Systematic Reviews of the Literature Are Not Always Either Useful Or the Best Way To Add To Science

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1 **REVIEW** 

# 2 Systematic Reviews of the Literature Are Not Always Either Useful Or the Best Way To Add To

- 3 Science
- 4

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14 Systematic reviews are becoming more popular as a way of doing research; however, not all

15 systematic reviews are clinically useful and sometimes another type of review (scoping, topical, or

16 critical) would be of greater value to the clinical and scientific community. The different types of

17 review and their use are described, illustrated by examples relevant to vascular surgery.

18

19 Keywords: Critical review, Scoping review, Systematic review, Topical review

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# 21 INTRODUCTION

22 Systematic reviews appear to have become increasingly popular in the published literature as often 23 they are considered as important sources of clinical evidence. The reason for performing systematic 24 reviews is to add information to what is known on a particular topic. However, all too often such 25 reviews are not clinically useful. Some have estimated that > 90% of systematic reviews are clinically useless, while others have suggested that the "gravy train" of systematic reviews constitutes 26 research waste.<sup>1,2</sup> The reasons for this include registered but unpublished reviews, duplicated or 27 28 poor methodological reviews, or those addressing questions which are not clinically useful. Do they 29 serve any other purpose? Perhaps they improve the citation or publication record of individual 30 researchers or clinicians or are considered a necessary starting point to a thesis? The former is not a laudable reason and the latter is a misplaced assumption. when perhaps another type of review, a 31 scoping review, would better summarise the field and identify the knowledge gaps and 32 33 opportunities for productive investigations. Other times they may add only incremental knowledge 34 rather than new knowledge. In contrast, good quality reviews are valuable to clinicians and guideline

committees, and can enhance the impact factor of a journal. This may be one of the reasons
underlying the steady increase in systematic reviews published in the *European Journal of Vascular and Endovascular Surgery* (Fig. 1). Scrutiny of the metrics of the 20 systematic reviews published in
2018 indicate the wide range of utility of the reviews to both journal impact factor and clinicians, but
six probably could be assigned to "the gravy train"<sup>2</sup> and take up journal pages at the expense of
original research.

41 The pyramids of clinical evidence all place systematic reviews of randomised controlled trials 42 (RCTs) at the apex, with reviews conducted with support of the Cochrane Collaboration on top. Such reviews are a special feature of the Cochrane Collaboration. For Cochrane reviews there is a 43 44 guaranteed rigorous search and review methodology, with emphasis on uncovering and reporting 45 potential sources of bias. Therefore, most Cochrane reviews focus on synthesising the data from 46 adequately powered RCTs and they can provide valuable additional information. The scientific 47 quality of any systematic review of RCTs is only as good as the quality of the included studies: small, 48 underpowered RCTs and those of poor methodological quality can provide very misleading results. 49 This risk of misleading information is even stronger in systematic reviews of observational studies where there is inherent patient selection, reporting bias, and overestimation of treatment effects is 50 51 common: these are the reviews most likely to be of limited clinical usefulness.<sup>3</sup> For reviews focusing 52 on long term outcomes, the number of patients lost to follow up should be assessed as part of the 53 study appraisal. Patient loss to follow up is usually higher in observational studies than in 54 randomised controlled trials, and this can limit the value of observational studies when synthesising evidence for longer term outcomes. 55

56 Here the discussion is when it is appropriate to consider undertaking a systematic review, 57 with a few tips for success, when a scoping review would be better than a systematic review, and 58 when a critical or topical review of recent evidence would be more appropriate.

59 WHAT IS A SYSTEMATIC REVIEW AND WHEN IS IT USEFUL?

A systematic review is used to marshal, appraise, and synthesise the evidence about a precise clinical
question, for example, comparing the clinical effectiveness of two different operations for the same
condition. A systematic review should follow standard rigorous methodology and reporting,<sup>4</sup> and
there are recommended approaches to minimise bias. The PICO (population, intervention,
comparator, outcomes) for the review all require clear and careful definition to ensure limited
clinical heterogeneity. Given these criteria, such reviews should not be considered as a quick and
easy way of doing research.

67 There are several different uses of a systematic review.

- 1 To synthesise the evidence from adequately powered (large) RCTs, these are likely to be Cochranereviews.
- 70 2 To synthesise the evidence from observational studies comparing the efficacy of treatments in
- 71 situations where randomised trials are not possible, for example, the efficacy of e-cigarettes to
- 72 promote and sustain smoking cessation.<sup>5</sup>
- 73 3 To synthesise the evidence from RCTs and observational studies about a clearly defined important
- clinical question, to which the answer is not already known and there is no evidence of a similar
- review being in progress or recently published (by checking PROSPERO and other research registries
- as well as conference abstracts), for example, is carotid artery stenting still associated with lower
- 57 stroke risk in asymptomatic patients given the advances in best medical therapy? Careful definition
- of the PICO in question and quality assessment of included studies are vital. If sufficient suitable
- real studies are identified, meta-analysis should be provided as well as sensitivity analysis for the best
- 80 quality studies, in cases where there is a wide range of study quality.
- 4 To investigate how outcomes have changed over time, to identify whether there has been
- 82 improvement in outcomes and patient benefit. Meta-regression can be a useful tool. An example
- 83 here is the recent updating of a 2010 systematic review evaluating the sex specific operative
- 84 mortality from intact abdominal aortic aneurysm repair, given the advances in both endograft and
- 85 imaging technology.<sup>6,7</sup>
- 5 To investigate how factors such as age, sex, ethnicity, and frailty influence clinical outcomes using
- 87 evidence from both RCTs and observational studies, for example, the influence of age, sex, and
- 88 contralateral occlusion on stroke and death after carotid endarterectomy or carotid stenting.<sup>8</sup>
- Presentation of sensitivity analyses to compare information obtained from RCTs *versus* observational
  studies may be illuminating.
- 91 6 To inform clinical practice guidelines about recent developments, for example, pre-emptive
- 92 procedures to limit type II endoleak after aneurysm repair using evidence from observational studies
- 93 and/or small RCTs. To avoid bias from small studies, it can be helpful to use a minimum threshold for
- 94 the number of patients as an inclusion criterion. In the absence of sufficient evidence, this would
- 95 convert to a topical review.
- 96 7 To obtain parameters for use in modelling studies or for estimating the sample size for a planned
- 97 large RCT. Examples might include recent changes in amputation rates to inform the provision of
- 98 services for amputees following the COVID 19 pandemic or current prevalence of abdominal aortic
- 99 aneurysm to inform the likely effectiveness of population screening programmes.<sup>9</sup>
- 100 8 To identify the range of reported outcomes, for example, for the development of Core Outcome
- 101 sets or to identify the full range of procedure associated complications.<sup>10</sup>

#### 102 THE TIMING OF SYSTEMATIC REVIEWS

- 103 The timing of systematic reviews is important, as the most cited and downloaded reviews address a
- 104 still controversial topic for which sufficient evidence is available but do not come too late to be
- 105 useful, after clinical practice has changed. Illustration of this point comes from analysis of the
- 106 citation and download rates of systematic reviews published in the European Journal of Vascular and
- 107 *Endovascular Surgery* in 2018. The most cited and downloaded review was on how to treat type II
- 108 endoleak. A review of thrombotic events after endothermal ablation of the great saphenous vein
- also appeared useful. These issues were clearly worries on many people's minds and were topics
- 110 likely to influence clinical practice. In contrast, the role of Nordic walking in exercise programmes
- 111 aroused little interest.

## 112 WHEN IS A SYSTEMATIC REVIEW EITHER NOT NEEDED OR UNHELPFUL?

113 There are several clear examples of when a systematic review is unlikely to be of clinical value.

- 114 1 When a recent systematic review is already available or in progress (check in PROSPERO<sup>11</sup> and
- 115 other research registries).
- 116 2 To answer questions that do not concern clinical effectiveness (PICO not applicable), for example
- 117 what is the best method of measuring the size of large venous ulcers? This needs an overview of
- 118 measurement methods.
- 119 3 To answer questions, where the answer is already known, for example, is the operative mortality
- 120 for intact AAA repair in women lower after EVAR or open repair? The answer here can be derived
- 121 from the interaction analyses in randomised trials and the systematic reviews of sex specific
- 122 differences.
- 4 To use observational data to answer questions that can be answered only by RCTs, for example,
- 124 what is the diameter threshold for repairing internal iliac aneurysms? This might be the subject of
- either a topical review if there are new data for the rupture of these aneurysms or a critical reviewof the literature.
- 127 5 To answer questions where there are no standard interventions or outcomes. An example is
- 128 provided by the recent review of pre-habilitation interventions before elective aneurysm repair.<sup>12</sup> A
- scoping review probably would have been more useful.

### 130 WHAT IS A SCOPING REVIEW AND WHEN IS IT NEEDED?

- 131 A scoping review is an exploratory but systematic literature search to find out how much is known
- about a broad topic or to discover gaps in the evidence, to provide a narrative review without formal
- 133 meta-analysis. The question(s) being addressed is broader and less specific and also may be more
- 134 complex and heterogeneous than that in a systematic review. Examples include "Do prisoners have
- adequate access to vascular services?" or "What is the evidence for shared decision making for

- Journal Pre-proof
- 136 critical limb ischaemia?". It might be used to identify whether a systematic review was necessary. A
- 137 scoping review can identify specific unanswered questions which can be addressed either with new
- 138 original research or some that can be answered by a systematic review. At the start of a thesis or
- 139 other piece of research work, a scoping review often is more useful and less labour intensive than a
- 140 systematic review.

### 141 WHAT IS A TOPICAL REVIEW AND WHEN IS IT NEEDED?

A topical review is an up to date overview of a current hot topic. Topical reviews may present areas 142 143 that are still developing rapidly and may provide an indication of the future direction of the field. 144 Examples might include the value of high sensitivity troponin assays to guide the management of peripheral arterial disease or methods for local, rather than systemic, antithrombotic therapy.<sup>13–15</sup> As 145 146 with the previous types of reviews they need to be systematic and thorough, but unlike systematic 147 or scoping reviews, they are guided by the literature and make more use of conference abstracts and 148 grey literature such as scientific and charitable foundation reports, government or industry reports. 149 The review should report not just specific outcomes but must include the key present and future 150 issues and/or challenges, which need to be addressed. Therefore, such reviews may be needed by government bodies and industry, as well as keeping clinicians informed about emerging technologies 151

and processes.

# 153 WHAT IS A CRITICAL REVIEW AND WHEN IS IT NEEDED?

154 A critical review is both an appraisal and a critique of new data on a topic, which may be either 155 controversial or inconsistent with earlier findings and guidelines. A potential example is if a new, 156 large series providing the diameter of ruptured internal iliac aneurysms indicates that the suggested 157 intervention diameter criterion in clinical guidelines needs to be revised. A real example is the recent 158 population based study from Denmark, which suggests that diabetes is not a factor that protects against the development of abdominal aortic aneurysm, although it may be protective of the 159 development of more proximal aneurysms.<sup>16</sup> The critical review then becomes a critique of the new 160 study set in the context of a critique of the previous evidence (which did not come from population 161 162 based studies).

### 163 HOW DO THE PROCESSES FOR THE VARIOUS REVIEW TYPES DIFFER?

- 164 The processes for these four different types of review are summarised and compared in Table 1. The
- 165 varying types of review described have different purposes and methodology but all should be
- thorough and systematic. There are some other specialist types of review, for example individual
- 167 patient meta-analyses of randomised trials but these require full access to original data and
- 168 specialist statistician input.
- 169 SO WHAT REVIEW DO YOU NEED?

- 170 The aim of this paper is to help you decide what kind of review you need to undertake and to
- discourage inappropriate systematic reviews, which are not likely to be clinically or scientifically
- useful and divert resources away from research productivity. Finding the gaps in the evidence, to
- 173 which you can contribute original research, may often be more rewarding than a systematic review.

## 174 CONFLICT OF INTEREST

- 175 None.
- 176 FUNDING
- 177 None.
- 178
- 179 Figure 1. The increasing number of systematic reviews published in European Journal of Vascular
- 180 *and Endovascular Surgery*. Data show the number of systematic reviews published from 2010 to
- 181 date on the vertical axis: the 2021 data show reviews recorded in Medline to end August 2021.



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Table 1. Methodology of systematic, scoping, topical, and critical reviews.						
Stages	Systematic	Scoping	Topical	Critical		
Question	Formulate the	Decide on the	What is the	Is the new evidence		
	precise question	broad topic	current knowledge	robust?		
			base?			
Checks	PROSPERO <sup>11</sup> and	Medline search for	Recent flagship	Recent flagship		
before you	other databases	reviews on the	scientific journals	scientific journals for		
start	for existing or	topic	for similar reviews	similar reviews		
	similar review					
Making the	Inclusion and	Not usually	Only after initial	Not applicable		
question	exclusion criteria	relevant	review of the key			
more	for relevant		literature			
detailed	studies					

Search for evidence	Use a minimum of two databases	Use a wide range of databases (to include nursing, social sciences, etc., as necessary)	By keywords in Medline, grey literature including conference and foundation reports	By keywords in Medline or scientific literature. conference proceedings for
				unpublished support
Select and	Use a minimum of	Use a minimum of	Guided by what	Guided by the new
extract	two researchers	two researchers	you find and limit	evidence
evidence			to the most	
			pertinent reports	
Evidence	Needs formal	Not assessed	Validity of	Must be assessed:
quality	assessment.		evidence needs	key part of the
	Sensitivity		discussion	critique
	analysis of good		C .	
	quality studies			
Outputs	Usually data	Tables of evidence	Key themes and	Narrative viewpoint
	synthesis with	with narrative	issues	and future data
	meta-analysis	synthesis		required
Reporting	PRISMA⁴	PRISMA extension	N/A	N/A
guidelines		for scoping reviews <sup>17</sup>	X	

185 N/A = not applicable.

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# 188 **REFERENCES**

189 1 Ionnadis J. The mass production of redundant, misleading and conflicted systematic reviews and

190 meta-analysis. *Millbank Q* 2106;**94**:485–514.

191 2 Roberts I, Ker K. How systematic reviews cause research waste. *Lancet* 2015;**386**:1536.

192 3 Jepsen P, Johnsen SP, Gillman MW, Sorensen HT. Interpretation of observational studies. *Heart* 

193 2004;**90**:956–60.

194 4 The PRISMA 2020 statement: an updated guideline for reporting systematic reviews. Available at:

195 <u>https://www.equator-network.org/reporting-guidelines/prisma/</u> [Accessed 10 August 2021].

196 5 Kalkhoran S, Glantz SA. E-cigarettes and smoking cessation in the real-word and clinical settings: a

197 systematic review and metanalysis. *Lancet Respir Med* 2016;**4**:116–28.

198 6 Grootenboer N, van Sambeek MRHM, Arends IR, Hendriks JM, Hunink MGM, Bosch JL. Systematic

review and meta-analysis of sex differences in outcome after intervention for abdominal aortic

200 aneurysm. Br J Surg 2010;97:1169–79.

201 7 Pouncey AL, David M, Morris RI, Ulug P, Martin G, Bicknell C, et al. Editor's Choice - Systematic

202 review and meta-analysis of sex specific differences in adverse events after open and endovascular

- 203 intact abdominal aortic aneurysm repair: consistently worse outcomes for women. *Eur J Vasc*
- 204 Endovasc Surg 2021;**62**:367–78.

- 205 8 Touze E, Trinquart L, Felgueiras R, Rerkasem K, Bonati LH, Meliksetyan G, et al. A clinical rule (sex,
- 206 contralateral occlusion, age, and restenosis) to select patients for stenting versus carotid
- 207 endarterectomy: systematic review of observational studies with validation in randomized trials.
- 208 Stroke 2013;**44**: 3394–400.
- 209 9 Chan WK, Yong E, Hong Q, Zhang L, Lingam P, Tan GWL, et al. Systematic review and meta-analysis
- of the prevalence of abdominal aortic aneurysm in Asian populations. *J Vasc Surg* 2021;**73**:1069–74.
- 211 10 Machin M, Ulug P, Pandirajan K, Bown MJ, Powell JT. Towards a core outcome set for abdominal
- 212 aortic aneurysm: systematic review of outcomes reported following intact and ruptured abdominal
- aortic aneurysm repair. *Eur J Vasc Endovasc Surg* 2021;**61**;909–18.
- 214 11 National Institute for Health Research. PROSPERO, International prospective register of
- 215 systematic reviews. Available at: https://www.crd.york.ac.uk/prospero/.
- 216 12 Bonner RJ, Wallace T, Jones AD, Scott DJ, Richards SH. The content of prehabilitative
- 217 interventions for patients undergoing repair of abdominal aortic aneurysms and their effect on post-
- 218 operative outcomes: a systematic review. *Eur J Vasc Endovasc Surg* 2021;**61**:756–65.
- 219 13 Matsushita K, Kwak L, Yang C, Pang Y, Ballew SW, Sang Y, et al. High-sensitivity cardiac troponin
- and natriuretic peptide with risk of lower-extremity peripheral artery disease: the Atherosclerosis
- Risk in Communities (ARIC) Study. *Eur Heart J* 2018;**39**:2412–9.
- 14 Degen H, Borst O, Ziegler M, Munoz A-K, Jamashi J, Walker B, et al. ADPase CD39 fused to
- glycoprotein VI-Fc boosts local antithrombotic effects at vascular lesions. JAHA 2017;6:e005991.
- 15 Ye W, Wang N, Hu K, Zhang L, Pan C, Gong Y, et al. Bio-inspired microcapsule for targeted
- antithrombotic drug delivery. *RSC Adv* 2018;**8**:27253–9.
- 226 16 Obel LM, Diederischen AC, Steffensen FH, Frost L, Lambrechtsen J, Busk M, et al. Population-
- 227 based risk factors for ascending, arch, descending and abdominal aortic dilations in 60-74 year-old
- individuals. J Am Coll Cardiol 2021;**78**:201–211.
- 229 17 Tricco AC, Lillie E, Zarin W, O'Brien KK, Colquhoun H, Levac D, et al. PRISMA Extension for scoping
- reviews (PRISMA-ScR): checklist and explanation. *Ann Intern Med* 2018;**167**:467–73.

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**Figure 1 The increasing number of systematic reviews published in European Journal of Vascular and Endovascular Surgery**. Data show number of systematic reviews published from 2010 to date on the vertical axis: the 2021 data show reviews recorded in Medline to end August 2021.

