

Endoleak detection after endovascular aortic aneurysm repair utilizing dynamic, time-resolved computed tomography angiography

Ph.D. Thesis (short version)

Márton Tibor Berczeli

Károly Rácz Doctoral School of Clinical Medicine
Semmelweis University



Supervisor: Péter Sótonyi, M.D., Ph.D.

Official reviewers: László Benkő, MD, Ph.D.

Dávid Korda, MD, Ph.D.

Head of the Complex Examination Committee:

György Wéber, M.D., Ph.D.

Members of the Complex Examination Committee:

Ferenc Alföldy, M.D., Ph.D.

Eszter Végh, M.D., Ph.D.

Budapest

2022

1. Introduction:

The widespread use of endovascular aortic aneurysm repair (EVAR) has become apparent in the past decades. The favorable perioperative mortality rate and the totally percutaneous nature of the procedure were the main driving factors of its success. However, major trials focusing on the long-term outcome of EVAR reported higher re-intervention rate with EVAR few years after the index procedure as compared to open repair. It is evident that as the number of EVARs grew over the years, a subsequent rise in the number of graft-related complications occurred. Thus, it was pertinent to adequately address the sequela of EVAR by utilizing advanced imaging techniques, such as dynamic, time-resolved computed tomography angiography (d-CTA) imaging. This imaging modality overcomes the limitation of having just two acquisition (time) points (early and delayed phase) after contrast injection. D-CTA involves multiple (~10-12) scan acquisitions after contrast injection enabling aneurysm sac imaging during the transit of contrast material across multiple time points. Having multiple scans along the contrast enhancement curve opens up a plethora of new aspects in characterizing endoleaks based on objective parameters such as temporal Hounsfield unit change (quantitative analysis).

2. Objectives:

After developing our dynamic CTA protocol for endoleak detection in patients who underwent EVAR, our first objective **(1)** was to explore if d-CTA has better endoleak type diagnosing capability compared to triphasic computed tomography angiography (t-CTA) when using DSA as baseline reference. In addition to determining endoleak type, our second aim **(2)** was to compare the radiation exposure during image acquisition with the two CT modalities. Our third objective **(3)** was to match d-CTA, t-CTA, and DSA in detecting the number of inflow vessels in patients diagnosed with type II endoleak. Time-resolved CTA acquisition opens up a plethora of new aspects to characterize endoleaks based on objective/quantitative parameters. After developing our quantitative image analysis protocol, our fourth objective **(4)** was to demonstrate the feasibility of a quantitative parameter aiding the differential diagnosis of endoleak types. In this study, similar to our third objective, **(5)** we matched d-CTA and DSA in detecting the number of inflow vessels in patients diagnosed with type II endoleak.

3. Methods:

3.1. Study I - Comparison of standard and dynamic computed tomography angiography in endoleak diagnosis

In this retrospective review, a total of 52 patients underwent d-CTA image acquisition after EVAR between 2019 March and 2021 July at the Cardiovascular Surgery Department of Houston Methodist Hospital. Amongst them, we selected patients with available triphasic and dynamic CTA images who had DSA acquisition in order to visualize potential endoleaks after EVAR within a three-month period and no interim interventions.

3.1.1. Qualitative image review

For the expert review of CTA images, two senior vascular imaging specialists and for the DSA image review two senior vascular surgeons were involved in the analysis of the images. Presence of endoleak, type of endoleak, and, if deemed necessary, inflow/target vessels contributing to type II endoleak were also recorded. Discrepancy between reviewers was solved by consensus.

3.2. Study II - Quantitative approach to characterize endoleak types

In this retrospective review patients undergone d-CTA and DSA image acquisition after EVAR between 2019 March and 2021 January at the Cardiovascular Surgery Department of Houston Methodist Hospital were selected (24). Qualitative analysis was the same as in Study I.

3.2.1. Quantitative analysis

Quantitative endoleak analysis being a new concept- to aid endoleak differential diagnosis based on temporal Hounsfield unit change, we selected two different regions of interest (ROI) for analysis: (1, ROI_{aorta}) in the aorta at the level of the proximal graft and (2, ROI_{endoleak}) one inside the aneurysm sac where contrast enhancement was detected. Then the software generated the corresponding time attenuation curves (TAC) for each ROI. TAC has multiple quantitative parameters that can be used for analysis, we selected the relative time difference between ROI_{aorta} and ROI_{endoleak} in reaching peak enhancement. Figure 1 demonstrates the quantitative image analysis of a time attenuation curve. Our practice focused on endoleak specific Δ time to peak values (Δ TTP) to highlight the difference in reaching peak enhancement between aorta and endoleak. In type II endoleak cases suspected target vessels can be selected for further analysis which can determine in- and outflow vessels contributing to the endoleak and aid future embolization. Quantitative analysis was performed using the dynamic CTA workflow of syngo.via (Syngo.via, VB30, Siemens Healthineers, Erlangen, Germany).

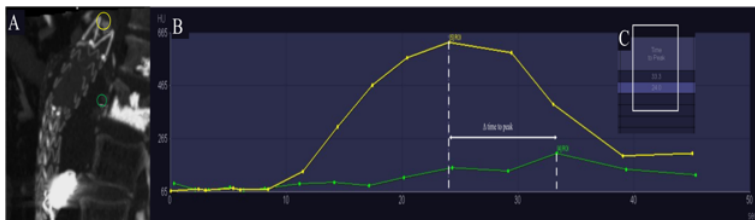


Figure 1. Example of a time attenuation curve analysis in a patient having a type II endoleak from a lumbar artery as inflow. (Fig. 1A) The selected region of interest (yellow above the stent-graft (ROI_{aorta}), green inside the aneurysm sac where endoleak is visualized (ROI_{endoleak})). Fig. 1B demonstrates the generated time-attenuation curves for the selected ROIs in Fig. 1A. Time difference between aortic and endoleak curves in reaching peak Hounsfield unit is recorded (Δ time to peak value - marked with white) (ROI=region of interest).

4. Results:

4.1. Study I - Comparison of standard and dynamic computed tomography angiography in endoleak diagnosis

From the 52 patients who underwent d-CTA acquisition in the study period, 19 met the criteria for selection. In this study population 15 out of 19 patients had conventional EVAR, 3 had fenestrated EVAR, and 1 had chimney EVAR. Mean (\pm Standard Deviation [SD]) age, serum creatinine, and body

mass index (BMI) before image acquisition were 78.8 (± 6.8) years, 97.3 (± 26.5) $\mu\text{mol/L}$, and 25.8 (± 5.8) m^2/kg , respectively.

D-CTA findings matched with DSA findings in 19 out of 19 cases (100%), while t-CTA findings matched 14/19 (73.7%). Figure 2 demonstrates the findings of the qualitative image review. Out of five patients who had a mismatch between t-CTA and DSA, two were discordant, and three were inconclusive for the type of endoleak.

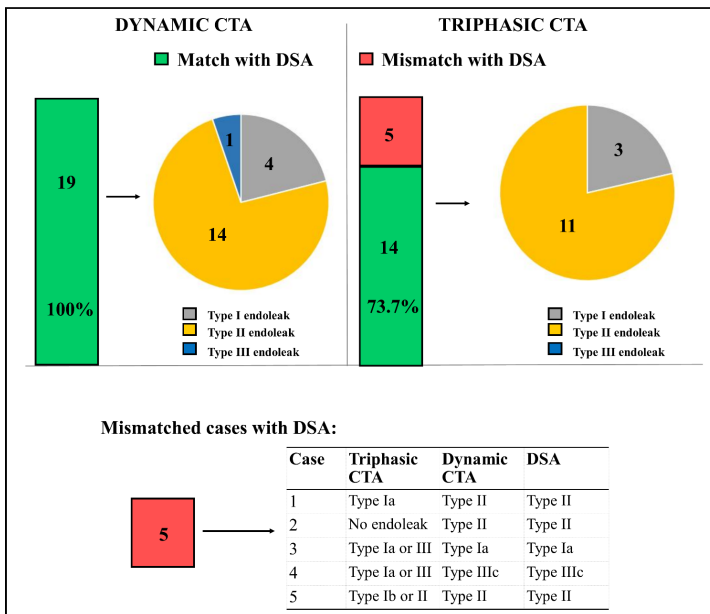


Figure 2. Findings of the qualitative image review. Dynamic CTA was compared to triphasic CTA, and DSA was used as reference standard.

In 11 patients, where type II endoleak was confirmed, the number of target arteries identified by d-CTA, t-CTA, and DSA were 23, 17, and 16 respectively ($p=0.009$), and d-CTA identified more target vessels than t-CTA ($p=0.034$).

The mean (\pm SD) dose-length product (DLP) for d-CTA and t-CTA were 1445 (\pm 550) and 1612 (\pm 530) mGy*cm, respectively ($p=0.255$). Although the DLP values showed no significant difference, it is important to emphasize the differences in acquisition protocols. Dynamic scans were acquired at 80 (interquartile range 70-97.5) kV tube voltage with the scan field of view covering the implanted stent graft only (23-33 cm), while triphasic scans were acquired at 120 kV covering the entire abdomen from diaphragm to pelvis.

4.2. Study II - Quantitative approach to characterize endoleaks

From the 24 patients 3 underwent chimney-EVAR, 3 had FEVAR, and 18 underwent conventional EVAR.

Mean(\pm SD) age, serum creatinine before d-CTA imaging, and BMI were 78(\pm 7.11) years, 99.9 (\pm 31.0) μ mol/L mg/dl, and 26.37 (\pm 5.31) m²/kg, respectively. The mean (\pm SD) dose-length product for d-CTA was 1038 (\pm 533) mGy*cm for a

total of 12 scans. Total iodinated contrast volume used for d-CTA was 77.1 (\pm 5.4) ml.

In 23/24 patients (95.8%), d-CTA findings correlated with DSA findings for endoleak type. In one patient, d-CTA demonstrated type III endoleak, whereas no evident endoleak was visible on DSA imaging (Table 1). This patient underwent an Ovation[®] (Endologix Inc., Irvine, USA) graft implantation and d-CTA imaging demonstrated the defect in one of the polymer sealing rings leading to type III endoleak, which was later relined. In the 16 cases of types II endoleak, d-CTA identified significantly more vessels (lumbar arteries) contributing to the endoleak as compared to DSA (33 vs 21, $p=0.010$).

Quantitative time-attenuation curve analysis was performed in 23/24 patients, while one patient in our study did not show any contrast enhancement in the aneurysm sac by d-CTA imaging. Amongst 23 patients, Δ TTP values (median, IQR) between ROI_{aorta} and ROI_{endoleak} was 1.64 (0.26-3.17) seconds for type I ($n=4$), 10.45 (8-12) seconds for type II ($n=16$), and 5 (4.78-6.05) seconds for type III endoleak ($n=3$), respectively ($p=0.002$) (Figure 15). Δ TTP was significantly shorter for type I ($p=0.003$) and III ($p=0.043$) endoleaks as compared to type II endoleak.

Table 1 summarizes findings on endoleak types and inflow vessels based on qualitative image review of d-CTA and DSA imaging. In type II endoleak cases, d-CTA identified more feeding vessels than DSA imaging (33 vs 21 vessels, p=0.010).

Endoleak type	Diagnostic Imaging Modality (n=24)		Inflow vessels of endoleak	
	Dynamic CTA	DSA	Dynamic CTA	DSA
I	4	4	Ia seal zone (1), Ia gutter endoleak (3)	Ia (4)
II	16	16	Lumbar a. (29) RRA (1) IMA (3)	Lumbar a. (17) RRA (1) IMA (3)
III	3	2	Component separation, graft defect, polymer sealing ring defect	Component separation, graft defect
No endoleak	1	2		

5. Conclusion:

Study I – Comparison of standard and dynamic computed tomography angiography in endoleak diagnosis

Based on our findings we conclude that dynamic computed tomography angiography imaging has better endoleak diagnosing capability as compared to standard triphasic computed tomography angiography when DSA was used as the reference standard.

There was no significant difference reported in radiation exposure between triphasic and dynamic CTA acquisitions.

In type II endoleak cases dynamic CTA identified more inflow vessels contributing to the pathomechanism of the endoleak as compared to triphasic CTA or DSA.

Study II - Quantitative approach to characterize endoleak types

Utilizing multiple CT scans, quantitative parameters such as Δ time to peak value demonstrated feasibility in objective endoleak analysis and can aid the differential diagnosis of endoleak types.

In type II endoleak cases dynamic CTA identified more inflow vessels contributing to the pathomechanism of the endoleak than DSA.

6. Bibliography of the candidate's publications:

Publications directly related to thesis:

1. **Berczeli M**, Chang SM, Lumsden AB, Chinnadurai P. Imaging aortic intramural hematoma with blood pools using time-resolved, dynamic computed tomography angiography. *J Vasc Surg.* 2022;76(4):1087-1088. **IF: 4.860**
2. **Berczeli M**, Chinnadurai P, Osztrogonácz P, Peden EK, Bavare CS, Sótonyi P, Chang SM, Lumsden AB. Dynamic Computed Tomography Angiography is More Accurate in Diagnosing Endoleaks than Standard Triphasic Computed Tomography Angiography and Enables Targeted Embolization. *Ann Vasc Surg.* 2023;88:318-326. **IF: 1.607**
3. **Berczeli M**, Chinnadurai P, Legeza P, Peden EK, Bavare CS, Chang SM, Lumsden AB. (2022) Dynamic, Time-Resolved CT Angiography After EVAR: (2022) A Quantitative Approach to Accurately Characterize Aortic Endoleak Type and Identify Inflow Vessels [published online ahead of print, 2022 Jan 23]. *J Endovasc Ther.* 2022 doi:10.1177/15266028211070970 **IF:3.089**
4. **5. Berczeli M**, Chinnadurai P, Ramirez-Giraldo JC, Garami Z, Lumsden AB, Atkins MD, Chang SM. Time-resolved, Cardiac-gated Computed Tomography After

Endovascular Ascending Aortic and Arch Repair. *Ann Thorac Surg.* 2022;113(5):1685-1691. **IF:5.113**

5. **5. Berczeli M**, Chinnadurai P, Chang SM, Lumsden AB. Time-Resolved, Dynamic Computed Tomography Angiography for Characterization of Aortic Endoleaks and Treatment Guidance via 2D-3D Fusion-Imaging. *J Vis Exp.* 2021; (178):10.3791/62958. **IF:1.424**
6. **Berczeli M**, Lumsden AB, Chang SM, Bavare CS, Chinnadurai P. Dynamic, Time-Resolved Computed Tomography Angiography Technique to Characterize Aortic Endoleak Type, Inflow and Provide Guidance for Targeted Treatment. *J Endovasc Ther.* 2022;29(1):11-22. **IF:3.089**
7. **7. Berczeli M**, Csobay-Novák C, Oláh Z, Sótónyi P. Multistage Endovascular Management of an Aortic Aneurysm Rupture into the Retroaortic Left Renal Vein. *Ann Vasc Surg.* 2021;73:509. e11-509.e14. **IF:1.607**

Publications directly not related to thesis:

1. **Berczeli M**, P Osztrogonác, L Hidi, and Z Szeberin. (2022) Percutan Endovascularis Aortarekonstrukcióval Szerzett Kezdeti Tapasztalataink. *ORVOSI HETILAP* 163 (33): 1318–1323. **IF:0.707**

2. **Berczeli M.**, P. Chinnadurai, R.G. McFall, O. Diaz, and A.B. Lumsden. (2022) Endovascular Treatment of Pancreaticoduodenal Aneurysm with Braided Stent-Assisted Coil Embolization Using Intraoperative Cone-Beam Computed Tomography Guidance. JOURNAL OF VASCULAR SURGERY CASES AND INNOVATIVE TECHNIQUES 8 (2): 265–270. **IF:-**
3. **Berczeli M** and David Garbaisz. (2022) Total Hip AND Femoral Artery Replacement. EUROPEAN JOURNAL OF VASCULAR AND ENDOVASCULAR SURGERY 2022;64(2-3):216 **IF: 6.427**
4. **Berczeli M**, Chinnadurai P, Veress DS, Diaz O, Bavare CS and Lumsden AB. (2022) Added Value of Selective Intra-Arterial Cone-Beam CT Angiography in the Management of Visceral Artery Aneurysms.” JOURNAL OF ENDOVASCULAR THERAPY In press: 152660282211185. **IF:3.089**
5. **Berczeli M**, Gavin W. Britz, Thomas Loh, and Alan B. Lumsden. (2022) Telerobotic Endovascular Interventions and Their Potential for Cerebrovascular Treatment. TEXAS HEART INSTITUTE JOURNAL 49 (2) **IF:1.103**
6. **Berczeli M**, Chinnadurai P, Legeza PT, Britz GW and Lumsden AB. (2022) Transcarotid Access for Remote

Robotic Endovascular Neurointerventions: A Cadaveric Proof-of-Concept Study. NEUROSURGICAL FOCUS 52 (1). **IF:4.332**

7. Hendricks W, Mecca J, Rahimi M, Rojo MR, Von Ballmoos MWC, McFall RG, Haddad P, **Berczeli M**, Sinha K, Barnes RG., Peden EK., Lumsden AB., MacGillivray TE, Corr S. (2022) Evaluation of a Novel System for RFID Intraoperative Cardiovascular Analytics. IEEE JOURNAL OF TRANSLATIONAL ENGINEERING IN HEALTH AND MEDICINE-JTEHM 10: 1–9. **IF:2.890**
8. Legeza, PT, Lettenberger AB, Murali B, Johnson LR, **Berczeli M**, Byrne MD, Britz G, O'Malley MK and Lumsden AB. (2022) Evaluation of Robotic-Assisted Carotid Artery Stenting in a Virtual Model Using Motion-Based Performance Metrics. JOURNAL OF ENDOVASCULAR THERAPY. **IF:3.089**
9. Sinha, K., **Berczeli M**, Lumsden AB, and Roy TL. (2022) Imaging: New Frontiers in Vascular Training. Methodist DeBakey Cardiovascular Journal 18 (3): 39–48. **IF:-**
10. Gyánó, M., **Berczeli M**, Csobay-Novák C, Szöllősi D, Óriás VI, Góg I, Kiss JP, Veres DS, Szigeti K, Osváth S, Pataki Á, Juhász V, Oláh Z, Sótónyi P, Nemes B. (2021)

Digital Variance Angiography Allows about 70% Decrease of DSA-Related Radiation Exposure in Lower Limb X-Ray Angiography. SCIENTIFIC REPORTS 11 (1). **IF:4.997**

11. Patelis N, Bisdas T, Jing Z, Feng J, Trenner M, Tri Nugroho N, Ocke Reis PE, Elkouri S, Lecis A, Karam L, Roux DL, Ionac M, **Berczeli M**, Jongkind V, Yeung KK, Katsargyris A, Avgerinos E, Moris D, Choong A, Ng JJ, Cvjetko I, Antoniou GA, Ghibu P, Svetlikov A, Pedrajas FG, Ebben H, Stepak H, Chornuy A, Kostiv S, Ancetti S, Tadayon N, Mekkar A, Magnitskiy L, Fidalgo-Domingos L, Matheiken S, Sarutte Rosello ES, Isik A, Kirkilesis G, Kakavia K, Georgopoulos S. (2021) Dataset of the Vascular E-Learning during the COVID-19 Pandemic (EL-COVID) Survey. DATA IN BRIEF 38. **IF:-**
12. Patelis N, Bisdas T, Jing Z, Feng J, Trenner M, Tri Nugroho N, Ocke Reis PE, Elkouri S, Lecis A, Karam L, Roux DL, Ionac M, **Berczeli M**, Jongkind V, Yeung KK, Katsargyris A, Avgerinos E, Moris D, Choong A, Ng JJ, Cvjetko I, Antoniou GA, Ghibu P, Svetlikov A, Pedrajas FG, Ebben H, Stepak H, Chornuy A, Kostiv S, Ancetti S, Tadayon N, Mekkar A, Magnitskiy L, Fidalgo-Domingos L, Matheiken S, Sarutte Rosello ES, Isik A, Kirkilesis G, Kakavia K, Georgopoulos S (2021) Vascular E-Learning

During the COVID-19 Pandemic: The EL-COVID Survey. ANNALS OF VASCULAR SURGERY 77: 63–70. **IF:1.607**

13. **Berczeli M**, Oláh Z, Szatai L, Daróczy L, Sótonyi P. (2020) Rupturált Óriás Thoracoabdominalis Aortaaneurysma Sikeres Kezelése Két Lépésben = Successful Two-Step Treatment of a Ruptured Giant Thoracoabdominal Aortic Aneurysm. ORVOSI HETILAP 161 (7): 269–274. **IF:0.707**
14. Gyánó, M, Csobay-Novák C, **Berczeli M**, Góg I, Kiss JP, Szigeti K, Osváth Sz, and Nemes B. (2020) Initial Operating Room Experience with Digital Variance Angiography in Carbon Dioxide-Assisted Lower Limb Interventions: A Pilot Study. CARDIOVASCULAR AND INTERVENTIONAL RADIOLOGY 43 (8): **IF:2.797**
15. Pál, D, Szilágyi B, **Berczeli M**, Szalay CsI, Sárdy B, Oláh Z, Székely T, Rácz G, Banga P, Czinege Z, Sótonyi P. (2020) Ruptured Aortic Aneurysm and Dissection Related Death: An Autopsy Database Analysis. PATHOLOGY AND ONCOLOGY RESEARCH 26 (4): 2391–2399. **IF:2.874**
16. Szilágyi B, **Berczeli M**, Lovas A, Oláh Z, Törő K, Sótonyi P. (2020) The Effects of Changing Meteorological

Parameters on Fatal Aortic Catastrophes. BMC
CARDIOVASCULAR DISORDERS 20 (1). **IF:2.174**

17. Tóth-Vajna Z, Tóth-Vajna G, Gombos Z, Szilágyi B, Járαι Z, **Berczeli M**, Sótonyi P. (2019) Screening of Peripheral Arterial Disease in Primary Health Care. VASCULAR HEALTH AND RISK MANAGEMENT 15: 355–363.
IF:-

18. Berczeli M, Szilágyi B, Lovas A, Pál D, Oláh Z, Törő K, Sótonyi P. (2018) Meteorológiai Paraméterek Változásának Hatása a Halálos Kimenetelű Aortaaneurysma-Rupturákra. ORVOSI HETILAP 159 (37): 1501–1505. **IF:0.707**

19. Nagy Z, Oláh Z, Kókai J, Molnár AB, Laczkó Á, Szabó GV, Juhász V, Garbaisz D, **Berczeli M**, Sztupinszky Z, Szeberin Z (2017) Homograftok Szerepe Az Alsó Végtagi Érrekonstrukciókban. MAGYAR SEBÉSZET 70 (1): 5–
12. **IF:-**