Review

## Longer term patient management following stroke: A systematic review

Christian Boehme<sup>1</sup>, Thomas Toell<sup>1</sup>, Wilfried Lang<sup>2</sup>, Michael Knoflach<sup>1,3</sup> and Stefan Kiechl<sup>1,3</sup>

## Abstract

**Background:** Tremendous progress in acute stroke therapy has improved short-term outcome but part of this achievement may be lost in the long run. Concepts for a better long-term management of stroke survivors are needed to address their unmet needs and to reduce the burden of post-stroke complications, residual deficits, and recurrent vascular events.

**Aims:** This review summarizes current knowledge on post-hospital care and the scientific evidence supporting individual programs.

**Summary of review:** A systematic search of electronic databases according to PRISMA guidelines identified 10,374 articles, 77 of which met the inclusion criteria. One large randomised controlled trial on a multifaceted care program delivered by the multidisciplinary stroke team reduced recurrent vascular events and improved quality of life and functional outcome one year after the event, while a number of studies offer solutions for individual components of post-hospital disease management like patient education, counselling, and self-management or the management of post-stroke complications and residual deficits. A majority of studies, however, was small in size and limited by a short follow-up. Most initiatives with a narrow focus on risk factor control failed to lower the risk of recurrent events. The caregivers' central role in post-stroke patient management is broadly neglected in research.

**Conclusions:** Over the past years, first knowledge on how to best organize post-hospital care of stroke patients has emerged. Comprehensive and pragmatic programs operated by the multidisciplinary stroke team hold promise to reduce the long-term health burden of stroke. There is a clear need for further high-quality studies with both clinical endpoints and patient-reported outcomes to establish sustainable solutions in different settings and regions to improve life after stroke, a key priority of the Stroke Action Plan for Europe 2018–2030.

### Keywords

Stroke, post-stroke care, disease management, complication, follow-up, review

Received: 26 March 2021; accepted: 21 April 2021

## Background

Structured follow-up programs are a well-established standard of care for myocardial infarction and cancer but so far not for stroke.<sup>1,2</sup> In coronary and cancer patients, there is compelling evidence that standardized disease pathways positively affect risk factor profiles, quality of life (QoL), and functional status,<sup>1,2</sup> whereas high-quality studies testing follow-up programs for stroke patients have not been available until recently.

Stroke is a leading cause of death and disability globally. The current lifetime risk of stroke is 25% for both men and women.<sup>3</sup> There are 14 million new strokes each year and over 80 million stroke survivors with a strong upwards trajectory given the continuous ageing of societies, population growth, and declining stroke fatality.<sup>4</sup> Exciting recent advances in acute stroke therapy translate into improved short-term outcome<sup>5</sup> which, however, may be lost in the long run through unmet needs in post-stroke care, post-stroke complications, residual deficits, and recurrent vascular events. Stroke is considered preventable by guideline-compliant control of risk factors and adequate vascular

#### **Corresponding author:**

Stefan Kiechl, Department of Neurology, Medical University of Innsbruck, Innsbruck, Austria.

Email: stefan.kiechl@i-med.ac.at

International Journal of Stroke 2021, Vol. 16(8) 917-926 © 2021 World Stroke Organization

## CC 0 S

Article reuse guidelines: sagepub.com/journals-permissions DOI: 10.1177/17474930211016963 journals.sagepub.com/home/wso





<sup>&</sup>lt;sup>1</sup>Department of Neurology, Medical University of Innsbruck, Innsbruck, Austria

<sup>&</sup>lt;sup>2</sup>Department of Neurology, Hospital St. John's of God, Vienna, Austria <sup>3</sup>VASCage, Research Centre on Vascular Ageing and Stroke, Innsbruck, Austria

Concepts for a better long-term management of stroke patients are urgently needed and the Stroke Action Plan for Europe 2018–2030 has declared life after stroke a key priority.<sup>7</sup> We herein report a systematic review of randomised controlled trials and rigorous scientific evaluations of structured multimodal intervention programs targeting longer term care of stroke patients.

## Methods

## Study design

This systematic review was conducted according to PRISMA guidelines.<sup>8</sup> It included peer-reviewed, qualitative and quantitative articles in English or German language without restrictions in time frame or countries.

## Search strategy and selection criteria

We systematically searched PubMed and the Cochrane Library and extracted articles from the start of each database until 29 January 2021. Articles were eligible if (a) they comprised multimodal post-stroke care programs, (b) the interventions were initiated within one year after stroke, and (c) the outcome was specified and assessed during follow-up. The review's focus is on stroke patients but several studies also enrolled TIApatients. We did not consider studies with a primary focus on rehabilitation programs or specific rehabilitation approaches, early supported discharge, individual post-stroke complications and prevention or therapy thereof, psychological support and social participation. The search strategy and terms are detailed in the Online Supplement (Supplementary Table 1). We used the following filters: clinical trial, randomised controlled trial (RCT), meta-analysis, review, systematic review, and cohort study. The clinicaltrials.gov website was scanned for the same search terms to detect ongoing and unpublished studies. All articles identified were screened for relevant content in the title and abstract by CB and TT. Duplicates and study protocols were excluded. The shortlist of remaining articles was read in full and the citations searched for additional relevant publications.

## Results

In total, 10,363 articles were retrieved from the database search and 11 additional records identified

through check of citations and the clinicaltrials.gov website. Of 10,374 records, 99 were appropriate for the full-text review and 77 met the eligibility criteria. (Figure 1) Of the studies included, 41 were RCTs and 24 other types of studies and previous reviews. (Figures 2 and 3, Supplementary Table 2) Twelve additional studies have not reported final results, most are still ongoing (Supplementary Figure 1).

## Large multifaceted RCTs

The SMART study (n = 3821, China) implemented a structured treatment and lifestyle modification algorithm involving stroke specialists. This in-hospital intervention did not show long-term effects on risk factor levels nor reductions in recurrent vascular events.9 Two large European RCTs have been finished **INSPIRE-TMS** (n = 2098,recently: Germany/ Denmark) employed a patient support program with up to eight outpatient visits including feedback and motivational interviews and repeated risk factor assessments to improve medication adherence. This trial included TIA and minor stroke patients and improved achievement of recommended prevention targets but failed to reduce recurrent vascular events during a follow-up of 3.6 years. In a post hoc analysis, patients attending more than half of scheduled appointments were less likely to suffer an outcome event.<sup>10</sup> The STROKE-CARD Study (n = 2149, Austria) included stroke and TIA patients except for patients with severe permanent disability. This study extended the responsibility of the multidisciplinary stroke team from acute therapy up to three months after the event, when a comprehensive outpatient re-evaluation was performed. It focused on residual deficits and additional rehabilitation demands, screening and treatment of post-stroke complications, management of vascular warning signs, re-evaluation of stroke aetiology, risk factor control, and patient/caregiver counselling. This study reduced recurrent vascular events and improved QoL (co-primary endpoints) as well as functional outcome (secondary endpoint) at 12 months.<sup>11</sup> However, it did not improve 12-month risk factor profiles compared with the control group, yet risk factor control was overall better than in real-world surveys from Europe.<sup>12</sup>

## Studies with a primary focus on risk factors

A majority of studies had their key focus on risk factor control which served as the primary study outcome. The ICARUSS program (n = 186, Australia) comprised four pre-scheduled three-monthly visits to the GP with constant advice by stroke specialists and improved



target level achievement and functional outcome.<sup>13,14</sup> The SOS project investigated a case manager-led pathway with educational sessions and regular check-ups. Pilot data on 45 patients revealed better risk factor control, less vascular events, and improved functional outcomes at 12 months. This pathway will be evaluated in a larger cohort study.<sup>15</sup> Another study (n = 70,Japan) enrolling minor stroke patients comprised salt restriction and exercise training offered by physical therapists once or twice per week for six months.<sup>16</sup> It improved risk factor control and reduced vascular events during a follow-up of almost three years. A Norwegian study (n = 195) implementing a comprehensive risk factor analysis and two lifestyle counselling visits reported improved risk factor control after one year.<sup>17</sup> In the NAILED trial (n = 537, Sweden), nurses performed a telephone-based risk factor assessment and counselling one month after discharge and a study physician adjusted pharmacological treatment if necessary. This procedure was repeated monthly until risk factor control was achieved. The program achieved sustained improvements in blood pressure and cholesterol levels up to three years.<sup>18,19</sup> In the PREVENTION trial (n=279, Canada), case management by pharmacists substantially improved risk factor control six months

after stroke (compared with nurse-led case management) but this did not translate into less recurrent vascular events.<sup>20</sup> The PRAISE trial (n = 600, USA) tested potential effects of a group workshop intervention after discharge. The program reduced blood pressure on short notice but had no effect on other risk factors or medication adherence.<sup>21</sup> The MIST trial (n = 386, New Zealand) reported improved medication adherence but no sustained improvement in risk profiles and no cardiovascular disease benefit.<sup>22</sup> The STANDFIRM trial (n = 563, Australia), a multidisciplinary team approach including nurse-led home visits, showed no effects on risk factor control but patients were randomised in the post-acute phase and protocol adherence was suboptimal.<sup>23</sup> A long-term follow-up of STANDFIRM accomplished lower LDL-cholesterol levels at 12 months.<sup>24</sup> The Stop Stroke trial (n = 523, UK), providing electronic advice to patients, carers, and GPs failed to demonstrate effects on risk factors<sup>25</sup> as did a nurse-led intervention (n = 349, Denmark) based on four home visits for blood pressure measurement and counselling.<sup>26</sup> Four previous reviews on lifestyle interventions after stroke/TIA reported moderate reductions in systolic blood pressure but less consistent effects on other risk factors. Most studies considered had short

	Study acronym or author	hor Year n Intervention lead			Intervention focus			Outcomes				
-					Risk factors	Compli- cations	Education or self manage- ment	Risk factor control	CVD	Quality of life	Func- tional outcome	Care- giver outcome
-	OSCARSS	2021	414	Trained researcher			x			x		x
	SHARE	2021	48	Nurse			x					x
	STROKE-CARD	2020	2,149	Multidisciplinary	x	x		x	x	x	x	
	INSPIRE-TMS	2020	2,098	Nurse/Physician	x			x	x			
	ICARUSS	2020	249	Physician/GP	x	x		x			x	
	iCaPPS	2020	151	Physician	x	x		x		x		
	TACAS	2020	400	Research clinician			x			x	x	
	BISC	2020	35	Psychologist			x			x	×	x
	GOTVED	2019	140	Multidisciplinary		x				x	x	
	EXTRAS	2019	573	Multidisciplinary						x		
	Fukuoka et al	2019	321	Nurse/GP	x		x	x	x			
	LAS-2	2019	280	Therapist		x				x		x
	NAILED follow-up	2018	660	Nurse	x			x				
	LAST	2018	380	Physical therapist			x		×	x	x	x
	SUSTAIN	2018	404	Nurse	x	x	x	x	x			
	Lo et al	2018	128	Nurse			x			x		
	STANDFIRM	2017	563	Nurse/GP	x			x	x			
	STANDFIRM extension	2017	563	Nurse/GP	x			х				
	Damush et al	2016	174	Nurse	x		x	x		x		
	CARE4STROKE	2016	63	Physical therapist		x				x	x	x
	NAILED	2015	537	Nurse	x			х				
	MIST	2015	386	Research assistant	x			x	x	x		
	Forster et al	2015	800	HCP		x				x	x	x
	Restore4Stroke	2015	113	Therapists		x	x			x		x
	McKenna et al	2015	25	Community team			x			×		
	Saal et al	2015	265	HCP		x	x		×	×		
	SMART	2014	3,821	Physician	×			x	x			
	Ihle-Hansen et al	2014	195	Physician/Nurse	×			x				
	PRAISE	2014	600	Research assistant	×	x		x				
	PREVENTION	2014	279	Pharmacist	x			х				
	Kono et al	2013	70	Physical therapist	x			x	x			
	YOU CALL-WE CALL	2013	186	HCP		x				x		
	Kim et al	2013	36	Research assistant	x	x	x	х				x
	TRACS	2013	928	Multidisciplinary			x			x		x
	Cadilhac et al	2011	143	HCP		x	x			x		
	Hornnes et al	2011	349	Nurse	×			x				
	Stop Stroke	2010	523	Trained researcher	×			x				
	ICARUSS pilot	2009	186	Physician/GP	×			x		x	×	
	Forster et al	2009	265	Nurse		x					x	x
	Allen et al	2009	380	Nurse	×	x		x		x		
	Sit et al	2007	190	Nurse			х			x		

## Figure 2. Summary of randomised controlled trials included in the review.

Note: Bold letters (X) indicate the primary outcome and standard letters (x) secondary outcomes. Green-coloured cells indicate significant effects of the intervention, yellow cells indicate mixed effects, and orange cells indicate no benefit. *n*: number of participants; CVD: cardiovascular events; GP: general practitioner; HCP: healthcare professional.

follow-ups that only exceptionally exceeded six months. Interventions lasting longer than four months and using multiple techniques to achieve behavioural change were more effective. (Supplementary Table 2)

# Studies targeting post-stroke complications and disability

A total of 23 studies employed standardized follow-up visits and screened for post-stroke complications

Study acronym or author	Pub. Year	n	Design	Intervention lead	Intervention Focus			Outcomes					
					Risk factors	Compli- cations	Education or self manage- ment	Risk factor control	CVD	Quality of life	Func- tional outcome	Care- giver outcome	
Struct-FU	2020	200	CS	Nurse		x		x			x		
AVERT	2020	668	CS	Physiotherapist Nurse							x		
Rudberg et al	2018	297	CS	Physician						x	x		
SOS	2016	45	CCS	Case manager	x			x	x	x	x		
Willeit et al	2015	4,947	CS	Multidisciplinary							x		
Fens et al	2014	144	СТ	Nurse			x			x		x	
Riks-Stroke	2008	15,959	CS	N/A							x		
Slot et al	2008	7,710	CS	Nurse							x		
Bravata et al	2007	2,603	CS	N/A					x				
Pedersen et al	2020	2,216	Review			x				x			
Zawawi et al	2020	102*	Review			x				x		x	
Chen et al	2019	19*	Review			x				x			
Denham et al	2019	13*	Review									x	
Vellipuram et al	2019	N/A	Review					x					
Bridgwood et al	2018	33,840	Review				x	x					
Wray et al	2018	24*	Review							x	x		
Hempler et al	2018	18*	Review									x	
Fryer et al	2016	1,863	Review							x			
Deijle et al	2016	2,574	Review					x	x				
Parke et al	2015	101*	Review							x	x	x	
Lawrence et al	2015	6,373	Review					x	x				
MacKay-Lyons et al	2013	1*	Review					x	x				
Fens et al	2013	14*	Review							x			
Redfern et al	2006	67*	Review			x		x		x		×	

#### Figure 3. Summary of other studies included in the review.

Note: Bold letters (X) indicate the primary outcome and standard letters (x) secondary outcomes. Green-coloured cells indicate significant effects of the intervention and orange cells indicate no benefit. \*Indicates number of studies included in a review. *n*: number of participants; CS: cohort study; CCS: case-control study; CT: controllectrial.

and/or residual deficits. Post-discharge interventions including nurse-led, telephone-based, or more complex multidisciplinary approaches showed positive effects on anxiety and functional outcomes. Benefits, however, were mainly modest and partly disappeared over time. The Struct-FU study (n = 200, Sweden)<sup>27</sup> systematically applied a modified post-stroke checklist<sup>28</sup> and provided tailored advice, further diagnostic work-up, and therapeutic referrals. Over 90% of participants reported stroke-related health problems and over 80% required new interventions underscoring the relevance of an early comprehensive follow-up.<sup>27</sup> Two reviews suggested that treatment recommendations are often not implemented but documented positive effects on

QoL especially if follow-up visits were combined with care and rehabilitation interventions. (Supplementary Table 2)

## Studies focused on educational support or self-management programs

Seven studies directed main attention on educational support. TACAS (n = 400, New Zealand) organized one or two home visits with an educational and motivational intervention to patients and family members six weeks after stroke. This simple measure resulted in better QoL and improved functional outcomes after 6–12 months. Interestingly, the intervention emerged

as particularly favourable in patients living alone or with a permanent support person and in female patients.<sup>29</sup> A nurse-led program early after discharge in patients with minor stroke (n = 190, Hong Kong) elicited positive effects on behavioural issues and medication compliance,<sup>30</sup> another study (n = 36, Republic of Korea) offering web-based education (regular use, 63%) showed benefits on health behaviours and caregiver mastery but not on risk factor control.<sup>31</sup> The LAST Study (n = 380, Norway) relied on individual coaching and observed a trend towards less vascular events during long-term follow-up.<sup>32</sup> Other studies relying on home visits or web-based interventions and outpatient group sessions showed no effects on QoL, functional status, and risk factor control.

Most initiatives offering self-management programs to stroke survivors reported on short-term outcomes and excluded patients with severe persistent deficits or aphasia. Five studies operating educational sessions, telephone calls, or group sessions showed modest short-term effects on mental health, QoL, and medication adherence. Reviews on self-management programs in specific settings (e.g. in-patient rehabilitation) found improvements in activities of daily living and QoL if the programs were initiated early after stroke.

### Studies reporting caregiver outcome

Few studies addressed caregiver outcomes. Three intervention trials (n < 150, each) with a duration up to 18 months demonstrated small favourable effects on caregiver depression and fatigue by continued rehabilitation in their homes with a multidisciplinary team, caregiver exercise programs, or regular home visits to screen for healthcare issues. LAS-2 (n = 280, Sweden) showed positive effects on depression even after five years.<sup>33</sup> Benefits for caregivers translated into less readmissions of the cared patients in one of the trials.<sup>34</sup> Several other trials, however, reported null effects on QoL and caregiver burden including one well-designed large RCT (TRACS, n = 928).<sup>35</sup> Reviews on caregivers' unmet needs agree that support by healthcare professionals is insufficient. More direct support in caring for the patients and for themselves, and a better access to training and healthcare services are warranted (Supplementary Table 2).

## Ongoing and unpublished studies

Three large trials with multifaceted post-stroke interventions, a variety of outcomes, and in part longer follow-up periods are expected to further inform about post-stroke care in near future (Supplementary Figure 1). The COMPASS-trial in the US aims to include 6,000 stroke patients with a follow-up of three months. The intervention involves a telephone call after two days and a visit to the post-acute stroke clinic after 7–14 days to assess early problems after discharge. Outcomes are QoL, post-stroke complications, mortality, functional status, and caregiver strain.<sup>36</sup> Several studies with a focus on various clinical outcomes are still recruiting and two trials have been completed but are still unpublished (Supplementary Figure 1).

## Discussion

Until now, no country in Europe is operating an evidence-based nationwide reimbursed post-stroke care pathway in spite of the many unmet needs for stroke survivors and their caregivers. Recurrent stroke and cardiovascular disease, post-stroke complications, residual neurological deficits, and inadequate medical and psychosocial care all contribute to long-term disability and OoL. A Swedish study revealed stroke-related health problems in 90% of stroke survivors necessitating medical interventions in more than 80%.27 Moreover, half of stroke patients are readmitted to hospital (or die) within the first year.<sup>37</sup> The AVERT trial reported disappointing rehabilitation referral in working-age patients with minor stroke and only 6 of 10 patients finally returned to work.<sup>38</sup> On the other hand, good functional outcome three to six months after stroke predicts long-term survival and results in better QoL and lower healthcare costs.39,40 These findings altogether highlight the huge burden of post-stroke morbidity and suggest a window of opportunity for post-stroke care pathways early after stroke.

The Stroke Action Plan for Europe 2018–2030 gives life after stroke a key priority and claims for (a) national care pathways encompassing the entire chain from primary prevention to long-term care (overarching goal), (b) obligatory reviews of rehabilitation and patient needs three to six months after stroke and annually thereafter, and (c) broad expert support to patients and carers.<sup>7</sup> Our systematic review summarizes current knowledge on post-stroke care and the evidence supporting individual programs. It indicates substantial under-research in this field compared to acute stroke therapy and long-term care of other major diseases but also emerging research interests in the last years. One large RCT (STROKE-CARD) successfully reduced the risk of recurrent stroke and cardiovascular disease and improved QoL and functional outcome.<sup>11</sup> This study considered multiple important aspects of post-stroke care (Supplementary Figure 2) and extended the responsibility of the multidisciplinary stroke team from acute management until the threemonth intervention. Low-threshold contact to the team enabled frequent detection of vascular warning signs in this early vulnerable period after stroke (5% of all patients) and contributed to the prevention of vascular events. The pragmatic STROKE-CARD concept is suitable for routine application and reimbursement has been applied for in Austria to enable nation-wide implementation. Limitations are the short follow-up of one year and exclusion of patients with permanent severe disability who may also benefit from intensified care. A long-term extension of the RCT is currently underway (NCT04205006) and the ongoing STROKE-CARD registry will create evidence patients with the most severe strokes for (NCT04582825). A number of other promising RCTs on multifaceted care programs like COMPASS<sup>36</sup> (Supplementary Figure 1) will finish soon and inform about the optimal organization of post-stroke care in various settings.

The largest share of previous studies had a primary focus on risk factor control and demonstrated improvements in risk profiles, especially systolic blood pressure. Interventions longer than four months with multiple contacts to patients, involvement of stroke experts, and use of multiple techniques to achieve behavioural change were more effective.<sup>10,13–15,18–20,41,42</sup> However, the few studies that reported effects on recurrent stroke and cardiovascular disease yielded disappointing findings.<sup>9,10,22,23,32,43</sup> It is important to emphasize that these studies do not disregard the relevance of secondary stroke prevention but rather reflect short follow-up periods (effects of preventive measure increase over time) and the fact that patient enrolment per se improves medication adherence in both the intervention and control group. Accordingly, risk factor levels in the control groups of these trials were usually better than in real-world observational studies rendering it difficult to achieve additional benefits.<sup>10-12</sup>

A growing number of studies moved beyond risk factor management and addressed additional aspects of post-stroke care. Systematic screening for poststroke complications and residual deficits has a considerable diagnostic yield. A majority of patients require tailored advice, further diagnostic work-up, and/or therapeutic and rehabilitation referrals.<sup>27</sup> In the STROKE-CARD trial, detection and management of post-stroke complications like anxiety, depression, fatigue, and attainment of additional rehabilitation demands contributed to improved functional outcome and QoL. Other studies revealed positive effects on anxiety, QoL, and activities of daily living, especially if follow-up visits were combined with care and rehabilitation interventions.<sup>29,33</sup> This evidence endorse the recommendation of the Stroke Action Plan for Europe 2018–2030 to establish an obligatory three to six month review of stroke patients.<sup>7</sup> Patient-reported outcomes on QoL and mental health should be included in this review. Collection of robust data on health

Educational support involving targeted patient counselling and self-management programs are further appealing components of post-stroke care. The TACAS trial made a proof of concept that a pragmatic intervention with one or two home visits to conduct a motivational interview and verbalize the unmet needs of patients and caregivers translates into long-term benefits regarding QoL and functional outcome.<sup>29</sup> Self-management programs initiated early after stroke accomplished short-term improvements in activities of daily living, QoL, and medication adherence but most of these programs were not delivered to stroke patients with severe disability or aphasia.

The caregivers' central role in patient management is broadly neglected in research. Few studies have addressed caregiver outcomes. Few studies including LAS-2 reported favourable effects on caregiver depression and fatigue<sup>33</sup> and in one of these trials, benefits for caregivers translated into less hospital readmissions of the cared patients.<sup>34</sup> However, the only large RCT (TRACS) failed to achieve improvements.<sup>35</sup> More information on support in caring for the patients and caring for themselves as well as a better access to training and healthcare services are warranted to improve the carers' situation. There are unmet service needs and insufficient support delivered by healthcare professionals. Furthermore, the combination of behavioural interventions with improvements on the organizational level (i.e. implementation of multidisciplinary poststroke care) seems to be crucial.<sup>41</sup>

## Limitations

We acknowledge that our selected search terms might not have detected all relevant articles, this may have affected the results. Though, by using two databases and the clinicaltrials.gov website as well as the reference lists, the risk for publication and reporting bias is low.

We did not consider programs targeting individual post-stroke complications including dementia, which is a major issue in stroke survivors,<sup>44</sup> and were not able to address all relevant aspects of post-stroke care.

Most intervention programs discussed herein were limited by short follow-ups compared to programs in myocardial infarction and cancer patients. Some studies might have yielded positive effects with longer follow-up periods, especially studies focusing on risk factor control.

## Summary

Stroke care pathways improved patient outcome and currently cover acute stroke therapy<sup>45</sup> or stroke care at

large including rehabilitation<sup>46</sup> but not post-stroke care. This systematic review identified one successful comprehensive post-stroke care program that extended the role of the multidisciplinary stroke team until three months after the event and a number of studies targeting individual important components of longer term post-stroke care like systematic screening for post-stroke complications and additional rehabilitation demands, refinement of secondary prevention, educational activities, patient counselling, and self-management programs. Most of these studies, however, were small in size, offered limited follow-up periods, and yielded partly inconsistent results. The carers' needs and interests are substantially under-researched and broadly neglected in clinical routine. There is an urgent need for further high-quality RCTs with both clinical and patient-reported outcomes to further advance post-stroke care, establish sustainable solutions in different settings and regions, and improve life after stroke in the long run. Implementation of pragmatic nationwide long-term care programs with proper re-imbursement, involvement of the multidisciplinary stroke team, and continuous quality control holds great promise to reduce poststroke morbidity and mortality and to meet the demands of stroke survivors and their caregivers, a key priority of the Stroke Action Plan for Europe 2018–2030.<sup>7</sup>

#### **Authors' contributions**

CB was involved in study design, screened papers for the review and prepared the manuscript. TT evaluated the screened articles and substantially revised the manuscript. WL and MK substantially revised the manuscript. SK was involved in study design, supervised the analysis and revised the final version of the manuscript.

#### **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.

#### **ORICID** iDs

Christian Boehme Dhttps://orcid.org/0000-0003-1369-418X Stefan Kiechl Dhttps://orcid.org/0000-0002-9836-2514

#### Supplemental material

Supplemental material for this article is available online.

#### References

1. Clark AM, Hartling L, Vandermeer B and McAlister FA. Meta-analysis: secondary prevention programs for patients with coronary artery disease. Ann Intern Med 2005; 143: 659-672.

- Jacobs LA and Shulman LN. Follow-up care of cancer survivors: challenges and solutions. *Lancet Oncol* 2017; 18: e19–29.
- GBD 2016 Lifetime Risk of Stroke Collaborators. Global, regional, and country-specific lifetime risks of stroke, 1990 and 2016. N Engl J Med 2018; 379: 2429–2437.
- GBD 2016 Neurology Collaborators. Global, regional, and national burden of neurological disorders, 1990-2016: a systematic analysis for the Global Burden of Disease Study 2016. *Lancet Neurol* 2019; 18: 459–480.
- Prabhakaran S, Ruff I and Bernstein RA. Acute stroke intervention: a systematic review. JAMA 2015; 313: 1451–1462.
- 6. Boehme C, Toell T, Mayer L, et al. The dimension of preventable stroke in a large representative patient cohort. *Neurology* 2019; 93: e2121–2132.
- Norrving B, Barrick J, Davalos A, et al. Action plan for stroke in Europe 2018–2030. *Eur Stroke J* 2018; 3: 309–336.
- 8. Liberati A, Altman DG, Tetzlaff J, et al. Preferred reporting items for systematic reviews and meta-analyses: the PRISMA statement. *BMJ* 2009; 339: b2700.
- Peng B, Ni J, Anderson CS, et al. Implementation of a structured guideline-based program for the secondary prevention of ischemic stroke in China. *Stroke* 2014; 45: 515–519.
- Ahmadi M, Laumeier I, Ihl T, et al. A support programme for secondary prevention in patients with transient ischaemic attack and minor stroke (INSPiRE-TMS): an open-label, randomised controlled trial. *Lancet Neurol* 2020; 19: 49–60.
- 11. Willeit P, Toell T, Boehme C, et al. STROKE-CARD care to prevent cardiovascular events and improve quality of life after acute ischaemic stroke or TIA: a randomised clinical trial. *EClinicalMedicine* 2020; 25: 100476.
- Heuschmann PU, Kircher J, Nowe T, et al. Control of main risk factors after ischaemic stroke across Europe: data from the stroke-specific module of the EUROASPIRE III survey. *Eur J Prev Cardiol* 2015; 22: 1354–1362.
- Joubert J, Davis SM, Donnan GA, et al. ICARUSS: an effective model for risk factor management in stroke survivors. *Int J Stroke* 2020; 15: 438–453.
- Joubert J, Reid C, Barton D, et al. Integrated care improves risk-factor modification after stroke: initial results of the Integrated Care for the Reduction of Secondary Stroke model. *J Neurol Neurosurg Psychiatry* 2009; 80: 279–284.
- Bodechtel U, Barlinn K, Helbig U, et al. The stroke east Saxony pilot project for organized post-stroke care: a case-control study. *Brain Behav* 2016; 6: e00455.
- Kono Y, Yamada S, Yamaguchi J, et al. Secondary prevention of new vascular events with lifestyle intervention in patients with noncardioembolic mild ischemic stroke: a single-center randomized controlled trial. *Cerebrovasc Dis* 2013; 36: 88–97.

- Ihle-Hansen H, Thommessen B, Fagerland MW, et al. Multifactorial vascular risk factor intervention to prevent cognitive impairment after stroke and TIA: a 12-month randomized controlled trial. *Int J Stroke* 2014; 9: 932–938.
- 18. Irewall A-L, Ögren J, Bergström L, Laurell K, Söderström L and Mooe T. Nurse-led, telephone-based, secondary preventive follow-up after stroke or transient ischemic attack improves blood pressure and LDL cholesterol: results from the first 12 months of the randomized, controlled NAILED stroke risk factor trial. *PLoS One* 2015; 10: e0139997.
- Ögren J, Irewall A-L, Söderström L and Mooe T. Longterm, telephone-based follow-up after stroke and TIA improves risk factors: 36-month results from the randomized controlled NAILED stroke risk factor trial. *BMC Neurol* 2018; 18: 153.
- McAlister FA, Majumdar SR, Padwal RS, Fradette M, Thompson A, Buck B, et al. Case management for blood pressure and lipid level control after minor stroke: PREVENTION randomized controlled trial. *CMAJ* 2014; 186: 577–584.
- Kronish IM, Goldfinger JZ, Negron R, et al. Effect of peer education on stroke prevention: the prevent recurrence of all inner-city strokes through education randomized controlled trial. *Stroke* 2014; 45: 3330–3336.
- 22. Barker-Collo S, Krishnamurthi R, Witt E, et al. Improving adherence to secondary stroke prevention strategies through motivational interviewing: randomized controlled trial. *Stroke* 2015; 46: 3451–3458.
- Olaiya MT, Kim J, Nelson MR, et al. Effectiveness of a shared team approach between nurses and doctors for improved risk factor management in survivors of stroke: a cluster randomized controlled trial. *Eur J Neurol* 2017; 24: 920–928.
- 24. Olaiya MT, Cadilhac DA, Kim J, et al. Communitybased intervention to improve cardiometabolic targets in patients with stroke: a randomized controlled trial. *Stroke* 2017; 48: 2504–2510.
- 25. Wolfe CDA, Redfern J, Rudd AG, Grieve AP, Heuschmann PU and McKevitt C. Cluster randomized controlled trial of a patient and general practitioner intervention to improve the management of multiple risk factors after stroke: stop stroke. *Stroke* 2010; 41: 2470–2476.
- 26. Hornnes N, Larsen K and Boysen G. Blood pressure 1 year after stroke: the need to optimize secondary prevention. *J Stroke Cerebrovasc Dis* 2011; 20: 16–23.
- Ullberg T, Månsson K, Berhin I and Pessah-Rasmussen H. Comprehensive and structured 3-month stroke followup using the post-stroke checklist (The Struct-FU study): a feasibility and explorative study. *J Stroke Cerebrovasc Dis* 2021; 30: 105482.
- Philp I, Brainin M, Walker MF, et al. Development of a poststroke checklist to standardize follow-up care for stroke survivors. *J Stroke Cerebrovasc Dis* 2013; 22: e173–180.
- 29. Fu V, Weatherall M, McPherson K, et al. Taking charge after stroke: a randomized controlled trial of a person-

centered, self-directed rehabilitation intervention. Int J Stroke 2020; 15: 954–964.

- Sit JWH, Yip VYB, Ko SKK, Gun APC and Lee JSH. A quasi-experimental study on a community-based stroke prevention programme for clients with minor stroke. *J Clin Nurs* 2007; 16: 272–281.
- Kim JI, Lee S and Kim JH. Effects of a web-based stroke education program on recurrence prevention behaviors among stroke patients: a pilot study. *Health Educ Res* 2013; 28: 488–501.
- 32. Askim T, Langhammer B, Ihle-Hansen H, et al. Efficacy and safety of individualized coaching after stroke: the LAST study (life after stroke): a pragmatic randomized controlled trial. *Stroke* 2018; 49: 426–432.
- 33. van den Berg M, Crotty M, Liu E, Killington M, Kwakkel G and van Wegen E. Early supported discharge by caregiver-mediated exercises and e-health support after stroke: a proof-of-concept trial. *Stroke* 2016; 47: 1885–1892.
- 34. Hedman A, Eriksson G, von Koch L and Guidetti S. Five-year follow-up of a cluster-randomized controlled trial of a client-centred activities of daily living intervention for people with stroke. *Clin Rehabil* 2019; 33: 262–276.
- Forster A, Dickerson J, Young J, et al. A structured training programme for caregivers of inpatients after stroke (TRACS): a cluster randomised controlled trial and cost-effectiveness analysis. *Lancet* 2013; 382: 2069–2076.
- Duncan PW, Bushnell CD, Rosamond WD, et al. The comprehensive post-acute stroke services (COMPASS) study: design and methods for a cluster-randomized pragmatic trial. *BMC Neurol* 2017; 17: 133.
- Bravata DM, Ho SY, Meehan TP, Brass LM and Concato J. Readmission and death after hospitalization for acute ischemic stroke: 5-year follow-up in the medicare population. *Stroke* 2007; 38: 1899–1904.
- 38. Walters R, Collier JM, Braighi Carvalho L, et al. Exploring post acute rehabilitation service use and outcomes for working age stroke survivors (≤65 years) in Australia, UK and South East Asia: data from the international AVERT trial. *BMJ Open* 2020; 10: e035850.
- Eriksson M, Norrving B, Terént A and Stegmayr B. Functional outcome 3 months after stroke predicts long-term survival. *Cerebrovasc Dis* 2008; 25: 423–429.
- Slot KB, Berge E, Dorman P, et al. Impact of functional status at six months on long term survival in patients with ischaemic stroke: prospective cohort studies. *BMJ* 2008; 336: 376–379.
- Bridgwood B, Lager KE, Mistri AK, Khunti K, Wilson AD and Modi P. Interventions for improving modifiable risk factor control in the secondary prevention of stroke. *Cochrane Database Syst Rev* 2018; 5: CD009103.
- 42. Deijle IA, Van Schaik SM, Van Wegen EEH, Weinstein HC, Kwakkel G and Van den Berg-Vos RM. Lifestyle interventions to prevent cardiovascular events after stroke and transient ischemic attack: systematic review and meta-analysis. *Stroke* 2017; 48: 174–179.

- 43. Fukuoka Y, Hosomi N, Hyakuta T, et al. Effects of a disease management program for preventing recurrent ischemic stroke. *Stroke* 2019; 50: 705–712.
- 44. Matz K, Teuschl Y, Firlinger B, et al. Multidomain lifestyle interventions for the prevention of cognitive decline after ischemic stroke: randomized trial. *Stroke* 2015; 46: 2874–2880.
- 45. Maas WJ, Lahr MMH, Buskens E, et al. Pathway design for acute stroke care in the era of endovascular

thrombectomy: a critical overview of optimization efforts. *Stroke* 2020; 51: 3452–3460.

46. Willeit J, Geley T, Schöch J, et al. Thrombolysis and clinical outcome in patients with stroke after implementation of the Tyrol Stroke Pathway: a retrospective observational study. *Lancet Neurol* 2015; 14: 48–56.