

Home-based moderate intensity cycling in community dwelling individuals with stroke: protocol of a randomised controlled study

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Abstract

Objective: To investigate the feasibility and preliminary effects of a 12-week home-based moderate intensity cycling programme in stroke survivors.

Design: Single-blinded, parallel group, Randomised Controlled Trial

Setting: Home-based

Participants: Participants will be survivors of first-ever stroke and randomly assigned to an Active Cycling Group (ACG) or control group.

Intervention: The ACG performed a 12-week home-based cycling programme, three sessions per week. Work intervals were executed at 75% of the maximal heart rate (HR_{max}), recovery intervals at 50% of HR_{max} . Work interval duration increased from three ten-minute intervals to 30 minutes continuous cycling. The control group completed a 12-week neurocognitive exercise programme.

Main outcome measurements: Primary outcome measures are compliance with training, adherence to protocol and safety as measured by adverse events. Secondary outcome measurements included a: simplified modified Rankin Scale questionnaire (stroke severity), Six-Minute Walking Test (walking ability), Ten-meter Walk Test (walking ability), Rivermead Mobility Index (walking ability), graded submaximal cycling test (exercise capacity), Physical Activity for Individuals with Physical Disability questionnaire (physical activity), EQ-5D-5L (quality of life), Stroke Specific Quality of Life questionnaire (quality of life), Montreal Cognitive Assessment (cognition), General Self-Efficacy Scale (self-efficacy), and Exercise Self-Regulation Questionnaire (self-regulation).

Keywords: stroke, aerobic exercise, cycling, moderate intensity, home-based

Introduction

Worldwide there are around 30 million stroke survivors, placing stroke second in causes of complex disability (1-4). Symptoms such as spasticity, apraxia, neglect and agnosia are well-known and extensively investigated (5, 6). Besides these symptoms, there are other consequences that recently receives more attention. Mackay-Lyons and Makrides (7) found that the average aerobic capacity of a person with stroke was only 60% of the age and gender related norm. This decreased aerobic capacity is one of the important risk factors in the development of recurrent stroke (8). Considering that recurrent stroke accounts for 20% of all stroke cases (9, 10).

The decreased aerobic capacity after stroke can be explained by a reduction in physical activity, changes in motivation and mood, fatigue, cognitive impairments, and neurological deficits like motor and sensory problems (11, 12). A decreased aerobic capacity may result in a vicious deconditioning cycle in which a reduced maximum oxygen uptake (VO_{2max}) leads to paucity of movements and less weight bearing activities. This can result in altered muscle properties and reduced central command. Both can contribute to a further decrease in VO_{2max} (11, 12). Stimulating people with stroke to be active can break the downward deconditioning spiral. Higher levels of self-efficacy and self-regulation have also been shown to be associated with increased physical activity in individuals (13-16).

Aerobic exercise has proven to be successful in the prevention of prolonged inactivity, avoidance of recurrent stroke and cardiorespiratory events, and improvement of aerobic capacity (17). A minimum of 20 minutes of aerobic exercise, three times a week during minimal 8 weeks, is recommended to achieve a clinically meaningful exercise effect in people with stroke (18).

Since 2011, low to moderate intensity aerobic exercise emerged in the stroke guidelines of the American Stroke Association (19) and in the AEROBICS guidelines (20) are a recommended treatment modality to improve VO_{2max} in people with stroke. In the years following these recommendations, more and more evidence accumulated, including evidence of the benefits of higher intensity training in a supervised, inpatient setting (21-23). Studies in an unsupervised home-based setting are still scarce. Nevertheless, recent guidelines state that aerobic exercise can be delivered in a hospital, in an outpatient clinic but also in an unsupervised home-based setting (18). Home-based training is the most convenient and accessible for most patients but raises concerns about feasibility including safety (19, 24-27).

The delivery of aerobic exercise in a supervised setting has several disadvantages that warrant a more thorough investigation of unsupervised exercise. One of the drawbacks of supervised training is the requirement for transport to the rehabilitation centre which is a frequently reported barrier for physical exercise (21-23, 28-31). It is also difficult to retain the benefits from training in a supervised setting in the long term (18).

In cardiac and obese patients, home-based aerobic exercises have been extensively investigated and have proven to be safe, well tolerated, and effective (32, 33). Here, the training intensities of home-based, unsupervised exercise programmes often vary between 60-80% of the maximal heart rate (HR_{max}) (1-3). As there is considerable overlap between the physical profile of cardiac and stroke survivors, the question arises whether similar exercise programmes would yield the same benefits in people with stroke.

In the stroke population, home-based physical exercises have shown to be as effective as conventional therapy for improving the activities of daily life (ADL) (34). In existing studies, the absence of supervision during functional exercises does not appear to impact therapy compliance (32). However, research that investigates the effect of unsupervised home-based training on aerobic capacity in people with stroke remains scarce (35, 36).

In one of the limitedly available studies, a 12-week home-based brisk walking programme in people with chronic stroke was researched and showed improved quality of life (37). Mayo, MacKay-Lyons (38), found no significant improvements in the 6-minute walk test after a one-year home-based stationary cycling programme. In 2019, the research group of Krawczyk, Vinther (39) investigated a home-based High Intensity Interval Training (HIIT) programme on a stationary bike in people with stroke. The programme consisted of 15-minute training sessions, five days a week for 12 weeks at an intensity of 77-93% of HR_{max} . Here HIIT proved to be safe and well-received in home-based people with stroke but did not translate into significant improvements in cardiorespiratory fitness and general well-being (39). Overall, existing home-based interventions either were performed at relatively low intensities, had large variations in intensity, used a short training duration and/or were carried out on device not suitable for participants with more severe impairments. Alternative devices such as recumbent bikes can enlarge the population of suitable participants.

This study therefore aims to investigate the feasibility and preliminary effectiveness of an individually tailored, home-based, 12-weeks moderate intensity training programme on a stationary recumbent bike in people with stroke.

Methods

Study design and setting

A single-blind, parallel group, randomised controlled design was used. Participants were randomised at a 1:1 ratio to a 12-week active cycling group (ACG) or a control group via sealed envelopes as illustrated in Figure 1. Group allocation occurred immediately after baseline testing. The intervention starts within one week of baseline testing. Post-intervention assessment is performed immediately after the 12-week programme and after a follow-up period of another 12 weeks. All assessments are carried out at RevArte, a neuro-rehabilitation clinic for outpatients and inpatients in the Antwerp

region (Belgium). All training sessions occurred at the participants’ homes. The reporting of this study conforms to the STROBE Guidelines (40).

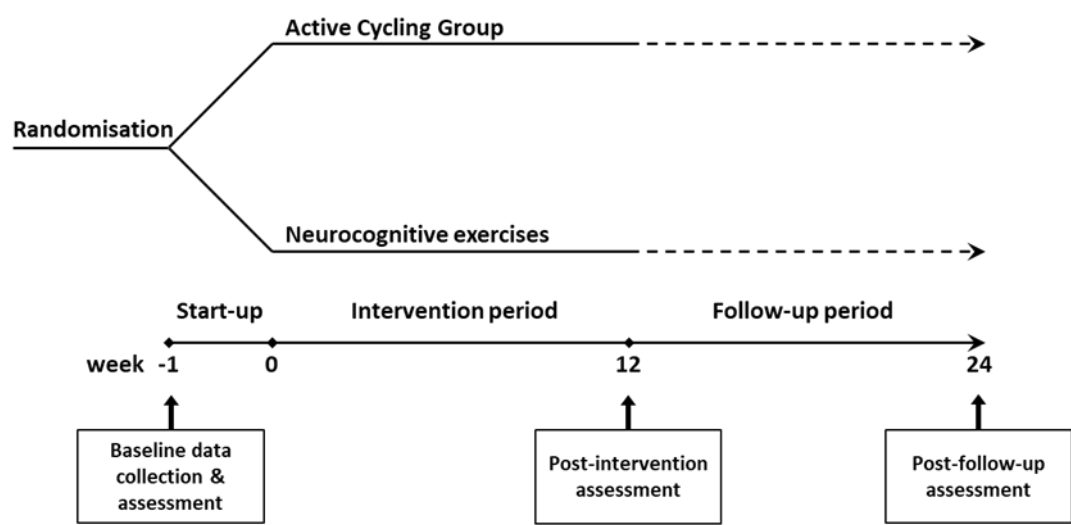


Figure 1 Study design and timeline

Participants

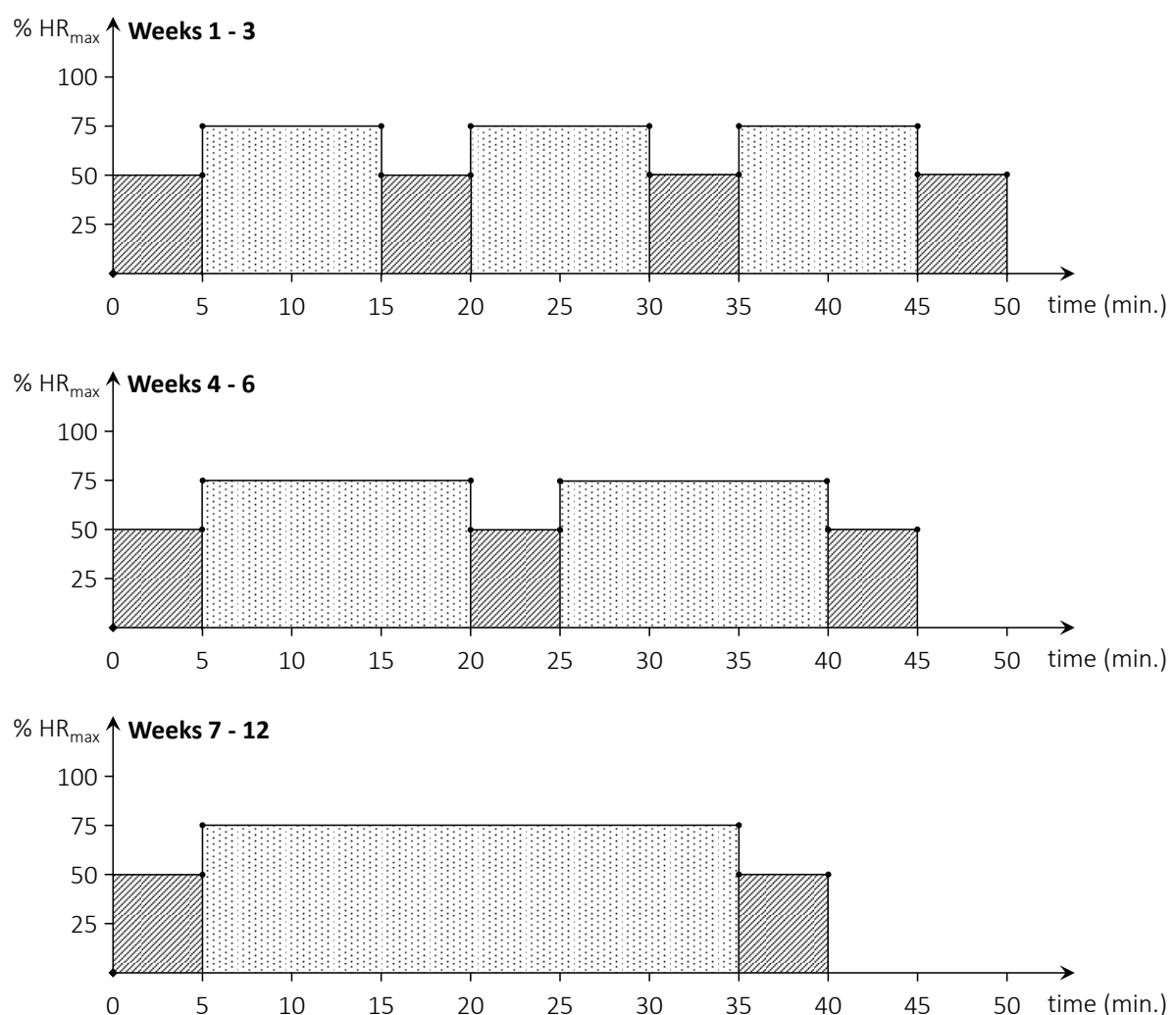
Participants will be recruited from the neuro-rehabilitation clinic RevArte by referral from their physical therapist. Participant inclusion criteria are: 1) first-ever stroke except subarachnoid haemorrhage (41), 2) age <80 years, 3) stroke onset between three months and two years, 4) community dwelling, 5) hemodynamically stable, 6) able to follow simple verbal instructions, and 7) able to pedal a Ergoselect 600 recumbent bike (Ergoline) >1 minute. The criteria for exclusion are: 1) history of substance abuse, psychological illnesses or neurological condition prior to stroke, 2) relative and absolute contra-indications for exercise testing (42), and 3) unable to install a recumbent bike at the participant’s home.

This study is approved by the Medical Ethics Committee of the Antwerp University Hospital (UZA; B300201939038). All subjects volunteer to participate and give prior written informed consent.

Treatment conditions

The ACG carries out a home-based cycling programme and the control group executes a home-based neurocognitive exercise programme. Both programmes are performed 3 times per week for a period of 12 weeks in addition to subjects’ usual care. Participants in the ACG executed an aerobic cycling programme and are provided with a recumbent bike (Tunturi E-80R, Tunturi New Fitness B.V., Almere, The Netherlands) and a compatible heart rate chest strap (5.3Khz) at their homes. Each training consists of 30 minutes of cycling in an interval mode (weeks 1-6) or continuously (weeks 7-12) as illustrated in Figure 2. The training heart rate is set at 75% of the theoretical HR_{max}, corresponding to

the higher end of moderate intensive activity (18). The HR_{max} is calculated as $[220 - \text{age (in years)}]$ (43, 44) or as $[168 - 0.51 \times \text{age (in years)}]$ for participants taking beta-blocker medication (45). The warm-up, cool-down, and active recovery interval heart rates are set at 50% of the HR_{max} . Each session started with a warm-up and ended with a cool-down of five minutes. Training consists of three working intervals of 10 minutes (weeks 1-3), two working intervals of 15 minutes (weeks 4-6), or continuous training of 30 minutes (weeks 7-12). An active recovery period of five minutes is programmed between two interval bouts. Participants are instructed to maintain a cycling cadence of 60 – 70 rpm throughout the training sessions. Participants are requested to keep an exercise diary and to export the data of each training session to a USB storage device connected to the bike. A researcher collects training data every four weeks by offloading the training data and monitor compliance, adherence, and adverse events. After the first week of training a researcher collects training data and verifies for malfunctions or problems.



HR_{max} = maximum heart rate; min. = minutes

Figure 2 Target training heart rate of the active cycling group over a period of 12 weeks

The **control group** carries out a neurocognitive exercise programme focused on executive functions, visual & spatial abilities, and visual attention. The programme was adapted from an existing visuospatial neglect test battery (46). During each training session, participants were requested to complete a workbook that contained several cancellation, bisection, drawing, visuospatial navigation, and other cognitive tasks. It is estimated that an average participant will take about 30 minutes to complete one workbook. The programme consists of 12 unique workbooks. Every 4 weeks, when 12 workbooks were completed, a researcher visited the participant to collect the workbooks and to leave 12 new workbooks for the next 4 weeks. After the first week of training a researcher collects training data and verifies for problems. Compliance and adverse events are also monitored. Participants are requested to keep an exercise diary to track progress.

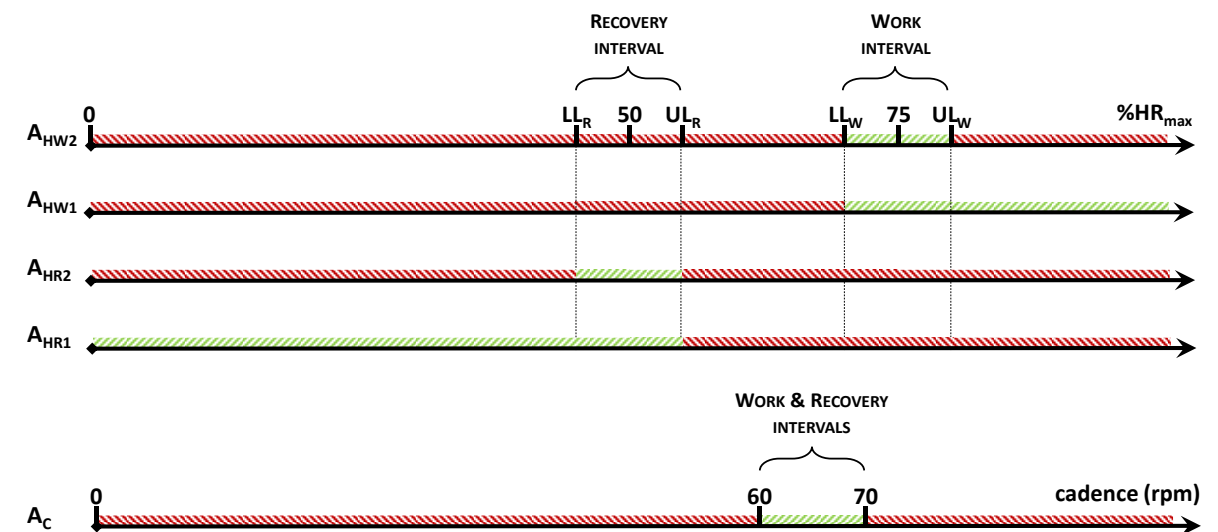
Outcome assessments

All outcome measures are assessed three times (Figure 1): 1) at baseline (approximately one week before the start of the training), 2) at post-intervention (within one week after the 12 weeks training), and 3) at the primary end-point 12 weeks after the end of the training. **Demographic and stroke information**, including medical history are collected (47). The **simplified modified Rankin Scale questionnaire (smRSq)** is used as a tool for measurement of baseline stroke severity (48)

Feasibility

Feasibility is split into three categories: compliance with training, adherence to protocol and safety as measured by the occurrence of adverse events. Each category is assessed continuously throughout the intervention in the ACG and control group. **Compliance with training** is measured by recording the number of training sessions completed in both groups. In the ACG, **adherence to protocol** is recorded by measuring the degree to which participants trained within the requested cycling cadence of 60 – 70 rpm (A_C), and the extent to which they stayed within a 5bpm range of the prescribed heart rate levels (A_H) (Figure 3). Separate adherence figures are calculated for adherence to the work interval heart rate (A_{HW}) and recovery interval heart rate (A_{HR}). For maximum standardisation, both upper (prescribed heart rate + 5bpm) and lower (prescribed heart rate - 5bpm) limits must be respected. In what follows, the latter is referred to as two-sided adherence for work heart rates (A_{HW2}) and recovery heart rates (A_{HR2}). To enable in-depth analysis and because of its clinical value, one-sided adherence is also calculated for heart rate levels by only considering the lower limit (work interval) or upper limit (recovery interval). Thus, one-sided adherence is the degree to which participants cycled *at or higher than* $75\% \times HR_{max} - 5bpm$ for work heart rates (A_{HW1}) and *at or lower than* $50\% \times HR_{max} + 5bpm$ for recovery heart rates (A_{HR1}). In the control group, completeness of workbooks was measured for

adherence. All participants were requested to write down any adverse events in their exercise diary. During each scheduled visit at a participant's home, a researcher also verified if any **adverse events** had occurred.



//// = heart rate or cadence within adherence range; \\\ = heart rate or cadence within non-adherence range; %HR_{max} = percentage of theoretical maximum heart rate; A_C = cadence adherence; A_{HR1} = one-sided recovery heart rate adherence; A_{HR2} = two-sided recovery heart rate adherence; A_{HW1} = one-sided work heart rate adherence; A_{HW2} = two-sided work heart rate adherence; LL_R = lower limit of prescribed heart rate during recovery interval (50% x HR_{max} - 5 bpm); LL_W = lower limit of prescribed heart rate during work interval (75% x HR_{max} - 5 bpm); rpm = revolutions per minute; UL_R = upper limit of prescribed heart rate during recovery interval (50% x HR_{max} + 5 bpm); UL_W = upper limit of prescribed heart rate during work interval (75% x HR_{max} + 5 bpm)

Figure 3 Visualization of adherence cut-off values for an individual participant in the active cycling group

Secondary outcome measures

The **6-Minute Walk Test (6MWT)** assesses walking endurance and was standardized according to the American Thoracic Society guidelines for the 6MWT (49). In addition to distance walked, heart rate throughout the test and step count are measured. Heart rate is measured using a Polar H7 Bluetooth chest strap (Polar Electro, Kempele, Finland). The test is filmed using a mobile phone camera. Step count is measured by post-test video analysis by two researchers. Although stroke-related impairments can strongly influence performance in the 6MWT (49), it is still a reliable (ICC 0.71 -0.99) (50, 51) and valid ($\rho = 0.99$; $p < 0.01$) (51) measure for walking ability post-stroke.

The **10-meter Walk Test (10mWT)** is performed by monitoring a participant walking 10m on a course of 14m, three times at self-selected (SSP) and three times at maximum (MP) pace (adapted from Cheng, Nelson (52)). Lines are marked at (A) the start (2m) and (B) the end (12m) of the measurement distance. The test execution is filmed in the sagittal plane using a mobile phone camera. The camera is held at a right angle to A until the participant crosses one foot over the line and then immediately held perpendicular to B until the participant again crosses a first foot. The time between these

moments is calculated by post-test video analysis. The averages at SSP and at MP are used to calculate the respective walking speeds. The 10mWT is a reliable (53, 54) and valid (52) test to measure gait speed in people with stroke. Mentiplay, Adair (55) found that gait speed has poor correlations with knee extensor ($r = 0.18 - 0.55$) and ankle plantar flexor strength ($r = 0.29 - 0.58$). However, moderate correlations exist for the strength of the ankle dorsiflexors of the paretic limb ($r = 0.50 - 0.73$).

The **Rivermead Mobility Index (RMI)** is used to measure mobility disability (56). The questionnaire contains 14 closed questions that address fundamental motor skills of increasing difficulty and is administered by a researcher. The Dutch version of the RMI has excellent intra-test reliability ($\rho = 0.97$) and is strongly correlated with the Barthel Index ($\rho = 0.84$) (57).

A **graded submaximal cycling test** measures exercise capacity as registered by power output in Watts (W). The test is performed on a stationary recumbent bike (Ergoline Ergoselect 600, Ergoline, Germany). The start workload of 20W is increased with 15W every minute until 1) 75% of HR_{max} is reached, 2) a cadence between 55 - 70 rpm can no longer be maintained or 3) the participant wants to stop, whichever occurs first. Followed by five minutes of cooling down at 20W. Heart rate is measured via a Polar H7 Bluetooth chest strap. Blood pressure is measured every minute at the non-paretic side via an arm blood pressure monitor (Panasonic Diagnostec® EW-BU15, Panasonic, Japan). Physical activity is measured via the **Dutch version of the Physical Activity Scale for Individuals with Physical Disabilities (PASIPD)**, a seven-day recall questionnaire for physical activity based on the Physical Activity Scale for the Elderly (58, 59). The Dutch PASIPD has a test-retest reliability of 0.77 and criterion validity of 0.30, which is in line with other physical activity questionnaires (59, 60).

Health related quality of life (HRQoL) is measured via two self-reported questionnaires. The **EuroQoL 5 Dimensions 5 Levels (EQ-5D-5L)** comprises five dimensions that are rated on a five-level scale (61). The EQ-5D-5L is a recommended core measure to include in every stroke recovery trial (47).

However, as generic HRQoL scales might underestimate the impact of stroke (62, 63), the **Stroke Specific Quality of Life Scale (SS-QoL)** is additionally included. It consists of 49 items, divided into 12 domains which are scored on a five-point Likert scale (63). Psychometric properties for the instrument have been documented by Lin, Fu (64) who established minimal detectable changes (MDC) in individuals with chronic stroke for the mobility (MDC = 1.5 points), self-care (MDC = 1.2 points) and upper extremity (MDC = 1.2 points) domains.

Cognitive health is measured via the **Montreal Cognitive Assessment (MoCA)** (65) which has been shown to be the most valid and clinically feasible screening instrument for cognitive impairments post stroke. Scores can vary from 0 to 30, with 26 or higher generally accepted as normal. In people with stroke, a cut-off score of 23-24 has been established for any degree of cognitive impairment within any domain with a sensitivity of 59-92% and a specificity of 67-85% (66).

Perceived self-efficacy is measured via the **Dutch General Self-Efficacy Scale (GSES)** (15, 67).

The **Exercise Self-Regulation Questionnaire (SRQ-E)** (16) is used to measure individual differences in the types of motivation or regulation for exercise. Scores on the individual items of the questionnaire are combined to form a Relative Autonomy Index (RAI). Low scores on the RAI suggest a more controlled regulatory style, whereas higher scores are indicative for more autonomous regulation. Psychometric properties of the GSES and SRQ-E have been established in multiple populations, however not in stroke. (16)

Sample size calculation

The sample size was calculated using PS Power and Sample Size (68). Oxygen uptake (VO_2) was chosen as an outcome variable, given several tests in the test battery are estimations for VO_2 . Based on data from Gjellesvik, Becker (25) an estimated mean (standard deviation) VO_2 of 2.27 (0.45) L·min⁻¹ at inclusion was selected as baseline value. According to the study, after 13 weeks of cardiorespiratory training in stroke patients an improvement of 16.25% can be expected. Based upon these assumptions an estimation of 14 participants was made to reach a power of 80% with a Type I error probability associated with the null hypothesis of 0.05. Taking a drop-out of 10% into account each group must contain at least 21 participants.

Statistical methods

The statistical program SPSS, version 27, is used for descriptive and non-parametrical statistical analysis (69). The Student t-test is used for comparing the homogeneity between the ACG and control group. Patient characteristics are described. As far as **feasibility** is concerned, linear mixed-effects models are used for analysing the compliance with training and adherence to protocol within a group. A graphical representation of the data is made. Safety is measured by the number and nature of adverse events that occurred. A descriptive comparison of the differences between pre-post values of **secondary outcome measures** for the ACG and control group is made. Results are compared to the clinical important minimal difference (CIMD). When no stroke specific normative values were available, a 10% measurement error (ME) is used based on comparison. Differences between changes in the ACG (pre – post-intervention assessment) and control group are also described using the MCID or ME. To determine differences between both groups over the three timepoints for continuous outcomes linear mixed-effects models are used with factors time and group, time-by-group interaction and subject-specific intercept. Time effects and group specific effects of training are compared between groups by means of post hoc testing with Bonferroni-Holm correction. These tests are conducted for all outcome measures. Statistical significance is set at $p < 0.05$ (two-tailed).

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