

# Physical activity as an effect modifier of lifestyle factors on small vessel disease burden

Andreas Gammelgaard Damsbo      Rolf Ankerlund Blauenfeldt  
Sigrid Breinholt Vestergaard      Niels Lech Pedersen  
Kim Morgenstjerne Østerskov      Mette Foldager Hindsholm  
Arzu Bilgin-Freiert      Claus Ziegler Simonsen      Rikke Beese Dalby  
Grethe Andersen      Janne Kaergaard Mortensen

**Background and aims** Physical activity (PA) may reduce the development of small vessel disease (SVD). The effect of physical activity and more classical vascular risk factors such as hypertension and diabetes in the development of SVD is debated, however. We aim to investigate the effect modification of physical activity on traditional vascular risk factors and the burden of small vessel disease among acute ischemic stroke patients.

**Methods** We have pooled patients from two clinical trials on acute ischemic stroke treatment. The main outcome is an ordinal scale score of quantified MR biomarkers of small vessel disease (SVD) burden based on visually assessed acute stroke scans (T2\* or SWI and FLAIR sequences). Biomarkers includes microbleeds, old lacunar infarcts, superficial siderosis, white matter hyperintensities and atrophy. Covariates includes age, sex, pre-stroke physical activity, diabetes, hypertension, atrial fibrillation and previous cardiovascular diseases. Pre-stroke PA was assessed with a questionnaire on inclusion within a few days after stroke onset. Data will be analyzed using bivariate and multivariate linear regression analysis.

**Results** We expect to include a total of around 1000 adult patients admitted to the comprehensive stroke centre at Aarhus University Hospital between 2013-2022. Preliminary results will be presented at ESOC 2024.

**Conclusions** Physical activity may be an important factor in modifying the risk of SVD development in stroke patients.

Source: [Article Notebook](#)

## Introduction

The correlation between physical activity, small vessel disease and classical risk factors is very much debated and not fully understood. (Moniruzzaman et al. 2020; Torres et al. 2019; Landman et al. 2021)

In this abstract, we present the preliminary results from our pooled SVD study, also presented at [ESOC 2024](#).

## Methods

This study is a cross-sectional study, based on a pooled dataset from two different randomised, clinical trials on patients with acute stroke.

## Results

Please refer to Figure 1 for an overview of subjects included for analysis.

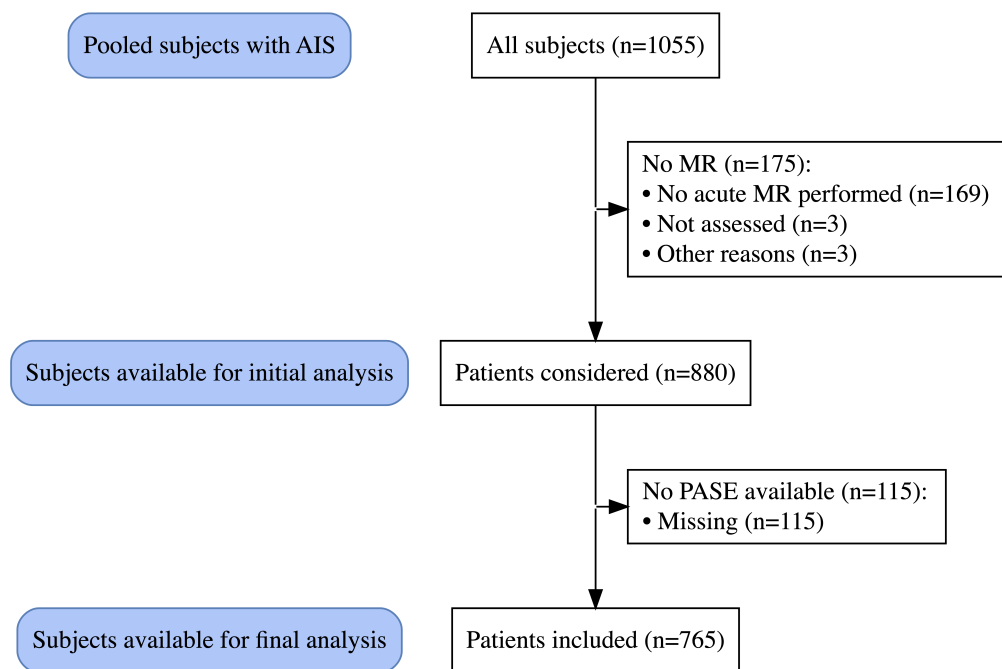


Figure 1: Flowchart of subject included for analysis

Source: [Article Notebook](#)

Baseline characteristics are included with the Table [1](#).

Table 1: Baseline values SVD burden score

Characteristic	Overall, N = 765	Female, N = 280	Male, N = 485
SVD score			
0	302 (39%)	105 (38%)	197 (41%)
1	216 (28%)	75 (27%)	141 (29%)
2	142 (19%)	61 (22%)	81 (17%)
3	71 (9.3%)	29 (10%)	42 (8.7%)
4	34 (4.4%)	10 (3.6%)	24 (4.9%)
Age	71 (62, 79)	75 (64, 80)	70 (61, 77)
Admission NIHSS	4.0 (2.0, 7.0)	4.0 (2.0, 8.0)	3.0 (2.0, 7.0)
Treated with tPA	460 (60%)	159 (57%)	301 (62%)
Treated with EVT	100 (13%)	30 (11%)	70 (14%)
Pre-stroke PASE score	108 (60, 161)	89 (55, 136)	116 (71, 175)
Living alone	203 (27%)	120 (43%)	83 (17%)

Source: [Article Notebook](#)

Scoring reliability between raters has been compared using different metrics, to show different nuances to the performance, see Table 2. The main performance measure is the intraclass correlation coefficient.

Table 2: Inter rater reliability testing

Variable	Agreement	Krippendorffs_Alpha	Fleiss_Kappa	Brennan_Predigers_Kappa	IntraclCorrC
microbleed	0.90	0.65	0.65	0.80	0
lacunes	0.83	0.56	0.56	0.66	0
wmh	0.88	0.72	0.72	0.75	0
atrophy	0.81	0.54	0.54	0.63	0
score	0.62	0.46	0.46	0.52	0

Source: [Article Notebook](#)

Below is the initial evaluation of possible PA effect modification on classical risk factors, Table 3. These results indicates no effect modification as odds ratios are largely unchanged, when PA is introduced in the model (on the right). This may not be the optimal method for this kind of evaluation, though.

Source: [Article Notebook](#)

Based on the preliminary SVD-scores, SVD score distribution stratified by PA quartile is presented in Figure 2.

Table 3: Multivariate, ordinal, logistic regression analysis without and with PASE score included

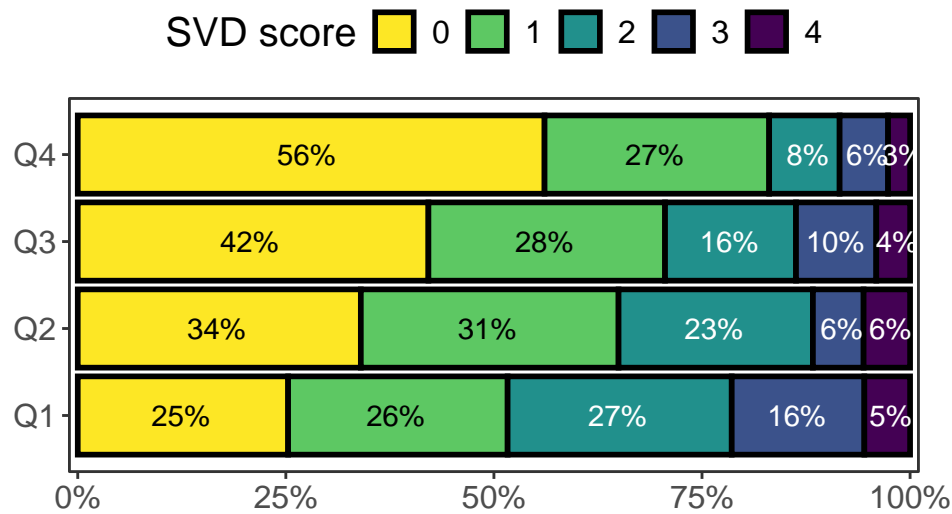
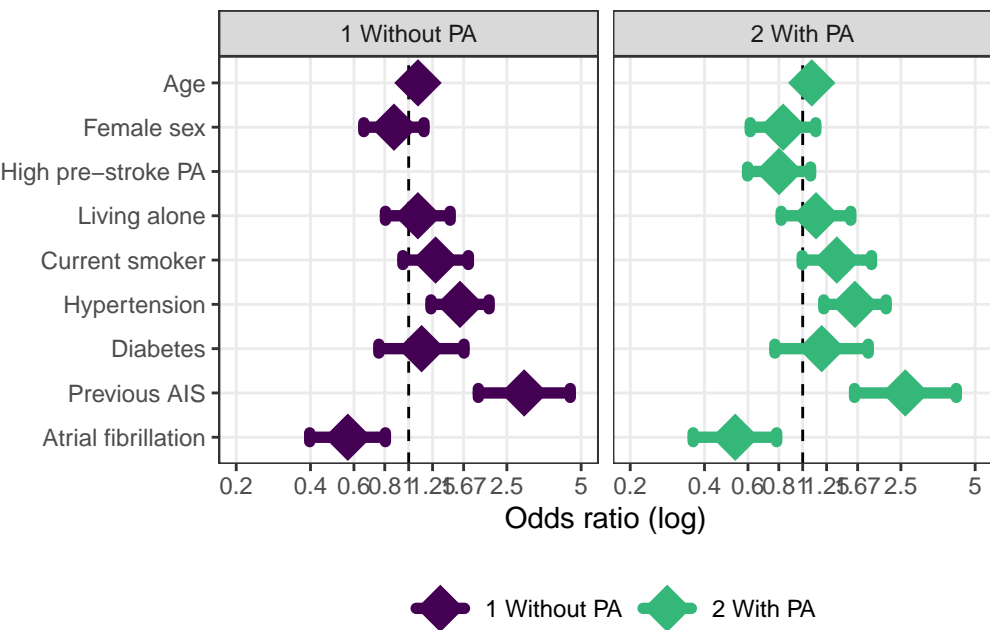


Figure 2

## Discussion

The numbers and figures presented here are very much preliminary and should only be used for discussion and inspiration. Also, if you have any interest in collaboration, please reach out!

- Landman, Thijs Rj, Dick Hj Thijssen, Anil M. Tuladhar, and Frank-Erik de Leeuw. 2021. "Relation between physical activity and cerebral small vessel disease: A nine-year prospective cohort study." *International Journal of Stroke: Official Journal of the International Stroke Society* 16 (8): 962–71. <https://doi.org/10.1177/1747493020984090>.
- Moniruzzaman, Mohammad, Aya Kadota, Hiroyoshi Segawa, Keiko Kondo, Sayuki Torii, Naoko Miyagawa, Akira Fujiyoshi, et al. 2020. "Relationship Between Step Counts and Cerebral Small Vessel Disease in Japanese Men." *Stroke* 51 (12): 3584–91. <https://doi.org/10.1161/STROKEAHA.120.030141>.
- Torres, Elisa R., Siobhan M. Hoscheidt, Barbara B. Bendlin, Vincent A. Magnotta, Gabriel D. Lancaster, Roger L. Brown, and Sergio Paradiso. 2019. "Lifetime Physical Activity and White Matter Hyperintensities in Cognitively-Intact Adults." *Nursing Research* 68 (3): 210–17. <https://doi.org/10.1097/NNR.0000000000000341>.