

Renal outcomes of suprarenal *vs.* infrarenal endograft fixation in endovascular abdominal aortic aneurysm repair: a narrative review

Alexander Geragotellis¹, Kofi Cox², Ho Cheung Anthony Yip², Matti Jubouri³, Ian M. Williams⁴, Damian M. Bailey⁵, Mohamad Bashir⁶

¹Faculty of Health Sciences, University of Cape Town, Cape Town, South Africa; ²St. George's University of London, London, UK; ³Hull York Medical School, University of York, York, UK; ⁴Department of Vascular Surgery, University Hospital of Wales, Heath Park, Cardiff, UK; ⁵Neurovascular Research Laboratory, Faculty of Life Sciences and Education, University of South Wales, Pontypridd, UK; ⁶Vascular & Endovascular Surgery, Velindre University NHS Trust, Health Education & Improvement Wales (HEIW), Cardiff, UK

Contributions: (I) Conception and design: M Bashir, A Geragotellis, M Jubouri; (II) Administrative support: M Bashir, M Jubouri; (III) Provision of study materials or patients: None; (IV) Collection and assembly of data: A Geragotellis, K Cox, HCA Yip, M Jubouri; (V) Data analysis and interpretation: A Geragotellis, K Cox, HCA Yip, M Jubouri; (VI) Manuscript writing: All authors; (VII) Final approval of manuscript: All authors. *Correspondence to:* Mohamad Bashir, MD, PhD, MRCS. Vascular & Endovascular Surgery, Velindre University NHS Trust, Health Education & Improvement Wales (HEIW), Cardiff, UK. Email: drmobashir@outlook.com.

Background and Objective: Abdominal aortic aneurysm (AAA) is a common pathology with a prevalence of 4.8%. AAA rupture is associated with significant mortality and so early diagnosis followed by regular monitoring is needed until treatment might be considered and plan intervention. Endovascular aneurysm repair (EVAR) is an established and effective alternative to open surgical repair (OSR) in the treatment of AAAs. Key parameters in defining conventional EVAR suitability include the infrarenal neck length and angulation for the fixation of the proximal graft component. Endograft fixation can be either suprarenal or infrarenal and much debate exists as to which approach is associated with optimum renal outcomes. This study aims to review the current literature with respect to the renal outcomes associated with conventional EVAR using suprarenal fixation (SRF) *vs.* infrarenal fixation (IRF).

Methods: A search was conducted from major search indices (PubMed, Google Scholar and EMBASE) to identify relevant literature pertaining to renal outcomes in EVAR. Recent papers comparing SRF and IRF were evaluated and their findings discussed.

Key Content and Findings: The mechanism of renal function decline (RFD) following EVAR is uncertain and likely multifactorial. Aortic morphology, endograft type and surgical technique may all contribute to RFD. There is a significant degree of heterogeneity within the literature regarding study design and definitions of RFD. Recent literature suggests that RFD is more acute for SRF than IRF in the first post-operative year, but the clinical significance of this decline in patients with normal kidney function is questionable. Studies indicate that SRF is associated with accelerated RFD at 5 years, and that the RFD is worse in patients who are female and who have pre-existing renal insufficiency.

Conclusions: SRF is associated with a greater decline in renal function than IRF in both short- and longterms. Although clinically insignificant in the short-term, the limited available long-term evidence suggests that SRF results in a relatively accelerated decline in renal function when compared to IRF, but it is possibly partially explained by the higher prevalence of advanced degenerative/atherosclerotic disease in SRF cohorts. These trends are noted particularly in female patients and in patients with baseline renal insufficiency.

Keywords: Endovascular; endovascular aneurysm repair (EVAR); abdominal aortic aneurysm (AAA); suprarenal; infrarenal

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Introduction

An abdominal aortic aneurysm (AAA) is a dilatation of the abdominal aorta beyond 1.5 times the normal diameter of the abdominal aorta at the level of the renal arteries (1). A diameter greater than 3.0 cm is typically classified as aneurysmal (2,3). AAA has a prevalence of approximately 4.8%, with men being affected more often than women (4). AAAs can be categorised as suprarenal, juxta renal or infrarenal depending on the location of the aneurysm with respect to the renal arteries (5). Rupture is a surgical emergency with a high mortality and so early diagnosis and treatment is crucial to prevent rupture and death. As aneurysm diameter increases from 5.0 to 6.0 cm, the risk of rupture rises sharply (3-15% annual risk of rupture at 5.0-6.0 cm diameter) and so intervention is considered to prevent aortic rupture and the associated morbidity and mortality (6).

Endovascular aneurysm repair (EVAR) has become a globally well-established treatment for AAA. It is an effective, safe and superior alternative to open surgical repair (OSR) (7-9). The anatomical morphology of an AAA can limit the suitability of many commercially available EVAR devices off-the-shelf. A key parameter in defining suitability to undergo standard EVAR is the length of nonaneurysmal infrarenal aorta, termed 'infrarenal neck length', for proximal device fixation (10). Most endografts that use infrarenal fixation (IRF) require an infrarenal neck length of at least 15 mm, a neck diameter of <32 mm and a neck angulation of less than 60 degrees. However, some earlier studies evaluated the suitability of standard EVAR for AAAs with suboptimal infrarenal anatomy (11-15).

In up to 66% of female and 46% of males with intact AAAs, the aneurysm morphology precludes conventional EVAR (16). However, post-EVAR, there are clinical data to indicate that the suprarenal and visceral segments of the aorta dilate less frequently and rapidly than the infrarenal neck (17). EVAR with suprarenal fixation (SRF) has been postulated as an alternative endovascular approach for the treatment of AAAs with hostile infrarenal neck anatomy (11,18,19). This review aims to summarise and evaluate the current literature with respect to the renal outcomes associated with conventional EVAR using suprarenal and IRF. We present the following article in accordance with the

Narrative Review reporting checklist (available at https:// cdt.amegroups.com/article/view/10.21037/cdt-22-196/rc).

Methods

A literature review was performed using major search indices (PubMed, Google Scholar and EMBASE) to search all scientific articles published to February 2022. The search terms used included: "Suprarenal", "Infrarenal", "EVAR", "Fixation", "Renal", "Abdominal Aortic Aneurysm". Additional sources were identified by reviewing reference lists of relevant publications. Publications with low reliability and non-English publications were excluded. Data were extracted based on their relevance to the topic instead of implementing a systematic approach to paper selection. We present the detail of our search strategy in *Table 1*.

Endovascular repair & endograft fixation

EVAR was first performed by Nicholas L. Volodos' in 1987 and was later popularised by the work of Juan Carlos Parodi and colleagues in 1991 (20,21). Endografts typically take one of three forms: tube, bifurcated and aorto-uni-iliac, though bifurcated grafts are most popular and are used in more than 90% of EVAR cases (22,23). Most AAAs are diagnosed using ultrasound, however thin-sliced contrast-enhanced computerised tomography (CT) imaging is needed to fully assess aneurysm morphology and to generate the 3D aortic models used in EVAR planning. Morphological parameters assessed using CT include the infrarenal aortic diameter and length proximally, and the suitability of the fixation points in the common iliac arteries distally (19). The major companies producing endografts have differing instructions for use (IFU) defining the exact anatomical measurements that are appropriate for their products. What is apparent is that EVAR devices can be used outside manufacturer IFUs without significantly affecting long-term mortality though this may be associated with higher rates of type 1 endoleak (T1E) (15).

Each device used has differing fixation strategies depending on the aortic anatomy being assessed (see *Table 2* and *Figures 1,2*) (24,25). Stent fixation can occur both

Items	Specification
Date of search	22/2/2022–29/2/2022
Databases and other	PubMed was the primary database used
sources searched	EMBASE and Scopus were also searched
Search terms used	"Suprarenal" [All Fields] AND "EVAR" [All Fields] AND "Renal" [All Fields]
	"Suprarenal" [All Fields] AND "EVAR" [All Fields] AND "Renal" [All Fields] AND "Fixation" [All Fields]
	"Infrarenal" [All Fields] AND "EVAR" [All Fields] AND "Renal" [All Fields]
	"Infrarenal" [All Fields] AND "EVAR" [All Fields] AND "Renal" [All Fields] AND "Fixation" [All Fields]
	"Endovascular" [All Fields] AND "Renal" [All Fields] AND "Abdominal Aortic Aneurysm" [All Fields]
Timeframe	1985–2022
	There was a focus on "recent" studies (published from 2010 onwards) that directly compared SRF and IRF
Inclusion and	Focus was placed on published original papers and reviews in English that directly compared SRF and IRF
exclusion criteria	The study excluded articles that were not relevant for the scope of the paper, or that did not directly compare SRF and IRF
Selection process	The search was conducted independently by AG, KC, HCAY, and MJ; data selection is the intersection of the search of these four authors

 Table 1 Summary of the literature search strategy

SRF, suprarenal fixation; IRF, infrarenal fixation.

passively, using the inherent radial force of the stent graft, and actively, using barbs and hooks to anchor the graft into the aortic wall. In order to generate an adequate radial force and proximal fixation into the available neck, the graft diameter is typically oversized by 10-20% (26). General, epidural, and local anaesthesia have all been successfully used for the endovascular repair of infrarenal AAAs (27,28). Whilst there are various endografts on the global market, the principles of endograft insertion remain similar. IRF, procedurally summarised elsewhere (29), is used where there is suitable aortic neck morphology and describes the positioning of the stent-graft immediately beneath the most inferiorly located renal artery. Where the graft is fixed in the suprarenal location, the procedure is similar but the graft is fixed more proximally with the metallic barbs that anchor the graft extending superiorly to the nonaneurysmal suprarenal aorta above the fabric-covered stentgraft (29). This theoretically allows perfusion to the coeliac, superior mesenteric artery and both renal arteries, which might partially be covered by the bare metallic stent.

Renal outcomes

Mechanisms of renal function decline (RFD)

One of the major complications of EVAR is a decline in RFD, possibly necessitating dialysis. Although EVAR avoids the renal insult highly associated with OSR (7), there can still be significant consequences. The full mechanism is unclear but suggestions have been made (30-32). The cause for renal injury is likely multifactorial, with the type of endograft, aortic morphology and surgical techniques all being contributors.

An inflammatory foreign body reaction may occur upon stent-graft introduction (31). It is also possible that the stent and graft material being present at the renal artery ostia result in renal artery occlusion or even parenchymal injury. Fixation barbs may interact with plaques at the renal artery origins, leading to luminal compromise. Trans-renal stent struts could cross the renal ostia, leading to a functional stenosis (33). It is also possible that thicker SRF stenting wires that cross the renal artery ostia may cause greater perturbations to

Table 2 A summary	of manufacturer	IFU detailing the	e various aneurysn	n dimensional re	quirement for com	mon endografts			
Parameters	Anaconda™ (Terumo Aortic)	Zenith [®] (Cook Medical Technologies) (non- fenestrated)	Zenith [®] Fenestrated (Cook Medical Technologies) (fenestrated)	Gore [®] Excluder [®] (Gore medical)	Endurant [™] II (Medtronic)	Aorfix TM (Lombard medical technologies)	Incraft [®] (Cordis [®])	Powerlink [®] (Endologix)	Ovation Prime [®] (TriVascular)
Neck angle relative to the axis of the suprarenal aorta (Figure 1A)	Unspecified	<45 degrees	<45 degrees	Unspecified	≤45 degrees (≤60 degrees if infrarenal neck length ≥15 mm)	Unspecified	<60 degrees	Unspecified	Unspecified
Neck angle relative to the long axis of the aneurysm (Figure 1B)	≤90 degrees	<60 degrees	<45 degrees	≤60 degrees	≤60 degrees (≤75 degrees if infrarenal neck length ≥15 mm)	≤90 degrees	<60 degrees	<60 degrees	≤60 degrees if proximal neck is ≥10 mm and ≤45 degrees if proximal neck is <10 mm
Neck diameter (Figure 1C)	17.5–31 mm	18-32 mm	19–31 mm	19–32 mm	19–32 mm	19–29 mm	17–31 mm	18–26 mm	16–30 mm
Infrarenal neck length (<i>Figure 1D</i>)	≥15 mm	≥15 mm	≥4 mm	≥15 mm	≥10 mm	≥15 mm	≥10 mm	≥15 mm	≥13 mm
CIA diameter (Figure 1E)	8.5–21 mm	8–20 mm	9–21 mm ipsilateral, 7–21 contralateral	8–25 mm	8–25 mm	9–19 mm	7–22 mm	10–14 mm	8–25 mm
CIA length (Figure 1F)	Unspecified	>10 mm	>10 mm	>10 mm	>15 mm	≥15 mm	≥15 mm	≥15 mm	≥10 mm
Femoral artery diameter	Unspecified	6.0 or 6.5 mm	7.7 or 8.5 mm	'Adequate'	'Adequate'	'Adequate'	'Adequate'	'Adequate'	'Adequate'
Fixation location	Infrarenal	Suprarenal	Suprarenal	Infrarenal	Suprarenal	Infrarenal or transrenal	Suprarenal	Infrarenal (but sometimes used as suprarenal)	Suprarenal
Sourced from man parameters column	ufacturer websit	tes and/or FDA is for use; CIA, c	patents. <i>Table 2</i> :ommon iliac arte.	is best interpriny.	eted alongside <i>Fi</i>	gure 1 which v	visually repre-	sents the measu	rements indicated in the



Figure 1 A representation of an AAA. Various angles and lengths are labelled and can be interpreted with reference to the manufacturer's required measurements in *Table 2*. A, suprarenal angle. B, infrarenal angle, relative to the long axis of the aneurysm. C, infrarenal neck diameter. D, infrarenal neck length. E, common iliac artery diameter. F, common iliac artery length. AAA, abdominal aortic aneurysm.

blood flow velocity of the renal arteries (34); it is likely that these mechanisms will become clearer after the anticipated publication of current hemodynamic studies (35).

The stenting wires may not cause a significant enough reduction in renal artery flow velocity to impair function. Alternatively, there could be inherent anatomic variability in the interaction between SR stent struts and renal artery ostia (36), as SR stent struts reportedly cross the renal artery ostia in 50% to 80% of patients (37,38). This may further complicate the understanding of the pathogenesis of RFD post-EVAR. Coverage of the origins of the renal arteries may impair blood flow and lead to RFD via occlusion and/or stenosis (31,32). Microembolism and dissections are possibilities (31,32). However, these sequelae are increasingly likely to be anticipated, detected, and avoided with the advent of new software systems being used during EVAR procedures (30,31).

The control of dyslipidaemia and hypertension is important among EVAR patients as these pathological processes can predispose to weakening of the aortic wall (39). These factors could contribute, to microembolisation of the renal arteries, leading to localised ischemia of the renal parenchyma (40). Ischaemia-reperfusion injury arising from compromised arterial flow to lower limbs



Figure 2 Sealant rings and stent-grafts frequently selected from the EVAR armoury. IRF: AneuRx (Medtronic); Excluder (W. L. Gore & Associates); Powerlink (Endologix); Aorfix (Lombard Medical); SRF: Zenith low profile (Cook Medical); Endurant/Talent (Medtronic); Incraft (Cordis Corp); Powerlink (Endologix); Aorfix (Lombard Medical). Reproduced via open access from Gozzo *et al.* (24). ePTFE, expanded polytetrafluoroethylene; EVAR, endovascular aneurysm repair; IRF, infrarenal fixation; SRF, suprarenal fixation.

by instrumentation in the femoral arteries has been suggested as a cause of RFD in EVAR (30,31,41). The exact mechanism is complex, but the oxidative stress formation and peroxidation of lipids seems to be key in promoting the inflammatory processes resulting in renal injury (42). Since EVAR requires contrast media for imaging throughout the treatment process, contrast-induced nephropathy (CIN) (43) could be another cause for RFD post-EVAR, although the exact mechanism is not well established (30,36).

Towards a standardised definition of renal injury

Despite the establishment of EVAR reporting standards (44,45), interpretation of the literature is limited by the heterogenous reporting of renal outcomes. Recent metaanalyses have highlighted this (46-48). It also remains difficult to grasp the essence of RFD as there are limited differences in dialysis or survival between SRF and IRF methods, which necessitates the development of more granular reporting strategies to quantify RFD. Serial serum creatinine (S-Cr) and creatinine clearance (CrCl, based on Cockcroft-Gault formula) are widely reported but are prone to inherent bias and inaccuracies (e.g., affected by nutritional intake, medications, race, age etc.) (49,50). Cystatin-C has been mentioned (51) but remains poorly characterised, perhaps due to its relative cost, hindering its uptake into the clinical realm. The most accurate measure of renal function is estimated glomerular filtration rate (eGFR, mL/min/1.73 m^2) (52,53) with a reduction of 20-30% considered clinically significant for medium- and longterm function (39,54). The formula used most commonly is the Chronic Kidney Disease Epidemiology Collaboration (CKD-EPI) or less commonly, the Modification of Diet in Renal Disease (MDRD). Some have argued the difference in eGFR between the 2 formulas is clinically insignificant (55). However, a recent systematic review and meta-analysis suggested that CKD-EPI is a more accurate estimate of GFR than MDRD (56).

Results from pooled meta-analyses

The effect of the type of stent graft fixation after EVAR on renal function remains uncertain, (18,36,48,57-61). Some studies have shown poorer renal function after SRF (18,19,58,59,61), whereas others suggest minimal differences between the fixation methods (60,62-64). Accepting that certain aortic anatomies are more favourable for SRF, no randomised controlled trials (RCTs) exist which compare the fixation strategies directly. Meta-analyses are therefore necessary to assess the existing literature.

Currently, Calderbank et al. (47) present the only metaanalysis that uses a uniform definition of renal dysfunction as the primary outcome which is a decrease in eGFR of greater than 20% at 1 year. A similar 1-year renal decline between SRF and IRF is observed, but over 5 years, SRF may accelerate renal impairment (47). The five series reporting an eGFR reduction at 1-year showed the weighted OR was 1.53 (SRF vs. IRF) (95% CI: 0.67-3.51; P=0.31; $I^2=53.2\%$) (47). Furthermore, 9.3% of those with SRF vs. 7.4% with IRF developed a 20% or higher drop of eGFR by 12 months (47). Secondary outcome was an eGFR drop of greater than 20% at 5 years. Pooling of various definitions of "renal dysfunction" (e.g., S-Cr changes) and variable follow-up time periods meant the analysis data was highly heterogeneous with respect to the secondary outcome. The duration of follow-up ranged from 72 h to 5 years and weighted OR (SRF vs. IRF) was 1.32 (95% CI: 1.01–1.71; P=0.03; I^2 =28.4%). The overall incidence of renal dysfunction was 5.1% in SRF compared to 4.6% in IRF (47).

Other meta-analyses combine the renal function outcomes (e.g., "Cr increase >20%", "CrCl decrease >20%" and "GFR decrease >20%") (36,48). However, these studies also corroborate that the method of aortic wall fixation does not influence earlier RFD at <1 year. However, SRF causes slightly worse outcomes in the longer term. Stather *et al.* (48) analysed 25 studies (SRF =16,634; IRF =38,198) and demonstrated insignificant differences between SRF and IRF in the short (30-day) or interim term (12 months) (48). The longer-term end points employed were variable between studies, ranging between 1–74 months, and were unable to undergo sensitivity analysis (48). Longer-term follow ups are encouraged in SRF *vs.* IRF comparative studies for more meaningful long-term outcomes in metaanalyses.

However, at these study endpoints, there were marginally worse RFD outcomes for SRF patients (SRF 5.98% vs. IRF 4.83%; OR =1.29; 95% CI: 1.18–1.40; P<0.001). There were also significant differences reported for renal infarcts (SRF 6.6% vs. IRF 2.3%; OR =2.78; 95% CI: 1.46–5.29; P=0.002), renal stenosis (SRF 2.4% vs. IRF 0.8%; OR =2.89; 95% CI: 1.00–8.38; P=0.05), and renal artery occlusion (SRF 2.4% vs. IRF 1.2%; OR =2.21; 95% CI: 1.15–4.25; P=0.02) (48). The relevance of these data findings must be carefully considered as there were insignificant differences in haemodialysis rates reported between SRF and IRF (P=0.58) (48). Some studies fail to identify these findings (19). The higher incidence of renal infarcts in the SRF group (48) may possibly be due in part to the use of older generations of endografts and the greater known radial fixation force with SRF (36,65). Earlier, Miller *et al.* (36) analysed 21 studies (SRF =1,949; IRF =2,525) and found statistically insignificant differences in the risk of postoperative renal complications between SRF and IRF over median follow-up of 1-year (36).

Longer-term evidence

Studies that feature 5-year follow-up durations are exclusively retrospective. These databases remain limited, by non-randomised selections which has implications on selection of SRF or IRF. Undoubtedly, SRF is preferred when certain anatomical characteristics are present such as shorter aortic neck length, increased angulation, and larger diameters (60,66). These anatomical characteristics are associated with more advanced degenerative/atherosclerotic disease of the aorta, which should be considered as contributors to the deterioration of renal function.

Saratzis et al. (54) reported on changes in eGFR (via CKD-EPI) in 242 patients following elective EVAR using both SRF and IRF at 5-year compared to baseline. This is the longest available cohort study reporting on renal outcomes with both SRF and IRF. A control group of 121 patients undergoing elective OSR [10 women (8%); mean age 72±6 years] and with available eGFR estimates at 5 years were also identified. They were matched with 242 SRF and 242 IRF patients for age, sex, smoking habits, diabetes, and eGFR at baseline (1:2 ratio). The volume of radiographic contrast (mL) was non-significant (SRF: 121±32 vs. IRF: 118±21, P=0.36) (54). Infrarenal neck anatomy was non-hostile, and the endografts used were Anaconda, Excluder, Endurant, and Zenith (distributions not reported) (54). Proportions of patients in whom an eGFR decrease of >30% from baseline eGFR was reported (18,39). Those undergoing SRF were significantly more likely to experience >30% eGFR decline at 1- and 5-year post-EVAR compared with the IRF population (1 year: IRF 19%, SRF 27%; 5 years: IRF 27%, SRF 47%). Secondary outcomes measured the change in eGFR at 1 year and stage of CKD at 1 and 5 years. There was a marked decline of eGFR during the first postoperative year (-10.7 for SRF and -2.2 for IRF) compared to remainder of follow-up (54). By 5 years, IRF patients had lost a mean of 8.2 mL/min/1.73 m² (95% CI: 6.5–10.8; P<0.001) compared

Other retrospective studies' mid- and long-term results also suggest poorer longer-term renal outcomes with SRF (39,57-59). Banno *et al.* (18) conducted a retrospective mid-term outcomes comparison of SRF (n=135, Zenith, Endurant or Incraft) *vs.* IRF (n=102, Excluder, Powerlink, Aorfix). Propensity matching of 87 pairs was conducted to exclude patients who had progressed to acute kidney injury (AKI) (18). This occurred significantly more frequently in the SRF group than in the IRF group (P=0.026) (18). Propensity scores were estimated using covariates of age, sex, baseline renal function, aortic aneurysm diameter, comorbidities, smoking habits, use of drugs impairing renal function, and frequency of contrast-enhanced CT scans performed (18).

Mid-term outcomes of 3-year were reported, though the median follow-up duration was 5.1 years (IQR: 4.0-6.4). Logistic regression analysis showed that SRF was independently predictive of >20% decline in the eGFR at 3 years post-EVAR (OR =2.06; 95% CI: 1.18-3.58; P=0.11) (18). The decrease in eGFR within the SRF group gradually increased annually and reached statistical difference at 3 years post-surgery compared with the IRF group (mean 17.8% vs. 11.6%, respectively; P=0.034) (18). Hahl et al. (39) retrospectively compared SRF (n=267, Zenith or Endurant) vs. IRF (n=91, Excluder) over 5-year. The baseline demographics, procedure duration and volume of contrast reagent used were insignificant between the groups, although the IRF group had higher rates of hypertension (78.0% vs. 58.4%, P=0.001) and dyslipidaemia (54.9% vs. 40.4%, P=0.020) (39). At 7-day post-intervention, 13.7% (n=36) of the SRF group vs. 3.5% (n=3) of the IRF group experienced $\geq 20\%$ decline in eGFR (P=0.009) (39). Median eGFR declined significantly from baseline levels to the 5-year time point in both groups (SRF 72.0 vs. 51.0, P<0.001; IRF 69.0 vs. 54.5, P=0.001). Mixed model analysis showing an annual rate of eGFR decline of -3.13 (95% CI: -3.40 to 2.85; P<0.01) (39). Furthermore, in both the SRF and IRF cohorts, the number with $\geq 20\%$ decline in eGFR did not differ significantly by 30-day and also in the follow-up years 1-5 [see Table 3, Hahl et al. (39)].

It is likely those with pre-existing renal insufficiency are prone to poorer EVAR outcomes when SRF is used (39,61,67,68). This may be complicated by non-randomised selection for SRF but possibly depends on surgical experience and the extent of aortic neck morphology (70,71). Hahl *et al.* (39) reported a sub-group analysis with preexisting renal insufficiency (eGFR \leq 60) and showed more

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First author	Publication date	Sample, n	Follow-up, years	Measurement used for renal function	Formula used for eGFR calculation	Main devices, % of total SRF + IRF cohort	Conclusion on renal outcomes over follow-up period
Hahl	2022	SRF: 267	5	eGFR decline ≥20%	CKD-EPI	Zenith; Endurant	Transient but immediately greater
et al. (39)		IRF: 91				Excluder	postoperative RFD in SRF; IRF is safer in the longer term, especially in patients with baseline renal insufficiency
Blecha <i>et al.</i> (67)		SRF: 76	Ŋ	eGFR decline ≥20%	Not reported	Powerlink (23%); Zenith (13%); Talent (6%);	Female sex and baseline renal insufficiency are significant risk factors
		IRF: 58				Excluder (39%); Aneurx (9%)	for RFD at 5 years (multivariate); SRF was significant risk factor for RFD at 5 years on univariate analysis and approached significance on multivariate analysis
Erben <i>et al.</i> (68)	2021	SRF: 670 IRF: 460	4.8±3.7	eGFR decline over follow up; rate of change of creatinine	MDRD	Not reported	Insignificant eGFR decline and rate of change of creatinine over follow up between SRF and IRF; longer hosoital stav in SRF patients (3.4+2.2
							vs. 2.3±1.0 days; P<0.001); SRF and female sex are predictors of longer hospital stay (multivariable regression)
Pujari <i>et al.</i> (61)		SRF: 3,225	30 days	Cr >2 mg/dL without dialysis or new dialysis	MDRD	Zenith (25%); Endurant (34%)	SRF is associated with more perioperative renal complications
		IRF: 2,309				Excluder (42%)	than IRF, especially in patients with baseline renal insufficiency
Sangtae <i>et al.</i> (60)		SRF: 114	3.8±2.6	<30% decrease in eGFR; >30% increase in follow-up	CKD-EPI; MDRD; Cockcroft-Gault	Endurant (42%); Talent (4%)	Insignificant difference in renal function and adverse events (P>0.107)
		IRF: 54	3.3±2.2	sCrª, dialysis needed		Excluder (24%); Powerlink (6%); Aneurx (2%);	between SRF and IRF
Banno <i>et al.</i> (18)	2020	SRF: 135	5.5±1.8 (stud) reported on	/ eGFR decline ≥20%	3-variable Japanese	Zenith (30%); Endurant/ Talent (24%); Incraft (3%)	SRF has worse mid-term outcomes than IRF based on mean eGFR decline
		IRF: 102	mid-term outcomes at 3-year)		equation ^b	Excluder (37%); Powerlink (5%)	(SRF: 17.8% decline; IRF: 11.6% decline; P=0.034)
Table 3 (conti	inued)						

Geragotellis et al. Renal outcomes of supra- vs. infra-renal fixation in endovascular AAA repair

Table 3 (cont.	inued)						
First author	Publication date	Sample, n	Follow-up, years	Measurement used for renal function deterioration	Formula used for eGFR calculation	Main devices, % of total SRF + IRF cohort	Conclusion on renal outcomes over follow-up period
Zettervall et al. (58)	2017	SRF: 2,273 (63%)	30 days	Cr >2 mg/dL without dialysis or new dialysis	Not reported	Zenith; Endurant	Overall rates of RFD in EVAR are low; SRF associated with higher likelihood
		IRF: 1,314 (37%)				Excluder	of perioperative RFD (OR =12.0; 95% CI: 1.6–91) and postoperative hospital stay duration >2 days (OR =1.4; 95% CI: 1.2–1.7)
Zettervall et al. (57)		SRF: 1,264	2.6	sCr >0.5 mg/dL or new perioperative haemodialysis		Zenith (31%); Endurant (12%); Talent (7%)	SRF associated with higher rates of RFD (OR =2.0; 95% CI: 1.2–3.4) and
		IRF: 1,310				Excluder (40%); AneuRx (11%)	postoperative hospital stay duration >2 days (OR =1.8; 95% CI: 1.4–2.2)
^a , if initial sC Serum Creat (Cordis Corp	ừ 1.2 mg/dL, inine – 1.094,	follow-up scar 1 × age – 0.287 × r, NJ, USA). IRF:	1.46 mg/dL w 0.739 (if fem Excluder (W.	as considered as renal impairr ale). SRF: Zenith (Cook Medica L. Gore & Associates, Flagsta	ment by calculating al, Bloomington, IN tff, AZ, USA); Powe	g 1.2+1.2×30%; ^b , 3-variabl , USA); Endurant/Talent (Me erlink (Endologix, Irvine, CA	e Japanese eGFR equation (69): 194 × dtronic, Minneapolis, MN, USA); Incraft USA); Aorfix (Lombard Medical, Irvine,

CA, USA); AneuRx (Medtronic, Minneapolis, MN, USA). (s)Cr, serum creatinine, mg/dL. SRF, suprarenal fixation; IRF, infrarenal fixation; CKD-FPI, Chronic Kidney Disease

renal function decline; MDRD, Modification of Diet in Renal Disease

Epidemiology Collaboration; RFD,

in the SRF (n=86) compared to the IRF group (n=25) had a \geq 20% eGFR decline (59.5% vs. 20.0%, P=0.036) (39). Pujari *et al.* (61) also included a subgroup analysis of patients with moderate kidney dysfunction (n=1,780; GFR =30–59). They reported higher rates of renal complications with SRF than IRF (SRF: 2.2% vs. IRF: 0.8%, P=0.02) (61). Blecha *et al.* (67) considered baseline renal insufficiency (multivariate OR =3.0, P=0.029) to be a significant predictor of >20% GFR decline by 5 years after EVAR. In addition, when using SRF in females; multivariate binary logistic regression analysis showed female sex (OR =3.9, P=0.023) to be a predictor of >20% GFR decline at 5 years (67).

Similarly, Erben *et al.* (68), using follow-up of $4.8\pm$ 3.7 years, found on Kaplan-Meier analysis that CKD patients undergoing SRF were more likely to progress to haemodialysis (P=0.039). Furthermore, the authors found on least square multivariable regression that SRF (Coef, 9.5; 95% CI: 0.11–1.11; P<0.0001) and female sex (Coef, 2.4; 95% CI: 0.17–0.41; P=0.02) were predictive of prolonged length of hospital stay (68). However, in the mid-term (3-year), sex of the patient has not been significantly associated with RFD (18).

Zettervall et al. (57) attempted to reduce any effect of patient selection by performing a sub-group analysis in those undergoing either SRF or IRF. SRF had a significantly increased risk of RFD (defined as S-Cr increase >0.5 mg/dL), irrespective of whether they were performed by surgeons of the Vascular Study Group of New England (VSGNE) (OR =2.0; 95% CI: 1.2-3.4) or those who routinely use SRF or IRF (SRF 3.7% vs. 1.3%, P=0.02; OR =2.9; 95% CI: 1.1–7.8) (57). However, as the authors argue, it is possible that routine users of IRF endografts still selected IRF grafts in preference of the straightforward anatomy and referred more complicated cases to those with more experience with SRF (57). Additionally, hostile aortic anatomies requiring SRF invariably require more contrast-medium which may confound study results. Those undergoing SRF received more contrast than those undergoing IRF (110 vs. 88 mL, P<0.01) (57). Other studies might report contrast volume differences as insignificant but possible effects on renal function should not be overlooked (39,54).

Future directions in clinical and academic practice

Those undergoing EVAR are often elderly with usually associated comorbidities. RFD following EVAR might be expected, irrespective of endograft fixation levels (62,64,72,73). SRF is regarded as safe in those with normal renal function in the short-term. However, there appears to be a trend towards greater RFD in those undergoing SRF over mid- and longer-term periods (18,39,67). Management of RFD requires regular blood level monitoring, especially those with preoperative baseline renal insufficiency. It is possible that biomarker-informed surveillance strategies for monitoring the development of RFD will grow in uptake, but at present, remain in their early stages (74,75).

An obvious clinical solution is to undertake additional periprocedural precautions. Procedurally, EVAR may cause renal impairment either through the procedure itself, or by associated CIN and dehydration. Atheromatous microembolisation can be minimised by reducing the manipulation to the patient (e.g., via catheterisation, wires and balloons) (30,40). It remains uncertain whether remote ischaemic preconditioning is protective against reperfusioninjuries of the kidneys (76,77). Obviously, to minimise CIN, postprocedural contrast use should be kept to a minimal level (30).

Contrast-precautions should be applied to all undergoing EVAR, rather than those with pre-existing renal dysfunction (30). Certain drugs such as fenoldopam may also be protective against CIN (78). Additionally, RenalGuard[®] and hemodynamic guided hydration have been proven as optimal strategies for preventing CIN in high-risk patients in a 2020 systematic review and meta-analysis (79). To mitigate EVAR-associated dehydration, bolus high-dose NaHCO₃ (1 mL/kg of 8.4% NaHCO₃) and crystalloid hydration has shown promise as a nephroprotective agent (80). By 30-day post EVAR, 7% of those in the intervention arm (aggressive rehydration with a single bolus of NaHCO₃) developed AKI compared to 33% in the control arm (aggressive rehydration only) (80). The proportion of EVARs undergoing SRF was 97% (56/58) (80).

Conclusions

The steeper short-term decline in renal function associated with SRF vs. IRF appears to have little clinical relevance in those with normal baseline renal function. The limited long-term evidence suggests that SRF is associated with an accelerated decline in renal function compared to IRF, although this may be partially due to a higher prevalence of advanced degenerative/atherosclerotic disease in SRF cohorts. Additional caution is necessary in females with baseline renal insufficiency. RFD with SRF has been emphasised in this population. In future practice, longerterm reporting on renal function outcomes must be performed to characterise the effects of SRF vs. IRF. Further multicentre prospective studies are necessary to improve the available evidence with further control for possible bias in patient-selection and surgical experience according to the anatomical features of the aorta and the aneurysms. It is realistic to expect that improvements in device design, surgical experience and clinical care would reflect improved renal outcomes in future studies.

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542

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544

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