# Proprioceptive impairment in unilateral neglect after stroke: A systematic review

SAGE Open Medicine Volume 8: 1–11 © The Author(s) 2020 Article reuse guidelines: sagepub.com/journals-permissions DOI: 10.1177/2050312120951073 journals.sagepub.com/home/smo

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## Abstract

**Introduction:** Unilateral neglect is a debilitating condition that can occur after stroke and can affect a variety of domains and modalities, including proprioception. Proprioception is a sensorimotor process essential to motor function and is thus important to consider in unilateral neglect. To date, there has not been a comprehensive review of studies examining the various aspects of proprioceptive impairment in unilateral neglect after stroke. This review aimed to determine if people with unilateral neglect have more severe proprioceptive impairments than those without unilateral neglect after stroke.

**Methods:** The MEDLINE, Embase, Scopus, CINAHL and Web of Science databases were searched from inception to September 2019 using an a priori search strategy. Two independent reviewers screened abstracts and full texts, and extracted data from the included full texts. A third reviewer resolved disagreements at each step. Risk of bias was assessed using the AXIS Quality Assessment tool.

**Results:** A total of 191 abstracts were identified, with 56 eligible for full-text screening. A total of 18 studies were included in the review and provided evidence that people with unilateral neglect have more severe proprioceptive impairment than people without unilateral neglect. This impairment is present in multiple subtypes of unilateral neglect and aspects of proprioception. Most studies had a moderate risk of bias.

**Conclusion:** People with unilateral neglect after stroke are more likely to have impaired processing of multiple types of proprioceptive information than those without unilateral neglect. However, the available evidence is limited by the large heterogeneity of assessment tools used to identify unilateral neglect and proprioception. Unilateral neglect and proprioception were rarely assessed comprehensively.

PROSPERO Registration: CRD42018086070.

#### **Keywords**

Rehabilitation, sensorimotor, neurology, unilateral neglect, proprioception, stroke, systematic review

Date received: 5 April 2020; accepted: 24 July 2020

# Introduction

Unilateral neglect (UN) is an umbrella term for a range of clinical presentations, characterised by the failure to report, respond, or orient to novel or meaningful stimuli presented on the side opposite to the brain lesion.<sup>1</sup> Depending on the method used for assessment, UN is estimated to affect  $23.5\%-67.8\%^{2,3}$  of people after stroke. However, the most commonly used assessments (cancellation tasks and National Institutes of Health Stroke Scale (NIHSS) Item 11) indicate the incidence is ~30%.<sup>4,5</sup> UN is linked to greater length of hospital stay, higher incidence

of falls, and a reduced likelihood of home as a discharge destination.<sup>2,6–9</sup> These negative sequelae are largely due to the poor functional outcomes that are associated with UN,

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Creative Commons Non Commercial CC BY-NC: This article is distributed under the terms of the Creative Commons Attribution-NonCommercial 4.0 License (https://creativecommons.org/licenses/by-nc/4.0/) which permits non-commercial use, reproduction and distribution of the work without further permission provided the original work is attributed as specified on the SAGE and Open Access pages (https://us.sagepub.com/en-us/nam/open-access-at-sage). which persist at 1-year post-stroke in 30%-60% of patients.<sup>10-12</sup> Despite the high incidence and negative functional consequences of UN, there is no consensus about effective yet clinically feasible assessment and treatment strategies for the condition.<sup>13-17</sup>

The lack of appropriate assessments and effective treatment strategies is an indication of the complexity of UN. UN is often associated with larger lesion volumes<sup>3</sup> and is not the result of lesions in a single location. Rather, it is the result of impaired functional connectivity between brain regions associated with attention, sensorimotor and visual processing, notably the frontoparietal networks.<sup>18–20</sup> As a result, symptoms of UN are varied and can present in a range of domains (visuospatial, auditory and sensorimotor) and spaces (personal, peri-personal and extra-personal).<sup>21,22</sup>

Clinical assessment of UN should target a combination of these domains and spaces, and yet, many current standardised clinical assessments are limited to only one.<sup>13</sup> The most common form of UN assessment uses pen-and-paper tasks, including line bisection, shape cancellation, or figure copying.<sup>14</sup> These tasks are unable to capture visuospatial neglect in the peri-personal and extra-personal spaces and do not account for auditory or sensorimotor domains.<sup>13</sup> Given this limited assessment scope, it is not surprising that UN is often considered a singular condition and associated solely with visuospatial impairment, neglecting other domains, and thus, the heterogeneous nature of the condition.<sup>9,23</sup>

Importantly, a sensorimotor impairment in UN that is often clinically neglected is proprioception.<sup>24</sup> Proprioception, as defined by Sherrington in 1906, derives from the Latin word proprius, meaning 'one's own', combined with the concept of perception and thus refers to 'perceiving one's own self'.<sup>25</sup> Proprioception is largely considered as the processes enabling joint movement, position detection and muscle force judgement (for review, see Hillier et al.<sup>25</sup>). However, information about the size and shape of body parts is crucial for proprioception.<sup>26</sup> Explicitly, to sense the position and movement of 'one's own self' is impossible without the information about what that 'self' is. Therefore, an updated and emerging definition of proprioception includes the body representation, defined as the stored internal model of the body and its parts.<sup>26-28</sup> Distinct methods of assessing body representation include judgement of laterality, body axes and body topography.<sup>29,30</sup>

Standardised clinical assessments of proprioception typically involve the detection or judgement of passively imposed joint movements in the absence of vision.<sup>31,32</sup> These capture a patient's ability to detect when and where a body part is moving, without accounting for body representation.<sup>33</sup> Skilled movement emerges from the judgement of movement direction, magnitude and the nature of the moving parts, provided by the body representation. Given that stroke rehabilitation focuses largely on the restoration of skilled movement, it is important to consider impairments in all aspects of proprioception. In the context of the complex presentation of UN, it is conceivable that multiple aspects of proprioception are impaired. However, this is unknown because, to date, no review examining proprioception in UN has been conducted.

Impaired proprioception is associated with poor motor and functional outcomes at all stages of stroke.<sup>34,35</sup> Thus, it should be an essential target in stroke rehabilitation, particularly in patients with UN. Therefore, the primary aim of this systematic review is to determine whether proprioception, including the body representation as part of the definition, is more impaired in people with UN compared to those without after a stroke. The secondary aims are to identify the assessments used to detect UN and its domains (e.g. spatial, extra-personal).

# Methods

## Protocol and registration

The protocol for this systematic review is registered in PROSPERO, under the registration number CRD42018086070 and can be accessed at the following link: https://www.crd. york.ac.uk/prospero/display\_record.php?RecordID=86070.

# Eligibility criteria

Studies were included if they met the following criteria: (1) cross-sectional design or intervention studies that provided cross-sectional data at baseline, (2) participants were adults aged 18 years and over, (3) participants had first-time stroke confirmed on medical imaging, (4) employed at least one standardised assessment of UN, and (4) at least one assessment of proprioception and (5) had data reported for participants with UN (UN+) and without (UN-). There was no restriction on publication year or language. Studies were excluded if they used clinical tests that assessed only balance and/or vestibular function and/or sensorimotor function that was not specific to proprioception.

#### Search strategy and study selection

This review followed Cochrane Methodology and Preferred Reporting Items for Systematic Reviews and Meta-Analysis (PRISMA) reporting guidelines.<sup>36</sup> The CINAHL, Embase, MEDLINE, Scopus and Web of Science electronic databases were searched from inception to 10 September 2019 using a pre-established search strategy developed by all members of the study team and reviewed by a university librarian (Supplemental Appendix 1). Two review authors (G.F. and D.K.) screened abstracts and full texts independently, and a third review author (C.Q.d.O.) resolved disagreements at each step of study selection. Screening was conducted using the Covidence software. Studies in languages other than English were translated using an online translation service.

### Data extraction

Two review authors (G.F. and C.Q.d.O.) independently extracted data using a standardised excel spreadsheet based on the Cochrane Data Extraction Template.<sup>37</sup> Extracted information included studies' authors, year, aims, setting, population, procedures; participant's recruitment, demographics and baseline characteristics, and completion rates; type of outcome measures used to assess UN and proprioception, assessment data for UN and proprioception with time points of collection; and the suggested mechanisms of interaction between proprioception and neglect. Determination of UN was based on standardised and validated clinical or laboratory tests developed to assess UN, such as Albert's Test,<sup>38</sup> Rivermead Behavioural Inattention Test (BIT)<sup>39</sup> and Catherine Bergego Scale (CBS)<sup>40</sup> (for summary, see Menon and Korner-Bitensky<sup>13</sup>). Assessment of proprioception included standardised tests in any of the following categories: (1) movement detection/direction discrimination, for example, the distal proprioception test;<sup>41</sup> (2) joint position judgement or reproduction for example, the Wrist Position Sense Test;<sup>42</sup> (3) force judgement or matching, for example, finger force reproduction as per Walsh et al.;43 and (4) tests of body representation, for example, the Recognise<sup>TM</sup> App for hand laterality.44 Assessments of proprioception that had not been formally validated were eligible for inclusion, given the lack of research into tests of certain aspects of proprioception. Where information was not available in the published study, details were requested from the corresponding author.

# Risk of bias assessment

Risk of bias was evaluated by two review authors (G.F. and C.Q.d.O.), independently using the Appraisal tool for Cross-Sectional Studies (AXIS), a 20-item scale developed using a Delphi panel consensus assessing 5 domains - Introduction (1 item), Methods (10 items), Discussion (5 items), Conclusion (2 items), and Other (2 items).45 The AXIS requires a Yes/No/Unsure assessment for each item, for example, 'Was the sample size justified?' and 'Were the results for the analyses described in the methods presented?' The full list of AXIS items is reported in Supplemental Appendix 4. Disagreements were resolved through discussion and when necessary, with a third review author (D.K.). Prior to discussion, percent agreement between the two reviewers on AXIS items was 90.28%. The AXIS acknowledges the issues with the summation of checklists for study quality,46,47 and as such, does not have published cut-off scores to categorise studies as low, medium, or high risk of bias.<sup>45</sup> Therefore, the authors used a quartiles system for categorising risk of bias according to the number of AXIS items met (low risk >15, moderate risk 10-15, high risk 4–9 and very high risk <4).<sup>48</sup>

### Summary measures and data synthesis

A descriptive synthesis of (1) between-group differences of UN+ and UN- stroke patients, and (2) types of assessments of UN and proprioception used in the included studies was conducted. The summary of study results was presented in separate tables for continuous (mean values and standard deviations, medians and interquartile range (IQR)) and dichotomous data (percentages and total number of participants). Effect size was calculated using Hedges g for continuous outcomes due to small and uneven group sizes<sup>49</sup> and odds ratios for dichotomous outcomes.<sup>50</sup> Risk of bias assessment was descriptively synthesised and presented in tabular format.

# Results

## Study selection

A total of 284 titles and abstracts were identified, with 191 eligible for abstract screening after duplicate removal, and 56 eligible for full-text screening. A total of 18 studies were included in the review<sup>23,51–67</sup> (see Figure 1). The predominant reasons for exclusion at full-text review were inadequate data reporting and a lack of a measure of proprioception. The full list of excluded studies and the reasons for their exclusion can be found in Supplemental Appendix 2.

## Study characteristics

**Population.** The complete characteristics of the participants included in the studies are summarised in Table 1. A combined total of 959 participants were included in the studies, 246 in the UN+ group and 713 in the UN- group. All studies had a small sample size (mean=50, SD= $\pm$ 68, range=6-281). The mean age and standard deviation were  $60.3 \pm 5.4$  years (UN+= $60.9 \pm 5.6$ , UN-= $59.8 \pm 5.4$ ), and the majority of participants were male (65%). Most studies recruited participants in the sub-acute phase (3 weeks to 6 months post-stroke); however, two studies<sup>59,63</sup> collected data from participants in the chronic phase (more than 6 months after stroke). Five studies<sup>23,53,64-66</sup> recruited stroke populations in different phases (chronic, acute, sub-acute), and one study<sup>67</sup> limited recruitment to the acute phase (less than 3 weeks post-stroke).

**Proprioception assessment.** Proprioception was assessed by 13 different methods across the studies, as described in Table 2. Three studies<sup>62,63,67</sup> assessed solely the detection or discrimination of movement. Five studies<sup>60–62,64,65</sup> assessed proprioception via a limb-matching task. Of the four studies<sup>51,54,59,67</sup> that examined laterality, three<sup>51,54,59</sup> used a hand laterality task and one<sup>67</sup> utilised a laterality task in which participants were required to point to the left or right hand of human stick figure drawings in various orientations. Five studies<sup>52,55,56,58,66</sup> used an assessment that required

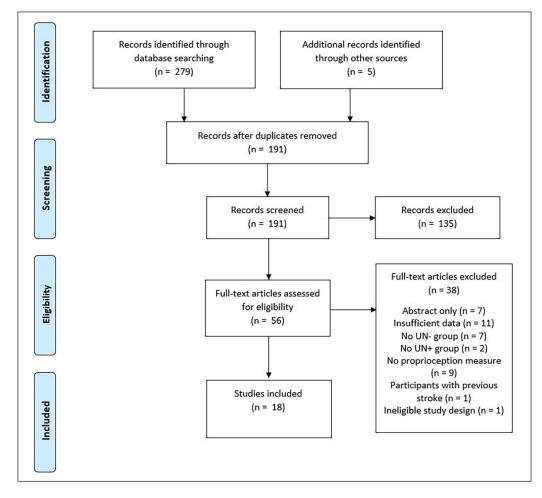


Figure 1. Flow of studies through review.

participants to identify the location of the body midline or a body axis. Finally, three studies<sup>23,53,57</sup> examined perceptions of body topography by asking participants to arrange tiles printed with body parts, detect targets placed on their body and localise points stimulated on their trunk, respectively. A single study<sup>67</sup> reported two distinct proprioception outcomes, laterality and movement detection. No study investigated force matching.

#### Risk of bias

According to the AXIS risk of bias assessment, 2 studies were rated as low risk of bias,<sup>52,66</sup> 12 as moderate risk of bias<sup>23,51,54,56,58–65,67</sup> and 3 as high risk of bias<sup>53,55,57</sup> (see Table 3, with full AXIS items and result reported in Supplemental Appendices 3 and 4). The two studies using the CBS as a measure of UN had an AXIS rating of low<sup>52</sup> and high risk of bias.<sup>57</sup> The main sources of risk of bias were sampling strategy, and consistency in reporting participant characteristics and results for all planned analyses. All studies defined their target population, reported internally consistent results, and justified their discussion and conclusions. Only one study<sup>52</sup> justified their sample size

and reported non-response to recruitment. Twelve studies<sup>23,51,53–58,60,61,65,66</sup> failed to report their methodological limitations and four studies<sup>53,57,60,61</sup> did not employ a validated assessment tool for UN. Five studies<sup>23,53–56</sup> did not present results for all planned analyses and six<sup>55,57–59,63,67</sup> did not provide sufficient participant demographic data. Finally, seven studies<sup>51,53,55,57,58,63,65,67</sup> used convenience samples.

# Synthesis of results

**Proprioceptive impairment in UN.** The majority of studies in this review support the existence of more severe proprioceptive impairment in UN+ compared to UN-. The two studies with low risk of bias reported conflicting results on the relationship between proprioceptive impairment and UN. One study<sup>52</sup> reported a body axis judgement error correlation of r=-0.61 and r=-0.56 for the CBS and Line Bisection Test, respectively, indicating that individuals with UN had more severe proprioceptive impairment. Another study<sup>66</sup> reported a mean difference of 2.4 cm in forearm bisection measurement between people with and without UN, with a small effect size indicating

Author	Setting	Time since stroke	Sampl	e size	Age (mean, SD)		Lesion side	
			UN+ UN-		UN+	UN-	UN+	UN-
Movement detection/	judgement							
Meyer et al. <sup>62</sup>	Inpatient rehab	Sub-acute	27	95	68 (60.2–77.7) <sup>a</sup>	66.7 (58.7–75.7) <sup>a</sup>	NR	NR
Schmidt et al.63	NR	Chronic	7	15	61.7 (14.8)	66.3 (12.2)	R=7	R = 15
van Stralen et al. <sup>67</sup>	ASU	Acute	9	47	58.9 (12.4)	61.5 (16.1)	R=8, I=NR	R=14, L=33
Joint position matchin	g							
Borde et al.61	NR	NR	6	3	63.3 (9.4)	62.33 (5.77)	R=6	R=3
Borde et al. <sup>60</sup>	Inpatient rehab	Sub-acute	10	20	63.4 (8.8)	62.5 (11.9) + 61.2 (15.5) <sup>b</sup>	R = 10	R=10, L=10
Semrau et al. <sup>65</sup>	ASU, Inpatient rehab	Acute, sub-acute	35	123	59 (20-86)	63 (18–89)	R=31, L=4	R = 67, L = 55, B = I
Semrau et al. <sup>64</sup>	ASU, Inpatient rehab	Acute, sub-acute	59	222	62.32 (15.19)	60.64 (14.46)	NR	NR
Laterality								
Baas et al.51	NR	NR	7	15	51.47 (13.62)	61.29 (7.89)	R=7	R = 15
Coslett <sup>54</sup>	NR	Sub-acute, chronic	3	3	63.67 (10.41)	58.67 (11.37)	NR	NR
van Stralen et al. <sup>67</sup>	ASU	Acute	9	47	58.9 (12.4)	61.5 (16.1)	R=8, I=NR	R = I4, L = 33
Vromen et al. <sup>59</sup>	Inpatient rehab	Chronic	8	12	55.3 (8.4)	59.5 (6.9)	R=8	R=12
Body axis/midline								
Barra et al.52	NR	Sub-acute	10	8	63.6 (7.53)	53.12 (18.26)	R=7, L=3	R=3, L=5
Heilman et al.55	NR	NR	5	5	48 (10.3)	58.6 (6.27)	R = 5	L=5
Richard et al. <sup>56</sup>	Inpatient rehab	Sub-acute, chronic	8	8	61.13 (12.45)	52.13 (13.61)	R=8	R=8
Saj et al. <sup>58</sup>	Inpatient rehab	NR	6	6	58 (12.7)	59.2 (11.2)	R=6	R=6
Tosi et al. <sup>66</sup>	Inpatient rehab	Acute, chronic	7	38	68.42 (7.23)	65.97 (12.5)	R=7	R=16, L=21, B=1
Body topography								, -
Cocchini et al. <sup>53</sup>	Inpatient rehab	Sub-acute	14	24	67.21 (8.43)	58.00 (10.7) + 59.45 (15.9) <sup>b</sup>	R=14	R=13, L=11
Di Vita et al. <sup>23</sup>	NR	Sub-acute, chronic	7	16	65.29 (10.29)	64.44 (13)	R=6, L=1	R=12, L=4
Rousseaux et al. <sup>57</sup>	NR	NR	9	6	53.1 (13.2)	46.3 (9.3)	R=9	R=6

**Table I.** Characteristics of included studies, by proprioceptive test type.

SD: standard deviation; UN: unilateral neglect; UN+: participants with UN; UN-: participants without UN; NR: not reported; ASU: acute stroke unit; R: right; L: left; B: bilateral.

<sup>a</sup>Median and IQR presented.

<sup>b</sup>Two UN- groups, first listed (R) side lesion, second (L) sided lesion. Acute defined as <3 weeks post-stroke, sub-acute 3 weeks to 6 months post-stroke, and chronic >6 months post-stroke.

greater proprioceptive impairment in UN+ group (Hedges g=0.22). Of the 13 studies with a moderate risk of bias, 77% (n=10) reported medium or large effect sizes indicating more severe proprioceptive impairment in people with UN.

The studies with the largest sample size<sup>64,65</sup> reported large and moderate effect sizes (Hedges g=1.59 and 1.43, and 0.63, respectively) indicating greater proprioceptive impairment in people with UN than those without. All three studies with a high risk of bias<sup>53,55,57</sup> reported significantly more severe proprioceptive impairment in UN+. A descriptive synthesis of the findings of the included studies' results is presented in Table 4 (continuous outcomes) and Table 5 (dichotomous outcomes) broken down by proprioceptive test subtype.

*UN* assessment. There were a total of 18 different assessment tools used to assess UN, with 12 studies<sup>23,51–54,56–61,63</sup>

using more than one outcome measure to assess UN (see Table 2). Nine studies<sup>53–56,58,60,62,63,67</sup> used pen-and-paper tests alone to identify UN, and two studies<sup>51,66</sup> used outcome measures that assessed personal neglect solely. Two studies<sup>59,61</sup> used a combination of a pen-and-paper task with a self-reported measure of UN and an environmental observation task, respectively. Two studies<sup>64,65</sup> used a behavioural assessment in isolation (the BIT), while two studies<sup>52,57</sup> added a functional assessment of UN (the CBS) to pen-and-paper cancellation and bisection tasks.

# Discussion

# Summary of evidence

We found moderate risk of bias in the majority of studies that demonstrated that people with UN after stroke have

Author	UN test(s)	Type of UN assessment	Proprioception test(s)	
Movement detection				
Meyer et al. <sup>62</sup>	SCT	Pen and paper	Em-NSA, TFT (0–3)	
Schmidt et al. <sup>63</sup>	chmidt et al. <sup>63</sup> LeCT, SCT, LBT, Figure Copying, Reading Test		Arm Position Test-Erro	
van Stralen et al. <sup>67</sup>	SCT	Pen and paper	RASP	
Joint position matching				
Borde et al. <sup>61</sup>	LCT, Observation, Environment Description, Double Letter Cancellation	Pen and paper, Extra- personal	Upper limb position reproduction, TFT	
Borde et al. <sup>60</sup>	LBT, LeCT	Pen and paper	Upper limb position reproduction	
Semrau et al. <sup>65</sup>	BIT	Behavioural	Robotic Arm Position Matching Task, TFT	
Semrau et al. <sup>64</sup>	Semrau et al. <sup>64</sup> BIT		Robotic Arm Position Matching Task	
Laterality			U U	
Baas et al.51	Fluff test (primary), LBT, BCT	Personal	Hand Laterality	
Coslett <sup>54</sup>	LBT, SCT, LCT	Pen and paper	Hand Laterality	
van Stralen et al. <sup>67</sup>	SCT	Pen and paper	Bergen Laterality Test	
Vromen et al. <sup>59</sup>	SCT, subjective neglect questionnaire	Pen and paper, self-report	Hand Laterality	
Body axis/midline				
Barra et al. <sup>52</sup>	BCT, CBS, LBT	Pen and paper, functional	Longitudinal Body Axis	
Heilman et al. <sup>55</sup>	LBT	Pen and paper	Pointing to body midline	
Richard et al.56	BCT, Scene Copy, LBT (2/3)	Pen and paper	Pointing to body midline	
Saj et al. <sup>58</sup>	BCT, LBT, Scene Copy	Pen and paper	Longitudinal Body Axis	
Tosi et al. <sup>66</sup>	Biasch's Test	Personal	Arm bisection task	
Body topography				
Cocchini et al. <sup>53</sup>	SCT, LCT	Pen and paper	Body Exploration Fluff Test	
Di Vita et al. <sup>23</sup>	LeCT, LCT, Use of Common Objects Test, Sentence Reading, Wundt– Jastrow Area Illusion	Personal	Body Topography	
Rousseaux et al. <sup>57</sup>	LBT, BCT, CBS	Pen and paper, functional	Tactile Stimulation Localisation	

Table 2. Assessment descriptions.

UN: unilateral neglect; SCT: star cancellation test; Em-NSA: Erasmus Modifications to the Nottingham Sensory Assessment; TFT: thumb finding test; RASP: Rivermead Assessment of Somatosensory Perception; LCT: line cancellation; LeCT: letter cancellation; LBT: line bisection test; BIT: behavioural inattention test; BCT: bell cancellation; CBS: Catherine Bergego Scale.

more severe proprioceptive impairments than those without. The assessment of UN was commonly limited to pen-and-paper tests designed to capture peri-personal hemi-spatial UN with minimal usage of UN tests that assess the impact of UN on functional activities. Multiple subtypes of proprioception were impaired, including movement detection, joint position matching, and the judgement of laterality, body axes and body topography. Deficits in spatial orientation and exploration in UN may be due to the disruption of distributed cortical networks controlling attention anchored in an egocentric frame of reference.<sup>68</sup> Importantly, this egocentric frame of reference depends extensively on proprioceptive, visual and vestibular inputs.<sup>69</sup> The results of this review suggest that proprioceptive deficits may underlie the disruption to the egocentric reference frame contributing to other impairments seen in UN.

# Proprioception assessment

Proprioception is a complex sensorimotor process, with multiple aspects that contribute to adequate motor function. The finding that multiple types of proprioceptive impairment are present in people with UN is important for two major reasons. First, despite its importance to function, the majority of Stroke Guidelines<sup>17,70–72</sup> do not include any recommendations for assessment or treatment of proprioceptive deficits, and subsequently, it is absent from national audits of stroke rehabilitation. Thus, it is likely that proprioceptive impairments are not being assessed and subsequently treated in this population. Second, the standardised clinical tools to test proprioception such as the Erasmus Modification of the Nottingham Sensory Assessment and the Rivermead Assessment of Somatosensory Perception solely assess simple movement detection. These tools use an ordinal grading system, defining

Author	Intro	Methods	Results	Conclusions	Other	Risk
Baas et al.51	100%	60%	75%	50%	50%	Moderate
Barra et al.52	100%	90%	80%	100%	100%	Low
Borde et al. <sup>61</sup>	100%	80%	90%	50%	0%	Moderate
Borde et al. <sup>60</sup>	100%	70%	100%	50%	100%	Moderate
Cocchini et al.53	0%	20%	50%	50%	0%	High
Coslett <sup>54</sup>	100%	70%	75%	50%	0%	Moderate
Di Vita et al. <sup>23</sup>	100%	70%	75%	50%	100%	Moderate
Heilman et al.55	100%	40%	25%	50%	50%	High
Meyer et al. <sup>62</sup>	100%	80%	75%	100%	100%	Moderate
Richard et al. <sup>56</sup>	100%	80%	50%	50%	50%	Moderate
Rousseaux et al. <sup>57</sup>	100%	60%	50%	50%	50%	High
Saj et al. <sup>58</sup>	100%	60%	50%	50%	100%	Moderate
Schmidt et al. <sup>63</sup>	100%	50%	50%	100%	100%	Moderate
Semrau et al.65	100%	70%	75%	50%	50%	Moderate
Semrau et al. <sup>64</sup>	100%	70%	75%	100%	50%	Moderate
Tosi et al. <sup>66</sup>	100%	90%	80%	50%	100%	Low
van Stralen et al. <sup>67</sup>	100%	70%	75%	100	100%	Moderate
Vromen et al. <sup>59</sup>	100%	80%	75%	100%	100%	Moderate

**Table 3.** AXIS risk of bias assessment summary – percentages of items satisfied.

the patient's proprioception as normal, impaired or absent (reducing sensitivity to change)<sup>31,32</sup> and have no correlation with patient function and activity.<sup>73</sup> In addition, over 75% of Australian physiotherapists and occupational therapists report using non-validated proprioception assessment tools.<sup>74</sup> Thus, standardised clinical assessments of proprioception fail to capture multiple aspects of the sense and have limited clinical utility. Furthermore, it is unknown what aspects of proprioception are assessed in clinical practice.

## UN assessment

The predominant bias of UN assessment to the visuospatial domain suggests that the processing of proprioceptive information is impaired in this type of UN. The studies in this review that did use assessments of other aspects of UN support the notion that proprioceptive information is also neglected in other domains of the condition. However, further evidence is required in order to draw conclusions about proprioceptive impairment outside the traditional definition of visuospatial UN.

This is relevant for two reasons. First, the tendency of researchers and clinicians to consider UN a *visuospatial* disorder alone means that rehabilitation targets impairments only in this domain and that proprioceptive deficits are rarely considered. Second, multiple issues such as poor reliability, lack of ability to detect change, and the allowance for compensation to skew results limit the usefulness of pen-and-paper assessments of UN.<sup>13,75</sup> Moreover, these tests correlate weakly with functional outcomes,<sup>76</sup> which is important given the negative impact of UN on patient functional capacity. Also, the use of pen-and-paper assessments fails to capture a subset of patients with milder presentations of UN, which still likely contribute to functional deficits. However, in the

acute stroke setting, there are more significant functional restrictions, and thus, these tests may be useful as a screening tool for UN at this stage. Where possible, assessment using an ecological tool such as the CBS would provide better insight on the impact of UN on function during the rehabilitation process. Importantly, only two included studies used the CBS, one rated as high and one as low on the AXIS scale.<sup>52,57</sup> Thus, the presence of proprioception impairment in ecologically defined UN is largely unknown.

# Proprioception and UN in clinical practice

The results of review suggest that higher levels of proprioceptive impairment could be a contributing factor in the poorer functional recovery seen in people with UN after stroke. However, despite being the most comprehensive tool available, the CBS does not directly measure the level of proprioceptive impairment present in UN. There is a clear need for a clinical assessment of UN that includes tests that are sensitive to impaired proprioception.

However, the clinical assessment of proprioception is currently inconsistent, frequently non-standardised and there is little data available on what constitutes a minimal clinically important difference (MCID). Importantly, assessment is typically focused solely on position detection. Thus, it is unsurprising that investigation of proprioceptive treatment often has a similar focus.<sup>77,78</sup> Broadening the scope of clinical assessment to include multiple aspects of proprioception is likely to broaden the scope of investigation of treatment strategies. For example, recent studies using somatosensory stimulation (mostly neuromuscular electrical stimulation)<sup>79,80</sup> to provide increased proprioceptive input show promising results in people after stroke. Improvements are thought to be due, in part, to the reintegration of the internal representations

Study	Proprioception outcome	UN+		UN-		Hedge's g	Impaired group	
		Mean (SD)	N	Mean (SD)	N		(effect size)	
Movement detection	n							
Meyer et al. <sup>62</sup>	Em-NSA (median, IQR)	7.0 (2.0-8.0)	27	8.0 (7.0-8.0)	95	UTD	UN+ (p < 0.05) <sup>a</sup>	
	TFT Score (0–3) (median, IQR)	1.0 (1.0-2.0)	27	0 (0-1.0)	95	UTD	$UN + (p < 0.05)^{2}$	
Schmidt et al.63	Arm Position Test–Error	7.5 (1.0)	7.0	4.5 (0.6)	15	3.88	UN+ (large)	
oint position match	ing			. ,				
Borde et al. <sup>61</sup>	Reproduction Error n – paretic upper limb	7.8 (2.5)	6	5 (5)	3	0.73	UN+ (medium)	
	Reproduction Error n – healthy upper limb	3.3 (3.5)	6	1.7 (4.6)	3	0.37	UN+ (small)	
	TFT Error	7.5 (2.7)	6	6.7 (2.9)	3	0.26	UN+ (small)	
Borde et al. <sup>60</sup>	Reproduction Error Total n	12.9 (7.7)	10	9.8 (7.8)	10 <sup>b</sup>	0.38	UN+ (small)	
	Reproduction Error No Vision n	7.2 (5.1)	10	6.2 (5.4)	10 <sup>b</sup>	0.18	Nil	
Semrau et al.65	TFT Score (0–3)	1.3 (1.1)	35	0.7 (0.9)	123	0.63	UN+ (medium)	
Semrau et al. <sup>64</sup>	Kinesthetic Score Vision (lower=better)	3.9 (1.7)	59	1.8 (1.2)	222	1.59	UN+ (large)	
	Kinesthetic Score No Vision (lower=better)	4.3 (1.4)	59	2.4 (1.3)	222	1.43	UN+ (large)	
Laterality								
Baas et al.51	Hand Laterality % Error	25 (5)	7	14 (3)	15	2.85	UN+ (large)	
Coslett <sup>54</sup>	Hand Laterality (L) % Error	41.7 (13.5)	3	6.7 (9.9)	3	2.37	UN+ (large)	
	Hand Laterality (R) % Error	16.3 (9.1)	3	7.3 (8.1)	3	0.84	UN+ (large)	
Vromen et al. <sup>59</sup>	Hand Laterality % Error	37.6 (21.5)	12	14.1 (14.7)	8	1.18	UN+ (large)	
Body axis / midline								
Heilman et al.55	Pointing to body midline–Midline Deviation	8.8 (NR)	5	-1.2 (NR)	5	UTD	UTD	
Richard et al. <sup>56</sup>	Pointing to body midline–Midline Deviation	9.4 (12.5)	8	1.6 (1.8)	8	0.83	UN+ (large)	
Saj et al. <sup>58</sup>	Longitudinal Body Axis Translation Head	2.3 (2.0)	6	-0.3 (1.4)	6	1.39	UN+ (large)	
	Longitudinal Body Axis Translation Trunk	5.9 (5.8)	6	-0.5 (1.1)	6	1.42	UN+ (large)	
	Longitudinal Body Axis Rotation Head	-4.6 (2.2)	6	-2.5 (1.5)	6	-1.03	UN+ (large)	
	Longitudinal Body Axis Rotation Trunk	-4.6 (3.3)	6	-2.3 (1.9)	6	-0.79	UN+ (medium)	
Tosi et al. <sup>66</sup>	Arm Bisection Task	69.7 (11.7)	7	67.3 (10.7)	37	0.22	UN+ (small)	
Body topography		· · /		× /			. /	
Di Vita et al. <sup>23</sup>	Body topography % Error	42.9 (27.5)	7	16 (17.2)	16	1.17	UN+ (large)	
Rousseaux et al. <sup>57</sup>	Localisation – Total Deviation	1.8 (11.4)	9	0.2 (7.8)	6	0.16	Nil	
	Localisation – Left Point Deviation	13.4 (13.2)	9	4.5 (9.5)	6	0.78	UN+ (medium)	

Table 4. Comparison of proprioceptive impairments between UN+ and UN- (continuous outcomes).

UN: unilateral neglect; UN+: participants with UN; UN-: participants without UN; SD: standard deviation; Em-NSA: Erasmus Modifications to the Nottingham Sensory Assessment; IQR: interquartile range; UTD: unable to determine; TFT: thumb finding test; L: left; R: right; NR: not reported.

Effect size determined using cut offs of 0.2 for small, 0.5 medium, and 0.8 large as reported by Lakens.<sup>49</sup>

<sup>a</sup>More impaired group determined by p values in study due to median and IQR reporting.

<sup>b</sup>Data reported for UN- group with right hemisphere damage only.

of the stimulated body parts. There is thus a clear need for a simple but comprehensive battery of proprioceptive tests to address the issues with clinical assessment of proprioception in general stroke populations. Once established, these tests could be incorporated into the clinical assessment of UN to enable identification of the multiple sensorimotor impairments present in this population.

# Limitations of the study

Adopting a broad definition of proprioception is both a strength and limitation of this review. On one hand, the unification of multiple components of an essential sensorimotor process allows more functional, and thus clinically relevant, conclusions to be drawn. However, it also limits the strength of the conclusions of this review, due to the heterogeneity in data. This is a strong argument for further research in UN that comprehensively defines proprioception.

A further limitation of this review is the small sizes of the UN+ groups in the included studies (mean=13  $SD = \pm 14$ , range=3-59). Eight studies with a collective sample size of 504 participants were excluded at full-text review due to insufficient reporting of data. All authors were contacted to request data, but the data were either unavailable or no reply was received. The inclusion of these data may change the strength of or the findings of the

Study	Proprioception outcome	N +		N-		Odds ratio (95% Cl)	More impaired group	
		% N		% N				
Movement detection								
Meyer et al. <sup>62</sup>	Em-NSA	48.I	27	15.8	95	4.95 (1.94–12.61)	UN+	
,	TFT (0–3)	77.8	27	47.4	95	3.89 (1.44–10.49)	UN+	
van Stralen et al. <sup>67</sup>	RASP-Impaired	100	9	25.7	35	53 (2.81–1001.40)	UN+	
Joint position matching								
Semrau et al. <sup>65</sup>	Robotic Arm Position Matching Task–Failure	100	35	59	123	48.78 (2.92–813.68)	UN+	
Semrau et al. <sup>64</sup>	Robotic Upper Limb Position Match–Impaired+/–vision	85	59	38	222	9.12 (4.27–19.51)	UN+	
Laterality								
van Stralen et al. <sup>67</sup>	Bergen Laterality Test Total Failure	25.7	9	11.1	35	2.21 (0.34–14.59)	Nil	
Body topography								
Cocchini et al. <sup>53</sup>	Body Exploration Fluff Test–Impaired	71.4	14	16.7	24	12.5 (2.57–60.70)	UN+	

Table 5. Comparison of proprioceptive impairments between UN+ and UN- (dichotomous outcomes).

UN: unilateral neglect; UN+: participants with UN; UN-: participants without UN; CI: confidence interval; Em-NSA: Erasmus Modifications to the Nottingham Sensory Assessment; TFT: thumb finding test; RASP: Rivermead Assessment of Somatosensory Perception.

present review. In addition, the data of this review come from studies with a predominately moderate risk of bias which limits the strength of the conclusions.

Finally, heterogeneity of studies and the reporting of data did not allow for meta-analysis. There is a clear need to establish consensus on standard assessments of both UN and proprioception in research and clinical settings to reduce heterogeneity, which would allow stronger conclusions in future reviews.

# Conclusion

We found that people with UN after stroke are more likely to have impaired processing of proprioceptive information than those without UN. These impairments occur across a variety of different subtypes of UN and aspects of proprioception. Assessment of both UN and proprioception is highly inconsistent, which likely reflects current clinical practice. Future investigations in this area should prioritise comprehensive and functional assessments of UN that include an assessment of proprioception.

# **Clinical messages**

- In UN after stroke:
  - Proprioceptive impairment is likely common and should be specifically assessed.
  - Proprioceptive impairment can present in multiple ways, including (but not limited to) deficits in movement detection and position matching but also in body representation.
  - Assessment of both UN and proprioception is frequently non-comprehensive, which likely reflects clinical practice.

#### Acknowledgements

The authors acknowledge associate professor Kris Rogers for his help in preparing this manuscript for publication.

#### **Declaration of conflicting interests**

The author(s) declared no potential conflicts of interest with respect to the research, authorship, and/or publication of this article.

#### Funding

The author(s) received no financial support for the research, authorship, and/or publication of this article.

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#### Supplemental material

Supplemental material for this article is available online.

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