

Supplementary File

Supplementary Figures

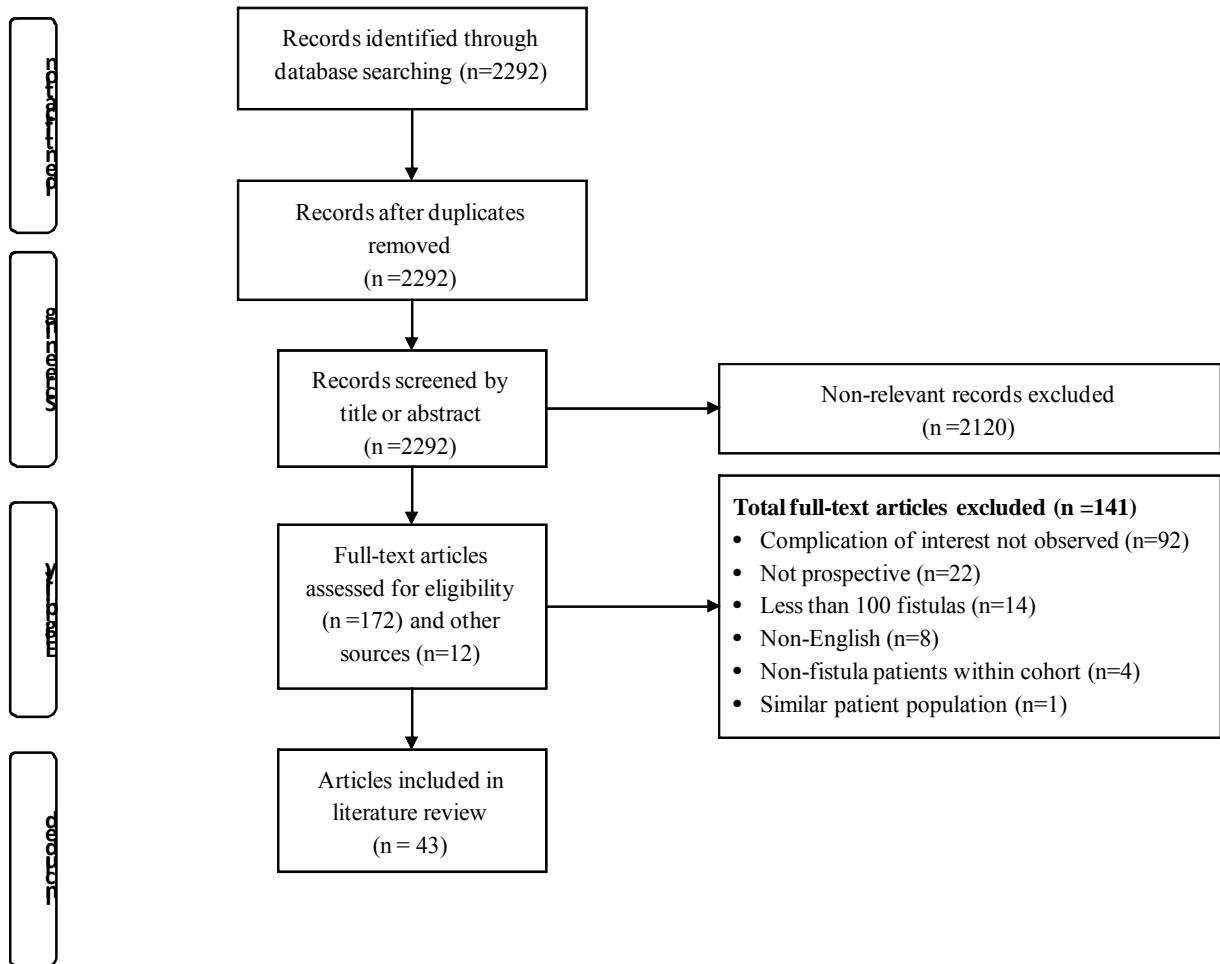


Figure S1: Flow diagram of study eligibility and inclusion.

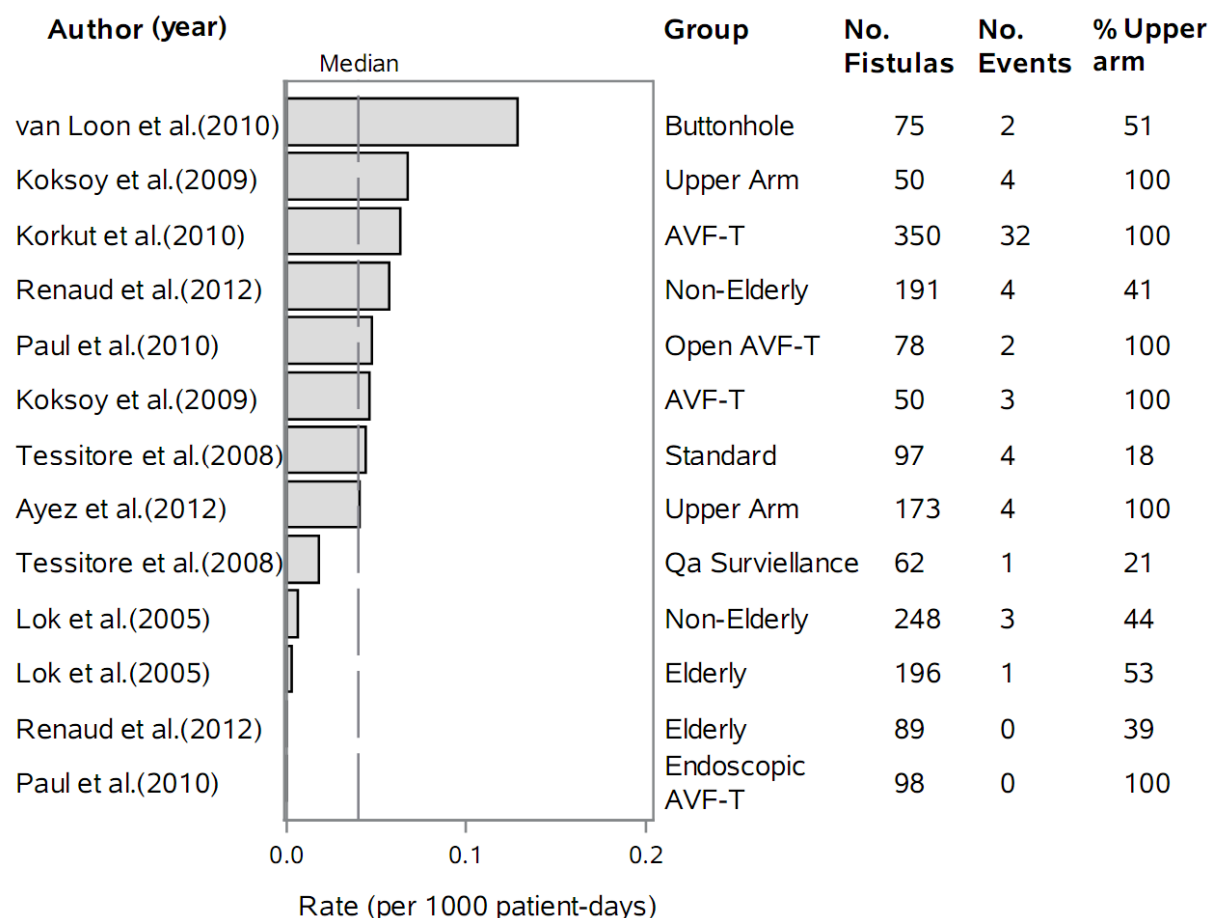


Figure S2: Studies reporting the incident rate for aneurysms per 1000 patient-days. Dashed line refers to the median event rate per 1000 patient-days. Please note, Van Loon et al. estimated an event rate of 3.01 Aneurysms per 1000 patient-days for patients cannulated using the rope ladder technique. We excluded this event rate because the figure became distorted when including this event rate.

AVF-T: transposed arteriovenous fistula; open AVF-T: AVF transposition of a deep vein through a long open incision; Endoscopic AVF-T: AVF transposition of a deep vein through endoscopic procedure; Buttonhole: All patients cannulated using a buttonhole technique; Upper Arm: cohort made up of all upper arm fistulas; Qa Surveillance: Access blood flow monitoring. Elderly: ≥ 65 years; Standard: no specific cohort followed.

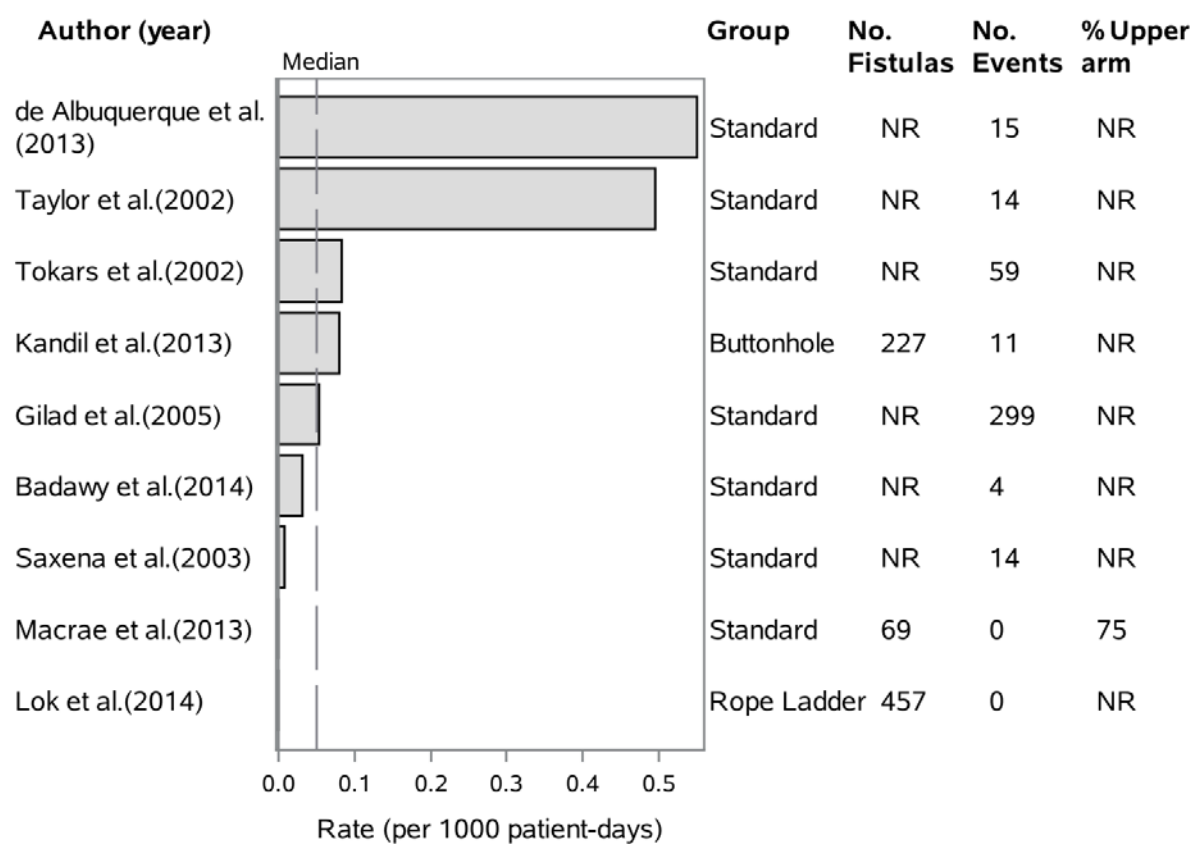


Figure S3: Studies reporting the incident rate for bloodstream infections per 1000 patient-days. Dashed line refers to the median event rate per 1000 patient-days.

Buttonhole: All patients cannulated using a buttonhole technique; Rope Ladder: All patients cannulated using a rope ladder technique; Standard: no specific cohort followed.

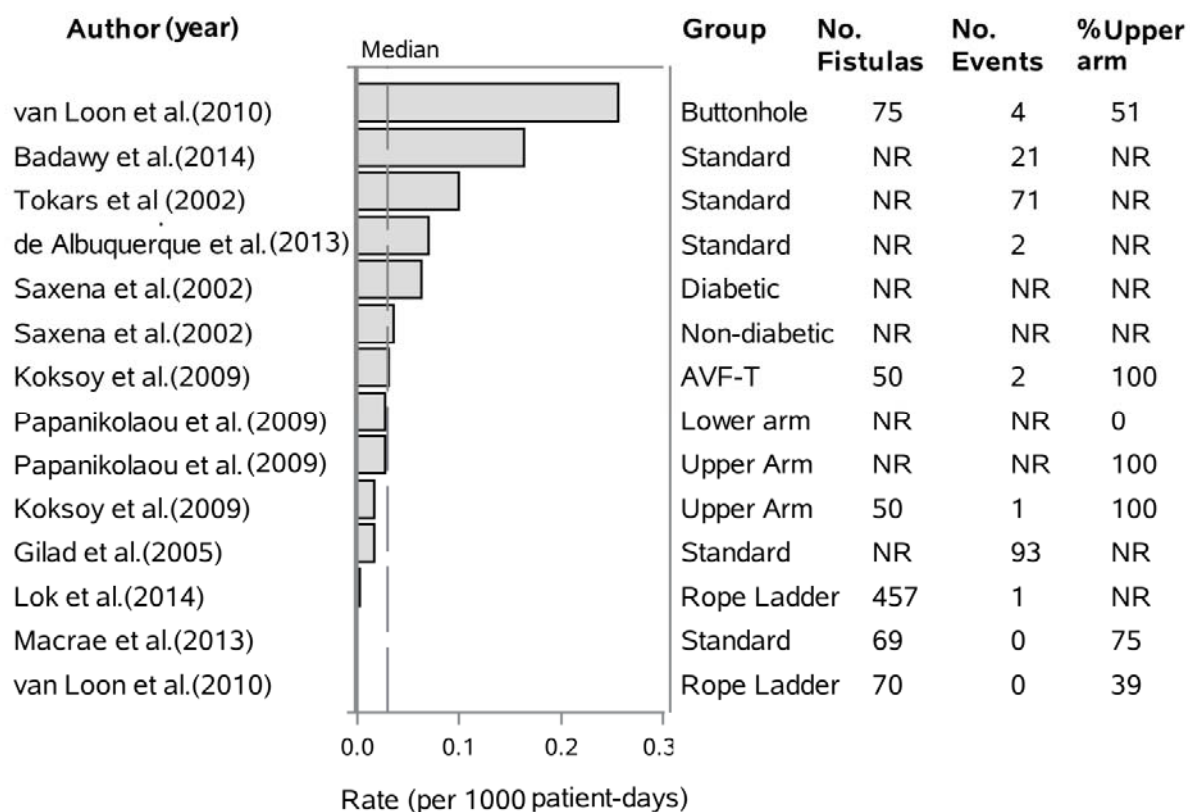


Figure S4: Studies reporting the incident rate for local access infections per 1000 patient-days. Dashed line refers to the median event rate per 1000 patient-days.

AVF-T: transposed arteriovenous fistula; Buttonhole: All patients cannulated using a buttonhole technique; Rope Ladder: All patients cannulated using a rope ladder technique; Diabetic: refers to diabetic nasal carriers of methicillin-resistant and methicillin-susceptible staphylococcus aureus (MRSA & MSSA); Non-diabetic: refers to non-diabetic nasal carriers of methicillin-resistant and methicillin-susceptible staphylococcus aureus (MRSA & MSSA); Upper Arm: cohort made up of all upper arm fistulas; Lower Arm: cohort made up of all lower arm fistulas; Standard: no specific cohort followed.

Table S1. PRISMA checklist

Section/topic		# Checklist item	Reported
TITLE			
Title	1	Identify the report as a systematic review, meta-analysis, or both.	Yes
ABSTRACT			
Structured summary	2	Provide a structured summary including, as applicable: background; objectives; data sources; study eligibility criteria, participants, and interventions; study appraisal and synthesis methods; results; limitations; conclusions and implications of key findings; systematic review registration number.	Yes
INTRODUCTION			
Rationale	3	Describe the rationale for the review in the context of what is already known.	Yes
Objectives	4	Provide an explicit statement of questions being addressed with reference to participants, interventions, comparisons, outcomes, and study design (PICOS).	Yes
METHODS			
Protocol and registration	5	Indicate if a review protocol exists, if and where it can be accessed (e.g., Web address), and, if available, provide registration information including registration number.	Yes
Eligibility criteria	6	Specify study characteristics (e.g., PICOS, length of follow-up) and report characteristics (e.g., years considered, language, publication status) used as criteria for eligibility, giving rationale.	Yes
Information sources	7	Describe all information sources (e.g., databases with dates of coverage, contact with study authors to identify additional studies) in the search and date last searched.	Yes
Search	8	Present full electronic search strategy for at least one database, including any limits used, such that it could be repeated.	Yes
Study selection	9	State the process for selecting studies (i.e., screening, eligibility, included in systematic review, and, if applicable, included in the meta-analysis).	Yes
Data collection process	10	Describe method of data extraction from reports (e.g., piloted forms, independently, in duplicate) and any processes for obtaining and confirming data from investigators.	Yes

Data items	11	List and define all variables for which data were sought (e.g., PICOS, funding sources) and any assumptions and simplifications made.	Yes
Risk of bias in individual studies	12	Describe methods used for assessing risk of bias of individual studies (including specification of whether this was done at the study or outcome level), and how this information is to be used in any data synthesis.	Yes
Summary measures	13	State the principal summary measures (e.g., risk ratio, difference in means).	Yes
Synthesis of results	14	Describe the methods of handling data and combining results of studies, if done, including measures of consistency (e.g., I^2) for each meta-analysis.	Yes
Risk of bias across studies	15	Specify any assessment of risk of bias that may affect the cumulative evidence (e.g., publication bias, selective reporting within studies).	Yes
Additional analyses	16	Describe methods of additional analyses (e.g., sensitivity or subgroup analyses, meta-regression), if done, indicating which were pre-specified.	N/A
RESULTS			
Study selection	17	Give numbers of studies screened, assessed for eligibility, and included in the review, with reasons for exclusions at each stage, ideally with a flow diagram.	Yes
Study characteristics	18	For each study, present characteristics for which data were extracted (e.g., study size, PICOS, follow-up period) and provide the citations.	Yes
Risk of bias within studies	19	Present data on risk of bias of each study and, if available, any outcome level assessment (see item 12).	Yes
Results of individual studies	20	For all outcomes considered (benefits or harms), present, for each study: (a) simple summary data for each intervention group (b) effect estimates and confidence intervals, ideally with a forest plot.	Yes
Synthesis of results	21	Present results of each meta-analysis done, including confidence intervals and measures of consistency.	N/A
Risk of bias across studies	22	Present results of any assessment of risk of bias across studies (see Item 15).	Yes
Additional analysis	23	Give results of additional analyses, if done (e.g., sensitivity or subgroup analyses, meta-regression [see Item 16]).	N/A
DISCUSSION			
Summary of evidence	24	Summarize the main findings including the strength of evidence for each main outcome; consider their relevance to key groups (e.g., healthcare providers, users, and policy makers).	Yes

Limitations	25	Discuss limitations at study and outcome level (e.g., risk of bias), and at review-level (e.g., incomplete retrieval of identified research, reporting bias).	Yes
Conclusions	26	Provide a general interpretation of the results in the context of other evidence, and implications for future research.	Yes
FUNDING			
Funding	27	Describe sources of funding for the systematic review and other support (e.g., supply of data); role of funders for the systematic review.	Yes

Table S2. Distribution of components describing study quality of observational studies

Was the conception of study question formed prior to data collection?	
No	21
Unclear	3
Yes	15
Recruitment Type	
Consecutive	31
Not reported	5
Random	3
Was enrolment based on pre-specified eligibility criteria?	
No	13
Yes	26
Was the exposure defined as the access planned (access intended) as opposed to the access in place prior to the study (access achieved)?	
No	20
Yes	19
Is it reported whether participants were eligible to different forms of fistula access?	
No	25
Unclear	1
Yes	13
Was the proportion lost to follow-up less or equal to 10%?	
Unclear	27
Yes	12
Aneurysm: Was the outcome definition the same as published standardized definition? 16,45	
No	1
Unclear	5

Yes	2
Not Applicable	31
Infection: Was the outcome definition the same as published standardized definition	
No	2
Unclear	7
Yes	13
Not Applicable	17
Ischemic Steal: Was the outcome definition the same as published standardized definition⁴⁴	
No	1
Unclear	9
Yes	2
Not Applicable	27
Venous Hypertension: Was the outcome definition the same as published standardized definition	
Unclear	1
Not Applicable	38
Thrombosis: Was the outcome definition the same as published standardized definition^{43,44}	
No	3
Unclear	11
Yes	1
Not Applicable	24
Were at least age, sex, diabetes, heart disease, and peripheral vascular disease considered or reported?	
No	32
Yes	7
Are reports of the study free of suggestion of selective outcome reporting?	

No	6
Unclear	3
Yes	30
Was the study apparently free of other problems that could put it at a risk of bias?	
No	17
Unclear	2
Yes	20

Table S3: Distribution of components describing study quality of randomized controlled trials

Was the allocation sequence adequately generated?	
Unclear	1
Yes	3
Was allocation adequately concealed?	
Unclear	1
Yes	3
Was knowledge of the allocated interventions adequately prevented during the study?	
No	1
Yes	3
Were incomplete outcome data adequately addressed?	
No	1
Yes	3
Are reports of the study free of suggestion of selective outcome reporting?	
No	1
Yes	3
Was the study apparently free of other problems that could put it at a risk of bias?	
No	1
Yes	3
Did the outcome assessor NOT have knowledge of the allocated intervention?	
No	2
Yes	2
Was the Proportion lost to follow-up less or equal to 10%?	
Unclear	1

Yes	3
Aneurysm: Was the outcome definition the same as published standardized definition? ^{16,45}	
Yes	1
Not Applicable	3
Infection: Was the outcome definition the same as published standardized definition?	
Yes	1
Not Applicable	3
Steal: Was the outcome definition the same as published standardized definition? ⁴⁴	
Not Applicable	4
Venous hypertension: Was the outcome definition the same as published standardized definition?	
Not Applicable	4
Thrombosis: Was the outcome definition the same as published standardized definition?	
Unclear	2
Yes	1
Not Applicable	1

*The number of cohorts reported rather than the number of studies.

Table S4: Complication rates reported by study and unique cohort.

Author (Year)	Group	Complication	No. Fistulas	No. Events	Reported Rate	Units	Rate (per 1000 patient-days)
Lok et al. (2005)	Non-Elderly (<65 years)	Aneurysm	248	3	0.0002	patient-month	0.01
Lok et al. (2005)	Elderly (≥65 years)	Aneurysm	196	1	0.00	patient-month	0.00
Tessitore et al. (2008)	Standard	Aneurysm	97	4	0.02	patient-year	0.04
Tessitore et al. (2008)	Qa Surveillance	Aneurysm	62	1	0.01	patient-year	0.02
Koksoy et al. (2009)	Upper Arm	Aneurysm	50	4	0.02	patient-year	0.07
Koksoy et al. (2009)	AVF-T	Aneurysm	50	3	0.02	patient-year	0.05
van Loon et al. (2010)	Buttonhole	Aneurysm	75	2	0.05	patient-year	0.13
van Loon et al. (2010)	Rope Ladder	Aneurysm	70	47	1.1	patient-year	3.01
Paul et al. (2010)	Open AVF-T	Aneurysm	78	2	0.02	patient-year	0.05
Paul et al. (2010)	Endovascular AVF-T	Aneurysm	98	0	0.00	patient-year	0.00
Korkut et al. (2010)	AVF-T	Aneurysm	350	32	0.02	patient-year	0.06
Renaud et al. (2012)	Non-Elderly (<65 years)	Aneurysm	191	4	0.02	patient-year	0.06
Renaud et al. (2012)	Elderly (≥65 years)	Aneurysm	89	0	0.00	patient-year	0.00
Ayez et al. (2012)	Upper Arm	Aneurysm	173	4	0.04	1000 patient-days	0.04
Elseviers et al. (2003)	Standard	Bleeding	1049	22	0.06	1000 patient-days	0.06
Koksoy et al. (2009)	AVF-T	Bleeding	50	2	0.01	patient-year	0.03
Koksoy et al. (2009)	Upper Arm	Bleeding	50	1	0.01	patient-year	0.02
Papanikolaou et al. (2009)	Upper Arm	Bleeding	.	.	0.04	patient-year	0.11
Papanikolaou et al. (2009)	Lower arm	Bleeding	.	.	0.03	patient-year	0.08
Korkut et al. (2010)	AVF-T	Bleeding	350	17	0.01	patient-year	0.03
Ayez et al. (2012)	Upper Arm	Bleeding	173	8	0.08	1000 patient-days	0.08
McCarley et al. (2001)	DVPM	Catheter Insertion	41	.	0.06	patient-year	0.16
McCarley et al. (2001)	VABFM	Catheter Insertion	43	.	0.07	patient-year	0.19
McCarley et al. (2001)	NM	Catheter Insertion	39	.	0.18	patient-year	0.49
Tessitore et al. (2008)	Standard	Catheter Insertion	97	22	0.10	patient-year	0.27
Tessitore et al. (2008)	Qa Surveillance	Catheter Insertion	62	4	0.03	patient-year	0.07
Ravani et al. (2002)	Standard	Death	197	78	0.23	patient-year	0.64
Astor et al. (2005)	Standard	Death	185	44	11.7	100 person-years	0.32

Jennings et al. (2006)	Standard	Death	134	4	0.03	patient-year	0.09
Tessitore et al. (2008)	Qa Surveillance	Death	62	21	0.14	patient-year	0.37
Tessitore et al. (2008)	Standard	Death	97	24	0.1	patient-year	0.26
Koksoy et al. (2009)	Upper Arm	Death	50	22	0.14	patient-year	0.37
Koksoy et al. (2009)	AVF-T	Death	50	10	0.06	patient-year	0.15
Paul et al. (2010)	Open AVF-T	Death	78	14	0.12	patient-year	0.33
Paul et al. (2010)	Endovascular AVF-T	Death	98	18	0.16	patient-year	0.44
Ng et al. (DOPPS) (2011)	Standard	Death	476	27	13	100 patient-years	0.36
Renaud et al. (2012)	Elderly (≥ 65 years)	Death	89	10	0.1	patient-year	0.28
Renaud et al. (2012)	Non-Elderly (< 65 years)	Death	191	13	0.07	patient-year	0.19
Ayez et al. (2012)	Upper Arm	Death	173	55	0.56	1000 patient-days	0.56
Tessitore et al. (2008)	Standard	Difficulty Cannulation	97	4	0.02	patient-year	0.04
Tessitore et al. (2008)	Qa Surveillance	Difficulty Cannulation	62	4	0.03	patient-year	0.07
Olsha et al. (2014)	Elderly (≥ 80 years)	Hematoma	128	2	0.01	patient-year	0.03
Lok et al. (2005)	Non-elderly (< 65 years)	High cardiac output	248	2	0.006	patient-year	0.017
Lok et al. (2005)	Elderly (≥ 65 years)	High cardiac output	196	0	0	patient-year	0
McCarley et al. (2001)	NM	Hospitalization (All-Cause)	39	.	0.72	patient-year	1.97
McCarley et al. (2001)	DVPM	Hospitalization (All-Cause)	41	.	0.47	patient-year	1.29
McCarley et al. (2001)	VABFM	Hospitalization (All-Cause)	43	.	0.10	patient-year	0.27
Ng et al. (DOPPS) (2011)	Standard	Hospitalization (All-Cause)	476	.	104	100 patient-years	2.85
Badawy et al. (2014)	Standard	Hospitalization (All-Cause)	.	121	2.83	100 patient-months	0.94
Ng et al. (DOPPS) (2011)	Standard	Hospitalization (Infection-Related)	476	.	12.6	100 patient-years	0.34
Lok et al. (2003)	Duplex US Monitoring	Hospitalization (VA-Related)	189	.	6	1000 access-days	6.00
Lok et al. (2003)	Transonic Surveillance	Hospitalization (VA-Related)	241	.	4	1000 access-days	4.00
Jennings et al. (2006)	Standard	Hospitalization (VA-Related)	134	0	0.00	patient-year	0.00
Ng et al. (DOPPS) (2011)	Standard	Hospitalization (VA-Related)	476	.	23.8	100 patient-years	0.65
Tokars et al. (2002)	Standard	Infection-Bacteremia	.	59	0.25	100 patient-months	0.08

Taylor et al. (2002)	Standard	Infection-Bacteremia	.	14	0.21	1000 dialysis sessions	0.50
Gilad et al. (2005)	Standard	Infection-Bacteremia	.	299	0.16	100 patient-months	0.05
Saxena et al. (2003)	Standard	Infection-Bacteremia	.	14	0.02	100 patient-months	0.01
Macrae et al. (2013)	Standard	Infection-Bacteremia	69	0	0.00	patient-year	0.00
de Albuquerque et al. (2013)	Standard	Infection-Bacteremia	.	15	0.55	1000 access-days	0.55
Kandil et al. (2013)	Buttonhole	Infection-Bacteremia	227	11	2.94	100 treatment years	0.08
Badawy et al. (2014)	Standard	Infection-Bacteremia	.	4	0.09	100 patient-months	0.03
Lok et al. (2014)	Rope Ladder	Infection-Bacteremia	457	0	0.00	1000 patient-days	0.00
Tokars et al. (2002)	Standard	Infection-Local	.	71	0.30	100 patient-months	0.10
Saxena et al. (2002)	Diabetic Nasel Carriers of MRSA/MSSA	Infection-Local	.	.	0.02	patient-year	0.06
Saxena et al. (2002)	Non-diabetic Nasel Carriers of MRSA/MSSA	Infection-Local	.	.	0.01	patient-year	0.04
Gilad et al. (2005)	Standard	Infection-Local	.	93	0.05	100 patient-months	0.02
Koksoy et al. (2009)	Upper Arm	Infection-Local	50	1	0.01	patient-year	0.02
Koksoy et al. (2009)	AVF-T	Infection-Local	50	2	0.01	patient-year	0.03
Papanikolaou et al. (2009)	Lower arm	Infection-Local	.	.	0.01	patient-year	0.03
Papanikolaou et al. (2009)	Upper Arm	Infection-Local	.	.	0.01	patient-year	0.03
van Loon et al. (2010)	Buttonhole	Infection-Local	75	4	0.09	patient-year	0.26
van Loon et al. (2010)	Rope Ladder	Infection-Local	70	0	0.00	patient-year	0.00
Macrae et al. (2013)	Standard	Infection-Local	69	0	0.00	patient-year	0.00
de Albuquerque et al. (2013)	Standard	Infection-Local	.	2	0.07	1000 access-days	0.07
Badawy et al. (2014)	Standard	Infection-Local	.	21	0.49	100 patient-months	0.16
Lok et al. (2014)	Rope Ladder	Infection-Local	457	2	0.00	1000 patient-days	0.00
Bonforte et al. (2004)	Lower arm	Ischemic steal syndrome	112	2	0.00	patient-month	0.02
Lok et al. (2005)	Elderly (≥ 65 years)	Ischemic steal syndrome	196	3	0.00	patient-month	0.01

Lok et al. (2005)	Non-Elderly (<65 years)	Ischemic steal syndrome	248	1	0.00	patient-month	0.00
Jennings et al. (2006)	Standard	Ischemic steal syndrome	134	2	0.02	patient-year	0.05
Huijbregts et al. (2008)	Standard	Ischemic steal syndrome	285	2	0.01	patient-year	0.03
Papanikolaou et al. (2009)	Lower arm	Ischemic steal syndrome	.	.	0.01	patient-year	0.03
Papanikolaou et al. (2009)	Upper Arm	Ischemic steal syndrome	.	.	0.10	patient-year	0.27
Paul et al. (2010)	Open AVF-T	Ischemic steal syndrome	78	2	0.02	patient-year	0.05
Paul et al. (2010)	Endovascular AVF-T	Ischemic steal syndrome	98	0	0.00	patient-year	0.00
Korkut et al. (2010)	AVF-T	Ischemic steal syndrome	350	25	0.02	patient-year	0.05
Jennings et al. (2011)	Elderly (≥ 65 years)	Ischemic steal syndrome	461	15	0.06	1000 patient-days	0.06
Renaud et al. (2012)	Non-Elderly (<65 years)	Ischemic steal syndrome	191	3	0.02	patient-year	0.04
Renaud et al. (2012)	Elderly (≥ 65 years)	Ischemic steal syndrome	89	2	0.02	patient-year	0.06
Ayez et al. (2012)	Upper Arm	Ischemic steal syndrome	173	12	0.12	1000 patient-days	0.12
Olsha et al. (2014)	Elderly (≥ 80 years)	Ischemic steal syndrome	128	11	0.06	patient-year	0.18
McCarley et al. (2001)	NM	Radiological Intervention	39	.	0.00	patient-year	0.00
McCarley et al. (2001)	VABFM	Radiological Intervention	43	.	0.21	patient-year	0.58
McCarley et al. (2001)	DVPM	Radiological Intervention	41	.	0.09	patient-year	0.25
Dixon et al. (2002)	Lower arm	Radiological Intervention	88	26	1.44	patient-year	3.95
Dixon et al. (2002)	Upper Arm	Radiological Intervention	117	76	2.71	patient-year	7.42
Lok et al. (2003)	Transonic Surveillance	Radiological Intervention	241	.	0.65	1000 access-days	0.65
Lok et al. (2003)	Transonic Surveillance	Radiological Intervention	241	.	0.10	1000 access-days	0.10
Lok et al. (2003)	Duplex US Monitoring	Radiological Intervention	189	.	0.73	1000 access-days	0.73
Lok et al. (2003)	Duplex US Monitoring	Radiological Intervention	189	.	0.21	1000 access-days	0.21
Lok et al. (2005)	Elderly (≥ 65 years)	Radiological Intervention	196	.	0.31	access-year	0.85
Lok et al. (2005)	Elderly (≥ 65 years)	Radiological Intervention	92	.	0.30	access-year	0.82

Lok et al. (2005)	Elderly (≥ 65 years)	Radiological Intervention	92	.	0.02	access-year	0.05
Lok et al. (2005)	Elderly (≥ 65 years)	Radiological Intervention	93	.	0.32	access-year	0.88
Lok et al. (2005)	Non-Elderly (< 65 years)	Radiological Intervention	93	.	0.26	access-year	0.71
Lok et al. (2005)	Non-Elderly (< 65 years)	Radiological Intervention	248	.	0.29	access-year	0.79
Lok et al. (2005)	Non-Elderly (< 65 years)	Radiological Intervention	93	.	0.09	access-year	0.25
Lok et al. (2005)	Non-Elderly (< 65 years)	Radiological Intervention	248	.	0.10	access-year	0.27
Lok et al. (2005)	Elderly (≥ 65 years)	Radiological Intervention	93	.	0.03	access-year	0.08
Lok et al. (2005)	Non-Elderly (< 65 years)	Radiological Intervention	139	.	0.09	access-year	0.25
Lok et al. (2005)	Non-Elderly (< 65 years)	Radiological Intervention	139	.	0.30	access-year	0.82
Lok et al. (2005)	Elderly (≥ 65 years)	Radiological Intervention	196	.	0.02	access-year	0.05
Shahin et al. (2005)	Qa Surveillance	Radiological Intervention	76	.	0.58	patient-year	1.59
Shahin et al. (2005)	Standard	Radiological Intervention	146	.	0.10	patient-year	0.27
Jennings et al. (2006)	Standard	Radiological Intervention	134	11	0.09	patient-year	0.25
Polkinghorne et al. (2006)	Standard	Radiological Intervention	68	1	0.01	patient-year	0.03
Polkinghorne et al. (2006)	Qa Surveillance	Radiological Intervention	69	5	0.05	patient-year	0.14
Huijbregts et al. (2008)	Standard	Radiological Intervention	285	49	0.24	patient-year	0.66
Tessitore et al. (2008)	Qa Surveillance	Radiological Intervention	62	33	0.21	patient-year	0.58
Tessitore et al. (2008)	Standard	Radiological Intervention	97	22	0.09	patient-year	0.24
Pflederer et al. (2008)	AVF-T	Radiological Intervention	161	.	0.31	patient-year	0.85
Pflederer et al. (2008)	Standard	Radiological Intervention	321	.	0.54	patient-year	1.48
Koksoy et al. (2009)	Upper Arm	Radiological Intervention	50	7	0.04	patient-year	0.12
Koksoy et al. (2009)	AVF-T	Radiological Intervention	50	4	0.02	patient-year	0.06
van Loon et al. (2010)	Buttonhole	Radiological Intervention	75	3	0.07	patient-year	0.19
van Loon et al. (2010)	Rope Ladder	Radiological Intervention	70	38	0.89	patient-year	2.44

Paul et al. (2010)	Open AVF-T	Radiological Intervention	78	70	0.61	patient-year	1.66
Renaud et al. (2012)	Elderly (≥ 65 years)	Radiological Intervention	89	32	0.33	patient-year	0.91
Renaud et al. (2012)	Non-Elderly (<65 years)	Radiological Intervention	191	59	0.31	patient-year	0.85
Ayez et al. (2012)	Upper Arm	Radiological Intervention	173	58	0.59	1000 patient-days	0.59
Lin et al. (2013)	Standard	Radiological Intervention	62	.	0.29	patient-year	0.79
Lin et al. (2013)	Infrared	Radiological Intervention	60	.	0.11	patient-year	0.30
Macrae et al. (2013)	Buttonhole	Radiological Intervention	70	135	0.90	patient-year	2.47
Macrae et al. (2013)	Standard	Radiological Intervention	69	82	0.72	patient-year	1.97
Agarwal et al. (2014)	AVF-T (2-Stage)	Radiological Intervention	83	.	2.15	patient-year	5.89
Agarwal et al. (2014)	AVF-T (1-Stage)	Radiological Intervention	61	.	1.84	patient-year	5.04
Olsha et al. (2014)	Elderly (≥ 80 years)	Radiological Intervention	128	198	0.9	patient-year	2.47
Lok et al. (2005)	Elderly (≥ 65 years)	Surgical Revisions	196	.	0.28	access-year	0.77
Lok et al. (2005)	Elderly (≥ 65 years)	Surgical Revisions	92	.	0.61	access-year	1.67
Lok et al. (2005)	Non-Elderly (<65 years)	Surgical Revisions	93	.	0.04	access-year	0.11
Lok et al. (2005)	Non-Elderly (<65 years)	Surgical Revisions	248	.	0.06	access-year	0.16
Lok et al. (2005)	Non-Elderly (<65 years)	Surgical Revisions	139	.	0.03	access-year	0.08
Lok et al. (2005)	Elderly (≥ 65 years)	Surgical Revisions	93	.	0.00	access-year	0.01
Jennings et al. (2006)	Standard	Surgical Revisions	134	4	0.03	patient-year	0.09
Polkinghorne et al. (2006)	Qa Surveillance	Surgical Revisions	69	7	0.07	patient-year	0.19
Polkinghorne et al. (2006)	Standard	Surgical Revisions	68	5	0.05	patient-year	0.15
Huijbregts et al. (2008)	Standard	Surgical Revisions	285	40	0.20	patient-year	0.55
Tessitore et al. (2008)	Qa Surveillance	Surgical Revisions	62	14	0.09	patient-year	0.25
Tessitore et al. (2008)	Standard	Surgical Revisions	97	6	0.02	patient-year	0.07
Koksoy et al. (2009)	Upper Arm	Surgical Revisions	50	1	0.01	patient-year	0.02

Koksoy et al. (2009)	AVF-T	Surgical Revisions	50	0	0.00	patient-year	0.00
van Loon et al. (2010)	Buttonhole	Surgical Revisions	75	3	0.07	patient-year	0.19
van Loon et al. (2010)	Rope Ladder	Surgical Revisions	70	3	0.07	patient-year	0.19
Paul et al. (2010)	Open AVF-T	Surgical Revisions	78	8	0.07	patient-year	0.19
Paul et al. (2010)	Endovascular AVF-T	Surgical Revisions	98	6	0.05	patient-year	0.15
Renaud et al. (2012)	Elderly (≥ 65 years)	Surgical Revisions	89	8	0.08	patient-year	0.23
Renaud et al. (2012)	Non-Elderly (< 65 years)	Surgical Revisions	191	21	0.11	patient-year	0.30
Ayez et al. (2012)	Upper Arm	Surgical Revisions	173	30	0.3	1000 patient-days	0.30
Lin et al. (2013)	Standard	Surgical Revisions	62	.	0.26	patient-year	0.71
Lin et al. (2013)	Infrared	Surgical Revisions	60	.	0.08	patient-year	0.22
Macrae et al. (2013)	Buttonhole	Surgical Revisions	70	.	0.09	patient-year	0.25
Macrae et al. (2013)	Standard	Surgical Revisions	69	.	0.11	patient-year	0.30
McCarley et al. (2001)	DVPM	Thrombosis	41	.	0.15	patient-year	0.41
McCarley et al. (2001)	VABFM	Thrombosis	43	.	0.07	patient-year	0.19
McCarley et al. (2001)	NM	Thrombosis	39	.	0.14	patient-year	0.38
Lok et al. (2003)	Transonic Surveillance	Thrombosis	241	.	0.33	1000 access-days	0.33
Lok et al. (2003)	Duplex US Monitoring	Thrombosis	189	.	0.44	1000 access-days	0.44
Elseviers et al. (2003)	Standard	Thrombosis	1049	55	0.15	1000 patient-days	0.15
Bonforte et al. (2004)	Lower arm	Thrombosis	112	12	0.00	patient-month	0.14
Mallamaci et al. (2005)	Standard	Thrombosis	205	78	0.39	1000 patient-days	0.39
Lok et al. (2005)	Elderly (≥ 65 years)	Thrombosis	196	25	0.00	patient-month	0.07
Lok et al. (2005)	Non-Elderly (< 65 years)	Thrombosis	248	34	0.00	patient-month	0.07
Shahin et al. (2005)	Qa Surveillance	Thrombosis	76	.	0.21	patient-year	0.57
Shahin et al. (2005)	Standard	Thrombosis	146	.	0.26	patient-year	0.71
Jennings et al. (2006)	Standard	Thrombosis	134	19	0.16	patient-year	0.43
Roosbeh et al. (2006)	Standard	Thrombosis	171	31	0.09	patient-year	0.26
Polkinghorne et al. (2006)	Qa Surveillance	Thrombosis	69	6	0.06	patient-year	0.17
Polkinghorne et al. (2006)	Standard	Thrombosis	68	4	0.04	patient-year	0.12

Huijbregts et al. (2008)	Standard	Thrombosis	285	29	0.14	patient-year	0.38
Tessitore et al. (2008)	Qa Surveillance	Thrombosis	62	5	0.03	patient-year	0.09
Tessitore et al. (2008)	Standard	Thrombosis	97	20	0.08	patient-year	0.22
Korkut et al. (2010)	AVF-T	Thrombosis	350	186	0.13	patient-year	0.37
Ayez et al. (2012)	Upper Arm	Thrombosis	173	25	0.25	1000 patient-days	0.25
Lin et al. (2013)	Standard	Thrombosis	62	11	0.18	patient-year	0.49
Lin et al. (2013)	Standard	Thrombosis	62	8	0.52	patient-year	1.43
Lin et al. (2013)	Infrared	Thrombosis	60	3	0.05	patient-year	0.14
Lin et al. (2013)	Infrared	Thrombosis	60	1	0.07	patient-year	0.19
Macrae et al. (2013)	Buttonhole	Thrombosis	70	6	0.04	patient-year	0.11
Macrae et al. (2013)	Standard	Thrombosis	69	7	0.05	patient-year	0.14
Elseviers et al. (2003)	Standard	Total Complications	1049	163	0.43	1000 patient-days	0.43
Lok et al. (2005)	Elderly (≥ 65 years)	Total Complications	196	57	0.00	patient-month	0.15
Lok et al. (2005)	Non-Elderly (<65 years)	Total Complications	248	61	0.00	patient-month	0.13
Papanikolaou et al. (2009)	Lower arm	Total Complications	.	.	0.25	patient-year	0.68
Papanikolaou et al. (2009)	Upper Arm	Total Complications	.	.	0.57	patient-year	1.56
Ayez et al. (2012)	Upper Arm	Total Complications	173	110	1.12	1000 patient-days	1.12
Tokars et al. (2001)	Standard	Total Infections	.	13	0.02	patient-month	0.60
Stevenson et al. (2002)	Standard	Total Infections	.	3	0.17	1000 dialysis sessions	0.39
Tokars et al. (2002)	Standard	Total Infections	.	130	0.56	100 patient-months	0.19
Elseviers et al. (2003)	Standard	Total Infections	1049	22	0.06	1000 patient-days	0.06
Jennings et al. (2006)	Standard	Total Infections	134	0	0.00	patient-year	0.00
Huijbregts et al. (2008)	Standard	Total Infections	285	8	0.04	patient-year	0.11
Qasaimieh et al. (2008)	Standard	Total Infections	104	24	0.36	patient-year	0.99
Pflederer et al. (2008)	AVF-T	Total Infections	161	.	0.07	patient-year	0.19
Pflederer et al. (2008)	Standard	Total Infections	321	.	0.05	patient-year	0.14
Labriola et al. (2011)	Standard	Total Infections	.	57	0.31	1000 access-days	0.31
Renaud et al. (2012)	Elderly (≥ 65 years)	Total Infections	89	0	0.00	patient-year	0.00
Renaud et al. (2012)	Non-Elderly (<65 years)	Total Infections	191	4	0.02	patient-year	0.06
Ravani et al. (DOPPS) (2013)	Standard	Total Infections	3352	.	3.00	1000 access-days	3.00
Ravani et al. (DOPPS) (2013)	Standard	Total Infections	3352	.	1.7	1000 access-days	1.70
Ravani et al. (DOPPS) (2013)	Standard	Total Infections	3352	.	0.9	1000 access-days	0.90

Macrae et al. (2013)	Standard	Total Infections	69	0	0.00	patient-year	0.00
Kandil et al. (2013)	Buttonhole	Total Infections	227	26	4.01	100 treatment years	0.11
Lok et al. (2014)	Rope Ladder	Total Infections	457	2	0.00	1000 patient-days	0.00
Korkut et al. (2010)	AVF-T	Venous hypertension	350	16	0.01	patient-year	0.03

Table S5: Median complication rate (min, max) by patient subgroup

Subgroup	Aneurysm	Infections		Steal	Thrombosis
		All Types	Local Access		
Elderly*	0.001 (0, 0.003)	NS	NS	0.06 (0.008 ,0.18)	NS
Non-Elderly*	0.03 (0.01, 0.06)	NS	NS	0.02 (0.002 ,0.04)	NS
Lower Arm	NS	NS	NS	0.03 (0.02 ,0.03)	NS
Upper Arm	0.047 (0, 0.07)	0.03 (0.02, 0.03)	0.03 (0.02, 0.03)	0.05 (0 ,0.27)	0.31 (0.25, 0.37)
Buttonhole	NS	0.17 (0.08, 0.26)	NS	NS	NS
Rope Ladder	NS	0.001 (0, 0.002)	0.001 (0, 0.002)	NS	NS
Any Surveillance	NS	NS	NS	NS	0.33 (0.09, 0.58)

*As defined in the study ; NS = Not Sufficient data (less than two groups)

Note: Subgroups are not mutually exclusive

Item S1 – Study Protocol

PROSPERO International prospective register of systematic reviews

Complication rates of the arteriovenous fistula: a systematic review

Ahmed A. Al-Jaishi, Matthew J. Oliver, Aiden R. Liu, Amit X. Garg, Joyce C. Zhang, Sonia M. Thomas, Louise M. Moist

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Review question(s)

For adult patients (≥18 years) on chronic hemodialysis, what is the complication rate of aneurysms, infections, ischemic steal syndrome, thrombosis, and venous hypertension among published literature between January 2000 and December 2013?

Searches

We designed and implemented a systematic literature search to identify all relevant published reports in Medline (PubMed) from January 1st 2000 to December 12th 2013. Our search strategy has been validated and has a sensitivity of 96.4% and specificity of 95.1% for capturing dialysis related published studies. We also used related articles feature in PubMed. Two investigators screened all English Language titles and abstracts obtained through the search syntax to identify potentially relevant articles. We retrieved the full-text of these articles to further assess their suitability for inclusion in this review. Bibliographies of selected articles were searched manually to identify any additional relevant studies.

Types of study to be included

We will include any study that collected data prospectively (observational cohort studies or randomized control trials) and followed patients for at least 3 months.

Condition or domain being studied

Currently, over 23,000 Canadians and 2 million people worldwide with kidney failure survive by receiving chronic hemodialysis. Hemodialysis cannot occur without a reliable connection between the dialysis machine and the patient's circulation – this connection is their “vascular access” (VA) and represents the patient's “lifeline”.

The VA can take one of three forms: i) a native arteriovenous fistula (fistula); ii) a synthetic arteriovenous graft (AV-graft); or iii) a central venous catheter (catheter). When the VA fails or has complications such as clotting or infection, a patient's life is put at risk. For this reason, the VA is also referred to as the “Achilles's heel” of hemodialysis. The ideal VA is one that facilitates adequate long-term dialysis with few complications and is not costly to create and maintain.

The three VA types are described below:

i) A fistula is surgically created by connecting an artery to a vein; the high blood flow from the artery to the vein distends and toughens the vein to the degree of maturation. After fistula maturation, the VA is ready for the repeated needle insertions (cannulations) needed for hemodialysis. Advanced planning is required for fistula creation and use due to variable wait times for surgical assessment, creation, and its maturation (typically 2-6 months after creation);

ii) An AV-graft (Figure 1-bottom) is a synthetic polytetrafluoroethylene (Gortex) tube that is surgically connected between an artery and a vein; this AV-graft can then be cannulated for hemodialysis. Unlike the fistula, the AV-graft does not need to mature but usually requires 2-3 weeks before use to allow the swelling and discomfort to subside.

iii) A central venous catheter is a plastic tube inserted into the internal jugular (most common), subclavian, or femoral vein; it may be subcutaneously tunnelled and can be used immediately.

Participants/ population

We will include any study that collected data prospectively (observational cohort studies or randomized control trials) and followed patients for at least 3 months. We will only deem a study eligible if it described ≥ 100 vascular accesses in patients with chronic kidney disease (CKD) and being treated with chronic hemodialysis. We chose a minimum sample size of 100 vascular access (i.e. fistula) because some of our outcomes of interest are rare events (e.g. ischemic steal syndrome has an event rate >0 to 5%) and would require a larger sample size to obtain a representative (and less spurious) estimate of event rates.

We include full-text articles in any language published after December 31st, 1999. Studies must have reported information on one or more of the following:

- a) aneurysms
- b) infections/sepsis/bacteremia
- c) ischemic steal syndrome
- d) AV access thrombosis or catheter dysfunction
- e) venous hypertension
- f) overall complications
- g) radiological interventions
- h) surgical revisions.

We will exclude studies of peritoneal dialysis and pediatric patients (<18 years).

Intervention(s), exposure(s)

No intervention will be studied. We aim to summarize the rates of fistula complications for all studies published between January 2000 and December 2013.

Comparator(s)/ control

None

Context

See above

Outcome(s)

Primary outcomes

Studies must have reported information on one or more of the following primary or secondary outcomes:

Primary Outcomes and definitions:

- a) Aneurysm: Diffuse and progressive degeneration of the vascular access site. Patient has signs of bleeding, infection, or ulceration.
- b) Infections: Definite or probable local vascular access infections, vascular access-related sepsis, bacteremia or a composite of these infections.
- c) Ischemic steal syndrome: One or more clinical manifestations of pain, ischemic neuropathy, ulceration, and gangrene felt to be related to a fistula diverting blood from the distal circulation resulting in a zone of arterial insufficiency in the tissues distal to the fistula.
- d) Thrombosis: Absence of bruit or thrill, using auscultation and palpation, throughout systole and diastole at least 8

cm proximal to the arteriovenous anastomosis.

e) Venous hypertension: High pressure in veins due to damage to venous system.

None

Secondary outcomes

Secondary Outcomes:

- a) Bleeding
- b) Catheter Insertion
- c) Death
- d) Difficulty Cannulation
- e) Hematoma
- f) Hospitalization (All-Cause)
- g) Hospitalization (Infection-Related)
- h) Hospitalization (VA-Related)
- i) Infection- Bacteremia
- j) Infection- Local
- k) Radiological Intervention
- l) Surgical Revisions
- m) Total Complications
- n) Total Infections

None

Data extraction, (selection and coding)

Two authors will abstract data from all selected studies. Any disagreements will be resolved by consensus with a third author.

Risk of bias (quality) assessment

We will assess risk of bias among included studies, exploring participation, patient selection, attrition, exposure and outcome measurements, confounding, and selective reporting using a previously validated methods [1,2]. The results from study quality will not be considered at the analysis stage and planned synthesis.

[1] Busse J, Guyatt G. Instrument for assessing risk of bias in cohort studies. Available at: <http://www.evidencepartners.com/resources/>.

[2] Higgins JPT, Green S, eds. Cochrane Handbook for Systematic Reviews of Interventions Version 5.1.0 [updated March 2011]. The Cochrane Collaboration; 2011. Available at: Available from www.cochrane-handbook.org.

Strategy for data synthesis

Data Analysis: depending on the quality of the data and our ability to pool information across studies, we will analyze our results in one of two ways: 1) Summarize results of systematic review using descriptive statistics; or 2) pool our

results using meta-analytic methods.

1. Summarizing results using descriptive statistics: If the data quality is poor and outcomes of interest are highly heterogeneous, we will not be able to pool the results using meta-analytic techniques. The heterogeneity of the data may be attribute to differences in sampled populations, outcome definitions, prevalence of co-morbid conditions, and variable sample selection criteria. Hence, it may not be appropriate to calculate a summary statistic based on the weighted average. In which case, we will report the median and range for the event rate (per 1000 access-days) for the outcome of interest. When incidence rates are not reported in a selected study, we will calculate the overall follow-up time (denominator) by multiplying the mean follow-up time by the number of patients. We will use the overall follow-up time to calculate the event rate per 1000 patient-days. It is important to note that in using this method we assume that the hazard rate of developing a particular outcome is constant across individuals and over time.

OR

2. Pooling results using meta-analytic methods: If the data permits pooling of results, we will calculate the 95% confidence interval (CI) for the study estimate of interest using the Wilson Score method. The Wilson Score interval has been shown to provide excellent coverage and has better performance than the standard Wald interval.

We will pool the rates of the complication of interest using a random effects meta-analysis and a linear mixed model. This method assumes that the observed rates follow a Normal distribution. We will account for correlation between subgroup estimates from the same study, as well as estimates from different articles but from the same dialysis facility.

We will use the I-squared statistic to measure the proportion of total variation in study estimates that is due to heterogeneity rather than sampling error. When reported, we will calculate the pooled estimate for pre-specified subgroups including site of vascular access (e.g. lower vs. upper arm fistula), age (elderly vs. non-elderly as defined in the selected study), as well as study location (North America vs. Europe).

We will perform the analyses using SAS 9.3 (SAS Institute Inc., Cary, NC, USA) PROC MIXED procedure. This method will allow us to specify covariates in the random effects univariable meta-regression. We will explore heterogeneity between risk estimates according to the mean patient age, proportions of men, diabetic patients, and patients with peripheral vascular disease, number of fistulas, recruitment start date, and publication year.

In sensitivity analyses, we will exclude all studies that are published after 2000 but recruited patients prior to 2000 and studies that asked study question after data collection (i.e. retrospective design). In order to justify our analyses, we will require at least three independent estimates per subgroup. We will use a two-sided p-value and consider a p-value <0.10 to be statistically significant.

Analysis of subgroups or subsets

In subgroup analyses, we will examine the effect of fistula location (lower vs. upper arm), age (elderly vs. non-elderly), and location of study (North America vs. Europe) on our primary outcomes.

Dissemination plans

We plan to submit our final result to an academic Nephrology Journal

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Details of any existing review of the same topic by the same authors

None to our knowledge

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None known

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Reference and/or URL for protocol

http://www.crd.york.ac.uk/PROSPEROFILES/10444_PROTOCOL_20140527.pdf

Stage of review

Ongoing

Date of registration in PROSPERO

07 July 2014

Date of publication of this revision

08 July 2014

Stage of review at time of this submission	Started	Completed
Preliminary searches	Yes	Yes
Piloting of the study selection process	Yes	Yes
Formal screening of search results against eligibility criteria	Yes	Yes
Data extraction	Yes	Yes
Risk of bias (quality) assessment	Yes	Yes

Data analysis

Yes

Yes

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