

OPEN ACCESS

EDITED BY

Valerie Moyra Pomeroy, University of East Anglia, United Kingdom

REVIEWED BY

Vahideh Moradi,

Iran-Helal Institute of Applied Science and Technology,

Iran

Meizhen Huang,

Hong Kong Polytechnic University, Hong Kong SAR, China

*CORRESPONDENCE

Yulong Wei

⊠ wyl_5128@163.com

[†]These authors share first authorship

SPECIALTY SECTION

This article was submitted to Stroke, a section of the journal Frontiers in Neurology

RECEIVED 09 November 2022 ACCEPTED 01 March 2023 PUBLISHED 03 April 2023

CITATION

Lyu T, Yan K, Lyu J, Zhao X, Wang R, Zhang C, Liu M, Xiong C, Liu C and Wei Y (2023) Comparative efficacy of gait training for balance outcomes in patients with stroke: A systematic review and network meta-analysis. Front. Neurol. 14:1093779. doi: 10.3389/fneur.2023.1093779

COPYRIGHT

© 2023 Lyu, Yan, Lyu, Zhao, Wang, Zhang, Liu, Xiong, Liu and Wei. This is an open-access article distributed under the terms of the Creative Commons Attribution License (CC BY).

The use, distribution or reproduction in other forums is permitted, provided the original author(s) and the copyright owner(s) are credited and that the original publication in this journal is cited, in accordance with accepted academic practice. No use, distribution or reproduction is permitted which does not comply with these terms.

Comparative efficacy of gait training for balance outcomes in patients with stroke: A systematic review and network meta-analysis

Tianyi Lyu^{1†}, Kang Yan^{1†}, Jiaxuan Lyu¹, Xirui Zhao¹, Ruoshui Wang¹, Chaoyang Zhang¹, Meng Liu¹, Chao Xiong², Chengjiang Liu³ and Yulong Wei^{1*}

¹School of Acupuncture-Moxibustion and Tuina, Beijing University of Chinese Medicine, Beijing, China, ²L3 & Maintenance Solutions, SUSE Software (Beijing) Co., Ltd., Beijing, China, ³Department of General Medicine, Affiliated Anging First People's Hospital of Anhui Medical University, HeFei, Anhui, China

Background: Growing evidence suggests that gait training can improve stroke patients' balance outcomes. However, it remains unclear which type of gait training is more effective in improving certain types of balance outcomes in patients with stroke. Thus, this network meta-analysis (NMA) included six types of gait training (treadmill, body-weight-supported treadmill, virtual reality gait training, robotic-assisted gait training, overground walking training, and conventional gait training) and four types of balance outcomes (static steady-state balance, dynamic steady-state balance, proactive balance, and balance test batteries), aiming to compare the efficacy of different gait training on specific types of balance outcomes in stroke patients and determine the most effective gait training.

Method: We searched PubMed, Embase, Medline, Web of Science, and Cochrane Library databases from inception until 25 April 2022. Randomized controlled trials (RCTs) of gait training for the treatment of balance outcomes after stroke were included. RoB2 was used to assess the risk of bias in the included studies. Frequentist random-effects network meta-analysis (NMA) was used to evaluate the effect of gait training on four categories of balance outcomes.

Result: A total of 61 RCTs from 2,551 citations, encompassing 2,328 stroke patients, were included in this study. Pooled results showed that body-weight-support treadmill (SMD=0.30, 95% CI [0.01, 0.58]) and treadmill (SMD=0.25, 95% CI [0.00, 0.49]) could improve the dynamic steady-state balance. Virtual reality gait training (SMD=0.41, 95% CI [0.10, 0.71]) and body-weight-supported treadmill (SMD=0.41, 95% CI [0.02, 0.80]) demonstrated better effects in improving balance test batteries. However, none of included gait training showed a significant effect on static steady-state balance and proactive balance.

Conclusion: Gait training is an effective treatment for improving stroke patients' dynamic steady-state balance and balance test batteries. However, gait training had no significant effect on static steady-state balance and proactive balance. To achieve maximum efficacy, clinicians should consider this evidence when recommending rehabilitation training to stroke patients. Considering bodyweight-supported treadmill is not common for chronic stroke patients in clinical practice, the treadmill is recommended for those who want to improve dynamic steady-state balance, and virtual reality gait training is recommended for those who want to improve balance test batteries.

Limitation: Missing evidence in relation to some types of gait training is supposed to be taken into consideration. Moreover, we fail to assess reactive balance in this NMA since few included trials reported this outcome.

Systematic Review Registration: PROSPERO, identifier CRD42022349965.

KEYWORDS

stroke, gait training, network meta-analysis, randomized controlled trials, balance outcomes

Introduction

Recently, stroke is the leading cause of death in China and the second leading cause of death worldwide (1, 2). Despite the fact that stroke mortality, prevalence, and incidence have decreased in the past 20 years, its prevalence is increasing in young individuals (3). Thus, the significance of stroke rehabilitation has grown. Balance disorders, one of the most common symptoms after a stroke, can affect patients' physiological and social functions (4, 5). Therefore, balance disorders place a heavy burden on both individuals and society.

Balance is one of the main functional goals of postural control and involves the coordination of movement strategies to stabilize the center of body mass during self-initiated and externally triggered stability perturbations (6). Balance disorders account for a series of gait-related disabilities, including problems with transferring, maintaining body posture, and locomotion (7, 8). Therefore, balance is an important component of gait to stabilize one's body during mobility. Meanwhile, growing evidence suggests that gait training can improve balance outcomes (9, 10).

Gait training refers to specific types of physical therapies that help individuals strengthen and improve their walking capacity (11). Treadmill, body-weight-supported treadmill, robot-assisted gait training, virtual reality gait training, conventional gait training, and overground walking training are common types of gait training that have the potential to improve balance capacity (details in Table 1). Several studies have suggested that the aforementioned gait training could counteract the balance dysfunction caused by various diseases, such as stroke and Parkinson's disease (9, 10). However, the effects of gait training on balance rehabilitation after stroke have been inconclusive. It remains unclear which type of gait training is the most effective. Canadian Guideline revealed that gait training (e.g., bodyweight-supported treadmill) might improve dynamic balance in the subacute phase after stroke (12). A recent meta-analysis suggested that overground walking training and robotic-assisted gait training showed no significant effect on balance outcomes while treadmill showed a significant effect on balance outcomes (13). Another systematic review and meta-analysis reported that no significant balance gains were obtained from gait training (e.g., body-weight-supported treadmill and robot-assisted gait training) (14).

According to Shumway-Cook and Woollacott (15), balance performance can be divided into four types, including dynamic steady-state balance, static steady-state balance, proactive balance, and reactive balance. In addition, there are only small-sized correlations between different types of balance performance (16). With reference to these findings, balance outcome measures are further subdivided into five types, including static steady-state balance, dynamic

steady-state balance, proactive balance, reactive balance, and balance test batteries (17). The first four types of balance outcome measures correspond, one by one, to the four types of balance performance, and the fifth type of balance outcome measure (balance test batteries) is added to assess the overall balance performance (18, 19). This classification has been used in several types of research to assess balance status and changes in response to exercise (20, 21). Thus, investigating the effect of gait training in specific type of balance outcomes might provide more comprehensive evidence in this field.

As mentioned earlier, evaluating the effects of gait training in post-stroke patients is of particular importance. However, selecting the optimal gait training to improve specific balance outcomes poses a challenge to clinicians. Because many gait training methods have not been directly compared in clinical trials, typical pairwise meta-analysis cannot be performed on them. Even when direct comparisons are available, the evidence is inadequate to make any conclusions. Therefore, we performed an NMA to compare the effects of different types of gait training on each type of balance outcome, thus identifying the optimal gait training for stroke survivors.

Materials and methods

Search strategy and selection criteria

The NMA was followed by the PRISMA statement (22), and the review protocol has been registered on PROSPERO (CRD42022349965). We searched PubMed, Embase, Medline, Web of Science, and Cochrane Library databases from inception until 25 April 2022, and the search strategy is described in Supplementary Table S1.

Inclusion criteria were based on the participants, interventions, comparators, outcomes, and study design (PICOS) approach (23). To be eligible for inclusion, studies had to meet the following criteria: (1) Population: adult stroke survivors who were considered suitable for gait training by the studies' authors. (2) Intervention: treadmill, bodyweight-supported treadmill, virtual reality gait training, robotic-assisted gait training, overground walking training, and conventional gait training. (3) Comparison: usual care, sham intervention, or no exercise intervention. (4) Outcome: balance outcome measurements (static steady-state balance, dynamic steady-state balance, proactive balance, and balance test batteries). In addition, we only assessed four types of balance outcomes due to the scarcity of studies on reactive balance. (5) Study design: randomized controlled trials.

The exclusion criteria were as follows: (1) intervention group including more than two types of gait training; (2) the type of physical

TABLE 1 Characteristics of included gait training.

Туре	Abbreviation	Description
Treadmill	TT	gait training requires participants to walk on motorized treadmills without any other device assisted
Body weight-supported treadmill	BWS-TT	gait training uses a harness to provide partial body weight support in conjunction with a motorized treadmill
Robot-assisted gait training	RA-GT	gait training with robotic-orthosis attached to patients' lower extremities, allowing the guidance force during ambulation
Virtual reality gait training	VR	gait training with VR systems, providing realistic environments in which people can have real-time interaction with objects and events
Conventional gait training	CGT	gait training administered by physiotherapist according to individual walking capacity of the participants
Overground walking training	OWT	gait training requires participants to walk around an outdoor circuit

exercise being unclear; (3) not reporting sufficient data to calculate the effect size; and (4) conference abstract without a fully published article.

Data extraction and processing

Two independent investigators (M-L and Y-X) searched databases for relevant articles and removed duplicate records, with discrepancies adjudicated by a third investigator (TY-L). After removing duplicate records, pairs of independent investigators (CY-Z and XR-Z) screened references and extracted study-level data, with discrepancies adjudicated by a third investigator (TY-L). We also extracted study characteristics (name of the first author, country of origin, year of publication, intervention groups, and the sample size), participant characteristics (mean age, gender trends, and duration of the disease), and physical exercise characteristics (type, duration, frequency, and time).

Outcomes

The main outcome data (expressed as mean and standard deviation) for changes in balance outcomes (static steady-state balance, dynamic steady-state balance, proactive balance, and balance test batteries) were extracted from initiation to the end of treatment. Only one representative outcome variable was included in the analysis when studies reported several variables under one of these outcome categories. In terms of dynamic steady-state balance, the highest priority was given to the usual walking speed. As a proxy for static steady-state balance, a single right leg stance with eyes opened was used. A functional reach test was preferably selected as a proxy for proactive balance. Berg balance scale was used as the most prominent balance test battery (17). If another test was used in a study, we chose to only include in our quantitative analysis those that had the most comparable temporal/spatial structure to the ones mentioned.

Assessment of heterogeneity and inconsistency

We assessed heterogeneity by reporting the I^2 statistic, as described in Section 9.5.2 of the Cochrane Handbook for Systematic Reviews of Interventions. Using this section, we used the following interpretation of the I^2 statistic: 0–40% might not be important; 30–60% may represent moderate heterogeneity; 50–90% may represent substantial

heterogeneity; and 75–100% may represent considerable heterogeneity (24). We used global and local methods to test the inconsistency of the research results. For global inconsistency, we evaluated inconsistency statistically using the design-by-treatment test (25). We assessed local inconsistency by splitting network estimates into the contribution of direct and indirect evidence (node-splitting test) (26). All heterogeneity and inconsistency analyses were performed with the statistical software R version 3.2.2.

Risk-of-bias assessment

Two independent investigators (CY-Z and XR-Z) assessed the risk of bias of the included RCTs by RoB 2 revised Cochrane risk-of-bias tool for randomized trials. The results were incorporated into the Confidence in Network Meta-Analysis (CINeMA) application to evaluate the credibility of each NMA's findings. CINeMA grade confidence was divided into four categories: high, moderate, low, or very low (27).

Sensitivity analysis

We assessed the sensitivity of our findings by repeating each NMA after excluding studies at an overall high risk of bias. However, because more than one-third of the studies' sample sizes were smaller than 30, sensitivity analysis was not performed on sample size (28).

Statistical analysis

Each type of balance outcome was drawn to the network graph. The random-effects frequentist NMA was fitted to assume a common random effect for all comparisons in the network. Balance outcomes for each parameter and each treatment comparison were estimated as standard mean difference (SMD) with 95% confidence intervals (CIs). The control group was used as the reference group in all forest plots. Meanwhile, the league tables were created to display the relative degree of balance outcomes for all comparisons among gait training. P-scores were applied to rank gait training on the basis of balance outcomes. P-scores ranged from 0 to 1, with a higher P-score indicating a greater effect. In addition, we compared the comparison-adjusted funnel plot and Egger's test to assess the risk of publication bias under specific circumstances, with Egger's test suggesting publication bias when p < 0.05. Finally, the network plots and

comparison-adjusted funnel plots were performed with StataSE version 16.0. Other network meta-analyses were performed with the statistical software R version 3.2.2.

Results

Literature search and characteristics of the included studies

After duplicate removal, we identified 1,353 relevant studies through a database searching strategy. After scanning titles and abstracts, we identified 148 articles. After full-text reading, we excluded 87 articles for reasons as follows: irrelevant study design (n=15), inappropriate control group (n=12), inappropriate intervention group (n=15), no outcomes of interest (n=23), incomplete data (n=12), and conference abstract (n=10). Finally, we identified 61 eligible RCTs (29-88) (Figure 1) containing 2,328 stroke patients, 1,157 (49.7%) of whom received gait training and 1,171 (50.3%) a control intervention. Overall, 1,318 (56.6%) of participants were men, and the mean age ranged from 44.2 to 74.8 years. The mean disease duration ranged from 19 days to 10.5 years. Supplementary Table S2 summarizes the characteristics of included RCTs.

Intervention

There were six different types of gait training included in this NMA: (1) treadmill; (2) body-weight-supported treadmill; (3) virtual reality gait training; (4) robotic-assisted gait training; (5) overground walking training; and (6) conventional gait training.

Of 66 eligible studies, treadmill was assessed in 24 studies (29–32, 34, 35, 37, 45, 54, 57, 59, 62, 63, 65, 67–70, 72–75, 77, 78); bodyweight-supported treadmill in 16 studies (32, 36, 38, 49, 59–62, 64, 66, 68, 71, 76, 83–85); conventional gait training in 9 studies (30, 38, 50, 51, 53, 56, 72, 82); overground walking training in 10 studies (36, 37, 60, 65, 67, 69–71, 74, 76); robotic-assisted gait training in 19 studies (31, 43, 46, 49–58, 66, 80–82, 87); and virtual reality gait training (33, 39–48, 73, 75, 77, 79, 86) in 16 studies.

Risk of bias

The RoB2 analysis showed that 18 (29.5%) studies had a low risk of bias, 38 (63.0%) had some concerns, and 5 (8.2%) had a high risk of bias (Figure 2; Supplementary Table S3). For the randomization process, nine (14.8%) studies had some concerns and two (3.3%) had a high risk of bias; for deviations from intended interventions, 37 (60.6%) studies had some concerns and 3 (4.9%) were at high risk of bias; for missing outcome data, four (6.6%) studies had some concerns and three (4.9%) had a high risk of bias; for measurement of the outcome, 61 (100%) studies had a low risk of bias; and for selection of the reported result, three (4.9%) studies had some concerns.

Dynamic steady-state balance

As shown in Figure 3A, the dynamic steady-state balance was available in 49 trials, comparing six types of gait training (517 patients)

with control intervention (540 patients). Only body-weight-supported treadmill and treadmill appeared superior to control intervention significantly. Ranking on the basis of p-score identified virtual reality gait training as the best and conventional gait training as the worst (Figure 4A). No significant difference was found between gait training (Table 2). The I^2 was 37.5%. Design-by-treatment test (p=0.46, Supplementary Table S4A) and node-splitting test (0% comparison p<0.05, Supplementary Table S5A) showed that no significant global and local inconsistency was observed. CINeMA grade confidence was low or very low in 14 (66.7%) comparisons (Supplementary Table S6A). Moreover, there was no publication bias in dynamic steady-state balance NMA (Egger's test: p=0.713, Supplementary Figure S1A).

Static steady-state balance

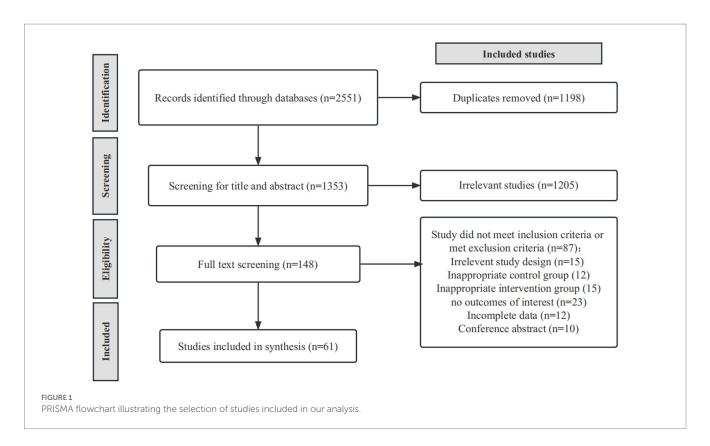
As shown in Figure 3B, the static steady-state balance outcome was available in four trials, comparing two types of gait training (53 patients) with the control intervention (52 patients). Compared with the control intervention, none of the included gait training (body-weight-supported treadmill, virtual reality gait training) showed any significant effect. Ranking on the basis of p-score identified virtual reality gait training as the best and body-weight-supported treadmill as the worst (Figure 4B). No significant difference was found between gait training (Table 3). The I^2 was 0%. Due to the lack of direct comparison, we could not observe inconsistency between direct and indirect comparisons. CINeMA grade confidence was low or very low in three (100%) comparisons (Supplementary Table S5B). Moreover, we could not assess publication bias for the static steady-state balance due to the scarcity of studies.

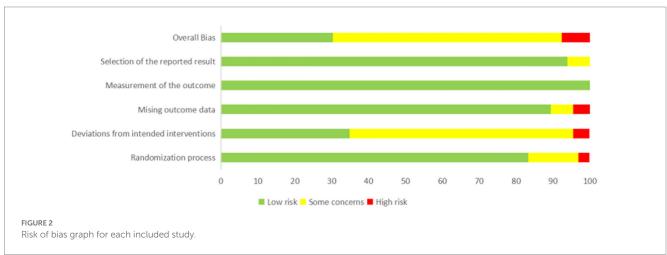
Proactive balance

As shown in Figure 3C, the proactive balance was available in 26 trials, comparing six types of gait training (437 patients) with control intervention (421 patients). Compared with the control intervention, none of the included gait training showed any significant effect. Ranking on the basis of p-score identified body-weight-supported treadmill as the best and overground walking training as the worst (Figure 4C). We did not find a significant difference in proactive balance between gait training according to the league table (Table 4). The I^2 was 0%. Designby-treatment test (p=0.72, Supplementary Table S3C) and node-splitting test (0% comparison p<0.05, Supplementary Table S4C) showed that no significant global and local inconsistency was observed. CINeMA grade confidence was low or very low in 14 (66.7%) comparisons. Moreover, we did not find publication bias in proactive balance NMA (Egger's test: p=0.909, Supplementary Figure S1B).

Balance test batteries

As shown in Figure 3D, the balance test batteries were available in 32 trials, which compared six types of gait training (494 patients) with control intervention (485 patients). Balance test battery analysis showed that virtual reality gait training and body-weight-supported treadmill appeared superior to control intervention significantly. Ranking on the basis of p-score identified virtual reality gait training as the best and conventional gait training as the worst (Figure 4D). Virtual reality gait training, body-weight-supported treadmill, and robotic-assisted gait training appeared

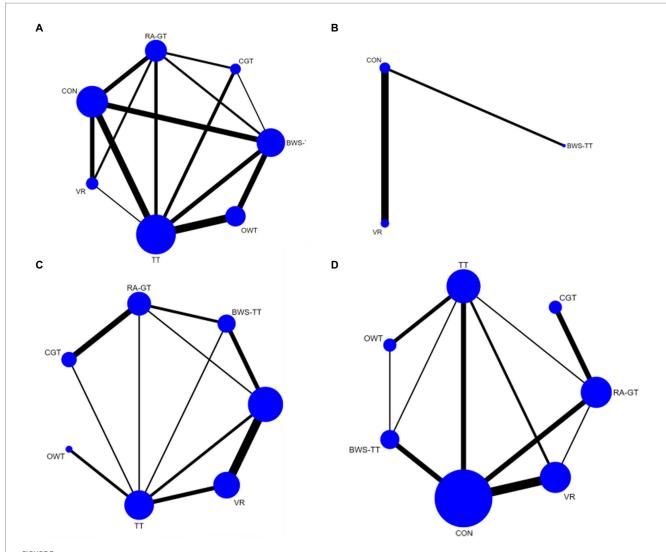




superior to conventional gait training (Table 5). The I^2 was 30%. Design-by-treatment test (p=0.74, Supplementary Table S3D) and node-splitting test (0% comparison p<0.05, Supplementary Table S4D) showed that no significant global and local inconsistency was observed. CINeMA grade confidence was low or very low in 17 (81.0%) comparisons. Moreover, there was no publication bias in balance test batteries NMA (Egger's test: p=0.503, Supplementary Figure S1C).

Sensitivity analysis

The findings essentially remained the same in static steadystate balance and proactive balance sensitivity analyses (Supplementary Figures S2B,C). With regard to dynamic steady-state balance sensitivity analysis (Supplementary Figure S2A), the order of body-weight-supported treadmill and virtual reality gait training was reversed. However, virtual reality gait training did not have a significant effect on dynamic steady-state balance, thus, the removal of these studies did not have a significant effect on our primary results. With regard to balance test battery sensitivity analysis (Supplementary Figure S2D), the body-weight-supported treadmill achieved a higher p-score than virtual reality gait training but did not differ significantly from the control group. Hence, the primary findings from NMA remained unchanged after the sensitivity analysis. Assessments of heterogeneity in all sensitivity analyses were also similar.



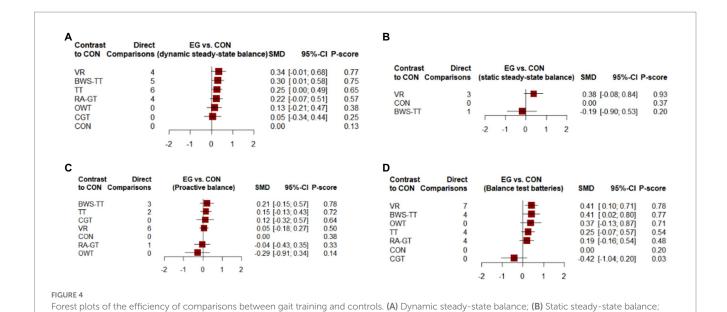
Network plots of the outcomes of the comparisons between gait training and controls in the NMA. (A) Dynamic steady-state balance; (B) Static steady-state balance; (C) Proactive balance; (D) Balance test batteries. TT, treadmill; BWS-TT, body-weight-supported treadmill training; RA-GT, robot-assisted gait training; VR, virtual-reality gait training; CGT, conventional gait training; OWT, overground walking training. Treatments with direct comparisons are linked with a line; the thickness of connecting lines corresponds to the number of trials evaluating the comparison; the size of the circle is proportional to the sample size.

Discussion

Previous studies suggested that gait training showed its potential to improve the balance outcomes of stroke patients (10, 11). However, it remains unclear which type of gait training is more effective in improving specific balance outcomes. Therefore, this study aimed to summarize the related articles to compare the effects of different gait training on different types of balance outcomes in stroke patients. Moreover, our study has some strengths. First, a large number of adult stroke survivors (n = 2,328) were included. Second, to ensure a good level of evidence, we only included RCTs. Third, assessing balance gains in specific types also strengthened this study. Finally, no evidence of inconsistency, substantial heterogeneity, or publication bias was found in any NMA, and sensitivity analyses showed similar results to the overall findings.

To assess optimal gait training, this study performed an NMA to combine data from 61 RCTs including six different types of gait training of 2,328 stroke patients. Our main findings indicated that body-weight-supported treadmill and virtual reality gait training were the most promising gait training to improve dynamic steady-state balance and balance test batteries, respectively.

In terms of dynamic steady-state balance, body-weight-supported treadmill and treadmill showed significant efficacy when compared to control intervention, while the other five types of gait training did not. This result was consistent with the previous studies (89). Gait speed could be improved by repetitive mass motor task practice. When compared to conventional therapy, the body-weight-supported treadmill has the advantage of dynamic balance measured by gait speed because it provides more intensive, repetitive, and task-oriented training during the same time span (90). Moreover, body-weight-supported devices have a positive effect on spatial and temporal gait characteristics by improving body weight distribution between paretic and non-paretic parts (91). The treadmill was the second better gait training to improve dynamic steady-state balance. One possible



control group significantly; P-scores were applied to rank gait training on the basis of balance outcome. P-scores ranged from 0 to 1, a higher P-core indicating a greater effect.

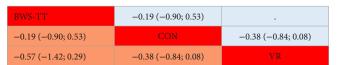
(C) Proactive balance; (D) Balance test batteries. TT, treadmill; BWS-TT, body-weight-supported treadmill training; RA-GT, robot-assisted gait training; VR, virtual-reality gait training; CGT, conventional gait training; OWT, overground walking training. 95% CI≥0 means intervention group is superior to

TABLE 2 League table for dynamic steady-state balance estimate gait training according to their relative effects and 95% credibility intervals (95% CIs).

VR	NA	0.29 (-0.74; 1.32)	0.47 (-0.22; 1.16)	NA	NA	0.18 (-0.22; 0.59)
0.04 (-0.38; 0.46)	BWS-TT	0.34 (-0.12; 0.81)	0.30 (-0.34; 0.94)	-0.06 (-0.42; 0.31)	0.09 (-0.64; 0.81)	0.30 (-0.13; 0.73)
0.09 (-0.30; 0.48)	0.05 (-0.21; 0.31)	TT	-0.29 (-0.89; 0.32)	0.31 (-0.03; 0.65)	0.28 (-0.19; 0.75)	0.29 (-0.03; 0.61)
0.12 (-0.28; 0.51)	0.08 (-0.25; 0.40)	0.03 (-0.27; 0.33)	RA-GT	NA	0.14 (-0.47; 0.75)	0.31 (-0.11; 0.73)
0.20 (-0.25; 0.66)	0.16 (-0.12; 0.45)	0.12 (-0.16; 0.39)	0.09 (-0.29; 0.47)	OWT	NA	NA
0.29 (-0.20; 0.78)	0.25 (-0.14; 0.63)	0.20 (-0.15; 0.55)	0.17 (-0.22; 0.56)	0.08 (-0.34; 0.51)	CGT	NA
0.34 (-0.01; 0.68)	0.30 (0.01; 0.58)	0.25 (0.00; 0.49)	0.22 (-0.07; 0.51)	0.13 (-0.21; 0.47)	0.05 (-0.34; 0.44)	CON

Bold values mean significant difference. Blue means relative degree of direct comparison between gait training. Light red means relative degree of combined comparison between gait training. Deep red means gait training. TT, treadmill; BWS-TT, body-weight-supported treadmill training; RA-GT, robot-assisted gait training; VR, virtual reality gait training; CGT, conventional gait training; OWT, overground walking training; CON, control group.

TABLE 3 League table for static steady-state balance estimate gait training according to their relative effects and 95% credibility intervals (95% CIs).



Blue means relative degree of direct comparison between gait training. Light red means relative degree of combined comparison between gait training. Deep red means gait training. BWS-TT, body-weight-supported treadmill training: VR. virtual reality gait training: CON, control group.

explanation was that treadmill and body-weight-supported treadmill shared the same training process. Both of them involved repetitive mass motor task practice. In addition, patients' characteristics, particularly stroke stage, should be taken into consideration when interpreting the effect of gait training on balance outcomes. Considering body-weight-supported treadmill is not a common intervention in the chronic stroke stage (92, 93), treadmill might be an alternative choice.

In terms of static steady-state balance, our findings showed that no gait training was superior to the control intervention significantly. This finding is consistent with previous studies assessing the effect of gait training on stroke patients (40, 94). For example, a randomized controlled trial suggested that gait training with various weight shifts might have no significant effect on static postural control (95). In other words, balance capacity improved by dynamic tasks might have no effect on static balance. Meanwhile, it has been common knowledge that gait training mainly requires dynamic balance. Therefore, unsurprisingly, no type of gait training showed any effect on static steady-state balance.

Another interesting finding of our study was that gait training did not show a significant effect on proactive balance. Thus, this finding did not support our hypothesis that gait training was effective to improve proactive balance. A previous study reported that stroke patients' capacity to recover effective mobility was only moderately associated with their capacity to improve proactive balance (96). This fact suggests that gait performance does not necessarily reflect poststroke patients' capacity to maintain proactive balance. However, the other studies reported significant improvement in the proactive balance of stroke patients after enhanced gait training (combining gait training with other task-oriented balance training) (66, 97). Therefore, the question of whether or not gait training has a positive effect on proactive balance is still open. The discrepancies between our results

NA 0.33 (-0.13; 0.79) -0.16 (-1.04; 0.72) NA 0.16(-0.36; 0.68)NA 0.08(-0.39; 0.56)NA NA 0.21 (-0.17; 0.59) -0.15 (-0.73; 0.42) NA 0.12 (-0.32; 0.57) NA 0.34 (-0.39; 1.06) -0.18 (-0.63; 0.27) 0.21(-0.15; 0.57)-0.04(-0.29; 0.22)0.49(-0.19; 1.18)0.41 (-0.29; 1.11) 0.29(-0.34; 0.91)NA -0.44 (-1.00; 0.12) NA 0.24 (-0.15; 0.64) 0.16 (-0.18; 0.49) 0.04 (-0.35; 0.43) -0.25 (-0.93; 0.43) 0.35 (-0.57; 1.27) NA 0.06 (-0.34; 0.45) -0.19 (-0.58; 0.20) 0.08(-0.27; 0.43)-0.03 (-0.45; 0.39) -0.15(-0.43; 0.13)-0.44(-1.00; 0.12)0.16 (-0.25; 0.56) 0.07 (-0.39; 0.54) -0.05 (-0.27; 0.18) -0.33 (-0.96; 0.29) -0.08 (-0.50; 0.34) 0.10 (-0.17; 0.38)

TABLE 4 League table for proactive balance estimate gait training according to their relative effects and 95% credibility intervals (95% CIs).

Blue means relative degree of direct comparison between gait training. Light red means relative degree of combined comparison between gait training. Deep red means gait training. TT, treadmill; BWS-TT, body-weight-supported treadmill training; RA-GT, robot-assisted gait training; VR, virtual reality gait training; CGT, conventional gait training; OWT, overground walking training; CON, control group.

TABLE 5 League table for balance test batteries estimates gait training according to their relative effects and 95% credibility intervals (95% CIs).

VR	NA	NA	0.22 (-0.41; 0.86)	-0.05 (-0.92; 0.81)	0.43 (0.07; 0.79)	NA
-0.01 (-0.49; 0.47)	BWS-TT	-0.01 (-0.71; 0.70)	-0.09 (-0.97; 0.79)	NA	0.51 (0.04; 0.98)	NA
0.03 (-0.52; 0.59)	0.04 (-0.46; 0.54)	OWT	0.10 (-0.41; 0.61)	NA	NA	NA
0.16 (-0.23; 0.54)	0.16 (-0.26; 0.59)	0.12 (-0.32; 0.56)	ТТ	-0.59 (-1.66; 0.48)	0.31 (-0.11; 0.72)	NA
0.22 (-0.21; 0.64)	0.22 (-0.29; 0.74)	0.18 (-0.41; 0.77)	0.06 (-0.38; 0.50)	RA-GT	0.05 (-0.35; 0.44)	0.61 (0.10; 1.13)
0.41 (0.10; 0.71)	0.41 (0.02; 0.80)	0.37 (-0.13; 0.87)	0.25 (-0.07; 0.57)	0.19 (-0.16; 0.54)	CON	NA
0.83 (0.16; 1.50)	0.84 (0.11; 1.56)	0.80 (0.02; 1.58)	0.67 (0.00; 1.35)	0.61 (0.10; 1.13)	0.42 (-0.20; 1.04)	CGT

Bold values mean significant difference. Blue means relative degree of direct comparison between gait training. Light red means relative degree of combined comparison between gait training. Deep red means gait training. TT, treadmill; BWS-TT, body-weight-supported treadmill training; RA-GT, robot-assisted gait training; VR, virtual reality gait training; CGT, conventional gait training; OWT, overground walking training; CON, control group.

and some of those previously documented might be the consequence of the fact that most gait training included in our study is a single task rather than a combined task, resulting in a reduction in efficacy.

Our result indicated that virtual reality gait training was the optimal gait training for balance test batteries. One possible explanation is that VR provides a greater number of weight transfer opportunities in different directions than other types of gait training. Another explanation is that virtual reality training could mimic real barriers in the screen image and, thus, enabled the reproduction of a rhythmic weight shift on the paretic side (98). Our results concerning balance test batteries are similar to published research suggesting that virtual reality gait training could significantly improve the berg balance scale score (99). Moreover, a body-weight-supported treadmill was the second best gait training to improve balance test batteries of stroke patients. An explanation is that the body-weight-supported treadmill provides partial weight support, with which related training can be conducted at higher intensities (100). Meanwhile, highintensity training is particularly effective in the rehabilitation of neurological diseases, including stroke (101).

There are several limitations to our study. First, missing evidence for some types of included gait training was identified in our NMA. For example, treadmill accounted for 26.8% of the intervention data. However, conventional gait training only accounted for 10.1%. Therefore, readers should take these results with caution due to inadequate data for direct comparisons for some types of gait training. Next, we failed to assess reactive balance in this NMA since few included trials reported reactive balance. Reactive balance is an important balance outcome to assess patients' capacity to stabilize one's body during perturbation. Future trials are supposed to consider reactive balance as an essential balance outcome when designing studies.

Conclusion

In summary, this NMA suggested that gait training was effective to improve some types of balance outcomes of stroke patients. Considering body- weight-supported treadmill is not a common intervention in the chronic stroke stage (92, 93), treadmill is recommended for those who want to improve their dynamic steady-state balance. Virtual reality gait training is recommended for those who want to improve their balance test batteries. This study emphasizes the importance of gait training to improve the balance outcomes of stroke patients and guides clinicians to formulate better gait training prescriptions.

Data availability statement

The original contributions presented in the study are included in the article/Supplementary material, further inquiries can be directed to the corresponding author.

Author contributions

TL and YW: conceptualization. TL, KY, and JL: methodology. TL and XZ: formal analysis. TL and RW: investigation. XZ, RW, and CZ: resources. XZ, RW, CZ, and ML: data curation. TL and KY: writing—original draft preparation and editing. CL and CX: writing—reviewing. CZ and ML: visualization. YW: supervision. All authors contributed to the article and approved the submitted version.

Funding

This study was supported by the National Key Research and Development Plan of China (2019YFC1710303).

Conflict of interest

CX is employed by SUSE Software (Beijing) Co., Ltd.

The remaining authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

References

- 1. World Health Organization. The top 10 causes of death. (2020). Available at: http://www.who.int/mediacentre/factsheets/fs310/en/
- 2. Wang W, Jiang B, Sun H, Ru X, Sun D, Wang L, et al. Prevalence, incidence, and mortality of stroke in China: results from a Nationwide population-based survey of 480 687 adults. *Circulation*. (2017) 135:759–71. doi: 10.1161/CIRCULATIONAHA.116.025250
- 3. Moradi V, Mafi H, Shariat A, Cleland JA, Nakhostin Ansari N, Savari S. Neurorehabilitation, the practical method of returning to work after stroke. *Iran J Public Health*. (2021) 50:209–10. doi: 10.18502/ijph.v50i1.5092
- 4. Huang S, Yu X, Lu Y, Qiao J, Wang H, Jiang LM, et al. Body weight support-tai chi footwork for balance of stroke survivors with fear of falling: a pilot randomized controlled trial. *Complement Ther Clin Pract.* (2019) 37:140–7. doi: 10.1016/j.ctcp.2019.101061
- 5. Nascimento LR, Rocha RJS, Boening A, Ferreira GP, Perovano MC. Home-based exercises are as effective as equivalent doses of Centre-based exercises for improving walking speed and balance after stroke: a systematic review. *J Physiother*. (2022) 68:174–81. doi: 10.1016/j.jphys.2022.05.018
- 6. Shumway-Cook A, Woollacott MH. Motor control: Translating research into clinical practice. Lippincott Williams & Wilkins (2006).
- 7. Gandolfi M, Valè N, Dimitrova E, Zanolin ME, Mattiuz N, Battistuzzi E, et al. Robot-assisted stair climbing training on postural control and sensory integration processes in chronic post-stroke patients: a randomized controlled clinical trial. *Front Neurosci.* (2019) 13:1143. doi: 10.3389/fnins.2019.01143
- 8. Dietz V, Colombo G. Influence of body load on the gait pattern in Parkinson's disease. *Mov Disord.* (1998) 13:255–61. doi: 10.1002/mds.870130210
- 9. Lorenzo-García P, Cavero-Redondo I, Torres-Costoso AI, Guzmán-Pavón MJ, Núñez de Arenas-Arroyo S, Álvarez-Bueno C. Body weight support gait training for patients with Parkinson disease: a systematic review and meta-analyses. *Arch Phys Med Rehabil.* (2021) 102:2012–21. doi: 10.1016/j.apmr.2021.02.016
- 10. Moucheboeuf G, Griffier R, Gasq D, Glize B, Bouyer L, Dehail P, et al. Effects of robotic gait training after stroke: a meta-analysis. *Ann Phys Rehabil Med.* (2020) 63:518–34. doi: 10.1016/j.rehab.2020.02.008
- 11. Pournajaf S, Calabrò RS, Naro A, Goffredo M, Aprile I, Tamburella F, et al. Robotic versus conventional Overground gait training in subacute stroke survivors: a multicenter controlled clinical trial. *J Clin Med.* (2023) 12:439. doi: 10.3390/jcm12020439
- 12. Stephan KM, Pérennou D. Mobility after stroke: relearning to walk In: T Platz, editor. Clinical pathways in stroke rehabilitation: evidence-based clinical practice recommendations. Cham (CH): Springer (2021). 123–47.
- 13. Nindorera F, Nduwimana I, Thonnard JL, Kossi O. Effectiveness of walking training on balance, motor functions, activity, participation and quality of life in people with chronic stroke: a systematic review with meta-analysis and meta-regression of recent randomized controlled trials. *Disabil Rehabil*. (2022) 44:3760–71. doi: 10.1080/09638288.2021.1894247
- 14. Hsu CY, Cheng YH, Lai CH, Lin YN. Clinical non-superiority of technology-assisted gait training with body weight support in patients with subacute stroke: a meta-analysis. *Ann Phys Rehabil Med.* (2020) 63:535–42. doi: 10.1016/j.rehab.2019.09.009
- $15. \, Shumway-Cook \, A, \, Woollacott \, MH. \, Motor \, control: \, translating \, research \, into \, clinical \, practice. \, 4th \, ed. \, Philadelphia, \, PA: \, Lippincott \, Williams \, \& \, Wilkins \, (2012).$
- 16. Kiss R, Schedler S, Muehlbauer T. Associations between types of balance performance in healthy individuals across the lifespan: a systematic review and meta-analysis. *Front Physiol.* (2018) 9:9. doi: 10.3389/fphys.2018.01366
- 17. Lesinski M, Hortobágyi T, Muehlbauer T, Gollhofer A, Granacher U. Effects of balance training on balance performance in healthy older adults: a systematic review and meta-analysis. *Sports medicine (Auckland, NZ)*. (2015) 45:1721–38. doi: 10.1007/s40279-015-0375-y

Publisher's note

All claims expressed in this article are solely those of the authors and do not necessarily represent those of their affiliated organizations, or those of the publisher, the editors and the reviewers. Any product that may be evaluated in this article, or claim that may be made by its manufacturer, is not guaranteed or endorsed by the publisher.

Supplementary material

The Supplementary material for this article can be found online at: https://www.frontiersin.org/articles/10.3389/fneur.2023.1093779/full#supplementary-material

- 18. Downs S. The Berg balance scale. J Physiother. (2015) 61:46. doi: 10.1016/j. jphys.2014.10.002
- 19. Berg KO, Wood-Dauphinee SL, Williams JI, Maki B. Measuring balance in the elderly: validation of an instrument. *Can J Public Health = Revue canadienne de sante publique.* (1992) 83:S7–S11.
- 20. Lacroix A, Hortobágyi T, Beurskens R, Granacher U. Effects of supervised vs. unsupervised training programs on balance and muscle strength in older adults: A systematic review and meta-analysis. *Sports Med.* (2017) 47:2341–61. doi: 10.1007/s40279-017-0747-6
- 21. Wang LC, Ye MZ, Xiong J, Wang XQ, Wu JW, Zheng GH. Optimal exercise parameters of tai chi for balance performance in older adults: a meta-analysis. *J Am Geriatr Soc.* (2021) 69:2000–10. doi: 10.1111/jgs.17094
- 22. Page MJ, Moher D, Bossuyt PM, Boutron I, Hoffmann TC, Mulrow CD, et al. PRISMA 2020 explanation and elaboration: updated guidance and exemplars for reporting systematic reviews. *BMJ*. (2021) 372:n160. doi: 10.1136/bmj.n160
- 23. Hutton B, Salanti G, Caldwell DM, Chaimani A, Schmid CH, Cameron C, et al. The PRISMA extension statement for reporting of systematic reviews incorporating network meta-analyses of health care interventions: checklist and explanations. *Ann Intern Med.* (2015) 162:777–84. doi: 10.7326/M14-2385
- 24. Higgins JPT, Thomas J, Chandler J, Cumpston M, Li T, Page MJ, (Eds.) et al. Cochrane Handbook for Systematic Reviews of Interventions version 6.3. Cochrane (2022). Available at: http://www.training.cochrane.org/handbook.
- 25. Higgins JP, Jackson D, Barrett JK, Lu G, Ades AE, White IR. Consistency and inconsistency in network meta-analysis: concepts and models for multi-arm studies. *Res Synth Methods*. (2012) 3:98–110. doi: 10.1002/jrsm.1044
- 26. Dias S, Welton NJ, Caldwell DM, Ades AE. Checking consistency in mixed treatment comparison meta-analysis. *Stat Med.* (2010) 29:932–44 . doi: 10.1002/sim.3767
- 27. Nikolakopoulou A, Higgins JPT, Papakonstantinou T, Chaimani A, Del Giovane C, Egger M, et al. CINeMA: an approach for assessing confidence in the results of a network meta-analysis. *PLoS Med.* (2020) 17:e1003082–19. doi: 10.1371/journal.pmed.1003082
- 28. Zeng C, li H, Yang T, Deng ZH, Yang Y, Zhang Y, et al. Electrical stimulation for pain relief in knee osteoarthritis: systematic review and network meta-analysis. *Osteoarthr Cartil.* (2015) 23:189–202. doi: 10.1016/j.joca.2014.11.014
- 29. Nave AH, Rackoll T, Grittner U, Bläsing H, Gorsler A, Nabavi DG, et al. Physical fitness training in patients with subacute stroke (PHYS-STROKE): multicentre, randomised controlled, endpoint blinded trial. *BMJ*. (2019) 366:l5101. doi: 10.1136/bmj. l5101
- 30. Pohl M, Mehrholz J, Ritschel C, Rückriem S. Speed-dependent treadmill training in ambulatory hemiparetic stroke patients: a randomized controlled trial. *Stroke*. (2002) 33:553–8. doi: 10.1161/hs0202.102365
- 31.Lee HJ, Lee SH, Seo K, Lee M, Chang WH, Choi BO, et al. Training for walking efficiency with a wearable hip-assist robot in patients with stroke: a pilot randomized controlled trial. *Stroke*. (2019) 50:3545–52. doi: 10.1161/STROKEAHA.119.025950
- 32. Graham SA, Roth EJ, Brown DA. Walking and balance outcomes for stroke survivors: a randomized clinical trial comparing body-weight-supported treadmill training with versus without challenging mobility skills. *J Neuroeng Rehabil*. (2018) 15:92. doi: 10.1186/s12984-018-0442-3
- 33. Cannell J, Jovic E, Rathjen A, Lane K, Tyson AM, Callisaya ML, et al. The efficacy of interactive, motion capture-based rehabilitation on functional outcomes in an inpatient stroke population: a randomized controlled trial. *Clin Rehabil.* (2018) 32:191–200. doi: 10.1177/0269215517720790
- 34. Gjellesvik TI, Becker F, Tjønna AE, Indredavik B, Lundgaard E, Solbakken H, et al. Effects of high-intensity interval training after stroke (the HIIT stroke study) on physical

and cognitive function: a multicenter randomized controlled trial. *Arch Phys Med Rehabil.* (2021) 102:1683–91. doi: 10.1016/j.apmr.2021.05.008

- 35. Globas C, Becker C, Cerny J, Lam JM, Lindemann U, Forrester LW, et al. Chronic stroke survivors benefit from high-intensity aerobic treadmill exercise: a randomized control trial. *Neurorehabil Neural Repair*. (2012) 26:85–95. doi: 10.1177/1545968311418675
- 36. DePaul VG, Wishart LR, Richardson J, Thabane L, Ma J, Lee TD. Varied overground walking training versus body-weight-supported treadmill training in adults within 1 year of stroke: a randomized controlled trial. *Neurorehabil Neural Repair*. (2015) 29:329–40. doi: 10.1177/1545968314546135
- 37. Langhammer B, Stanghelle JK. Exercise on a treadmill or walking outdoors? A randomized controlled trial comparing effectiveness of two walking exercise programmes late after stroke. *Clin Rehabil*. (2010) 24:46–54. doi: 10.1177/0269215509343328,
- 38. Høyer E, Jahnsen R, Stanghelle JK, Strand LI. Body weight supported treadmill training versus traditional training in patients dependent on walking assistance after stroke: a randomized controlled trial. *Disabil Rehabil.* (2012) 34:210–9. doi: 10.3109/09638288.2011.593681
- 39. In T, Lee K, Song C. Virtual reality reflection therapy improves balance and gait in patients with chronic stroke: randomized controlled trials. *Med Sci Monit.* (2016) 22:4046–53. doi: 10.12659/msm.898157
- 40. Lloréns R, Gil-Gómez JA, Alcañiz M, Colomer C, Noé E. Improvement in balance using a virtual reality-based stepping exercise: a randomized controlled trial involving individuals with chronic stroke. *Clin Rehabil*. (2015) 29:261–8. doi: 10.1177/0269215514543333
- 41. Park DS, Lee DG, Lee K, Lee G. Effects of virtual reality training using Xbox Kinect on motor function in stroke survivors: a preliminary study. *J Stroke Cerebrovasc Dis.* (2017) 26:2313–9. doi: 10.1016/j.jstrokecerebrovasdis
- 42. McEwen D, Taillon-Hobson A, Bilodeau M, Sveistrup H, Finestone H. Virtual reality exercise improves mobility after stroke: an inpatient randomized controlled trial. Stroke. (2014) 45:1853–5. doi: 10.1161/STROKEAHA.114.005362
- 43. Mirelman A, Bonato P, Deutsch JE. Effects of training with a robot-virtual reality system compared with a robot alone on the gait of individuals after stroke. *Stroke*. (2009) 40:169–74. doi: 10.1161/STROKEAHA.108.516328
- 44. Worthen-Chaudhari L. Effectiveness, usability, and cost-benefit of a virtual reality-based telerehabilitation program for balance recovery after stroke: a randomized controlled trial. *Arch Phys Med Rehabil.* (2015) 96:1544. doi: 10.1016/j.apmr.2015.03.025
- 45. de Rooij IJM, van de Port IGL, Punt M, Abbink-van Moorsel PJM, Kortsmit M, van Eijk RPA, et al. Effect of virtual reality gait training on participation in survivors of subacute stroke: a randomized controlled trial. *Phys Ther*. (2021) 101:pzab051. doi: 10.1093/ptj/pzab051
- 46. Kayabinar B, Alemdaroğlu-Gürbüz İ, Yilmaz Ö. The effects of virtual reality augmented robot-assisted gait training on dual-task performance and functional measures in chronic stroke: a randomized controlled single-blind trial. *Eur J Phys Rehabil Med.* (2021) 57:227–37. doi: 10.23736/S1973-9087.21.06441-8
- 47. Lee MM, Lee KJ, Song CH. Game-based virtual reality canoe paddling training to improve postural balance and upper extremity function: a preliminary randomized controlled study of 30 patients with subacute stroke. *Med Sci Monit.* (2018) 24:2590–8. doi: 10.12659/MSM.906451
- 48. Tramontano M, Bergamini E, Iosa M, Belluscio V, Vannozzi G, Morone G. Vestibular rehabilitation training in patients with subacute stroke: a preliminary randomized controlled trial. *NeuroRehabilitation*. (2018) 43:247–54. doi: 10.3233/NRF-185427
- 49. Kim H, Park G, Shin JH, You JH. Neuroplastic effects of end-effector robotic gait training for hemiparetic stroke: a randomised controlled trial. *Sci Rep.* (2020) 10:12461. doi: 10.1038/s41598-020-69367-3
- 50. Yu D, Yang Z, Lei L, Chaoming N, Ming W. Robot-assisted gait training plan for patients in Poststroke recovery period: a single blind randomized controlled trial. *Biomed Res Int.* (2021) 2021:1–7. doi: 10.1155/2021/5820304
- 51. Belas Dos Santos M, Barros de Oliveira C, Dos Santos A, Garabello Pires C, Dylewski V, Arida RM. A comparative study of conventional physiotherapy versus robot-assisted gait training associated to physiotherapy in individuals with ataxia after stroke. *Behav Neurol.* (2018) 2018:2892065–6. doi: 10.1155/2018/2892065
- 52. Yun N, Joo MC, Kim SC, Kim MS. Robot-assisted gait training effectively improved lateropulsion in subacute stroke patients: a single-blinded randomized controlled trial. *Eur J Phys Rehabil Med.* (2018) 54:827–36. doi: 10.23736/S1973-9087.18.05077-3
- 53. Aprile I, Iacovelli C, Goffredo M, Cruciani A, Galli M, Simbolotti C, et al. Efficacy of end-effector robot-assisted gait training in subacute stroke patients: clinical and gait outcomes from a pilot bi-Centre study. *NeuroRehabilitation*. (2019) 45:201–12. doi: 10.3233/NRE-192778
- 54. Bang DH, Shin WS. Effects of robot-assisted gait training on spatiotemporal gait parameters and balance in patients with chronic stroke: a randomized controlled pilot trial. *NeuroRehabilitation*. (2016) 38:343–9. doi: 10.3233/NRE-161325
- 55. Li DX, Zha FB, Long JJ, Liu F, Cao J, Wang YL. Effect of robot assisted gait training on motor and walking function in patients with subacute stroke: a random controlled study. *J Stroke Cerebrovasc Dis.* (2021) 30:105807. doi: 10.1016/j.jstrokecerebrovasdis

- 56. Tanaka N, Matsushita S, Sonoda Y, Maruta Y, Fujitaka Y, Sato M, et al. Effect of stride management assist gait training for Poststroke hemiplegia: a single center, openlabel, randomized controlled trial. *J Stroke Cerebrovasc Dis.* (2019) 28:477–86. doi: 10.1016/j.istrokecerebrovasdis.2018.10.025
- 57. Ogino T, Kanata Y, Uegaki R, Yamaguchi T, Morisaki K, Nakano S, et al. Effects of gait exercise assist robot (GEAR) on subjects with chronic stroke: a randomized controlled pilot trial. *J Stroke Cerebrovasc Dis.* (2020) 29:104886. doi: 10.1016/j. jstrokecerebrovasdis.2020.104886
- 58. Han EY, Im SH, Kim BR, Seo MJ, Kim MO. Robot-assisted gait training improves brachial-ankle pulse wave velocity and peak aerobic capacity in subacute stroke patients with totally dependent ambulation: randomized controlled trial. *Medicine (Baltimore)*. (2016) 95:e5078. doi: 10.1097/MD.00000000000005078
- 59. Srivastava A, Taly AB, Gupta A, Kumar S, Murali T. Bodyweight-supported treadmill training for retraining gait among chronic stroke survivors: a randomized controlled study. *Ann Phys Rehabil Med.* (2016) 59:235–41. doi: 10.1016/j. rehab.2016.01.014
- 60. Gama GL, Celestino ML, Barela JA, Forrester L, Whitall J, Barela AM. Effects of gait training with body weight support on a treadmill versus Overground in individuals with stroke. *Arch Phys Med Rehabil.* (2017) 98:738–45. doi: 10.1016/j.apmr.2016.11.022
- 61. Yen CL, Wang RY, Liao KK, Huang CC, Yang YR. Gait training induced change in corticomotor excitability in patients with chronic stroke. *Neurorehabil Neural Repair.* (2008) 22:22–30. doi: 10.1177/1545968307301875
- 62. Kang HK, Kim Y, Chung Y, Hwang S. Effects of treadmill training with optic flow on balance and gait in individuals following stroke: randomized controlled trials. *Clin Rehabil.* (2012) 26:246–55. doi: 10.1177/0269215511419383
- 63. Kuys SS, Brauer SG, Ada L. Higher-intensity treadmill walking during rehabilitation after stroke in feasible and not detrimental to walking pattern or quality: a pilot randomized trial. *Clin Rehabil*. (2011) 25:316–26. doi: 10.1177/0269215510382928,
- 64. Palmcrantz S, Wall A, Vreede KS, Lindberg P, Danielsson A, Sunnerhagen KS, et al. Impact of intensive gait training with and without electromechanical assistance in the chronic phase after stroke-a multi-arm randomized controlled trial with a 6 and 12 months follow up. Front Neurosci. (2021) 15:660726. doi: 10.3389/fnins.2021.660726
- 65. Aguiar LT, Nadeau S, Britto RR, Teixeira-Salmela LF, Martins JC, Samora GAR, et al. Effects of aerobic training on physical activity in people with stroke: a randomized controlled trial. *NeuroRehabilitation*. (2020) 46:391–401. doi: 10.3233/NRE-193013
- 66. Manji A, Amimoto K, Matsuda T, Wada Y, Inaba A, Ko S. Effects of transcranial direct current stimulation over the supplementary motor area body weight-supported treadmill gait training in hemiparetic patients after stroke. *Neurosci Lett.* (2018) 662:302–5. doi: 10.1016/j.neulet.2017.10.049
- 67. Lamberti N, Straudi S, Malagoni AM, Argirò M, Felisatti M, Nardini E, et al. Effects of low-intensity endurance and resistance training on mobility in chronic stroke survivors: a pilot randomized controlled study. *Eur J Phys Rehabil Med.* (2017) 53:228–39. doi: 10.23736/S1973-9087.16.04322-7
- 68. Kim KH, Lee KB, Bae YH, Fong SSM, Lee SM. Effects of progressive backward body weight suppoted treadmill training on gait ability in chronic stroke patients: a randomized controlled trial. *Technol Health Care.* (2017) 25:867–76. doi: 10.3233/THC-160720
- 69. Danks KA, Pohlig R, Reisman DS. Combining fast-walking training and a step activity monitoring program to improve daily walking activity after stroke: a preliminary study. *Arch Phys Med Rehabil.* (2016) 97:S185–93. doi: 10.1016/j.apmr.2016.01.039
- 70. Hollands KL, Pelton TA, Wimperis A, Whitham D, Tan W, Jowett S, et al. Feasibility and preliminary efficacy of visual Cue training to improve adaptability of walking after stroke: multi-Centre, single-blind randomised control pilot trial. *PLoS One.* (2015) 10:e0139261. doi: 10.1371/journal.pone.0139261
- 71. Combs-Miller SA, Kalpathi Parameswaran A, Colburn D, Ertel T, Harmeyer A, Tucker L, et al. Body weight-supported treadmill training vs. overground walking training for persons with chronic stroke: a pilot randomized controlled trial. *Clin Rehabil.* (2014) 28:873–84. doi: 10.1177/0269215514520773
- 72. Baer GD, Salisbury LG, Smith MT, Pitman J, Dennis M. Treadmill training to improve mobility for people with sub-acute stroke: a phase II feasibility randomized controlled trial. *Clin Rehabil.* (2018) 32:201–12. doi: 10.1177/0269215517720486
- 73. Cho KH, Lee WH. Virtual walking training program using a real-world video recording for patients with chronic stroke: a pilot study. *Am J Phys Med Rehabil.* (2013) 92:371–84. doi: 10.1097/PHM.0b013e31828cd5d3
- 74. Olawale OA, Jaja SI, Anigbogu CN, Appiah-Kubi KO, Jones-Okai D. Exercise training improves walking function in an African group of stroke survivors: a randomized controlled trial. *Clin Rehabil*. (2011) 25:442–50. doi: 10.1177/0269215510389199
- 75. Yang YR, Tsai MP, Chuang TY, Sung WH, Wang RY. Virtual reality-based training improves community ambulation in individuals with stroke: a randomized controlled trial. *Gait Posture*. (2008) 28:201–6. doi: 10.1016/j.gaitpost.2007.11.007
- 76. Nilsson L, Carlsson J, Danielsson A, Fugl-Meyer A, Hellström K, Kristensen L, et al. Walking training of patients with hemiparesis at an early stage after stroke: a comparison of walking training on a treadmill with body weight support and walking training on the ground. Clin Rehabil. (2001) 15:515–27. doi: 10.1191/026921501680425234
- 77. Miclaus RS, Roman N, Henter R, Caloian S. Lower extremity rehabilitation in patients with post-stroke sequelae through virtual reality associated with Mirror therapy. *Int J Environ Res Public Health.* (2021) 18:2654. doi: 10.3390/ijerph18052654

- 78. Dobkin BH, Nadeau SE, Behrman AL, Wu SS, Rose D, PhDet al. Prediction of responders for outcome measures of locomotor experience applied post stroke trial. *J Rehabil Res Dev.* (2014) 51:39–50. doi: 10.1682/JRRD.2013.04.0080
- 79. Song YB, Chun MH, Kim W, Lee SJ, Yi JH, Park DH. The effect of virtual reality and tetra-ataxiometric posturography programs on stroke patients with impaired standing balance. *Ann Rehabil Med.* (2014) 38:160–6. doi: 10.5535/arm.2014.38.2.160
- 80. Kang CJ, Chun MH, Lee J, Lee JY. Effects of robot (SUBAR)-assisted gait training in patients with chronic stroke: randomized controlled trial. *Medicine (Baltimore)*. (2021) 100:e27974. doi: 10.1097/MD.000000000027974
- 81. Song KJ, Chun MH, Lee J, Lee C. The effect of robot-assisted gait training on cortical activation in stroke patients: a functional near-infrared spectroscopy study. *NeuroRehabilitation*. (2021) 49:65–73. doi: 10.3233/NRE-210034
- 82. Park C, Oh-Park M, Dohle C, Bialek A, Friel K, Edwards D, et al. Effects of innovative hip-knee-ankle interlimb coordinated robot training on ambulation, cardiopulmonary function, depression, and fall confidence in acute hemiplegia. *NeuroRehabilitation*. (2020) 46:577–87. doi: 10.3233/NRE-203086
- 83. Sczesny-Kaiser M, Trost R, Aach M, Schildhauer TA, Schwenkreis P, Tegenthoff M. A randomized and controlled crossover study investigating the improvement of walking and posture functions in chronic stroke patients using HAL exoskeleton-the HALESTRO study (HAL-exoskeleton STROke study). Front Neurosci. (2019) 13:259. doi: 10.3389/fnins.2019.00259
- 84. Mustafaoğlu R, Erhan B, Yeldan İ, Ersöz Hüseyinsinoğlu B, Gündüz B, Razak Özdinçler A. The effects of body weight-supported treadmill training on static and dynamic balance in stroke patients: a pilot, single-blind, randomized trial. *Turk J Phys Med Rehabil.* (2018) 64:344–52. doi: 10.5606/tftrd.2018.2672
- 85. Peurala SH, Tarkka IM, Pitkänen K, Sivenius J. The effectiveness of body weight-supported gait training and floor walking in patients with chronic stroke. *Arch Phys Med Rehabil.* (2005) 86:1557–64. doi: 10.1016/j.apmr.2005.02.005
- 86. Marques-Sule E, Arnal-Gómez A, Buitrago-Jiménez G, Suso-Martí L, Cuenca-Martínez F, Espí-López GV. Effectiveness of Nintendo Wii and physical therapy in functionality, balance, and daily activities in chronic stroke patients. *J Am Med Dir Assoc.* (2021) 22:1073–80. doi: 10.1016/j.jamda.2021.01.076
- 87. Kim J, Kim DY, Chun MH, Kim SW, Jeon HR, Hwang CH, et al. Effects of robot-(morning walk $^{\textcircled{o}}$) assisted gait training for patients after stroke: a randomized controlled trial. *Clin Rehabil.* (2019) 33:516–23. doi: 10.1177/0269215518806563
- 88. Gangopadhyay S, Saha S, Sengupta M, Maity B, Chakrabarti D. Effect of body weight support treadmill training on gait recovery, lower limb function and dynamic balance in patients with chronic stroke: a randomised controlled trial. *J Clin Diagn Res.* (2021) 15:KC09-KC13. doi: 10.7860/JCDR/2021/50063.15535
- 89. Combs SA, Dugan EL, Passmore M, Riesner C, Whipker D, Yingling E, et al. Balance, balance confidence, and health-related quality of life in persons with chronic

- stroke after body weight-supported treadmill training. Arch Phys Med Rehabil. (2010) 91:1914–9. doi: 10.1016/j.apmr.2010.08.025
- 90. Mao YR, Lo WL, Lin Q, Li L, Xiao X, Raghavan P, et al. The effect of body weight support treadmill training on gait recovery, proximal lower limb motor pattern, and balance in patients with subacute stroke. *Biomed Res Int.* (2015) 2015:175719:1–10. doi: 10.1155/2015/175719
- 91. Kosak MC, Reding MJ. Comparison of partial body weight-supported treadmill gait training versus aggressive bracing assisted walking post stroke. *Neurorehabil Neural Repair.* (2000) 14:13–9. doi: 10.1177/154596830001400102
- 92. Middleton A, Merlo-Rains A, Peters DM, Greene JV, Blanck EL, Moran R, et al. Body weight-supported treadmill training is no better than overground training for individuals with chronic stroke: a randomized controlled trial. *Top Stroke Rehabil.* (2014) 21:462–76. doi: 10.1310/tsr2106-462
- 93. Duncan PW, Sullivan KJ, Behrman AL, Azen SP, Wu SS, Nadeau SE, et al. Bodyweight-supported treadmill rehabilitation after stroke. N Engl J Med. (2011) 364:2026–36. doi: $10.1056/\rm NEJMoa1010790$
- 94. Kim A, Darakjian N, Finley JM. Walking in fully immersive virtual environments: an evaluation of potential adverse effects in older adults and individuals with Parkinson's disease. *J Neuroeng Rehabil.* (2017) 14:16. doi: 10.1186/s12984-017-0225-2
- 95. Cho KH, Lee WH. Effect of treadmill training based real-world video recording on balance and gait in chronic stroke patients: a randomized controlled trial. *Gait Posture*. (2014) 39:523–8. doi: 10.1016/j.gaitpost.2013.09.003
- 96. States RA, Pappas E, Salem YCochrane Stroke Group. Overground physical therapy gait training for chronic stroke patients with mobility deficits. *Cochrane Database Syst Rev.* (2009) 2009:CD006075. doi: 10.1002/14651858.CD006075.pub2
- 97. Kim J, Jung S, Song C. The effects of auditory feedback gait training using smart insole on stroke patients. *Brain* sciences.2021;11:1377, doi: 10.3390/brainsci11111377.
- 98. Sarasso E, Gardoni A, Tettamanti A, Agosta F, Filippi M. Virtual reality balance training to improve balance and mobility in Parkinson's disease: a systematic review and meta-analysis. *J Neurol.* (2022) 269:1873–88. doi: 10.1007/s00415-021-10857-3
- 99. Lee HC, Huang CL, Ho SH, Sung WH. The effect of a virtual reality game intervention on balance for patients with stroke: a randomized controlled trial. *Games for health journal.* (2017) 6:303–11. doi: 10.1089/g4h.2016.0109
- 100. Mackay-Lyons M, McDonald A, Matheson J, Eskes G, Klus MA. Dual effects of body-weight supported treadmill training on cardiovascular fitness and walking ability early after stroke: a randomized controlled trial. *Neurorehabil Neural Repair*. (2013) 27:644–53. doi: 10.1177/1545968313484809
- 101. Apte S, Plooij M, Vallery H. Influence of body weight unloading on human gait characteristics: a systematic review. *J Neuroeng Rehabil.* (2018) 15:53. doi: 10.1186/s12984-018-0380-0