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Title: COLONIC ISCHEMIA AFTER STANDARD ENDOVASCULAR ABDOMINAL AORTIC ANEURYSM REPAIR, A RARE BUT DANGEROUS COMPLICATION

Annals of Vascular Surgery

Dear Dr. Berchiolli.

I am pleased to inform you that your paper "COLONIC ISCHEMIA AFTER STANDARD ENDOVASCULAR ABDOMINAL AORTIC ANEURYSM REPAIR, A RARE BUT DANGEROUS COMPLICATION" has been accepted for publication in Annals of Vascular Surgery

It is accepted with the understanding that the contents have not been published elsewhere, and is subject to minor editorial changes. When editing is complete galley proofs will be available for your attention online, an email notification will be sent to you with instructions on how to access them.

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Yours sincerely, Prof. O. Goeau-Brissonniere Editor Annals of Vascular Surgery

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COLONIC ISCHEMIA AFTER STANDARD ENDOVASCULAR ABDOMINAL AORTIC ANEURYSM REPAIR, A RARE 1 2 **BUT DANGEROUS COMPLICATION** 3 4 Berchiolli R, Adami D, Marconi M, Mari M, Puta B, Ferrari M 5 Vascular Surgery Unit, Cardio Thoracic and Vascular department, A.O.U.P., University of Pisa, Pisa, Italy 6 7 Author names and affiliations: 8 Berchiolli Raffaella, MD, Vascular Surgery Unit, University of Pisa 9 Adami Daniele, MD, Vascular Surgery Unit, University of Pisa 10 Marconi Michele, MD, PhD, Vascular Surgery Unit, University of Pisa 11 Mari Marta, MD, Vascular Surgery Unit, University of Pisa 12 Puta Besjona, MD, Vascular Surgery Unit, University of Pisa Ferrari Mauro, MD, Full Professor, Vascular Surgery Unit, University of Pisa 13 14 15 Corresponding author: 16 Berchiolli Raffaella, MD, Vascular Surgery Unit, University of Pisa 17 r.berchiolli@ao-pisa.toscana.it 18 tel. 0039050995498 19

20	ABSTRACT
21	Colonic Ischemia (CI) after abdominal aortic aneurysm (AAA) repair, although rare, is associated with severe
22	prognosis. Endovascular Aneurysm Repair (EVAR) is becoming the standard of practice in most vascular
23	centers, and it also may reduce CI incidence in comparison to conventional open repair.
24	We report 2 cases of fatal CI after 636 standard EVAR procedures performed in our institution, from
25	January 1998 to December 2017. Both patients were electively treated by high skilled operators.
26	In one patient, presenting early CI, EVAR procedure was complicated by intraoperative common iliac artery
27	rupture. The other one, presenting CI in 7th postoperative day, had an history of previous left
28	hemicolectomy. In both patients, CI with leakage of fecal material in the abdominal cavity was confirmed by
29	surgical exploration.
30	Only few cases of CI after EVAR have been reported in Literature, and the etiology of this complication
31	remains uncertain. While saving the inferior mesenteric artery is almost impossible during standard EVAR,
32	the preservation of hypogastric arteries could play an important role, especially after colonic surgery, but
33	other factors should be considered. Our preliminary, although limited experience, seem to suggest that in
34	CI developing, intraoperative persistent hypotension and hypogastric branches distal embolization have
35	both a role that should be better addressed.
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37	INTRODUCTION
38	Endovascular repair of abdominal aortic aneurysms (EVAR) has proven to be an effective alternative to
39	traditional open repair (OR) in properly selected abdominal aortic aneurysm (AAA) patients, by reducing
40	postoperative morbidity and hospital stay, and by ensuring a faster postoperative recovery [1].
41	Colonic Ischemia (CI), although rare, is a known and life-threatening complication after both OR and EVAR.
42	CI Incidence is similar between the two treatment modalities, ranging from 1 to 3% after OR (10% in
43	emergent cases), and from 1.5 to 3% after EVAR [2,3].
44	Several possible physio-pathological mechanisms were suspected to contribute to CI, including non-

occlusive visceral ischemia due to haemorrhagic shock or vasopressor drugs, inferior mesenteric artery

(IMA) and hypogastric arteries (HA) occlusion, intraprocedural embolization, and previous colonic surgery

We report 2 cases of CI after standard elective EVAR performed at our center.

CASE 1

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On April 2011, an 85-year-old man underwent scheduled duplex-ultrasound scan (DUS) showing a non-51 ruptured AAA (maximum diameter, 58 mm). The patient was referred to our unit for AAA treatment. During 52 hospitalization, preoperative computed tomographic angiography (CTA) confirmed the AAA. Patient 53 54 presented a 25 mm proximal aortic neck of 25 mm in length with thrombosis. Iliac vessels were angulated and highly calcified on both sides (Figure 1). Despite all those anatomical features challenging for EVAR, 55 patient was judged unfit for open repair because of a medical history positive for coronary arteries disease 56 57 with severe left ventricular dysfunction (ejection fraction 25%). Consequently, an EVAR procedure was carried out under epidural anesthesia. After surgical femoral access, 58 59 a bifurcated standard endograft (Zenith LP main body ZALB-28-84; left extension ZALL-16-48; right extension ZALL-16-48; Cook Medical Inc - Bloomington, IN, USA) was implanted. Devices progression 60 61 through patient's anatomy was difficult due to the extensively diseased iliac vessels. During controlateral limb shaft retrieval, arterial pressure dropped down under 70mmHg. Aorto-iliac angiography showed 62 complete left common iliac artery disruption with contrast extravasation. Patient was immediately treated 63 by HA embolization (four IDCTM Interlocking Detachable Coils 8 x 20 mm; Boston Scientific - Marlborough, 64 MA, USA) and endografts extension in to the external iliac artery (Zenith LP ZALL-16-60; Cook Medical Inc -65 Bloomington, IN, USA). Completion angiography showed a satisfactory result with complete AAA exclusion, 66 patency of right HA and both renal arteries, without endoleak or contrast medium extravasation. Estimated 67 68 intraoperative blood loss was 1000ml. After EVAR patient was recovered in intensive care unit. Twenty-four hours after EVAR, patient experienced 69 severe abdominal pain, associated with increase in lactate (15 mmol/l, preoperative 2 mmol/l) and 70 creatinine (2,6 mg/dl, preoperative 1,8 mg/dl) serum levels. Immediate laparotomy was carried out 71 72 showing severe CI, and leakage of faecal material. Despite successful left colectomy and colostomy, patient 73 died in fourth post-operative day for multi-organ failure (MOF).

CASE 2

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79 80 On April 2008, an 84-year-old man was referred to our unit after non-ruptured AAA was found during CTA (maximum diameter, 61 mm). The AAA presented a growth of 7 mm in 6 months. He had a past history of chronic obstructive pulmonary disease (COPD) with reduced FEV1/FVC ratio (40% on pre-assessment), atrial fibrillation in oral anticoagulant therapy and severe left ventricular dysfunction (ejection fraction 29%), previous open left colectomy for cancer.

AAA presented a 27 mm proximal aortic neck of 21 mm in length. Iliac vessels were highly calcified on both sides. Patient was judged unfit for open repair because of comorbidities and hostile abdomen.

An EVAR procedure was carried out under general anaesthesia. After surgical femoral access, a bifurcated standard endograft (Zenith Flex main body TFB-1-30; left extension TFLE-18-71; right extension TFLE-22-37; Cook Medical Inc - Bloomington, IN, USA) was implanted. Completion angiography showed a satisfactory result with complete AAA exclusion, patency of both hypogastric and renal arteries, without endoleak or contrast medium extravasation. Estimated intraoperative blood loss was 250 ml. After EVAR patient was recovered in normal ward. In 7th postoperative day after EVAR, patient experienced abdominal pain, diarrhoea and High White Blood Cell Count (25 \times 109/l, preoperative 8 \times 109/l), associated with increase in lactate (13 mmol/l, preoperative 1,6 mmol/l) serum levels. Sigmoidoscopy and subsequent laparotomy confirmed CI and leakage of faecal material. After total colectomy and colostomy, patient died in tenth post-operative day for multi-organ failure (MOF).

DISCUSSION

EVAR has gained acceptance over OR and multiple studies have documented the benefits of EVAR because of its minimal invasiveness [1]. Despite a relatively low incidence ranging from 1% to 3%, CI remains a rare but potential life-threatening complication following AAA repair [2,6].

Relatively small reports showed a CI incidence similar for both, EVAR and OR [2,7]. In present series the rate of post-EVAR CI was found to be 0.3% (2/636 pts), in line with previous reported results, while incidence of CI after OR was 0.6% [1,7,8]. Despite its very low incidence in our series, CI was associated with a hundred percent of emergent colectomy and exitus.

Dadian et al reported a 2.9% incidence of CI in a series of 278 patients treated by EVAR [7]. Out of eight patients presenting CI in their series four required colectomy, and three died after surgery.

Presumed causes of CI after AAA repair are several, including HAs and IMA artery occlusion, micro-embolization, haemorrhagic shock and hypoperfusion [1,7,8,9,10,11]. However, CI frequently occurs even with no single identifiable risk factors or when multiple possible causes are simultaneously present, making its occurrence difficult to predict.

A potential role of HA occlusion in CI development has been debated [1]. Indeed, after HA embolization, the most frequent complication is buttock claudication. The reported risk ranges from 11% to 50% after single or bilateral, HA coil or plug embolization. While CI appears to be uncommon after HA(s) embolization [9,12,13,14].

Mehta et al. reported 107 patients were treated by EVAR and HA occlusion, and no patient presented CI requiring laparotomy. Therefore, they concluded that HA embolization could be performed safely, without significant risk of severe pelvic ischemia [4]. Despite those encouraging results, other authors reported a higher incidence of CI associated with HA occlusion and emphasized the importance of preserving the hypogastric circulation during AAA repair [2,7]. Branched stent grafts aiming to maintain anterograde perfusion of hypogastric arteries are currently under clinical evaluation. Early reports demonstrated encouraging short-term results. Unfortunately, the use of these devices is limited to favourable anatomic conditions, which are not frequently encountered [15]. In present series, however, only one patient developed CI after mono-lateral HA occlusion, while 77 single HA exclusion were performed in absence of complications. CI developed in the only patient who presented a severe intraoperative blood loss, could be the latter a potential risk factor by itself. HA exclusion associated with severe intraoperative blood loss (Table I), a potential risk factor by itself [16]. So, it should be admitted that CI development a more complex, and probably multifactorial pathogenesis. Another, theoretically, important predictor of CI is IMA interruption [1,2]. During EVAR, however, IMA occlusion is an unavoidable part of the AAA exclusion procedure. Moreover, Miller et al reported a 10-year series in which 10 out 11 patients presenting post-operative CI had an occluded IMA before EVAR [8]. This suggests that preserving IMA patency could not offer real advantage to the patients. Similar results were found in a randomized study on patient submitted to OR [10]. Indeed, inadequate mesenteric collateral vessels more than IMA patency by itself, either as a result of intrinsic disease (preoperative occlusion of the large arteries to the digestive system, such as superior mesenteric artery and/or celiac axis) or iatrogenic (e.g. after colectomy), might be predisposing factors for Cl. According with this theory, Maldonado and Dadian reported previous colectomy as potential risk factor for CI development. Also in our experience, a patient without other recognized risk factor but previous colectomy, suffered fatal CI (Table I) [1,7]. Lastly, IMA and HA distal embolization has been recognized as potential cause of CI after EVAR in many studies [7,8,11]. Dadian documented atheroemboli in pathologic specimens from 4 of 8 patients with CI after EVAR [7]. During EVAR, indeed, difficult devices navigation in severe narrowed and calcified arteries, especially in presence of extensive thrombosis, may lead to an increased risk of embolization [11]. Moreover, a recent paper by Mansour et al showed a greater risk distal embolization in case of HA, when using coils rather than plugs [17]. In our experience, the role of distal embolization was not fully investigated, but both patients had a thrombus in the AAA sac and delivery of the endografts was difficult and time-consuming. Moreover, one patients had unplanned coils HA embolization. From a speculative

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point of view, we should admit that embolization could be recognized as important CI developing factor at least in that patient (Table I).

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CONCLUSION

- Although rare, CI still remains a serious complication following AAA repair even in the endovascular era.
- 150 Several possible physio-pathological mechanisms were suspected to contribute to CI, including non
- occlusive visceral ischemia due to haemorrhagic shock or vasopressor drugs, IMA and HA occlusion,
- intraprocedural embolization, and previous colonic surgery [7,8].
- Larger studies with longer follow-up could confirm our hypothesis.

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193	TABLE LEGEND
194	Table 1. Demographic, anatomical and intra-procedural details of patients presenting CI after EVAR
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196	FIGURE LEGEND
197	Figure 1. Computed tomographic angiography case 1
198	Figure 2. Computed tomographic angiography case 2

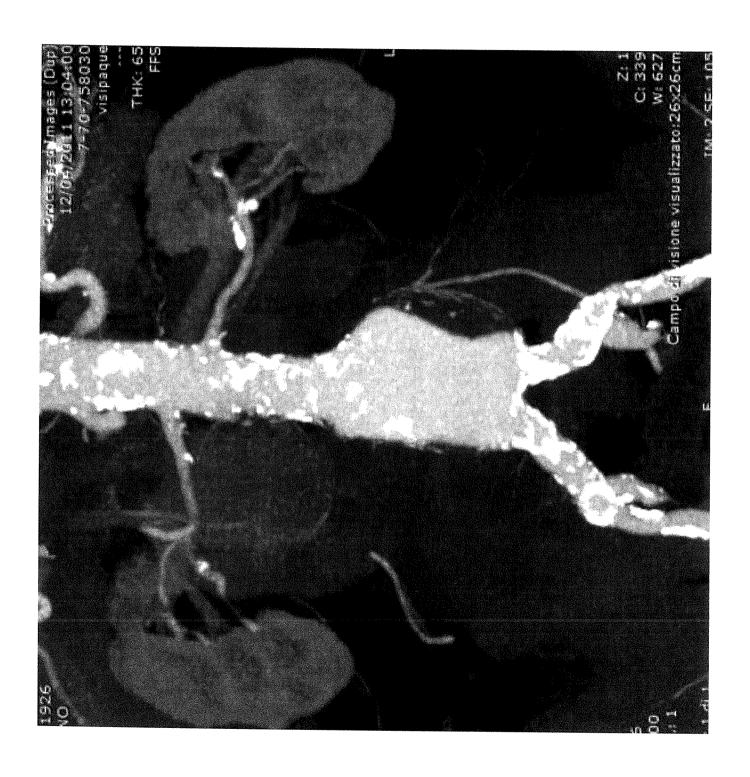


Figure 2

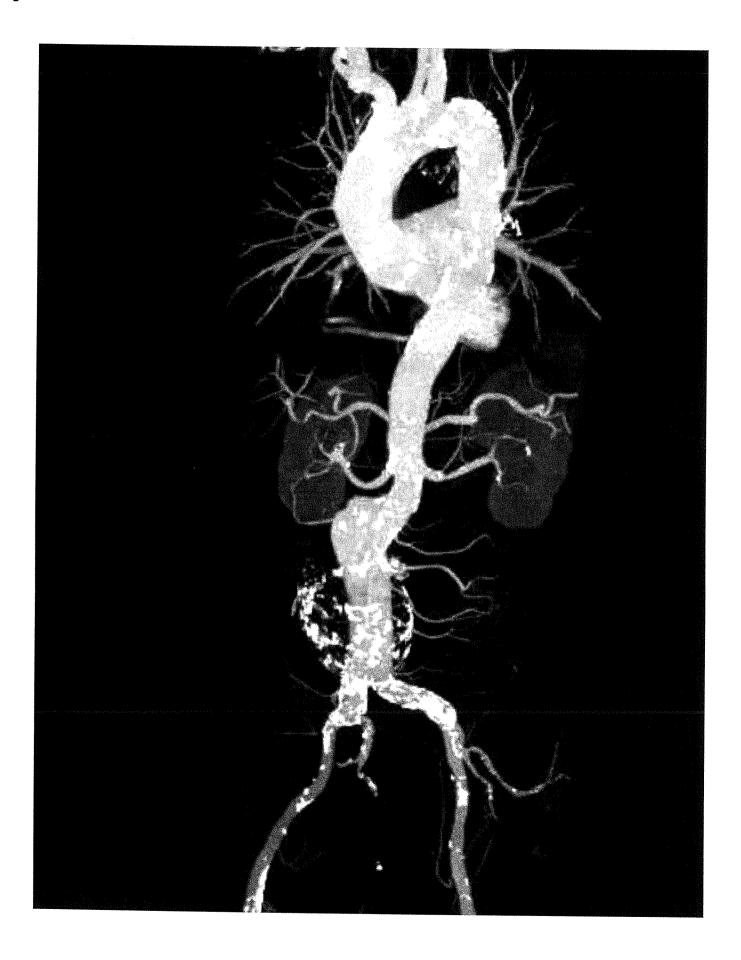


Table I Demographic, anatomical and intra-procedural details of patients presenting CI after EVAR

			M		·				Intraoperative		
		Previous	Aortic Neck	Iliac	MA	200			Sever	Estimated	:
Age	Gender	Colonic Surgery	Thrombosis	Vessels Thrombosis	>	۵	Graft	Procedural Time (min)	Hypotension (SAP	Blood Loss (ml)	Adjunctive procedures
									<70mmHg)		
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)	Colectomy	2	<u>S</u>	ກັກ	tes	Zenith	180	o N	250	None