

REVIEW

# Arterial Stiffness and Aortic Aneurysmal Disease – A Narrative Review

Konstantinos G Moulakakis (1)<sup>1</sup>,\*, Christos F Pitros (1)<sup>1</sup>,\*, Ioannis T Theodosopoulos<sup>2</sup>, Spyridon N Mylonas (1)<sup>3</sup>, John D Kakisis<sup>2</sup>, Christos Manopoulos<sup>4</sup>, Nikolaos PE Kadoglou<sup>5</sup>

<sup>1</sup>Department of Vascular Surgery, Patras University Hospital, University of Patras, Patras, Greece; <sup>2</sup>Department of Vascular Surgery, "Attikon" University Hospital, School of Medicine, National and Kapodistrian University of Athens, Athens, Greece; <sup>3</sup>Department of Vascular and Endovascular Surgery, Faculty of Medicine and University Hospital of Cologne, University of Cologne, Cologne, 50937, Germany; <sup>4</sup>Biofluid Mechanics and Biomedical Engineering Laboratory, Fluids Section, School of Mechanical Engineering, National Technical University of Athens, Athens, Greece; <sup>5</sup>Medical School, University of Cyprus, Nicosia, Cyprus

Correspondence: Konstantinos G Moulakakis, Associate Professor of Vascular Surgery, Vascular Surgery Department, Patras University Hospital, Patras, Greece, Tel +0030 6937357508, Email konmoulakakis@yahoo.gr

Abstract: It has been documented that large-artery stiffness is independently associated with increased cardiovascular risk and may potentially lead to heart and kidney failure and cerebrovascular disease. A systematic review of studies investigating changes in arterial stiffness in patients undergoing endovascular repair of aortic disease was conducted. In addition, a review of the available literature was performed, analyzing findings from studies using the cardio-ankle vascular index (CAVI) as a marker of arterial stiffness. Overall, 26 studies were included in the present analysis. Our research revealed a high heterogeneity of included studies regarding the techniques used to assess the aortic stiffness. Aortic stiffness was assessed by pulse wave velocity (PWV), elastic modulus (Ep), and augmentation index (AI). Currently a few studies exist investigating the role of CAVI in patients having an aortic aneurysm or undergoing endovascular aortic repair. The majority of studies showed that the treatment of an abdominal aortic aneurysm (AAA) either with open repair (OR) or endovascular aortic repair (EVAR) reduces aortic compliance significantly. Whether EVAR reconstruction might contribute a higher effect on arterial stiffness compared to OR needs further focused research. An increase of arterial stiffness was uniformly observed in studies investigating patients following thoracic endovascular aortic repair (TEVAR), and the effect was more pronounced in young patients. The effects of increased arterial stiffness after EVAR and TEVAR on the heart and the central hemodynamic, and an eventual effect on cardiac systolic function, need to be further investigated and evaluated in large studies and special groups of patients.

**Keywords:** arterial stiffness, pulse wave velocity, PWV, cardio-ankle vascular index, CAVI, augmentation index, AI, aortic aneurysm, endovascular aortic aneurysm repair, EVAR, thoracic endovascular aortic aneurysm repair, TEVAR

#### Introduction

It is well known and documented that large-artery stiffness increases with age, atherosclerosis, and in certain disease states, and is independently associated with increased cardiovascular risk.<sup>1,2</sup> Large-artery stiffening plays a significant role in hemodynamic dysfunction characterized by excess pulsatility and may potentially lead to heart and kidney failure and cerebrovascular disease.<sup>1</sup> Moreover, guidelines from the Japanese Society of Hypertension state that an increase of pulse wave velocity (PWV) is an indicator of organ damage and could be used for further evaluation of risk assessment.<sup>2</sup>

Several methods have been suggested to assess arterial stiffness, such as pressure-strain elastic modulus (Ep), stiffness parameter ( $\beta$ ), pulse wave velocity (PWV), and vascular compliance (Cv). PWV is regarded as the simplest and most widely applied technique.<sup>3–7</sup> Cardio-ankle vascular index (CAVI) is a technique used to assess the PWV, based on the stiffness parameter  $\beta$ . CAVI reflects the stiffness of the entire aorta and leg arteries (from the ascending segment to the tibial arteries) and has the advantage of being less affected by blood pressure at the time of measurement.<sup>8</sup> However, it was recently

<sup>\*</sup>These authors contributed equally to this work

suggested that CAVI is still intrinsically affected by blood pressure, leading to the introduction of CAVI0. It is believed to be effective as a CAVI enhancement in improving the pressure-independent evaluation of arterial stiffness.<sup>9</sup>

In recent decades, the endovascular repair for the treatment of thoracic (TEVAR) and abdominal aortic aneurysms (EVAR) has emerged as a competitive alternative to open surgery in both acute and elective settings, and is often the approach of choice for high-risk patients.<sup>10</sup> The currently available stent-grafts used for aortic aneurysm exclusion consist of a metallic skeleton (stainless steel, nitinol) and a fabric cover (ePTFE, polyester). Clinical studies focusing on measuring and documenting the variations in arterial stiffness in patients with thoracic or abdominal aortic aneurysms, the impact of the treatment option, as well as the association with cardiovascular outcomes and events are scarce. The aim of the present study is a systematic presentation of the currently available literature focusing on: (1) Studies assessing arterial stiffness in patients with abdominal aortic aneurysm (AAA) and the impact of treatment option on its changes; and (2) Studies assessing arterial stiffness before and after thoracic endovascular aneurysm repair (TEVAR).

#### **Materials and Methods**

## Search Strategy

Studies were identified by searching electronic databases and scanning bibliographic references of articles. The National Library of Medicine's Medline database was searched using the PubMed interface and SCOPUS from 1992 to 2023. No language constraints were used. The last search was run on July 20, 2023. Keywords were selected using medical subject headings (MeSH) for PubMed and MeSH/Emtree for Scopus. The keywords "arterial stiffness", "pulse wave velocity", "PWV", and "CAVI" "cardio-ankle vascular index" were combined with "AAA", "TAAA", "aortic aneurysm", "EVAR", and "TEVAR". The databases were searched with an unrestricted search strategy, applying exploded MeSH and keywords combined with the Boolean operators AND or OR to retrieve relevant reports as reported in Table 1. A second-level search included a manual screen of the reference lists of the articles identified through the electronic search. Eligibility assessment was performed independently in an unblinded standardized manner by 2 reviewers; disagreements between reviewers were resolved by consensus.

#### Results

The search identified 431 records in total after application of the inclusion criteria and the control for duplicates. The literature search strategy is outlined in a study flow diagram (Figure 1). A total of 26 studies fulfilled the inclusion criteria. The selected studies were published between 2004 and 2023, reflecting study periods extending from 2002 to 2022.

Table I	Search Strategy	and Medical Su	oject Headings	(MeSH)	) Used for the Analysis

	Search Strategy	Results				
	MEDLINE (PubMed)					
I	("CAVI"[All Fields] OR "cardio-ankle vascular index"[All Fields] OR "PWV"[All Fields] OR "pulse-wave velocity"[All Fields] OR ("vascular stiffness"[MeSH Terms] OR ("vascular"[All Fields] AND "stiffness"[All Fields]) OR "vascular stiffness"[All Fields]) OR "arterial stiffness"[All Fields])	23,793				
2	("TAAA"[All Fields] OR "AAA"[All Fields] OR ("aortic aneurysm"[MeSH Terms] OR ("aortic"[All Fields] AND "aneurysm"[All Fields]) OR "aortic aneurysm"[All Fields]) OR "EVAR"[All Fields] OR ("endovascular aneurysm repair"[MeSH Terms] OR ("endovascular"[All Fields] AND "aneurysm"[All Fields] AND "repair"[All Fields]) OR "endovascular aneurysm repair"[All Fields] OR "tevar"[All Fields] OR "tevars"[All Fields])	90,242				
3	#I AND #2	407				
	SCOPUS					
TITLE	TITLE-ABS-KEY: (PWV OR pulse-wave velocity OR CAVI OR cardio-ankle vascular index OR arterial stiffness) AND (aortic aneurysm)					

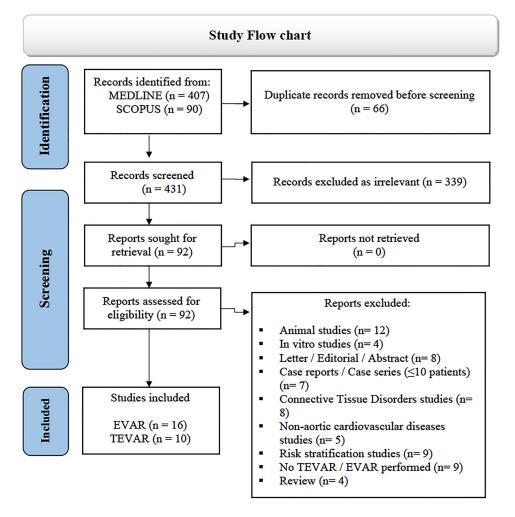


Figure I Study flow chart.

Aortic stiffness was assessed by pulse wave velocity (PWV). The research revealed, however, heterogeneity among studies regarding the techniques used. Various levels of PWV measurement were used; between carotid and femoral (carotid-femoral), carotid and brachial (carotid-brachial), carotid and radial (carotid-radial), ankle and brachial (brachial-ankle), and heart central and carotid. CAVI was investigated in only a few studies in patients with an aortic aneurysm.

## Studies Assessing Arterial Stiffness in Patients with AAA, Before and After Treatment

The review identified 16 studies (Table 2) reporting the influence of abdominal endografts, following EVAR, on arterial stiffness. 5,11-25 Development of an aortic aneurysm could affect the arterial stiffness resulting in increased cardiovascular risk. Aykan et al investigated arterial stiffness by measuring the CAVI, in a cross-sectional study involving 59 subjects with abdominal aortic aneurysm (AAA) and 32 healthy subjects. The CAVI was significantly higher in AAA patients than in controls, positively correlated with AAA diameter, and negatively correlated with left ventricular ejection fraction. Moreover, in a recent original study CAVI values obtained from patients with AAA were found to be statistically higher than in control subjects (healthy adults, adjusted for age and gender). 14

Regarding the changes in arterial stiffness associated with AAA treatment, it has been described that EVAR is related to increased arterial stiffness as a result of the endograft implantation.<sup>5,21,22</sup> When compared to healthy controls, patients undergoing EVAR seem to have higher measured levels of PWV<sup>5</sup> and AI.<sup>21</sup> Sekhri et al, in 2004, were the first to correlate endograft implantation for aortic aneurysm repair to aortic stiffness.<sup>11</sup> In this pilot study, of a 38-patient cohort, it was shown that the treatment of the aortic aneurysm with open or endovascular technique reduces aortic compliance significantly. Also,

Moulakakis et al Dovepress

 Table 2 Studies Assessing Arterial Stiffness in Patients with AAA, Before and After Treatment

Author, Publication Year	Total N	Subgroups (n)	Groups of Interest	Techniques for Arterial Stiffness Assessment	Adjunct meas.	Conclusions
Sekhri et al 2004 <sup>11</sup>	38	Pre-repair (20) Post-OR (6) Post-EVAR (18) Endoleaks (6) No endoleaks (12)	OR vs EVAR, pre- and post-EVAR Endoleaks vs no endoleaks	Ep (elastic modulus) β (stiffness)	NR	<ul> <li>Ep and beta were significantly higher in successful EVAR than in open repair</li> <li>A massive increase in Ep and β was observed in the presence of detectable endoleak.</li> <li>Sacs with endoleaks were significantly less compliant</li> </ul>
Van Herwaarden et al 2006 <sup>18</sup>	11	EVAR (II)	Pre- and post-EVAR	Dynamic MRA Ep, β	NR	<ul> <li>EVAR results in increased aneurysm sac Ep and β</li> <li>Stent-graft design seems to alter Ep and β within the aneurysm neck</li> <li>The presence of an endoleak does not seem to have an effect on Ep or β</li> </ul>
Lantelme et al 2009 <sup>19</sup>	50	OR (39) EVAR (11)	OR vs EVAR	Pulse wave velocity (PWV) (carotid-radial and carotid- femoral) Al	NR	Al increased after stent-graft implantation and decreased after graft-prosthesis placement     Stent-grafts (EVAR) increase reflected waves more than graft-prostheses (OR)
Moloney et al 2011 <sup>20</sup>	19	OR (12) EVAR (7)	OR vs EVAR	Applanation tonometry laser Doppler fluximetry (LDF) Al	NR	Aortic grafting resulted in a decreased AI
Kadoglou et al 2012 <sup>5</sup>	81	EVAR (48) Controls (31)	EVAR vs controls Pre- and post-EVAR	PWV (carotid- femoral)	Biomarkers	Patients with AAA present with significantly elevated PWV levels compared to controls     EVAR was related to an increase in PWV
Lee et al 2013 <sup>21</sup>	102	EVAR (51) Controls (51)	EVAR vs controls Pre- and post-EVAR	PWV (carotid- femoral, brachial-ankle, carotid- brachial)	Wave reflection (carotid AI (cAI), carotid augmentation pressure, backward pressure wave (Pb), forward pressure wave)	<ul> <li>ba-PWV is less likely influenced by the presence of AAA</li> <li>AAA does not increase but may actually slightly decrease the magnitude of wave reflections at the central aorta</li> <li>Pb is less likely influenced by the presence of AAA than cAI</li> <li>Stent-graft might slightly increase wave reflections by inducing a positive reflection</li> <li>EVAR may correct the impacts of AAA on cf-PWV and cAI</li> </ul>

(Continued)

Table 2 (Continued).

Author, Publication Year	Total N	Subgroups (n)	Groups of Interest	Techniques for Arterial Stiffness Assessment	Adjunct meas.	Conclusions
Kadoglou et al 2014 <sup>22</sup>	124	EVAR (118) - PTFE (46) - Polyester (72)	Pre- and post-EVAR PTFE vs polyester fabric	PWV (carotid- femoral)	Biomarkers	Arterial stiffness was positively associated with mean blood pressure, age, OPG, and AAA size     Endograft implantation seemed to further increase PWV     The latter effect was independently associated with the polyester endograft type and accompanied by deteriorated inflammatory response
Takeda et al 2014 <sup>23</sup>	40	EVAR (30) TEVAR (10)	Pre- and post-EVAR Pre- and post-TEVAR	PWV (brachial- ankle)	Echocardiography	EVAR raised aortic vascular stiffness, induced LV hypertrophy, and impaired LV diastolic function without an elevation of BP in the short term, and I year after     Low LV distensibility at baseline may be related to the impairment of exercise tolerance after EVAR
Gray et al 2016 <sup>24</sup>	34	OR (15) EVAR (19)	OR vs EVAR	PWV (carotid- femoral)	NR	EVAR patients have a significantly higher postopera- tive PWV than those undergoing OR
Hori et al 2019 <sup>25</sup>	25	EVAR (25)	Pre- and post-EVAR	PWV (brachial- ankle)	NR	High postoperative PWV was independently associated with sac enlargement, whereas low preoperative PWV was asso- ciated with sac shrinkage
Valdivia et al 2019 <sup>12</sup>	44	OR (19) EVAR (25)	OR vs EVAR	PWV (carotid- radial) Al Central aortic pressure	Biomarkers	No significant changes in PWV and central blood pressures
Marketou et al 2021 <sup>13</sup>	73	OR (12) EVAR (61)	OR vs EVAR	PWV (carotid- femoral)	Echocardiography – GLS (global longitudinal strain)	Both EVAR and open AAA repair are followed by an immediate deterioration of LV GLS that is related to an increase in aortic stiffness     AAA repair leads not only to an increase in aortic stiffness, as measured by the increase in PWV, but also to reduced cardiac systolic function

(Continued)

Moulakakis et al Dovepress

Table 2 (Continued).

Author, Publication Year	Total N	Subgroups (n)	Groups of Interest	Techniques for Arterial Stiffness Assessment	Adjunct meas.	Conclusions
Mylonas et al 2021 <sup>14</sup>	213	AAA (110) - OR (54) - EVAR (56) Controls (103)	AAA vs controls OR vs EVAR	Cardio-ankle vascular index (CAVI)	NR	Increased PWV, expressed with CAVI values, for patients with an AAA. Significant increase of arterial stiffness in EVAR and open repair, 48 h postoperatively Significant increase of arterial stiffness at 6 months in EVAR patients compared to the baseline
Holewijn et al 2021 <sup>15</sup>	20	EVAR (20)	Pre- and post-EVAR	PWV (carotid- femoral) Al subendocardial viability ratio (SEVR)	NR	EVAR caused an increase in PWV compared with baseline, which remained elevated through I-year follow-up     No differences in central pressure, AI, and SEVR were observed during follow-up
Nishibe et al 2021 <sup>16</sup>	119	EVAR (119)	Pre- and post-EVAR	PWV (brachial- ankle)	NR	Preoperative PWV was indepen- dently associated with sac shrinkage after EVAR
Ugajin et al 2022 <sup>17</sup>	222	EVAR (222)	Pre- and post-EVAR	PWV (brachial- ankle)	NR	PWV before EVAR was asso- ciated with significant sac growth after EVAR

Abbreviations: OR, open repair; PWV, pulse wave velocity; Al, augmentation index; NR, not reported.

EVAR appeared to cause significantly higher levels of elastic modulus (Ep) and aortic stiffness ( $\beta$ ) compared to OR. Moreover, when EVAR was associated with the presence of endoleak a massive increase was reported in Ep and  $\beta$ , leading to the conclusion that sacs with endoleak were significantly less compliant. A few years later, in 2006, a study utilizing dynamic magnetic resonance angiography (MRA) for arterial stiffness measurement reported that the presence of an endoleak would not affect Ep or  $\beta$ . Nevertheless, these studies agreed that EVAR results in Ep and  $\beta$  increase at the level of the aneurysmal sac.

Increased aortic vascular stiffness after EVAR, as measured by PWV, was also reported in a 2014 study that additionally documented an induced left ventricular hypertrophy and impaired diastolic dysfunction.<sup>23</sup> Holewijn et al highlighted the cardiovascular risk due to PWV increase after EVAR compared to baseline, which remained increased during the 1-year follow-up.<sup>15</sup> Nevertheless, in this study, no differences in central pressure, augmentation index, or subendocardial viability ratio were reported. Furthermore, the type of the fabric cover has been found to play a role in the increase of arterial stiffness after EVAR. In a previous study at our center, polyester grafts were found to further increase PWV, when compared to PTFE grafts, and were associated to worse inflammatory response.<sup>22</sup>

Recently, PWV has been utilized to investigate its effect on sac behavior following EVAR. <sup>16,17,25</sup> In a 2018 study of a 25-patient cohort, low preoperative PWV was associated to sac shrinkage after EVAR, whilst high postoperative PWV was an independent factor associated to sac enlargement. <sup>25</sup> More recent studies with larger cohorts corroborate these findings, indicating that arterial stiffness is one of the key factors influencing sac behavior postoperatively <sup>16</sup> and that PWV may be a useful tool for assessing the risk of future sac growth following EVAR. <sup>17</sup>

Another important question is whether the treatment strategy applied (EVAR vs OR) has an impact on arterial stiffness changes observed after AAA repair. In the attempt to provide an answer to this question, contradictory findings were revealed underlining the importance of the arterial stiffness measurement method applied. A recent study assessed

the cardio-ankle vascular index (CAVI) before and after surgery in 110 patients undergoing EVAR and open repair. This study showed that at 48 hours postoperatively the CAVI values were increased in both groups (EVAR and OR) when compared to baseline values. At 6 months of follow-up the CAVI values returned to the baseline for the patients of the open repair group. However, in the endovascular group CAVI values remained higher when compared with the baseline values. <sup>14</sup> Marketou et al found increased arterial stiffness (PWV) in both groups (EVAR and OR), and they also recorded a deterioration of left ventricular total longitudinal strain (GLS) which is an important diagnostic and prognostic marker of early systolic dysfunction. <sup>13</sup> The authors concluded that AAA repair leads not only to an increase in aortic stiffness, measured by the increase in pulse wave velocity, but also to reduced cardiac systolic function. <sup>13</sup>

Further studies showed that the EVAR conferred a higher effect on arterial stiffness compared to OR. Sekhri et al found that elastic modulus (Ep) and stiffness beta were significantly higher in successful EVARs than in open repair, and sacs with endoleaks were significantly less compliant.<sup>11</sup> Gray et al documented a significantly higher postoperative PWV measurement in EVAR patients compared to OR, posing the concern of the long-term effects on cardiovascular morbidity in patients undergoing endovascular repair.<sup>24</sup> In addition, Lantelme et al found that the augmentation index increased after endograft implantation and decreased after graft prosthesis placement, concluding that stent-grafts (EVAR) increase reflected waves more than graft-prostheses (OR).<sup>19</sup> Moloney et al showed in their study that, in both EVAR and OR, replacement of the aneurysmal aorta resulted in a decreased AI.<sup>20</sup>

In contrast, Valdivia et al did not find significant changes in carotid-radial pulse wave velocity (PWVCR) and central blood pressures for both EVAR and OR in the early term. <sup>12</sup> However, they mentioned their concern regarding the use of PWVCR as a tool of arterial stiffness measuring post-AAA repair, since the site of repair was off this method's route.

## Studies Assessing Arterial Stiffness in Patients with TAA, Before and After Treatment The review identified 10 studies (Table 3) reporting the influence of thoracic endografts, following TEVAR, on arterial stiffness. <sup>27–36</sup> Most studies on TEVAR utilized PWV to measure arterial stiffness. <sup>27,28,30,33,35</sup>

In 2017, PWV was used for measuring the arterial stiffness in patients following TEVAR for descending thoracic aorta disease, showing remarkable increase 6 months postoperatively.<sup>27</sup> Hori et al suggest that endoskeleton (stents are mounted inside graft fabric) stent-graft design may allow for aneurysm repair with minimal impact on PWV, thereby reducing the risk of end-organ damage following endovascular repair.<sup>28</sup> Adding to their suggestion, in 2020<sup>30</sup> they compared exoskeleton (metallic stents are mounted outside graft fabric) to endoskeleton endografts, highlighting that postoperatively exoskeleton endografts significantly increased PWV in patients undergoing TEVAR for aortic arch aneurysm, while patients who received an endoskeleton-type endograft experienced a significantly lower risk of cardiac and cerebrovascular events. This was also linked to changes of diastolic dysfunction markers. However, treatment length was not correlated to changes in PWV; this was found also in the Yamashita et al study.<sup>31</sup> At this point, it should be stated that the method used by Yamashita et al was brachial-ankle PWV and thus the site of repair was off the route examined by this method. In contrast, a study has shown that, when the endograft is contained in the ascending aorta, there was no deterioration of the whole aorta's viscoelastic characteristics.<sup>32</sup> In addition, it was demonstrated more recently that treatment length was an independent factor related to PWV increase.<sup>35</sup> The same study also showed that treatment site as a sole factor did not have a different effect on PWV.

Another useful tool that emerged recently is four-dimensional (4D) flow magnetic resonance imaging (MRI), utilized by two of the most recent studies on TEVAR once again proving the impact of endograft implantation on the hemodynamics of the ascending aorta. Gil-Sala et al, via their long-term follow-up (median 126 months), showed that, when compared to matched controls, previously healthy patients who received TEVAR implantation after blunt traumatic thoracic aortic injury exhibited increased ascending aortic diameter, length, and volume, as well as increased aortic stiffness and aberrant flow patterns throughout the whole thoracic aorta.

In their study, Tobey et al pioneered the use of intravascular ultrasound (IVUS) for the measurement of aortic compliance of the ascending aorta and the aortic arch, following TEVAR, which is inextricably linked to the stiffening of the aorta. They demonstrated that thoracic aortic compliance can be properly evaluated by IVUS, and that decreased proximal aortic compliance following endograft implantation may be connected to long-term proximal dilatation.<sup>29</sup>

Since the thoracic aorta is more compliant in younger patients, <sup>29</sup> studies speculate that young patients are more likely to be affected by the endograft implantation<sup>31</sup> and, therefore, congestive cardiac failure and dilated cardiomyopathy post-TEVAR. <sup>37</sup>

 Table 3 Studies Assessing Aortic Stiffness in Patients with TAA, Before and After Treatment

Author, Publication Year	Total N	Subgroups	Groups of Interest, Comparison	Techniques for Arterial Stiffness Assessment	Adjunct Meas.	Conclusions
Moulakakis et al 2017 <sup>27</sup>	27	TEVAR (27)	Pre- and post- TEVAR (descending TAA)	PWV (carotid- femoral)	NT-proBNP (N-terminal pro- brain natriuretic peptide)	<ul> <li>Endovascular treatment of descending thoracic aortic aneurysms is associated with increased NT-proBNP levels and arterial stiffness</li> <li>No correlation between number of stent-grafts and the changes in PWV</li> </ul>
Hori et al 2017 <sup>28</sup>	74	TEVAR (74)	Pre- and post- TEVAR (arch AA)	PWV (brachial- ankle)	NR	Endoskeleton stent-graft design may provide aneurysm repair with mini- mum effect on PWV, which may result in reduced risk of end-organ damage after endovascular repair
Tobey et al 2019 <sup>29</sup>	79	TEVAR (66)	Pre- and post- TEVAR (arch and ascending AA)	IVUS - Dynamic compliance (Cd)	NR	Reduced proximal aortic compli- ance after endograft placement may be related to long-term proximal dilation
Hori et al 2020 <sup>30</sup>	86	TEVAR (86) Exoskeleton (26) Endoskeleton (60)	Pre- and post- TEVAR (arch AA) Endoskeleton vs exoskeleton graft	PWV (brachial- ankle) Echocardiography	NR	<ul> <li>A significant increase in PWV after surgery was observed in patients receiving an exoskeleton-type stent-graft, which was also associated with increase in left ventricular volume and decrease in e', which is a measure for diastolic dysfunction</li> <li>No positive correlation between the treatment length and the changes in PWV</li> </ul>
Yamashita et al 2021 <sup>31</sup>	64	TEVAR (64)	Pre- and post- TEVAR	PWV (brachial- ankle)	NR	PWV increased significantly after TEVAR, especially in older patients with CAD, whereas treatment length or device type was not a predictor of PWV progression after TEVAR
Salvi et al 2022 <sup>32</sup>	60	TEVAR (30) Controls (30)	Ascending AA repair vs controls	PWV (carotid- femoral) Al GRC = bP/fP (backward and forward wave amplitude ratio)	NR	No worsening of the viscoelastic properties of the total aorta was observed after insertion of a rigid prosthesis in the ascending aorta     The pressure curves recorded in the medium term after surgery are comparable to those of the subjects of the control group for both PWV and morphology

(Continued)

Dovepress Moulakakis et al

Table 3 (Continued).

Author, Publication Year	Total N	Subgroups	Groups of Interest, Comparison	Techniques for Arterial Stiffness Assessment	Adjunct Meas.	Conclusions
Youssef et al 2020 <sup>33</sup>	28	TEVAR (14) Controls (14)	TEVAR vs controls	PWV	NT-proBNP Echocardiography	<ul> <li>In young patients, TEVAR after blunt traumatic aortic injury (BTAI) may cause adverse cardiovascular complications due to increased aortic stiffness</li> <li>Screening for arterial hypertension during follow-up is recommended</li> </ul>
Hiraoka et al 2020 <sup>34</sup>	46	TEVAR (46)	Pre- and post- TEVAR	4D flow MRI	NR	The rate of enlargement of the ascending aorta may be affected by the change in wall shear stress or aortic stiffening after TEVAR
Hori et al 2021 <sup>35</sup>	183	TEVAR (183)	Pre- and post- TEVAR (device/site/ treatment length)	PWV (brachial- ankle)	NR	<ul> <li>Najuta thoracic stent-graft did not show a significant increase in PWV, while other commercially available devices showed a significant increase</li> <li>The treatment site did not have a different effect on PWV</li> <li>The treatment length was an independent factor associated with an increase in PWV</li> </ul>
Gil-Sala et al 2021 <sup>36</sup>	52	TEVAR (26) Controls (26)	TEVAR vs control	4D cardiovascular magnetic resonance (CMR)	NR	At long-term follow-up, previously healthy patients who underwent TEVAR implantation after blunt traumatic aortic injury (BTAI) had increased diameter, length, and volume of the ascending aorta, and increased aortic stiffness and abnormal flow patterns in the whole thoracic aorta compared with matched controls

Abbreviations: AA, aortic aneurysm; PWV, pulse wave velocity; AI, augmentation index; NR, not reported.

In a recent study the role of aortic stiffening after TEVAR in young patients is pointed out and, therefore, screening for arterial hypertension during follow-up is suggested.<sup>32</sup> Endograft placement has a known impact on heart remodeling increasing the stress that the left ventricle (LV) receives with each stroke.<sup>23,37</sup> N-terminal pro-brain natriuretic peptide (NT-proBNP)<sup>27,33</sup> and echocardiography<sup>23,30</sup> are useful tools for assessing the impact of arterial stiffening on the heart.

#### Conclusions

Our review showed that the treatment of an abdominal aortic aneurysm either with open repair or endovascularly is associated with an increase of arterial stiffness. An increase of arterial stiffness was uniformly observed in studies investigating patients following thoracic endovascular aortic repair (TEVAR), and the effect was more pronounced in young patients. The effects of increased arterial stiffness after EVAR and TEVAR on the heart and the induced heart remodeling need to be further investigated and evaluated in large studies and special groups of patients. Assessing the association of aortic aneurysmal disease with arterial stiffness may help prevent future cardiovascular events in these patients.

Moulakakis et al Dovepress

### **Funding**

There is no funding to report.

#### **Disclosure**

The authors report no conflicts of interest in this work.

#### References

- 1. Miyoshi T, Ito H. Arterial stiffness in health and disease: the role of cardio-ankle vascular index. *J Cardiol*. 2021;78(6):493–501. doi:10.1016/j. jjcc.2021.07.011
- Umemura S, Arima H, Arima S, et al. The Japanese Society of Hypertension Guidelines for the Management of Hypertension (JSH 2019). Hypertens Res. 2019;42(9):1235–1481. doi:10.1038/s41440-019-0284-9
- 3. O'Rourke MF, Staessen JA, Vlachopoulos C, Duprez D, Plante G. E. Clinical applications of arterial stiffness; definitions and reference values. *Am J Hypertens*. 2002;15(5):426–444. doi:10.1016/S0895-7061(01)02319-6
- 4. Tanaka H, Munakata M, Kawano Y, et al. Comparison between carotid-femoral and brachial-ankle pulse wave velocity as measures of arterial stiffness. *J Hypertens*. 2009;27(10):2022–2027. doi:10.1097/HJH.0b013e32832e94e7
- Kadoglou NPE, Moulakakis KG, Papadakis I, et al. Changes in aortic pulse wave velocity of patients undergoing endovascular repair of abdominal aortic aneurysms. J Endovasc Ther. 2012;19(5):661–666. doi:10.1583/JEVT-12-3916MR.1
- Xuereb RA, Magri CJ, Xuereb RG. Arterial Stiffness and its Impact on Cardiovascular Health. Curr Cardiol Rep. 2023;25(10):1337–1349. doi:10.1007/s11886-023-01951-1
- 7. Hayashi K, Yamamoto T, Takahara A, Shirai K. Clinical assessment of arterial stiffness with cardio-ankle vascular index. *J Hypertens*. 2015;33 (9):1742–1757. doi:10.1097/HJH.0000000000000651
- 8. Saiki A, Sato Y, Watanabe R, et al. The role of a novel arterial stiffness parameter, cardio-ankle vascular index (CAVI), as a surrogate marker for cardiovascular diseases. *J Atheroscler Thromb*. 2016;23(2):155–168. doi:10.5551/jat.32797
- 9. Giudici A, Khir AW, Reesink KD, Delhaas T, Spronck B. Five years of cardio-ankle vascular index (CAVI) and CAVI0: how close are we to a pressure-independent index of arterial stiffness? *J Hypertens*. 2021;39(11):2128–2138. doi:10.1097/HJH.0000000000002928
- Wanhainen A, Verzini F, Van Herzeele I, et al. Editor's Choice European Society for Vascular Surgery (ESVS) 2019 Clinical Practice Guidelines on the Management of Abdominal Aorto-iliac Artery Aneurysms. Eur J Vasc Endovasc Surg. 2019;57(1):8–93. doi:10.1016/j.ejvs.2018.09.020
- 11. Sekhri AR, Lees WR, Adiseshiah M. Measurement of aortic compliance in abdominal aortic aneurysms before and after open and endoluminal repair: preliminary results. *J Endovasc Ther*. 2004;11(4):472–482. doi:10.1583/04-1243.1
- 12. Valdivia AR, Fuente MF, Santos AD, et al. Impact of the Aortic Graft on Arterial Stiffness and Inflammatory Biomarkers after Endovascular Aortic Repair or Open Surgical Repair in Abdominal Aortic Aneurysm Disease. *Ann Vasc Surg*. 2019;59:84–90. doi:10.1016/j.avsg.2018.12.092
- 13. Marketou M, Papadopoulos G, Kontopodis N, et al. Early Left Ventricular Global Longitudinal Strain Deterioration After Aortic Aneurysm Repair: impact of Aortic Stiffness. *J Endovasc Ther*. 2021;28(2):352–359. doi:10.1177/1526602820976636
- Mylonas SN, Moulakakis KG, Kadoglou N, et al. Arterial Stiffness Assessed by Cardio-Ankle Vascular Index in Patients With Abdominal Aortic Aneurysm and Its Alterations After Treatment. Vasc Endovascular Surg. 2021;55(8):804–810. doi:10.1177/15385744211023281
- 15. Holewijn S, Vermeulen JJM, van Helvert M, van de Velde L, Reijnen MMPJ. Changes in Noninvasive Arterial Stiffness and Central Blood Pressure After Endovascular Abdominal Aneurysm Repair. *J Endovasc Ther.* 2021;28(3):434–441. doi:10.1177/15266028211007460
- 16. Nishibe T, Kano M, Maekawa K, et al. Association of preoperative pulse wave velocity to aneurysm sac shrinkage after endovascular aneurysm repair. *Int Angiol.* 2021;40(5):409–415. doi:10.23736/S0392-9590.21.04691-5
- 17. Ugajin A, Iwakoshi S, Ichihashi S, et al. Prediction of Abdominal Aortic Aneurysm Growth After Endovascular Aortic Repair by Measuring Brachial-Ankle Pulse Wave Velocity. *Ann Vasc Surg.* 2022;81:163–170. doi:10.1016/j.avsg.2021.08.056
- 18. Van Herwaarden JA, Muhs BE, Vincken KL, et al. Aortic compliance following EVAR and the influence of different endografts: determination using dynamic MRA. *J Endovasc Ther*. 2006;13(3):406–414. doi:10.1583/06-1848.1
- 19. Lantelme P, Dzudie A, Milon H, et al. Effect of abdominal aortic grafts on aortic stiffness and central hemodynamics. *J Hypertens*. 2009;27 (6):1268–1276. doi:10.1097/HJH.0b013e3283299b22
- 20. Moloney MA, McHugh S, O'Donnell DH, et al. Comparison of arterial stiffness and microcirculatory changes following abdominal aortic aneurysm grafting. *Ir J Med Sci*. 2011;180(2):375–378. doi:10.1007/s11845-010-0513-3
- 21. Lee CW, Sung SH, Chen CK, et al. Measures of carotid-femoral pulse wave velocity and augmentation index are not reliable in patients with abdominal aortic aneurysm. *J Hypertens*. 2013;31(9):1853–1860. doi:10.1097/HJH.0b013e328362360a
- 22. Kadoglou NPE, Moulakakis KG, Papadakis I, et al. Differential effects of Stent-graft fabrics on arterial stiffness in patients undergoing endovascular aneurysm repair. *J Endovasc Ther.* 2014;21(6):850–858. doi:10.1583/14-4772MR.1
- 23. Takeda Y, Sakata Y, Ohtani T, et al. Endovascular aortic repair increases vascular stiffness and alters cardiac structure and function. *Circ J.* 2014;78 (2):322–328. doi:10.1253/circj.CJ-13-0877
- 24. Gray C, Goodman P, Badger SA, O'Malley MK, O'Donohoe MK, McDonnell CO. Endovascular aneurysm repair increases aortic arterial stiffness when compared to open repair of abdominal aortic aneurysms. *Vasc Endovascular Surg.* 2016;50(5):317–320. doi:10.1177/1538574416647503
- 25. Hori D, Nomura Y, Yamauchi T, et al. Perioperative factors associated with aneurysm sac size changes after endovascular aneurysm repair. *Surg Today*. 2019;49(2):130–136. doi:10.1007/s00595-018-1714-z
- 26. Aykan AÇ, Çetin M, Kalaycıoğlu E, Menteşe Ü. Assessment of cardio-ankle vascular index in patients with abdominal aortic aneurysm: an observational study. *Vascular*. 2021;29(2):190–195. doi:10.1177/1708538120946549
- 27. Moulakakis KG, Kadoglou NPE, Antonopoulos CN, et al. Changes in Arterial Stiffness and N-terminal pro-brain natriuretic peptide Levels after Endovascular Repair of Descending Thoracic Aorta. *Ann Vasc Surg.* 2017;38:220–226. doi:10.1016/j.avsg.2016.04.025
- 28. Hori D, Akiyoshi K, Yuri K, et al. Effect of endoskeleton stent graft design on pulse wave velocity in patients undergoing endovascular repair of the aortic arch. *Gen Thorac Cardiovasc Surg.* 2017;65(9):506–511. doi:10.1007/s11748-017-0787-8

**Dove**press Moulakakis et al

29. Tobey DJ, Reynolds TS, Kopchok GE, Donayre CE, Khoynezhad A, White RA. In Vivo Assessment of Ascending and Arch Aortic Compliance. Ann Vasc Surg. 2019;57:22-28. doi:10.1016/j.avsg.2018.12.060

- 30. Hori D, Yuri K, Kusadokoro S, Shimizu T, Kimura N, Yamaguchi A. Effect of endoprostheses on pulse wave velocity and its long-term outcomes after thoracic endovascular aortic repair. Gen Thorac Cardiovasc Surg. 2020;68(10):1134-1141. doi:10.1007/s11748-020-01343-0
- 31. Yamashita Y, Oishi Y, Matsuyama S, et al. Determinants of Changes in Arterial Stiffness after Thoracic Endovascular Aortic Repair. Ann Vasc Surg. 2021;70:474-480. doi:10.1016/j.avsg.2020.06.062
- 32. Salvi L, Alfonsi J, Grillo A, et al. Postoperative and mid-term hemodynamic changes after replacement of the ascending aorta. J Thorac Cardiovasc Surg. 2022;163(4):1283–1292. doi:10.1016/j.jtcvs.2020.05.031
- 33. Youssef A, Kalaja I, Alkomi U, et al. Aortic stiffness and related complications after endovascular repair of blunt thoracic aortic injury in young patients. Vasa - Eur J Vasc Med. 2020;49(4):295-300. doi:10.1024/0301-1526/a000858
- 34. Hiraoka T, Komiya T, Shimamoto T. Enlargement Rate of the Ascending Aorta After Thoracic Endovascular Aortic Repair. Semin Thorac Cardiovasc Surg. 2020;32(2):211–217. doi:10.1053/j.semtcvs.2019.09.010
- 35. Hori D, Fujimori T, Kusadokoro S, Yamamoto T, Kimura N, Yamaguchi A. Treatment Site Does Not Affect Changes in Pulse Wave Velocity but Treatment Length and Device Selection Are Associated With Increased Pulse Wave Velocity After Thoracic Endovascular Aortic Repair. Front Physiol. 2021;12:1-11. doi:10.3389/fphys.2021.739185
- 36. Gil-Sala D, Guala A, Garcia Reyes ME, et al. Geometric, Biomechanic and Haemodynamic Aortic Abnormalities Assessed by 4D Flow Cardiovascular Magnetic Resonance in Patients Treated by TEVAR Following Blunt Traumatic Thoracic Aortic Injury. Eur J Vasc Endovasc Surg. 2021;62(5):797–807. doi:10.1016/j.ejvs.2021.07.016
- 37. Sultan S, Acharya Y, Soliman O, Parodi JC, Hynes N. TEVAR and EVAR, the unknown knowns of the cardiovascular hemodynamics; and the immediate and long-term consequences of fabric material on major adverse clinical outcome. Front Surg. 2022;9. doi:10.3389/fsurg.2022.940304

Vascular Health and Risk Management

## **Dove**press

#### Publish your work in this journal

Vascular Health and Risk Management is an international, peer-reviewed journal of therapeutics and risk management, focusing on concise rapid reporting of clinical studies on the processes involved in the maintenance of vascular health; the monitoring, prevention and treatment of vascular disease and its sequelae; and the involvement of metabolic disorders, particularly diabetes. This journal is indexed on PubMed Central and MedLine. The manuscript management system is completely online and includes a very quick and fair peer-review system, which is all easy to use. Visit http://www.dovepress.com/testimonials.php to read real quotes from published authors.

Submit your manuscript here: https://www.dovepress.com/vascular-health-and-risk-management-journal



