neck factors were commonly used factors. Severe angulation was 126 defined as infrarenal proximal neck angulation greater than 60 degrees. Severe 127 calcification was defined as a  $\geq$ 50% calcified proximal neck, and severe thrombus was 128 defined as a  $\geq$ 50% circumferential proximal neck thrombus ( $\geq$ 2 mm). The term "hostile 129 neck" was used if 1 or more of three proximal neck factors (short proximal neck, severe 130 angulation, or severe calcification/thrombus) were present. A conical neck was defined as 131 gradual neck dilatation  $\geq 2$  mm within the first 10 mm below the lowest renal artery. A 132 large neck was indicated when the patient was treated with a main body  $\geq 31$  mm in 133 proximal diameter.

#### 134 **Data analysis**

135 All statistical analyses were performed using JMP pro statistical software 136 version 15 (SAS Institute Inc., Cary, NC, USA). For comparisons, categorical variables 137 were analysed using a chi-square test or Fisher's exact test as appropriate. Continuous 138 variables were analysed using Student's *t*-test or the Mann-Whitney test as appropriate. 139 Cox proportional hazard analysis was applied for the univariate analysis of risk factors 140 for the development of late T1AEL. Analysis of the receiver operating characteristic 141 (ROC) curves was used to determine the cutoff value of "neck hostility" for late T1AEL, 142 and the area under the ROC curve (AUC) provides a measure of the factor "neck hostility" 143 to discriminate between patients who developed late T1AEL and those who did not. 144 Freedom from late T1AEL was assessed using Kaplan-Meier life-table analysis, and a 145 log-rank test was performed. All p values were two-sided, with p < 0.05 regarded as 146 indicative of statistical significance.

147

#### 148 **RESULTS**

During the study period, a total of 655 patients underwent EVAR at our institution. Among them, 115 patients were excluded from the study, namely, due to complex EVAR (n=14), treatment with aorto-uni-iliac devices (n=6) or aortic cuffs (n=7), primary indications for iliac aneurysms (n=86), or other reasons (n=2). In addition, three patients who developed primary T1AEL were excluded because the primary endpoint of this study was late proximal type I endoleak. As a result, 537 patients were ultimately included in this study. Patient demographics are shown in **Table 1**. There was no in156 hospital death, and the primary technical success rate was 100% (Note: three patients with 157 primary type I endoleaks who were observed conservatively were excluded from this 158 study). The preoperative morphological characteristics are detailed in Table 2. 159 Approximately 30% of patients had hostile neck anatomy, and of those, ten patients had 160 two hostile neck criteria. Implanted devices are also shown in Table 2. The median 161 follow-up duration was 4.0 (interquartile range [IQR]; 2.1, 6.0) years. Among the study 162 cohort, twelve patients (2.2%) developed late T1AELs a median of 3.2 (IQR; 3.0, 5.4) 163 years after EVAR (one was due to distal migration of endografts). As a result, one patient 164 died of AAA rupture, and another patient was observed to have late T1AEL because of 165 the recurrence of malignant disease. Of the remaining ten patients, nine were treated 166 successfully by proximal aortic extension, and one required open prosthetic graft 167 replacement (Table 3).

#### 168 **ROC curve analysis**

169ROC curve analysis of neck hostility revealed that the AUC for the predicted170probabilities was 0.79 (95% confidence interval [CI]: 0.66-0.87). At the optimal cutoff171value of 8, the sensitivity of the minimum neck hostility for predicting late T1AEL was17275.0% with 78.1% specificity (Figure 2). Accordingly, we determined a neck hostility173value of 8 or less to be a hostile neck factor and compared it with other popular hostile174neck criteria. There were 124 (23.1%) patients with neck hostility values  $\leq 8$  in the study175cohort.

#### 176 Univariate analysis

177

Univariate Cox regression analysis revealed a conical neck anatomy (P = .013,

178hazard ratio [HR]: 4.269, 95% CI: 1.352-13.478) and neck hostility value  $\leq 8$  (P =.0003,179HR: 11.168, 95% CI: 3.019-41.308) to be significant preoperative morphological risk180factors for late T1AEL (**Table 4**). Well-known hostile neck criteria such as short neck,181severe angulation, severe calcification/thrombus, conical neck, and large neck were not182statistically significant as discriminators of late T1AEL. Subsequent multivariate analysis183could not be performed owing to the small number of cases that developed late T1AELs.

184 Freedom from late proximal type I endoleaks

The Kaplan-Meier survival curve of the freedom from late T1AEL is shown in Figure 2. At the 4-year follow-up, the estimated rate of freedom from late T1AEL was 99.2% (n = 2; standard error [SE], 0.0055) in the neck hostility group with values >8 and 92.0% (n = 6; SE, 0.081) in the neck hostility group with values  $\leq 8$  (log rank test; P  $\leq 0.001$ ) (Figure 3).

190

#### 191 **DISCUSSION**

192 This is the first report advocating for the use of the factor "neck hostility", 193 which multiplies two factors, angulation distance and neck length, as a predictor of late 194 T1AEL. The term "hostile neck" was coined by Dillavou et al in their study to determine 195 a suitable neck anatomy for successful EVAR.<sup>4</sup> In current clinical settings, the term is 196 used when the proximal aortic anatomy is excluded from the eligibility criteria for 197 manufacturers' regulatory clinical trials (IFUs). However, these criteria are based on 198 benchtop and clinical research by manufacturers only for securing optimal immediate 199 outcomes after EVAR, and whether these hostile neck criteria impact long-term outcomes

200 remains controversial.

201 In terms of proximal neck length, there is no room for discussion about its 202 hostility. However, most surgeons do not seem to face much difficulty when using the 203 latest endograft for severe angulation, which is a factor of the so-called hostile neck 204 anatomy. In fact, even if the angulation is severe, the most recently developed endografts 205 conform well, and the long-term results are good when the distance to the angulation 206 point is long enough (Figure 1C, D). If the distance to the angulation point is short and 207 there is a short neck length, even the most recently developed endografts cannot conform 208 to it, and proximal sealing becomes poor. In contrast, if the distance to the angulation 209 point is short but the proximal neck length is long enough, good sealing can be achieved. 210 Based on the above, we hypothesized that the factor obtained by multiplying the neck 211 length by the distance to the angulation point might have an impact on proximal sealing 212 in the late period, which was evaluated in this study.

213 Regarding the angulation point, we initially determined the angulation point 214 based on the centerline and started studying the distance to it. However, even if the 215 centerline is actually angled, proximal sealing can be improved by aligning the stent graft 216 with the proximal neck when there is sufficient space on the greater curve of the aortic 217 lumen. In contrast, if the greater curve is also severely angulated, it becomes difficult to 218 place the stent graft coaxially in the aorta proximal to the angulation point. Therefore, we 219 determined the angulation point of the greater curve line and found that the angulation 220 distance to that point was important.

221

In the 3D workstation, the greater curve line can be drawn automatically. The

degree of change from the perpendicular plane at the lowest renal artery level to the angulation point was examined preliminarily at 30, 45, and 60 degrees, and we concluded that 45 degrees was the degree of clinical difficulty. As a result, we determined 45 degrees to be the threshold value in this study.

226 The hostile neck criteria are essentially anatomic criteria for clinical decision 227 making to obtain primary success. Although there are several studies investigating the 228 impacts of hostile neck anatomy on post-EVAR outcomes, most of them are studies of 229 intraoperative or short-term outcomes.<sup>11–13</sup> The impacts of "so-called" hostile neck 230 anatomy or each factor of challenging neck on late outcomes are diverse. Jordan et al 231 investigated a study population with challenging neck anatomy enrolled in the ANCHOR 232 trial and concluded that neck diameter and length were predictive of T1AEL (including 233 primary T1AEL, though), but mural thrombus was protective.<sup>14</sup> Antoniou et al conducted 234 a meta-analysis of the outcomes after EVAR in patients with hostile versus favourable 235 neck anatomy and reported that patients with hostile anatomy had a fourfold increased 236 risk of developing T1AEL<sup>8</sup>. However, this analysis was of events within 30 days and at 237 most 1 year. In contrast, AbuRahma reported that there was no significant difference in 238 the incidence of late T1AEL between hostile and favourable neck anatomy. Limited to 239 neck length, Bastos Goncalves et al reported that a neck length <10 mm was a significant 240 risk factor for postoperative neck-related adverse events corresponding to T1AEL. 241 However, most events occurred within one year after surgery.<sup>15</sup> AbuRahma et al also 242 reported that neck length <10 mm was an independent risk factor for late T1AEL.<sup>7</sup> 243 Several studies have reported the association between a large neck and postoperative

244 T1AEL.<sup>6,16–18</sup> They reported that a large proximal neck was associated with not only 245 delayed T1AEL but also lower survival. A conical neck is also a well-known risk factor 246 for T1AEL, and this was the second strongest risk factor for late T1AEL in our study. 247 Pitoulias et al also reported that this anatomy is strongly associated with proximal neck 248 failure after EVAR, but most T1AELs in their study developed within one month after 249 EVAR. Although there are some reports that the development of T1AEL is not influenced 250 by neck angulation (similar to our findings),<sup>19</sup> others have suggested that aortic curvature 251 is a better parameter than angulation to predict late T1AEL.<sup>20</sup>

252 Even in the endovascular era, we have OSR as an alternative, and accordingly, 253 we should aim to eradicate late aneurysm-related death after EVAR (which is extremely 254 rare after OSR). There are several types of endoleaks that cause aneurysm sac growth 255 after EVAR, and at present, an increase in aneurysm diameter occurs at a constant rate, 256 which is difficult to predict before surgery. Among those endoleaks, proximal type I 257 endoleak is the most dangerous and difficult to repair. If a morphological factor, such as 258 "neck hostility" in this study, that can predict late T1AEL, is identified, patients with good 259 long-term outcome after EVAR can be identified preoperatively. As a result, this less 260 invasive treatment for AAA can be applied to more patients, with better long-term 261 outcomes. Therefore, the factor "neck hostility" might be helpful in achieving better late 262 outcomes when surgeons are making decisions related to repair planning.

The present study had several limitations. First, our study was a nonrandomized, retrospective, observational study conducted at a single institution. Thus, the findings presented herein are susceptible to unmeasured confounding factors, referral bias, and other potential biases. Second, because the number of patients who developed late T1AEL was quite small, multivariate analysis was not performed. In addition, the cut-off value of 8 for "neck hostility" was calculated based on a small sample size. The primary purpose of this study is not to propose a cutoff value for neck hostility of 8, but rather it is to focus on the use of the product two factors, neck length and angulation distance, as a predictor of late T1AEL after EVAR. Larger scale data and image analysis are required in future work.

273

#### 274 CONCLUSIONS

In this study, we presented a new morphological factor, namely, "neck hostility", which multiples two factors, the angulation distance and neck length, and demonstrated a correlation between this factor and the development of late T1AEL. To the best of our knowledge, the present study is the first to use the distance to the angulation point in the greater curve of the proximal aortic neck as a factor in predicting late T1AEL. Further studies utilizing similar approaches are expected to improve late outcomes after EVAR.

281

Funding: This research did not receive any specific grant from funding agencies in thepublic, commercial, or not-for-profit sectors.

284

Declaration of conflicts of interest: All authors declare that they have no conflicts ofinterest.

287 This research did not receive any specific grant from funding agencies in the public,

288 commercial, or not-for-profit sectors.

291	1	Becquemin JP, Pillet JC, Lescalie F, Sapoval M, Goueffic Y, Lermusiaux P, et al.
292		A randomized controlled trial of endovascular aneurysm repair versus open
293		surgery for abdominal aortic aneurysms in low- to moderate-risk patients. J Vasc
294		Surg 2011; <b>53</b> (5):1167-1173.e1. Doi: 10.1016/j.jvs.2010.10.124.
295	2	Blankensteijn JD, de Jong SECA, Prinssen M, van der Ham AC, Buth J, van
296		Sterkenburg SMM, et al. Two-Year Outcomes after Conventional or
297		Endovascular Repair of Abdominal Aortic Aneurysms. N Engl J Med
298		2005; <b>352</b> (23):2398–405. Doi: 10.1056/NEJMoa051255.
299	3	Participants ET. Comparison of endovascular aneurysm repair with open repair in
300		patients with abdominal aortic aneurysm (EVAR trial 1), 30-day operative
301		mortality results: randomised controlled trial. Lancet 2004;364:843-8. Doi:
302		10.1016/S0140-6736(04)16979-1.
303	4	Dillavou ED, Muluk SC, Rhee RY, Tzeng E, Woody JD, Gupta N, et al. Does
304		hostile neck anatomy preclude successful endovascular a ortic aneurysm repair? ${\cal J}$
305		Vasc Surg 2003;38(4):657-63. Doi: 10.1016/S0741-5214(03)00738-9.
306	5	Hoshina K, Ishimaru S, Sasabuchi Y, Yasunaga H, Komori K. Outcomes of
307		Endovascular Repair for Abdominal Aortic Aneurysms. Ann Surg
308		2019; <b>269</b> (3):564–73. Doi: 10.1097/sla.00000000002508.
309	6	Stather PW, Sayers RD, Cheah A, Wild JB, Bown MJ, Choke E. Outcomes of
310		endovascular aneurysm repair in patients with hostile neck anatomy. Eur J Vasc
311		Endovasc Surg 2012;44(6):556-61. Doi: 10.1016/j.ejvs.2012.10.003.
312	7	AbuRahma AF, Yacoub M, Mousa AY, Abu-Halimah S, Hass SM, Kazil J, et al.

313 Aortic Neck Anatomic Features and Predictors of Outcomes in Endovascular 314 Repair of Abdominal Aortic Aneurysms Following vs Not Following Instructions 315 for Use. J Am Coll Surg 2016;222(4):579–89. Doi: 316 10.1016/j.jamcollsurg.2015.12.037. 317 8 Antoniou GA, Georgiadis GS, Antoniou SA, Kuhan G, Murray D. A meta-318 analysis of outcomes of endovascular abdominal aortic aneurysm repair in 319 patients with hostile and friendly neck anatomy. J Vasc Surg 2013;57(2):527-38. 320 Doi: 10.1016/j.jvs.2012.09.050. 321 9 Cerini P, Guzzardi G, Divenuto I, Parziale G, Brustia P, Carriero A, et al. Are 322 abdominal aortic aneurysms with hostile neck really unsuitable for EVAR? Our 323 experience. Radiol Medica 2016;121(6):528-35. Doi: 10.1007/s11547-016-0620-324 y. 325 10 Yamamoto K, Komori K, Banno H, Narita H, Kodama A, Sugimoto M. 326 Validation of patient selection for endovascular aneurysm repair or open repair of 327 abdominal aortic aneurysm: Single-center study. Circ J 2015;79(8):1699-705. 328 Doi: 10.1253/circj.CJ-14-1160. 329 11 Chaikof EL, Fillinger MF, Matsumura JS, Rutherford RB, White GH, Blankensteijn JD, et al. Identifying and grading factors that modify the outcome 330 331 of endovascular aortic aneurysm repair. J Vasc Surg 2002;35(5):1061-6. Doi: 332 10.1067/mva.2002.123991. 333 12 Grisafi JL, Rahbar R, Nelms J, Detschelt EL, Chess BA, Benckart DH, et al. 334 Challenging neck anatomy is associated with need for intraoperative

335		endovascular adjuncts during Endovascular Aortic Aneurysm Repair (EVAR).
336		Ann Vasc Surg 2011;25(6):729-34. Doi: 10.1016/j.avsg.2011.02.028.
337	13	Torsello G, Troisi N, Donas KP, Austermann M. Evaluation of the Endurant stent
338		graft under instructions for use vs off-label conditions for endovascular aortic
339		aneurysm repair. J Vasc Surg 2011;54(2):300–6. Doi: 10.1016/j.jvs.2010.12.062.
340	14	Jordan WD, Ouriel K, Mehta M, Varnagy D, Moore WM, Arko FR, et al.
341		Outcome-based anatomic criteria for defining the hostile aortic neck. J Vasc Surg
342		2015; <b>61</b> (6):1383-1390.e1. Doi: 10.1016/j.jvs.2014.12.063.
343	15	Bastos Goncalves F, Hoeks SE, Teijink JA, Moll FL, Castro JA, Stolker RJ, et al.
344		Risk factors for proximal neck complications after endovascular aneurysm repair
345		using the endurant stentgraft. Eur J Vasc Endovasc Surg 2015;49(2):156-62.
346		Doi: 10.1016/j.ejvs.2014.10.003.
347	16	Gargiulo M, Gallitto E, Wattez H, Verzini F, Bianchini Massoni C, Loschi D, et
348		al. Outcomes of endovascular aneurysm repair performed in abdominal aortic
349		aneurysms with large infrarenal necks. J Vasc Surg 2017;66(4):1065–72. Doi:
350		10.1016/j.jvs.2017.01.066.
351	17	Howard DPJ, Marron CD, Sideso E, Puckridge PJ, Verhoeven ELG, Spark JI.
352		Editor's Choice – Influence of Proximal Aortic Neck Diameter on Durability of
353		Aneurysm Sealing and Overall Survival in Patients Undergoing Endovascular
354		Aneurysm Repair. Real World Data from the Gore Global Registry for
355		Endovascular Aortic Treatment (GREAT. Eur J Vasc Endovasc Surg
356		2018; <b>56</b> (2):189–99. Doi: 10.1016/j.ejvs.2018.03.027.

357	18	Oliveira NFG, Gonçalves FB, Ultee K, Pinto JP, Josee van Rijn M, Raa S Ten, et
358		al. Patients with large neck diameter have a higher risk of type IA endoleaks and
359		aneurysm rupture after standard endovascular aneurysm repair. J Vasc Surg
360		2019; <b>69</b> (3):783–91. Doi: 10.1016/j.jvs.2018.07.021.
361	19	Oliveira NFG, Bastos Gonçalves FM, De Vries JPPM, Ultee KHJ, Werson DAB,
362		Hoeks SE, et al. Mid-term results of EVAR in severe proximal aneurysm neck
363		angulation. Eur J Vasc Endovasc Surg 2015;49(1):19–27. Doi:
364		10.1016/j.ejvs.2014.10.001.
365	20	Schuurmann RCL, Van Noort K, Overeem SP, Ouriel K, Jordan WD, Muhs BE,
366		et al. Aortic Curvature Is a Predictor of Late Type Ia Endoleak and Migration
367		after Endovascular Aneurysm Repair. J Endovasc Ther 2017;24(3):411–7. Doi:
368		10.1177/1526602817700378.
369		

370

	Total (n = 537)
Age, years	79 (74, 82)
Male sex	446 (83.1)
Comorbidities	
HT	396 (73.7)
DL	230 (42.8)
Diabetes	60 (11.2)
CAD	164 (30.5)
CVD	79 (14.7)
CKD≥G3b*	124 (23.1)
Dialysis	13 (2.4)
COPD	255 (47.5)
Current smoker	81 (15.1)
Medicines	
Antiplatelet agent	203 (37.8)
Anticoagulation agent	38 (7.1)
Statin	204 (38.0)
β-blocker	105 (19.6)
CCB	272 (50.7)
ACEI/ARB	220 (41.0)

371 **Table 1.** Patient demographics

372 ACEI, angiotensin-converting enzyme inhibitor; ARB, angiotensin receptor blocker;

373 CAD, coronary artery disease; CCB, calcium-channel blocker; CI, confidence interval;

374 CKD, chronic kidney disease; COPD, chronic obstructive pulmonary disease; CVD,

375 cerebrovascular disease; DL, dyslipidaemia; HT, hypertension.

376 Data are presented as the median (interquartile range) or number (%).

377 \*CKD  $\geq$  G3b indicates eGFR < 45 ml/min/1.73 m<sup>2</sup>.

## 

### 

## Table 2. Preoperative morphological characteristics and implanted devices

	Total (n = 537)			
Aneurysm diameter (mm)	52 (50, 58)			
Proximal neck length (mm)	27 (20, 35)			
Hostile neck	157 (29.2)			
No. of hostile neck criteria				
1	147 (27.3)			
2	10 (1.9)			
Hostile neck factors				
Short neck	38 (7.1)			
Severe angulation	124 (23.1)			
Severe calcification/thrombus	5 (0.9)			
Conical neck	82 (15.3)			
Large neck	66 (12.3)			
Proximal adjunctive procedure	85 (15.8)			
Angulation distance (mm)	50 (50, 50)			
Neck hostility	1250 (850, 1650)			
Device type				
Gore Excluder	199 (37.1)			
Cook Zenith	156 (29.1)			
Medtronic Endurant	136 (25.3)			
Endologix Powerlink/AFX	22 (4.1)			
Lombard Aorfix	11 (2.0)			
Cordis Incraft	8 (1.5)			
Medtronic Talent	5 (0.9)			

381 The definitions of the morphological factors are detailed in the main body of the382 manuscript.

<sup>383</sup> Data are presented as the median (interquartile range) or number (%).

No.	Aortic neck diameter (mm)	Diameter of endograft (mm)	Sac behaviour*	Type of repair	Outcomes	
1	28	32	Growth	Observation***	Death	
2	20	23	Growth	PE	Success	
3	24	28	Shrinkage**	PE	Success	
4	27	30	Growth	PE	Success	
5	25	28	Growth	Open conversion	Success	
6	23	26	Growth	PE	Success	
7	30	23	Growth	PE	Success	
8	25	28	Shrinkage**	PE with chimney	Success	
9	20	25	Growth	PE with fenestration	Success	
10	32	25	Growth	Growth PE with chimney		
11	23	28	Growth	Open conversion	Death	
12	32	36	Growth	PE	Success	

**Table 3.** Details of cases developed late type Ia endoleak

387 \*Sac behaviour at the time of late type Ia endoleak development.

388 \*\*These two cases developed type Ia endoleak when the significantly reduced aneurysm

389 expanded.

390 \*\*\*This case was observed to have late type Ia endoleak because of the recurrence of

- 391 malignant disease.
- 392 PE, proximal extension
- 393

394 Table 4. Univariate Cox regression analysis of the preoperative morphological risk395 factors for late proximal type I endoleak.

396

Factors	P value	HR	95% CI
Hostile neck (#1 or #2 or #3)	.120	2.499	.788 – 7.923
Short neck (#1)	.634	1.647	.212 - 12.820
Severe neck angulation (#2)	.129	2.546	.761 - 8.510
Severe calcification/thrombus (#3)	.999	0.000	.000
Conical neck	.013	4.269	1.352 - 13.478
Large neck	.473	1.744	.382 – 7.963
Proximal adjunctive procedure	.999	0.000	.000
Neck hostility value $\leq 8$	.0003	11.168	3.019 - 41.308

397

398 Hostile neck anatomy was indicated when one or more of three proximal neck factors 399 (short neck, severe infrarenal neck angulation, and severe neck calcification/thrombus) 400 were present. A short neck was defined as an infrarenal neck length less than 15 mm. 401 Severe neck angulation was defined as infrarenal neck angulation greater than 60°. Severe 402 neck calcification/thrombus was defined as either calcified proximal neck 50% or more 403 or circumferential proximal neck thrombus ( $\geq 2 \text{ mm}$ ) 50% or more. A conical neck was 404 indicated when neck dilatation was 2 mm or more within the first 10 mm after the lowest 405 renal artery. A large neck was indicated when the patient was treated with a main body 31 406 mm or more in the proximal neck diameter. 407 Boldface *P* values represent statistical significance (P < .05) 408

409 **Figure legends** 

410

411 Figure 1. Illustration (A) and CT angiography images (B, C, D, E) of the angulation 412 distance. (A) The angulation point was defined as the point perpendicular to the greater 413 curve at 45° just below the lowest renal artery. The angulation distance was measured as 414 the greater curve length from the lowest renal artery to the angulation point. An angulation 415 distance greater than 50 mm and aortic angulation less than 45° was set to 50 mm as the 416 angulation distance. Figure 1B and C shows a case with a short angulation distance (23 417 mm). This case developed late type Ia endoleak, which resulted in death by rupture five 418 years after the initial surgery. Figure 1D and 1E shows another case with a relatively short 419 and severe angulated neck but a long angulation distance (80 mm). This case obtained 420 significant sac shrinkage with sustained good proximal sealing.

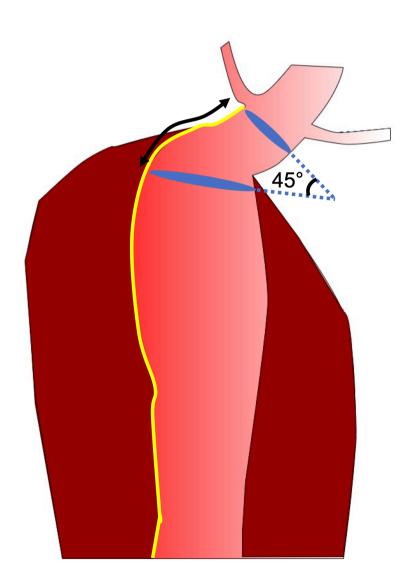
421

Figure 2. Receiver operating characteristic (ROC) curve analysis of neck hostility for the prediction of late type Ia endoleak. The area under the curve (AUC) for the predicted probabilities was 0.79. The sensitivity was 75.0% and the specificity was 78.1% at the optimal cutoff value of 8.

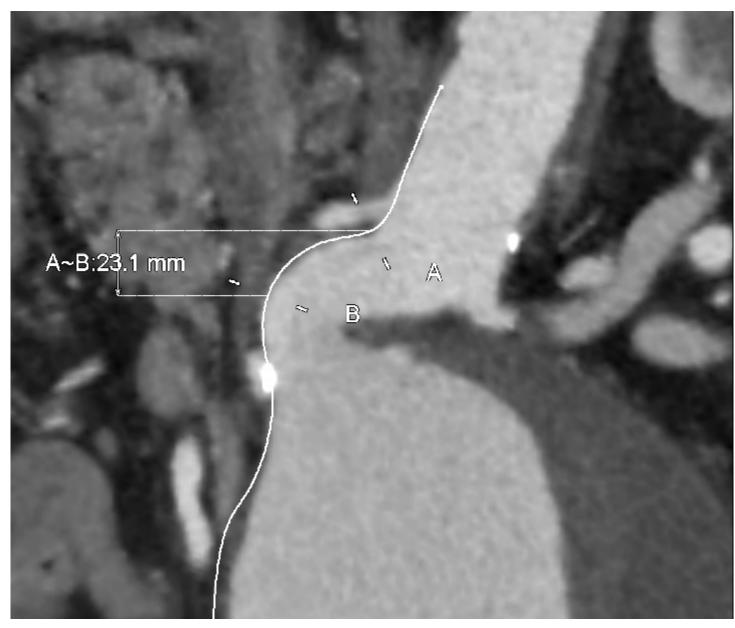
426

Figure 3. Kaplan-Meier survival curve for freedom from late type Ia endoleak. At the 4year follow-up, the estimated rate of freedom from late T1AEL was 99.2% in the neck hostility group with values >8 and 92.0% in the neck hostility group with values  $\leq 8$  (logrank test; *P* <.0001).

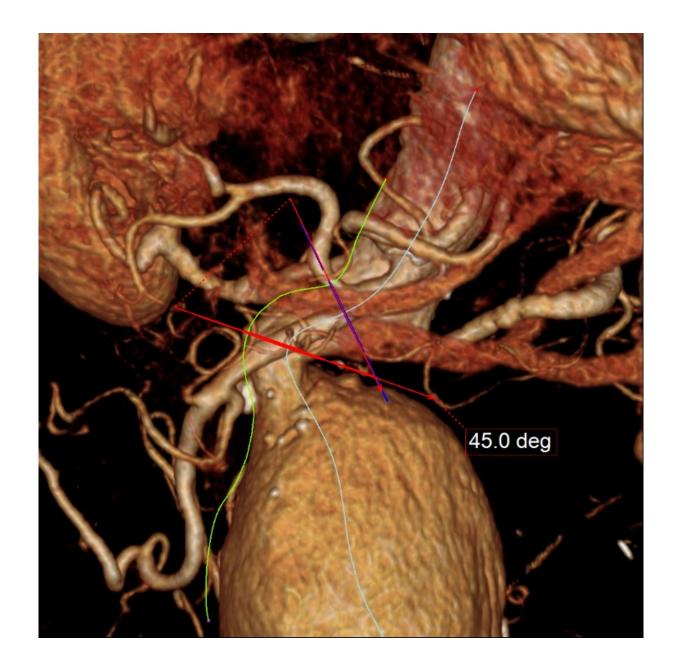




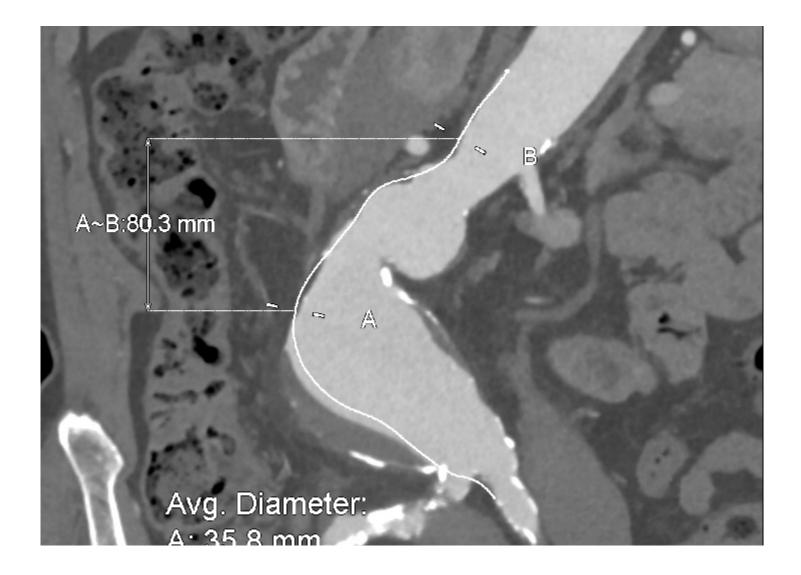
# Figure 1B



# Figure 1C



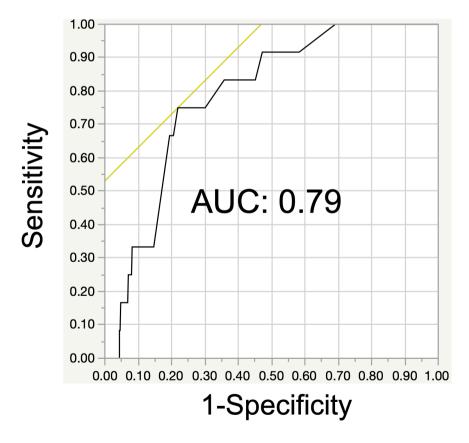
## Figure 1D



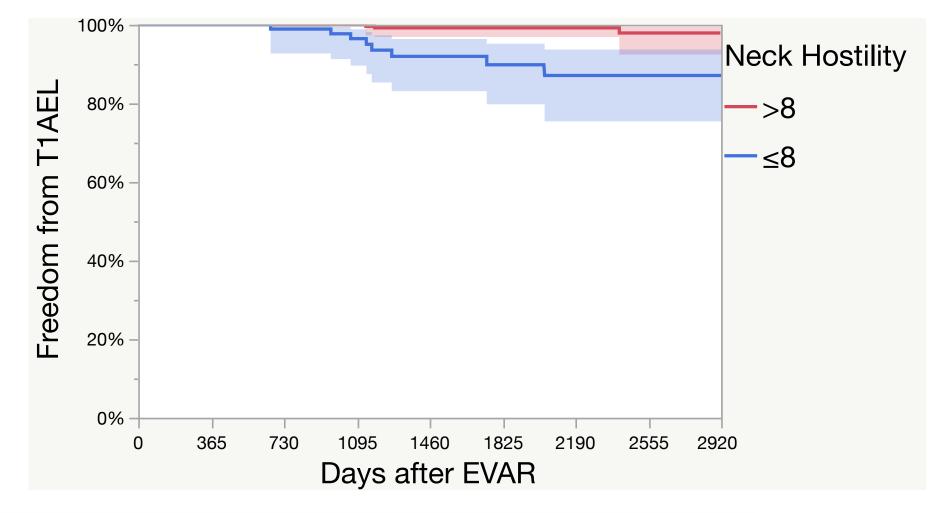
# Figure 1E



## Figure 2



## Figure 3



Neck hostility >8	413	369	325	281	224	168	107	69	49
Neck hostility ≤8	124	108	88	76	52	43	29	22	17

No. at risks