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Ultrasound measurement of traumatic scar and skin thickness: A scoping review of evidence across the translational pipeline of research-to-practice

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1	Ultrasound measurement of traumatic scar and skin thickness: A scoping review of evidence
2	across the translational pipeline of research-to-practice*
3	Running Title: Review of scar thickness measurement with ultrasound
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- 29 (ANZBA) 2022 Annual Scientific Meeting, the 2022 Centre for Children's Health Research
- 30 Symposium, Child Health Research Centre, The University of Queensland, and the 2023
- 31 British Burn Association Annual Conference.

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32 ABSTRACT:

Objectives: To identify the ultrasound methods used in the literature to measure traumatic
 scar thickness, and map gaps in the translation of these methods using evidence across the
 research-to-practice pipeline.

Design: Scoping review

Data Sources: Electronic database searches of Ovid MEDLINE, Embase, Cumulative Index
of Nursing and Allied Health Literature (CINAHL) and Web of Science. Grey literature
searches were conducted in Google. Searches were conducted from inception (date last
searched 27/05/2022).

Data Extraction: Records using B-mode ultrasound to measure scar and skin thickness across the research-to-practice pipeline of evidence were included. Data was extracted from included records pertaining to: methods used; reliability and measurement error; clinical, health service, implementation and feasibility outcomes; factors influencing measurement methods; strengths and limitations; and use of measurement guidelines and/or frameworks. **Results:** Of the 9309 records identified, 118 were included for analysis (n = 82 journal articles, n = 36 abstracts) encompassing 5213 participants. Reporting of methods used was poor. B-mode, including high-frequency (i.e., > 20 MHz) ultrasound was the most common type of ultrasound used (n = 72; 61%), and measurement of the combined epidermal and dermal thickness (n = 28; 24%) was more commonly measured than the epidermis or dermis alone (n = 7, 6%). The scar characteristics most commonly reported to be measured were epidermal oedema, and dermal fibrosis and hair follicle density. Most records analysed (n = 115; 97%) pertained to the early stages of the research-to-practice pipeline, as part of research initiatives.

Page 5 of 77

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55 **Conclusions:** The lack of evaluation of measurement initiatives in routine clinical practice 56 was identified as an evidence gap. Considerations for the ultrasound measurement of 57 cutaneous traumatic scarring in research and clinical practice are presented based on the 58 review findings. Standardising the core methodological components of ultrasound 59 measurement is recommended based on poor methodological reporting in some records.

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STRENGTHS AND LIMITATIONS OF THIS STUDY:

Evidence pertaining to the implementation of ultrasound measurement in routine
 clinical practice and research-to-practice gaps were determined by categorising
 included records into one of the four Australian Government Department of Health
 and Aged Care Medical Research Future Fund research-to-practice pipeline phases.
 Clinical, health service, implementation and feasibility outcomes related to ultrasound
 measurement in included records were summarised to determine what is needed to
 close the research-to-practice gap for ultrasound measurement of scar thickness.

- The reported methods compiled in this review were used to inform the development of nine methodological considerations to guide health practitioners and researchers using ultrasound to measure scar and skin thickness.
- A limitation is that only articles available in English or with an English abstract were considered for inclusion and data extraction, although the large number of included records means this is unlikely to have changed the review findings.

75 INTRODUCTION:

Traumatic cutaneous injury, caused by sharp object penetration (e.g., surgery or vaccination) or burns (including thermal, chemical and friction) may result in the formation of hypertrophic scarring.¹ While major injuries to non-fetal skin heal through the formation of scar tissue, most resultant scars are small, linear and/or barely visible.²⁻⁵ Hypertrophic scars, however, result from an aberrant healing response that leads to the formation of red, raised scars, often accompanied by pruritus and skin tightening, which remain within the boundaries of the initial injury.⁶⁻¹¹ The sequelae of hypertrophic scars have the potential to impact patient's physical and psychosocial quality of life. ^{12 13}

A characteristic of hypertrophic scarring that both patients and clinicians have identified as being important, and which has subsequently been used as a way to measure clinical and treatment outcomes, is scar thickness.¹³⁻²¹ Scar thickness can be measured both subjectively, through clinician assessment and patient-reported outcomes, or objectively, utilising medical imaging methods. ^{22 23} The pathological complexity of hypertrophic scars means that they generally extend below the level of the surrounding skin, supporting the use of medical imaging modalities such as ultrasound for thickness quantification, as these are capable of providing information about subcutaneous structures and processes. ^{23 24} Scar thickness measurement using ultrasound can be conducted in both clinical and research contexts. For example, ultrasound is regularly used in our own clinical practice at the Pegg Leditschke Children's Burn Centre to measure scar thickness, ²⁵ particularly prior to treatment with ablative fractional CO₂ laser, where scar thickness measurement is used to determine the required depth of penetration.²⁶ The routine use of measurements like ultrasound to guide clinical decision-making has been termed measurement-based care.²⁷

98 There are several clinical skills that assessors or practitioners are required to master to
99 effectively conduct measurement of scar thickness with ultrasound. Proficiency in these skills

Page 7 of 77

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BMJ Open

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may aid in bridging the gap between its use predominantly in research methods, to more 00 widespread use in routine clinical practice. These skills have previously been described as 01 part of a training curriculum in point-of-care ultrasound, ²⁸ but can equally be applied to scar 02 thickness measurement using ultrasound. These include: 1) understanding when to conduct 03 scar thickness measurement with ultrasound; 2) the ability to operate an ultrasound machine 04 to obtain useful images; 3) the ability to recognise physiological and pathological skin 05 06 features (i.e., epidermis, dermis) on ultrasound images, and be able to measure the thickness of each; and 4) successfully utilising the thickness measurement as the basis of measurement-07 based care, including quantifying changes in scar thickness in response to treatment.²⁸ For)8 example, a study of ultrasound measurement in people with systematic sclerosis identified)9 that ultrasound may be capable of differentiating between early oedema and fibrosis, and 10 detect thickening before it is observed clinically, thus providing opportunities to prevent or 11 treat fibrosis early.²⁹ Whilst it is ideal for all ultrasound assessors to have the skills 12 mentioned, the number of assessors required to be proficient in these skills differs in research 13 14 and clinical practice. In research, a small number of researchers are generally responsible for conducting ultrasound measurement, whereas members across the entire multi-disciplinary 15 team may be required to conduct these measurements in routine clinical practice, and thus 16 17 require some level of proficiency.³⁰ Ultrasound, itself, is a popular, safe, non-invasive and largely cost-effective (compared to 18

other imaging modalities) imaging method with measurement utility in both adult and paediatric populations. $^{25 31 32}$ Modern B-mode (brightness mode) ultrasound, particularly high- (i.e., ≥ 20 MHz) or ultra-high frequency (30-100 MHz) 33 ultrasonography, has the capacity to allow differentiation between the epidermis and dermis, permitting quantification of skin layer-specific scar characteristics. This allows assessors to observe and understand the pathological mechanisms of individual scars and adjust treatment protocols accordingly. $^{25 34-}$

³⁹ Indeed, measurement of scar thickness using these methods may allow quantification of
fibrosis and oedema within the scar, and can also be used to distinguish scar tissue from
uninjured skin by measuring the presence and density of hair follicles. ⁴⁰⁻⁴³ B-mode
ultrasound is also commonly used as the basis for other imaging methods, such as colour
Doppler ultrasound or elastography, which can allow quantification of additional scar
characteristics, such as their elastic properties. ^{34-37 44 45}

Despite the clinical advantages of B-mode ultrasound for scar thickness measurement, methods utilised in the literature are poorly reported and lack standardisation. This casts doubt on the validity of clinical decision-making in measurement-based care initiatives (e.g., setting depth of AFCO₂ penetration) informed by research findings (e.g., response to treatment) where ultrasound measurements are used. ⁴⁶ Lack of standardisation also makes between-study comparison, such as systematic reviews and meta-analyses, difficult, ⁴⁷ and poor methodological reporting hampers the ability to accurately replicate findings. This scoping review focusses on mapping and identifying gaps in ultrasound methods and evaluation reported in the current literature along the research-to-clinical practice pipeline. Methodological considerations for assessors or practitioners performing scar thickness measurements using ultrasound are presented based on the review findings.

METHODS:

Protocol Publication and Review Structure:

The protocol for this review has been published *a priori*. ⁴⁸ This scoping review was conducted and is reported according to the Arksey and O'Malley (2005) ⁴⁹ framework. The steps outlined in this framework are: 1) identifying the research question; 2) identifying relevant records; 3) selecting appropriate records; 4) charting extracted data; and 5) collating, summarising and reporting the results. ⁴⁹

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Research Question:

The primary question of this scoping review was: "What do we know and not know about the measurement of traumatic cutaneous scar thickness using ultrasound?" This question was addressed through exploration of: methods used; reliability and measurement error; clinical, health service, implementation and feasibility outcomes; factors influencing ultrasound imaging and measurement methods; strengths and limitations of measurement methods; and use of measurement guidelines and/or frameworks. While the focus of this review was the measurement of traumatic cutaneous scar thickness with ultrasound, methods used to measure the thickness of unscarred skin were reported where these were used in combination with measurement of scar thickness (e.g., as control or comparator measurements).

159 Identifying Relevant Records:

A standardised search strategy was developed and piloted with the assistance of a medical
librarian using the concepts 'ultrasound', 'skin', 'thickness' and 'measure', with associated
terms and truncations (supplementary box 1). Ovid MEDLINE, Embase, Cumulative Index of
Nursing and Allied Health Literature (CINAHL) and Web of Science electronic databases
were searched from conception to identify original studies (date last searched 27th May
2022).

The phrase 'ultrasound scar thickness measurement' was used to conduct additional searches
in 1) Google Scholar, and 2) Google to identify original studies in grey literature, and studies
not identified in database searches. Title and abstract searches in Google Scholar and Google
were limited to the first 200 results. ⁵⁰

Record Selection:

Following de-duplication, six reviewers screened records using Covidence (Veritas Health
Innovation, Melbourne, Australia; available at www.covidence.org) for eligibility according

Page **8** of **45**

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to the inclusion criteria (Table 1). During both title and abstract and ful	l text screening, one
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174 researcher (BM) screened all records as a single reviewer, while other researchers (MS, TM,

175 TR, BD and ZT) screened records as a second reviewer. Conflicts were resolved through

- 176 discussion between at least two authors to reach agreement. A third author was used as a
- 177 tiebreaker where agreement could not be reached.

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178 Table 1. Inclusion and exclusion criteria for studies included in the scoping review.

 Traumatic scars measured with ultrasound based on B-mode ultrasound (including high-frequency, ultra-high-frequency and Doppler) Measurements taken of living, human individuals Measurement of traumatic cutaneous scarring arising from penetration of the skin with sharp objects (including thermal, chemical or friction) Articles written in English, or with English abstracts Charting the Data: The data extraction table was developed in Microsoft Excel and piloted by two authors (BM and ZT) through independent extraction and comparison of data from two records. The table was then modified to include the scar characteristics (e.g., fibrosis, oedema) measured, measurer/assessor training, and the number of measurements taken (Supplementary Table 1) Full text data extraction was completed by four authors (BM, MS, TM and ZT). An additional author (BD) independently extracted data from five randomly selected records, which was compared to data extracted by other authors. Minimal differences between data extracted by the independent author and that by other authors were observed, thus further 		Inclusion	Exclusion
English abstracts 179 180 Charting the Data: 181 The data extraction table was developed in Microsoft Excel and piloted by two authors (BM 182 and ZT) through independent extraction and comparison of data from two records. The table 183 was then modified to include the scar characteristics (e.g., fibrosis, oedema) measured, 184 measurer/assessor training, and the number of measurements taken (Supplementary Table 1) 185 Full text data extraction was completed by four authors (BM, MS, TM and ZT). An 186 additional author (BD) independently extracted data from five randomly selected records, 187 which was compared to data extracted by other authors. Minimal differences between data 188 extracted by the independent author and that by other authors were observed, thus further		 Traumatic scars measured with ultrasound based on B-mode ultrasound (including high-frequency, ultra-high-frequency and Doppler) Measurements taken of living, human individuals Measurement of traumatic cutaneous scarring arising from penetration of the skin with sharp objects (including surgery or vaccination), or as a result of burns, (including thermal, chemical or friction) Articles written in English, or with 	 Reviews, discussion papers, opinion pieces Measurement of non-traumatic scars (e.g., acne scars) Measurement of skin thickness in non-traumatic conditions (e.g., diabetes) Measurement of skin thickness where there is no cutaneous involvement in the trauma (e.g., traumatic brain injury) Measurement using A-mode ultrasound
 Charting the Data: The data extraction table was developed in Microsoft Excel and piloted by two authors (BM and ZT) through independent extraction and comparison of data from two records. The table was then modified to include the scar characteristics (e.g., fibrosis, oedema) measured, measurer/assessor training, and the number of measurements taken (Supplementary Table 1) Full text data extraction was completed by four authors (BM, MS, TM and ZT). An additional author (BD) independently extracted data from five randomly selected records, which was compared to data extracted by other authors. Minimal differences between data extracted by the independent author and that by other authors were observed, thus further 	179	English abstracts	
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Page 11 of 77

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BMJ Open

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independent extraction was not performed. As is typical in scoping reviews, the certainty or 189 quality of evidence was not appraised. 49 190

The research-to-practice pipeline published by the Australian Government Department of 191 Health and Aged Care Medical Research Future Fund (figure 1) was used to categorise each 192 included record based on their stated aims into one of the four phases. ⁵¹ The final phase of 193 this pipeline (phase 4) indicates initiatives used in routine clinical practice. 194

Where clinical (e.g., treatment satisfaction, scar symptoms), health service (e.g., efficiency, 195 196 safety, effectiveness, equity, patient-centredness and timeliness) and implementation (e.g., acceptability, adoption, appropriateness, fidelity, cost, penetration and sustainability) 197 outcomes were addressed, they were reported according to Proctor et al. 52. Measurement 198 instrument-specific feasibility outcomes defined by Prinsen et al. 53 are reported in the current 199 200 review. These outcomes included ease of administration, standardisation, completion time, instrument cost and availability, and ease of score calculation. ⁵³ Reliability and measurement 201 error were defined according to COnsensus-based Standards for the selection of health 202 Measurement INstruments (COSMIN) tools. 54 55 Measurements with an intraclass correlation 203 coefficient (ICC) of 0.7 or greater were considered reliable.⁵⁵ Measurement error was 204 assessed by comparing the reported standard error of the measurement (SEM) with the 205 reported smallest detectable change (SDC). Where the reported measurement error was 206 207 smaller than the reported smallest detectable change, it was interpreted as indicating real change or variance can be detected, and that change or variance is not a result of error.⁵⁵ 208 **Patient and Public Involvement** 209 210 There was no patient and/or public involvement in the design, conduct, reporting or dissemination of information in this scoping review. 211 **RESULTS:** 212

Electronic database searches identified 9309 records. After removal of 3703 duplicate records, the titles and abstracts of 5606 records were screened for relevance according to the inclusion criteria (Table 1). Following full-text screening, 104 records proceeded to data extraction. Searches in Google and Google Scholar identified an additional 14 records, providing a total of 118 records for data extraction. Search and screening results are presented according to the Preferred Reporting Items for Systematic reviews and Meta-Analyses (PRISMA) flow diagram (figure 2).⁵⁶ **Record Characteristics:** Of the 118 records included in this review, 82 were journal articles (69%) and 36 were abstracts (31%) (Table 2), representing a total of 5213 participants. The majority (n = 44; 37% of included records) measured adults aged 8 years or older, ^{21 26 34 36-38 42 43 57-94} and were measurements of burn scars (n = 69 records; 58%) (Table 2). 21 25 26 31 32 35 36 38 42 59 64 $^{66-71}$ $^{74-78}$ 84-87 91 92 94-133 Most identified records used ultrasound measurement of scar thickness as part of research initiatives, and were categorised as either phase 2 (n= 70; 59%)^{21 31 34 39 42 43 57 59-63} $66\ 67\ 69\ 70\ 73\ 74\ 77-80\ 83\ 84\ 87-90\ 92\ 93\ 95-98\ 100-102\ 105-107\ 113\ 116-122\ 124-126\ 128\ 133-151$ or phase 3 (n = 45; 38%) 25 26 32 35-38 58 64 65 68 71 72 75 76 81 82 85 86 91 94 103 108 110-112 114 115 123 127 129-132 152-161 on the research-to-practice pipeline. ⁵¹ Three records (3%)^{99 104 109} used ultrasound to measure treatment response to an intervention already used in routine clinical practice (phase 4), including compression garments99 109 and CO2 fractional laser. 104

Table 2. Characteristics of records included in this review

First Author	Sample	Population Type	Scar	Translational
(year)	Size (n)		Aetiology	Pipeline Phase*
Journal articles				
Agabalyan (2017)	10	Adult	Not	2
			specified	
Alsharnoubi (2018)	15	Paediatric	Burn	2
Alsharnoubi (2018)	15	Paediatric	Burn	2
Alshehari (2015)	30	Not reported	Mixed	2

Page 11 of 45

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3	Avetikov (2018)	50	Paediatric & adult	Not	3
4				specified	
5	Berry (1985)	16	Paediatric & adult	Burn	4
6 7	Blome-Eberwein	16	Paediatric & adult	Burn	2
8	(2012)	10		2	-
9	Blome-Eberwein	36	Adult	Not	2
10	(2016)	50	1 Idult	specified	2
11	(2010) Blome-Eberwein	10	A dult	Burn	2
12	(2010)	17	Auun	Dum	2
13	(2019) Coi (2010)	51	A dult	Not	2
14	Cal (2019)	51	Auun	not	Z
15	$C_{andry}(2010)$	17	A .J. 14	Specified	2
16 17	Candy (2010)	1/	Adult	NOT	2
17	C^{1} (2017)		4 1 1	specified	2
19	Chae (2016)	23	Adult	Not	3
20				specified	_
21	Chang (2014)	60	Adult	Surgical	2
22	Chan (2004)	56	Paediatric & adult	Burn	2
23	Cheng (2001)	58	Paediatric	Burn	3
24	Cho (2014)	146	Not reported	Burn	2
25	Danin (2012)	22	Paediatric & adult	Burn	3
26	Deng (2019)	20	Adult	Not	2
2/				specified	
20	Deng (2021)	31	Adult	Not	2
30				specified	
31	Deng (2021)	45	Adult	Not	2
32				specified	
33	Dunkin (2007)	113	Adult	Surgical	2
34	Elrefaie (2007)	22	Paediatric & adult	Not	2
35	Eliciaie (2020)			specified	-
36	Engrav (2010)	67	Paediatric & adult	Burn	4
37	Englav (2010) Fabbrocini (2016)	20	$\Delta dult$	Mixed	2
38	Faborocini (2010) Fong (1007)	20 16	Dadiatric & adult	Burn	2 3
39 40	$\frac{1997}{1000}$	10	Paodiatria & adult	Mixed	3
41	Fraccalvieri (2013)	5		Mixed	2
42	$\frac{1}{2011}$	3		Niixed	2
43	Gankande (2014)	30	Adult	Burn	3
44	Ge (2022)	21	Paediatric & adult	Mixed	3
45	Gee Kee (2016)	43	Paediatric	Burn	2
46	Guo (2020)	87	Paediatric & adult	Not	3
47				specified	
48	Huang (2017)	1	Adult	Burn	3
49	Huang (2021)	5	Adult	Burn	3
50	Huang (2020)	43	Adult	Not	3
52				specified	
53	Issler-Fisher (2021)	187	Adult	Burn	2
54	Issler-Fisher (2020)	78	Adult	Burn	3
55	Issler-Fisher (2017)	47	Paediatric & adult	Burn	3
56	Joo (2020)	48	Adult	Not	2
57		- 0		specified	-
58	Katz (1985)	4	Not reported	Burn	3
59	Kemn Bohan (2021)	21	Not reported	Burn	3
60	1.2021)	<i>L</i> 1	rotropolica	L'UIII	5

2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18	Kim (2018) Lacarrubba (2008) Lau (2005) Lee (2020) Lee (2019) Li (2013) Li (2020) Li (2021) Li (2021) Li (2021) Li (2021) Li-Tsang (2005) Li-Tsang (2006)
19 20	Lobos (2017)
21 22 23	Mamdouh (2021)
24	Meirte (2016)
25	Miletta (2021)
26	Nedelec (2014)
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20	Nedelec (2019)
30	Nedelec (2020)
31	Nicoletti (2015)
32	Niessen (1998)
33	Reinholz (2020)
34	Reinholz (2016)
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37	Schwaiger (2018)
38	Simons (2017)
39	Soykan (2014)
40	Timar-Banu (2011
41	Ud-Din (2019)
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44	van den Kerckhov
45	(2005)
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48	van der Veer (201
49 50	Wang (2009)
50	Wang (2010)
52	Wiseman (2020,
53	2021)
54	Wood (1996)
55	Xuan (2021)
56 57	
57 58	Yeol Lee (2022)
59	Yim (2010)
60	Zadkowski (2016)

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n (2018)	148	Not reported	Burn	2	3
arrubba (2008)	8	Paediatric & adult	Mixed		>
(2005)	100	Paediatric & adult	Burn	-	>
e(2020)	55	Adult	Burn		2
e(2019)	55	Adult	Burn	2	2
(2013)	7	Adult	Burn	-	>
(2020)	21	Paediatric & adult	Mixed	-	>
(2021)	165	Paediatric	Mixed	-	>
(2018)	34	Adult	Burn	-	3
(2021)	105	Adult	Burn		2
(2001) Tsang (2005)	101	Adult	Surgical	-	3
Tsang (2006)	45	Adult	Not		,
1 Sung (2000)	15	riduit	specified	4	-
Tsang (2010)	104	Paediatric & adult	Mixed	2	,
13 ang (2010)	35	Paediatric & adult	Not	2	ž
005 (2017)			specified		,
mdouh (2021)	40	A dult	Not	~	,
indouii (2021)	+0	nun	specified	2	-
irte (2016)	Q	A dult	Burn	~	,
(2010)	20	Paediatric & adult	Burn	4	>
delec(2021)	16	Adult	Burn	4	2
delec (2014)	32	Adult	Burn	-	, 2
delec (2008)	52 70	Adult	Burn	-	, ,
$\frac{1}{2} \left(2019 \right)$	70 51	A dult	Burn	2	-)
(2020)	31 27	Auun Dadiatric & adult	Surgical	4	ے ک
(1008)	145	Paodiatria & adult	Surgical	2	ב ז
(1990)	145	A dult	Mixed	4	ב ר
$\frac{111012}{2020}$	23	Adult	Not	4	<u>بَ</u>
111012 (2010)	0	Adult	not	:	,
(2019)	15	A duilt	Mixed	~	,
$\frac{1}{2018}$	13	Adult Deadiatria	Durn	4	<u>_</u>
1011S(2017)	49	A dult	Dulli) >
(2014)	8/	Adult	Surgical) 7
Din (2010)	30 62	Adult	Mixed) >
-Din (2019)	62	Adult	NOT	-	>
dan Vanalıkasıa	(0	A	specified	~	`
den Kercknove	60	Adult	Burn	2	<u> </u>
US)	ſ	A 1 1/	D	~	`
den Kercknove	6	Adult	Burn	-	5
(3)		A 1 1/	0 · 1	~	
der Veer (2010)	44	Adult	Surgical	2	2
ng (2009)	22	Adult	Burn	4	2
ng (2010)	21	Paediatric	Burn	-	5
seman (2020, 21)	153	Paediatric	Burn	2	2
od (1996)	1	Paediatric	Burn		3
an (2021)	72	Not reported	Not	2	2
			specified		
ol Lee (2022)	16	Adult	Mixed		3
n (2010)	31	Paediatric & adult	Burn	2	2
lkowski (2016)	47	Paediatric	Burn	2	2

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3	Abstracts				
4	Agabalyan (2016)	10	Not reported	Burn	2
5	Anthonissen (2015)	N.R.	Not reported	Burn	3
7	Bajouri (2018)	20	Not reported	Burn	2
, 8	$\frac{1}{2010}$	438	Not reported	Not	3
9	Dezugiy (2017)	150	rotreponed	specified	5
10	$\mathbf{B}_{\mathbf{a}}$	103	Not reported	Mixed	3
11	Dezugiy (2014)	103	Dadiatria & adult	Mixed	2 2
12	(2011, 2012)	10	Paediautic & adult	wiixed	Z
13	(2011, 2012)	10		D	•
14	Blome-Eberwein	19	Adult	Burn	3
15	(2012)				
16	Blome-Eberwein	66	Not reported	Burn	2
17	(2014)				
18	Cho (2012)	30	Not reported	Burn	4
19	Cho (2012)	60	Paediatric & adult	Burn	2
20	Comstock (2018)	1	Adult	Burn	2
21	Cooper (2021)	25	Not reported	Burn	2
22	Du (2006)	1	A dult	Burn	2
23	Edger Lacourgière	1	Not reported	Durn	2
24		44	Not reported	Bulli	3
26	(2022)				•
27	El-Zawhary (2007)	57	Not reported	Mixed	2
28	George (2019)	11	Not reported	Burn	3
29	Jacobs (2016)	6	Paediatric & adult	Burn	2
30	Jang (2009)	20	Not reported	Not	2
31				specified	
32	Kim (2009)	5	Paediatric & adult	Burn	2
33	Li (2016)	34	Not reported	Burn	3
34	Li-Tsang (2011)	4	Not reported	Not	2
35	21 10ung (2011)			specified	-
36	Li-Tsang (2010)	45	Not reported	Not	2
3/	LI-13alig (2010)	TJ	Not reported	specified	2
38	Maari (2017)	10	Not reported	Not	C
39 40	Maari (2017)	12	Not reported		2
40		10		specified	2
42	Moortgat (2020)	10	Not reported	Burn	2
43	Nedelec (2018)	60	Not reported	Burn	2
44	Peters (2018)	5	Not reported	Burn	2
45	Seo (2011)	48	Not reported	Burn	3
46	Siwy (2016)	15	Not reported	Burn	2
47	Timina (2013)	49	Not reported	Not	3
48			1	specified	
49	Tu (2014)	59	Not reported	Not	2
50	14 (2011)	0,7	riorreponeu	specified	-
51	Ud Din (2017)	20	Not reported	Surgical	2
52	Ud Din (2017)	20	Not reported	Surgical	2
53	Ud-Din(2017)	20	Not reported	Surgical	2
54	Ud-Din(2018)	62	Not reported	Surgical	3
55 56	Zuccaro (2021)	20	Paediatric	Burn	3
57	Zuccaro (2019)	13	Paediatric	Burn	3
58	Zuccaro (2021)	20	Paediatric	Burn	3
59	Legend: Paediatric: mea	surement of	patients under the ag	e of 18; Adult: measurement of	of
60	patients aged 18 years of	older; N.R.:	Not reported; Burn:	scars caused by thermal,	

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chemical or friction injury; Surgical: scars caused by surgical procedures (including biopsies); Mixed: participant scars caused by mixed trauma (e.g., burn and acne) **Footnotes:** *Stage in the research to clinical practice translational pipeline, based on the Australian Government Department of Health and Aged Care⁵¹

234 Methods used to measure traumatic cutaneous scar thickness:

- B-mode, including high-frequency (i.e., ≥ 20 MHz) B-mode ultrasound was the most
- commonly reported ultrasound type (n = 72; 61%) (Table 3). Specialised B-mode ultrasound
- 237 devices, including the Tissue Ultrasound Palpation System (TUPS; a B-mode ultrasound
 - transducer in-series with a load cell to allow measured compression of the skin), ^{72 117 141 144}
- and colour Doppler ultrasound, ^{61 156} were used in six records (Table 3).

First Author (year)	Ultrasound Type	Ultrasound Frequency (MHz)	Measurement Parameters	Scar Characteristic Measured	Scar Relocation
Journal articles					
Agabalyan (2017)	High-frequency	20	Epidermal, dermal & combined	N.R.	Not relevant – single measurement
Alsharnoubi (2018)	Midrange ultrasound	N.R.	N.R.	Fibrosis	N.R.
Alsharnoubi (2018)	Midrange ultrasound	N.R.	N.R.	Fibrosis [†]	N.R.
Alshehari (2015)	N.R.	N.R.	Maximum elevation above normal skin	N.R.	N.R.
Avetikov (2018)	B-mode	N.R.	Combined epidermal & dermal	N.R.	Not relevant – single measurement
Berry (1985)	N.R.	N.R.	N.R.	N.R.	N.R. [‡]
Blome- Eberwein (2012)	B-mode	N.R.	Combined epidermal & dermal [§]	N.R.	N.R. [‡]
Blome- Eberwein (2016)	High-frequency	50	N.R.	Fibrosis [†]	N.R. [‡]
Blome- Eberwein (2019)	High-frequency	35	Dermal	Fibrosis, hair follicle density	N.R.
Cai (2019)	High-frequency	50	Dermal	N.R.	N.R.‡
Candy (2010)	B-mode	N.R.	N.R.	N.R.	Scar boundaries traced
Chae (2016)	N.R.	N.R	Combined epidermal & dermal	N.R.	Not relevant – single measurement
Chang (2014)	N.R.	12	N.R.	N.R.	N.R.

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Page **16** of **45**

Chan (2004)	N.R.	N.R.	N.R.	N.R.	Tracing
Cheng (2001)	B-mode	5-10	Combined epidermal & dermal	N.R.	Tracing & cutting out paper Photographs
Cho (2014)	High-frequency	7.5	N.R.	N.R.	N.R.
Danin (2012)	B-mode	20	Epidermal & dermal	N.R.	N.R.
Deng (2019)	N.R.	N.R.	N.R.	N.R.	N.R.
Deng (2021)	Colour Doppler	4-15	Dermal	Fibrosis [†]	N.R.
Deng (2021)	B-mode	8-12	Epidermal & dermal	Fibrosis [†]	Photographs
Dunkin (2007)	High-frequency	N.R.	N.R.	Fibrosis & oedema [†]	Measurements taken at set linear distances along scar
Elrefaie (2020)	High-frequency	13	N.R.	Fibrosis & oedema [†]	N.R [‡]
Engrav (2010)	N.R.	N.R.	N.R.	N.R.	N.R.
Fabbrocini (2016)	N.R.	N.R.	N.R.	Fibrosis & oedema [†]	N.R [‡]
Fong (1997)	B-mode	7.5	N.R.	Fibrosis [†]	Tracing
Fraccalvieri (2013)	High-frequency	7-10 & 10-13	N.R.	Fibrosis & oedema [†]	N.R.
Fraccalvieri (2011)	High-frequency	10-13	Combined epidermal & dermal	Fibrosis [†]	N.R.
Gankande (2014)	High-frequency	20	Combined epidermal & dermal	N.R.	Scar marked & photographed
Ge (2022)	N.R.	N.R.	N.R.	N.R.	N.R.
Gee Kee (2016)	B-mode	8-18	Combined epidermal & dermal	N.R.	Transducer in centre of original burn site where no scar present
Guo (2020)	N.R.	2-15 & 4-15	Combined epidermal & dermal ^c	Fibrosis [†]	Thickest site on peripheral regions
Huang (2017)	N.R.	N.R.	Combined epidermal & dermal	N.R.	Marked & linear measurements from bony landmarks

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3 4	Huang (2021)	B-mode	5-12	N.R.	Oedema [†]	Not relevant – single measurement
5	Huang (2020)	B-mode	5-12	Combined epidermal & dermal	N R	N R
6 7 8	Issler-Fisher (2021)	N.R.	N.R.	N.R.	N.R.	Photograph & measurement of thickest area
9 10	Issler-Fisher (2020)	N.R.	N.R.	N.R.	N.R.	N.R.
11 12 13	Issler-Fisher (2017)	N.R.	N.R.	N.R.	Fibrosis [†]	Scar mapped with drawing Thickest area measured
13	Joo (2020)	N.R.	N.R.	N.R.	Fibrosis [†]	N.R.
15	Katz (1985)	B-mode	10	Combined epidermal & dermal	N.R.	N.R.
16 17	Kemp Bohan (2021)	High-frequency	12	N.R.	Fibrosis [†]	Tracing – thickest area & adjacent landmarks marked
18 19	Kim (2018)	N.R.	22	Combined epidermal & dermal	N.R.	Not relevant – single measurement
20 21 22	Lacarrubba (2008)	B-mode	20	Combined epidermal & dermal	N.R.	N.R.
23 24 25	Lau (2005)	Tissue Ultrasound Palpation System	5 (burn) & 10 (surgical)	N.R.	N.R.	Tracing – most severe/prominent site
26 27	Lee (2020)	High-frequency	20	Combined epidermal & dermal	Fibrosis [†]	Not relevant – single measurement
28	Lee (2019)	High-frequency	20	Combined epidermal & dermal	Fibrosis [†]	Marked with pen
29 30	Li (2013)	High-frequency	12	Combined epidermal & dermal	Fibrosis [†]	Tracing
31	Li (2020)	N.R.	10	N.R.	Fibrosis [†]	N.R.
32	Li (2021)	High-frequency	20	N.R.	N.R.	Thickest area
33	Li (2021)	High-frequency	20	N.R. [§]	Fibrosis [†]	Thickest area
34	Li (2018)	N.R.	N.R.	Combined epidermal & dermal	N.R.	N.R.
35 36 37	Li-Tsang (2005)	Tissue Ultrasound Palpation System	N.R.	N.R.	N.R.	N.R.
38 39 40	Li-Tsang (2006)	B-mode	N.R.	N.R.	N.R.	N.R [‡]

Li-Tsang (2010)	B-mode	N.R.	N.R.	Fibrosis [†]	N.R.
Lobos (2017)	B-mode & colour Doppler	18	N.R.	Fibrosis [†]	Not relevant – single measurement
Mamdouh (2021)	High-frequency	N.R.	Combined epidermal & dermal§	Fibrosis [†]	N.R.
Meirte (2016)	High-frequency	22	Dermal	Fibrosis & oedema [†]	Marked with surgical pen, including boundaries of probe. Photograph of body position & probe location
Miletta (2021)	N.R.	50	N.R.	Fibrosis [†]	Tracing – worst scar
Nedelec (2014)	High-frequency	20	Combined epidermal & dermal	N.R.	Tracing including notable landmarks. Measurement site circled. Photograph
Nedelec (2008)	High-frequency	20	Combined epidermal & dermal	N.R.	Tracing including notable landmarks. Measurement site circled. Photograph
Nedelec (2019)	High-frequency	20	Combined epidermal & dermal	Fibrosis & oedema [†]	Tracing. Hole cut over measurement area
Nedelec (2020)	High-frequency	20	Combined epidermal & dermal	N.R.	Photograph
Nicoletti (2015)	N.R.	22	Epidermis to fascia	N.R.	N.R.
Niessen (1998)	B-mode	N.R.	N.R.	Fibrosis & oedema [†]	3cm border marked with tape – measurements lateral
Reinholz (2020)	B-mode	11	Combined epidermal & dermal	Fibrosis & oedema [†]	N.R.
Reinholz (2016)	B-mode	11	Combined epidermal & dermal§	Fibrosis & oedema [†]	N.R.
Schwaiger (2018)	B-mode	11	N.R.	Fibrosis & oedema†	N.R.
Simons (2017)	B-mode	8-18	Combined epidermal & dermal	N.R.	Tracing – scar & anatomical landmarks

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3	Sovkan (2014)	N.R.	3-9	N.R.	Fibrosis [†]	N.R.
4 5	Timar-Banu (2001)	High-frequency	20	Combined epidermal & dermal	Fibrosis [†]	N.R.
7	Ud-Din (2019)	High-frequency	50	Combined epidermal & dermal	Fibrosis	Defined anatomical location
8 9 10	van den Kerckhove (2003)	High-frequency	20	Combined epidermal & dermal	N.R.	Test sites marked. Thermoplastic splints created with space for transducer
11 12 13 14	van den Kerckhove (2005)	High-frequency	20	Combined epidermal & dermal	N.R.	Test site boundaries marked & traced
15 16	van der Veer (2010)	N.R.	7.5	N.R.	Fibrosis [†]	Standardised linear measurement points
17	Wang (2009)	High-frequency	N.R.	N.R.	Fibrosis [†]	N.R.
18 19	Wang (2010)	B-mode	N.R.	Combined epidermal & dermal	N.R.	Tracing – scar & anatomical landmarks
20 21 22	Wiseman (2020, 2021)	B-mode	N.R.	Combined epidermal & dermal	Fibrosis [†]	Centrally site of interest
23 24	Wood (1996)	B-mode	7 & 10	N.R.	N.R.	Transducer affixed to tracking arm
25	Xuan (2021)	High-frequency	20	N.R.	 Fibrosis[†] 	N.R.
26 27 28	Yeol Lee (2022)	B-mode	7-16	N.R.	N.R.	N.R.
20	Yim (2010)	High-frequency	12	N.R.	N.R.	N.R.
30 31	Zadkowski (2016)	B-mode	N.R.	Combined epidermal & dermal	N.R.	N.R.
32 33	Abstracts					
34 35 36	Agabalyan (2016)	N.R.	20	Epidermal, dermal & combined	N.R.	N.R.
37 38	Anthonissen (2015)	N.R.	22	Epidermal & dermal	N.R.	N.R.
39 40	Bajouri (2018)	N.R.	N.R.	Epidermal & dermal	N.R.	N.R.
41 42						Page 20 of 45

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Bezugly (2019) Bezugly (2014) Blome- Eberwein (2011, 2012)	High-frequency High-frequency N.R.	22, 33 & 75 33 & 75 N.R.	Epidermal & dermal Epidermal & dermal N.R.	N.R. N.R. N.R.	N.R. N.R. N.R.
Eberwein (2012)	High-frequency	N.R.	N.R.	Fibrosis	N.R.
Blome- Eberwein (2014)	High-frequency	N.R.	N.R.	N.R.	N.R.
Cho (2012)	N.R.	N.R.	N.R.	N.R.	N.R.
Cho (2012)	N.R.	N.R.	N.R.	N.R.	N.R.
Comstock	N.R.	N.R.	N.R.	N.R.	N.R.
(2018)					
Cooper (2021)	N.R.	N.R.	N.R.	N.R.	N.R.
Du (2006)	B-mode	15	N.R.	N.R.	N.R.
Edgar-	N.R.	N.R.	N.R.	N.R.	N.R.
Lacoursière (2022)					
El-Zawhary (2007)	N.R.	N.R.	N.R.	N.R.	N.R.
George (2019)	N.R.	N.R.	N.R.	N.R.	N.R.
Jacobs (2016)	N.R.	N.R.	N.R.	N.R.	N.R.
Jang (2009)	N.R.	N.R.	N.R.	N.R.	N.R.
Kim (2009)	N.R.	N.R.	N.R.	N.R.	N.R.
Li (2016)	N.R.	N.R.	N.R.	N.R.	N.R.
Li-Tsang	Tissue Ultrasound	N.R.	N.R.	N.R.	N.R.
(2011)	Palpation System				
Li-Tsang	Tissue Ultrasound	N.R.	N.R.	N.R.	N.R.
(2010)	Palpation System				
Maari (2017)	N.R.	N.R.	N.R.	N.R.	N.R.

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Moortgat	High-frequency	N.R.	Dermal	N.R.	N.R.
(2020)					
Nedelec (2018)	N.R.	N.R.	N.R.	N.R.	N.R.
Peters (2018)	High-frequency	22	N.R.	N.R.	N.R.
Seo (2011)	N.R.	7.5	N.R.	N.R.	Thickest point
Siwy (2016)	N.R.	N.R.	N.R.	N.R.	N.R.
Timina (2013)	N.R.	20-40	N.R.	N.R.	N.R.
Tu (2014)	High-frequency	N.R.	N.R.	N.R.	N.R.
	ultrasound				
	biomicroscopy				
Ud-Din (2017)	N.R.	N.R.	N.R.	N.R.	N.R.
Ud-Din (2017)	High-frequency	50	N.R.	N.R.	N.R.
Ud-Din (2018)	High-frequency	N.R.	N.R.	Fibrosis [†]	N.R.
Zuccaro (2021)	N.R.	N.R.	N.R.	N.R.	N.R.
Zuccaro (2019)	B-mode	N.R.	N.R.	N.R.	N.R.
Zuccaro (2021)	B-mode	6-18	Combined epidermal & dermal	N.R.	Scar outlined &
					photographed

Legend: B-mode: brightness-mode ultrasound (< 20 MHz); High-frequency: high-frequency B-mode ultrasound (> 20 MHz); N.R.: Not reported

Footnotes: [†]Indirect reference made in record (e.g. in introduction or discussion); [‡]Photographs taken of the scar but not specified whether used for relocation; [§]Not stated in methods, so images provided in record used by authors of this review to provide subjective judgement

The type of scar and skin thickness measurement (i.e., thickness of the dermis, epidermis, or combined epidermal and dermal measurement) was reported in 39 records (33%) (Table 3). Where reported, combined measurement of epidermal and dermal thickness was the most common method (n = 28; 24%).^{21 25 31 32 35 37 58 63 65 69 70 75 76 78 82 86 87 91 94 103 108 112 125 126 128 129} ¹⁴⁵ ¹⁴⁹ Separate epidermal and dermal thickness measurements were reported in seven records (6%).^{34 74 89 95 98 132 153 154} Of these records, two authors provided a rationale for this decision: each skin layer provided different information on the scar; ³⁴ or responded differently to treatment. 74 92 Four records (3%) directly reported that fibrosis was the scar characteristic targeted by the measurement. ^{36 42 96 160} One of these records also quantified hair follicle density to assess the difference between scared and unscarred skin.⁴² An additional 26 records (22%) made indirect reference (i.e., within the introduction or discussion) to the measurement of fibrosis. 21 61 63 67-70 73 81-84 89 92 93 97 110 115 119 125 126 142 150 151 155 156 160 Ten records (8%) made indirect reference to the measurement of both oedema and fibrosis, 39 43 62 74 77 79 80 139 146 157 and one record made indirect reference to the measurement of oedema.⁶⁴ Additional objective and subjective measurement methods were employed alongside ultrasound measurement in 115 records (97%) (Supplementary Table 2). All three phase 4 studies involving implementation in routine clinical practice utilised additional measurements. ^{99 104 109} The additional objective measurements included elastography (elasticity), cutometric assessment (pliability) and Doppler ultrasound (vascularity). The additional subjective measurements were conducted using clinician-based rating scales (e.g., Vancouver Scar Scale, used in 46 records^{21 26 39 42 43 58 61 66 67 69 71 72 76 85 88 90-93 96-98 100 101 103 104}

264 ^{108 113-115 117 127-129 134-137 139 141-144 147 150}) or patient-reported outcome measures (PROMs). ^{59 63}

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Page 23 of 45

Page 25 of 77

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BMJ Open

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Methods used to relocate the scar for repeated measurements were reported in 34 records 266 (29%) (Table 3). The most common relocation method was tracing the outline or boundaries 267 of the scar on a transparent or translucent sheet (n = 14; 12%), ^{32 70 77 87 88 102 103 110 119 129 141} 268 occasionally including prominent or bony landmarks close to the scar. ^{25 68 75} Photographs (n 269 =11; 9%) and linear measurements from defined points or anatomical landmarks on or around 270 the scar (n = 4; 3%) were also used for scar relocation. The 'worst' or 'thickest' part of the 271 272 scar, as determined by patients or assessors, was chosen as the measurement site in 15 records (13%).26 32 39 61 62 66 91 92 115 123 125 126 139 143 155 273

Measurement of unscarred skin, either contralateral or adjacent to the scar, was performed in 274 29 records (25%). These measurements were primarily used as controls or comparators to 275 scar measurements (n = 27, 23%).²¹ $\frac{31}{32}$ $\frac{36}{37}$ $\frac{37}{42}$ $\frac{57}{60}$ $\frac{63}{65}$ $\frac{69}{68}$ $\frac{76}{82}$ $\frac{85}{86}$ $\frac{89}{91}$ $\frac{91}{92}$ $\frac{108}{110}$ $\frac{112}{127}$ $\frac{129}{130}$ $\frac{138}{138}$ 276 ¹⁵² 155 Additionally, three records (3%) evaluating treatment efficacy measured both 277 unaffected skin thickness and the thickness of a 'control' or untreated scar. 93 100 131 All 278 instances where additional ultrasound measurements were taken of unscarred skin or 279 untreated scars were reported as part of research initiatives aligning with phases 2 and 3 of 280 the research-to-practice pipeline (figure 1).⁵¹ 281

282 Reliability and measurement error

Reliability was calculated for both scarred and unscarred skin in 14 records (12%) and was generally acceptable (Supplementary Table 2). This included inter-rater reliability (n = 5; 4% of included records), $^{62} ^{69} ^{76} ^{110} ^{137}$ intra-rater reliability (n = 3; 2% of included records), $^{31} ^{32} ^{70}$ and both inter- and intra-rater reliability (n = 7; 6%) $^{21} ^{25} ^{91} ^{94} ^{123} ^{132} ^{141}$. The intraclass correlation coefficient (ICC) was the most commonly reported reliability statistic (n = 10; 8% of included records), $^{21} ^{25} ^{69} ^{70} ^{76} ^{91} ^{91} ^{94} ^{110} ^{132} ^{141}$ where it was reported for both scar and unscarred skin measurements in four records (3%). $^{21} ^{25} ^{76} ^{91}$ The reported combined thickness

(i.e., epidermal and dermal) ICCs for inter-rater reliability of scarred skin ranged from 0.82 to 0.985, while the inter-rater ICC for the measurement of unscarred skin ranged from 0.33 to 0.98, with one of the four records reporting an ICC below the threshold value of 0.7 (ICC = 0.33)²⁵ and one record simply reported that the inter-rater ICC for scarred skin was "acceptable to high".⁶⁹ The reported intra-rater reliability for combined thickness measurements of scarred skin ranged from 0.89 to 0.983, and for unscarred skin ranged from 0.61 to 0.982, with one record reporting an ICC below the threshold of 0.7 (ICC = 0.61).²⁵ One record reported both the inter- and intra-rater ICCs for individual epidermal (inter-rater ICC = 0.297; intra-rater ICC = 0.809) and dermal (inter-rater ICC = 0.991; intra-rater ICC = 0.991) scar thickness measurement.¹³² Four records (3% of included records) reporting reliability used Pearson's R, an undisclosed method, or description (e.g., high) as detailed in supplementary table 2. ^{31 62 123 137}.

Measurement error for inter-rater and intra-rater reliability of combined, epidermal or dermal thickness was reported in five records (4%) using standard error of the measurement (SEM). The inter-rater SEM for the combined epidermal and dermal thickness of scarred skin ranged from 0.11 mm to 0.5 mm, and the intra-rater SEMs ranged from 0.18 to 0.52 mm. Individual records reported SEM values for unscarred skin, and separate epidermal and dermal measurements, available in supplementary table 2. ^{21 25 32 94 132} Only one record reported calculation of the smallest detectable change (SDC). In that record the inter-and intra-rater SDC was calculated for both scarred and unscarred skin. The scarred skin SDCs were 1.4 mm (inter-rater) and 0.6 mm (intra-rater), and unscarred skin SDCs were 0.8 mm (inter-rater) and 0.5 mm (intra-rater).²⁵ The reported SEMs were all close to or below the largest SDC value reported. This finding may indicate that ultrasound can detect true variance in scar thickness above measurement error for traumatic scar and skin thickness.

50 314 Clinical, health service, implementation and feasibility outcomes:

Page 27 of 77

BMJ Open

No record specifically investigated clinical, health service, implementation or feasibility outcomes of ultrasound as a measurement-based-care initiative. Ultrasound was used to assess the clinical outcomes of scar treatment initiatives in all included records. Clinical, health service, implementation and feasibility outcomes related to ultrasound measurement were, however, reported in 67 records that focused on scar treatments. ^{21 25 31 32 34-36 39 42 57 58 60} 62-71 73-78 82 86 90 91 93-95 99 100 102 103 109-112 115 120 123 127 129 132 133 138 139 141 145-147 150 153-160 The clinical outcome of patient satisfaction related to ultrasound measurement was only reported in one record. Whilst patient satisfaction was not directly measured in that record, a proxy measure of satisfaction was reported by the authors stating that no paediatric patient or their caregiver refused ultrasound measurement once the purpose was explained.²⁵ Timeliness was the only reported health service outcome, reported as the time required to take ultrasound measurements. Where reported, this was short, taking between one to five minutes, ^{25 35 112}. The most common implementation outcomes reported in the identified records were fidelity, acceptability and appropriateness. Fidelity to the measurement method was reported through the use of experienced or trained assessors (n = 7; 6%), $25 \times 86 \times 91 \times 132 \times 146 \times 150 \times 155$ and/or utilising the same assessor/s for all measurement sessions (n = 5; 4%).^{25 66 139 146 155} Differences between intended and actual measurement methods were not discussed. The training and/or experience of the assessors was discussed in 24 records (20%),^{21 25 32 35 60 63 64 67-71 74 76 86 91 102 103 110 129 139} ¹⁴¹¹⁴⁶¹⁵⁶ where measurements were either taken by a clinician (n = 13; 11%), ²¹²⁵³²⁶⁴⁶⁹⁻⁷¹⁷⁴⁸⁶ $^{92\ 110\ 141\ 143}$ members of the research team (n = 6; 5%), $^{67\ 68\ 76\ 91\ 102\ 146}$ or by specialist sonographers and/or radiologists (n = 5; 4%).^{63 103 129 139 156} Only one record reported on fidelity in the context of routine clinical practice. In this instance, ultrasound was conducted in the department of radiology, however the role or training of the staff was not reported.¹⁰⁹

The acceptability and appropriateness of the ultrasound methods used in individual records were generally based on author opinion and outlined in the discussion. Acceptability was reported in 25 records (21%), ^{21 25 31 32 34-36 39 69 73 77 78 82 91 94 95 103 109-112 115 141 145 156 157} including for paediatric populations, where one record reported potential difficulty in measuring this population, ³¹ contrasting that which reported that measurement was acceptable to both children and their caregivers.²⁵ One record reported acceptability where the intervention being analysed by ultrasound was already part of routine clinical practice. In this instance, the authors referenced additional publications which stated that ultrasound had an accuracy of 0.5 mm, which was judged by the authors to be sufficient for assessment of scar thickness. ^{109 25} ^{35 112} Potential difficulty was identified in the measurement of open wounds, ²⁵ and traditionally hard-to-reach areas (such as the axillae or groin).³¹ The appropriateness of the ultrasound methods was reported in 46 records (39%), and was generally addressed in the discussion. 25 31 34 35 39 42 57 58 62 65 66 69-71 75-78 82 90 91 93-95 99 100 103 109 110 112 115 120 123 127 132 138 141 147 153-160 Of these records, two (2%) determined that ultrasound was not appropriate for scar measurement. The first stated that it was too inaccurate and complex; ⁹⁵ and the second, which reported on initiatives within routine clinical practice, determined that the minimum resolution of the Diasonography ultrasonic scanner (Nuclear Enterprises, Edinburgh, UK) precluded its use in scars thinner than 3mm.⁹⁹ The feasibility of ultrasound was reported in 12 records (10%).^{25 31 34 73 82 91 93 109 110 133 141} Five records considered ultrasound not feasible for scar measurements. The rationale presented included high-frequency 20 MHz ultrasound having an inadequate penetration depth; ^{34 91} and ultrasound measurement and training of investigators requiring too much time (as reported in

361 one record in phase 4 of the research-to-practice pipeline). ^{31 109 110} Another factor identified

362 as precluding feasibility was the inability to consistently relocate the measurement site.²⁵

363 Conversely, one record reported ultrasound to be feasible in combination with Vancouver

Page 29 of 77

BMJ Open

Scar Scale (VSS) measurement, ⁷³ and another stated that ultrasound was able to distinguish between subcutaneous fat and muscle, which was interpreted by the authors of that record to mean that skin thickness measurements were accurate. ¹³³ The majority (n = 11; 92%) of the records reporting feasibility were research initiatives in phase 2 or 3 of the research to practice pipeline. One record examined feasibility in the context of routine clinical practice (i.e., phase 4; figure 1), ¹⁰⁹ where it was determined that ultrasound was not suitable for use in their twelve-year longitudinal study due to changes in staff, equipment and software over such a long time period, which introduced additional variables to the measurement process that were impossible to control. 109 Factors influencing ultrasound images and measurement methods: The only factor that was reported to influence the imaging and measurement methods was the measurement of scars with open wounds. This was reported in one record, which determined that ultrasound and ultrasound gel was unsuitable in this instance. The authors of that record suggested the use of a flexible transparent plastic wrap, which is placed over the measurement area prior to measurement with ultrasound.²⁵ **Reported strengths and limitations of the measurement methods:** The safety, practicality, objectivity, versatility, reliability and non-invasive nature of ultrasound were all reported as strengths of the measurement method. ^{31 35-37 42 58 66 68 69 80 82 91} 94 109 111 123 132 133 138 141 147 149 153 155 157 159 When compared to other subjective or clinical measurement methods (e.g., VSS) and 3D camera, ultrasound was viewed as the superior measurement method of scar and skin thickness, due to its improved accuracy, greater sensitivity to change and objectivity. ^{25 69 76 103 110} The ability of ultrasound to differentiate between scarred and unscarred skin was also highlighted (n = 4; 3%).^{42 65 75 112} as was the

versatility of ultrasound in its ability to measure a variety of anatomical areas and be used with child participants (i.e., <18 years) (n = 2; 2%).^{31 156}

The poor correlation between ultrasound and histological thickness measurements, ⁹⁵ and the established inverse relationship between ultrasound penetration depth and the resolution of superficial structures were identified as limitations of ultrasound in the measurement of scar thickness. ^{34 35 82 99 153 156 157}. One record, reporting on a longitudinal study that was conducted over twelve years, reported that the continuous development of ultrasound software and hardware over that time limited the usefulness of ultrasound. ¹⁰⁹ Despite being reported elsewhere as acceptable (i.e., between one to five minutes^{25 35 112}), one record reported that the time-consuming nature of measurement and the requirement for assessors to be trained in the operation of, and techniques required for, ultrasonography was a limitation of the method. ¹¹⁰ Methodologically, concerns were raised around the pressure caused by application of the ultrasound transducer to the skin, and how that may influence thickness measurement. ^{26 66 68} ¹⁴¹ The size of the transducer head relative to the size of scars was also considered a potential limitation, as multiple measurements are required for quantification of larger scars.⁹¹ Finally, it was recognised that there may be a difference between changes to the scar that can be measured by ultrasound, and what is felt and/or experienced by the patient. ^{78 82 125 126} It was suggested that changes that are detectable by ultrasound may be smaller than those able to be detected by patients. A minimum change in scar thickness as measured by ultrasound of between 1 to 6 mm is required before a patient may report noticing any difference to their scar thickness, ^{25 78} indicating that a holistic approach to scar thickness using the patient's opinion as well as objective measurement through ultrasound may be beneficial.

409 Guidelines or frameworks used to guide the measurement methods:

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No records reported using any guidelines or frameworks to inform their measurement
methods. One record utilised suggestions from The American Wound Healing Society to
support the measurement of contralateral, unscarred skin thickness on the same individual as
a control or comparator.⁷⁸

414 Methodological Considerations:

Based on the ultrasound methods and outcomes identified in this review, a list of
methodological considerations have been compiled (Supplementary Table 4). These are
intended to guide the decision-making and methodological reporting of researchers and/or
clinicians undertaking scar or skin thickness ultrasound measurement.

DISCUSSION:

This review mapped the methods used in the published literature to measure traumatic scar thickness using ultrasound across the research-to-practice translational pipeline. No record reported their methods with sufficient detail to allow them to be independently replicated. Overall, there was a lack of consistent rationale underpinning which skin layers (i.e., epidermis, dermis and combined) were measured, and little consideration was given to the training and experience required by assessors. The included records mainly aligned with the second and third phases of the research-to-practice pipeline (figure 1), with only three reporting the use of ultrasound in routine clinical practice. ^{99 104 109} This suggests a research translational gap, where ultrasound is either most commonly used as an outcome measure for research initiatives and is not regularly used to evaluate care once treatments are implemented into routine clinical practice, or that use in routine clinical practice is not reported or evaluated.

432 While efforts have been made to standardise ultrasound measurement procedures elsewhere
 433 in dermatology (including tumours, cancers, vascular anomalies, and systemic sclerosis^{46 47}),

this same effort has not yet extended to the measurement of traumatic scarring. Methodological standardisation has the potential to increase confidence in the use of ultrasound as the basis of measurement-based care initiatives for clinical decision-making, allowing patient care and scar treatments to be tailored towards individual needs. ^{26 161 162} Standardising the core methodological components of ultrasound measurement of scar thickness, or at the very least, creating a standardised framework for methodological decision-making, may support implementation of ultrasound measurement into routine clinical practice, supported by strategies to overcome barriers to implementation at local sites.

This review identified novel insights into the identification of the composition of cutaneous scars using ultrasound, and highlighted the apparent lack of consistent understanding of, or rationale behind, what scar thickness characteristics were being measured. Fibrosis is generally understood to be the primary cause of scar thickness through the deposition of excessive extracellular matrix proteins such as collagen.^{164 165} This has been confirmed through histological analysis, which has shown the presence of excess collagen and other extracellular matrix proteins in the dermis of hypertrophic scars.^{40 41} An additional method for assessing the effects of scarring on the dermis, as identified by one record in this review, ⁴² is through quantification of the presence and density of hair follicles. This quantification may serve as a method of differentiation between scarred and physiological skin, and may also serve as a measure of skin function.⁴² What is less understood, and perhaps largely overlooked, is the function of the epidermis in scar thickness. In the one record identified in this review that directly report the measurement of the epidermis, the authors noted that the measurement quantified the presence of oedema.⁴³ This was further supported by two records that noted that the epidermis and dermis responded differently to treatment, ^{74 92} indicating that there is likely a difference in the composition of the scar between these skin layers.

Page 33 of 77

BMJ Open

Cutaneous oedema has been observed using high-frequency ultrasound in other pathologies, including atopic dermatitis and skin ageing, where it is characterised by the presence of a sub-epidermal low echogenic band (SLEB), which is a hyperechoic band at the dermoepidermal junction.¹⁶⁶ Understanding the interplay between epidermal oedema, dermal fibrosis and the presence and density of hair follicles may result in an increased understanding of the mechanisms and treatment responses of cutaneous scarring. With better understanding, more targeted scar treatments that inform a greater understanding of scar responsivity may arise.

Another important, but potential limiting factor for the use of ultrasound to measure scar thickness raised in this review is the training and/or experience required of assessors, and the ramifications this likely has on the reliability of measurements and interpretation.¹⁶⁷ This review identified 24 records where assessor experience was discussed, however none made any recommendations on the optimal training and/or experience. Identifying the training requirements of assessors may prove an important step towards more widespread implementation of reliable ultrasound scar thickness measurement in research trials and as the basis for measurement-based care in routine clinical practice.²⁸ A panel of dermatological and ultrasound experts has previously recommended that a physician with a minimum of 300 examinations per year should hold responsibility for ultrasound measurements.⁴⁶ It has also been suggested that training existing members of clinical teams and standardising measurement method/s may be the most effective way to achieve minimum reliability standards under clinical conditions. This could allow measurement to be reliably conducted within an outpatient clinic setting by a number of healthcare providers assisting workflow, negating the requirement for patients to wait for an experienced radiographer. ^{25 28} In the current review reliability estimates were generally acceptable but were tested under research conditions.

484 Study Limitations:

Only articles available in English or with an English abstract were considered for inclusion and data extraction, which may have resulted in the omission of eligible information. Data extraction was completed on the translated English abstracts of two non-English articles, however the non-English articles themselves were not available to the authors, and thus could not be analysed. Based on the number of records included in this review, however, it is unlikely that this would have impacted review findings. An additional limitation was that authors of included records were not contacted to provide clarification or further information, as this was not feasible given the number of results identified. It should also be acknowledged that the included records were not designed to align with the specific aims of this review, which likely explains some of the lack of reporting on outcomes of interest in our review, particularly clinical, health service and implementation outcomes. Furthermore, as this review relied on published information (including grey literature), routine practices employed within organisations may not have been considered and unpublished industry sponsored reports may not have been identified. It is also important to consider the limitations of ultrasound itself for the holistic quantification of cutaneous scarring. Ultrasound transducers are generally small, meaning that it is difficult to assess the entirety of a scar, necessitating multiple measurements.¹⁶⁸

502 Additionally, thickness is often not the only scar parameter of clinical or research interest. It

503 has therefore been recommended that multi-modal measurement techniques are employed,

which include both subjective and objective measurements. ^{169 170} However, use of these

505 methods may be challenging in routine clinical practice, due to the length of time and training

506 required. Thus, feasibility and implementation outcomes are of importance in evaluating

507 measurement-based care initiatives involving ultrasound alone or multimodal measurement

59 tools in scar care practice -a field in its infancy based on this review.
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509 Future Directions:

It is intended that the results of this review will be used to inform the creation of a Delphi consensus study, leading to the formation of a guideline for the measurement of traumatic scar thickness using ultrasound. This guideline can then be used by researchers and clinicians to standardise the measurement of scars. In preparation for this study, we have provided a list of methodological considerations for assessors or practitioners when planning to conduct scar thickness measurements with ultrasound (Supplementary Table 4). Future research could also investigate aspects that were beyond the scope of this review including factors influencing the implementation of ultrasound-based care initiatives, strategies to support implementation, and how research-based initiatives could be applied in practice. Further studies are needed that compare SDCs to SEMs to interpret reliability estimates to confirm our interpretation that ultrasound may have the ability to detect true change or variance in scar thickness above measurement error, which was based on the SDC reported by a single study. Our interpretation is supported by mostly acceptable reliability estimates of ultrasound thickness for other cutaneous conditions. ^{29 171}

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The authors declare no competing interests. The research presented in this publication was
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0	907	therapy for hurn scar prevention and management in children: a randomized controlled trial				
10	908	Clin Rehabil 2020:31(1):120-31 doi: 10.1177/0269215519877516				
11	900	127 Wood FM Currie K Backman B et al Current difficulties and the possible future directions in				
12	010	scar accossment Pures 1006:22/6):455 59, doi: 10.1016/0205.4170/05/00169.0				
13	910 011	129 Żadkowski T. Nachulowicz D. Mazgai M. et al. A pow CO2 laser technique for the treatment of				
14	911 012	120. Ząukowski T, Nachulewicz P, Mazgaj M, et al. A new CO2 laser technique for the treatment of				
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21 22	918	Burn Care Res 2019;40(Supplement_1):S215-S15. doi: 10.1093/jbcr/irz013.3/4				
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57	950	dve laser in the treatment of hypertronhic hurn scars in Chinese children with Fitzpatrick				
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25	1020	10 1007/DBS 0b0120218106046b
26	1021	160 Dowors DS Sarkar S Coldgef DB et al Scar accessment: current problems and future solutions
27	1022	The Journal of hum care & rehabilitation 1000/20(1 Dt 1)/54 60, doi: 10.1007/00004620
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Figure 2. Preferred Reporting Items for Systematic reviews and Meta-Analyses (PRISMA) flow diagram for this study.

159x102mm (220 x 220 DPI)

BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

Supplementary Box 1. Full search strategy for Ovid MEDLINE.

((ultrasound.ti,ab. OR ultra sound.ti,ab. OR sonograph*.ti,ab. OR ultrasonic.ti,ab. OR high-frequency.ti,ab. OR high frequency.ti,ab. OR hfus.ti,ab. OR ultrasonog*.ti,ab. OR exp Ultrasonography/)

AND

((skin.ti,ab. OR epiderm*.ti,ab. OR derm*.ti,ab. OR cutaneous.ti,ab OR scar*.ti,ab OR keloid*.ti,ab OR cicatri*.ti,ab OR exp Skin/ OR exp Dermatology/ OR exp Cicatrix/)

AND

(thickness*.ti,ab. OR thicken*.ti,ab. OR depth.ti,ab. OR volume.ti,ab. OR height.ti,ab. OR vancouver scar scale.ti,ab)

ADJ10

(measure*.ti,ab. OR quantif*.ti,ab. OR calculat*.ti,ab OR estimat*.ti,ab OR assess*.ti,ab. OR determin*.ti,ab. OR evaluat*.ti,ab OR imag*.ti,ab OR exam*.ti,ab)))

NOT (exp animals/ NOT exp humans/)

Legend: ab, abstract (searches the abstract of the publication); adj10, adjacency (search terms must be located within 10 words of one another); exp, explode (used to include all subheadings when searching MeSH headings); ti, title (searches the title of the publication)



BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

Supplementary	Table 1	Extraction	categories	and fields
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Extraction category	Extraction field
Publication details	First author
	Year of publication
	Title of publication
	Country (first author)
	Country (study)
	Country (recruited)
	Publication type (e.g., peer-reviewed journal article, abstract)
	Journal name
	Corresponding author contact details
	Use of scar thickness measurement (e.g., longitudinal study, response to
	treatment)
Study details	Aim/objective
	Research questions
	Target population/topics
	 Study design (e.g., RCT, mixed methods)
	Data and analysis (i.e., statistical methods)
	Removal of scar treatments before ultrasound measurement (e.g., length of
	time before measurement)
	Reason for measurement (e.g., research, clinical initiative)
	Inclusion/exclusion criteria
	Dates of data collection
	Ultrasound thickness collection methods (e.g., direct collection, collected
	from medical records)
	Contralateral/unaffected/comparator skin thickness measurement
	Other methods used
	Use of guidelines/frameworks for measurement methods
	How previously published methods/guidelines were used
	Research pipeline stage
	Setting (e.g., inpatient/outpatient clinics)
	Scar type (e.g., burn scar, surgical scar)
Participant details	Number of participants
	Population type (e.g., adult/paediatric)
	Gender ratio
	Fatient involvement in unckness determination
I litrogound mothoda	Iltracound mode
Offiasound methods	Device name and monufacturer
	Frequency used
	Number of measurements taken
	What did researchers report they were measuring (e.g. fibrosis ordema)
	Anatomical locations/functional measurement units measured
	Patient orientation
	Illtrasound transducer orientation
	Methods used to prevent skin compression
	Measurement site relocation strategies
	Type of skin measurement (i.e. enidermis/dermis/combined)
	Measurer training
Psychometric properties*	Reliability
r sychometric properties	Measurement error
Feasibility [†] outcomes	Time taken for measurement
-	Availability of measurement method

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3		Ease of administration
4		Number of steps required
5		Number of people required to conduct measurements
6		Considerations for special populations
7	Implementation [†] outcomes	A acontability
8	implementation* outcomes	Acceptaolity
9		Adoption
10		Appropriateness
11		Cost
12		Feasibility
12		Fidelity
17		Sustainability
14	Strengths and limitations of	Subminuome
15	manufacture mathematical manufacture manufacture mathematical mathematicas mathematicas mathemat	Limitations
10	measurement methods	Deminations
17		Barriers
18		Enablers
19	Findings	Ultrasound-related findings
20	*Psychometric properties as	outlined in the COSMIN Risk of Bias tool to assess the quality of studies on
21	reliability or measurement e	error of outcome measurement instruments ¹
22	[†] Feasibility outcomes as per	Prinsen <i>et al.</i> ²
23	[‡] Implementation outcomes	as per Proctor <i>et al</i> ³
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First author (year)	Objective measurement methods	Clinician-based rating scale	PROM
Journal articles			
Agabalyan (2017)	Histology	-	-
Alsharnoubi (2018)	Laser Doppler perfusion	VSS	-
Alsharnoubi (2018)	Laser Doppler perfusion	VSS	-
Alshehari (2015)	-	VSS	-
Avetikov (2018)		-	-
Berry (1985)	Transcutaneous oxygen measurement	Scar redness and hypertrophy	Scar redness and hypertrophy ratin
• • •		rating scale (0-5 Likert scale)	scale (0-5 Likert scale)
Blome-Eberwein (2012)	Doppler flowmeter – vascularity	VSS	POSAS-P
· · · · · · · · · · · · · · · · · · ·	Cutometer – pliability	POSAS-O	
	Semmes-Weinstein monofilament		
	Aesthesiometer testing set –		
	sensation		
Blome-Eberwein (2016)	Cutometer – pliability	VSS	POSAS-P
	Dermaspectrometer – colour	POSAS-O	
	Semmes-Weinstein Aesthesiometer		
	Monofilament Testing Set –		
	sensation		
Blome-Eberwein (2019)	-	VSS	-
Cai (2019)	_	Clinical evaluation	~ -
Candy (2010)	Spectrocolorimeter – colour	VSS	-
Chae (2016)	Spectrophotometer – pigmentation	VSS	POSAS-P
0	speer op not on the promotion	POSAS-O	
Chang (2014)	-	VSS	_
()		Photographic evaluation (0-10	
		VAS)	
Chan (2004)	Cutometer – viscoelasticity		-
	Spectrophotometer – pigmentation		
Cheng (2001)	-	VSS	-
Cho (2014)	Mexameter – colour	Treatment efficacy (0-10 VAS)	Itching scale (0-4 Likert scale)
		110minut 01110my (0 10 7110)	Trening Source (Series Source)

Supplementary Table 2. Additional measurement methods used alongside ultrasound in included studies

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	Tewameter - trans-epidermal water		
	loss		
	Sebumeter – sebum		
	Cutometer – elasticity		
Danin (2012)	Cutometer – elasticity	VSS	-
Deng (2019)	DermaLab Combo – colour	POSAS-O	-
	Dermoscopy – vascularity		
Deng (2021)	-	VSS	-
Deng (2021)	Doppler – blood perfusion	POSAS-O	POSAS
5 ()	Dermlite Foto IIPro – erythema		
Dunkin (2007)	- Uk	-	-
Elrefaie (2020)	Ultrasound – echogenicity.	VSS	-
	compressibility & vascularity		
Engrav (2010)	Durometer – hardness	Clinical appearance based on	-
Engla((2010)	Chromameter – colour	nhotographs	
Fabbrocini (2016)	-	mVSS (vascularity nigmentation	_
1 ubbioenni (2010)		nliability)	
Fong (1997)	Cutometer – elasticity	Clinical rating – colour change	_
1011g (1997)	Cutometer clusterty	consistent itch hypersensitivity	
		blistering	
Fraccalvieri (2013)	Colour power Doppler –	VSS	
Traceatviett (2015)	vascularisation	Visual analogue scale pain and	
	vascularisation	itch	
Fraccalvieri (2011)	Histology		
Traceatvien (2011)	Echocontrastography		-
	neovoscularisation		
Conkondo (2014)	Dorm I ab combo arytheme &	mVSS (como porticipanto)	
Galikalide (2014)	electicity	mv SS (some participants)	-
$C_{2}(2022)$	elasticity	DOGAGO	DOGAG
Ge (2022)	-	POSAS-O	PUSAS
		Subjective reports on patient	
C K (2017)		range of movement	
Gee Kee (2016)	3D photography – thickness	POSAS-O	POSAS
Guo (2020)	Ultrasound – blood flow grade	-	-
	Shear wave elastography – scar		
	stiffness		

1144419 (2021)	-	-	-
Huang (2020)	Shear wave elastography – scar	-	-
	stiffness		
Issler-Fisher (2021)	-	VSS	POSAS-P
		POSAS-O	
Issler-Fisher (2020)	-	VSS	POSAS-P
		POSAS-O	Patient pain & itch scales
Issler-Fisher (2017)	-	VSS	POSAS-P
		POSAS-O	Patient pain, itch & quality of life
			rating scales
Joo (2020)	- 6	VSS	Pain severity (0-10 VAS)
Katz (1985)	Cicatrometer – firmness	-	-
Kemp Bohan (2021)	-	-	-
Kim (2018)	- ~ ~ ~ ~	-	-
Lacarrubba (2008)	-	Clinical evaluation of lesion size	-
Lau (2005)	-	VSS	-
Lee (2020)	-	mVSS (height, pliability,	POSAS-P
		vascularity, pigmentation)	
		POSAS-O	
Lee (2019)	-	mVSS (height, pliability,	POSAS-P
		vascularity, pigmentation)	
		POSAS-O	
Li (2013)	Micrometer – tissue thickness	- 06	-
	Force/torque sensor – load applied to		
	scar		
Li (2020)	Cutometer – elasticity	VSS	Quality of life questionnaire
	Mexameter – colour		
	PeriCam PSI system and mexameter		
	– blood supply		
Li (2021)	Laser Doppler flowmetry – perfusion	VSS	-
Li (2018)	Spectrocolourimeter – scar colour	VSS	Pain & itch (0-10 VAS)
Li (2021)	-	VSS	Treatment satisfaction
	Spectrocolourimeter – scar colour	VSS	Pain & itch (VAS scale not specifie
Li-Tsang (2005)		VCC	Doin & itch $(V \land S)$

Page 55 of 77

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Li-Tsang (2010) Lobos (2017)	Spectrocolorimeter – colour -	VSS (pliability) Modified Seattle Scar Scale	Pain & itch (10-point VAS)
Mamdouh (2021)			Patient satisfaction $(V \land S)$
Maintooun (2021)	-	V 55	Tatient satisfaction (VAS)
Miletta (2021)	- Colourmeter scor colour	- Unalgar likely DOSAS O	- Unalear likely DOSAS D
Wiletta (2021)	Dermal torque meter – scar	Olicical, likely 105A5-0	Short Form 36 Quality of Life Survey
Nedelec (2014)	Cutometer electicity		
Nedelee (2014)	Maxamatar colour	-	-
Nadalaa (2008)	Cutamatar alasticity	mVSS	
Nedelec (2008)	Maxamatar calour	111 v 55	-
Nadalaa (2010)	Mexameter – colour		
Nedelec (2019)	Cutometer – elasticity	-	-
N. 1.1. (2020)	Mexameter – colour		D^{+}_{-} 0 $(10 1)^{+}_{-}$ VAC
Nedelec (2020)	Cutometer – elasticity	-	Pain & itch (10cm line VAS)
	Mexameter – colour		
Nicoletti (2015)	-		-
Niessen (1998)	Histology		-
Reinholz (2020)	3D topographic imaging device	POSAS-O	Dermatology Quality of Life Index POSAS-P
Reinholz (2016)	Optical coherence tomography – thickness	POSAS-O	Dermatology Quality of Life Index POSAS-P
Schwaiger (2018)	3D topographic imaging device	-	-
Simons (2017)	3D camera – scar height	POSAS-O	-
Soykan (2014)	Slide calliper – dimensions	POSAS-O	POSAS-P
Timar-Banu (2001)	Metric ruler – dimensions	Validated 3-point scoring system	- /
× ,		for redness, hardness, itching &	
		pain	
Ud-Din (2019)	Optical coherence tomography –	-	-
	thickness		
	Histology		
van den Kerckhove (2005)	Chromameter – ervthema	-	-
van der Veer (2010)	Slide calliner – dimensions	-	-
Wang (2009)	Histology	_	-
Wang(200)	-	_	_
11 ung (2010)			

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(-	POSAS-O	Numeric rating scale for itch Toronto Paediatric Itch Scale CH-9D BBSIP
Wood (1996)	-	VSS	-
Xuan (2021)	Histology	-	-
Yeol Lee (2022)	Cutometer – elasticity Elastography	mVSS	-
Yim (2010)	Cutometer – elasticity Tewameter – trans-epidermal water loss Mexameter – colour	-	-
Zadkowski (2016)		VSS	-
Abstracts	Cr.		
Agabalyan (2016)	Histology		-
Bajouri (2018)	-	VSS	-
Bezugly (2019)	Clinical or histopathological diagnosis		-
Bezugly (2014)	-	-	-
Blome-Eberwein (2011, 2012)	Doppler vascularity, elasticity and sensation	VSS	Pain and itching scale (0-10 Likert scale)
Blome-Eberwein (2012)	-	-	-
Blome-Eberwein (2014)	Doppler flowmeter – vascularity Cutometer – pliability Semmes-Weinstein monofilament aesthesiometer testing set – sensation	VSS	POSAS-P
Cho (2012)	-	VSS	-
Cho (2012)	CK-MPA Multi-Probe adaptor – pigmentation, erythema and trans- epidermal water loss Cutometer – elasticity	-	-
Comstock (2018)	Computer-based tools – Thickness & pliability	Unclear, likely POSAS-O	Unclear, likely POSAS-P

Page 57 of 77

 BMJ Open

Cooper (2021)	Colorimeter – pigmentation	Unclear, likely POSAS-O	Unclear, likely POSAS-P
Du (2006)	-	-	-
Edgar-Lacoursière (2022)	Cutometer – elasticity	-	-
	Mexameter – colour		
El-Zawhary (2007)	Histology	-	-
George (2019)	-	-	-
Jacobs (2016)	Cutometer – pliability	POSAS-O	-
	Colorimeter – colour		
Jang (2009)	Mexameter – pigmentation	-	-
	Tewameter – trans-epidermal water		
	loss		
	Sebumeter – sebum		
	Cutometer – elasticity		
	Laser Doppler – perfusion		
Kim (2009)	Histology	VSS	-
Li (2016)	Spectrocolourimeter – scar colour	VSS	Patient report of pain & itch
Li-Tsang (2011)	-	VSS (thickness, pliability and	
()		pigmentation)	
Li-Tsang (2010)	Histology	VSS	Self-report questionnaire
	Spectrocolourimeter – scar colour		
Maari (2017)	Cutometer – elasticity		_
(2017)	Mexameter – nigmentation		
Moortgat (2020)	Cutometer – elasticity	Unclear likely POSAS-O	Unclear likely POSAS-P
10011gat (2020)	Chromameter – colour	onerear, nikely i obrid-o	Oncical, intery i Obrid-i
	Tawamatar trans anidermal water		
	1055 Cornoomotor hydration		
Nadalaa (2018)	Conconnector electicity		
Nedelec (2018)	Cutometer – elasticity	-	-
D ((2010)	Mexameter – colour	DOGAGO	
Peters (2018)	Cutometer – elasticity	POSAS-O	POSAS-P
A A A A A	Colourimeter – colour		
Seo 2011	Cutometer – elasticity		
Siwy (2016)	Colourimeter – colour	-	SF-36 Quality of Life Measurement
	Torque meter – pliability & elasticity		POSAS-P
Timina (2013)	-	-	-

Tu (2014)	-	VSS	-	
Ud-Din (2017)	Laser perfusion imaging	-	-	
	Optical coherence tomography –			
	thickness			
	Histology			
Ud-Din (2017)	Optical coherence tomography –	-	-	
	thickness			
Ud-Din (2018)	Optical coherence tomography –	-	-	
	thickness			
	Histology			
Zuccaro (2021)	Multi-parameter skin analysis device	VSS	Unclear, likely POSAS-P	
		Unclear, likely POSAS-O	· · ·	
Zuccaro (2019)	Acoustic radiation force impulse	-	-	
× /	ultrasound elastography			
Zuccaro (2021)	Acoustic radiation force impulse –	VSS	POSAS-P	
	stiffness	POSAS-O (did not include		
	DermLab Combo elasticity probe –	surface area and relief subscales)		
	elasticity	NI .		
	DermLab Combo colour probe –			
	colour			
POSAS-P: POSAS patien	t scale); VAS: Visual Analogue Scale; CHU-9D:	Child Health Utility-9D; BBSIP: Bri	isbane Burn Scar Impact Pro	file
				Page 10 of 28
	For peer review only - http://bmjop	en.bmj.com/site/about/guidelines.xht	ml	

BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

Supplementary Table 3: Reliability of ultrasound methods reported in each included study

First Author (year)	Reliability Test & Measurement Error	Reliability & Measurement Error Test Statistics & Details
Inter-rater reliability		
Anthonissen (2015)	ICC; SEM	Epidermal – 0.297; 0.02mm
		Dermal – 0.991; 0.13mm
Chang (2014)	Pearson correlation	R=0.90, p<0.001
Dunkin (2007)	N.R.	N.R.
Fong (1997)	ICC	0.93, p=0.146
Gankande (2014)	ICC (95% CI)	Individual site:
	× ,	Rater 1 vs rater 2
		'Best scar' – 0.95 (0.92, 0.96)
		'Worst scar' $-0.95(0.91, 0.97)$
		'Normal skin' – 0.94 (0.91, 0.96)
		Rater 1 vs rater 3:
		'Best scar' $-0.86(0.78, 0.91)$
		'Worst scar' $-0.91(0.85, 0.95)$
		'Normal skin' -0.92 (0.88, 0.95)
		Rater 2 vs rater 3:
		'Best scar' $= 0.93 (0.89, 0.95)$
		'Worst scar' $= 0.96 (0.92, 0.93)$
		(Normal skin' - 0.95 (0.92, 0.97))
		Average site:
		Rater 1 vs rater 2
		'Best scar' $= 0.97 (0.94, 0.99)$
		'Worst scar' $= 0.98 (0.96, 0.99)$
		(Normal skin' - 0.97 (0.93, 0.99))
		Rater 1 vs rater 3
		'Best scar' $= 0.90 (0.77, 0.95)$
		Worst scar' = 0.97 (0.91, 0.98)
		(Normal skin' = 0.96 (0.92, 0.98))
		$\mathbf{R}_{\text{other}} = 0.90 (0.92, 0.98)$
		(Rate 2 vs rate 2 (Rest scar' $0.05(0.88, 0.08)$)
		$\frac{1}{2} \frac{1}{2} \frac{1}$
		$(Normal skin^2 - 0.98 (0.94, 0.99))$
$I_{00}(2005)$	ICC	0.84 p < 0.01
Lau(2003)		0.04, p > 0.01
Lee (2020)	ICC (059/ CI): SEM	Acceptable to high
Lee (2019)	ICC (95% CI), SEM	$\frac{5\text{Cal.}}{\text{Single: 0.057 (0.024, 0.072)}}$
		Single. $0.957(0.954-0.975)$
		Average: $0.983 (0.977-0.991)$
		SEIVI. U. IU IIIII Unscorred skin:
		$\frac{OIISCAITCU SKIII.}{Single: 0.067 (0.040, 0.080)}$
		Siligie. 0.907 (0.949-0.980)
		Average. 0.969 (0.982-0.993)
Nadalaa (2000)	ICC (0.5% CI)	SEIVI. 0.04 IIIII Most source coer: 0.00 (0.84.0.05)
$\frac{1}{2008}$	ICC (95% CI)	$\frac{1}{10000000000000000000000000000000000$
		Less severe scar. $0.91 (0.83 - 0.93)$
		Donot site: $0.85 (0.82 - 0.94)$
$Q_{}(2011)$	ND	Normal skin: $0.85 (0.75-0.92)$
Seo (2011)	N.K.	"High"
Simons (2017)	ICC (95% CI); SEM	Scar: 0.82 (0.7-0.89); 0.05 cm
		Normal skin: $0.33 (0.08-0.54)$; 0.03 cm
Van Den Kerckhove	ICC (95% CI); SEM	One day:
(2003)		0.88 (0.81-0.95); 0.29 mm

BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

		$\frac{\text{Day-to-day:}}{2}$
		0.94 (0.90-0.98); 0.21mm
Intra-rater reliability		
Anthonissen (2015)	ICC; SEM	Epidermal = 0.809; 0.01mm
		Dermal – 0.991; 0.13mm
Gankande (2014)	ICC (95% CI)	'Best scar' – 0.97 (0.89, 0.94)
		'Worst scar' – 0.92 (0.88, 0.95)
		'Normal skin' – 0.86 (0.81, 0.89)
Gee Kee (2016)	N.R.	N.R.
Lau (2005)	ICC	Intra-rater: 0.98, p<0.01
Lee (2019)	ICC (95% CI)	Scar:
		Single: 0.951 (0.871-0.987)
		Average: 0.983 (0.953-0.966)
		SEM: 0.10 mm
		Unscarred skin:
		Single: 0.948 (0.881-0.976)
		Average: 0.982 (0.954-0.993)
		SEM: 0.04 mm
Li (2013)	ICC	0.89
Seo (2011)	NR	"High"
Simons (2017)	ICC (95% CD ⁻ SEM	Scar: $0.95 (0.91-0.97) \cdot 0.02 \text{ cm}$
		Normal skin: $0.61 (0.41-0.75)$: 0.02 cm
Van Den Kerckhove	ICC (95% CD: SEM	$0.98 (0.97-0.99) \cdot 0.11 \text{mm}$
(2003)		0.90 (0.97 0.99), 0.111111
Wang(2010)	SE	Peak: 0.032
		3 months: 0.018
		6 months: 0 309
		9 months: 0.353
		7 monuis. 0.333

Abbreviations used in tables: N.R.: Not reported; ICC: Intraclass Correlation Coefficient; 95% CI: 95% Confidence Interval; SEM: Standard Error of Measurement; SE: Standard Error

Summary of findings for measurement error:

The reported inter-rater SEM measurements for the combined (i.e., epidermal and dermal) thickness measurement of scars was reported in two records as 0.11 mm^4 and 0.5 mm^5 The inter-rater SEM for the combined thickness measurement of unscarred skin was also calculated in one record (SEM = 0.3 mm).⁵ The inter-rater SEM was calculated in one record for the measurement of epidermal (SEM = 0.02 mm) and dermal (0.13) measurements⁶, and one record reported only the dermal SEM for scar thickness (SEM = 0.1 mm) and unscarred skin (0.04 mm).⁷ The intra-rater SEM for the combined thickness measurement of scarred skin ranged from 0.18 mm to 0.52 mm, and was measured at 0.2 mm for unscarred skin in one record.⁵ One record reported the intra-rater SEM for epidermal (0.01 mm) and dermal (0.12 mm),⁶ and one record reported the intra-rater SEM for dermal scar (0.1 mm) and unscarred skin (0.04).⁷

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Supplementary Table 4. Methodological considerations for researchers and/or clinicians undertaking measurement of scar thickness using ultrasound.

Consideration	Details & examples of considerations	Publications in our review addressing the consideration	Details reported in included review records
Preventing skin	Using standoff methods	5,8-12	- Use of ultrasound gel to prevent contact
compression	(e.g., ultrasound gel,		between ultrasound transducer and skin surface
during measurement	water bath) to prevent transducer touching the		to minimise compression applied by direct application of transducer ^{5,8-11}
	skin		- Silicone pad placed underneath transducer ¹²
	Application of minimal pressure by transducer	13-17	- Transducer held to maintain minimal pressure on scar ^{13,14,16}
	1 5		- Training users to apply minimal force on transducer to prevent scar or skin distortion ^{15,17}
	Deliberately	18-20	- Measurement of thickness with and without
	compressing skin to		compression with transducer ^{18,20}
	quantify scar		- Thickness measurements taken using TUPS,
	compressibility		which uses controlled and metered compression during measurement ¹⁹
)rienting the	Orienting the patient	7,17,21	- Patient supine throughout measurement to
oatient	during measurement		allow measurement to be taken in the same
	(e.g., upright, supine,		position ^{7,17,21}
	prone or seated)		
	Maintaining patient	8	Patients asked to hold breath during
	stillness during		measurement of scars on the chest to allow
	measurement		shear-wave ultrasound ⁸
Placing	Orientating ultrasound	22	- Direction of transducer recorded to ensure
ultrasound	transducer [e.g.,		consistency ²²
transducer	vertical (superior to		
	inferior/cranial to		
	caudal), horizonal		
	(medial to lateral)]		

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Orienting the transducer in relation to the scar (e.g.,	8,14,16,17,21,23-25	- Transducer oriented perpendicular to the skin surface to provide optimal image ^{8,14,17,21,23-25}
perpendicular) Measuring difficult/tight areas (e.g., axillae or other	5	- Exclusion of fingers and toes in paediatric measurements due to size of measurement area and thin skin ⁵
Mapping measurement area (e.g., tracing,	5,11,15,17,19,21,26-31	- Scars traced using translucent paper 17,19,21,26,28,30,31
schematic diagram)		 Scars and surrounding anatomical landmarks traced using translucent paper ¹⁵ Scar mapped on transparent paper, which was then cut out ²⁷ Scar mapped with drawing, no elaboration
Photographing	23,25,32	 Scar mapped with drawing, no claboration provided ²⁹ Scars traced using Visitrak (Smith & Nephew Medical Limited, England) ^{5,11} Assessed area marked and photograph taken in
measurement area		initial consultation ^{23,32}
Measuring specific scar locations (e.g., centre of scar, worst area of scar, counting transducer lengths)	5,7,8,12,18-20,22,29,32-36	 Photographs of scars taken ²⁵ Measurement taken at standardised transducer lengths along surgically created scars of prespecified dimensions ³³ Measurements taken at thickest/most severe point ^{18-20,29,32,34,36}
		as determined by the patient and/or clinician ⁷ - Transducer placed on thickest site on periphera regions ⁸ - Transducer placed on area initially identified to
	transducer in relation to the scar (e.g., perpendicular) Measuring difficult/tight areas (e.g., axillae or other joints) Mapping measurement area (e.g., tracing, schematic diagram) Photographing measurement area Measuring specific scar locations (e.g., centre of scar, worst area of scar, counting transducer lengths)	transducer in relation to the scar (e.g., perpendicular) Measuring5difficult/tight areas (e.g., axillae or other joints) Mapping measurement area (e.g., tracing, schematic diagram)5,11,15,17,19,21,26-31 area (e.g., tracing, schematic diagram)Photographing measurement area23,25,32Photographing measurement area5,7,8,12,18-20,22,29,32-36Measuring specific scar locations (e.g., centre of scar, worst area of scar, counting transducer lengths)5,7,8,12,18-20,22,29,32-36

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Conducting linear measurements from nearby anatomical landmarks Acclimatising scar to measurement conditions Removing scar to ultrasound measurement

Acclimatising patient to 4,17,21,28,40-45 room prior to measurement - Measurement area selected by the measurer with -selected area marked with tape ¹²

- Measurements taken at set linear distances from cranial/caudal border of linear sternal scar ³⁵

- Linear measurements from anatomical landmark to measurement site ¹⁶

- Transducer placement mapped in 3dimensional space using a surgical precision tracking arm ³⁷

- Pressure garments removed 10 minutes before measurement ²⁷

- Pressure garments removed 15 minutes before measurement to regain original (uncompressed) scar thickness or to reduce blanching effects on measurement ^{19,39}

- Pressure garments/gels/moisturisers removed 20 minutes before measurement ^{7,21,28}

- Pressure garments removed 30 minutes before measurement ^{11,24,25,38}

- Sequential measurement of scars following direct treatment with vacuum massage at 5, 30, 60 and 120 minutes to monitor effect of treatment ²³

- Patients rested for minimum 5 minutes before measurement ^{4,17,21}

- Scar exposed to room conditions for 10 minutes ²⁸ to allow equilibrium to be reached with surrounding environment ⁴⁰

- Patients resting in room with constant temperature for 15 mins ⁴¹ to allow scar to stabilise ⁴³

5.53 Maintaining patient position before measurement

Measuring different skin and/or dermis lavers individually

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44 45 46 Measuring epidermis

Measuring both epidermis/dermis combined (no

4,5,7,10,11,14,16,17,21,22,25,27,34,39,54-67

- Patients rested for 20 minutes prior to measurement 28,44

- Patients resting for 10 minutes before repeated measurements taken ⁴²

- Patients wait in testing room holding position for 5 min before measurement to stabilise cutaneous blood flow ⁴

- Patients allowed to adapt in controlled room to exclude external variables 45

- Patients remained supine for at least 5 minutes before measurement to avoid artefacts on Doppler imaging ¹²

- Patients allowed to acclimatise to room and assumed a supine position for a minimum of 10 minutes before measurements of biophysical parameters ¹⁰

- Measurement of epidermal, dermal and combined epidermal and dermal thickness to allow comparison with histological measurement 46.47

- Measurement of the epidermal and dermal thickness ^{44,48}, combined with layer acoustic density⁶

- Measurement of the epidermal, dermal and subcutaneous thickness, combined with acoustic density ^{49,50}

- Measurement of dermal thickness as treatment thought to affect/target the dermis ^{23,36,51-53}

- Combined epidermal and dermal thickness measurement to provide information on the full thickness of the scar 4,5,7,10,11,14,16,17,21,22,25,27,34,39,54-67

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Measurement objective	individual measurement) Measuring fibrosis/oedema/hair follicles	7,9,10,12,13,15,16,23,24,28-31,33,35,36,44,53,57,59,60,62,63,68-81	- Measurement of fibrosis or collagen architecture ^{7,10,16,23,28-31,33,35,36,44,53,57,60,62,63,68,69,71- 73,76-78,81}
			 Measurement of inflammation/oedema ¹³ Quantification of the sub epidermal low echogenic band, indicating oedema ⁵⁹ Measurement of both fibrosis and oedema ^{9,12,15,24,57,70,74,75,79,80}
Factors influencing scar site measurement	Measuring contralateral skin/control scar	8,13,14,22,28,29,51,54-57,82-87 5,7,11,17,21,24,37,42,53,58-60,65,88,89 38,39,44,78,80,81	 Measurement of the presence and density of hair follicles to differentiate scarred and unscarred skin⁵³ Measurement of additional, non-scarred subjects ^{54,78} Measurement of unscarred/unaffected skin on same subject as scar measurement contralaterally or at anatomically similar location to provide normative measurements for skin thickness 5,7,8,11,13,14,17,21,22,28,29,37-39,42,44,51,53,55-60,65,80,84-89
			 Measurement of both untreated scar and unaffected skin ⁸¹⁻⁸³ Measurement of a control scar subjected to care as usual treatment on the same individual ²⁴
	Measuring open wounds or sores in the scar	5	- Use of flexible transparent plastic wrap placed over the measurement area to prevent contact between ultrasound gel and transducer with the
	Operator training and/or experience	5,7,11,13,15,17,19,23,26-28,30,38,39,57,60,65,71,72,86,90-92	 Trained outcome assessor ^{5,12,15,17,26,71} Measurements taken by radiologist/sonographer ^{27,65,72,91}

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4,5,7,8,10,11,19,22,24,25,30,33,36,39,43,44,46,51,53,56,59,60,65,67,78.84,91.93

measurements per scar

.3,44,46,...

- Assessors with burn experience ^{86,92}

- Ultrasound located in department of radiology 90

- Measurements conducted by trained therapist/doctor under guidance of experienced radiologist 11,13,28,38

- Measurements conducted by trained clinicians who use device regularly and received training by company representative of devices 7,60

- Device-specific training provided: 1 week ¹⁹; 3 sessions of 3 hours for 3 weeks, plus 10 independent assessments of scars using study protocol ³⁹; training provided over 3 months ³⁰; physical therapist trained in ultrasound application ²³

- 3 ultrasound images taken from each patient 8,10,25,30,36,43,44,46,51,53,56,59,78,84

- Clearest of 3 measurements used ¹¹

- 3 measurements in 3 locations across scar used. Individual and average measurements reported ³⁹

- Measurements performed in duplicate ^{33,93}

- Measurements taken at different points of the

scar, thickest used for analysis ⁹¹

- 5 measurements of each site 5,22

- 9 measurements taken, removal of maximum and minimum, 7 measurements used for average 19

- Measurements taken by 3 assessors at 3 different time points during day 7,60

- Measurement of 2 sites on the same scar²⁴

- Single ultrasound image taken for analysis 67

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3 4	Use of	Using additional	5,8-10,12,14,16,17,20-22,24-26,28,30,31,34,35,39-47,49,52,55-58,65,67-69,74-	- Histology/immunohistochemistry
+ 5	additional	objective assessment	79,81-83,85-91,94-110	12,16,46,47,49,57,77,78,87,99,102,107,109
6	measurement	instruments (e.g.,		- Blood flow and blood perfusion measurement
7	tools as well as	histology, colour		using laser Doppler perfusion imaging,
8	ultrasound	Doppler ultrasound		flowmetry or PeriCam and scar colour and
9	measurements	cutometer		micro-vessel percentage using dermosconvolour
10	measurements	eutometer,		and micro voggal parcentage
11		colourimeter)		and micro-vesser percentage.
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13				- Oximeter ⁴⁰
14				- Infra-red camera ⁴⁰
15				- Measurement of scar stiffness or
16				pliability/elasticity using elastography or
17				cutometer 8,14,17,20,21,24-26,28,42,45,52,56,65,81-
18				83,85,88,89,95,97,98,100,103-105
19				- Measurement of sensation using Semmes-
20				Weinstein filaments ^{81-83,85}
21				- Measurement of scar colour (including
22				nigmentation and erythema) using
23				spactronhotomator, colourimator, chromomotor
24				spectrophotometer, colourinteter, chromatheter,
25				26 31 41 43 45 52 55 65 67 70 81 86 80 00 95 98 100 106 110
27				20,51,41,45-45,52,55,05,07,77,81,00,67,70,55-96,100-100,110
28				- Measurement of trans-epidermal water loss
29				using Tewameter or scar hydration using
30				Corneometer ^{45,52,95,98}
31				Measurement of sebum level using sebumeter
32				95,98
33				- Measurement of hardness using durometer ⁹⁰
34				- Measurement of neovascularisation using
35				echocontrastography 57
0C 7C				- Measurement of scar dimensions (e.g., scar
2/ 20				height and volume) using 3D camera 3D
30				imaging methods ruler or calliner 5.9.10.22.35.74.76
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44 45 46 Using subjective assessment instruments (e.g., clinical rating scales, PROMs) 18,19,22,27-29,32,36,39,40,43,44,48,51,55,56,60,65,66,68-71,79-83,85,86,90,91,93-97,99,110-114 - Measurement of skin thickness using micrometer or optical coherence tomography 16,30,58,75,107-109

- Measurement of scar firmness or deformation using cicatrometer, force/torque sensor (in line with ultrasound to measure load applied) or torque meter ^{30,31,106}

- Multi-parameter skin analysis device ⁶⁵

- Measurement of erythema and elasticity using probes of DermaLab Combo ³⁹

- Multi-probe adaptor taking multiple measurements (pigmentation, erythema, transepidermal water loss) ⁹⁵ PROMs:

- Measurement of scar quality using POSAS patient report ^{7,22,29,32,44,55,60,62,63,65,74-}76,81,85,94,96,105,106,113,114

- Subjective rating scales for scar symptoms (e.g., pain, itch) or subjective scar severity ratings ^{25,29,40,41,52,62,63,71,79,82,83,92,101,102,110,114}

- Patient quality of life questionnaires 74,75,100,106
- Measurement of generic health-related quality of life using CHU-9D ^{62,63}

- Measurement of scar-specific health-related quality of life using BBSIP ^{62,63}

- subjective evaluation of response to treatment/treatment satisfaction ^{80,115} Clinical rating scales:

- Measurement of scar quality using POSAS observer report 7,22,29,32,44,52,55,60,62,63,65,74-76,81,85,86,96,97,105,113-115

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Determining the order ⁵

of measurement

- Measurement of physical scar characteristics using VSS or modified versions of the VSS ^{7,17-} 19,27,29,32,34,36,37,39,41-43,48,55,56,60,64,65,68-71,79-85,91-94,99-102,110-112,114,116,117

- Measurement of scar characteristics in relation to unscarred skin using Seattle Scar Scale or modified Seattle Scar Scale ⁷²

- Subjective rating scales for scar symptoms (e.g., pain, itch) as assessed by the clinician and/or researcher and/or clinical evaluation of scar severity ^{10,28,40,51,56,66,72,90,91,93,95}

Standardised order of measurement: 3D photograph, POSAS-O, then ultrasound ⁵
 Order of device use not specified 34,68,69,82,83,85,86,91,98,100,107

Abbreviations: TUPS: Tissue Ultrasound Palpation System; 3D: three-dimensional; POSAS: Patient and Observer Scar Assessment Scale; CHU-9D: Child Health Utility 9D; BBSIP: Brisbane Burn Scar Impact Profile; VSS: Vancouver Scar Scale; mVSS: Modified Vancouver Scar Scale; POSAS-O: Patient and Observer Scar Assessment Scale, observer measure

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Preferred Reporting Items for Systematic reviews and Meta-Analyses extension for Scoping Reviews (PRISMA-ScR) Checklist

SECTION	ITEM	PRISMA-ScR CHECKLIST ITEM	REPORTED ON PAGE #
TITLE			
Title	1	Identify the report as a scoping review.	1
ABSTRACT			
Structured summary	2	Provide a structured summary that includes (as applicable): background, objectives, eligibility criteria, sources of evidence, charting methods, results, and conclusions that relate to the review questions and objectives.	3-4
INTRODUCTION			
Rationale	3	Describe the rationale for the review in the context of what is already known. Explain why the review questions/objectives lend themselves to a scoping review approach.	5-7
Objectives	4	Provide an explicit statement of the questions and objectives being addressed with reference to their key elements (e.g., population or participants, concepts, and context) or other relevant key elements used to conceptualize the review questions and/or objectives.	7
METHODS			
Protocol and registration	5	Indicate whether a review protocol exists; state if and where it can be accessed (e.g., a Web address); and if available, provide registration information, including the registration number.	7
Eligibility criteria	6	Specify characteristics of the sources of evidence used as eligibility criteria (e.g., years considered, language, and publication status), and provide a rationale.	8-10
Information sources*	7	Describe all information sources in the search (e.g., databases with dates of coverage and contact with authors to identify additional sources), as well as the date the most recent search was executed.	8
Search	8	Present the full electronic search strategy for at least 1 database, including any limits used, such that it could be repeated.	9
Selection of sources of evidence†	9	State the process for selecting sources of evidence (i.e., screening and eligibility) included in the scoping review.	9
Data charting process‡	10	Describe the methods of charting data from the included sources of evidence (e.g., calibrated forms or forms that have been tested by the team before their use, and whether data charting was done independently or in duplicate) and any processes for obtaining and confirming data from investigators.	10-11
Data items	11	List and define all variables for which data were sought and any assumptions and simplifications made.	10-11 and supplementary table 1
Critical appraisal of individual	12	If done, provide a rationale for conducting a critical appraisal of included sources of evidence; describe	N/A





SECTION	ITEM	PRISMA-ScR CHECKLIST ITEM	REPORTED ON PAGE #
sources of evidence§		the methods used and how this information was used in any data synthesis (if appropriate).	
Synthesis of results	13	Describe the methods of handling and summarizing the data that were charted.	10-11
RESULTS			
Selection of sources of evidence	14	Give numbers of sources of evidence screened, assessed for eligibility, and included in the review, with reasons for exclusions at each stage, ideally using a flow diagram.	11-12
Characteristics of sources of evidence	15	For each source of evidence, present characteristics for which data were charted and provide the citations.	12-15
Critical appraisal vithin sources of evidence	16	If done, present data on critical appraisal of included sources of evidence (see item 12).	N/A
Results of individual sources of evidence	17	For each included source of evidence, present the relevant data that were charted that relate to the review questions and objectives.	Results section (11-46)
Synthesis of results	18	Summarize and/or present the charting results as they relate to the review questions and objectives.	Results section (11-46)
DISCUSSION			
Summary of evidence	19	Summarize the main results (including an overview of concepts, themes, and types of evidence available), link to the review questions and objectives, and consider the relevance to key groups.	47-49
Limitations	20	Discuss the limitations of the scoping review process.	49-50
Conclusions	21	Provide a general interpretation of the results with respect to the review questions and objectives, as well as potential implications and/or next steps.	50-51
FUNDING			
Funding	22	Describe sources of funding for the included sources of evidence, as well as sources of funding for the scoping review. Describe the role of the funders of the scoping review.	51

JBI = Joanna Briggs Institute; PRISMA-ScR = Preferred Reporting Items for Systematic reviews and Meta-Analyses extension for Scoping Reviews.

* Where *sources of evidence* (see second footnote) are compiled from, such as bibliographic databases, social media platforms, and Web sites.

† A more inclusive/heterogeneous term used to account for the different types of evidence or data sources (e.g., quantitative and/or qualitative research, expert opinion, and policy documents) that may be eligible in a scoping review as opposed to only studies. This is not to be confused with *information sources* (see first footnote).
‡ The frameworks by Arksey and O'Malley (6) and Levac and colleagues (7) and the JBI guidance (4, 5) refer to the process of data extraction in a scoping review as data charting.

§ The process of systematically examining research evidence to assess its validity, results, and relevance before using it to inform a decision. This term is used for items 12 and 19 instead of "risk of bias" (which is more applicable to systematic reviews of interventions) to include and acknowledge the various sources of evidence that may be used in a scoping review (e.g., quantitative and/or qualitative research, expert opinion, and policy document).

From: Tricco AC, Lillie E, Zarin W, O'Brien KK, Colquhoun H, Levac D, et al. PRISMA Extension for Scoping Reviews (PRISMAScR): Checklist and Explanation. Ann Intern Med. 2018;169:467–473. doi: 10.7326/M18-0850.



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Ultrasound measurement of traumatic scar and skin thickness: A scoping review of evidence across the translational pipeline of research-to-practice

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Ultrasound measurement of traumatic scar and skin thickness: A scoping review of evidence across the translational pipeline of research-to-practice*

<u>Running Title:</u> Review of scar thickness measurement with ultrasound

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ABSTRACT:

 Objectives: To identify the ultrasound methods used in the literature to measure traumatic scar thickness, and map gaps in the translation of these methods using evidence across the research-to-practice pipeline.

Design: Scoping review

Data Sources: Electronic database searches of Ovid MEDLINE, Embase, Cumulative Index of Nursing and Allied Health Literature (CINAHL) and Web of Science. Grey literature searches were conducted in Google. Searches were conducted from inception (date last searched 27/05/2022).

Data Extraction: Records using B-mode ultrasound to measure scar and skin thickness across the research-to-practice pipeline of evidence were included. Data was extracted from included records pertaining to: methods used; reliability and measurement error; clinical, health service, implementation and feasibility outcomes; factors influencing measurement methods; strengths and limitations; and use of measurement guidelines and/or frameworks.

Results: Of the 9309 records identified, 118 were analysed (n = 82 articles, n = 36 abstracts) encompassing 5213 participants. Reporting of methods used was poor. B-mode, including high-frequency (i.e., > 20 MHz) ultrasound was the most common type of ultrasound used (n = 72 records; 61% of records), and measurement of the combined epidermal and dermal thickness (n = 28; 24%) was more commonly measured than the epidermis or dermis alone (n = 7, 6%). Reliability of ultrasound measurement was poorly reported (n=14; 12%). The scar characteristics most commonly reported to be measured were epidermal oedema, dermal fibrosis and hair follicle density. Most records analysed (n = 115; 97%) pertained to the early stages of the research-to-practice pipeline, as part of research initiatives.

Conclusions: The lack of evaluation of measurement initiatives in routine clinical practice was identified as an evidence gap. Diverse methods used in the literature identified the need for greater standardisation of ultrasound thickness measurements. Findings have been used to develop nine methodological considerations for practitioners to guide methods and reporting.

STRENGTHS AND LIMITATIONS OF THIS STUDY:

- Use of the Australian Government Department of Health and Aged Care Medical Research Future Fund research-to-practice pipeline phases to categorise records allowed identification of gaps in the use of ultrasound for clinical practice.
- Clinical, health service, implementation and feasibility outcomes related to ultrasound measurement in included records were summarised to determine what is needed to close the research-to-practice gap for ultrasound measurement of scar thickness.
- A limitation is that only articles available in English or with an English abstract were considered for inclusion and data extraction, thus findings are likely most relevant to English speaking countries.

INTRODUCTION:

Traumatic cutaneous injury, caused by sharp object penetration (e.g., surgery or vaccination) or burns (including thermal, chemical and friction) may result in the formation of hypertrophic scarring. (1) Hypertrophic scars result from an aberrant cutaneous healing response that leads to the formation of red, raised scars, often accompanied by pruritus and skin tightening, which remain within the boundaries of the initial injury. (2-7) The sequelae of hypertrophic scars can impact on patient's physical and psychosocial quality of life. (8, 9)

A characteristic of hypertrophic scarring that both patients and clinicians have identified as being important, and which has subsequently been used as a way to measure clinical and treatment outcomes, is scar thickness. (9-17) Scar thickness can be measured both subjectively, through clinician assessment and patient-reported outcomes, or objectively, utilising medical imaging methods. (18, 19) The pathological complexity of hypertrophic scars means that they generally extend below the level of the surrounding skin, supporting the use of medical imaging modalities such as ultrasound for thickness quantification, as these are capable of providing information about subcutaneous structures and processes. (19, 20) Scar thickness measurement using ultrasound can be conducted in both clinical and research contexts. Where routine measurements like ultrasound are used to guide clinical decisionmaking and treatment, this practice is known as measurement-based care. (21)

Ultrasound is a safe, non-invasive and largely cost-effective (compared to other imaging modalities) imaging method with measurement utility in both adult and paediatric populations. (22-24) Modern B-mode (brightness mode) ultrasound, particularly high- (i.e., ≥20 MHz) or ultra-high frequency (30-100 MHz) (25) ultrasonography, allows differentiation between the epidermis and dermis, which permits quantification of skin layer-specific scar characteristics. This differentiation may allow assessors to observe and understand the pathological mechanisms of individual scars and adjust treatment protocols accordingly. (24,

Page 7 of 83

BMJ Open

26-31) Additionally, B-mode ultrasound is commonly used as the basis for other imaging methods, such as colour Doppler ultrasound or elastography, which can allow quantification of additional scar characteristics, such as their elastic properties. (26-29, 32, 33)

Despite the clinical advantages of B-mode ultrasound for scar thickness measurement, methods are poorly reported and lack standardisation in the literature. This casts doubt on the validity of clinical decision-making in measurement-based care initiatives (e.g., setting depth of AFCO₂ penetration) informed by research findings (e.g., response to treatment) where ultrasound measurements are used. (34) Lack of standardisation also makes between-study comparison, such as systematic reviews and meta-analyses, difficult, (35) and poor methodological reporting hampers the ability to accurately replicate findings. This scoping review focusses on mapping and identifying gaps in ultrasound methods and evaluation reported in the current literature along the research-to-clinical practice pipeline. (36) Methodological considerations for people performing ultrasound scar thickness measurements, including practitioners (herein termed assessors) using ultrasound in clinical practice are presented based on the review findings.

METHODS:

Protocol Publication and Review Structure:

The protocol for this review has been published *a priori*. (37) This scoping review was conducted and is reported according to the Arksey and O'Malley (2005) (38) framework. The steps outlined in this framework are: 1) identifying the research question; 2) identifying relevant records; 3) selecting appropriate records; 4) charting extracted data; and 5) collating, summarising and reporting the results. (38)

Research Question:

The primary question of this scoping review was: "What do we know and not know about the measurement of traumatic cutaneous scar thickness using ultrasound?" This question was addressed through exploration of: methods used; reliability and measurement error; clinical, health service, implementation and feasibility outcomes; factors influencing ultrasound imaging and measurement methods; strengths and limitations of measurement methods; and use of measurement guidelines and/or frameworks. While the focus of this review was the measurement of traumatic cutaneous scar thickness with ultrasound, methods used to measure the thickness of unscarred skin were reported where these were used in combination with measurement of scar thickness (e.g., as control or comparator measurements).

Identifying Relevant Records:

A standardised search strategy was developed and piloted with the assistance of a medical librarian using the concepts 'ultrasound', 'skin', 'thickness' and 'measure', with associated terms and truncations (supplementary box 1). Ovid MEDLINE, Embase, Cumulative Index of Nursing and Allied Health Literature (CINAHL) and Web of Science electronic databases were searched from conception to identify original studies (date last searched 27th May 2022).

The phrase 'ultrasound scar thickness measurement' was used to conduct additional searches in 1) Google Scholar, and 2) Google to identify original studies in grey literature, and studies not identified in database searches. Title and abstract searches in Google Scholar and Google were limited to the first 200 results. (39)

Record Selection:

Following de-duplication, six reviewers screened records using Covidence (Veritas Health Innovation, Melbourne, Australia; available at <u>www.covidence.org</u>) for eligibility according to the inclusion criteria (Table 1). Both peer-reviewed journal articles and abstracts were

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included to ensure that all the available and most recent methodological information was obtained. (40) Data collected from peer-reviewed journal articles was considered the primary source of data, with information from abstracts used to confirm or extend the journal data. The inclusion of abstracts will assist future authors to further investigate the information presented as full texts may become available. During both title and abstract and full text screening, one researcher (BM) screened all records as a single reviewer, while other researchers (MS, TM, TR, BD and ZT) screened records as a second reviewer. Conflicts were resolved through discussion between at least two authors to reach agreement. A third author was used as a tiebreaker where agreement could not be reached.

Table 1. Inclusion and exclusion criteria for studies included in the scoping review.

Inclusion	Exclusion
 Traumatic scars measured with ultrasound based on B-mode ultrasound (including high-frequency, ultra-high- frequency and Doppler) Measurements taken of living, human individuals Measurement of traumatic cutaneous scarring arising from penetration of the skin with sharp objects (including surgery or vaccination), or as a result of burns, (including thermal, chemical or friction) Articles written in English, or with English abstracts 	 Reviews, discussion papers, opinion pieces Measurement of non-traumatic scars (e.g., acne scars). Non-traumatic scars measured along with burn scars were included Measurement of skin thickness in non-traumatic conditions (e.g., diabetes) Measurement of skin thickness where there is no cutaneous involvement in the trauma (e.g., traumatic brain injury) Measurement using A-mode ultrasound

Charting the Data:

The data extraction table was developed in Microsoft Excel and piloted by two authors (BM and ZT) through independent extraction and comparison of data from two records. The table was then modified to include the scar characteristics (e.g., fibrosis, oedema) measured, measurer/assessor training, the number of measurements taken and funding sources

(Supplementary Table 1). Full text data extraction was completed by four authors (BM, MS, TM and ZT). An additional author (BD) independently extracted data from five randomly selected records, which was compared to data extracted by other authors. Minimal differences between data extracted by the independent author and that by other authors were observed, thus further independent extraction was not performed. As is typical in scoping reviews, the certainty or quality of evidence was not appraised. (38)

The research-to-practice pipeline published by the Australian Government Department of Health and Aged Care Medical Research Future Fund (figure 1) was used to categorise each included record based on their stated aims into one of the four phases. (36) Studies related to phase 1 of this pipeline, basic research, were only included in this review when data on scar or skin thickness pertained to human participants (table 1). Phase 2 of this pipeline included randomised controlled trials, while phase 3 included pragmatic and observational studies conducted outside randomised controlled trials. The final phase of this pipeline (phase 4) indicates initiatives used in routine clinical practice.

Where clinical (e.g., treatment satisfaction, scar symptoms), health service (e.g., efficiency, safety, effectiveness, equity, patient-centredness and timeliness) and implementation (e.g., acceptability, adoption, appropriateness, fidelity, cost, penetration and sustainability) outcomes were addressed, they were reported and defined according to Proctor *et al.* (41). For example, in the context of this scoping review, acceptability is defined as the level to which ultrasound is palatable amongst stakeholders (e.g., assessors), appropriateness is the perceived fit of ultrasound within regular clinical practice, and fidelity is the degree to which ultrasound is used in the way it was initially described. (41) Measurement instrument-specific feasibility outcomes defined by Prinsen *et al.* (42) are reported in the current review. These outcomes included ease of administration, standardisation, completion time, instrument cost and availability, and ease of score calculation. (42) Reliability and measurement error were

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defined according to COnsensus-based Standards for the selection of health Measurement INstruments (COSMIN) tools. (43, 44) Measurements with an intraclass correlation coefficient (ICC) of 0.7 or greater were considered reliable. (44) Measurement error was assessed by comparing the reported standard error of the measurement (SEM) with the reported smallest detectable change (SDC). Where the reported measurement error was smaller than the reported smallest detectable change, it was interpreted as indicating real change or variance can be detected, and that change or variance is not a result of error. (44)

Patient and Public Involvement

There was no patient and/or public involvement in the design, conduct, reporting or dissemination of information in this scoping review.

RESULTS:

Electronic database searches identified 9309 records. After removal of 3703 duplicate records, the titles and abstracts of 5606 records were screened for relevance according to the inclusion criteria (Table 1). Following full-text screening, 104 records proceeded to data extraction. Searches in Google and Google Scholar identified an additional 14 records, providing a total of 118 records for data extraction. Search and screening results are presented according to the Preferred Reporting Items for Systematic reviews and Meta-Analyses (PRISMA) flow diagram (supplementary figure 1). (45)

Record Characteristics:

Of the 118 records included in this review, 82 were journal articles (69%) and 36 were abstracts (31%) (Table 2), representing a total of 5213 participants (range 1-438; mode 20 participants per record). Adults aged 18 years and older were most commonly targeted in articles (n = 43 articles; 52% of articles), (17, 26, 29, 46-85) while most abstracts did not report the age group measured (n = 25 abstracts; 69% of abstracts). (86-110) The most

common scar type measured was burn scars in both journal articles ($n = 43$ articles; 52% of
articles), (17, 22-24, 27, 47, 57-59, 61, 62, 64-67, 71-75, 81, 82, 84, 111-130) and abstracts (n
= 23 abstracts; 64% of abstracts) (28, 30, 86-88, 91-94, 96, 98, 102-106, 131-135) (Table 2).
Most identified articles used ultrasound measurement of scar thickness as part of research
initiatives, and were categorised as either phase 2 ($n = 50$ articles; 61% of articles) (17, 22,
26, 31, 46-49, 51-56, 61, 63-65, 67, 69-71, 74-76, 78, 81, 83, 84, 111, 112, 114, 115, 117,
124-127, 129, 130, 136-145) or phase 3 (n = 30 articles; 37% of articles). (23, 24, 27, 29, 50,
57-60, 62, 66, 68, 72, 73, 77, 79, 80, 82, 85, 116, 118, 120-123, 128, 146-149) on the
research-to-practice pipeline. (36) Phase 2 was also the most common phase represented by
abstracts (n = 21; 58% of abstracts), (86, 88, 91, 93, 95, 97, 99-104, 106-108, 131-134, 150,
151) followed by phase 3 (n = 15 abstracts; 42% of abstracts). (28, 30, 87, 89, 90, 94, 96, 98,
105, 109, 110, 135, 152-154) Phase 4 was addressed by two articles (2% of articles) (113,
119) and one abstract (2% of abstracts), (92) which used ultrasound to measure treatment
response to an intervention already used in routine clinical practice, including compression
garments (113, 119) and CO_2 fractional laser. (92) No records pertained to phase 1.

Page 11 of 39

Characteristic	Category	Number of Records (Translational Pipeline Phase 2*)	Number of Records (Translational Pipeling Phase 3*)	Number of Records (Translational Pipeling Phase 4*)
Journal Articles		Tipenne Thase 2)	Tipenne Thase 5)	Tipenne Thase 4)
Funding	Commercial	2	1	1
Source	Non-	23	13	0
	commercial		10	Ŭ
	Commercial	2	1	1
	& Non-	2	1	1
	commercial			
	No funding	6	3	0
	Not reported	16	12	0 0
Population	Adult	27	12	0
Type	Paediatric	6	10 4	0
rype	Paediatric	13	7	2
	and Adult	15	,	2
	Not reported	3	3	0
Scar Actiology	Burn	22	18	1
Seal Trettology	Surgical [†]	5	2	0
	Mixed	10	3	0 0
	Not specified		7	0
Abstracts			· · · ·	
Funding	Commercial	0	0	0
Source	Non-	3	1	0
	commercial			
	Commercial	0	0	0
	& Non-			
	commercial			
	No funding	0	0	0
	Not reported	17	14	1
Population	Adult	1	2	0
Type	Paediatric	0	3	0
rype	D 1' / '	4	1	0
	Paediatric	4		0
	Paediatric and Adult	4		0
	and Adult Not reported	4		1
Scar Aetiology	and Adult Not reported Burn	4 15 12	9 10	1
Scar Aetiology	Paediatric and Adult Not reported Burn Surgical [†]	4 15 12 1	9 10 2	1 1 0
Scar Aetiology	Paediatric and Adult Not reported Burn Surgical [†] Mixed	4 15 12 1 2	1 9 10 2 1	1 1 0 0

Table 2. Summary of characteristics of records included in this review*

Legend: Paediatric: measurement of patients under the age of 18; Adult: measurement of patients aged 18 years or older; Burn: scars caused by thermal, chemical or friction injury; Surgical: scars caused by surgical procedures (including biopsies); Mixed: scars of included record were of mixed origin (e.g., burn and acne)

Footnotes: *Stage in the research to clinical practice translational pipeline, as defined by the Australian Government Department of Health and Aged Care (36); [†]Type of surgery defined in supplementary table 2

* A breakdown of each characteristic per record is presented in Supplementary Table 2

Methods used to measure traumatic cutaneous scar thickness:

B-mode, including high-frequency (i.e., ≥ 20 MHz) B-mode ultrasound was the most

commonly reported ultrasound type used in both articles (n = 56; 68% of articles) (17, 22-24,

26, 29, 31, 46-49, 53, 54, 56, 57, 59, 60, 64, 65, 67, 69-78, 80-82, 84, 85, 111, 112, 114, 116-

118, 120, 122, 123, 126-130, 138, 139, 141, 142, 144-146, 149), while most abstracts did not

- report the type of ultrasound used (n = 22; 61% of abstracts) (86, 87, 92-98, 101, 103, 105,
- 106, 108, 131-134, 150-153) (Table 3). Specialised B-mode ultrasound devices, including the

Tissue Ultrasound Palpation System (TUPS; a B-mode ultrasound transducer in-series with a

load cell to allow measured compression of the skin), (68, 99, 100, 124) and colour Doppler

ultrasound, (52, 149) were used in six records (Table 3).

Page 13 of 39

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12 Table 3. Summary of measurement methods used in included record*

The type of scar and skin thickness measurement (i.e., thickness of the dermis, epidermis, or combined epidermal and dermal measurement) was reported in 39 records (33%) (Table 3). Where reported, combined measurement of epidermal and dermal thickness was the most common method used in articles (n = 32; 76% of articles reporting skin measurement type). (17, 22-24, 27, 29, 50, 53, 56-58, 60, 64-66, 70, 72-77, 80-82, 114, 116, 118, 122, 126, 127, 130, 139, 146, 148) Separate epidermal and/or dermal thickness measurements were reported in seven journal articles (17% of articles reporting skin thickness measurement type). (26, 47, 48, 52, 53, 71, 118) Of these records, two authors provided a rationale for this decision: each skin layer provided different information on the scar; (26) or responded differently to treatment. (67, 71) Most abstracts did not report the type of skin measurement used (n = 30; 83% of abstracts). (28, 30, 91-101, 103-110, 131-134, 150-154) Three articles (4% of articles) (47, 110, 111) and one abstract (3% of abstracts) (28) directly

reported that fibrosis was the scar characteristic targeted by the measurement. One of these records also quantified hair follicle density to assess the difference between scared and unscarred skin. (47) An additional 25 articles (30% of articles) (17, 46, 52, 53, 56, 63-65, 67, 70, 79, 80, 83, 84, 112, 120, 123, 125-127, 140, 142, 145, 148, 149, 155) and one abstract (3% of abstracts) (110) made indirect reference (i.e., within the introduction or discussion) to the measurement of fibrosis. Ten journal articles (12%) made indirect reference to the measurement of both oedema and fibrosis, (31, 54, 55, 71, 74, 76-78, 138, 144) and one record made indirect reference to the measurement of oedema. (59)

Additional objective and/or subjective measurement methods were employed alongside
ultrasound measurement in 72 articles (88% of articles) (17, 22, 24, 26, 29, 31, 46-53, 55-57,
60-70, 72-81, 83-85, 111-122, 124-130, 136-142, 144, 145, 147-149) and 31 abstracts (86%
of abstracts) (86, 88, 89, 91-95, 97-110, 131-134, 150, 151, 153, 154) (Supplementary Table
All three phase 4 studies involving implementation in routine clinical practice utilised

Page 17 of 83

BMJ Open

additional measurements. (92, 113, 119) The additional objective measurements used in included records were elastography (elasticity), cutometric assessment (pliability) and Doppler ultrasound (vascularity). The additional subjective measurements were conducted using clinician-based rating scales (e.g., Vancouver Scar Scale or modified Vancouver Scar Scale) or Patient Reported Outcome Measures. The Vancouver Scar Scale was used in 35 articles (43% of articles) (17, 31, 46, 47, 49, 50, 52, 55, 57, 61-64, 66-70, 73, 85, 111, 112, 114, 116, 118, 121, 124, 128, 130, 136-138, 140-142) and 11 abstracts (31% of abstracts) (88, 91, 92, 98-100, 107, 134, 150, 151, 153). Patient-reported outcome measures (PROMs) were used in 27 articles (33% of articles) and 11 abstracts (31% of abstracts). (46, 53, 56, 57, 60, 72-75, 85, 91, 94, 97, 101-106, 111, 112, 114, 115, 117, 118, 120, 122, 129, 131-133, 138, 140, 141, 148, 150, 151, 153, 154) Of the records that reported using PROMs, the most commonly used was the patient report of the Patient and Observer Scar Assessment Scale (POSAS), used in 17 articles (63% of articles reporting use of PROMs) (17, 22, 46, 50, 53, 61, 62, 64, 76, 77, 79, 114, 121, 125-127, 147) and 8 abstracts (73% of abstracts reporting use of PROMs) (91, 93, 102, 104, 106, 132, 153) (Supplementary Table 4). In most cases, additional measurement methods were used to supplement ultrasound thickness measurements as research outcomes. In some records (n = 16; 14% of records), however, ultrasound was compared with histology, POSAS, dermoscopy, VSS and modified VSS, clinical assessment, modified Seattle Scar Scale, high-definition optical coherence tomography, 3D camera, immunohistochemistry, and immunohistomorphometry. (17, 24, 26, 29, 31, 50, 51, 64, 73, 77, 86, 95, 110, 120, 124, 149) Where the effectiveness of ultrasound was judged against other methods, it was only found to be inadequate against histology. (26, 86)

Methods used to relocate the scar for repeated measurements were reported in 34 records
(29%) (Supplementary Table 3). The most common relocation method was tracing the outline

64	or boundaries of the scar on a transparent or translucent sheet ($n = 14$ articles; 35% of articles
65	reporting scar relocation), (23, 49, 65, 74, 81, 115, 116, 120, 124, 125, 153) occasionally
66	including prominent or bony landmarks close to the scar. (23, 24, 72, 73, 123) Photographs (n
67	= 10 articles; 25% of articles reporting relocation and $n = 1$ abstract) and linear measurements
68	from defined points or anatomical landmarks on or around the scar ($n = 4$ articles; 10% of
69	articles reporting relocation) were also used for scar relocation. The 'worst' or 'thickest' part
70	of the scar, as determined by patients or assessors, was chosen as the measurement site in 14
71	journal articles (35% of journal articles reporting relocation) (23, 31, 52, 54, 57, 61, 62, 67,
72	126, 127, 138, 141, 148, 155) and one abstract. (105)
73	Measurement of unscarred skin either contralateral or adjacent to the scar, was performed in
75	Wedstreinent of unsearred skin, entier contrancerar of adjacent to the sear, was performed in
74	32 articles (39% of articles%) (17, 22-24, 27, 29, 46-48, 50, 51, 53, 56-60, 64, 72, 73, 80, 81,
75	85, 114, 118, 120-122, 128, 145, 146, 148) and 7 abstracts (19% of abstracts) (28, 94, 95,
76	150, 151, 153, 154) These measurements were primarily used as controls or comparators to
77	scar measurements (n = 27, 69% of records reporting unscarred skin measurement). (17, 22,
78	23, 28, 29, 47, 48, 51, 53, 56-60, 64, 67, 73, 80, 85, 95, 118, 120, 122, 128, 146, 148, 153,
79	154) Additionally, four records (10% of records reporting unscarred skin measurement)
80	evaluating treatment efficacy measured both unaffected skin thickness and the thickness of a
81	'control' or untreated scar. (46, 74, 94, 114) All instances where additional ultrasound
82	measurements were taken of unscarred skin or untreated scars were reported as part of
83	research initiatives aligning with phases 2 and 3 of the research-to-practice pipeline (figure
84	1). (36)
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85 Reliability and measurement error

Reliability was calculated for both scarred and unscarred skin in 13 articles (16% of articles)
and two abstracts (5% of abstracts), and was generally considered acceptable (Supplementary

Page 19 of 83

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BMJ Open

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88	Table 5). This included inter-rater reliability ($n = 5$; 4% of articles), (54, 64, 73, 120, 137)
89	intra-rater reliability ($n = 3$; 4% of journal articles), (22, 23, 65) and both inter- and intra-rater
90	reliability (n = 7; 6%; including 2 abstracts) (17, 24, 57, 82, 87, 105, 124). The intraclass
91	correlation coefficient (ICC) was the most commonly reported reliability statistic ($n = 10$; 8%
92	of records, including one abstract), (17, 24, 57, 64, 65, 73, 82, 87, 120, 124) where it was
93	reported for both scar and unscarred skin measurements in four articles (5% of articles). (17,
94	24, 57, 73) The reported combined thickness (i.e., epidermal and dermal) ICCs for inter-rater
95	reliability of scarred skin ranged from 0.82 to 0.985, while the inter-rater ICC for the
96	measurement of unscarred skin ranged from 0.33 to 0.98, with one of the four records
97	reporting an ICC below the threshold value of 0.7 (ICC = 0.33) (24) and one record simply
98	reported that the inter-rater ICC for scarred skin was "acceptable to high". (64) The reported
99	intra-rater reliability for combined thickness measurements of scarred skin ranged from 0.89
100	to 0.983, and for unscarred skin ranged from 0.61 to 0.982, with one record reporting an ICC
101	below the threshold of 0.7 (ICC = 0.61). (24) One record reported both the inter- and intra-
102	rater ICCs for individual epidermal (inter-rater ICC = 0.297 ; intra-rater ICC = 0.809) and
103	dermal (inter-rater ICC = 0.991 ; intra-rater ICC = 0.991) scar thickness measurement. (87)
104	Four articles (5% of articles) reporting reliability used Pearson's R, an undisclosed method,
105	or description (e.g., high) as detailed in supplementary table 2. (22, 54, 105, 137)
106	Measurement error for inter-rater and intra-rater reliability of combined, epidermal or dermal
107	thickness was reported in four articles (5% of articles) and one abstract using standard error
108	of the measurement (SEM). The inter-rater SEM for the combined epidermal and dermal
109	thickness of scarred skin ranged from 0.11 mm to 0.5 mm, and the intra-rater SEMs ranged
110	from 0.18 to 0.52 mm. Individual records reported SEM values for unscarred skin, and
111	separate epidermal and dermal measurements, available in Supplementary Table 5. (17, 23,
112	24, 82, 87) Only one record reported calculation of the smallest detectable change (SDC). In

that record the inter-and intra-rater SDC was calculated for both scarred and unscarred skin.
The scarred skin SDCs were 1.4 mm (inter-rater) and 0.6 mm (intra-rater), and unscarred skin
SDCs were 0.8 mm (inter-rater) and 0.5 mm (intra-rater). (24) The reported SEMs were all
close to or below the largest SDC value reported. This finding may indicate that ultrasound
can detect true variance in scar thickness above measurement error for traumatic scar and skin
thickness.

Of the records that reported reliability and measurement error, measurements were taken by
practitioners with varying clinical expertise and roles within the treating team. These
included therapists, nurses and doctors, sometimes under the supervision of trained
radiologists. One record reported that 3 assessors received 3 hours of training, and conducted
10 assessments using the study protocol before the study began. (57)

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Clinical, health service, implementation and feasibility outcomes:

No record specifically investigated clinical, health service, implementation or feasibility outcomes of ultrasound as a measurement-based-care initiative. Ultrasound was used to assess the clinical outcomes of scar treatment initiatives in all included records. Clinical, health service, implementation and feasibility outcomes related to ultrasound measurement were, however, reported in 53 journal articles (17, 22-24, 26, 27, 31, 46-48, 50, 51, 54, 56-61, 63-66, 69-75, 77, 80, 82, 113-116, 119, 120, 122-124, 128, 129, 138, 142-144, 148, 149, 155) and 14 abstracts (28, 86, 87, 89, 90, 95, 96, 102, 105, 107, 109, 110, 152, 153) that focused on scar treatments.

The clinical outcome of patient satisfaction related to ultrasound measurement was only reported in one journal article. Whilst patient satisfaction was not directly measured in that record, a proxy measure of satisfaction was reported by the authors stating that no paediatric

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patient or their caregiver refused ultrasound measurement once the purpose was explained.(24)

Timeliness was the only reported health service outcome, reported as the time required to
take ultrasound measurements. Where reported in three journal articles, this was short, taking
between one to five minutes. (24, 27, 122)

The most common implementation outcomes reported in the identified records were fidelity, 141 acceptability and appropriateness. Fidelity to the measurement method was reported through 142 143 the use of experienced or trained assessors (n = 6 journal articles; n = 1 abstract), (24, 57, 58, 87, 142, 144, 148) and/or utilising the same assessor/s for all measurement sessions (n = 5) 144 journal articles; 6% of included journal articles). (24, 61, 138, 144, 148) Differences between 145 146 intended and actual measurement methods were not discussed. The training and/or experience of the assessors was discussed in 24 records (23 journal articles and 1 abstract), (17, 23, 24, 147 27, 51, 56-59, 63-66, 71, 73, 115, 116, 120, 123, 124, 138, 144, 149, 153) where 148 measurements were either taken by a clinician (n = 13; 54% of records reporting training), 149 (17, 23, 24, 58, 59, 64-67, 71, 120, 124, 141) members of the research team (n = 6; 25%) of 150 records reporting training), (57, 63, 73, 115, 123, 144) or by specialist sonographers and/or 151 radiologists (n = 5, including one abstract; 21% of records reporting training). (56, 116, 138, 152 149, 153) Only one record reported on fidelity in the context of routine clinical practice. In 153 154 this instance, ultrasound was conducted in the department of radiology, however the role or training of the staff was not reported. (119) 155

The acceptability and appropriateness of the ultrasound methods used in individual records
were generally based on author opinion and outlined in the discussion. Acceptability was
reported in 26 records (23 journal articles and 3 abstracts), (17, 22-24, 26-28, 31, 57, 64, 70,
74, 75, 77, 80, 82, 86, 96, 116, 119, 120, 122, 124, 143, 149, 155) including for paediatric

Page 22 of 83

BMJ Open

populations, where one record reported potential difficulty in measuring this population, (22) contrasting that which reported that measurement was acceptable to both children and their caregivers. (24) One record reported acceptability where the intervention being analysed by ultrasound was already part of routine clinical practice. In this instance, the authors referenced additional publications which stated that ultrasound had an accuracy of 0.5 mm, which was judged by the authors to be sufficient for assessment of scar thickness. (24, 27, 119, 122) Potential difficulty was identified in the measurement of open wounds, (24) and traditionally hard-to-reach areas (such as the axillae or groin). (22) The appropriateness of the ultrasound methods was reported in 35 journal articles (43% of included journal articles) (22, 24, 26, 27, 31, 46-48, 50, 54, 57, 60, 61, 64-66, 69, 72-75, 77,

80, 82, 113, 114, 116, 119, 120, 122, 124, 128, 148, 149, 155) and 11 abstracts (31% of included abstracts) (86, 87, 89, 90, 95, 102, 105, 107, 109, 110, 152), where it was generally addressed in the discussion. Of these records, two (4% of records reporting appropriateness) determined that ultrasound was not appropriate for scar measurement. The first stated that it was too inaccurate and complex; (86) and the second, which reported on initiatives within routine clinical practice, determined that the minimum resolution of the Diasonography ultrasonic scanner (Nuclear Enterprises, Edinburgh, UK) precluded its use in scars thinner than 3mm. (113)

The feasibility of ultrasound was reported in 12 journal articles (15% of included journal articles). (22, 24, 26, 46, 57, 70, 80, 119, 120, 124, 129) Five records considered ultrasound not feasible for scar measurements. The rationale presented included high-frequency 20 MHz ultrasound having an inadequate penetration depth; (26, 57) and ultrasound measurement and training of investigators requiring too much time (as reported in one record in phase 4 of the research-to-practice pipeline). (22, 119, 120) Another factor identified as precluding feasibility was the inability to consistently relocate the measurement site. (24) Conversely,

Page 21 of 39

Page 23 of 83

BMJ Open

one record reported ultrasound to be feasible in combination with Vancouver Scar Scale (VSS) measurement, (70) and another stated that ultrasound was able to distinguish between subcutaneous fat and muscle, which was interpreted by the authors of that record to mean that skin thickness measurements were accurate. (129) The majority (n = 11; 92%) of the records reporting feasibility were research initiatives in phase 2 or 3 of the research to practice pipeline. One record examined feasibility in the context of routine clinical practice (i.e., phase 4; figure 1), (119) where it was determined that ultrasound was not suitable for use in their twelve-year longitudinal study due to changes in staff, equipment and software over such a long time period, which introduced additional variables to the measurement process that were impossible to control. (119)

195 Factors influencing ultrasound images and measurement methods:

The only factor that was reported to influence the imaging and measurement methods was the measurement of scars with open wounds. This was reported in one record, which determined that ultrasound and ultrasound gel was unsuitable in this instance. The authors of that record suggested the use of a flexible transparent plastic wrap, which is placed over the measurement area prior to measurement with ultrasound. (24)

201 Reported strengths and limitations of the measurement methods:

The safety, practicality, objectivity, versatility, reliability and non-invasive nature of
ultrasound were all reported as strengths of the measurement method. (22, 27-29, 47, 50, 57,
61, 64, 77, 78, 80, 82, 87, 89, 95, 96, 105, 107, 109, 119, 123, 124, 129, 139, 148) When
compared to other subjective or clinical measurement methods (e.g., VSS) and 3D camera,
ultrasound was viewed as the superior measurement method of scar and skin thickness, due to
its improved accuracy, greater sensitivity to change and objectivity. (24, 64, 73, 116, 120)
The ability of ultrasound to differentiate between scarred and unscarred skin was also

highlighted (n = 4; 3%), (47, 60, 72, 122) as was the versatility of ultrasound in its ability to measure a variety of anatomical areas and be used with child participants (i.e., <18 years) (n = 2; 2%). (22, 149)

The poor correlation between ultrasound and histological thickness measurements, (86) and the established inverse relationship between ultrasound penetration depth and the resolution of superficial structures were identified as limitations of ultrasound in the measurement of scar thickness. (26, 27, 77, 80, 89, 113, 149) This may be an evidence gap worth exploring in more depth. One record, reporting on a longitudinal study that was conducted over twelve years, reported that the continuous development of ultrasound software and hardware over that time limited the usefulness of ultrasound. (119) Despite being reported elsewhere as acceptable (i.e., between one to five minutes (24, 27, 122)), one record reported that the timeconsuming nature of measurement and the requirement for assessors to be trained in the operation of, and techniques required for, ultrasonography was a limitation of the method. (120) Methodologically, concerns were raised around the pressure caused by application of the ultrasound transducer to the skin, and how that may influence thickness measurement. (61, 62, 123, 124) The size of the transducer head relative to the size of scars was also considered a potential limitation, as multiple measurements are required for quantification of larger scars. (57) Finally, it was recognised that there may be a difference between changes to the scar that can be measured by ultrasound, and what is felt and/or experienced by the patient. (75, 80, 126, 127) It was suggested that changes that are detectable by ultrasound may be smaller than those able to be detected by patients. In patients with burn scars, a minimum change in scar thickness of between 1 to 6 mm measured by ultrasound, has been reported to be required before a patient may report noticing any difference to their scar thickness. (24, 75) While further research is required to allow generalisation of these findings

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to other scar aetiologies, this indicates that a holistic approach to scar thickness using the

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50 51 52	252
53 54	253
55 56	254
57 58	255
59 60	256

patient's opinion as well as objective measurement through ultrasound may be beneficial.
Guidelines or frameworks used to guide the measurement methods:
No records reported using any guidelines or frameworks to inform their measurement
methods. One record utilised suggestions from The American Wound Healing Society to
support the measurement of contralateral, unscarred skin thickness on the same individual as
a control or comparator. (75)
Methodological Considerations:
Based on the ultrasound methods and outcomes identified in this review, a list of

methodological considerations have been compiled (Supplementary Table 6). These are
intended to guide the decision-making and methodological reporting of researchers and/or
clinicians undertaking scar or skin thickness ultrasound measurement.

245 **DISCUSSION:**

6 This review mapped the methods used in the published literature to measure traumatic scar 7 thickness using ultrasound across the research-to-practice translational pipeline. No record 8 reported their methods with sufficient detail to allow them to be independently replicated. 9 Overall, there was a lack of consistent rationale underpinning which skin layers (i.e., 0 epidermis, dermis and combined) were measured, and little consideration was given to the training and experience required by assessors. The included records mainly aligned with the 1 second and third phases of the research-to-practice pipeline (figure 1), with only three records 2 3 (2 articles and 1 abstract) reporting the use of ultrasound in routine clinical practice (phase 4). 4 (92, 113, 119). The paucity of records aligning with phase four studies (use in clinical 5 practice) suggests a translational gap from research to regular clinical practice. There are two likely explanations for this: 1) that ultrasound is most commonly used as an outcome measure 256

for research initiatives and is not regularly used to evaluate care once treatments are implemented into routine clinical practice; or 2) that use of ultrasound in routine clinical practice is not reported or evaluated, as routine clinical practice is rarely published. Searching of grey literature was conducted in an attempt to identify clinical practice documents, however none were located. Surveys of health service departments may be the best method of identifying ultrasound methods used in regular clinical practice as part of future research. While some records reported using additional subjective and objective measurement methods in addition to ultrasound, none used these methods to determine the criterion validity of the ultrasound for scar thickness measurement. This is another evidence gap that should be addressed. While efforts have been made to standardise ultrasound measurement procedures elsewhere in dermatology (including tumours, cancers, vascular anomalies, and systemic sclerosis (34, 35)), this same effort has not yet extended to the measurement of traumatic scarring. Methodological standardisation has the potential to increase confidence in the use of ultrasound as the basis of measurement-based care initiatives for clinical decision-making, allowing patient care and scar treatments to be tailored towards individual needs. (62, 147, 156) Standardising the core methodological components of ultrasound measurement of scar thickness, or at the very least, creating a standardised framework for methodological decision-making, may support implementation of ultrasound measurement into routine clinical practice, supported by strategies to overcome barriers to implementation at local sites. (157)This review identified novel insights into the identification of the composition of cutaneous scars using ultrasound, and highlighted the apparent lack of consistent understanding of, or

rationale behind, what scar thickness characteristics were being measured. Fibrosis is

Page 25 of 39

Page 27 of 83

BMJ Open

generally understood to be the primary cause of scar thickness through the deposition of excessive extracellular matrix proteins such as collagen. (158, 159) This has been confirmed through histological analysis, which has shown the presence of excess collagen and other extracellular matrix proteins in the dermis of hypertrophic scars. (160, 161) An additional method for assessing the effects of scarring on the dermis, as identified by one record in this review, (47) is through quantification of the presence and density of hair follicles. This quantification may serve as a method of differentiation between scarred and physiological skin, and may also serve as a measure of skin function. (47) What is less understood, and perhaps largely overlooked, is the function of the epidermis in scar thickness. In the one record identified in this review that directly report the measurement of the epidermis, the authors noted that the measurement quantified the presence of oedema. (55) This was further supported by two records that noted that the epidermis and dermis responded differently to treatment, (67, 71) indicating that there is likely a difference in the composition of the scar between these skin layers. Cutaneous oedema has been observed using high-frequency ultrasound in other pathologies, including atopic dermatitis and skin ageing, where it is characterised by the presence of a sub-epidermal low echogenic band (SLEB), a hyperechoic band at the dermoepidermal junction. (162) Understanding the interplay between epidermal oedema, dermal fibrosis and the presence and density of hair follicles may result in an increased understanding of the mechanisms and treatment responses of cutaneous scarring. With better understanding, more targeted scar treatments that inform a greater understanding of scar responsivity may arise.

Another important, but potential limiting factor for the use of ultrasound to measure scar thickness raised in this review is the training and/or experience required of assessors, and the ramifications this likely has on the reliability of measurements and interpretation. (163) This review identified 24 records where assessor experience was discussed, however none made
any recommendations on the optimal training and/or experience. Identifying the training requirements of assessors may prove an important step towards more widespread implementation of reliable ultrasound scar thickness measurement in research trials and as the basis for measurement-based care in routine clinical practice. (164) A panel of dermatological and ultrasound experts has previously recommended that a physician with a minimum of 300 examinations per year should hold responsibility for ultrasound measurements. (34) It has also been suggested that training existing members of clinical teams and standardising measurement method/s may be the most effective way to achieve minimum reliability standards under clinical conditions. This could allow measurement to be reliably conducted within an outpatient clinic setting by a number of healthcare providers assisting workflow, negating the requirement for patients to wait for an experienced radiographer. (24, 164) In the current review, reliability estimates were generally acceptable but were tested under research conditions. The diverse experience and expertise of assessors, where reported for the reliability estimates, means that the acceptable reliability results should be generalisable to most clinical teams, as therapists, doctors and nurses were all included. The cumulative sample size of all reliability studies also supports this generalisation; however each team should perform their own reliability estimates before conducting ultrasound thickness measurements.

324 Study Limitations:

Only articles available in English or with an English abstract were considered for inclusion
and data extraction, which may have resulted in the omission of eligible information. Data
extraction was completed on the English abstracts of two non-English articles that were
available electronically, however the non-English articles themselves were not available to
the authors, and thus could not be analysed. Based on the number of records included in this
review, however, it is unlikely that this would have impacted the review findings. It is

Page 27 of 39

Page 29 of 83

BMJ Open

acknowledged that methods reported in included abstracts may not be fully reproducible, due to their brevity. Thus, findings were reported separately to articles. An additional limitation was that authors of included records were not contacted to provide clarification or further information, as this was not feasible given the number of results identified. It should also be acknowledged that the included records were not designed to align with the specific aims of this review, which likely explains some of the lack of reporting on outcomes of interest in our review, particularly clinical, health service and implementation outcomes. Furthermore, as this review relied on published information (including grey literature), routine practices employed within organisations may not have been considered and unpublished industry sponsored reports may not have been identified. It is also important to consider the limitations of ultrasound itself for the holistic quantification of cutaneous scarring. Ultrasound transducers are generally small, meaning that it is difficult to assess the entirety of a scar, necessitating multiple measurements. (165) Additionally, thickness is often not the only scar parameter of clinical or research interest. It has therefore been recommended that multi-modal measurement techniques are employed, which include both subjective and objective measurements. (166, 167) However, use of these methods may be challenging in routine clinical practice, due to the length of time and training required. Thus, feasibility and implementation outcomes are of importance in evaluating measurement-based care initiatives involving ultrasound alone or multimodal measurement tools in scar care practice – a field in its infancy based on this review. **Future Directions:**

352 It is intended that the results of this review will be used to inform the creation of a Delphi
353 consensus study, leading to the formation of a guideline for the measurement of traumatic
354 scar thickness using ultrasound. This guideline can then be used by researchers and clinicians

to standardise the measurement of scars. In preparation for this study, we have provided a list of methodological considerations for assessors or practitioners when planning to conduct scar thickness measurements with ultrasound (Supplementary Table 6). Future research could also investigate aspects that were beyond the scope of this review including factors influencing the implementation of ultrasound-based care initiatives, strategies to support implementation, and how research-based initiatives could be applied in practice. Further studies are needed that compare SDCs to SEMs to interpret reliability estimates to confirm our interpretation that ultrasound may have the ability to detect true change or variance in scar thickness above measurement error, which was based on the SDC reported by a single study. Our interpretation is supported by mostly acceptable reliability estimates of ultrasound thickness for other cutaneous conditions. (168, 169) Additional investigations should also be conducted to determine the criterion validity of ultrasound as a measure for scar thickness.

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371 COMPETING INTERESTS:

The authors declare no competing interests. The research presented in this publication was conducted as part of BM's PhD, and will be included in his thesis for submission to The University of Queensland.

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Page 29 of 39

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379	AUTHOR CONTRIBUTIONS
380	BM and ZT conceived the project after identifying this area as a knowledge gap in existing
381	literature. BM developed the research questions and study methodology, conducted the
382	literature search, screened all articles and extracted data. Record screening and data
383	extraction was completed by BM, MS, TM, and TR, with additional extraction completed by
384	BD to assess consistency. MS, TM, TR and RK provided advice to BM on the clinical
385	implications of ultrasound measurement. MS, RK and ZT contributed to the supervision of
386	BM as a PhD student. BM drafted the paper, and ZT and MS provided critical appraisal of
387	the drafted manuscript, with further advice provided by TM, TR, BD and RK.
388	DATA SHARING STATEMENT:
389	Not applicable
390	ETHICS APPROVAL STATEMENT:
391	This study does not involve human participants. No ethics approval was required.
392	FIGURE LEGENDS:
393	Figure 1: Research to clinical practice pipeline.
394	

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Supplementary Box 1. Full search strategy for Ovid MEDLINE.

((ultrasound.ti,ab. OR ultra sound.ti,ab. OR sonograph*.ti,ab. OR ultrasonic.ti,ab. OR highfrequency.ti,ab. OR high frequency.ti,ab. OR hfus.ti,ab. OR ultrasonog*.ti,ab. OR exp Ultrasonography/)

AND

((skin.ti,ab. OR epiderm*.ti,ab. OR derm*.ti,ab. OR cutaneous.ti,ab OR scar*.ti,ab OR keloid*.ti,ab OR cicatri*.ti,ab OR exp Skin/ OR exp Dermatology/ OR exp Cicatrix/)

AND

(thickness*.ti,ab. OR thicken*.ti,ab. OR depth.ti,ab. OR volume.ti,ab. OR height.ti,ab. OR vancouver scar scale.ti,ab)

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(measure*.ti,ab. OR quantif*.ti,ab. OR calculat*.ti,ab OR estimat*.ti,ab OR assess*.ti,ab. OR determin*.ti,ab. OR evaluat*.ti,ab OR imag*.ti,ab OR exam*.ti,ab)))

NOT (exp animals/ NOT exp humans/)

Legend: ab, abstract (searches the abstract of the publication); adj10, adjacency (search terms must be located within 10 words of one another); exp, explode (used to include all subheadings when searching MeSH headings); ti, title (searches the title of the publication)



BMJ Open

BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

Supplementary Figure 1. Preferred Reporting Items for Systematic reviews and Meta-

Analyses (PRISMA) flow diagram for this study.



BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

Supplementary Table 1: Extraction categories and fields

Extraction category	Extraction field			
Publication details	First author			
	Year of publication			
	Title of publication			
	Country (first author)			
	Country (study)			
	Country (recruited)			
	Publication type (e.g., peer-reviewed journal article, abstract)			
	Journal name			
	Corresponding author contact details			
	Funding source (e.g., commercial, non-commercial)			
	Use of scar thickness measurement (e.g., longitudinal study, response to			
	treatment)			
Study details	Aim/objective			
	Research questions			
	Target population/topics			
	Study design (e.g., RCT, mixed methods)			
	Data and analysis (i.e., statistical methods)			
	Removal of scar treatments before ultrasound measurement (e.g., length of			
	time before measurement)			
	Reason for measurement (e.g., research, clinical initiative)			
	Inclusion/exclusion criteria			
	Dates of data collection			
	Ultrasound thickness collection methods (e.g., direct collection, collected			
	from medical records)			
	Contralateral/unaffected/comparator skin thickness measurement			
	Other methods used			
	Use of guidelines/frameworks for measurement methods			
	How previously published methods/guidelines were used			
	Research pipeline stage			
	Setting (e.g., inpatient/outpatient clinics)			
	Scar type (e.g., burn scar, surgical scar)			
Participant details	Number of participants			
	Population type (e.g., adult/paediatric)			
	Gender ratio			
	Patient involvement in thickness determination			
	How patients were involved in thickness determination			
Ultrasound methods	Ultrasound mode			
	Device name and manufacturer			
	Frequency used			
	Number of measurements taken			
	What did researchers report they were measuring (e.g. fibrosis ordema)			
	Anatomical locations/functional measurement units measured			
	Patient orientation			
	Illtrasound transducer orientation			
	Methods used to prevent skin compression			
	Measurement site relocation strategies			
	Type of skin measurement (i.e. enidermis/dermis/combined)			
	Measurer training			
Psychometric properties*	Reliability			
r sychometric properties	Measurement error			
Feasibility [†] outcomes	Time taken for measurement			

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3		Availability of measurement method
4		Ease of administration
5		Number of steps required
6		Number of people required to conduct measurements
7		Considerations for special populations
8	Implementation [‡] outcomes	Accentability
9	implementation outcomes	Adoption
10		Appropriateness
11		Cost
12		Eost
13		Fidality
14		Fluenty
15	Strongthe and limitations of	Sustainability
16	Strengths and minitations of	
17	measurement methods	
18		Barriers
19		Enablers
20	Findings	Ultrasound-related findings
21	[*] Psychometric properties as outl	ined in the COSMIN Risk of Bias tool to assess the quality of studies on
22	reliability or measurement error	of outcome measurement instruments ¹
23	Feasibility outcomes as per Prin	nsen <i>et al.</i> ²
24	[‡] Implementation outcomes as pe	er Proctor <i>et al.</i> ³
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Supplementary Table 2. Characteristics of records included in this review. Studies are listed alphabetically by author within the translational
pipeline phase.

First Author (year)	Country of Publication	Funding Sources	Sample Size (n)	Population Type	Scar Aetiology	Translational Pipeline Phase [*]
Journal articles						
Agabalyan (2017)	Canada	Non-commercial	10	Adult	Not specified	2
Alsharnoubi (2018)	Egypt	No funding	15	Paediatric	Burn	2
Alsharnoubi (2018)	Egypt	Not reported	15	Paediatric	Burn	2
Alshehari (2015)	Egypt	Not reported	30	Not reported	Mixed	2
Blome-Eberwein (2012)	United States	Non-commercial	16	Paediatric & adult	Burn	2
Blome-Eberwein (2016)	United States	Not reported	36	Adult	Not specified	2
Blome-Eberwein (2019)	United States	Non-commercial	19	Adult	Burn	2
Cai (2019)	China	Non-commercial	51	Adult	Not specified	2
Candy (2010)	Hong Kong	Not reported	17	Adult	Not specified	2
Chan (2004)	China	Non-commercial	56	Paediatric & adult	Burn	2
Chang (2014)	Taiwan	Non-commercial	60	Paediatric & adult	Surgical (cleft	2
-					lip repair)	
Cho (2014)	Korea	Non-commercial	146	Not reported	Burn	2
Deng (2019)	China	Not reported	20	Adult	Not specified	2
Deng (2021)	China	No funding	31	Adult	Not specified	2
Deng (2021)	Hong Kong and China	Non-commercial	45	Adult	Not specified	2
Dunkin (2007)	England	Non-commercial	113	Adult	Surgical (dermal	2
					scratch)	
Elrefaie (2020)	Not specified	Not reported	22	Paediatric & adult	Not specified	2
Fabbrocini (2016)	Not specified	Not reported	20	Adult	Mixed	2
Fraccalvieri (2011)	Italy	No funding	5	Adult	Mixed	2
Fraccalvieri (2013)	Italy	Not reported	3	Paediatric & adult	Mixed	2
Gee Kee (2016)	Australia	Commercial	43	Paediatric	Burn	2
Issler-Fisher (2021)	Australia	Commercial	187	Adult	Burn	2
Joo (2020)	Korea	Non-commercial	48	Adult	Not specified	2
Lacarrubba (2008)	Not specified	Not reported	8	Paediatric & adult	Mixed	2

BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

I (2005)	11 17		100		P
Lau (2005)	Hong Kong	Not reported	100	Paediatric & adult	Burn
Lee (2019)	United Kingdom	Non-commercial	55	Adult	Burn
Lee (2020)	United Kingdom	Non-commercial	55	Adult	Burn
Li (2013)	China	Non-commercial	7	Adult	Burn
Li (2020)	China	Not reported	21	Paediatric & adult	Mixed
Li (2021)	China	Non-commercial	165	Paediatric	Mixed
Li (2021)	China	Non-commercial	105	Adult	Burn
Li-Tsang (2006)	Not specified	Non-commercial	45	Adult	Not specified
Li-Tsang (2010)	China	Non-commercial	104	Paediatric & adult	Mixed
Mamdouh (2021)	Egypt	Not reported	40	Adult	Not specified
Meirte (2016)	Belgium	Non-commercial	9	Adult	Burn
Miletta (2021)	United States	Non-commercial	29	Paediatric & adult	Burn
Nedelec (2019)	Canada	Non-commercial	70	Adult	Burn
Nedelec (2020)	Canada	Non-commercial	51	Adult	Burn
Nicoletti (2015)	Italy	Not reported	27	Paediatric & adult	Surgical (scar
	-				reconstruction)
Niessen (1998)	The Netherlands	Commercial & Non-	145	Paediatric & adult	Surgical (breast
		commercial			reduction)
Reinholz (2020)	Germany	No funding	25	Adult	Mixed
Schwaiger (2018)	Germany	No funding	15	Adult	Mixed
van den Kerckhove	Belgium	Not reported	60	Adult	Burn
(2005)	C	1			
van der Veer (2010)	The Netherlands	Non-commercial	44	Adult	Surgical
					(cardiothoracic
					surgery)
Wang (2009)	China	Non-commercial	22	Adult	Burn
Wiseman (2020, 2021)	Australia	Commercial & Non-	153	Paediatric	Burn
		commercial			
Xuan (2021)	Not specified	Not reported	72	Not reported	Not specified
Yim (2010)	Korea	No funding	31	Paediatric & adult	Burn
Zadkowski (2016)	Not specified	Not reported	47	Paediatric	Burn
Avetikov (2018)	Not specified	Not reported	50	Paediatric & adult	Not specified
	riet speemed	rior reported	50		rest op control

BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

			• •		
Chae (2016)	Korea	Non-commercial	23	Adult	Not specified
Cheng (2001)	Hong Kong	Not reported	58	Paediatric	Burn
Danin (2012)	France	Not reported	22	Paediatric & adult	Burn
Fong (1997)	Not specified	Not reported	16	Paediatric & adult	Burn
Gankande (2014)	Australia	Non-commercial	30	Adult	Burn
Ge (2022)	China	Not reported	21	Paediatric & adult	Mixed
Guo (2020)	China	Non-commercial	87	Paediatric & adult	Not specified
Huang (2017)	Taiwan	Not reported	1	Adult	Burn
Huang (2020)	China	Non-commercial	43	Adult	Not specified
Huang (2021)	Taiwan	Not reported	5	Adult	Burn
Issler-Fisher (2017)	Australia	No funding	47	Paediatric & adult	Burn
Issler-Fisher (2020)	Australia	No funding	78	Adult	Burn
Katz (1985)	United States	Not reported	4	Not reported	Burn
Kemp Bohan (2021)	United States	No funding	21	Not reported	Burn
Kim (2018)	Not specified	Not reported	148	Not reported	Burn
Li (2018)	China	Non-commercial	34	Adult	Burn
Li-Tsang (2005)	China	Non-commercial	101	Adult	Surgical
					(orthopaedic
					surgery)
Lobos (2017)	Not specified	Not reported	35	Paediatric & adult	Not specified
Nedelec (2008)	Canada	Non-commercial	32	Adult	Burn
Nedelec (2014)	Not specified	Non-commercial	46	Adult	Burn
Reinholz (2016)	Not specified	Commercial	8	Adult	Not specified
Simons (2017)	Australia	Non-commercial	49	Paediatric	Burn
Soykan (2014)	The Netherlands	Non-commercial	87	Adult	Surgical
					(cardiothoracic
					surgerv)
Timar-Banu (2011)	Canada	Non-commercial	30	Adult	Mixed
Ud-Din (2019)	United Kingdom	Non-commercial	62	Adult	Not specified
van den Kerckhove	Not specified	Not reported	6	Adult	Burn
(2003)	r		Ũ		

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3 4	Wang (2010)	Australia	Commercial & Non- commercial	21	Paediatric	Burn	3
5	Wood (1996)	Not specified	Not reported	1	Paediatric	Burn	3
0 7	Yeol Lee (2022)	Korea	Non-commercial	16	Adult	Mixed	3
, 8	Berry (1985)	Not specified	Commercial	16	Paediatric & adult	Burn	4
9	Engrav (2010)	Not specified	Commercial & Non-	67	Paediatric & adult	Burn	4
10	g (· · ·)		commercial				-
11	Abstracts						
12	Agabalyan (2016)	Not specified	Non-commercial	10	Not reported	Burn	2
13	Bajouri (2018)	Not specified	Not reported	20	Not reported	Burn	2
15	Blome-Eberwein (2011,	Not specified	Not reported	16	Paediatric & adult	Mixed	2
16	2012)		1				
17	Blome-Eberwein (2014)	Not specified	Not reported	66	Not reported	Burn	2
18	Cho (2012)	Not specified	Not reported	60	Paediatric & adult	Burn	2
19	Comstock (2018)	Not specified	Not reported	1	Adult	Burn	2
20 21	Cooper (2021)	Not specified	Not reported	25	Not reported	Burn	2
22	El-Zawhary (2007)	Not specified	Not reported	57	Not reported	Mixed	2
23	Jacobs (2016)	Not specified	Not reported	6	Paediatric & adult	Burn	2
24	Jang (2009)	Not specified	Not reported	20	Not reported	Not specified	2
25	Kim (2009)	Not specified	Not reported	5	Paediatric & adult	Burn	2
26	Li-Tsang (2010)	Not specified	Not reported	45	Not reported	Not specified	2
2/	Li-Tsang (2011)	Not specified	Not reported	4	Not reported	Not specified	2
20 29	Maari (2017)	Not specified	Non-commercial	12	Not reported	Not specified	2
30	Moortgat (2020)	Not specified	Not reported	10	Not reported	Burn	2
31	Nedelec (2018)	Not specified	Not reported	60	Not reported	Burn	2
32	Peters (2018)	Not specified	Not reported	5	Not reported	Burn	2
33	Siwy (2016)	Not specified	Non-commercial	15	Not reported	Burn	2
34	Tu (2014)	Not specified	Not reported	59	Not reported	Not specified	2
35 36	Ud-Din (2017)	Not specified	Not reported	20	Not reported	Surgical (tissue	2
37		L	1			biopsies)	
38	Anthonissen (2015)	Not specified	Not reported	N.R.	Not reported	Burn	3
39	Bezugly (2014)	Not specified	Not reported	103	Not reported	Mixed	3
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BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

Bezugly (2019)	Not specified	Not reported	438	Not reported	Not specified	3
Blome-Eberwein (2012)	Not specified	Not reported	19	Adult	Burn	3
Du (2006)	Not specified	Not reported	1	Adult	Burn	3
Edgear-Lacoursière	Canada	Not reported	44	Not reported	Burn	3
(2022)		-		-		
George (2019)	Not specified	Not reported	11	Not reported	Burn	3
Li (2016)	Not specified	Not reported	34	Not reported	Burn	3
Seo (2011)	Korea	Not reported	48	Not reported	Burn	3
Timina (2013)	Not specified	Not reported	49	Paediatric & adult	Not specified	3
Ud-Din (2017)	Not specified	Not reported	20	Not reported	Surgical (tissue	3
		-		-	biopsies)	
Ud-Din (2018)	Not specified	Not reported	62	Not reported	Surgical (tissue	3
		-		-	biopsies)	
Zuccaro (2019)	Canada	Not reported	13	Paediatric	Burn	3
Zuccaro (2021)	Not specified	Not reported	20	Paediatric	Burn	3
Zuccaro (2021)	Canada	Non-commercial	20	Paediatric	Burn	3
Cho (2012)	Not specified	Not reported	30	Not reported	Burn	4

Legend: Paediatric: measurement of patients under the age of 18; Adult: measurement of patients aged 18 years or older; N.R.: Not reported; Burn: scars caused by thermal, chemical or friction injury; Surgical: scars caused by surgical procedures (including biopsies); Mixed: participant scars caused by mixed trauma (e.g., burn and acne)

Footnotes: *Stage in the research to clinical practice translational pipeline, based on the Australian Government Department of Health and Aged Care⁴

Page 51 of 83

BMJ Open

First Author (year)	Ultrasound Type	Ultrasound Frequency (MHz)	Measurement Parameters	Scar Characteristic Measured	Scar Relocation
Journal articles					
Agabalyan (2017)	High-frequency	20	Epidermal, dermal & combined	N.R.	Not relevant – single measurement
Alsharnoubi (2018)	Midrange ultrasound	N.R.	N.R.	Fibrosis	N.R.
Alsharnoubi (2018)	Midrange ultrasound	N.R.	N.R.	Fibrosis [†]	N.R.
Alshehari (2015)	N.R.	N.R.	Maximum elevation above normal skin	N.R.	N.R.
Avetikov (2018)	B-mode	N.R.	Combined epidermal & dermal	N.R.	Not relevant – single measurement
Berry (1985)	N.R.	N.R.	N.R.	N.R.	N.R. [‡]
Blome- Eberwein (2012)	B-mode	N.R.	Combined epidermal & dermal [§]	N.R.	N.R. [‡]
Blome- Eberwein (2016)	High-frequency	50	N.R.	Fibrosis [†]	N.R. [‡]
Blome- Eberwein (2019)	High-frequency	35	Dermal	Fibrosis, hair follicle density	N.R.
Cai (2019)	High-frequency	50	Dermal	N.R.	N.R. [‡]
Candy (2010)	B-mode	N.R.	N.R.	N.R.	Scar boundaries traced
Chae (2016)	N.R.	N.R	Combined epidermal & dermal	N.R.	Not relevant – single measurement
Chang (2014)	N.R.	12	N.R.	N.R.	N.R.

Chan (2004)	N.R.	N.R.	N.R.	N.R.	Tracing
Cheng (2001)	B-mode	5-10	Combined epidermal & dermal	N.R.	Tracing & cutting out paper Photographs
Cho (2014)	High-frequency	7.5	N.R.	N.R.	N.R.
Danin (2012)	B-mode	20	Epidermal & dermal	N.R.	N.R.
Deng (2019)	N.R.	N.R.	N.R.	N.R.	N.R.
Deng (2021)	Colour Doppler	4-15	Dermal	Fibrosis [†]	N.R.
Deng (2021)	B-mode	8-12	Epidermal & dermal	Fibrosis [†]	Photographs
Dunkin (2007)	High-frequency	N.R.	N.R.	Fibrosis & oedema [†]	Measurements taken at set linear distances along scar
Elrefaie (2020)	High-frequency	13	N.R.	Fibrosis & oedema [†]	N.R [‡]
Engrav (2010)	N.R.	N.R.	N.R.	N.R.	N.R.
Fabbrocini (2016)	N.R.	N.R.	N.R.	Fibrosis & oedema [†]	N.R [‡]
Fong (1997)	B-mode	7.5	N.R.	Fibrosis [†]	Tracing
Fraccalvieri (2013)	High-frequency	7-10 & 10-13	N.R.	Fibrosis & oedema [†]	N.R.
Fraccalvieri (2011)	High-frequency	10-13	Combined epidermal & dermal	Fibrosis [†]	N.R.
Gankande (2014)	High-frequency	20	Combined epidermal & dermal	N.R.	Scar marked & photographed
Ge (2022)	N.R.	N.R.	N.R.	N.R.	N.R.
Gee Kee (2016)	B-mode	8-18	Combined epidermal & dermal	N.R.	Transducer in centre of original burn site where no scar present
Guo (2020)	N.R.	2-15 & 4-15	Combined epidermal & dermal ^c	Fibrosis [†]	Thickest site on peripheral regions
Huang (2017)	N.R.	N.R.	Combined epidermal & dermal	N.R.	Marked & linear measurements from bony landmarks

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Huang (2021)	B-mode	5-12	N.R.	Oedema [†]	Not relevant – single
Huang (2020)	B-mode	5-12	Combined epidermal & dermal	N.R.	N.R.
Issler-Fisher (2021)	N.R.	N.R.	N.R.	N.R.	Photograph & measurement of thickest area
Issler-Fisher (2020)	N.R.	N.R.	N.R.	N.R.	N.R.
Issler-Fisher (2017)	N.R.	N.R.	N.R.	Fibrosis [†]	Scar mapped with drawing Thickest area measured
Joo (2020)	N.R.	N.R.	N.R.	Fibrosis [†]	N.R.
Katz (1985)	B-mode	10	Combined epidermal & dermal	N.R.	N.R.
Kemp Bohan (2021)	High-frequency	12	N.R.	Fibrosis [†]	Tracing – thickest area & adjacent landmarks marked
Kim (2018)	N.R.	22	Combined epidermal & dermal	N.R.	Not relevant – single measurement
Lacarrubba (2008)	B-mode	20	Combined epidermal & dermal	N.R.	N.R.
Lau (2005)	Tissue Ultrasound Palpation System	5 (burn) & 10 (surgical)	N.R.	N.R.	Tracing – most severe/prominent site
Lee (2020)	High-frequency	20	Combined epidermal & dermal	Fibrosis [†]	Not relevant – single measurement
Lee (2019)	High-frequency	20	Combined epidermal & dermal	Fibrosis [†]	Marked with pen
Li (2013)	High-frequency	12	Combined epidermal & dermal	Fibrosis [†]	Tracing
Li (2020)	N.R.	10	N.R.	Fibrosis [†]	N.R.
Li (2021)	High-frequency	20	N.R.	N.R.	Thickest area
Li (2021)	High-frequency	20	N.R. [§]	Fibrosis [†]	Thickest area
Li (2018)	N.R.	N.R.	Combined epidermal & dermal	N.R.	N.R.
Li-Tsang (2005)	Tissue Ultrasound Palpation System	N.R.	N.R.	N.R.	N.R.
Li-Tsang (2006)	B-mode	N.R.	N.R.	N.R.	N.R [‡]

Li-Tsang	B-mode	N.R.	N.R.	Fibrosis [†]	N.R.
(2010) Lobos (2017)	B-mode & colour	18	N.R.	Fibrosis [†]	Not relevant – single measurement
Mamdouh (2021)	High-frequency	N.R.	Combined epidermal & dermal [§]	Fibrosis [†]	N.R.
Meirte (2016)	High-frequency	22	Dermal	Fibrosis & oedema [†]	Marked with surgical pen, including boundaries of probe. Photograph of body position & probe location
Miletta (2021)	N.R.	50	N.R.	Fibrosis [†]	Tracing – worst scar
Nedelec (2014)	High-frequency	20	Combined epidermal & dermal	N.R.	Tracing including notable landmarks. Measurement site circled. Photograph
Nedelec (2008)	High-frequency	20	Combined epidermal & dermal	N.R.	Tracing including notable landmarks. Measurement site circled. Photograph
Nedelec (2019)	High-frequency	20	Combined epidermal & dermal	Fibrosis & oedema [†]	Tracing. Hole cut over measurement area
Nedelec (2020)	High-frequency	20	Combined epidermal & dermal	N.R.	Photograph
Nicoletti (2015)	N.R.	22	Epidermis to fascia	N.R.	N.R.
Niessen (1998)	B-mode	N.R.	N.R.	Fibrosis & oedema [†]	3cm border marked with tape – measurements lateral
Reinholz (2020)	B-mode	11	Combined epidermal & dermal	Fibrosis & oedema [†]	N.R.
Reinholz (2016)	B-mode	11	Combined epidermal & dermal [§]	Fibrosis & oedema [†]	N.R.
Schwaiger (2018)	B-mode	11	N.R.	Fibrosis & oedema [†]	N.R.
Simons (2017)	B-mode	8-18	Combined epidermal & dermal	N.R.	Tracing – scar & anatomical landmarks

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BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

2						
3	Soykan (2014)	N.R.	3-9	N.R.	Fibrosis [†]	N.R.
4	Timar-Banu	High-frequency	20	Combined epidermal & dermal	Fibrosis [†]	N.R.
5	(2001)		_0		11010015	
6	(2001) Ud_Din (2019)	High-frequency	50	Combined enidermal & dermal	Fibrosis	Defined anatomical location
/	von den	Ligh frequency	20	Combined epidermal & dermal	N D	Test sites marked
0	Vall Util	Ingli-frequency	20	Combined epidermar & dermar	1 \.I\.	The state of the s
9 10	(2002)					Thermoplastic spinits created
10	(2003)	TT: 1 C	20		ND	with space for transducer
12	van den	High-frequency	20	Combined epidermal & dermal	N.K.	Test site boundaries marked
13	Kerckhove					& traced
14	(2005)					
15	van der Veer	N.R.	7.5	N.R.	Fibrosis [†]	Standardised linear
16	(2010)					measurement points
17	Wang (2009)	High-frequency	N.R.	N.R.	Fibrosis [†]	N.R.
18	Wang (2010)	B-mode	N.R.	Combined epidermal & dermal	N.R.	Tracing – scar & anatomical
19	U V V					landmarks
20	Wiseman	B-mode	N.R.	Combined epidermal & dermal	Fibrosis [†]	Centrally site of interest
21	(2020, 2021)					
22	(2020, 2021) Wood (1996)	B-mode	7 & 10	NR	NR	Transducer affixed to
23	(1990)	D mode	/ @ 10		10.10.	tracking arm
25	X_{uan} (2021)	High_frequency	20	NR	Fibrosist	N R
26	Xual (2021)	P mode	20	N.R. N D	N D	N.R. N D
27	(2022)	D-IIIOUE	/-10	N.K.	IN.K.	IN.K.
28	(2022)	TT 1 C	10	ND	ND	ND
29	Y1m (2010)	High-frequency	12	N.K.	N.K.	N.K.
30	Zadkowski	B-mode	N.R.	Combined epidermal & dermal	N.R.	N.R.
31	(2016)					
32	Abstracts					
33						
24 25	Agabalyan	N.R.	20	Epidermal, dermal & combined	N.R.	N.R.
36	(2016)					
37	Anthonissen	N.R.	22	Epidermal & dermal	N.R.	N.R.
38	(2015)			•		
39	Bajouri (2018)	High-frequency	N.R.	Epidermal & dermal	N.R.	N.R.
40		-o 1		r ······		

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Bezugly (2019)	High-frequency	22, 33 & 75	Epidermal & dermal	N.R.	N.R.
Bezugly (2014)	High-frequency	33 & 75	Epidermal & dermal	N.R.	N.R.
Blome-	N.R.	N.R.	N.R.	N.R.	N.R.
Eberwein					
(2011, 2012)	High fraguency	ND	ND	Fibrosia	ND
Eberwein	High-frequency	N.K.	N.K.	FIDIOSIS	N.K.
(2012)					
Blome-	High-frequency	N.R.	N.R.	N.R.	N.R.
Eberwein		0 _k			1 11 11
(2014)					
Cho (2012)	N.R.	N.R.	N.R.	N.R.	N.R.
Cho (2012)	N.R.	N.R.	N.R.	N.R.	N.R.
Comstock	N.R.	N.R.	N.R.	N.R.	N.R.
(2018)					
Cooper (2021)	N.R.	N.R.	N.R.	N.R.	N.R.
Du (2006)	B-mode	15	N.R.	N.R.	N.R.
Edgar-	N.R.	N.R.	N.R.	N.R.	N.R.
Lacoursière					
(2022)	ND	ND			ND
El-Zawhary	N.R.	N.R.	N.R.	N.R.	N.R.
(2007)	ND	ND	ND	ND	ND
George (2019)	N.K.	N.R.	N.K.	N.R.	N.K.
Jacobs (2016)	N.K.	N.K.	N.K.	N.K.	N.K.
Jang (2009) $Kim (2000)$	N.K. N D	N.K. N D	N.K.	N.K.	N.K. N D
KIIII (2009)	N.K. N D	IN.K. N D	N.K. N.D	N.K. N D	N.K.
LI(2010)	IN.K. Tiagua Illtragound	IN.K. N D	N.K. N.D	N.K. N D	N.K.
L_1 - I sang	Delection System	N.K.	N.K.	N.K.	N.K.
(2011)	Tiggue Ultrogound	ND	ND	ND	ND
(2010)	Palnation System	IN. K .	IN.R.	1 N.I X.	IN. K .
(2010) Maari (2017)	N R	NR	NR	NR	NR
(2017)	11.11.	± 1.1\.	11.11.	11.11.	11.11.

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Moortgat	High-frequency	N.R.	Dermal	N.R.	N.R.
(2020)					
Nedelec (2018)	N.R.	N.R.	N.R.	N.R.	N.R.
Peters (2018)	High-frequency	22	N.R.	N.R.	N.R.
Seo (2011)	N.R.	7.5	N.R.	N.R.	Thickest point
Siwy (2016)	N.R.	N.R.	N.R.	N.R.	N.R.
Timina (2013)	N.R.	20-40	N.R.	N.R.	N.R.
Tu (2014)	High-frequency	N.R.	N.R.	N.R.	N.R.
	ultrasound				
	biomicroscopy				
Ud-Din (2017)	N.R.	N.R.	N.R.	N.R.	N.R.
Ud-Din (2017)	High-frequency	50	N.R.	N.R.	N.R.
Ud-Din (2018)	High-frequency	N.R.	N.R.	Fibrosis [†]	N.R.
Zuccaro (2021)	N.R.	N.R.	N.R.	N.R.	N.R.
Zuccaro (2019)	B-mode	N.R.	N.R.	N.R.	N.R.
Zuccaro (2021)	B-mode	6-18	Combined epidermal & dermal	N.R.	Scar outlined &
					photographed

Legend: Scar relocation: Methods used by assessors to relocate the measured scar for sequential measurements; B-mode: brightness-mode ultrasound (< 20 MHz); High-frequency: high-frequency B-mode ultrasound (> 20 MHz); N.R.: Not reported

Footnotes: [†]Indirect reference made in record (e.g. in introduction or discussion); [‡]Photographs taken of the scar but not specified whether used for relocation; [§]Not stated in methods, so images provided in record used by authors of this review to provide subjective judgement

First author (year)	Objective measurement methods	Clinician-based rating scale	PROM
Journal articles			
Agabalyan (2017)	Histology	-	-
Alsharnoubi (2018)	Laser Doppler perfusion	VSS	-
Alsharnoubi (2018)	Laser Doppler perfusion	VSS	-
Alshehari (2015)	-	VSS	-
Avetikov (2018)		-	-
Berry (1985)	Transcutaneous oxygen measurement	Scar redness and hypertrophy	Scar redness and hypertrophy rating
	, and the second s	rating scale (0-5 Likert scale)	scale (0-5 Likert scale)
Blome-Eberwein (2012)	Doppler flowmeter – vascularity	VSS	POSAS-P
	Cutometer - pliability	POSAS-O	
	Semmes-Weinstein monofilament		
	Aesthesiometer testing set –		
	sensation		
Blome-Eberwein (2016)	Cutometer – pliability	VSS	POSAS-P
Bioline Eber wein (2010)	Dermaspectrometer – colour	POSAS-0	
	Semmes-Weinstein Aesthesiometer		
	Monofilament Testing Set –		
	sensation		
Blome-Eberwein (2019)	-	VSS	_
Cai (2019)	_	Clinical evaluation	_
Candy (2019)	Spectrocolorimeter – colour	VSS	_
Chae (2016)	Spectrophotometer – nigmentation	VSS	POSAS-P
Chae (2010)	Specifophotometer prementation	POSAS-O	100/10-1
Chang (2014)		VSS	
Chang (2014)	-	Photographic avaluation (0, 10	-
		VAS)	
Chan (2004)	Cutomator viscoalecticity	(AS)	
Chan (2004)	Spectrophotometer pigmentation	-	-
Chang (2001)	Specifophotometer – pigmentation	VSS	
Cheng (2001)	- Maxamatar colour	Treatment officient (0, 10, VAS)	- Itahing scale (0, 4 Likort scale)
Cho(2014)	viexanceier - coloni	Treatment enreacy (0-10 vAS)	number scale (0-4 Likeri scale)

Supplementary Table 4. Additional measurement methods used alongside ultrasound in included studies

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	Tewameter – trans-epidermal water		
	loss		
	Sebumeter – sebum		
	Cutometer – elasticity		
Danin (2012)	Cutometer – elasticity	VSS	-
Deng (2019)	DermaLab Combo – colour	POSAS-O	-
-	Dermoscopy – vascularity		
Deng (2021)	-	VSS	-
Deng (2021)	Doppler – blood perfusion	POSAS-O	POSAS
	Dermlite Foto IIPro – erythema		
Dunkin (2007)		-	-
Elrefaie (2020)	Ultrasound – echogenicity,	VSS	-
	compressibility & vascularity		
Engray (2010)	Durometer – hardness	Clinical appearance based on	_
	Chromameter – colour	photographs	
Fabbrocini (2016)	-	mVSS (vascularity, pigmentation,	_
1 4001001111 (2010)		pliability)	
Fong (1997)	Cutometer – elasticity	Clinical rating – colour change.	-
1 0118 (1227)	Culoniciti Chastienty	consistent itch hypersensitivity	
		blistering	
Fraccalvieri (2013)	Colour power Doppler –	VSS	
Theedivien (2013)	vascularisation	Visual analogue scale – pain and	
	vascularisation	itch	
Fraccalvieri (2011)	Histology	-	-
	Fchocontrastography –		
	neovascularisation		
Gankande (2014)	DermI ab combo $-$ erythema &	mVSS (some participants)	
Gankande (2014)	elasticity	m v 55 (some participants)	-
$G_{e}(2022)$		$POSAS_O$	POSAS
00 (2022)	-	Subjective reports on patient	I OBAD
		range of movement	
$C_{22} K_{22} (2016)$	2D photography thistrass		
Gee Kee (2016)	SD photography – thickness	POSAS-O	POSAS
Guo (2020)	Ultrasound – blood flow grade	-	-
	Shear wave elastography – scar		
	stiffness		

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ear wave elastography – scar fness catrometer – firmness	- VSS POSAS-O VSS POSAS-O VSS POSAS-O VSS - - - Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	- POSAS-P POSAS-P Patient pain & itch scales POSAS-P Patient pain, itch & quality of life rating scales Pain severity (0-10 VAS) - - POSAS-P POSAS-P
catrometer – firmness	VSS POSAS-O VSS POSAS-O VSS POSAS-O VSS - - - Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	POSAS-P POSAS-P Patient pain & itch scales POSAS-P Patient pain, itch & quality of life rating scales Pain severity (0-10 VAS) - - - - POSAS-P POSAS-P
eatrometer – firmness	VSS POSAS-O VSS POSAS-O VSS POSAS-O VSS - - - Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	POSAS-P POSAS-P Patient pain & itch scales POSAS-P Patient pain, itch & quality of life rating scales Pain severity (0-10 VAS) - - - - POSAS-P POSAS-P
catrometer – firmness	POSAS-O VSS POSAS-O VSS POSAS-O VSS - - - Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	POSAS-P Patient pain & itch scales POSAS-P Patient pain, itch & quality of life rating scales Pain severity (0-10 VAS) - - - - POSAS-P POSAS-P
catrometer – firmness	VSS POSAS-O VSS POSAS-O VSS - - Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	POSAS-P Patient pain & itch scales POSAS-P Patient pain, itch & quality of life rating scales Pain severity (0-10 VAS) - - - - POSAS-P POSAS-P
eatrometer – firmness	POSAS-O VSS POSAS-O VSS - - Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	Patient pain & itch scales POSAS-P Patient pain, itch & quality of life rating scales Pain severity (0-10 VAS) - - - - POSAS-P POSAS-P
catrometer – firmness	VSS POSAS-O VSS - - Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	POSAS-P Patient pain, itch & quality of life rating scales Pain severity (0-10 VAS) - - - - POSAS-P POSAS-P
catrometer – firmness	POSAS-O VSS - - Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	Patient pain, itch & quality of life rating scales Pain severity (0-10 VAS) - - - - - POSAS-P POSAS-P
catrometer – firmness	VSS - - - Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	Pain severity (0-10 VAS) - - - - - POSAS-P POSAS-P
catrometer – firmness	- - Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	- - - - POSAS-P POSAS-P
atrometer – firmness	- - Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	- - - - POSAS-P POSAS-P
	- Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	- - - POSAS-P POSAS-P
	- Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	- - POSAS-P POSAS-P
	VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	- - POSAS-P POSAS-P
	vSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	- POSAS-P POSAS-P
	mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	POSAS-P POSAS-P
	vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	POSAS-P
	POSAS-O mVSS (height, pliability, vascularity, pigmentation)	POSAS-P
	mVSS (height, pliability, vascularity, pigmentation)	POSAS-P
	vascularity, pigmentation)	
	POSAS-O	
crometer – tissue thickness	-	-
rce/torque sensor – load applied to		
r		
tometer – elasticity	VSS	Quality of life questionnaire
exameter – colour		
iCam PSI system and mexameter		
lood supply		
ser Doppler flowmetry – perfusion	VSS	-
ectrocolourimeter – scar colour	VSS	Pain & itch (0-10 VAS)
	VSS	Treatment satisfaction
ectrocolourimeter – scar colour	VSS	Pain & itch (VAS scale not specified
ectrocolorimeter – colour	VSS	Pain & itch (VAS)
	ce/torque sensor – load applied to ometer – elasticity xameter – colour Cam PSI system and mexameter ood supply er Doppler flowmetry – perfusion ctrocolourimeter – scar colour ctrocolourimeter – scar colour	ce/torque sensor – load applied to ometer – elasticity VSS xameter – colour Cam PSI system and mexameter ood supply er Doppler flowmetry – perfusion VSS ctrocolourimeter – scar colour VSS vSS ctrocolourimeter – scar colour VSS ctrocolourimeter – colour VSS

Page 61 of 83

 BMJ Open

Li-Tsang (2010)	Spectrocolorimeter – colour	VSS (pliability)	Pain & itch (10-point VAS)
Lobos (2017)	-	Modified Seattle Scar Scale	-
Mamdouh (2021)	-	VSS	Patient satisfaction (VAS)
Meirte (2016)	_	-	-
Miletta (2021)	Colourmeter – scar colour	Unclear, likely POSAS-O	Unclear, likely POSAS-P
1110tuu (2021)	Dermal torque meter – scar		Short Form 36 Quality of Life Survey
$N_{2} = 1 + 1 + 2 = (2014)$	Compliance		
Nedelec (2014)	Cutometer – elasticity	-	-
	Mexameter – colour		
Nedelec (2008)	Cutometer – elasticity Mexameter – colour	mVSS	-
Nedelec (2019)	Cutometer – elasticity	-	-
	Mexameter – colour		
Nedelec (2020)	Cutometer – elasticity	-	Pain & itch (10cm line VAS)
	Mexameter – colour		×
Nicoletti (2015)	-		-
Niessen (1998)	Histology		-
Reinholz (2020)	3D topographic imaging device	POSAS-O	Dermatology Quality of Life Index
			POSAS-P
Reinholz (2016)	Optical coherence tomography – thickness	POSAS-O	Dermatology Quality of Life Index POSAS-P
Schwaiger (2018)	3D topographic imaging device	-	-
Simons (2017)	3D camera – scar height	POSAS-O	-
Soykan (2014)	Slide calliper – dimensions	POSAS-O	POSAS-P
Timar-Banu (2001)	Metric ruler – dimensions	Validated 3-point scoring system	-
		for redness, hardness, itching &	
		pain	
Ud-Din (2019)	Optical coherence tomography –	-	-
	thickness		
	Histology		
van den Kerckhove (2005)	Chromameter – erythema	-	-
van der Veer (2010)	Slide calliper – dimensions	-	-
Wang (2009)	Histology	-	-
Wang (2010)	-	-	-

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Wiseman (2020, 2021)	-	POSAS-O	POSAS-P Numeric rating scale for itch Toronto Paediatric Itch Scale CH-9D BBSID
Wood (1996)	_	VSS	-
Xuan (2021)	Histology	-	-
Yeol Lee (2022)	Cutometer – elasticity	mVSS	-
× ,	Elastography		
Yim (2010)	Cutometer – elasticity	-	-
	Tewameter – trans-epidermal water loss		
	Mexameter – colour		
Zadkowski (2016)		VSS	-
Abstracts			
Agabalyan (2016)	Histology		-
Bajouri (2018)	-	VSS	-
Bezugly (2019)	Clinical or histopathological diagnosis		-
Bezugly (2014)	-	-	-
Blome-Eberwein (2011, 2012)	Doppler vascularity, elasticity and sensation	VSS	Pain and itching scale (0-10 Likert scale)
Blome-Eberwein (2012)	-	-	-
Blome-Eberwein (2014)	Doppler flowmeter – vascularity Cutometer – pliability Semmes-Weinstein monofilament	VSS	POSAS-P
	aesthesiometer testing set – sensation		
Cho (2012)	-	VSS	-
Cho (2012)	CK-MPA Multi-Probe adaptor – pigmentation, erythema and trans- epidermal water loss Cutometer – elasticity	-	-
Comstock (2018)	Computer-based tools – Thickness & pliability	Unclear, likely POSAS-O	Unclear, likely POSAS-P
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Page 63 of 83

 BMJ Open

Cooper (2021)	Colorimeter – pigmentation	Unclear, likely POSAS-O	Unclear, likely POSAS-P
Du (2006)	-	-	-
Edgar-Lacoursiere (2022)	Cutometer – elasticity	-	-
	Mexameter – colour		
El-Zawhary (2007)	Histology	-	-
George (2019)	-	-	-
Jacobs (2016)	Cutometer – pliability	POSAS-O	-
L (2000)	Colorimeter – colour		
Jang (2009)	Mexameter – pigmentation	-	-
	Tewameter – trans-epidermai water		
	loss Sahumatan sahum		
	Sebullieler – sebulli		
	Lesen Deppler – parfusion		
$V_{im}(2000)$	Laser Doppier – perfusion Histology	VCC	
$\mathbf{K}_{1111}(2009)$	Spectrocolourimeter scor colour		- Detionst report of pain & itch
Li(2010) Li Teong (2011)	spectrocolourimeter – scar colour	VSS (thickness plicbility and	Fatient report of pain & nen
L1-1 sang (2011)	-	pigmontation)	-
\mathbf{L} i Teong (2010)	Histology	VSS	Salf report questionnaire
LI-1 sang (2010)	Spectrocolourimeter scor colour	v 33	Sen-report questionnaire
Maari (2017)	Cutometer electicity	191	
	Mexameter - nigmentation		-
Moortgat (2020)	Cutometer = elasticity	Unclear likely POSAS-O	Unclear likely POSAS-P
1001tgat (2020)	Chromameter – colour	Olicical, likely 1 OSAS-O	Olicical, likely 1 OSAS-1
	Tewameter – trans-enidermal water		
	Corneometer – hydration		
Nedelec (2018)	Cutometer – elasticity	_	_
	Mexameter – colour		
Peters (2018)	Cutometer - elasticity	POSAS-O	POSAS-P
	Colourimeter - colour		
Seo 2011	Cutometer – elasticity		
Siwy (2016)	Colourimeter – colour	-	SF-36 Quality of Life Measuremer
	Torque meter – pliability & elasticity		POSAS-P
Timina (2013)		_	
BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

Tu (2014)	-	VSS	-	
Ud-Din (2017)	Laser perfusion imaging	-	-	
	Optical coherence tomography –			
	thickness			
	Histology			
Ud-Din (2017)	Optical coherence tomography –	-	-	
	thickness			
Ud-Din (2018)	Optical coherence tomography –	-	-	
	thickness			
	Histology			
Zuccaro (2021)	Multi-parameter skin analysis device	VSS	Unclear, likely POSAS-P	
	6	Unclear, likely POSAS-O		
Zuccaro (2019)	Acoustic radiation force impulse	-	-	
	ultrasound elastography			
Zuccaro (2021)	Acoustic radiation force impulse –	VSS	POSAS-P	
	stiffness	POSAS-O (did not include		
	DermLab Combo elasticity probe –	surface area and relief subscales)		
	elasticity			
	DermLab Combo colour probe –			
	colour			
Legend: (m)VSS: (Modified) Va	ancouver Scar Scale; POSAS: Patient and	Observer Scar Assessment Scale (PO	SAS-O: POSAS observer sca	ıle;
POSAS-P: POSAS patient scale)	; VAS: Visual Analogue Scale; CHU-9D:	Child Health Utility-9D; BBSIP: Bri	sbane Burn Scar Impact Profi	le
	, <u> </u>			Page 23 of 41
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Supplementary Table 5: Reliability of ultrasound methods reported in each included study

First Author (year)	Reliability Test & Measurement Error	Reliability & Measurement Error Test Statistics & Details
Inter-rater reliability		
Anthonissen (2015)	ICC; SEM	Epidermal – 0.297; 0.02mm
		Dermal – 0.991; 0.13mm
Chang (2014)	Pearson correlation	R=0.90, p<0.001
Dunkin (2007)	N.R.	N.R.
Fong (1997)	ICC	0.93, p=0.146
Gankande (2014)	ICC (95% CI)	Individual site:
		Rater 1 vs rater 2
		'Best scar' – 0.95 (0.92, 0.96)
		'Worst scar' – 0.95 (0.91, 0.97)
		'Normal skin' – 0.94 (0.91, 0.96)
		Rater 1 vs rater 3:
		'Best scar' $-0.86(0.78, 0.91)$
		'Worst scar' -0.91 (0.85, 0.95)
		'Normal skin' -0.92 (0.88, 0.95)
		Rater 2 vs rater 3:
		'Best scar' $= 0.93 (0.89, 0.95)$
		'Worst scar' $-0.96(0.92, 0.97)$
		'Normal skin' $= 0.95 (0.92, 0.97)$
		Average site:
		Rater 1 vs rater 2
		'Best scar' $= 0.97 (0.94, 0.99)$
		'Worst scar' $= 0.98 (0.94, 0.99)$
		(Normal skin' - 0.97 (0.93, 0.99))
		Rater 1 vs rater 3
		'Best scar' $= 0.90(0.77, 0.95)$
		(0.17, 0.93)
		(Normal skin' - 0.96 (0.92, 0.98))
		Rotar 2 vs rater 2
		(Best scar' $= 0.95 (0.88, 0.98)$
		(Normal skin2 - 0.98 (0.94, 0.99))
$L_{00}(2005)$	ICC	0.84 m < 0.01
Lau(2003)		0.04, p<0.01 "A acomtable to high"
Lee (2020)	ICC (05% CD) SEM	Acceptable to high
Lee (2019)	ICC (95% CI), SEM	$\frac{5021}{2}$
		$\begin{array}{c} \text{Single. 0.957} \\ (0.954-0.973) \\ \text{Averages 0.085} \\ (0.077, 0.001) \\ \end{array}$
		Average: 0.985 (0.977-0.991)
		SEM: 0.10 mm
		Unscarred SKIII: Single: 0.067 (0.040, 0.090)
		Single: 0.967 (0.949-0.980)
		Average: 0.989 (0.982-0.993)
N 1.1 (2000)		SEM: 0.04 mm
Nedelec (2008)	ICC (95% CI)	Most severe scar: 0.90 (0.84-0.95)
		Less severe scar: 0.91 (0.85-0.95)
		Donor site: 0.89 (0.82-0.94)
~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~		Normal skin: 0.85 (0.75-0.92)
Seo (2011)	N.R.	"High"
Simons (2017)	ICC (95% CI); SEM	Scar: 0.82 (0.7-0.89); 0.05 cm
		Normal skin: 0.33 (0.08-0.54); 0.03 cm
Van Den Kerckhove	ICC (95% CI); SEM	<u>One day:</u>
(2003)		0.88 (0.81-0.95); 0.29 mm

BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

		<u>Day-to-day:</u> 0.04 (0.00, 0.08): 0.21mm
Intra rator reliability		0.94 (0.90-0.98), 0.2111111
Anthonissen (2015)	ICC: SEM	Enidermal 0.800: 0.01mm
Anthomssen (2013)	ICC, SEIVI	Dermal = 0.001; 0.12mm
C_{an} transfer (2014)		Definal = 0.991, 0.1511111
Gankande (2014)	ICC (93% CI)	$\frac{1}{10000000000000000000000000000000000$
		Worst scar $= 0.92 (0.88, 0.95)$
	ND	Normal skin' $-0.86(0.81, 0.89)$
Gee Kee (2016)	N.R.	N.R.
Lau (2005)	ICC	Intra-rater: 0.98, p<0.01
Lee (2019)	ICC (95% CI)	<u>Scar:</u>
		Single: 0.951 (0.871-0.987)
		Average: 0.983 (0.953-0.966)
		SEM: 0.10 mm
		Unscarred skin:
		Single: 0.948 (0.881-0.976)
		Average: 0.982 (0.954-0.993)
		SEM: 0.04 mm
Li (2013)	ICC	0.89
Seo (2011)	N.R.	"High"
Simons (2017)	ICC (95% CI): SEM	Scar: 0.95 (0.91-0.97): 0.02 cm
		Normal skin: $0.61 (0.41-0.75)$: 0.02 cm
Van Den Kerckhove (2003)	ICC (95% CI); SEM	0.98 (0.97-0.99); 0.11mm
Wang (2010)	SE	Peak: 0.032
		3 months: 0.018
		6 months: 0.399
		9 months: 0.353

Abbreviations used in tables: N.R.: Not reported; ICC: Intraclass Correlation Coefficient; 95% CI: 95% Confidence Interval; SEM: Standard Error of Measurement; SE: Standard Error

Summary of findings for measurement error:

The reported inter-rater SEM measurements for the combined (i.e., epidermal and dermal) thickness measurement of scars was reported in two records as 0.11 mm^5 and 0.5 mm.⁶ The inter-rater SEM for the combined thickness measurement of unscarred skin was also calculated in one record (SEM = 0.3 mm).⁶ The inter-rater SEM was calculated in one record for the measurement of epidermal (SEM = 0.02 mm) and dermal (0.13) measurements⁷, and one record reported only the dermal SEM for scar thickness (SEM = 0.1 mm) and unscarred skin (0.04 mm).⁸ The intra-rater SEM for the combined thickness measurement of scarred skin ranged from 0.18 mm to 0.52 mm, and was measured at 0.2 mm for unscarred skin in one record.⁶ One record reported the intra-rater SEM for epidermal (0.01 mm) and dermal (0.12 mm),⁷ and one record reported the intra-rater SEM for dermal scar (0.1 mm) and unscarred skin (0.04).⁸

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Supplementary Table 6. Methodological considerations for researchers and/or clinicians undertaking measurement of scar thickness using ultrasound.

Consideration	Details & examples of considerations	Publications in our review addressing the consideration	Details reported in included review records
Preventing skin	Using standoff methods	6,9-13	- Use of ultrasound gel to prevent contact
compression	(e.g., ultrasound gel,		between ultrasound transducer and skin surface
during	water bath) to prevent		to minimise compression applied by direct
measurement	transducer touching the		application of transducer ^{6,9-12}
	skin		- Silicone pad placed underneath transducer ¹³
	Application of minimal	14-18	- Transducer held to maintain minimal pressure
	pressure by transducer		on scar 14,15,17
			- Training users to apply minimal force on
			transducer to prevent scar or skin distortion ^{10,18}
	Deliberately	19-21	- Measurement of thickness with and without
	compressing skin to		compression with transducer ^{13,21}
	quantify scar		- Thickness measurements taken using TUPS,
	compressibility		which uses controlled and metered compression
Oniontino the	Orienting the nations	8.18.22	Definit suring through out measurement to
Orienting the	during massurement		- Patient supine throughout measurement to
patient	(a g upright suping		now measurement to be taken in the same
	(e.g., upright, suprie,		position
	Maintaining patient	9	- Patients asked to hold breath during
	stillness during		measurement of scars on the chest to allow
	measurement		shear-wave ultrasound ⁹
Placing	Orientating ultrasound	23	- Direction of transducer recorded to ensure
ultrasound	transducer [e.g.,		consistency ²³
transducer	vertical (superior to		5
	inferior/cranial to		
	caudal), horizonal		
	(medial to lateral)]		

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	Orienting the transducer in relation to the scar (e.g.,	9,15,17,18,22,24-26	- Transducer oriented perpendicular to the skin surface to provide optimal image ^{9,15,18,22,24-26}
	perpendicular) Measuring difficult/tight areas (e.g., axillae or other	6	- Exclusion of fingers and toes in paediatric measurements due to size of measurement area and thin skin ⁶
Relocating scars for	Mapping measurement area (e.g., tracing,	6,12,16,18,20,22,27-32	- Scars traced using translucent paper 18,20,22,27,29,31,32
longitudinal measurement	schematic diagram)		 Scars and surrounding anatomical landmarks traced using translucent paper ¹⁶ Scar mapped on transparent paper, which was then cut out ²⁸ Scar mapped with drawing, no elaboration provided ³⁰ Scars traced using Visitrak (Smith & Nephew Medical Limited, Franke, 0.612)
	Photographing measurement area	24,26,33	- Assessed area marked and photograph taken in initial consultation ^{24,33}
	Measuring specific scar locations (e.g., centre of scar, worst area of scar, counting transducer lengths)	6,8,9,13,19-21,23,30,33-37	 Photographs of scars taken ²⁶ Measurement taken at standardised transducer lengths along surgically created scars of prespecified dimensions ³⁴ Measurements taken at thickest/most severe point ^{19-21,30,33,35,37}, as determined by the patient and/or clinician ⁸
			 Transducer placed on thickest site on peripheral regions ⁹ Transducer placed on area initially identified to have greatest burn depth ²³
	For p	eer review only - http://bmjopen.bmj.com/site/about/guideline	s.xhtml Page 27 of 41

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BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

17,38 Conducting linear measurements from nearby anatomical landmarks .2,24-26,28,29,39,40 Acclimatising Removing scar treatments prior to scar to ultrasound measurement conditions measurement 5,18,22,29,41-46 Acclimatising patient to room prior to measurement

- Measurement area selected by the measurer with -selected area marked with tape ¹³

- Measurements taken at set linear distances from cranial/caudal border of linear sternal scar ³⁶

- Linear measurements from anatomical landmark to measurement site ¹⁷

- Transducer placement mapped in 3dimensional space using a surgical precision tracking arm ³⁸

- Pressure garments removed 10 minutes before measurement $^{\rm 28}$

- Pressure garments removed 15 minutes before measurement to regain original (uncompressed) scar thickness or to reduce blanching effects on measurement ^{20,40}

- Pressure garments/gels/moisturisers removed 20 minutes before measurement ^{8,22,29}

- Pressure garments removed 30 minutes before measurement ^{12,25,26,39}

- Sequential measurement of scars following direct treatment with vacuum massage at 5, 30, 60 and 120 minutes to monitor effect of treatment ²⁴

- Patients rested for minimum 5 minutes before measurement ^{5,18,22}

- Scar exposed to room conditions for 10 minutes ²⁹ to allow equilibrium to be reached with surrounding environment ⁴¹

- Patients resting in room with constant temperature for 15 mins ⁴² to allow scar to stabilise ⁴⁴

Maintaining patient position before measurement

Measuring and/or dermis different skin individually layers

Measuring epidermis

Measuring both epidermis/dermis combined (no

5,6,8,11,12,15,17,18,22,23,26,28,35,40,55-68

- Patients rested for 20 minutes prior to measurement 29,45

- Patients resting for 10 minutes before repeated measurements taken 43

- Patients wait in testing room holding position for 5 min before measurement to stabilise cutaneous blood flow ⁵

- Patients allowed to adapt in controlled room to exclude external variables 46

- Patients remained supine for at least 5 minutes before measurement to avoid artefacts on Doppler imaging ¹³

- Patients allowed to acclimatise to room and assumed a supine position for a minimum of 10 minutes before measurements of biophysical parameters ¹¹

- Measurement of epidermal, dermal and combined epidermal and dermal thickness to allow comparison with histological measurement 47,48

- Measurement of the epidermal and dermal thickness ^{45,49}, combined with layer acoustic density⁷

- Measurement of the epidermal, dermal and subcutaneous thickness, combined with acoustic density 50,51

- Measurement of dermal thickness as treatment thought to affect/target the dermis ^{24,37,52-54}

- Combined epidermal and dermal thickness measurement to provide information on the full thickness of the scar 5,6,8,11,12,15,17,18,22,23,26,28,35,40,55-68

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Measurement objective	individual measurement) Measuring fibrosis/oedema/hair follicles	8,10,11,13,14,16,17,24,25,29-32,34,36,37,45,54,58,60,61,63,64,69-82	- Measurement of fibrosis or collagen architecture ^{8,11,17,24,29-32,34,36,37,45,54,58,61,63,64,69,70,72-} 74,77-79,82
			 Measurement of inflammation/oedema ¹⁴ Quantification of the sub epidermal low echogenic band, indicating oedema ⁶⁰ Measurement of both fibrosis and oedema ^{10,13,16,25,58,71,75,76,80,81}
Factors influencing scar site measurement	Measuring contralateral skin/control scar	9,14,15,23,29,30,52,55-58,83-88 6,8,12,18,22,25,38,43,54,59-61,66,89,90 39,40,45,79,81,82	 Measurement of the presence and density of hair follicles to differentiate scarred and unscarred skin⁵⁴ Measurement of additional, non-scarred subjects ^{55,79} Measurement of unscarred/unaffected skin on same subject as scar measurement contralaterally or at anatomically similar location to provide normative measurements for skin thickness 6,8,9,12,14,15,18,22,23,29,30,38-40,43,45,52,54,56-61,66,81,85-90
			 Measurement of both untreated scar and unaffected skin ⁸²⁻⁸⁴ Measurement of a control scar subjected to care as usual treatment on the same individual ²⁵
	Measuring open wounds or sores in the scar	6	- Use of flexible transparent plastic wrap placed over the measurement area to prevent contact between ultrasound gel and transducer with the open wound/sore ⁶
	Operator training and/or experience	6,8,12,14,16,18,20,24,27-29,31,39,40,58,61,66,72,73,87,91-93	 Trained outcome assessor ^{6,13,16,18,27,72} Measurements taken by radiologist/sonographer 28,66,73,92

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5,6,8,9,11,12,20,23,25,26,31,34,37,40,44,45,47,52,54,57,60,61,66,68,79,85,92,94

measurements per scar

- Assessors with burn experience ^{87,93} - Ultrasound located in department of radiology

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- Measurements conducted by trained therapist/doctor under guidance of experienced radiologist 12,14,29,39

- Measurements conducted by trained clinicians who use device regularly and received training by company representative of devices ^{8,61}

- Device-specific training provided: 1 week ²⁰; 3 sessions of 3 hours for 3 weeks, plus 10 independent assessments of scars using study protocol ⁴⁰; training provided over 3 months ³¹; physical therapist trained in ultrasound application 24

- 3 ultrasound images taken from each patient 9,11,26,31,37,44,45,47,52,54,57,60,79,85

- Clearest of 3 measurements used ¹²

- 3 measurements in 3 locations across scar used. Individual and average measurements reported 40

- Measurements performed in duplicate ^{34,94}

- Measurements taken at different points of the

scar, thickest used for analysis ⁹²

- 5 measurements of each site 6,23

- 9 measurements taken, removal of maximum and minimum, 7 measurements used for average 20

- Measurements taken by 3 assessors at 3 different time points during day 8,61

- Measurement of 2 sites on the same scar²⁵

- Single ultrasound image taken for analysis ⁶⁸

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2				
3 4	Use of additional	Using additional objective assessment	6,9-11,13,15,17,18,21-23,25-27,29,31,32,35,36,40-48,50,53,56-59,66,68-70,75- 80,82-84,86-92,95-111	- Histology/immunohistochemistry 13,17,47,48,50,58,78,79,88,100,103,108,110
5	measurement	instruments (e.g.,		- Blood flow and blood perfusion measurement
7	tools as well as	histology, colour		using laser Doppler perfusion imaging.
7 8	ultrasound	Doppler ultrasound		flowmetry or PeriCam and scar colour and
9	magguramante	cutometer		micro vassal percentage using dermoscopyolour
10	measurements	cutometer,		and micro vessel percentage using definoscopyolodi
11		colourimeter)		and micro-vesser percentage. 35 69 70 83 84 86 87 92 99 101 108
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13				- Oximeter 4
14				- Infra-red camera ⁴¹
15				- Measurement of scar stiffness or
16				pliability/elasticity using elastography or
17				cutometer ^{9,15,18,21,22,25-27,29,43,46,53,57,66,82-}
18				84,86,89,90,96,98,99,101,104-106
19				- Measurement of sensation using Semmes-
20				Weinstein filaments ^{82-84,86}
21				- Measurement of scar colour (including
22				nigmentation and erythema) using
23				spectrophotometer colourimeter chromameter
25				mexameter or Dermlite Foto IIPro ^{18,22,25-}
26				27,32,42,44-46,53,56,66,68,80,82,87,90,91,96-99,101-107,111
27				Maggymemont of theme anidownal yester loss
28				- Measurement of trans-epidermai water loss
29				using Tewameter of scar hydration using
30				Corneometer 40,55,70,77
31				- Measurement of sebum level using sebumeter
32				90,99
33				- Measurement of hardness using durometer ⁹¹
34 25				- Measurement of neovascularisation using
35				echocontrastography ⁵⁸
37				- Measurement of scar dimensions (e.g., scar
38				height and volume) using 3D camera, 3D
39				imaging methods, ruler or calliner ^{6,10,11,23,36,75,77}
40				mono monous, raier or earriper
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43		For	peer review only - http://hmionen.hmi.com/cite/about/quidelines.x	Page 32 of 41

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44 45 46 Using subjective assessment instruments (e.g., clinical rating scales, PROMs) 19,20,23,28-30,33,37,40,41,44,45,49,52,56,57,61,66,67,69-72,80-84,86,87,91,92,94-98,100,111-115 - Measurement of skin thickness using micrometer or optical coherence tomography 17,31,59,76,108-110

- Measurement of scar firmness or deformation using cicatrometer, force/torque sensor (in line with ultrasound to measure load applied) or torque meter ^{31,32,107}

- Multi-parameter skin analysis device ⁶⁶

- Measurement of erythema and elasticity using probes of DermaLab Combo $^{\rm 40}$

- Multi-probe adaptor taking multiple measurements (pigmentation, erythema, transepidermal water loss) ⁹⁶

PROMs:

- Measurement of scar quality using POSAS patient report ^{8,23,30,33,45,56,61,63,64,66,75-}77,82,86,95,97,106,107,114,115

- Subjective rating scales for scar symptoms (e.g., pain, itch) or subjective scar severity ratings ^{26,30,41,42,53,63,64,72,80,83,84,93,102,103,111,115}

- Patient quality of life questionnaires ^{75,76,101,107}
- Measurement of generic health-related quality of life using CHU-9D ^{63,64}

- Measurement of scar-specific health-related quality of life using BBSIP ^{63,64}

- subjective evaluation of response to treatment/treatment satisfaction ^{81,116} Clinical rating scales:

- Measurement of scar quality using POSAS observer report ^{8,23,30,33,45,53,56,61,63,64,66,75-}77,82,86,87,97,98,106,114-116

- Measurement of physical scar characteristics using VSS or modified versions of the VSS ^{8,18-} 20,28,30,33,35,37,38,40,42-44,49,56,57,61,65,66,69-72,80-86,92-95,100-103.111-113.115.117.118 - Measurement of scar characteristics in relation to unscarred skin using Seattle Scar Scale or modified Seattle Scar Scale ⁷³ - Subjective rating scales for scar symptoms (e.g., pain, itch) as assessed by the clinician and/or researcher and/or clinical evaluation of scar severity 11,29,41,52,57,67,73,91,92,94,96 Determining the order - Standardised order of measurement: 3D of measurement photograph, POSAS-O, then ultrasound ⁶ - Order of device use not specified 35,69,70,83,84,86,87,92,99,101,108 Abbreviations: TUPS: Tissue Ultrasound Palpation System; 3D: three-dimensional; POSAS: Patient and Observer Scar Assessment Scale; CHU-9D: Child Health Utility 9D; BBSIP: Brisbane Burn Scar Impact Profile; VSS: Vancouver Scar Scale; mVSS: Modified Vancouver Scar Scale; POSAS-O: Patient and Observer Scar Assessment Scale, observer measure

BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

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BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

Preferred Reporting Items for Systematic reviews and Meta-Analyses extension for Scoping Reviews (PRISMA-ScR) Checklist

SECTION	ITEM	PRISMA-ScR CHECKLIST ITEM	REPORTED ON PAGE #
TITLE			·
Title	1	Identify the report as a scoping review.	1
ABSTRACT			
Structured summary	2	Provide a structured summary that includes (as applicable): background, objectives, eligibility criteria, sources of evidence, charting methods, results, and conclusions that relate to the review questions and objectives.	3-4
INTRODUCTION			
Rationale	3	Describe the rationale for the review in the context of what is already known. Explain why the review questions/objectives lend themselves to a scoping review approach.	5-7
Objectives	4	Provide an explicit statement of the questions and objectives being addressed with reference to their key elements (e.g., population or participants, concepts, and context) or other relevant key elements used to conceptualize the review questions and/or objectives.	7
METHODS			
Protocol and registration	5	Indicate whether a review protocol exists; state if and where it can be accessed (e.g., a Web address); and if available, provide registration information, including the registration number.	7
Eligibility criteria	6	Specify characteristics of the sources of evidence used as eligibility criteria (e.g., years considered, language, and publication status), and provide a rationale.	8-10
Information sources*	7	Describe all information sources in the search (e.g., databases with dates of coverage and contact with authors to identify additional sources), as well as the date the most recent search was executed.	8
Search	8	Present the full electronic search strategy for at least 1 database, including any limits used, such that it could be repeated.	9
Selection of sources of evidence†	9	State the process for selecting sources of evidence (i.e., screening and eligibility) included in the scoping review.	9
Data charting process‡	10	Describe the methods of charting data from the included sources of evidence (e.g., calibrated forms or forms that have been tested by the team before their use, and whether data charting was done independently or in duplicate) and any processes for obtaining and confirming data from investigators.	10-11
Data items	11	List and define all variables for which data were sought and any assumptions and simplifications made.	10-11 and supplementary table 1
Critical appraisal of individual	12	If done, provide a rationale for conducting a critical appraisal of included sources of evidence; describe	N/A



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SECTION	ITEM	PRISMA-ScR CHECKLIST ITEM	REPORTED ON PAGE #
sources of evidence§		the methods used and how this information was used in any data synthesis (if appropriate).	
Synthesis of results	13	Describe the methods of handling and summarizing the data that were charted.	10-11
RESULTS			
Selection of sources of evidence	14	Give numbers of sources of evidence screened, assessed for eligibility, and included in the review, with reasons for exclusions at each stage, ideally using a flow diagram.	11-12
Characteristics of sources of evidence	15	For each source of evidence, present characteristics for which data were charted and provide the citations.	12-15
Critical appraisal vithin sources of evidence	16	If done, present data on critical appraisal of included sources of evidence (see item 12).	N/A
Results of individual sources of evidence	17	For each included source of evidence, present the relevant data that were charted that relate to the review questions and objectives.	Results section (11-46)
Synthesis of results	18	Summarize and/or present the charting results as they relate to the review questions and objectives.	Results section (11-46)
DISCUSSION			
Summary of evidence	19	Summarize the main results (including an overview of concepts, themes, and types of evidence available), link to the review questions and objectives, and consider the relevance to key groups.	47-49
Limitations	20	Discuss the limitations of the scoping review process.	49-50
Conclusions	21	Provide a general interpretation of the results with respect to the review questions and objectives, as well as potential implications and/or next steps.	50-51
FUNDING			
Funding	22	Describe sources of funding for the included sources of evidence, as well as sources of funding for the scoping review. Describe the role of the funders of the scoping review.	51

JBI = Joanna Briggs Institute; PRISMA-ScR = Preferred Reporting Items for Systematic reviews and Meta-Analyses extension for Scoping Reviews.

* Where *sources of evidence* (see second footnote) are compiled from, such as bibliographic databases, social media platforms, and Web sites.

† A more inclusive/heterogeneous term used to account for the different types of evidence or data sources (e.g., quantitative and/or qualitative research, expert opinion, and policy documents) that may be eligible in a scoping review as opposed to only studies. This is not to be confused with *information sources* (see first footnote).
‡ The frameworks by Arksey and O'Malley (6) and Levac and colleagues (7) and the JBI guidance (4, 5) refer to the process of data extraction in a scoping review as data charting.

§ The process of systematically examining research evidence to assess its validity, results, and relevance before using it to inform a decision. This term is used for items 12 and 19 instead of "risk of bias" (which is more applicable to systematic reviews of interventions) to include and acknowledge the various sources of evidence that may be used in a scoping review (e.g., quantitative and/or qualitative research, expert opinion, and policy document).

From: Tricco AC, Lillie E, Zarin W, O'Brien KK, Colquhoun H, Levac D, et al. PRISMA Extension for Scoping Reviews (PRISMAScR): Checklist and Explanation. Ann Intern Med. 2018;169:467–473. doi: 10.7326/M18-0850.



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Ultrasound measurement of traumatic scar and skin thickness: A scoping review of evidence across the translational pipeline of research-to-practice

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Ultrasound measurement of traumatic scar and skin thickness: A scoping review of evidence across the translational pipeline of research-to-practice*

<u>Running Title:</u> Review of scar thickness measurement with ultrasound

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ABSTRACT:

 Objectives: To identify the ultrasound methods used in the literature to measure traumatic scar thickness, and map gaps in the translation of these methods using evidence across the research-to-practice pipeline.

Design: Scoping review

Data Sources: Electronic database searches of Ovid MEDLINE, Embase, Cumulative Index of Nursing and Allied Health Literature (CINAHL) and Web of Science. Grey literature searches were conducted in Google. Searches were conducted from inception (date last searched 27/05/2022).

Data Extraction: Records using B-mode ultrasound to measure scar and skin thickness across the research-to-practice pipeline of evidence were included. Data was extracted from included records pertaining to: methods used; reliability and measurement error; clinical, health service, implementation and feasibility outcomes; factors influencing measurement methods; strengths and limitations; and use of measurement guidelines and/or frameworks.

Results: Of the 9309 records identified, 118 were analysed (n = 82 articles, n = 36 abstracts) encompassing 5213 participants. Reporting of methods used was poor. B-mode, including high-frequency (i.e., > 20 MHz) ultrasound was the most common type of ultrasound used (n = 72 records; 61% of records), and measurement of the combined epidermal and dermal thickness (n = 28; 24%) was more commonly measured than the epidermis or dermis alone (n = 7, 6%). Reliability of ultrasound measurement was poorly reported (n=14; 12%). The scar characteristics most commonly reported to be measured were epidermal oedema, dermal fibrosis and hair follicle density. Most records analysed (n = 115; 97%) pertained to the early stages of the research-to-practice pipeline, as part of research initiatives.

Conclusions: The lack of evaluation of measurement initiatives in routine clinical practice was identified as an evidence gap. Diverse methods used in the literature identified the need for greater standardisation of ultrasound thickness measurements. Findings have been used to develop nine methodological considerations for practitioners to guide methods and reporting.

STRENGTHS AND LIMITATIONS OF THIS STUDY:

- Use of the Australian Government Department of Health and Aged Care Medical Research Future Fund research-to-practice pipeline phases to categorise records allowed identification of gaps in the use of ultrasound for clinical practice.
- Clinical, health service, implementation and feasibility outcomes related to ultrasound measurement in included records were summarised to determine what is needed to close the research-to-practice gap for ultrasound measurement of scar thickness.
- A limitation is that only articles available in English or with an English abstract were considered for inclusion and data extraction, thus findings are likely most relevant to English speaking countries.

INTRODUCTION:

Traumatic cutaneous injury, caused by sharp object penetration (e.g., surgery or vaccination) or burns (including thermal, chemical and friction) may result in the formation of hypertrophic scarring. (1) Hypertrophic scars result from an aberrant cutaneous healing response that leads to the formation of red, raised scars, often accompanied by pruritus and skin tightening, which remain within the boundaries of the initial injury. (2-7) The sequelae of hypertrophic scars can impact on patient's physical and psychosocial quality of life. (8, 9)

A characteristic of hypertrophic scarring that both patients and clinicians have identified as being important, and which has subsequently been used as a way to measure clinical and treatment outcomes, is scar thickness. (9-17) Scar thickness can be measured both subjectively, through clinician assessment and patient-reported outcomes, or objectively, utilising medical imaging methods. (18, 19) The pathological complexity of hypertrophic scars means that they generally extend below the level of the surrounding skin, supporting the use of medical imaging modalities such as ultrasound for thickness quantification, as these are capable of providing information about subcutaneous structures and processes. (19, 20) Scar thickness measurement using ultrasound can be conducted in both clinical and research contexts. Where routine measurements like ultrasound are used to guide clinical decisionmaking and treatment, this practice is known as measurement-based care. (21)

Ultrasound is a safe, non-invasive and largely cost-effective (compared to other imaging modalities) imaging method with measurement utility in both adult and paediatric populations. (22-24) Modern B-mode (brightness mode) ultrasound, particularly high- (i.e., ≥20 MHz) or ultra-high frequency (30-100 MHz) (25) ultrasonography, allows differentiation between the epidermis and dermis, which permits quantification of skin layer-specific scar characteristics. This differentiation may allow assessors to observe and understand the pathological mechanisms of individual scars and adjust treatment protocols accordingly. (24,

Page 7 of 83

BMJ Open

26-31) Additionally, B-mode ultrasound is commonly used as the basis for other imaging methods, such as colour Doppler ultrasound or elastography, which can allow quantification of additional scar characteristics, such as their elastic properties. (26-29, 32, 33)

Despite the clinical advantages of B-mode ultrasound for scar thickness measurement, methods are poorly reported and lack standardisation in the literature. This casts doubt on the validity of clinical decision-making in measurement-based care initiatives (e.g., setting depth of AFCO₂ penetration) informed by research findings (e.g., response to treatment) where ultrasound measurements are used. (34) Lack of standardisation also makes between-study comparison, such as systematic reviews and meta-analyses, difficult, (35) and poor methodological reporting hampers the ability to accurately replicate findings. This scoping review focusses on mapping and identifying gaps in ultrasound methods and evaluation reported in the current literature along the research-to-clinical practice pipeline. (36) Methodological considerations for people performing ultrasound scar thickness measurements, including practitioners (herein termed assessors) using ultrasound in clinical practice are presented based on the review findings.

METHODS:

Protocol Publication and Review Structure:

The protocol for this review has been published *a priori*. (37) This scoping review was conducted and is reported according to the Arksey and O'Malley (2005) (38) framework. The steps outlined in this framework are: 1) identifying the research question; 2) identifying relevant records; 3) selecting appropriate records; 4) charting extracted data; and 5) collating, summarising and reporting the results. (38)

Research Question:

The primary question of this scoping review was: "What do we know and not know about the measurement of traumatic cutaneous scar thickness using ultrasound?" This question was addressed through exploration of: methods used; reliability and measurement error; clinical, health service, implementation and feasibility outcomes; factors influencing ultrasound imaging and measurement methods; strengths and limitations of measurement methods; and use of measurement guidelines and/or frameworks. While the focus of this review was the measurement of traumatic cutaneous scar thickness with ultrasound, methods used to measure the thickness of unscarred skin were reported where these were used in combination with measurement of scar thickness (e.g., as control or comparator measurements).

Identifying Relevant Records:

A standardised search strategy was developed and piloted with the assistance of a medical librarian using the concepts 'ultrasound', 'skin', 'thickness' and 'measure', with associated terms and truncations (supplementary box 1). Ovid MEDLINE, Embase, Cumulative Index of Nursing and Allied Health Literature (CINAHL) and Web of Science electronic databases were searched from conception to identify original studies (date last searched 27th May 2022).

The phrase 'ultrasound scar thickness measurement' was used to conduct additional searches in 1) Google Scholar, and 2) Google to identify original studies in grey literature, and studies not identified in database searches. Title and abstract searches in Google Scholar and Google were limited to the first 200 results. (39)

Record Selection:

Following de-duplication, six reviewers screened records using Covidence (Veritas Health Innovation, Melbourne, Australia; available at <u>www.covidence.org</u>) for eligibility according to the inclusion criteria (Table 1). Both peer-reviewed journal articles and abstracts were

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included to ensure that all the available and most recent methodological information was obtained. (40) Data collected from peer-reviewed journal articles was considered the primary source of data, with information from abstracts used to confirm or extend the journal data. The inclusion of abstracts will assist future authors to further investigate the information presented as full texts may become available. During both title and abstract and full text screening, one researcher (BM) screened all records as a single reviewer, while other researchers (MS, TM, TR, BD and ZT) screened records as a second reviewer. Conflicts were resolved through discussion between at least two authors to reach agreement. A third author was used as a tiebreaker where agreement could not be reached.

Table 1. Inclusion and exclusion criteria for studies included in the scoping review.

Inclusion	Exclusion
 Traumatic scars measured with ultrasound based on B-mode ultrasound (including high-frequency, ultra-high-frequency and Doppler) Measurements taken of living, human individuals Measurement of traumatic cutaneous scarring arising from penetration of the skin with sharp objects (including surgery or vaccination), or as a result of burns, (including thermal, chemical or friction) Articles written in English, or with English abstracts 	 Reviews, discussion papers, opinion pieces Measurement of non-traumatic scars (e.g., acne scars). Where non-traumatic scars measured along with burn scars, these were included Measurement of skin thickness in non-traumatic conditions (e.g., diabetes) Measurement of skin thickness where there is no cutaneous involvement in the trauma (e.g., traumatic brain injury) Measurement using A-mode ultrasound

Charting the Data:

The data extraction table was developed in Microsoft Excel and piloted by two authors (BM and ZT) through independent extraction and comparison of data from two records. The table was then modified to include the scar characteristics (e.g., fibrosis, oedema) measured, measurer/assessor training, the number of measurements taken and funding sources

(Supplementary Table 1). Full text data extraction was completed by four authors (BM, MS, TM and ZT). An additional author (BD) independently extracted data from five randomly selected records, which was compared to data extracted by other authors. Minimal differences between data extracted by the independent author and that by other authors were observed, thus further independent extraction was not performed. As is typical in scoping reviews, the certainty or quality of evidence was not appraised. (38)

The research-to-practice pipeline published by the Australian Government Department of Health and Aged Care Medical Research Future Fund (figure 1) was used to categorise each included record based on their stated aims into one of the four phases. (36) Studies related to phase 1 of this pipeline, basic research, were only included in this review when data on scar or skin thickness pertained to human participants (table 1). Phase 2 of this pipeline included randomised controlled trials, while phase 3 included pragmatic and observational studies conducted outside randomised controlled trials. The final phase of this pipeline (phase 4) indicates initiatives used in routine clinical practice.

Where clinical (e.g., treatment satisfaction, scar symptoms), health service (e.g., efficiency, safety, effectiveness, equity, patient-centredness and timeliness) and implementation (e.g., acceptability, adoption, appropriateness, fidelity, cost, penetration and sustainability) outcomes were addressed, they were reported and defined according to Proctor *et al.* (41). For example, in the context of this scoping review, acceptability is defined as the level to which ultrasound is palatable amongst stakeholders (e.g., assessors), appropriateness is the perceived fit of ultrasound within regular clinical practice, and fidelity is the degree to which ultrasound is used in the way it was initially described. (41) Measurement instrument-specific feasibility outcomes defined by Prinsen *et al.* (42) are reported in the current review. These outcomes included ease of administration, standardisation, completion time, instrument cost and availability, and ease of score calculation. (42) Reliability and measurement error were

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defined according to COnsensus-based Standards for the selection of health Measurement INstruments (COSMIN) tools. (43, 44) Measurements with an intraclass correlation coefficient (ICC) of 0.7 or greater were considered reliable. (44) Measurement error was assessed by comparing the reported standard error of the measurement (SEM) with the reported smallest detectable change (SDC). Where the reported measurement error was smaller than the reported smallest detectable change, it was interpreted as indicating real change or variance can be detected, and that change or variance is not a result of error. (44)

Patient and Public Involvement

There was no patient and/or public involvement in the design, conduct, reporting or dissemination of information in this scoping review.

RESULTS:

Electronic database searches identified 9309 records. After removal of 3703 duplicate records, the titles and abstracts of 5606 records were screened for relevance according to the inclusion criteria (Table 1). Following full-text screening, 104 records proceeded to data extraction. Searches in Google and Google Scholar identified an additional 14 records, providing a total of 118 records for data extraction. Search and screening results are presented according to the Preferred Reporting Items for Systematic reviews and Meta-Analyses (PRISMA) flow diagram (supplementary figure 1). (45)

Record Characteristics:

Of the 118 records included in this review, 82 were journal articles (69%) and 36 were abstracts (31%) (Table 2), representing a total of 5213 participants (range 1-438; mode 20 participants per record). Adults aged 18 years and older were the most highly represented age group reported in articles (n = 43 articles; 52% of articles), (17, 26, 29, 46-85) while most abstracts did not report the age group measured (n = 25 abstracts; 69% of abstracts). (86-110)

The most common scar type measured was burn scars in both journal articles (n = 43 articles; 52% of articles), (17, 22-24, 27, 47, 57-59, 61, 62, 64-67, 71-75, 81, 82, 84, 111-130) and abstracts (n = 23 abstracts; 64% of abstracts) (28, 30, 86-88, 91-94, 96, 98, 102-106, 131-135) (Table 2). Most identified articles used ultrasound measurement of scar thickness as part of research initiatives, and were categorised as either phase 2 (n = 50 articles; 61% of articles) (17, 22, 26, 31, 46-49, 51-56, 61, 63-65, 67, 69-71, 74-76, 78, 81, 83, 84, 111, 112, 114, 115, 117, 124-127, 129, 130, 136-145) or phase 3 (n = 30 articles; 37% of articles). (23, 24, 27, 29, 50, 57-60, 62, 66, 68, 72, 73, 77, 79, 80, 82, 85, 116, 118, 120-123, 128, 146-149) on the research-to-practice pipeline. (36) Phase 2 was also the most common phase represented by abstracts (n = 21; 58% of abstracts), (86, 88, 91, 93, 95, 97, 99-104, 106-108, 131-134, 150, 151) followed by phase 3 (n = 15 abstracts; 42% of abstracts). (28, 30, 87, 89, 90, 94, 96, 98, 105, 109, 110, 135, 152-154) Phase 4 was addressed by two articles (2% of articles) (113, 119) and one abstract (2% of abstracts), (92) which used ultrasound to measure treatment response to an intervention already used in routine clinical practice, including compression garments (113, 119) and CO₂ fractional laser. (92) No records pertained to phase 1.

Characteristic	Category	Number of Records (Translational Pipeline Phase 2*)	Number of Records (Translational Pipeline Phase 2*)	Number of Records (Translational Pipeling Phase 4*)
Journal Articles		Tipenne Thase 2)	Tipenne Thase 5)	Tipenne Thase 4)
Funding	Commercial	2	1	1
Source	Non-	23^{-}	13	0
	commercial		10	· ·
	Commercial	2	1	1
	& Non-	-	-	-
	commercial			
	No funding	6	3	0
	Not reported	16	12	ů 0
Population	Adult	27	12	0
Type	Paediatric	21 6	4	0
Гурс	Paediatric	13	7	2
	and Adult	15	7	2
	Not reported	3	3	0
Scar Aetiology	Burn	22	18	1
	Surgical [†]		2	0
	Mixed	10	23	0
	Not specified	10	7	0
Abstracts	rtorspecifica		· · · · · · · · · · · · · · · · · · ·	
Funding	Commercial	0	0	0
Source	Non-	3	1	0
	commercial			
	Commercial	0	0	0
	& Non-			-
	commercial			
	No funding	0	0	0
	Not reported	17	14	1
Population	Adult	1	2	0
Туре	Paediatric	0	3	0
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	Paediatric	4		· · · · · · · · · · · · · · · · · · ·
	and Adult	4		0
	and Adult Not reported	4	9	1
Scar Aetiology	and Adult Not reported Burn	4 15 12	9 10	1
Scar Aetiology	Paediatric and Adult Not reported Burn Surgical [†]	4 15 12 1	9 10 2	1 1 0
Scar Aetiology	Paediatric and Adult Not reported Burn Surgical [†] Mixed	15 12 1 2	9 10 2 1	1 1 0 0

Table 2. Summary of characteristics of records included in this review*

Legend: Paediatric: measurement of patients under the age of 18; Adult: measurement of patients aged 18 years or older; Burn: scars caused by thermal, chemical or friction injury; Surgical: scars caused by surgical procedures (including biopsies); Mixed: scars of included record were of mixed origin (e.g., burn and acne)

Footnotes: *Stage in the research to clinical practice translational pipeline, as defined by the Australian Government Department of Health and Aged Care (36); [†]Type of surgery defined in supplementary table 2

* A breakdown of each characteristic per record is presented in Supplementary Table 2

Methods used to measure traumatic cutaneous scar thickness:

B-mode, including high-frequency B-mode ultrasound (i.e., ≥ 20 MHz) was the most

commonly reported ultrasound type in the included articles (n = 56; 68% of articles) (17, 22-

24, 26, 29, 31, 46-49, 53, 54, 56, 57, 59, 60, 64, 65, 67, 69-78, 80-82, 84, 85, 111, 112, 114,

116-118, 120, 122, 123, 126-130, 138, 139, 141, 142, 144-146, 149), while most abstracts did

- not report the type of ultrasound used (n = 22; 61% of abstracts) (86, 87, 92-98, 101, 103,
- 105, 106, 108, 131-134, 150-153) (Table 3). Specialised B-mode ultrasound devices,

including the Tissue Ultrasound Palpation System (TUPS; a B-mode ultrasound transducer

in-series with a load cell to allow measured compression of the skin), (68, 99, 100, 124) and

colour Doppler ultrasound, (52, 149) were used in six records (Table 3). 9)...
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reported	42
node	3
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h-frequency	9
er	3
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ler	1
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lema	0
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reported	34
	er reported hode drange h-frequency er reported dermal mal dermal & dermal mbined epidermal & dermal er reported rosis lema rosis & oedema er reported dess-mode ultrasound (<20 MHz); H

12 Table 3. Summary of measurement methods used in included record*

The type of scar and skin thickness measurement (i.e., thickness of the dermis, epidermis, or combined epidermal and dermal measurement) was reported in 39 records (33%) (Table 3). Where reported, combined measurement of epidermal and dermal thickness was the most common method used in articles (n = 32; 76% of articles reporting skin measurement type). (17, 22-24, 27, 29, 50, 53, 56-58, 60, 64-66, 70, 72-77, 80-82, 114, 116, 118, 122, 126, 127, 130, 139, 146, 148) Separate epidermal and/or dermal thickness measurements were reported in seven journal articles (17% of articles reporting skin thickness measurement type). (26, 47, 48, 52, 53, 71, 118) Of these records, two authors provided a rationale for this decision: each skin layer provided different information on the scar; (26) or responded differently to treatment. (67, 71) Most abstracts did not report the type of skin measurement used (n = 30; 83% of abstracts). (28, 30, 91-101, 103-110, 131-134, 150-154) Three articles (4% of articles) (47, 110, 111) and one abstract (3% of abstracts) (28) directly

reported that fibrosis was the scar characteristic targeted by the measurement. One of these records also quantified hair follicle density to assess the difference between scared and unscarred skin. (47) An additional 25 articles (30% of articles) (17, 46, 52, 53, 56, 63-65, 67, 70, 79, 80, 83, 84, 112, 120, 123, 125-127, 140, 142, 145, 148, 149, 155) and one abstract (3% of abstracts) (110) made indirect reference (i.e., within the introduction or discussion) to the measurement of fibrosis. Ten journal articles (12%) made indirect reference to the measurement of both oedema and fibrosis, (31, 54, 55, 71, 74, 76-78, 138, 144) and one record made indirect reference to the measurement of oedema. (59)

Additional objective and/or subjective measurement methods were employed alongside
ultrasound measurement in 72 articles (88% of articles) (17, 22, 24, 26, 29, 31, 46-53, 55-57,
60-70, 72-81, 83-85, 111-122, 124-130, 136-142, 144, 145, 147-149) and 31 abstracts (86%
of abstracts) (86, 88, 89, 91-95, 97-110, 131-134, 150, 151, 153, 154) (Supplementary Table
All three phase 4 studies involving implementation in routine clinical practice utilised

Page 17 of 83

BMJ Open

additional measurements. (92, 113, 119) The additional objective measurements used in included records were elastography (elasticity), cutometric assessment (pliability) and Doppler ultrasound (vascularity). The additional subjective measurements were conducted using clinician-based rating scales (e.g., Vancouver Scar Scale or modified Vancouver Scar Scale) or Patient Reported Outcome Measures. The Vancouver Scar Scale was used in 35 articles (43% of articles) (17, 31, 46, 47, 49, 50, 52, 55, 57, 61-64, 66-70, 73, 85, 111, 112, 114, 116, 118, 121, 124, 128, 130, 136-138, 140-142) and 11 abstracts (31% of abstracts) (88, 91, 92, 98-100, 107, 134, 150, 151, 153). Patient-reported outcome measures (PROMs) were used in 27 articles (33% of articles) and 11 abstracts (31% of abstracts). (46, 53, 56, 57, 60, 72-75, 85, 91, 94, 97, 101-106, 111, 112, 114, 115, 117, 118, 120, 122, 129, 131-133, 138, 140, 141, 148, 150, 151, 153, 154) Of the records that reported using PROMs, the most commonly used was the patient report of the Patient and Observer Scar Assessment Scale (POSAS), used in 17 articles (63% of articles reporting use of PROMs) (17, 22, 46, 50, 53, 61, 62, 64, 76, 77, 79, 114, 121, 125-127, 147) and 8 abstracts (73% of abstracts reporting use of PROMs) (91, 93, 102, 104, 106, 132, 153) (Supplementary Table 4). In most cases, additional measurement methods were used to supplement ultrasound thickness measurements as research outcomes. In some records (n = 16; 14% of records), however, ultrasound was compared with histology, POSAS, dermoscopy, VSS and modified VSS, clinical assessment, modified Seattle Scar Scale, high-definition optical coherence tomography, 3D camera, immunohistochemistry, and immunohistomorphometry. (17, 24, 26, 29, 31, 50, 51, 64, 73, 77, 86, 95, 110, 120, 124, 149) Where the effectiveness of ultrasound was judged against other methods, it was only found to be inadequate against histology. (26, 86)

Methods used to relocate the scar for repeated measurements were reported in 34 records
(29%) (Supplementary Table 3). The most common relocation method was tracing the outline

64	or boundaries of the scar on a transparent or translucent sheet ($n = 14$ articles; 35% of articles
65	reporting scar relocation), (23, 49, 65, 74, 81, 115, 116, 120, 124, 125, 153) occasionally
66	including prominent or bony landmarks close to the scar. (23, 24, 72, 73, 123) Photographs (n
67	= 10 articles; 25% of articles reporting relocation and $n = 1$ abstract) and linear measurements
68	from defined points or anatomical landmarks on or around the scar ($n = 4$ articles; 10% of
69	articles reporting relocation) were also used for scar relocation. The 'worst' or 'thickest' part
70	of the scar, as determined by patients or assessors, was chosen as the measurement site in 14
71	journal articles (35% of journal articles reporting relocation) (23, 31, 52, 54, 57, 61, 62, 67,
72	126, 127, 138, 141, 148, 155) and one abstract. (105)
73	Measurement of unscarred skin either contralateral or adjacent to the scar, was performed in
75	Wedstreinent of unsearred skin, entier contrancerar of adjacent to the sear, was performed in
74	32 articles (39% of articles%) (17, 22-24, 27, 29, 46-48, 50, 51, 53, 56-60, 64, 72, 73, 80, 81,
75	85, 114, 118, 120-122, 128, 145, 146, 148) and 7 abstracts (19% of abstracts) (28, 94, 95,
76	150, 151, 153, 154) These measurements were primarily used as controls or comparators to
77	scar measurements (n = 27, 69% of records reporting unscarred skin measurement). (17, 22,
78	23, 28, 29, 47, 48, 51, 53, 56-60, 64, 67, 73, 80, 85, 95, 118, 120, 122, 128, 146, 148, 153,
79	154) Additionally, four records (10% of records reporting unscarred skin measurement)
80	evaluating treatment efficacy measured both unaffected skin thickness and the thickness of a
81	'control' or untreated scar. (46, 74, 94, 114) All instances where additional ultrasound
82	measurements were taken of unscarred skin or untreated scars were reported as part of
83	research initiatives aligning with phases 2 and 3 of the research-to-practice pipeline (figure
84	1). (36)
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85 Reliability and measurement error

Reliability was calculated for both scarred and unscarred skin in 13 articles (16% of articles)
and two abstracts (5% of abstracts), and was generally considered acceptable (Supplementary

Page 19 of 83

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BMJ Open

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88	Table 5). This included inter-rater reliability ($n = 5$; 4% of articles), (54, 64, 73, 120, 137)
89	intra-rater reliability ($n = 3$; 4% of journal articles), (22, 23, 65) and both inter- and intra-rater
90	reliability (n = 7; 6%; including 2 abstracts) (17, 24, 57, 82, 87, 105, 124). The intraclass
91	correlation coefficient (ICC) was the most commonly reported reliability statistic ($n = 10$; 8%
92	of records, including one abstract), (17, 24, 57, 64, 65, 73, 82, 87, 120, 124) where it was
93	reported for both scar and unscarred skin measurements in four articles (5% of articles). (17,
94	24, 57, 73) The reported combined thickness (i.e., epidermal and dermal) ICCs for inter-rater
95	reliability of scarred skin ranged from 0.82 to 0.985, while the inter-rater ICC for the
96	measurement of unscarred skin ranged from 0.33 to 0.98, with one of the four records
97	reporting an ICC below the threshold value of 0.7 (ICC = 0.33) (24) and one record simply
98	reported that the inter-rater ICC for scarred skin was "acceptable to high". (64) The reported
99	intra-rater reliability for combined thickness measurements of scarred skin ranged from 0.89
100	to 0.983, and for unscarred skin ranged from 0.61 to 0.982, with one record reporting an ICC
101	below the threshold of 0.7 (ICC = 0.61). (24) One record reported both the inter- and intra-
102	rater ICCs for individual epidermal (inter-rater ICC = 0.297 ; intra-rater ICC = 0.809) and
103	dermal (inter-rater ICC = 0.991 ; intra-rater ICC = 0.991) scar thickness measurement. (87)
104	Four articles (5% of articles) reporting reliability used Pearson's R, an undisclosed method,
105	or description (e.g., high) as detailed in supplementary table 2. (22, 54, 105, 137)
106	Measurement error for inter-rater and intra-rater reliability of combined, epidermal or dermal
107	thickness was reported in four articles (5% of articles) and one abstract using standard error
108	of the measurement (SEM). The inter-rater SEM for the combined epidermal and dermal
109	thickness of scarred skin ranged from 0.11 mm to 0.5 mm, and the intra-rater SEMs ranged
110	from 0.18 to 0.52 mm. Individual records reported SEM values for unscarred skin, and
111	separate epidermal and dermal measurements, available in Supplementary Table 5. (17, 23,
112	24, 82, 87) Only one record reported calculation of the smallest detectable change (SDC). In

that record the inter-and intra-rater SDC was calculated for both scarred and unscarred skin.
The scarred skin SDCs were 1.4 mm (inter-rater) and 0.6 mm (intra-rater), and unscarred skin
SDCs were 0.8 mm (inter-rater) and 0.5 mm (intra-rater). (24) The reported SEMs were all
close to or below the largest SDC value reported. This finding may indicate that ultrasound
can detect true variance in scar thickness above measurement error for traumatic scar and skin
thickness.

Of the records that reported reliability and measurement error, measurements were taken by
practitioners with varying clinical expertise and roles within the treating team. These
included therapists, nurses and doctors, sometimes under the supervision of trained
radiologists. One record reported that 3 assessors received 3 hours of training, and conducted
10 assessments using the study protocol before the study began. (57)

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Clinical, health service, implementation and feasibility outcomes:

No record specifically investigated clinical, health service, implementation or feasibility outcomes of ultrasound as a measurement-based-care initiative. Ultrasound was used to assess the clinical outcomes of scar treatment initiatives in all included records. Clinical, health service, implementation and feasibility outcomes related to ultrasound measurement were, however, reported in 53 journal articles (17, 22-24, 26, 27, 31, 46-48, 50, 51, 54, 56-61, 63-66, 69-75, 77, 80, 82, 113-116, 119, 120, 122-124, 128, 129, 138, 142-144, 148, 149, 155) and 14 abstracts (28, 86, 87, 89, 90, 95, 96, 102, 105, 107, 109, 110, 152, 153) that focused on scar treatments.

The clinical outcome of patient satisfaction related to ultrasound measurement was only reported in one journal article. Whilst patient satisfaction was not directly measured in that record, a proxy measure of satisfaction was reported by the authors stating that no paediatric

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patient or their caregiver refused ultrasound measurement once the purpose was explained.(24)

Timeliness was the only reported health service outcome, reported as the time required to
take ultrasound measurements. Where reported in three journal articles, this was short, taking
between one to five minutes. (24, 27, 122)

The most common implementation outcomes reported in the identified records were fidelity, 141 acceptability and appropriateness. Fidelity to the measurement method was reported through 142 143 the use of experienced or trained assessors (n = 6 journal articles; n = 1 abstract), (24, 57, 58, 87, 142, 144, 148) and/or utilising the same assessor/s for all measurement sessions (n = 5) 144 journal articles; 6% of included journal articles). (24, 61, 138, 144, 148) Differences between 145 146 intended and actual measurement methods were not discussed. The training and/or experience of the assessors was discussed in 24 records (23 journal articles and 1 abstract), (17, 23, 24, 147 27, 51, 56-59, 63-66, 71, 73, 115, 116, 120, 123, 124, 138, 144, 149, 153) where 148 measurements were either taken by a clinician (n = 13; 54% of records reporting training), 149 (17, 23, 24, 58, 59, 64-67, 71, 120, 124, 141) members of the research team (n = 6; 25%) of 150 records reporting training), (57, 63, 73, 115, 123, 144) or by specialist sonographers and/or 151 radiologists (n = 5, including one abstract; 21% of records reporting training). (56, 116, 138, 152 149, 153) Only one record reported on fidelity in the context of routine clinical practice. In 153 154 this instance, ultrasound was conducted in the department of radiology, however the role or training of the staff was not reported. (119) 155

The acceptability and appropriateness of the ultrasound methods used in individual records
were generally based on author opinion and outlined in the discussion. Acceptability was
reported in 26 records (23 journal articles and 3 abstracts), (17, 22-24, 26-28, 31, 57, 64, 70,
74, 75, 77, 80, 82, 86, 96, 116, 119, 120, 122, 124, 143, 149, 155) including for paediatric

Page 22 of 83

BMJ Open

populations, where one record reported potential difficulty in measuring this population, (22) contrasting that which reported that measurement was acceptable to both children and their caregivers. (24) One record reported acceptability where the intervention being analysed by ultrasound was already part of routine clinical practice. In this instance, the authors referenced additional publications which stated that ultrasound had an accuracy of 0.5 mm, which was judged by the authors to be sufficient for assessment of scar thickness. (24, 27, 119, 122) Potential difficulty was identified in the measurement of open wounds, (24) and traditionally hard-to-reach areas (such as the axillae or groin). (22) The appropriateness of the ultrasound methods was reported in 35 journal articles (43% of included journal articles) (22, 24, 26, 27, 31, 46-48, 50, 54, 57, 60, 61, 64-66, 69, 72-75, 77,

80, 82, 113, 114, 116, 119, 120, 122, 124, 128, 148, 149, 155) and 11 abstracts (31% of included abstracts) (86, 87, 89, 90, 95, 102, 105, 107, 109, 110, 152), where it was generally addressed in the discussion. Of these records, two (4% of records reporting appropriateness) determined that ultrasound was not appropriate for scar measurement. The first stated that it was too inaccurate and complex; (86) and the second, which reported on initiatives within routine clinical practice, determined that the minimum resolution of the Diasonography ultrasonic scanner (Nuclear Enterprises, Edinburgh, UK) precluded its use in scars thinner than 3mm. (113)

The feasibility of ultrasound was reported in 12 journal articles (15% of included journal articles). (22, 24, 26, 46, 57, 70, 80, 119, 120, 124, 129) Five records considered ultrasound not feasible for scar measurements. The rationale presented included high-frequency 20 MHz ultrasound having an inadequate penetration depth; (26, 57) and ultrasound measurement and training of investigators requiring too much time (as reported in one record in phase 4 of the research-to-practice pipeline). (22, 119, 120) Another factor identified as precluding feasibility was the inability to consistently relocate the measurement site. (24) Conversely,

Page 21 of 39

Page 23 of 83

BMJ Open

one record reported ultrasound to be feasible in combination with Vancouver Scar Scale (VSS) measurement, (70) and another stated that ultrasound was able to distinguish between subcutaneous fat and muscle, which was interpreted by the authors of that record to mean that skin thickness measurements were accurate. (129) The majority (n = 11; 92%) of the records reporting feasibility were research initiatives in phase 2 or 3 of the research to practice pipeline. One record examined feasibility in the context of routine clinical practice (i.e., phase 4; figure 1), (119) where it was determined that ultrasound was not suitable for use in their twelve-year longitudinal study due to changes in staff, equipment and software over such a long time period, which introduced additional variables to the measurement process that were impossible to control. (119)

195 Factors influencing ultrasound images and measurement methods:

The only factor that was reported to influence the imaging and measurement methods was the measurement of scars with open wounds. This was reported in one record, which determined that ultrasound and ultrasound gel was unsuitable in this instance. The authors of that record suggested the use of a flexible transparent plastic wrap, which is placed over the measurement area prior to measurement with ultrasound. (24)

201 Reported strengths and limitations of the measurement methods:

The safety, practicality, objectivity, versatility, reliability and non-invasive nature of
ultrasound were all reported as strengths of the measurement method. (22, 27-29, 47, 50, 57,
61, 64, 77, 78, 80, 82, 87, 89, 95, 96, 105, 107, 109, 119, 123, 124, 129, 139, 148) When
compared to other subjective or clinical measurement methods (e.g., VSS) and 3D camera,
ultrasound was viewed as the superior measurement method of scar and skin thickness, due to
its improved accuracy, greater sensitivity to change and objectivity. (24, 64, 73, 116, 120)
The ability of ultrasound to differentiate between scarred and unscarred skin was also

highlighted (n = 4; 3%), (47, 60, 72, 122) as was the versatility of ultrasound in its ability to measure a variety of anatomical areas and be used with child participants (i.e., <18 years) (n = 2; 2%). (22, 149)

The poor correlation between ultrasound and histological thickness measurements, (86) and the established inverse relationship between ultrasound penetration depth and the resolution of superficial structures were identified as limitations of ultrasound in the measurement of scar thickness. (26, 27, 77, 80, 89, 113, 149) This may be an evidence gap worth exploring in more depth. One record, reporting on a longitudinal study that was conducted over twelve years, reported that the continuous development of ultrasound software and hardware over that time limited the usefulness of ultrasound. (119) Despite being reported elsewhere as acceptable (i.e., between one to five minutes (24, 27, 122)), one record reported that the timeconsuming nature of measurement and the requirement for assessors to be trained in the operation of, and techniques required for, ultrasonography was a limitation of the method. (120) Methodologically, concerns were raised around the pressure caused by application of the ultrasound transducer to the skin, and how that may influence thickness measurement. (61, 62, 123, 124) The size of the transducer head relative to the size of scars was also considered a potential limitation, as multiple measurements are required for quantification of larger scars. (57) Finally, it was recognised that there may be a difference between changes to the scar that can be measured by ultrasound, and what is felt and/or experienced by the patient. (75, 80, 126, 127) It was suggested that changes that are detectable by ultrasound may be smaller than those able to be detected by patients. In patients with burn scars, a minimum change in scar thickness of between 1 to 6 mm measured by ultrasound, has been reported to be required before a patient may report noticing any difference to their scar thickness. (24, 75) While further research is required to allow generalisation of these findings

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to other scar aetiologies, this indicates that a holistic approach to scar thickness using the

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46 47	250
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53 54	253
55 56	254
57 58	255
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patient's opinion as well as objective measurement through ultrasound may be beneficial.
Guidelines or frameworks used to guide the measurement methods:
No records reported using any guidelines or frameworks to inform their measurement
methods. One record utilised suggestions from The American Wound Healing Society to
support the measurement of contralateral, unscarred skin thickness on the same individual as
a control or comparator. (75)
Methodological Considerations:
Based on the ultrasound methods and outcomes identified in this review, a list of

methodological considerations have been compiled (Supplementary Table 6). These are
intended to guide the decision-making and methodological reporting of researchers and/or
clinicians undertaking scar or skin thickness ultrasound measurement.

245 **DISCUSSION:**

6 This review mapped the methods used in the published literature to measure traumatic scar 7 thickness using ultrasound across the research-to-practice translational pipeline. No record 8 reported their methods with sufficient detail to allow them to be independently replicated. 9 Overall, there was a lack of consistent rationale underpinning which skin layers (i.e., 0 epidermis, dermis and combined) were measured, and little consideration was given to the training and experience required by assessors. The included records mainly aligned with the 1 second and third phases of the research-to-practice pipeline (figure 1), with only three records 2 3 (2 articles and 1 abstract) reporting the use of ultrasound in routine clinical practice (phase 4). 4 (92, 113, 119). The paucity of records aligning with phase four studies (use in clinical 5 practice) suggests a translational gap from research to regular clinical practice. There are two likely explanations for this: 1) that ultrasound is most commonly used as an outcome measure 256

for research initiatives and is not regularly used to evaluate care once treatments are implemented into routine clinical practice; or 2) that use of ultrasound in routine clinical practice is not reported or evaluated, as routine clinical practice is rarely published. Searching of grey literature was conducted in an attempt to identify clinical practice documents, however none were located. Surveys of health service departments may be the best method of identifying ultrasound methods used in regular clinical practice as part of future research. While some records reported using additional subjective and objective measurement methods in addition to ultrasound, none used these methods to determine the criterion validity of the ultrasound for scar thickness measurement. This is another evidence gap that should be addressed. While efforts have been made to standardise ultrasound measurement procedures elsewhere in dermatology (including tumours, cancers, vascular anomalies, and systemic sclerosis (34, 35)), this same effort has not yet extended to the measurement of traumatic scarring. Methodological standardisation has the potential to increase confidence in the use of ultrasound as the basis of measurement-based care initiatives for clinical decision-making, allowing patient care and scar treatments to be tailored towards individual needs. (62, 147, 156) Standardising the core methodological components of ultrasound measurement of scar thickness, or at the very least, creating a standardised framework for methodological decision-making, may support implementation of ultrasound measurement into routine clinical practice, supported by strategies to overcome barriers to implementation at local sites. (157)This review identified novel insights into the identification of the composition of cutaneous scars using ultrasound, and highlighted the apparent lack of consistent understanding of, or

rationale behind, what scar thickness characteristics were being measured. Fibrosis is

Page 25 of 39

Page 27 of 83

BMJ Open

generally understood to be the primary cause of scar thickness through the deposition of excessive extracellular matrix proteins such as collagen. (158, 159) This has been confirmed through histological analysis, which has shown the presence of excess collagen and other extracellular matrix proteins in the dermis of hypertrophic scars. (160, 161) An additional method for assessing the effects of scarring on the dermis, as identified by one record in this review, (47) is through quantification of the presence and density of hair follicles. This quantification may serve as a method of differentiation between scarred and physiological skin, and may also serve as a measure of skin function. (47) What is less understood, and perhaps largely overlooked, is the function of the epidermis in scar thickness. In the one record identified in this review that directly report the measurement of the epidermis, the authors noted that the measurement quantified the presence of oedema. (55) This was further supported by two records that noted that the epidermis and dermis responded differently to treatment, (67, 71) indicating that there is likely a difference in the composition of the scar between these skin layers. Cutaneous oedema has been observed using high-frequency ultrasound in other pathologies, including atopic dermatitis and skin ageing, where it is characterised by the presence of a sub-epidermal low echogenic band (SLEB), a hyperechoic band at the dermoepidermal junction. (162) Understanding the interplay between epidermal oedema, dermal fibrosis and the presence and density of hair follicles may result in an increased understanding of the mechanisms and treatment responses of cutaneous scarring. With better understanding, more targeted scar treatments that inform a greater understanding of scar responsivity may arise.

Another important, but potential limiting factor for the use of ultrasound to measure scar thickness raised in this review is the training and/or experience required of assessors, and the ramifications this likely has on the reliability of measurements and interpretation. (163) This review identified 24 records where assessor experience was discussed, however none made

any recommendations on the optimal training and/or experience. Identifying the training requirements of assessors may prove an important step towards more widespread implementation of reliable ultrasound scar thickness measurement in research trials and as the basis for measurement-based care in routine clinical practice. (164) A panel of dermatological and ultrasound experts has previously recommended that a physician with a minimum of 300 examinations per year should hold responsibility for ultrasound measurements. (34) It has also been suggested that training existing members of clinical teams and standardising measurement method/s may be the most effective way to achieve minimum reliability standards under clinical conditions. This could allow measurement to be reliably conducted within an outpatient clinic setting by a number of healthcare providers assisting workflow, negating the requirement for patients to wait for an experienced radiographer. (24, 164) In the current review, reliability estimates were generally acceptable but were tested under research conditions. The diverse experience and expertise of assessors, where reported for the reliability estimates, means that the acceptable reliability results should be generalisable to most clinical teams, as therapists, doctors and nurses were all included. The cumulative sample size of all reliability studies also supports this generalisation; however each team should perform their own reliability estimates before conducting ultrasound thickness measurements.

324 Study Limitations:

Only articles available in English or with an English abstract were considered for inclusion
and data extraction, which may have resulted in the omission of eligible information. Data
extraction was completed on the English abstracts of two non-English articles that were
available electronically, however the non-English articles themselves were not available to
the authors, and thus could not be analysed. Based on the number of records included in this
review, however, it is unlikely that this would have impacted the review findings. It is

Page 27 of 39

Page 29 of 83

BMJ Open

acknowledged that methods reported in included abstracts may not be fully reproducible, due to their brevity. Thus, findings were reported separately to articles. An additional limitation was that authors of included records were not contacted to provide clarification or further information, as this was not feasible given the number of results identified. It should also be acknowledged that the included records were not designed to align with the specific aims of this review, which likely explains some of the lack of reporting on outcomes of interest in our review, particularly clinical, health service and implementation outcomes. Furthermore, as this review relied on published information (including grey literature), routine practices employed within organisations may not have been considered and unpublished industry sponsored reports may not have been identified. It is also important to consider the limitations of ultrasound itself for the holistic quantification of cutaneous scarring. Ultrasound transducers are generally small, meaning that it is difficult to assess the entirety of a scar, necessitating multiple measurements. (165) Additionally, thickness is often not the only scar parameter of clinical or research interest. It has therefore been recommended that multi-modal measurement techniques are employed, which include both subjective and objective measurements. (166, 167) However, use of these methods may be challenging in routine clinical practice, due to the length of time and training required. Thus, feasibility and implementation outcomes are of importance in evaluating measurement-based care initiatives involving ultrasound alone or multimodal measurement tools in scar care practice – a field in its infancy based on this review. **Future Directions:**

352 It is intended that the results of this review will be used to inform the creation of a Delphi
353 consensus study, leading to the formation of a guideline for the measurement of traumatic
354 scar thickness using ultrasound. This guideline can then be used by researchers and clinicians

to standardise the measurement of scars. In preparation for this study, we have provided a list of methodological considerations for assessors or practitioners when planning to conduct scar thickness measurements with ultrasound (Supplementary Table 6). Future research could also investigate aspects that were beyond the scope of this review including factors influencing the implementation of ultrasound-based care initiatives, strategies to support implementation, and how research-based initiatives could be applied in practice. Further studies are needed that compare SDCs to SEMs to interpret reliability estimates to confirm our interpretation that ultrasound may have the ability to detect true change or variance in scar thickness above measurement error, which was based on the SDC reported by a single study. Our interpretation is supported by mostly acceptable reliability estimates of ultrasound thickness for other cutaneous conditions. (168, 169) Additional investigations should also be conducted to determine the criterion validity of ultrasound as a measure for scar thickness.

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371 COMPETING INTERESTS:

The authors declare no competing interests. The research presented in this publication was conducted as part of BM's PhD, and will be included in his thesis for submission to The University of Queensland.

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Page 29 of 39

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379	AUTHOR CONTRIBUTIONS
380	BM and ZT conceived the project after identifying this area as a knowledge gap in existing
381	literature. BM developed the research questions and study methodology, conducted the
382	literature search, screened all articles and extracted data. Record screening and data
383	extraction was completed by BM, MS, TM, and TR, with additional extraction completed by
384	BD to assess consistency. MS, TM, TR and RK provided advice to BM on the clinical
385	implications of ultrasound measurement. MS, RK and ZT contributed to the supervision of
386	BM as a PhD student. BM drafted the paper, and ZT and MS provided critical appraisal of
387	the drafted manuscript, with further advice provided by TM, TR, BD and RK.
388	DATA SHARING STATEMENT:
389	Not applicable
390	ETHICS APPROVAL STATEMENT:
391	This study does not involve human participants. No ethics approval was required.
392	FIGURE LEGENDS:
393	Figure 1: Research to clinical practice pipeline.
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BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

Supplementary Box 1. Full search strategy for Ovid MEDLINE.

((ultrasound.ti,ab. OR ultra sound.ti,ab. OR sonograph*.ti,ab. OR ultrasonic.ti,ab. OR high-frequency.ti,ab. OR high frequency.ti,ab. OR hfus.ti,ab. OR ultrasonog*.ti,ab. OR exp Ultrasonography/)

AND

((skin.ti,ab. OR epiderm*.ti,ab. OR derm*.ti,ab. OR cutaneous.ti,ab OR scar*.ti,ab OR keloid*.ti,ab OR cicatri*.ti,ab OR exp Skin/ OR exp Dermatology/ OR exp Cicatrix/)

AND

(thickness*.ti,ab. OR thicken*.ti,ab. OR depth.ti,ab. OR volume.ti,ab. OR height.ti,ab. OR vancouver scar scale.ti,ab)

ADJ10

(measure*.ti,ab. OR quantif*.ti,ab. OR calculat*.ti,ab OR estimat*.ti,ab OR assess*.ti,ab. OR determin*.ti,ab. OR evaluat*.ti,ab OR imag*.ti,ab OR exam*.ti,ab)))

NOT (exp animals/ NOT exp humans/)

Legend: ab, abstract (searches the abstract of the publication); adj10, adjacency (search terms must be located within 10 words of one another); exp, explode (used to include all subheadings when searching MeSH headings); ti, title (searches the title of the publication)



BMJ Open

BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

Supplementary Figure 1. Preferred Reporting Items for Systematic reviews and Meta-

Analyses (PRISMA) flow diagram for this study.



BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

Supplementary Table 1: Extraction categories and fields

Extraction category	Extraction field
Publication details	First author
	Year of publication
	Title of publication
	Country (first author)
	Country (study)
	Country (recruited)
	Publication type (e.g., peer-reviewed journal article, abstract)
	Journal name
	Corresponding author contact details
	Funding source (e.g., commercial, non-commercial)
	Use of scar thickness measurement (e.g., longitudinal study, response to
	treatment)
Study details	Aim/objective
	Research questions
	Target population/topics
	Study design (e.g., RCT, mixed methods)
	Data and analysis (i.e., statistical methods)
	Removal of scar treatments before ultrasound measurement (e.g., length of
	time before measurement)
	Reason for measurement (e.g., research, clinical initiative)
	Inclusion/exclusion criteria
	Dates of data collection
	Ultrasound thickness collection methods (e.g., direct collection, collected
	from medical records)
	Contralateral/unaffected/comparator skin thickness measurement
	Other methods used
	Use of guidelines/frameworks for measurement methods
	How previously published methods/guidelines were used
	Research pipeline stage
	Setting (e.g., inpatient/outpatient clinics)
	Scar type (e.g., burn scar, surgical scar)
Participant details	Number of participants
	Population type (e.g., adult/paediatric)
	Gender ratio
	Patient involvement in thickness determination
	How patients were involved in thickness determination
Ultrasound methods	Ultrasound mode
	Device name and manufacturer
	Frequency used
	Number of measurements taken
	What did researchers report they were measuring (e.g. fibrosis ordema)
	Anatomical locations/functional measurement units measured
	Patient orientation
	Illtrasound transducer orientation
	Methods used to prevent skin compression
	Measurement site relocation strategies
	Type of skin measurement (i.e. enidermis/dermis/combined)
	Measurer training
Psychometric properties*	Reliability
r sychometric properties	Measurement error
Feasibility [†] outcomes	Time taken for measurement

BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

2		
3		Availability of measurement method
4		Ease of administration
5		Number of steps required
6		Number of people required to conduct measurements
7		Considerations for special populations
8	Implementation [‡] outcomes	Accentability
9	implementation outcomes	Adoption
10		Appropriateness
11		Cost
12		Eost
13		Fidality
14		Fluenty
15	Strongthe and limitations of	Sustainability
16	Strengths and minitations of	
17	measurement methods	
18		Barriers
19		Enablers
20	Findings	Ultrasound-related findings
21	[*] Psychometric properties as outl	ined in the COSMIN Risk of Bias tool to assess the quality of studies on
22	reliability or measurement error	of outcome measurement instruments ¹
23	Feasibility outcomes as per Prin	nsen <i>et al.</i> ²
24	[‡] Implementation outcomes as pe	er Proctor <i>et al.</i> ³
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Supplementary Table 2. Characteristics of records included in this review. Studies are listed alphabetically by author within the translational
pipeline phase.

First Author (year)	Country of Publication	Funding Sources	Sample Size (n)	Population Type	Scar Aetiology	Translational Pipeline Phase [*]
Journal articles						
Agabalyan (2017)	Canada	Non-commercial	10	Adult	Not specified	2
Alsharnoubi (2018)	Egypt	No funding	15	Paediatric	Burn	2
Alsharnoubi (2018)	Egypt	Not reported	15	Paediatric	Burn	2
Alshehari (2015)	Egypt	Not reported	30	Not reported	Mixed	2
Blome-Eberwein (2012)	United States	Non-commercial	16	Paediatric & adult	Burn	2
Blome-Eberwein (2016)	United States	Not reported	36	Adult	Not specified	2
Blome-Eberwein (2019)	United States	Non-commercial	19	Adult	Burn	2
Cai (2019)	China	Non-commercial	51	Adult	Not specified	2
Candy (2010)	Hong Kong	Not reported	17	Adult	Not specified	2
Chan (2004)	China	Non-commercial	56	Paediatric & adult	Burn	2
Chang (2014)	Taiwan	Non-commercial	60	Paediatric & adult	Surgical (cleft	2
-					lip repair)	
Cho (2014)	Korea	Non-commercial	146	Not reported	Burn	2
Deng (2019)	China	Not reported	20	Adult	Not specified	2
Deng (2021)	China	No funding	31	Adult	Not specified	2
Deng (2021)	Hong Kong and China	Non-commercial	45	Adult	Not specified	2
Dunkin (2007)	England	Non-commercial	113	Adult	Surgical (dermal	2
					scratch)	
Elrefaie (2020)	Not specified	Not reported	22	Paediatric & adult	Not specified	2
Fabbrocini (2016)	Not specified	Not reported	20	Adult	Mixed	2
Fraccalvieri (2011)	Italy	No funding	5	Adult	Mixed	2
Fraccalvieri (2013)	Italy	Not reported	3	Paediatric & adult	Mixed	2
Gee Kee (2016)	Australia	Commercial	43	Paediatric	Burn	2
Issler-Fisher (2021)	Australia	Commercial	187	Adult	Burn	2
Joo (2020)	Korea	Non-commercial	48	Adult	Not specified	2
Lacarrubba (2008)	Not specified	Not reported	8	Paediatric & adult	Mixed	2

BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

I (2005)	11 17		100		P
Lau (2005)	Hong Kong	Not reported	100	Paediatric & adult	Burn
Lee (2019)	United Kingdom	Non-commercial	55	Adult	Burn
Lee (2020)	United Kingdom	Non-commercial	55	Adult	Burn
Li (2013)	China	Non-commercial	7	Adult	Burn
Li (2020)	China	Not reported	21	Paediatric & adult	Mixed
Li (2021)	China	Non-commercial	165	Paediatric	Mixed
Li (2021)	China	Non-commercial	105	Adult	Burn
Li-Tsang (2006)	Not specified	Non-commercial	45	Adult	Not specified
Li-Tsang (2010)	China	Non-commercial	104	Paediatric & adult	Mixed
Mamdouh (2021)	Egypt	Not reported	40	Adult	Not specified
Meirte (2016)	Belgium	Non-commercial	9	Adult	Burn
Miletta (2021)	United States	Non-commercial	29	Paediatric & adult	Burn
Nedelec (2019)	Canada	Non-commercial	70	Adult	Burn
Nedelec (2020)	Canada	Non-commercial	51	Adult	Burn
Nicoletti (2015)	Italy	Not reported	27	Paediatric & adult	Surgical (scar
	-				reconstruction)
Niessen (1998)	The Netherlands	Commercial & Non-	145	Paediatric & adult	Surgical (breast
		commercial			reduction)
Reinholz (2020)	Germany	No funding	25	Adult	Mixed
Schwaiger (2018)	Germany	No funding	15	Adult	Mixed
van den Kerckhove	Belgium	Not reported	60	Adult	Burn
(2005)	C	1			
van der Veer (2010)	The Netherlands	Non-commercial	44	Adult	Surgical
					(cardiothoracic
					surgery)
Wang (2009)	China	Non-commercial	22	Adult	Burn
Wiseman (2020, 2021)	Australia	Commercial & Non-	153	Paediatric	Burn
		commercial			
Xuan (2021)	Not specified	Not reported	72	Not reported	Not specified
Yim (2010)	Korea	No funding	31	Paediatric & adult	Burn
Zadkowski (2016)	Not specified	Not reported	47	Paediatric	Burn
Avetikov (2018)	Not specified	Not reported	50	Paediatric & adult	Not specified
	riet speemed	ristiepoited	50		rest op control

BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

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Chae (2016)	Korea	Non-commercial	23	Adult	Not specified
Cheng (2001)	Hong Kong	Not reported	58	Paediatric	Burn
Danin (2012)	France	Not reported	22	Paediatric & adult	Burn
Fong (1997)	Not specified	Not reported	16	Paediatric & adult	Burn
Gankande (2014)	Australia	Non-commercial	30	Adult	Burn
Ge (2022)	China	Not reported	21	Paediatric & adult	Mixed
Guo (2020)	China	Non-commercial	87	Paediatric & adult	Not specified
Huang (2017)	Taiwan	Not reported	1	Adult	Burn
Huang (2020)	China	Non-commercial	43	Adult	Not specified
Huang (2021)	Taiwan	Not reported	5	Adult	Burn
Issler-Fisher (2017)	Australia	No funding	47	Paediatric & adult	Burn
Issler-Fisher (2020)	Australia	No funding	78	Adult	Burn
Katz (1985)	United States	Not reported	4	Not reported	Burn
Kemp Bohan (2021)	United States	No funding	21	Not reported	Burn
Kim (2018)	Not specified	Not reported	148	Not reported	Burn
Li (2018)	China	Non-commercial	34	Adult	Burn
Li-Tsang (2005)	China	Non-commercial	101	Adult	Surgical
					(orthopaedic
					surgery)
Lobos (2017)	Not specified	Not reported	35	Paediatric & adult	Not specified
Nedelec (2008)	Canada	Non-commercial	32	Adult	Burn
Nedelec (2014)	Not specified	Non-commercial	46	Adult	Burn
Reinholz (2016)	Not specified	Commercial	8	Adult	Not specified
Simons (2017)	Australia	Non-commercial	49	Paediatric	Burn
Soykan (2014)	The Netherlands	Non-commercial	87	Adult	Surgical
					(cardiothoracic
					surgery)
Timar-Banu (2011)	Canada	Non-commercial	30	Adult	Mixed
Ud-Din (2019)	United Kingdom	Non-commercial	62	Adult	Not specified
van den Kerckhove	Not specified	Not reported	6	Adult	Burn
(2003)			Ũ		

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BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

3 4	Wang (2010)	Australia	Commercial & Non- commercial	21	Paediatric	Burn	3
5	Wood (1996)	Not specified	Not reported	1	Paediatric	Burn	3
0 7	Yeol Lee (2022)	Korea	Non-commercial	16	Adult	Mixed	3
, 8	Berry (1985)	Not specified	Commercial	16	Paediatric & adult	Burn	4
9	Engrav (2010)	Not specified	Commercial & Non-	67	Paediatric & adult	Burn	4
10	g (· · ·)		commercial				-
11	Abstracts						
12 13	Agabalyan (2016)	Not specified	Non-commercial	10	Not reported	Burn	2
13	Bajouri (2018)	Not specified	Not reported	20	Not reported	Burn	2
15	Blome-Eberwein (2011,	Not specified	Not reported	16	Paediatric & adult	Mixed	2
16	2012)		I.				
17	Blome-Eberwein (2014)	Not specified	Not reported	66	Not reported	Burn	2
18	Cho (2012)	Not specified	Not reported	60	Paediatric & adult	Burn	2
19	Comstock (2018)	Not specified	Not reported	1	Adult	Burn	2
20 21	Cooper (2021)	Not specified	Not reported	25	Not reported	Burn	2
21	El-Zawhary (2007)	Not specified	Not reported	57	Not reported	Mixed	2
23	Jacobs (2016)	Not specified	Not reported	6	Paediatric & adult	Burn	2
24	Jang (2009)	Not specified	Not reported	20	Not reported	Not specified	2
25	Kim (2009)	Not specified	Not reported	5	Paediatric & adult	Burn	2
26	Li-Tsang (2010)	Not specified	Not reported	45	Not reported	Not specified	2
2/	Li-Tsang (2011)	Not specified	Not reported	4	Not reported	Not specified	2
20 29	Maari (2017)	Not specified	Non-commercial	12	Not reported	Not specified	2
30	Moortgat (2020)	Not specified	Not reported	10	Not reported	Burn	2
31	Nedelec (2018)	Not specified	Not reported	60	Not reported	Burn	2
32	Peters (2018)	Not specified	Not reported	5	Not reported	Burn	2
33	Siwy (2016)	Not specified	Non-commercial	15	Not reported	Burn	2
34	Tu (2014)	Not specified	Not reported	59	Not reported	Not specified	2
35	Ud-Din (2017)	Not specified	Not reported	20	Not reported	Surgical (tissue	2
37	× /	1	1		1	biopsies)	
38	Anthonissen (2015)	Not specified	Not reported	N.R.	Not reported	Burn	3
39	Bezugly (2014)	Not specified	Not reported	103	Not reported	Mixed	3
40		1	1		L		

BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

Bezugly (2019)	Not specified	Not reported	438	Not reported	Not specified	3
Blome-Eberwein (2012)	Not specified	Not reported	19	Adult	Burn	3
Du (2006)	Not specified	Not reported	1	Adult	Burn	3
Edgear-Lacoursière	Canada	Not reported	44	Not reported	Burn	3
(2022)		-		-		
George (2019)	Not specified	Not reported	11	Not reported	Burn	3
Li (2016)	Not specified	Not reported	34	Not reported	Burn	3
Seo (2011)	Korea	Not reported	48	Not reported	Burn	3
Timina (2013)	Not specified	Not reported	49	Paediatric & adult	Not specified	3
Ud-Din (2017)	Not specified	Not reported	20	Not reported	Surgical (tissue	3
	5	-		-	biopsies)	
Ud-Din (2018)	Not specified	Not reported	62	Not reported	Surgical (tissue	3
		-		-	biopsies)	
Zuccaro (2019)	Canada	Not reported	13	Paediatric	Burn	3
Zuccaro (2021)	Not specified	Not reported	20	Paediatric	Burn	3
Zuccaro (2021)	Canada	Non-commercial	20	Paediatric	Burn	3
Cho (2012)	Not specified	Not reported	30	Not reported	Burn	4

Legend: Paediatric: measurement of patients under the age of 18; Adult: measurement of patients aged 18 years or older; N.R.: Not reported; Burn: scars caused by thermal, chemical or friction injury; Surgical: scars caused by surgical procedures (including biopsies); Mixed: participant scars caused by mixed trauma (e.g., burn and acne)

Footnotes: *Stage in the research to clinical practice translational pipeline, based on the Australian Government Department of Health and Aged Care⁴
Page 51 of 83

BMJ Open

First Author (year)	Ultrasound Type	Ultrasound Frequency (MHz)	Measurement Parameters	Scar Characteristic Measured	Scar Relocation
Journal articles					
Agabalyan (2017)	High-frequency	20	Epidermal, dermal & combined	N.R.	Not relevant – single measurement
Alsharnoubi (2018)	Midrange ultrasound	N.R.	N.R.	Fibrosis	N.R.
Alsharnoubi (2018)	Midrange ultrasound	N.R.	N.R.	Fibrosis [†]	N.R.
Alshehari (2015)	N.R.	N.R.	Maximum elevation above normal skin	N.R.	N.R.
Avetikov (2018)	B-mode	N.R.	Combined epidermal & dermal	N.R.	Not relevant – single measurement
Berry (1985)	N.R.	N.R.	N.R.	N.R.	N.R. [‡]
Blome- Eberwein (2012)	B-mode	N.R.	Combined epidermal & dermal [§]	N.R.	N.R. [‡]
Blome- Eberwein (2016)	High-frequency	50	N.R.	Fibrosis [†]	N.R. [‡]
Blome- Eberwein (2019)	High-frequency	35	Dermal	Fibrosis, hair follicle density	N.R.
Cai (2019)	High-frequency	50	Dermal	N.R.	N.R. [‡]
Candy (2010)	B-mode	N.R.	N.R.	N.R.	Scar boundaries traced
Chae (2016)	N.R.	N.R	Combined epidermal & dermal	N.R.	Not relevant – single measurement
Chang (2014)	N.R.	12	N.R.	N.R.	N.R.

Chan (2004)	N.R.	N.R.	N.R.	N.R.	Tracing
Cheng (2001)	B-mode	5-10	Combined epidermal & dermal	N.R.	Tracing & cutting out paper Photographs
Cho (2014)	High-frequency	7.5	N.R.	N.R.	N.R.
Danin (2012)	B-mode	20	Epidermal & dermal	N.R.	N.R.
Deng (2019)	N.R.	N.R.	N.R.	N.R.	N.R.
Deng (2021)	Colour Doppler	4-15	Dermal	Fibrosis [†]	N.R.
Deng (2021)	B-mode	8-12	Epidermal & dermal	Fibrosis [†]	Photographs
Dunkin (2007)	High-frequency	N.R.	N.R.	Fibrosis & oedema [†]	Measurements taken at set linear distances along scar
Elrefaie (2020)	High-frequency	13	N.R.	Fibrosis & oedema [†]	N.R [‡]
Engrav (2010)	N.R.	N.R.	N.R.	N.R.	N.R.
Fabbrocini (2016)	N.R.	N.R.	N.R.	Fibrosis & oedema [†]	N.R [‡]
Fong (1997)	B-mode	7.5	N.R.	Fibrosis [†]	Tracing
Fraccalvieri (2013)	High-frequency	7-10 & 10-13	N.R.	Fibrosis & oedema [†]	N.R.
Fraccalvieri (2011)	High-frequency	10-13	Combined epidermal & dermal	Fibrosis [†]	N.R.
Gankande (2014)	High-frequency	20	Combined epidermal & dermal	N.R.	Scar marked & photographed
Ge (2022)	N.R.	N.R.	N.R.	N.R.	N.R.
Gee Kee (2016)	B-mode	8-18	Combined epidermal & dermal	N.R.	Transducer in centre of original burn site where no scar present
Guo (2020)	N.R.	2-15 & 4-15	Combined epidermal & dermal ^c	Fibrosis [†]	Thickest site on peripheral regions
Huang (2017)	N.R.	N.R.	Combined epidermal & dermal	N.R.	Marked & linear measurements from bony landmarks

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Huang (2021)	B-mode	5-12	N.R.	Oedema [†]	Not relevant – single
Huang (2020)	B-mode	5-12	Combined epidermal & dermal	N.R.	N.R.
Issler-Fisher (2021)	N.R.	N.R.	N.R.	N.R.	Photograph & measurement of thickest area
Issler-Fisher (2020)	N.R.	N.R.	N.R.	N.R.	N.R.
Issler-Fisher (2017)	N.R.	N.R.	N.R.	Fibrosis [†]	Scar mapped with drawing Thickest area measured
Joo (2020)	N.R.	N.R.	N.R.	Fibrosis [†]	N.R.
Katz (1985)	B-mode	10	Combined epidermal & dermal	N.R.	N.R.
Kemp Bohan (2021)	High-frequency	12	N.R.	Fibrosis [†]	Tracing – thickest area & adjacent landmarks marked
Kim (2018)	N.R.	22	Combined epidermal & dermal	N.R.	Not relevant – single measurement
Lacarrubba (2008)	B-mode	20	Combined epidermal & dermal	N.R.	N.R.
Lau (2005)	Tissue Ultrasound Palpation System	5 (burn) & 10 (surgical)	N.R.	N.R.	Tracing – most severe/prominent site
Lee (2020)	High-frequency	20	Combined epidermal & dermal	Fibrosis [†]	Not relevant – single measurement
Lee (2019)	High-frequency	20	Combined epidermal & dermal	Fibrosis [†]	Marked with pen
Li (2013)	High-frequency	12	Combined epidermal & dermal	Fibrosis [†]	Tracing
Li (2020)	N.R.	10	N.R.	Fibrosis [†]	N.R.
Li (2021)	High-frequency	20	N.R.	N.R.	Thickest area
Li (2021)	High-frequency	20	N.R. [§]	Fibrosis [†]	Thickest area
Li (2018)	N.R.	N.R.	Combined epidermal & dermal	N.R.	N.R.
Li-Tsang (2005)	Tissue Ultrasound Palpation System	N.R.	N.R.	N.R.	N.R.
Li-Tsang (2006)	B-mode	N.R.	N.R.	N.R.	N.R [‡]

Li-Tsang	B-mode	N.R.	N.R.	Fibrosis [†]	N.R.
(2010) Lobos (2017)	B-mode & colour	18	N.R.	Fibrosis [†]	Not relevant – single measurement
Mamdouh (2021)	High-frequency	N.R.	Combined epidermal & dermal [§]	Fibrosis [†]	N.R.
Meirte (2016)	High-frequency	22	Dermal	Fibrosis & oedema [†]	Marked with surgical pen, including boundaries of probe. Photograph of body position & probe location
Miletta (2021)	N.R.	50	N.R.	Fibrosis [†]	Tracing – worst scar
Nedelec (2014)	High-frequency	20	Combined epidermal & dermal	N.R.	Tracing including notable landmarks. Measurement site circled. Photograph
Nedelec (2008)	High-frequency	20	Combined epidermal & dermal	N.R.	Tracing including notable landmarks. Measurement site circled. Photograph
Nedelec (2019)	High-frequency	20	Combined epidermal & dermal	Fibrosis & oedema [†]	Tracing. Hole cut over measurement area
Nedelec (2020)	High-frequency	20	Combined epidermal & dermal	N.R.	Photograph
Nicoletti (2015)	N.R.	22	Epidermis to fascia	N.R.	N.R.
Niessen (1998)	B-mode	N.R.	N.R.	Fibrosis & oedema [†]	3cm border marked with tape – measurements lateral
Reinholz (2020)	B-mode	11	Combined epidermal & dermal	Fibrosis & oedema [†]	N.R.
Reinholz (2016)	B-mode	11	Combined epidermal & dermal [§]	Fibrosis & oedema [†]	N.R.
Schwaiger (2018)	B-mode	11	N.R.	Fibrosis & oedema [†]	N.R.
Simons (2017)	B-mode	8-18	Combined epidermal & dermal	N.R.	Tracing – scar & anatomical landmarks

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BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

2						
3	Soykan (2014)	N.R.	3-9	N.R.	Fibrosis [†]	N.R.
4	Timar-Banu	High-frequency	20	Combined epidermal & dermal	Fibrosis [†]	N.R.
5	(2001)				11010015	
6	(2001) Ud_Din (2019)	High-frequency	50	Combined enidermal & dermal	Fibrosis	Defined anatomical location
/	von den	Ligh frequency	20	Combined epidermal & dermal	N D	Test sites marked
0	Vall Util	Ingli-frequency	20	Combined epidermar & dermar	1 \.I\.	The state of the s
9 10	(2002)					Thermoplastic spinits created
10	(2003)	TT: 1 C	20		ND	with space for transducer
12	van den	High-frequency	20	Combined epidermal & dermal	N.K.	Test site boundaries marked
13	Kerckhove					& traced
14	(2005)					
15	van der Veer	N.R.	7.5	N.R.	Fibrosis [†]	Standardised linear
16	(2010)					measurement points
17	Wang (2009)	High-frequency	N.R.	N.R.	Fibrosis [†]	N.R.
18	Wang (2010)	B-mode	N.R.	Combined epidermal & dermal	N.R.	Tracing – scar & anatomical
19	U V V					landmarks
20	Wiseman	B-mode	N.R.	Combined epidermal & dermal	Fibrosis [†]	Centrally site of interest
21	(2020, 2021)					
22	(2020, 2021) Wood (1996)	B-mode	7 & 10	NR	NR	Transducer affixed to
23	(1990)	D mode	/ @ 10		10.10.	tracking arm
25	X_{uan} (2021)	High_frequency	20	NR	Fibrosist	N R
26	Xual (2021)	P mode	20	N.R. N D	N D	N.R. N D
27	(2022)	D-IIIOUE	/-10	N.K.	IN.K.	IN.K.
28	(2022)	TT 1 C	10	ND	ND	ND
29	Y1m (2010)	High-frequency	12	N.K.	N.K.	N.K.
30	Zadkowski	B-mode	N.R.	Combined epidermal & dermal	N.R.	N.R.
31	(2016)					
32	Abstracts					
33						
24 25	Agabalyan	N.R.	20	Epidermal, dermal & combined	N.R.	N.R.
36	(2016)					
37	Anthonissen	N.R.	22	Epidermal & dermal	N.R.	N.R.
38	(2015)			•		
39	Bajouri (2018)	High-frequency	N.R.	Epidermal & dermal	N.R.	N.R.
40		-o 1		r ······		

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Bezugly (2019)	High-frequency	22, 33 & 75	Epidermal & dermal	N.R.	N.R.
Bezugly (2014)	High-frequency	33 & 75	Epidermal & dermal	N.R.	N.R.
Blome-	N.R.	N.R.	N.R.	N.R.	N.R.
Eberwein					
(2011, 2012)	High fraguency	ND	ND	Fibrosia	ND
Eberwein	High-frequency	N.K.	N.K.	FIDIOSIS	N.K.
(2012)					
Blome-	High-frequency	N.R.	N.R.	N.R.	N.R.
Eberwein		0 _k			1 11 11
(2014)					
Cho (2012)	N.R.	N.R.	N.R.	N.R.	N.R.
Cho (2012)	N.R.	N.R.	N.R.	N.R.	N.R.
Comstock	N.R.	N.R.	N.R.	N.R.	N.R.
(2018)					
Cooper (2021)	N.R.	N.R.	N.R.	N.R.	N.R.
Du (2006)	B-mode	15	N.R.	N.R.	N.R.
Edgar-	N.R.	N.R.	N.R.	N.R.	N.R.
Lacoursière					
(2022)	ND	ND			ND
El-Zawhary	N.R.	N.R.	N.R.	N.R.	N.R.
(2007)	ND	ND	ND	ND	ND
George (2019)	N.K.	N.R.	N.K.	N.R.	N.K.
Jacobs (2016)	N.K.	N.K.	N.K.	N.K.	N.K.
Jang (2009) Kim (2000)	N.K. N D	N.K. N D	N.K.	N.K.	N.K. N D
KIIII (2009)	N.K. N D	IN.K. N D	N.K. N.D	N.K. N D	N.K.
LI(2010)	IN.K. Tiagua Illtragound	IN.K. N D	N.K. N.D	N.K. N D	N.K.
L_1 - I sang	Delection System	N.K.	N.K.	N.K.	N.K.
(2011)	Tiggue Ultrogound	ND	N D	ND	ND
(2010)	Palnation System	IN. K .	IN.R.	1 N.I X.	IN. K .
(2010) Maari (2017)	N R	NR	NR	NR	NR
(2017)	11.11.	± 1.1\.	11.11.	11.11.	11.11.

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BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplen

Moortgat	High-frequency	N.R.	Dermal	N.R.	N.R.
(2020)					
Nedelec (2018)	N.R.	N.R.	N.R.	N.R.	N.R.
Peters (2018)	High-frequency	22	N.R.	N.R.	N.R.
Seo (2011)	N.R.	7.5	N.R.	N.R.	Thickest point
Siwy (2016)	N.R.	N.R.	N.R.	N.R.	N.R.
Timina (2013)	N.R.	20-40	N.R.	N.R.	N.R.
Tu (2014)	High-frequency	N.R.	N.R.	N.R.	N.R.
	ultrasound				
	biomicroscopy				
Ud-Din (2017)	N.R.	N.R.	N.R.	N.R.	N.R.
Ud-Din (2017)	High-frequency	50	N.R.	N.R.	N.R.
Ud-Din (2018)	High-frequency	N.R.	N.R.	Fibrosis [†]	N.R.
Zuccaro (2021)	N.R.	N.R.	N.R.	N.R.	N.R.
Zuccaro (2019)	B-mode	N.R.	N.R.	N.R.	N.R.
Zuccaro (2021)	B-mode	6-18	Combined epidermal & dermal	N.R.	Scar outlined &
					photographed

Legend: Scar relocation: Methods used by assessors to relocate the measured scar for sequential measurements; B-mode: brightness-mode ultrasound (< 20 MHz); High-frequency: high-frequency B-mode ultrasound (> 20 MHz); N.R.: Not reported

Footnotes: [†]Indirect reference made in record (e.g. in introduction or discussion); [‡]Photographs taken of the scar but not specified whether used for relocation; [§]Not stated in methods, so images provided in record used by authors of this review to provide subjective judgement

First author (year)	Objective measurement methods	Clinician-based rating scale	PROM
Journal articles			
Agabalyan (2017)	Histology	-	-
Alsharnoubi (2018)	Laser Doppler perfusion	VSS	-
Alsharnoubi (2018)	Laser Doppler perfusion	VSS	-
Alshehari (2015)	-	VSS	-
Avetikov (2018)		-	-
Berry (1985)	Transcutaneous oxygen measurement	Scar redness and hypertrophy	Scar redness and hypertrophy rating
	, and the second s	rating scale (0-5 Likert scale)	scale (0-5 Likert scale)
Blome-Eberwein (2012)	Doppler flowmeter – vascularity	VSS	POSAS-P
	Cutometer - pliability	POSAS-O	
	Semmes-Weinstein monofilament		
	Aesthesiometer testing set –		
	sensation		
Blome-Eberwein (2016)	Cutometer – pliability	VSS	POSAS-P
Bioline Eber wein (2010)	Dermaspectrometer – colour	POSAS-0	
	Semmes-Weinstein Aesthesiometer		
	Monofilament Testing Set –		
	sensation		
Blome-Eberwein (2019)	-	VSS	_
Cai (2019)	_	Clinical evaluation	_
Candy (2019)	Spectrocolorimeter – colour	VSS	_
Chae (2016)	Spectrophotometer – nigmentation	VSS	POSAS-P
Chae (2010)	Specifophotometer prementation	POSAS-O	100/10-1
Chang (2014)		VSS	
Chang (2014)	-	Photographic avaluation (0, 10	-
		VAS)	
Chan (2004)	Cutomator viscoalecticity	(AS)	
Chan (2004)	Spectrophotometer pigmentation	-	-
Chang (2001)	Specifophotometer – pigmentation	VSS	
Cheng (2001)	- Maxamatar colour	Treatment officient (0, 10, VAS)	- Itahing scale (0, 4 Likort scale)
Cho (2014)	viexanceier - coloni	Treatment enreacy (0-10 vAS)	number scale (0-4 Likeri scale)

Supplementary Table 4. Additional measurement methods used alongside ultrasound in included studies

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	Tewameter – trans-epidermal water		
	loss		
	Sebumeter – sebum		
	Cutometer – elasticity		
Danin (2012)	Cutometer – elasticity	VSS	-
Deng (2019)	DermaLab Combo – colour	POSAS-O	-
-	Dermoscopy – vascularity		
Deng (2021)	-	VSS	-
Deng (2021)	Doppler – blood perfusion	POSAS-O	POSAS
	Dermlite Foto IIPro – erythema		
Dunkin (2007)		-	-
Elrefaie (2020)	Ultrasound – echogenicity,	VSS	-
	compressibility & vascularity		
Engray (2010)	Durometer – hardness	Clinical appearance based on	_
	Chromameter – colour	photographs	
Fabbrocini (2016)	-	mVSS (vascularity, pigmentation,	_
1 4001001111 (2010)		pliability)	
Fong (1997)	Cutometer – elasticity	Clinical rating – colour change.	-
1 0118 (1227)	Culoniciti Chastienty	consistent itch hypersensitivity	
		blistering	
Fraccalvieri (2013)	Colour power Doppler –	VSS	
Theedivien (2013)	vascularisation	Visual analogue scale – pain and	
	vascularisation	itch	
Fraccalvieri (2011)	Histology	-	-
	Fchocontrastography –		
	neovascularisation		
Gankande (2014)	DermI ab combo $-$ erythema &	mVSS (some participants)	
Gankande (2014)	elasticity	m v 55 (some participants)	-
$G_{e}(2022)$		$POSAS_O$	POSAS
Ge (2022)	-	Subjective reports on patient	I OBAD
		range of movement	
$C_{22} K_{22} (2016)$	2D photography thistrass		
Gee Kee (2016)	SD photography – thickness	POSAS-O	POSAS
Guo (2020)	Ultrasound – blood flow grade	-	-
	Shear wave elastography – scar		
	stiffness		

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ear wave elastography – scar fness catrometer – firmness	- VSS POSAS-O VSS POSAS-O VSS POSAS-O VSS - - - Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	- POSAS-P POSAS-P Patient pain & itch scales POSAS-P Patient pain, itch & quality of life rating scales Pain severity (0-10 VAS) - - POSAS-P POSAS-P
catrometer – firmness	VSS POSAS-O VSS POSAS-O VSS POSAS-O VSS - - - Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	POSAS-P POSAS-P Patient pain & itch scales POSAS-P Patient pain, itch & quality of life rating scales Pain severity (0-10 VAS) - - - - POSAS-P POSAS-P
eatrometer – firmness	VSS POSAS-O VSS POSAS-O VSS POSAS-O VSS - - - Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	POSAS-P POSAS-P Patient pain & itch scales POSAS-P Patient pain, itch & quality of life rating scales Pain severity (0-10 VAS) - - - - POSAS-P POSAS-P
catrometer – firmness	POSAS-O VSS POSAS-O VSS POSAS-O VSS - - - Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	POSAS-P Patient pain & itch scales POSAS-P Patient pain, itch & quality of life rating scales Pain severity (0-10 VAS) - - - - POSAS-P POSAS-P
catrometer – firmness	VSS POSAS-O VSS POSAS-O VSS - - Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	POSAS-P Patient pain & itch scales POSAS-P Patient pain, itch & quality of life rating scales Pain severity (0-10 VAS) - - - - POSAS-P POSAS-P
eatrometer – firmness	POSAS-O VSS POSAS-O VSS - - Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	Patient pain & itch scales POSAS-P Patient pain, itch & quality of life rating scales Pain severity (0-10 VAS) - - - - POSAS-P POSAS-P
catrometer – firmness	VSS POSAS-O VSS - - Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	POSAS-P Patient pain, itch & quality of life rating scales Pain severity (0-10 VAS) - - - - POSAS-P POSAS-P
catrometer – firmness	POSAS-O VSS - - Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	Patient pain, itch & quality of life rating scales Pain severity (0-10 VAS) - - - - - POSAS-P POSAS-P
catrometer – firmness	VSS - - - Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	Pain severity (0-10 VAS) - - - - - POSAS-P POSAS-P
catrometer – firmness	- - - Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	- - - - POSAS-P POSAS-P
atrometer – firmness	- - Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	- - - - POSAS-P POSAS-P
	- Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	- - - POSAS-P POSAS-P
	- Clinical evaluation of lesion size VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	- - POSAS-P POSAS-P
	VSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	- - POSAS-P POSAS-P
	vSS mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	- POSAS-P POSAS-P
	mVSS (height, pliability, vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	POSAS-P POSAS-P
	vascularity, pigmentation) POSAS-O mVSS (height, pliability, vascularity, pigmentation)	POSAS-P
	POSAS-O mVSS (height, pliability, vascularity, pigmentation)	POSAS-P
	mVSS (height, pliability, vascularity, pigmentation)	POSAS-P
	vascularity, pigmentation)	
	POSAS-O	
crometer – tissue thickness	-	-
rce/torque sensor – load applied to		
r		
tometer – elasticity	VSS	Quality of life questionnaire
exameter – colour		
iCam PSI system and mexameter		
lood supply		
ser Doppler flowmetry – perfusion	VSS	-
ectrocolourimeter – scar colour	VSS	Pain & itch (0-10 VAS)
	VSS	Treatment satisfaction
ectrocolourimeter – scar colour	VSS	Pain & itch (VAS scale not specified
ectrocolorimeter – colour	VSS	Pain & itch (VAS)
	ce/torque sensor – load applied to ometer – elasticity xameter – colour Cam PSI system and mexameter ood supply er Doppler flowmetry – perfusion ctrocolourimeter – scar colour ctrocolourimeter – scar colour	ce/torque sensor – load applied to ometer – elasticity VSS xameter – colour Cam PSI system and mexameter ood supply er Doppler flowmetry – perfusion VSS ctrocolourimeter – scar colour VSS vSS ctrocolourimeter – scar colour VSS ctrocolourimeter – colour VSS

Page 61 of 83

 BMJ Open

Li-Tsang (2010)	Spectrocolorimeter – colour	VSS (pliability)	Pain & itch (10-point VAS)
Lobos (2017)	-	Modified Seattle Scar Scale	-
Mamdouh (2021)	-	VSS	Patient satisfaction (VAS)
Meirte (2016)	_	-	-
Miletta (2021)	Colourmeter – scar colour	Unclear, likely POSAS-O	Unclear, likely POSAS-P
1110tuu (2021)	Dermal torque meter – scar		Short Form 36 Quality of Life Survey
$N_{2} = 1 + 1 + 2 = (2014)$	Compliance		
Nedelec (2014)	Cutometer – elasticity	-	-
	Mexameter – colour		
Nedelec (2008)	Cutometer – elasticity Mexameter – colour	mVSS	-
Nedelec (2019)	Cutometer – elasticity	-	-
	Mexameter – colour		
Nedelec (2020)	Cutometer – elasticity	-	Pain & itch (10cm line VAS)
	Mexameter – colour		×
Nicoletti (2015)	-		-
Niessen (1998)	Histology		-
Reinholz (2020)	3D topographic imaging device	POSAS-O	Dermatology Quality of Life Index
			POSAS-P
Reinholz (2016)	Optical coherence tomography – thickness	POSAS-O	Dermatology Quality of Life Index POSAS-P
Schwaiger (2018)	3D topographic imaging device	-	-
Simons (2017)	3D camera – scar height	POSAS-O	-
Soykan (2014)	Slide calliper – dimensions	POSAS-O	POSAS-P
Timar-Banu (2001)	Metric ruler – dimensions	Validated 3-point scoring system	-
		for redness, hardness, itching &	
		pain	
Ud-Din (2019)	Optical coherence tomography –	-	-
	thickness		
	Histology		
van den Kerckhove (2005)	Chromameter – erythema	-	-
van der Veer (2010)	Slide calliper – dimensions	-	-
Wang (2009)	Histology	-	-
Wang (2010)	-	-	-

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Wiseman (2020, 2021)	-	POSAS-O	POSAS-P Numeric rating scale for itch Toronto Paediatric Itch Scale CH-9D BBSID
Wood (1996)	_	VSS	-
Xuan (2021)	Histology	-	-
Yeol Lee (2022)	Cutometer – elasticity	mVSS	-
× ,	Elastography		
Yim (2010)	Cutometer – elasticity	-	-
· · ·	Tewameter – trans-epidermal water loss		
	Mexameter – colour		
Zadkowski (2016)		VSS	-
Abstracts			
Agabalyan (2016)	Histology		-
Bajouri (2018)	-	VSS	-
Bezugly (2019)	Clinical or histopathological diagnosis		-
Bezugly (2014)	-	-	-
Blome-Eberwein (2011, 2012)	Doppler vascularity, elasticity and sensation	VSS	Pain and itching scale (0-10 Likert scale)
Blome-Eberwein (2012)	-	-	-
Blome-Eberwein (2014)	Doppler flowmeter – vascularity Cutometer – pliability Semmes-Weinstein monofilament	VSS	POSAS-P
	aesthesiometer testing set – sensation		
Cho (2012)	-	VSS	-
Cho (2012)	CK-MPA Multi-Probe adaptor – pigmentation, erythema and trans- epidermal water loss Cutometer – elasticity	-	-
Comstock (2018)	Computer-based tools – Thickness & pliability	Unclear, likely POSAS-O	Unclear, likely POSAS-P
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Page 63 of 83

 BMJ Open

Cooper (2021)	Colorimeter – pigmentation	Unclear, likely POSAS-O	Unclear, likely POSAS-P
Du (2006)	-	-	-
Edgar-Lacoursiere (2022)	Cutometer – elasticity	-	-
	Mexameter – colour		
El-Zawhary (2007)	Histology	-	-
George (2019)	-	-	-
Jacobs (2016)	Cutometer – pliability	POSAS-O	-
Lan a (2000)	Colorimeter – colour		
Jang (2009)	Mexameter – pigmentation	-	-
	Tewameter – trans-epidermai water		
	loss Sahumatan sahum		
	Sebullieler – sebulli		
	Lesen Deppler – parfusion		
$V_{im}(2000)$	Laser Doppier – perfusion Histology	VCC	
$\mathbf{K}_{1111}(2009)$	Spectrocolourimeter scor colour	VSS VSS	- Detionst report of pain & itch
Li(2010) Li Teong (2011)	spectrocolourimeter – scar colour	VSS (thickness plicbility and	Fatient report of pain & nen
LI-1 Salig (2011)	-	pigmontation)	-
\mathbf{L} i Teong (2010)	Histology	VSS	Salf report questionnaire
LI-1 sang (2010)	Spectrocolourimeter scor colour	v 33	Sen-report questionnaire
Maari (2017)	Cutometer electicity	191	
Widdi'i (2017)	Mexameter - nigmentation		-
Moortgat (2020)	Cutometer = elasticity	Unclear likely POSAS-O	Unclear likely POSAS-P
1001tgat (2020)	Chromameter – colour	Olicical, likely 1 OSAS-O	Olicical, likely 1 OSAS-1
	Tewameter – trans-enidermal water		
	Corneometer – hydration		
Nedelec (2018)	Cutometer – elasticity	_	_
	Mexameter – colour		
Peters (2018)	Cutometer - elasticity	POSAS-O	POSAS-P
	Colourimeter - colour		
Seo 2011	Cutometer – elasticity		
Siwy (2016)	Colourimeter – colour	-	SF-36 Quality of Life Measuremer
	Torque meter – pliability & elasticity		POSAS-P
Timina (2013)		_	

Tu (2014)	-	VSS	-	
Ud-Din (2017)	Laser perfusion imaging	-	-	
	Optical coherence tomography –			
	thickness			
	Histology			
Ud-Din (2017)	Optical coherence tomography –	-	-	
	thickness			
Ud-Din (2018)	Optical coherence tomography –	-	-	
	thickness			
	Histology			
Zuccaro (2021)	Multi-parameter skin analysis device	VSS	Unclear, likely POSAS-P	
	6	Unclear, likely POSAS-O		
Zuccaro (2019)	Acoustic radiation force impulse	-	-	
	ultrasound elastography			
Zuccaro (2021)	Acoustic radiation force impulse –	VSS	POSAS-P	
	stiffness	POSAS-O (did not include		
	DermLab Combo elasticity probe –	surface area and relief subscales)		
	elasticity			
	DermLab Combo colour probe –			
	colour			
Legend: (m)VSS: (Modified) Va	ancouver Scar Scale; POSAS: Patient and	Observer Scar Assessment Scale (PO	SAS-O: POSAS observer sca	ıle;
POSAS-P: POSAS patient scale)	; VAS: Visual Analogue Scale; CHU-9D:	Child Health Utility-9D; BBSIP: Bri	sbane Burn Scar Impact Profi	le
	, <u> </u>			Page 23 of 41
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Supplementary Table 5: Reliability of ultrasound methods reported in each included study

First Author (year)	Reliability Test & Measurement Error	Reliability & Measurement Error Test Statistics & Details
Inter-rater reliability		
Anthonissen (2015)	ICC; SEM	Epidermal – 0.297; 0.02mm
		Dermal – 0.991; 0.13mm
Chang (2014)	Pearson correlation	R=0.90, p<0.001
Dunkin (2007)	N.R.	N.R.
Fong (1997)	ICC	0.93, p=0.146
Gankande (2014)	ICC (95% CI)	Individual site:
		Rater 1 vs rater 2
		'Best scar' – 0.95 (0.92, 0.96)
		'Worst scar' – 0.95 (0.91, 0.97)
		'Normal skin' – 0.94 (0.91, 0.96)
		Rater 1 vs rater 3:
		'Best scar' $-0.86(0.78, 0.91)$
		'Worst scar' -0.91 (0.85, 0.95)
		'Normal skin' -0.92 (0.88, 0.95)
		Rater 2 vs rater 3:
		'Best scar' $= 0.93 (0.89, 0.95)$
		'Worst scar' $-0.96(0.92, 0.97)$
		'Normal skin' $= 0.95 (0.92, 0.97)$
		Average site:
		Rater 1 vs rater 2
		'Best scar' $= 0.97 (0.94, 0.99)$
		'Worst scar' $= 0.98 (0.94, 0.99)$
		(Normal skin' - 0.97 (0.93, 0.99))
		Rater 1 vs rater 3
		'Best scar' $= 0.90(0.77, 0.95)$
		(0.17, 0.93)
		(Normal skin' - 0.96 (0.92, 0.98))
		Rotar 2 vs rater 2
		'Best scar' $= 0.95 (0.88, 0.98)$
		(Normal skin2 - 0.98 (0.94, 0.99))
$L_{00}(2005)$	ICC	0.84 m < 0.01
Lau(2003)		0.04, p<0.01 "A acomtable to high"
Lee (2020)	ICC (05% CD) SEM	Acceptable to high
Lee (2019)	ICC (95% CI), SEM	$\frac{5021}{2}$
		$\begin{array}{c} \text{Single. 0.957} \\ (0.954-0.973) \\ \text{Averages 0.085} \\ (0.077, 0.001) \\ \end{array}$
		Average: 0.985 (0.977-0.991)
		SEM: 0.10 mm
		Unscarred SKIII: Single: 0.067 (0.040, 0.090)
		Single: 0.967 (0.949-0.980)
		Average: 0.989 (0.982-0.993)
N 1.1 (2000)		SEM: 0.04 mm
Nedelec (2008)	ICC (95% CI)	Most severe scar: 0.90 (0.84-0.95)
		Less severe scar: 0.91 (0.85-0.95)
		Donor site: 0.89 (0.82-0.94)
~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~		Normal skin: 0.85 (0.75-0.92)
Seo (2011)	N.R.	"High"
Simons (2017)	ICC (95% CI); SEM	Scar: 0.82 (0.7-0.89); 0.05 cm
		Normal skin: 0.33 (0.08-0.54); 0.03 cm
Van Den Kerckhove	ICC (95% CI); SEM	<u>One day:</u>
(2003)		0.88 (0.81-0.95); 0.29 mm

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		<u>Day-to-day:</u> 0.04 (0.00, 0.08): 0.21mm
Intra rator reliability		0.94 (0.90-0.98), 0.2111111
Anthonissen (2015)	ICC: SEM	Enidermal 0.800: 0.01mm
Anthomssen (2013)	ICC, SEIVI	Dermal = 0.001; 0.12mm
C_{an} transfer (2014)		Definal = 0.991, 0.1511111
Gankande (2014)	ICC (93% CI)	$\frac{1}{10000000000000000000000000000000000$
		Worst scar $= 0.92 (0.88, 0.95)$
	ND	Normal skin' $-0.86(0.81, 0.89)$
Gee Kee (2016)	N.R.	N.R.
Lau (2005)	ICC	Intra-rater: 0.98, p<0.01
Lee (2019)	ICC (95% CI)	<u>Scar:</u>
		Single: 0.951 (0.871-0.987)
		Average: 0.983 (0.953-0.966)
		SEM: 0.10 mm
		Unscarred skin:
		Single: 0.948 (0.881-0.976)
		Average: 0.982 (0.954-0.993)
		SEM: 0.04 mm
Li (2013)	ICC	0.89
Seo (2011)	N.R.	"High"
Simons (2017)	ICC (95% CI): SEM	Scar: 0.95 (0.91-0.97): 0.02 cm
		Normal skin: $0.61 (0.41-0.75)$: 0.02 cm
Van Den Kerckhove (2003)	ICC (95% CI); SEM	0.98 (0.97-0.99); 0.11mm
Wang (2010)	SE	Peak: 0.032
		3 months: 0.018
		6 months: 0.399
		9 months: 0.353

Abbreviations used in tables: N.R.: Not reported; ICC: Intraclass Correlation Coefficient; 95% CI: 95% Confidence Interval; SEM: Standard Error of Measurement; SE: Standard Error

Summary of findings for measurement error:

The reported inter-rater SEM measurements for the combined (i.e., epidermal and dermal) thickness measurement of scars was reported in two records as 0.11 mm^5 and 0.5 mm.⁶ The inter-rater SEM for the combined thickness measurement of unscarred skin was also calculated in one record (SEM = 0.3 mm).⁶ The inter-rater SEM was calculated in one record for the measurement of epidermal (SEM = 0.02 mm) and dermal (0.13) measurements⁷, and one record reported only the dermal SEM for scar thickness (SEM = 0.1 mm) and unscarred skin (0.04 mm).⁸ The intra-rater SEM for the combined thickness measurement of scarred skin ranged from 0.18 mm to 0.52 mm, and was measured at 0.2 mm for unscarred skin in one record.⁶ One record reported the intra-rater SEM for epidermal (0.01 mm) and dermal (0.12 mm),⁷ and one record reported the intra-rater SEM for dermal scar (0.1 mm) and unscarred skin (0.04).⁸

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Supplementary Table 6. Methodological considerations for researchers and/or clinicians undertaking measurement of scar thickness using ultrasound.

Consideration	Details & examples of considerations	Publications in our review addressing the consideration	Details reported in included review records
Preventing skin	Using standoff methods	6,9-13	- Use of ultrasound gel to prevent contact
compression	(e.g., ultrasound gel,		between ultrasound transducer and skin surface
during	water bath) to prevent		to minimise compression applied by direct
measurement	transducer touching the		application of transducer ^{6,9-12}
	skin		- Silicone pad placed underneath transducer ¹³
	Application of minimal	14-18	- Transducer held to maintain minimal pressure
	pressure by transducer		on scar 14,15,17
			- Training users to apply minimal force on
			transducer to prevent scar or skin distortion ^{10,18}
	Deliberately	19-21	- Measurement of thickness with and without
	compressing skin to		compression with transducer ^{13,21}
	quantify scar		- Thickness measurements taken using TUPS,
	compressibility		which uses controlled and metered compression
Oniontino the	Orienting the nations	8.18.22	Definit suring through out measurement to
Orienting the	during massurement		- Patient supine throughout measurement to
patient	(a g upright suping		now measurement to be taken in the same
	(e.g., upright, suprie,		position
	Maintaining patient	9	- Patients asked to hold breath during
	stillness during		measurement of scars on the chest to allow
	measurement		shear-wave ultrasound ⁹
Placing	Orientating ultrasound	23	- Direction of transducer recorded to ensure
ultrasound	transducer [e.g.,		consistency ²³
transducer	vertical (superior to		5
	inferior/cranial to		
	caudal), horizonal		
	(medial to lateral)]		

	Orienting the transducer in relation to the scar (e.g.,	9,15,17,18,22,24-26	- Transducer oriented perpendicular to the skin surface to provide optimal image ^{9,15,18,22,24-26}
	perpendicular) Measuring difficult/tight areas (e.g., axillae or other	6	- Exclusion of fingers and toes in paediatric measurements due to size of measurement area and thin skin ⁶
Relocating scars for	Mapping measurement area (e.g., tracing,	6,12,16,18,20,22,27-32	- Scars traced using translucent paper 18,20,22,27,29,31,32
longitudinal measurement	schematic diagram)		 Scars and surrounding anatomical landmarks traced using translucent paper ¹⁶ Scar mapped on transparent paper, which was then cut out ²⁸ Scar mapped with drawing, no elaboration provided ³⁰ Scars traced using Visitrak (Smith & Nephew Medical Limited, Franke, 0.612)
	Photographing measurement area	24,26,33	- Assessed area marked and photograph taken in initial consultation ^{24,33}
	Measuring specific scar locations (e.g., centre of scar, worst area of scar, counting transducer lengths)	6,8,9,13,19-21,23,30,33-37	 Photographs of scars taken ²⁶ Measurement taken at standardised transducer lengths along surgically created scars of prespecified dimensions ³⁴ Measurements taken at thickest/most severe point ^{19-21,30,33,35,37}, as determined by the patient and/or clinician ⁸
			 Transducer placed on thickest site on peripheral regions ⁹ Transducer placed on area initially identified to have greatest burn depth ²³
	For p	eer review only - http://bmjopen.bmj.com/site/about/guideline	s.xhtml Page 27 of 41

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BM, MS, TM, TR, BD, RK, ZT – Ultrasound Scoping Review: Supplement

17,38 Conducting linear measurements from nearby anatomical landmarks .2,24-26,28,29,39,40 Acclimatising Removing scar treatments prior to scar to ultrasound measurement conditions measurement 5,18,22,29,41-46 Acclimatising patient to room prior to measurement

- Measurement area selected by the measurer with -selected area marked with tape ¹³

- Measurements taken at set linear distances from cranial/caudal border of linear sternal scar ³⁶

- Linear measurements from anatomical landmark to measurement site ¹⁷

- Transducer placement mapped in 3dimensional space using a surgical precision tracking arm ³⁸

- Pressure garments removed 10 minutes before measurement $^{\rm 28}$

- Pressure garments removed 15 minutes before measurement to regain original (uncompressed) scar thickness or to reduce blanching effects on measurement ^{20,40}

- Pressure garments/gels/moisturisers removed 20 minutes before measurement ^{8,22,29}

- Pressure garments removed 30 minutes before measurement ^{12,25,26,39}

- Sequential measurement of scars following direct treatment with vacuum massage at 5, 30, 60 and 120 minutes to monitor effect of treatment ²⁴

- Patients rested for minimum 5 minutes before measurement ^{5,18,22}

- Scar exposed to room conditions for 10 minutes ²⁹ to allow equilibrium to be reached with surrounding environment ⁴¹

- Patients resting in room with constant temperature for 15 mins ⁴² to allow scar to stabilise ⁴⁴

Maintaining patient position before measurement

Measuring and/or dermis different skin individually layers

Measuring epidermis

Measuring both epidermis/dermis combined (no

5,6,8,11,12,15,17,18,22,23,26,28,35,40,55-68

- Patients rested for 20 minutes prior to measurement 29,45

- Patients resting for 10 minutes before repeated measurements taken 43

- Patients wait in testing room holding position for 5 min before measurement to stabilise cutaneous blood flow ⁵

- Patients allowed to adapt in controlled room to exclude external variables 46

- Patients remained supine for at least 5 minutes before measurement to avoid artefacts on Doppler imaging ¹³

- Patients allowed to acclimatise to room and assumed a supine position for a minimum of 10 minutes before measurements of biophysical parameters ¹¹

- Measurement of epidermal, dermal and combined epidermal and dermal thickness to allow comparison with histological measurement 47,48

- Measurement of the epidermal and dermal thickness ^{45,49}, combined with layer acoustic density⁷

- Measurement of the epidermal, dermal and subcutaneous thickness, combined with acoustic density 50,51

- Measurement of dermal thickness as treatment thought to affect/target the dermis ^{24,37,52-54}

- Combined epidermal and dermal thickness measurement to provide information on the full thickness of the scar 5,6,8,11,12,15,17,18,22,23,26,28,35,40,55-68

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Measurement objective	individual measurement) Measuring fibrosis/oedema/hair follicles	8,10,11,13,14,16,17,24,25,29-32,34,36,37,45,54,58,60,61,63,64,69-82	- Measurement of fibrosis or collagen architecture ^{8,11,17,24,29-32,34,36,37,45,54,58,61,63,64,69,70,72-} 74,77-79,82
			 Measurement of inflammation/oedema ¹⁴ Quantification of the sub epidermal low echogenic band, indicating oedema ⁶⁰ Measurement of both fibrosis and oedema ^{10,13,16,25,58,71,75,76,80,81}
Factors influencing scar site measurement	Measuring contralateral skin/control scar	9,14,15,23,29,30,52,55-58,83-88 6,8,12,18,22,25,38,43,54,59-61,66,89,90 39,40,45,79,81,82	 Measurement of the presence and density of hair follicles to differentiate scarred and unscarred skin⁵⁴ Measurement of additional, non-scarred subjects ^{55,79} Measurement of unscarred/unaffected skin on same subject as scar measurement contralaterally or at anatomically similar location to provide normative measurements for skin thickness 6,8,9,12,14,15,18,22,23,29,30,38-40,43,45,52,54,56-61,66,81,85-90
			 Measurement of both untreated scar and unaffected skin ⁸²⁻⁸⁴ Measurement of a control scar subjected to care as usual treatment on the same individual ²⁵
	Measuring open wounds or sores in the scar	6	- Use of flexible transparent plastic wrap placed over the measurement area to prevent contact between ultrasound gel and transducer with the open wound/sore ⁶
	Operator training and/or experience	6,8,12,14,16,18,20,24,27-29,31,39,40,58,61,66,72,73,87,91-93	 Trained outcome assessor ^{6,13,16,18,27,72} Measurements taken by radiologist/sonographer 28,66,73,92

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5,6,8,9,11,12,20,23,25,26,31,34,37,40,44,45,47,52,54,57,60,61,66,68,79,85,92,94

measurements per scar

- Assessors with burn experience ^{87,93} - Ultrasound located in department of radiology

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- Measurements conducted by trained therapist/doctor under guidance of experienced radiologist 12,14,29,39

- Measurements conducted by trained clinicians who use device regularly and received training by company representative of devices ^{8,61}

- Device-specific training provided: 1 week ²⁰; 3 sessions of 3 hours for 3 weeks, plus 10 independent assessments of scars using study protocol ⁴⁰; training provided over 3 months ³¹; physical therapist trained in ultrasound application ²⁴

- 3 ultrasound images taken from each patient 9,11,26,31,37,44,45,47,52,54,57,60,79,85

- Clearest of 3 measurements used ¹²

- 3 measurements in 3 locations across scar used. Individual and average measurements reported 40

- Measurements performed in duplicate ^{34,94}

- Measurements taken at different points of the

scar, thickest used for analysis ⁹²

- 5 measurements of each site 6,23

- 9 measurements taken, removal of maximum and minimum, 7 measurements used for average 20

- Measurements taken by 3 assessors at 3 different time points during day 8,61

- Measurement of 2 sites on the same scar²⁵

- Single ultrasound image taken for analysis ⁶⁸

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3 4	Use of additional	Using additional objective assessment	6,9-11,13,15,17,18,21-23,25-27,29,31,32,35,36,40-48,50,53,56-59,66,68-70,75- 80,82-84,86-92,95-111	- Histology/immunohistochemistry 13,17,47,48,50,58,78,79,88,100,103,108,110
5	measurement	instruments (e.g.,		- Blood flow and blood perfusion measurement
7	tools as well as	histology, colour		using laser Doppler perfusion imaging.
7 8	ultrasound	Doppler ultrasound		flowmetry or PeriCam and scar colour and
9	magguramante	cutometer		micro vassal percentage using dermoscopyolour
10	measurements			and micro vessel percentage using definoscopyolodi
11		colourimeter)		and micro-vesser percentage. 35 69 70 83 84 86 87 92 99 101 108
12				
13				- Oximeter 4
14				- Infra-red camera ⁴¹
15				- Measurement of scar stiffness or
16				pliability/elasticity using elastography or
17				cutometer ^{9,15,18,21,22,25-27,29,43,46,53,57,66,82-}
18				84,86,89,90,96,98,99,101,104-106
19				- Measurement of sensation using Semmes-
20				Weinstein filaments ^{82-84,86}
21				- Measurement of scar colour (including
22				nigmentation and erythema) using
23				spectrophotometer colourimeter chromameter
25				mexameter or Dermlite Foto IIPro ^{18,22,25-}
26				27,32,42,44-46,53,56,66,68,80,82,87,90,91,96-99,101-107,111
27				Maggymemont of theme anidownal yester loss
28				- Measurement of trans-epidermai water loss
29				using Tewameter of scar hydration using
30				Corneometer 40,55,70,77
31				- Measurement of sebum level using sebumeter
32				90,99
33				- Measurement of hardness using durometer ⁹¹
34 25				- Measurement of neovascularisation using
35				echocontrastography ⁵⁸
37				- Measurement of scar dimensions (e.g., scar
38				height and volume) using 3D camera, 3D
39				imaging methods, ruler or calliner ^{6,10,11,23,36,75,77}
40				mono monous, raier or earriper
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43		For	peer review only - http://hmionen.hmi.com/cite/about/quidelines.x	Page 32 of 41

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44 45 46 Using subjective assessment instruments (e.g., clinical rating scales, PROMs) 19,20,23,28-30,33,37,40,41,44,45,49,52,56,57,61,66,67,69-72,80-84,86,87,91,92,94-98,100,111-115 - Measurement of skin thickness using micrometer or optical coherence tomography 17,31,59,76,108-110

- Measurement of scar firmness or deformation using cicatrometer, force/torque sensor (in line with ultrasound to measure load applied) or torque meter ^{31,32,107}

- Multi-parameter skin analysis device ⁶⁶

- Measurement of erythema and elasticity using probes of DermaLab Combo $^{\rm 40}$

- Multi-probe adaptor taking multiple measurements (pigmentation, erythema, transepidermal water loss) ⁹⁶

PROMs:

- Measurement of scar quality using POSAS patient report ^{8,23,30,33,45,56,61,63,64,66,75-}77,82,86,95,97,106,107,114,115

- Subjective rating scales for scar symptoms (e.g., pain, itch) or subjective scar severity ratings ^{26,30,41,42,53,63,64,72,80,83,84,93,102,103,111,115}

- Patient quality of life questionnaires ^{75,76,101,107}
- Measurement of generic health-related quality of life using CHU-9D ^{63,64}

- Measurement of scar-specific health-related quality of life using BBSIP ^{63,64}

- subjective evaluation of response to treatment/treatment satisfaction ^{81,116} Clinical rating scales:

- Measurement of scar quality using POSAS observer report ^{8,23,30,33,45,53,56,61,63,64,66,75-}77,82,86,87,97,98,106,114-116

- Measurement of physical scar characteristics using VSS or modified versions of the VSS ^{8,18-} 20,28,30,33,35,37,38,40,42-44,49,56,57,61,65,66,69-72,80-86,92-95,100-103.111-113.115.117.118 - Measurement of scar characteristics in relation to unscarred skin using Seattle Scar Scale or modified Seattle Scar Scale ⁷³ - Subjective rating scales for scar symptoms (e.g., pain, itch) as assessed by the clinician and/or researcher and/or clinical evaluation of scar severity 11,29,41,52,57,67,73,91,92,94,96 Determining the order - Standardised order of measurement: 3D of measurement photograph, POSAS-O, then ultrasound ⁶ - Order of device use not specified 35,69,70,83,84,86,87,92,99,101,108 Abbreviations: TUPS: Tissue Ultrasound Palpation System; 3D: three-dimensional; POSAS: Patient and Observer Scar Assessment Scale; CHU-9D: Child Health Utility 9D; BBSIP: Brisbane Burn Scar Impact Profile; VSS: Vancouver Scar Scale; mVSS: Modified Vancouver Scar Scale; POSAS-O: Patient and Observer Scar Assessment Scale, observer measure

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Preferred Reporting Items for Systematic reviews and Meta-Analyses extension for Scoping Reviews (PRISMA-ScR) Checklist

SECTION	ITEM	PRISMA-ScR CHECKLIST ITEM	REPORTED ON PAGE #	
TITLE				
Title	1	Identify the report as a scoping review.	1	
ABSTRACT				
Structured summary	2	Provide a structured summary that includes (as applicable): background, objectives, eligibility criteria, sources of evidence, charting methods, results, and conclusions that relate to the review questions and objectives.	3-4	
INTRODUCTION				
Rationale	3	Describe the rationale for the review in the context of what is already known. Explain why the review questions/objectives lend themselves to a scoping review approach.	5-7	
Objectives	4	Provide an explicit statement of the questions and objectives being addressed with reference to their key elements (e.g., population or participants, concepts, and context) or other relevant key elements used to conceptualize the review questions and/or objectives.	7	
METHODS				
Protocol and registration	5	Indicate whether a review protocol exists; state if and where it can be accessed (e.g., a Web address); and if available, provide registration information, including the registration number.	7	
Eligibility criteria	6	Specify characteristics of the sources of evidence used as eligibility criteria (e.g., years considered, language, and publication status), and provide a rationale.	8-10	
Information sources*	7	Describe all information sources in the search (e.g., databases with dates of coverage and contact with authors to identify additional sources), as well as the date the most recent search was executed.	8	
Search	8	Present the full electronic search strategy for at least 1 database, including any limits used, such that it could be repeated.	9	
Selection of sources of evidence†	9	State the process for selecting sources of evidence (i.e., screening and eligibility) included in the scoping review.	9	
Data charting process‡	10	Describe the methods of charting data from the included sources of evidence (e.g., calibrated forms or forms that have been tested by the team before their use, and whether data charting was done independently or in duplicate) and any processes for obtaining and confirming data from investigators.	10-11	
Data items	11	List and define all variables for which data were sought and any assumptions and simplifications made.	10-11 and supplementary table 1	
Critical appraisal of individual	12	If done, provide a rationale for conducting a critical appraisal of included sources of evidence; describe	N/A	



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SECTION	ITEM	PRISMA-ScR CHECKLIST ITEM	REPORTED ON PAGE #
sources of evidence§		the methods used and how this information was used in any data synthesis (if appropriate).	
Synthesis of results	13	Describe the methods of handling and summarizing the data that were charted.	10-11
RESULTS			
Selection of sources of evidence	14	Give numbers of sources of evidence screened, assessed for eligibility, and included in the review, with reasons for exclusions at each stage, ideally using a flow diagram.	11-12
Characteristics of sources of evidence	15	For each source of evidence, present characteristics for which data were charted and provide the citations.	12-15
Critical appraisal vithin sources of evidence	16	If done, present data on critical appraisal of included sources of evidence (see item 12).	N/A
Results of individual sources of evidence	17	For each included source of evidence, present the relevant data that were charted that relate to the review questions and objectives.	Results section (11-46)
Synthesis of results	18	Summarize and/or present the charting results as they relate to the review questions and objectives.	Results section (11-46)
DISCUSSION			
Summary of evidence	19	Summarize the main results (including an overview of concepts, themes, and types of evidence available), link to the review questions and objectives, and consider the relevance to key groups.	47-49
Limitations	20	Discuss the limitations of the scoping review process.	49-50
Conclusions	21	Provide a general interpretation of the results with respect to the review questions and objectives, as well as potential implications and/or next steps.	50-51
FUNDING			
Funding	22	Describe sources of funding for the included sources of evidence, as well as sources of funding for the scoping review. Describe the role of the funders of the scoping review.	51

JBI = Joanna Briggs Institute; PRISMA-ScR = Preferred Reporting Items for Systematic reviews and Meta-Analyses extension for Scoping Reviews.

* Where *sources of evidence* (see second footnote) are compiled from, such as bibliographic databases, social media platforms, and Web sites.

† A more inclusive/heterogeneous term used to account for the different types of evidence or data sources (e.g., quantitative and/or qualitative research, expert opinion, and policy documents) that may be eligible in a scoping review as opposed to only studies. This is not to be confused with *information sources* (see first footnote).
‡ The frameworks by Arksey and O'Malley (6) and Levac and colleagues (7) and the JBI guidance (4, 5) refer to the process of data extraction in a scoping review as data charting.

§ The process of systematically examining research evidence to assess its validity, results, and relevance before using it to inform a decision. This term is used for items 12 and 19 instead of "risk of bias" (which is more applicable to systematic reviews of interventions) to include and acknowledge the various sources of evidence that may be used in a scoping review (e.g., quantitative and/or qualitative research, expert opinion, and policy document).

From: Tricco AC, Lillie E, Zarin W, O'Brien KK, Colquhoun H, Levac D, et al. PRISMA Extension for Scoping Reviews (PRISMAScR): Checklist and Explanation. Ann Intern Med. 2018;169:467–473. doi: 10.7326/M18-0850.



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