Epidemiology and Psychiatric Sciences

cambridge.org/eps

Original Article

Cite this article: Cheval B, Csajbók Z, Formánek T, Sieber S, Boisgontier MP, Cullati S, Cermakova P (2021). Association between physical-activity trajectories and cognitive decline in adults 50 years of age or older. Epidemiology and Psychiatric Sciences 30, e79, 1–10. https://doi.org/10.1017/S2045796021000688

Received: 14 June 2021 Accepted: 17 November 2021

Key words:

Aging; cognition; longitudinal studies; physical activity

Author for correspondence:

Boris Cheval, E-mail: boris.cheval@unige.ch
*Corresponding author for statistical analyses:
Pavla Cermakova, E-mail: Pavla.Cermakova@
Ifmotol.cuni.cz

© The Author(s), 2021. Published by Cambridge University Press. This is an Open Access article, distributed under the terms of the Creative Commons Attribution licence (http://creativecommons.org/licenses/by/4.0/), which permits unrestricted re-use, distribution and reproduction, provided the original article is properly cited.



Association between physical-activity trajectories and cognitive decline in adults 50 years of age or older

Boris Cheval^{1,2} , Zsófia Csajbók^{3,4,5}, Tomáš Formánek^{3,6} , Stefan Sieber^{7,8}, Matthieu P. Boisgontier^{9,10}, Stéphane Cullati^{11,12} and Pavla Cermakova^{3,4*}

¹Swiss Center for Affective Sciences, University of Geneva, Geneva, Switzerland; ²Laboratory for the Study of Emotion Elicitation and Expression (E3Lab), Department of Psychology, University of Geneva, Geneva, Switzerland; ³National Institute of Mental Health, Klecany, Czech Republic; ⁴Second Faculty of Medicine, Charles University, Prague, Czech Republic; ⁶EpiCentre, Department of Psychiatry, University of Cambridge, Cambridge, UK; ⁷LIVES Centre, Swiss Centre of Expertise in Life Course Research, Carouge, Switzerland; ⁸Center for the Interdisciplinary Study of Gerontology and Vulnerability, University of Geneva, Geneva, Switzerland; ⁹School of Rehabilitation Sciences, Faculty of Health Sciences, University of Ottawa, Ottawa, Canada; ¹⁰Bruyère Research Institute, Ottawa, Canada; ¹¹Population Health Laboratory, University of Fribourg, Fribourg, Switzerland and ¹²Department of Readaptation and Geriatrics, University of Geneva, Geneva, Switzerland

Abstract

Aims. To investigate the associations of physical-activity trajectories with the level of cognitive performance (CP) and its decline in adults 50 years of age or older.

Methods. We studied 38 729 individuals $(63 \pm 9 \text{ years}; 57\% \text{ women})$ enrolled in the Survey of Health, Ageing and Retirement in Europe (SHARE). Physical activity was self-reported and CP was assessed based on immediate recall, verbal fluency and delayed recall. Physical-activity trajectories were estimated using growth mixture modelling and linear mixed-effects models were used to investigate the associations between the trajectories and CP.

Results. The models identified two trajectories of physical activity: constantly high physical activity ($N = 27\,634$: 71%) and decreasing physical activity ($N = 11\,095$; 29%). Results showed that participants in the decreasing physical-activity group exhibited a lower level of CP compared to the high physical-activity group (immediate recall: $\beta = 0.94$; 95% confidence interval [CI] = 0.92–0.95; verbal fluency: $\beta = 0.98$; 95% CI = 0.97–0.98; delayed recall: $\beta = 0.95$; 95% CI = 0.94–0.97). Moreover, compared with participants in the constantly high physical-activity group, participants in the decreasing physical-activity group showed a steeper decline in all cognitive measures (immediate recall: $\beta = -0.04$; 95% CI = -0.05 to -0.04; verbal fluency: $\beta = -0.22$; 95% CI = -0.24 to -0.21; delayed recall: $\beta = -0.04$; 95% CI = -0.05 to -0.04).

Conclusions. Physical-activity trajectories are associated with the level and evolution of CP in adults over 50 years. Specifically, our findings suggest that a decline in physical activity over multiple years is associated with a lower level and a steeper decline in CP.

Introduction

Physical activity (PA) and cognitive performance (CP) are strongly linked and have been shown to decline with ageing (Cheval *et al.*, 2018*b*, 2020*c*; Sebastiani *et al.*, 2020). Multiple observational and interventional studies have demonstrated that a higher level of PA is associated with better CP in old age (Colcombe and Kramer, 2003; Angevaren *et al.*, 2008; Hamer *et al.*, 2018; Cheval *et al.*, 2020*b*) However, most of these studies focused on the association between PA levels and CP. Yet, disregarding the intra-individual evolution of PA over time may have biased the observed association with CP (Mok *et al.*, 2019). Because PA is a complex behaviour that evolves over time (Mok *et al.*, 2019), examining the associations between life-course PA trajectories and CP is warranted.

Recent studies have attempted to address this gap by considering the evolution of PA over time (Wannamethee *et al.*, 1998; Gregg *et al.*, 2003; Stessman *et al.*, 2009; Almeida *et al.*, 2014; Jefferis *et al.*, 2014; Elhakeem *et al.*, 2018; Laddu *et al.*, 2018; Saint-Maurice *et al.*, 2019; Aggio *et al.*, 2020; Sanchez-Sanchez *et al.*, 2020). However, most of those studies were based on conventional methods to identify groups with different PA trajectories, which typically use clinical or empirical cut-points (Wannamethee *et al.*, 1998; Gregg *et al.*, 2003; Stessman *et al.*, 2009; Almeida *et al.*, 2014; Jefferis *et al.*, 2014; Elhakeem *et al.*, 2018). To reduce the reliance on these cut-points, data-driven approaches such as growth mixture modelling have been proposed (Nagin and Odgers, 2010). These models do not require a priori trajectory classifications as the trajectories emerge from the data (Aggio *et al.*, 2020; Formánek *et al.*, 2020). Using this

approach (Barnett *et al.*, 2008; Laddu *et al.*, 2017), studies have shown PA trajectories in older adults were associated with multiple health outcomes including disability, cardiovascular diseases and all-cause mortality (Laddu *et al.*, 2018, 2020; Saint-Maurice *et al.*, 2019; Aggio *et al.*, 2020; Sanchez-Sanchez *et al.*, 2020). However, the associations of PA trajectories with CP and is unknown.

In this study, we aimed to investigate the associations of PA trajectories with CP level and its decline in adults 50 years of age or older. We hypothesised that unfavourable profiles (e.g., individuals with decreasing PA over time) would be associated with lower levels and steeper decline of CP compared to more favourable profiles (e.g., individuals maintaining high PA levels).

Methods

Design and population

Data were drawn from the Survey of Health, Ageing and Retirement in Europe (SHARE), an ongoing population-based study of health, social network and economic conditions of community-dwelling middle-aged and older individuals, living in 28 European countries and Israel (N = 139556). SHARE was described in detail elsewhere (Börsch-Supan et al., 2013). Briefly, participants were sampled based on probability selection methods. Individuals eligible for the study were people aged 50 years or older and their partners, irrespective of age. Data were collected using computer-assisted personal interviewing (CAPI) in participants' homes. The study was initiated in 2004 and followed by six subsequent waves with approximately 2-year intervals and wave 7 being completed in 2017. Wave 3 was devoted to data collection related to childhood histories (SHARELIFE). This wave did not contain any data related to PA and CP and was therefore not used in the current study. This study was carried out in accordance with the Declaration of Helsinki. SHARE has been approved by the Ethics Committee of the University of Mannheim (waves 1-4) and the Ethics Council of the Max Plank Society (waves 4-7). All participants provided written informed consent. Data were pseudo-anonymised and participants were informed about the storage and use of the data and their right to withdraw consent.

In the present study, we included a total of 38 729 participants who fulfilled the following criteria: (1) age = 50 years or older at baseline, (2) measures of PA in at least three waves, (3) measures of cognition in at least two waves and (4) no report of diagnosed dementia (Fig. 1).

Measures

Physical activity

PA was assessed as part of CAPI in waves 1, 2, 4, 5, 6 and 7 using two questions. 'How often do you engage in vigorous physical activity, such as sports, heavy housework, or a job that involves physical labour?' and 'How often do you engage in activities that require a low or moderate level of energy such as gardening, cleaning the car, or doing a walk?' Participants answered on a fourpoint scale (0 = Hardly ever, or never; 1 = One to three times a month; 2 = Once a week; 3 = More than once a week). An overall score of PA ranging from 0 to 6 was created by summing up the scores on the two questions, with higher scores reflecting greater PA.

Cognitive tests

CP was assessed in waves 1, 2, 4, 5, 6 and 7 using validated tests of verbal fluency, immediate recall and delayed recall. In the verbal fluency test (Rosen, 1980), participants were instructed to name as many different animals as they could think of in 1 minute. The score was the total number of correctly named animals, with a higher score indicating higher verbal fluency. Immediate and delayed recall were assessed using an adapted 10-word delayed-recall test (Harris and Dowson, 1982). In the immediate-recall test, participants first listened to a 10-word list that was read out loud by the interviewer. Then, immediately after the reading of this list, participants were asked to recall as many words as possible. At the end of the cognitive testing session, the participants were asked again to recall any of the words from the list, which captured the delayed recall score. Both scores ranged from 0 to 10, with a higher score indicating greater performance.

Covariates

Models were adjusted for sociodemographic and health-related characteristics identified as potentially confounding and mediating factors in the association between PA and CP level and decline (Trost et al., 2002; Kirk and Rhodes, 2011; Bauman et al., 2012; Choi et al., 2019; Cheval et al., 2021). The value of the covariates was the one at baseline if all PA and CP data were available. In case of missing data at baseline, the value of the covariate was taken from the closest available wave. The selected sociodemographic factors were area of Europe (Western Europe, Scandinavia, Southern Europe, Central and Eastern Europe), age (years), sex (male, female), education (seven categories based on the International Standard Classification of Education 1997), (United Nations Educational, 2006) residence (big city, suburbs or outskirts of a big city, large town, small town, rural area or village), household size (number of members), a partner in a household (yes, no), household net worth (standardised difference between household gross financial assets and financial liabilities), current job situation (working, not working), number of children, number of grandchildren and attrition (non-participation in all the waves or death during the follow-up; yes, no). Health-related characteristics were the number of limitations in instrumental activities of daily living (IADL), number of depressive symptoms assessed with the EURO-D scale (Prince et al., 1999), number of chronic diseases, body mass index (continuous variable), mobility limitations index (number of limitations), smoking (ever smoked daily, never smoked daily), alcohol use (more than two glasses of alcohol almost every day, less), frequency of eating fruits and vegetables (every day, 3-6 times a week, twice a week, once a week, less than once a week).

Statistical analyses

PA trajectories

As in our previous study on trajectories of depressive symptoms (Formánek *et al.*, 2020), growth mixture modelling with maximum likelihood estimation was used to identify latent trajectories of PA (Jung and Wickrama, 2008). This approach estimates latent classes following similar trajectories of PA over time with a high probability. Consistent with previous guidelines and literature (Van De Schoot *et al.*, 2017; Formánek *et al.*, 2020), we first freely estimated time slopes in a latent basis growth model that was entered to the growth mixture model (i.e., the

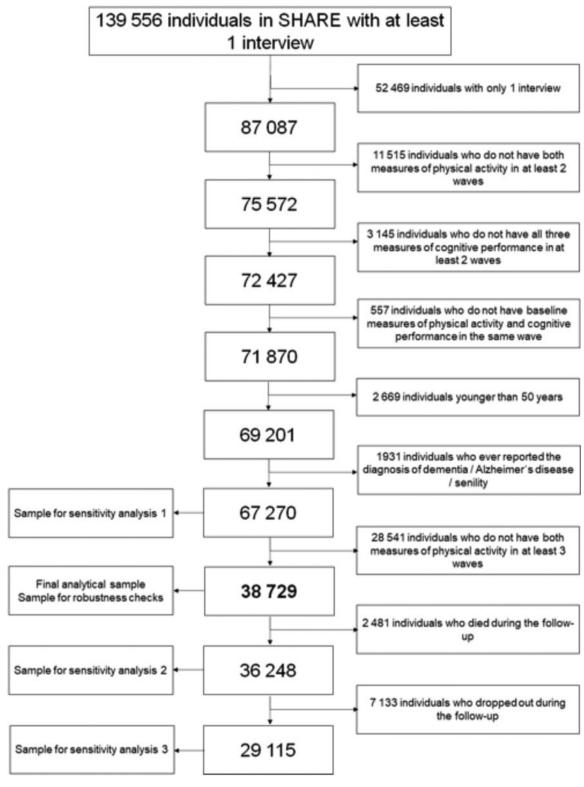


Fig. 1. Selection of the study sample.

classification). Second, the most parsimonious model among those with a different number of PA trajectories (i.e., classes) was determined using the following indicators: Akaike information criterion, Bayesian information criterion, sample-size adjusted Bayesian information criterion, Vuong-Lo-Mendell-

Rubin likelihood ratio test, Luo–Mendell–Rubin adjusted likelihood ratio test and bootstrap likelihood ratio test (see Supplementary Methods). Finally, we checked the number of individuals within each PA trajectory group to ensure an adequate sample size.

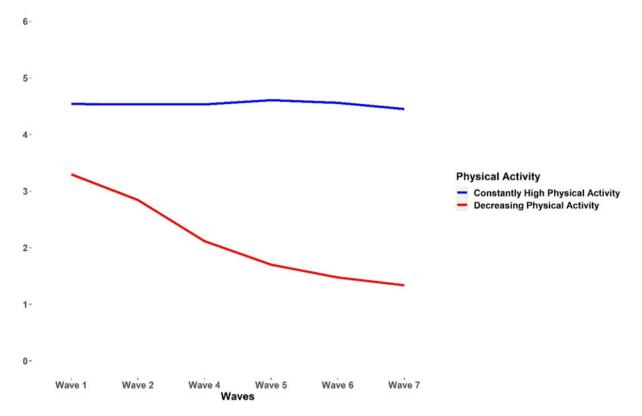


Fig. 2. Physical activity trajectories.

Association between PA trajectories and baseline characteristics First, independent samples t-tests and χ^2 tests were used to compare the differences in baseline characteristics between PA trajectories. Second, a multivariable analysis using logistic regression was performed to estimate the odds ratio (OR) with a 95% confidence interval (CI) for the association between participants' baseline characteristics and PA trajectories.

Association between PA trajectories and the level and rate of decline of CP

We used linear mixed-effects models to investigate the associations of PA trajectories with the level of CP and its rate of decline over time. The models included time in years since baseline, PA trajectory, their interaction term (PA trajectory × time) and covariates. Model 1 was adjusted for age and sex. Other sociodemographic covariates (birth cohort, region, education, residence, household size, partner in household, household net worth, current job situation, number of children, number of grandchildren and attrition) were included in Model 2. Health-related characteristics (limitations in IADL, depressive symptoms, number of chronic diseases, body mass index, mobility limitations index, smoking, alcohol use and eating behaviour) were included in Model 3. We adjusted for covariates group-wise in three steps to assess whether sociodemographic and health-related characteristics may explain the observed associations. The model random structure encompassed random intercepts for participants and random linear slopes for time. In addition, we stratified the dataset according to PA trajectories, and (1) fitted a crude model containing only time and (2) Models 1-3 (except interaction term) for each cognitive test, separately. Based on the fit of the crude stratified model, we visualised the cognitive decline across trajectories of PA and per cognitive tests.

Sensitivity and robustness analyses

We performed three sensitivity analyses (SA), in which we replicated the growth mixture modelling on (1) a sample of participants having at least two measures of PA, two measures of CP and no diagnosis of dementia (SA1: $N=67\ 270$), (2) a sample only including surviving participants (SA2: $N=36\ 248$) and (3) a sample of participants who neither dropped out nor died during the follow-up of the study (SA3: $N=29\ 115$). We conducted two robustness analyses, in which we performed the growth mixture modelling on (a) the moderate physical activity measure only and (b) on the vigorous physical activity measure only.

Data availability

Access to the SHARE data is provided free of charge on the basis of a release policy that gives quick and convenient access to all scientific users worldwide after individual registration. All details about the application and registration process can be found on this website: http://www.share-project.org. The study protocol and syntax of the statistical analysis will be shared upon request from the corresponding author of this study.

Results

PA trajectories

The study sample included 38 729 adults 50 years of age or older (63 years on average, 57% women). Two PA trajectory groups emerged from the growth mixture modelling as the best model for describing the longitudinal data: constantly high PA (N = 27 634: 71%) and decreasing PA (N = 11 095; 29%) (Fig. 2). This

Table 1. Baseline characteristics of the participants across PA trajectories

	Constantly high physical activity (n = 27 634)	Decreasing physical activity ($n = 11095$)	p Value
Cognition			
Immediate recall, mean ± s.p.	5.5 ± 1.6	4.7 ± 1.8	<0.001
Verbal fluency, mean ± s.b.	21.5 ± 7.3	17.9 ± 7.1	<0.001
Delayed recall, mean ± s.b.	4.1 ± 2.0	3.2 ± 2.0	<0.001
Sociodemographic characteristics			
Age, mean ± s.d.	61.2 ± 8.0	66.8 ± 9.6	<0.001
Women, <i>n</i> (%)	15 095 (55)	6819 (62)	<0.001
High education ^a , <i>n</i> (%)	6 601 (24)	1 487 (13)	<0.001
Highest decile of household net worth, n (%)	3 228 (12)	644 (6)	<0.001
Urban residence ^b , <i>n</i> (%)	7 667 (28)	3 061 (28)	0.76
Employed or self-employed, n (%)	11 166 (40)	1 750 (16)	<0.001
Two or more household members, n (%)	22 958 (83)	8 418 (76)	<0.001
Partner in household, n (%)	21 488 (78)	7 693 (69)	<0.001
Two or more children, n (%)	20 721 (75)	8 128 (73)	<0.001
Two or more grandchildren, n (%)	13 154 (48)	6 727 (61)	<0.001
Region, n (%)			
Western Europe	11 618 (42)	4 107 (37)	<0.001
Scandinavia	3 334 (12)	743 (7)	
Southern Europe	4 459 (16)	2 870 (26)	
Central and Eastern Europe	7 204 (26)	2 820 (25)	
Israel	1 019 (4)	555 (5)	
Health-related characteristics			
High depressive symptoms ^c , <i>n</i> (%)	5 909 (21)	3 945 (36)	<0.001
One or more limitations in IADL, n (%)	2 026 (7)	2 682 (24)	<0.001
One or more chronic diseases, n (%)	19 536 (71)	9 542 (86)	<0.001
One or more mobility limitations, n (%)	10 269 (37)	7 326 (66)	<0.001
Smoking, n (%)	18 997 (69)	7 762 (70)	0.02
Alcohol, n (%)	4 960 (18)	1 567 (14)	<0.001
Obesity ^d , n (%)	4 984 (18)	3 202 (29)	<0.001
Poor diet ^e , n (%)	6 032 (22)	2 950 (27)	<0.001

s.d., standard deviation; IADL, instrumental activities of daily living.

two-trajectory solution showed an acceptable fit to the data, although with relatively low entropy (0.603), thereby indicating some overlap between the two classes. Results from the process of model selection, including the results of the sensitivity and robustness analyses, are provided in the supplementary materials.

PA trajectories and baseline characteristics of the participants

Table 1 summarises the characteristics of the participants stratified by PA trajectory. Compared with the constantly high PA group, individuals with decreasing PA showed lower baseline level in the three measures of CP (mean verbal fluency $17.9 \ \nu$.

21.5; mean immediate recall 4.7 ν . 5.5; mean delayed recall 3.2 ν . 4.1; ps < 0.001). In addition, individuals with decreasing PA were older (p < 0.001) and predominantly women (p < 0.001).

Table 2 shows the results of the logistic regression models testing the association between PA trajectories and participants' sociodemographic and health-related characteristics. Compared with individuals with the constantly high PA, decreasing PA was associated with older age, lower education, lower household wealth, non-urban residence, not working, living without a partner, greater depressive symptoms, more limitations in IADL and mobility, more chronic diseases, smoking, obesity and poor diet. When compared to Western Europe, decreasing PA was

^aInternational Standard Classification of Education level 5 or 6.

^bBig city, its suburbs or outskirts.

^c4 or more points on EURO-D scale.

^dBody mass index 30 and more.

^eFruits or vegetables less than every day.

Table 2. Associations of participants' characteristics with decreasing physical activity

OR (95% CI)
1.08 (1.07-1.08)**
1.01 (0.95-1.07)
0.60 (0.56-0.65)**
0.61 (0.55-0.67)**
0.94 (0.89-1.00)*
0.70 (0.65-0.75)**
1.05 (0.93-1.19)
0.82 (0.74-0.92)*
0.95 (0.89–1.01)
1.04 (0.98-1.11)
Reference
0.58 (0.53-0.65)**
2.60 (2.41–2.79)**
0.82 (0.76-0.88)**
1.84 (1.61-2.11)**
1.50 (1.42-1.60)**
2.28 (2.12–2.46)**
1.28 (1.20-1.38)**
1.80 (1.70-1.91)**
1.57 (1.48-1.66)**
0.95 (0.88–1.02)
1.63 (1.53-1.73)**
1.44 (1.35–1.53)**

OR, odds ratio; CI, confidence interval.

All characteristics were entered into the model. In addition, the model included attrition and cognition (mean of z-scores of all three cognitive tests). Because of collinearity, birth cohort was not included into the model.

more likely to occur in Southern Europe and Israel and less likely in Scandinavia and Central and Eastern Europe.

Physical-activity trajectories and cognitive performance

Table 3 shows the results of the linear mixed-effects models. Compared with individuals from the constantly high PA group, individuals in the decreasing PA group had a significantly lower level of baseline CP, independently of all covariates (p < 0.001 for all three measures in Model 1, 2 and 3). Individuals in the decreasing PA group also had a steeper decline of CP in all CP measures, independently from covariates, as shown by the PA trajectory × time interaction (p < 0.001 for all three measures in Models 1, 2 and 3).

When stratified by the PA trajectory, individuals from the constantly high PA group had negligible rates of decline in immediate

recall, verbal fluency and none in delayed recall, while individuals from the decreasing PA group showed steeper rates of decline in all three cognitive measures (Fig. 3). Adjustment for covariates only slightly attenuated the rates of decline. In the fully adjusted models, individuals from the decreasing PA group had a steeper decline in all cognitive measures (immediate recall: $\beta = -0.04$; 95% CI = -0.05 to -0.04; verbal fluency: $\beta = -0.22$; 95% CI = -0.24 to -0.21; delayed recall: $\beta = -0.04$; 95% CI = -0.05 to -0.04), in comparison to the other group. In contrast, individuals from the constantly high PA group only demonstrated a small decline in immediate recall ($\beta = -0.01$; 95% CI -0.01 to -0.01) and verbal fluency ($\beta = -0.05$; 95% CI -0.06 to -0.04) and none in delayed recall ($\beta = 6.0 \times 10^{-4}$; 95% CI -2.2×10^{-3} to 3.4×10^{-3}).

Sensitivity and robustness analyses

The SA and robustness analyses (Supplementary tables S4, S5 and S6) yielded similar results as the main analysis. Specifically, we replicated the 2-class solution (see supplementary methods). Furthermore, the linear mixed-effects models showed that, compared with individuals from the constantly high PA group, individuals from the decreasing PA group had a significantly lower baseline level of the three CP measures and exhibited a steeper decline of those measures. The only difference is that the associations between the PA groups and the decline of immediate and delayed recall over time did not remain significant after the adjustment for all the covariates.

Discussion

In the present large-scale longitudinal multicentric study of adults aged 50 years and older, we identified two distinct PA trajectories: constantly high PA and decreasing PA. These profiles of PA trajectories were both associated with the level of CP and the rate of its decline over time. Compared with the constantly high PA profile, the decreasing PA profile was significantly associated with lower baseline levels of CP, and with a significantly steeper decline of CP over time. These associations were only slightly attenuated after adjustment for the sociodemographic and health-related characteristics. These findings suggest that the CP decline over time is mirrored in the longitudinal changes of PA.

Our results complement the literature on the association of PA and CP during ageing from a longitudinal perspective. Using growth mixture modelling, we identified two distinct PA trajectories over time. One PA trajectory showed PA levels that remained constant across time, a finding that is consistent with previous studies that observed stable PA trajectories in old age (Laddu et al., 2018; Aggio et al., 2020). The second PA trajectory that emerged showed a decreased PA across time. This result is in line with previous studies that have observed a reduction in PA at old age (Lounassalo et al., 2019; Sanchez-Sanchez et al., 2020). However, in addition to these two PA trajectories, other studies found increasing PA trajectories in middle-aged and older adults (Lounassalo et al., 2019; Saint-Maurice et al., 2019). Moreover, studies often reported three or four PA trajectory groups, which contrasts with the two PA trajectories that have emerged from the current data (Lounassalo et al., 2019). This discrepancy may be explained by the features of the scale that was used to measure PA (i.e., 2-items measuring the frequency of moderate and vigorous PA), which was associated with low variance. As such, it was difficult to identify clearly distinct PA trajectories as indicated by

^aInternational Standard Classification of Education level 5 or 6.

^bBig city, its suburbs or outskirts.

^c4 or more points on EURO-D scale.

^dBody mass index 30 and more.

^eFruits or vegetables less than every day.

^{*}p < 0.05; **p < 0.001.

Table 3. Level of cognitive performance and rate of cognitive decline per year across trajectories of physical activity

	Model 1	Model 2	Model 3
	ß (95% CI)		
Association of decreasing	g physical activity with the level of cognitive p	erformance	
Immediate recall	-0.47 (-0.50; -0.44)**	-0.18 (-0.21; -0.15)**	-0.10 (-0.13; -0.06)**
Verbal fluency	-2.58 (-2.73; -2.43)**	-1.03 (-1.17; -0.90)**	-0.71 (-0.85; -0.57)**
Delayed recall	-0.51 (-0.55; -0.47)**	-0.17 (-0.21; -0.14)**	-0.08 (-0.12; -0.04)**
Association of decreasing	g physical activity with the rate of cognitive de	cline (physical activity trajectory×time)	
Immediate recall	-0.03 (-0.04; -0.03)**	-0.03 (-0.04; -0.03)**	-0.03 (-0.04; -0.03)**
Verbal fluency	-0.17 (-0.19; -0.15)**	-0.15 (-0.17; -0.14)**	-0.16 (-0.17; -0.14)**
Delayed recall	-0.04 (-0.05; -0.04)**	-0.04 (-0.05; -0.04)**	-0.04 (-0.05; -0.04)**
Rate of cognitive decline	stratified by the trajectory of physical activity		
Decreasing physical activ	rity		
Immediate recall	-0.05 (-0.05; -0.04)**	-0.04 (-0.05; -0.04)**	-0.04 (-0.05; -0.04)**
Verbal fluency	-0.24 (-0.26; -0.23)**	-0.23 (-0.24; -0.21)**	-0.22 (-0.24; -0.21)**
Delayed recall	-0.05 (-0.05; -0.04)**	-0.04 (-0.05; -0.04)**	-0.04 (-0.05; -0.04)**
Constantly high physical	activity		
Immediate recall	-0.01 (-0.01; -0.01)**	-0.01 (-0.01; -0.01)**	-0.01 (-0.01; -0.01)**
Verbal fluency	-0.07 (-0.08; -0.06)**	-0.06 (-0.06; -0.05) **	-0.05 (-0.06; -0.04)**
Delayed recall	$-2.2 \times 10^{-3} \ (-4.9 \times 10^{-3}; \ 5.4 \times 10^{-4})$	$-2.2 \times 10^{-4} \ (-2.5 \times 10^{-3}; \ 3.0 \times 10^{-3})$	$6.0 \times 10^{-4} \ (-2.2 \times 10^{-3}; \ 3.4 \times 10^{-4})$

 $^{{\}sf CI},$ confidence interval.

Results are derived from linear mixed-effects models.

Model 1: adjusted for age and sex.

Model 2: adjusted for age, sex, birth cohort, region, education, residence, household size, a partner in household, household net worth, current job situation, number of children, number of grandchildren and attrition.

Model 3: adjusted for age, sex, birth cohort, region, education, residence, household size, a partner in household, household net worth, current job situation, number of children, number of grandchildren, attrition, limitations in IADL, depressive symptoms, number of chronic diseases, body mass index, mobility limitations index, smoking, alcohol use and eating behaviour.

*p < 0.05: **p < 0.001.

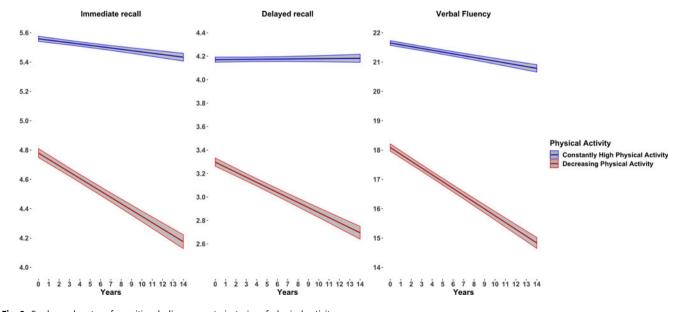


Fig. 3. Crude yearly rates of cognitive decline across trajectories of physical activity.

the low level of entropy observed between our two selected PA groups.

In the present study, 71% of our sample were classified in the more favourable profile (i.e., constantly high PA), while only 29%

were classified in the unfavourable profile (i.e., decreasing PA). This result is rather consistent with previous studies observing that about 30% of older adults showed a decreasing PA trajectory across time (Saint-Maurice *et al.*, 2019; Aggio *et al.*, 2020;

Sanchez-Sanchez et al., 2020). However, while this finding is very encouraging in terms of public health, it must be interpreted with caution given that, in our study, PA was assessed using a self-reported questionnaire, which may have biased the estimation of participants' PA behaviours. Furthermore, when age is used as the time scale instead of the wave of measurement (i.e., using an accelerated longitudinal design assessing PA evolution from 65 to 100 years), results showed a clear decline in PA across ageing in all the PA groups (Laddu et al., 2018, 2020).

We observed that a number of sociodemographic and health-related characteristics are associated with the PA trajectories. In particular, older age, worse socioeconomic status, poor lifestyle profile and worse health were associated with increased odds of belonging to the unfavourable PA group. Our findings are consistent with previous studies that have investigated the multiple correlates of higher engagement in PA (Trost et al., 2002; Bauman et al., 2012). For example, chronic conditions (Cheval et al., 2021), depressive symptoms (Choi et al., 2019), or lower-level education (Kirk and Rhodes, 2011), have been found to be associated with a lower level of engagement in PA. Our results confirm these associations but also reveal that these factors are not only associated with the level of PA engagement, but also with its evolution across time. Even though causality cannot be established due to the observational design of this study, these findings indicate that decline in PA and CP could be predicted. Focusing health care and preventive efforts on subgroups of individuals with low socioeconomic status and poor health profile could contribute to the prevention of decreasing PA and CP.

The strong relationship between higher levels of PA and better cognitive function are well-established (Cheval et al., 2018b, 2020c; Sebastiani et al., 2020), and our results confirm this association. To the best of our knowledge, our study was the first to investigate this association using a data-driven approach to identify PA trajectories. One study has estimated trajectories of PA over 28 years in people with or without dementia and found that PA started to decline 9 years before the diagnosis of dementia (Sabia et al., 2017). A result that rather suggests that changes in PA may simply result from the decline in CP, which is consistent with other studies that have observed an association from changes in CP to changes in PA (Cheval et al., 2020c). Moreover, intervention studies in older adults observed a protective effect of an increased PA on CP (Colcombe and Kramer, 2003; Angevaren et al., 2008), although other studies did not (Sink et al., 2015; Young et al., 2015). However, these studies focused on relatively short-term changes in PA (i.e., from one time of measurement to another one; or before and after an intervention), but disregarded the long-term trajectories of PA across time. Overall, our results are in line with previous studies showing that changes in PA are associated with changes in CP.

Several mechanisms have been suggested to explain the association between a higher level of PA and a maintained level of CP (Cotman and Berchtold, 2002; Colcombe and Kramer, 2003; Cotman et al., 2007; Hillman et al., 2008; Raichlen and Alexander, 2017). For example, PA has been associated with increased brain plasticity, angiogenesis, synaptogenesis and neurogenesis primarily through the release of growth factors such as brain-derived neurotrophic factor, insulin-like growth factor-1 and vascular endothelial growth factor (Cotman and Berchtold, 2002; Cotman et al., 2007; Hillman et al., 2008). Moreover, the cognitive demands inherent in many types of PA, which may include planning, reasoning, decision-making

and multitasking, may also have positive effects on the brain (Raichlen and Alexander, 2017; Singh *et al.*, 2019). Alternatively, additional mechanisms have been suggested to explain the association in the other direction – i.e., from cognitive functioning to higher engagement in physical activity (Cheval *et al.*, 2018a, 2019, 2020a). In particular, anchored within the theory of effort minimisation in physical activity (TEMPA) (Cheval and Boisgontier, 2021), these studies showed that cognitive functions are critical to counteract the automatic tendency to effort minimisation, thereby favouring engagement in PA.

Among the strengths of the present study are the large sample size, the longitudinal design, the data-driven approach to identify PA trajectories across time and the reliance on three measures of CP based on established procedures. However, some limitations should be noted. First, PA was assessed using a self-reported questionnaire, which may have reduced measurement validity. Second, the two PA groups differed in both the initial PA levels (i.e., high v. moderate) and the evolution of PA across time (i.e., maintaining v. decreasing). As such, it was not possible to disentangle the influence of these two factors (i.e., level or slope) on CP. Future research should rely on a more detailed and reliable questionnaire or on device-based measures of PA to be able to better discriminate between PA trajectories, and their associations with CP. Likewise, as the PA groups differed regarding overall health-related condition, we cannot exclude that a steeper cognitive decline observed in the unfavourable PA profile can result from this poorer health condition rather than the change in PA. Furthermore, as it cannot be avoided in long-term longitudinal studies, we cannot exclude a selection bias due to attrition. Yet, to minimise this bias, our statistical analyses were adjusted for attrition and we conducted SA excluding participants who dropped out or died during the follow-up. Finally, the correlational nature of the SHARE design cannot guarantee causal links between PA trajectories and CP decline.

In conclusion, these findings confirm the hypothesis that an unfavourable PA profile may be associated with weaker CP in old age, thereby supporting the need to promote effective strategies to help individuals to maintain their PA levels over the life course.

Supplementary material. The supplementary material for this article can be found at https://doi.org/10.1017/S2045796021000688.

Data. This SHARE dataset is available at http://www.share-project.org/data-access.html.

Acknowledgements. This paper uses data from SHARE Waves 1, 2, 3 (SHARELIFE), 4, 5, 6, 7 and 8 (DOIs: 10..6103/SHARE.w1.600, 10..6103/SHARE.w2.600, 10..6103/SHARE.w3.600, 10..6103/SHARE.w4.600, 10..6103/SHARE.w5.600, 10..6103/SHARE.w6.600, 10..6103/SHARE.w7.711, 10.6103/SHARE.w8cabeta.001). The SHARE data collection was primarily funded by the European Commission through FP5 (QLK6-CT-2001-00360), FP6 (SHARE-I3: RII-CT-2006-062193, COMPARE: CIT5-CT-2005-028857, SHARELIFE: CIT4-CT-2006-028812) and FP7 (SHARE-PREP: no.211909, SHARE-LEAP: no.227822, SHARE M4: no.261982). Additional funding from the German Ministry of Education and Research, the Max Planck Society for the Advancement of Science, the U.S. National Institute on Aging (U01_AG09740-13S2, P01_AG005842, P01_AG08291, P30_AG12815, R21_AG025169, Y1-AG-4553-01, IAG_BSR06-11, OGHA_04-064, HHSN271201300071C) and from various national funding sources is gratefully acknowledged (see www.share-project.org).

Author contributions. B.C.: conceptualisation, writing – original draft; Z.C.: conceptualisation, formal analysis, writing – original draft; T.F.: formal analysis, writing – review & editing; S.S.: writing – review & editing; M.P.B.:

writing – review & editing; S.C.: supervision, writing – review & editing; P.C.: conceptualisation, formal analysis, writing – original draft.

Financial support. B.C. is supported by an Ambizione grant (PZ00P1_180040) from the Swiss National Science Foundation (SNSF). Z.C. is supported by Charles University Research Centre programme UNCE/HUM/025 (204056) and PRIMUS grant (247066). P.C. is supported by grant PRIMUS (247066) conducted at Charles University and the Ministry of Health of the Czech Republic (grant NU20J-04-00022). M.P.B. is supported by the Natural Sciences and Engineering Research Council of Canada (RGPIN-2021-03153) and by a grant from The Banting Research Foundation.

Ethical standards. This study was part of the SHARE study, approved by the relevant research ethics committees in the participating countries.

Informed consent. All participants provided written informed consent.

References

- Aggio D, Papachristou E, Papacosta O, Lennon LT, Ash S, Whincup P, Wannamethee SG and Jefferis BJ (2020) Trajectories of physical activity from midlife to old age and associations with subsequent cardiovascular disease and all-cause mortality. *Journal of Epidemiology and Community Health* 74, 130–136.
- Almeida OP, Khan KM, Hankey GJ, Yeap BB, Golledge J and Flicker L (2014) 150 min of vigorous physical activity per week predicts survival and successful ageing: a population-based 11-year longitudinal study of 12 201 older Australian men. British Journal of Sports Medicine 48, 220–225.
- Angevaren M, Aufdemkampe G, Verhaar H, Aleman A and Vanhees L (2008) Physical activity and enhanced fitness to improve cognitive function in older people without known cognitive impairment. The Cochrane database of systematic reviews 3, CD005381.
- Barnett TA, Gauvin L, Craig CL and Katzmarzyk PT (2008) Distinct trajectories of leisure-time physical activity and predictors of trajectory class membership: a 22 year cohort study. *International Journal of Behavioral Nutrition and Physical Activity* 5, 1–8.
- Bauman AE, Reis RS, Sallis JF, Wells JC, Loos RJ, Martin BW and Group LPASW (2012) Correlates of physical activity: why are some people physically active and others not? *The Lancet* 380, 258–271.
- Börsch-Supan A, Brandt M, Hunkler C, Kneip T, Korbmacher J, Malter F, Schaan B, Stuck S and Zuber S (2013) Data resource profile: the Survey of Health, Ageing and Retirement in Europe (SHARE). *International Journal of Epidemiology* 42, 992–1001.
- **Cheval B and Boisgontier MP** (2021) The theory of effort minimization in physical activity. *Exercise and Sport Sciences Reviews* **49**, 168–178.
- Cheval B, Radel R, Neva JL, Boyd LA, Swinnen SP, Sander D and Boisgontier MP (2018a) Behavioral and neural evidence of the rewarding value of exercise behaviors: a systematic review. Sports Medicine 48, 1389–1404.
- Cheval B, Sieber S, Guessous I, Orsholits D, Courvoisier DC, Kliegel M, Stringhini S, Swinnen S, Burton-Jeangros C, Cullati S and Boisgontier MP (2018b) Effect of early- and adult-life socioeconomic circumstances on physical inactivity. Medicine and Science in Sports and Exercise 50, 476–485.
- Cheval B, Rebar AL, Miller MM, Sieber S, Orsholits D, Baranyi G, Courvoisier DC, Cullati S, Sander D and Boisgontier MP (2019) Cognitive resources moderate the adverse impact of poor neighborhood conditions on physical activity. Preventive Medicine 126, 105741.
- Cheval B, Daou M, Cabral DAR, Bacelar M, Parma JO, Forestier C, Orsholits D, Sander D, Boisgontier MP and Miller MW (2020a) Higher inhibitory control is required to escape the innate attraction to effort minimization. *Psychology of Sport and Exercise* 51, 101781.
- Cheval B, Darrous L, Choi K, Klimentidis Y, Raichlen D, Alexander G, Cullati S, Kutalik Z and Boisgontier MP (2020b) Physical activity and general cognitive functioning: A Mendelian Randomization study. BioRxiv.
- Cheval B, Orsholits D, Sieber S, Courvoisier DC, Cullati S and Boisgontier MP (2020c) Relationship between decline in cognitive resources and physical activity. *Health Psychology* **39**, 519–528.

- Cheval B, Maltagliati S, Sieber S, Beran D, Chalabaev A, Sander D, Cullati S and Boisgontier MP (2021) Why are individuals with diabetes less active? The mediating role of physical, emotional, and cognitive factors. *Annals of Behavioral Medicine* 55, 904–917.
- Choi KW, Chen C-Y, Stein MB, Klimentidis YC, Wang M-J, Koenen KC and Smoller JW (2019) Assessment of bidirectional relationships between physical activity and depression among adults: a 2-sample Mendelian randomization study. JAMA Psychiatry 76, 399–408.
- Colcombe S and Kramer AF (2003) Fitness effects on the cognitive function of older adults: a meta-analytic study. *Psychological Science* **14**, 125–130.
- Cotman CW and Berchtold NC (2002) Exercise: a behavioral intervention to enhance brain health and plasticity. *Trends in Neurosciences* **25**, 295–301.
- Cotman CW, Berchtold NC and Christie L-A (2007) Exercise builds brain health: key roles of growth factor cascades and inflammation. *Trends in Neurosciences* 30, 464–472.
- Elhakeem A, Murray ET, Cooper R, Kuh D, Whincup P and Hardy R (2018) Leisure-time physical activity across adulthood and biomarkers of cardio-vascular disease at age 60–64: a prospective cohort study. *Atherosclerosis* **269**, 279–287.
- Formánek T, Csajbók Z, Wolfová K, Kučera M, Tom S, Aarsland D and Cermakova P (2020) Trajectories of depressive symptoms and associated patterns of cognitive decline. *Scientific Reports* 10, 1–11.
- Gregg EW, Cauley JA, Stone K, Thompson TJ, Bauer DC, Cummings SR and Ensrud KE and Group SoOFR. (2003) Relationship of changes in physical activity and mortality among older women. *JAMA* 289, 2379–2386.
- Hamer M, Terrera GM and Demakakos P (2018) Physical activity and trajectories in cognitive function: English longitudinal study of ageing. *Journal of Epidemiology and Community Health* 72, 477–483.
- Harris S and Dowson J (1982) Recall of a 10-word list in the assessment of dementia in the elderly. *The British Journal of Psychiatry* **141**, 524–527.
- Hillman CH, Erickson KI and Kramer AF (2008) Be smart, exercise your heart: exercise effects on brain and cognition. *Nature Reviews Neuroscience* 9, 58–65.
- Jefferis BJ, Whincup PH, Lennon LT, Papacosta O and Goya Wannamethee S (2014) Physical activity in older men: longitudinal associations with inflammatory and hemostatic biomarkers, N-terminal pro-brain natriuretic peptide, and onset of coronary heart disease and mortality. *Journal of the American Geriatrics Society* **62**, 599–606.
- Jung T and Wickrama KA (2008) An introduction to latent class growth analysis and growth mixture modeling. Social and Personality Psychology Compass 2, 302–317.
- Kirk MA and Rhodes RE (2011) Occupation correlates of adults' participation in leisure-time physical activity: a systematic review. American Journal of Preventive Medicine 40, 476–485.
- Laddu DR, Cawthon PM, Parimi N, Hoffman AR, Orwoll E, Miljkovic I and Stefanick ML (2017) Trajectories of the relationships of physical activity with body composition changes in older men: the MrOS study. BMC Geriatrics 17, 1–10.
- Laddu D, Parimi N, Cauley JA, Cawthon PM, Ensrud KE, Orwoll E, Stefanick M and Langsetmo L and Group OFiMSR (2018) The association between trajectories of physical activity and all-cause and cause-specific mortality. *Journals of Gerontology Series A Biological Sciences and Medical Sciences* 73, 1708–1713.
- Laddu DR, Parimi N, Stone KL, Lapidus J, Hoffman AR, Stefanick ML and Langsetmo L (2020) Physical activity trajectories and associated changes in physical performance in older men: the MrOS study. The Journals of Gerontology – Series A Biological Sciences and Medical Sciences 75, 1967– 1973.
- Lounassalo I, Salin K, Kankaanpää A, Hirvensalo M, Palomäki S, Tolvanen A, Yang X and Tammelin TH (2019) Distinct trajectories of physical activity and related factors during the life course in the general population: a systematic review. BMC Public Health 19, 1–12.
- Mok A, Khaw K-T, Luben R, Wareham N and Brage S (2019) Physical activity trajectories and mortality: population-based cohort study. *BMJ* 365, 13222
- Nagin DS and Odgers CL (2010) Group-based trajectory modeling in clinical research. Annual Review of Clinical Psychology 6, 109–138.

- Prince MJ, Reischies F, Beekman AT, Fuhrer R, Jonker C, Kivela S-L, Lawlor BA, Lobo A, Magnusson H and Fichter M (1999) Development of the EURO-D scale-a European Union initiative to compare symptoms of depression in 14 European centres. *The British Journal of Psychiatry* 174, 330–338.
- Raichlen DA and Alexander GE (2017) Adaptive capacity: an evolutionary neuroscience model linking exercise, cognition, and brain health. *Trends in Neurosciences* 40, 408–421.
- Rosen WG (1980) Verbal fluency in aging and dementia. Journal of Clinical and Experimental Neuropsychology 2, 135–146.
- Sabia S, Dugravot A, Dartigues J-F, Abell J, Elbaz A, Kivimäki M and Singh-Manoux A (2017) Physical activity, cognitive decline, and risk of dementia: 28 year follow-up of Whitehall II cohort study. *British Medical Journal* 357, j2709.
- Saint-Maurice PF, Coughlan D, Kelly SP, Keadle SK, Cook MB, Carlson SA, Fulton JE and Matthews CE (2019) Association of leisure-time physical activity across the adult life course with all-cause and cause-specific mortality. JAMA Network Open 2, e190355.
- Sanchez-Sanchez JL, Izquierdo M, Carnicero-Carreño JA, García-García FJ and Rodríguez-Mañas L (2020) Physical activity trajectories, mortality, hospitalization, and disability in the Toledo study of healthy aging. *Journal of Cachexia, Sarcopenia and Muscle* 11, 1007–1017.
- Sebastiani P, Andersen SL, Sweigart B, Du M, Cosentino S, Thyagarajan B, Christensen K, Schupf N and Perls TT (2020) Patterns of multi-domain cognitive aging in participants of the long life family study. *Geroscience* 42, 1335–1350.
- Singh AS, Saliasi E, Van Den Berg V, Uijtdewilligen L, De Groot RH, Jolles J, Andersen LB, Bailey R, Chang Y-K and Diamond A (2019) Effects of

- physical activity interventions on cognitive and academic performance in children and adolescents: a novel combination of a systematic review and recommendations from an expert panel. *British Journal of Sports Medicine* **53**, 640–647.
- Sink KM, Espeland MA, Castro CM, Church T, Cohen R, Dodson JA, Guralnik J, Hendrie HC, Jennings J and Katula J (2015) Effect of a 24-month physical activity intervention vs health education on cognitive outcomes in sedentary older adults: the LIFE randomized trial. *JAMA* 314, 781–790.
- Stessman J, Hammerman-Rozenberg R, Cohen A, Ein-Mor E and Jacobs JM (2009) Physical activity, function, and longevity among the very old. Archives of Internal Medicine 169, 1476–1483.
- Trost SG, Owen N, Bauman AE, Sallis JF and Brown W (2002) Correlates of adults' participation in physical activity: review and update. *Medicine and Science in Sports and Exercise* 34, 1996–2001.
- United Nations Educational (2006) International Standard Classification of Education 1997. UNESCO, Paris.
- Van De Schoot R, Sijbrandij M, Winter SD, Depaoli S and Vermunt JK (2017) The GRoLTS-checklist: guidelines for reporting on latent trajectory studies. Structural Equation Modeling: A Multidisciplinary Journal 24, 451–467.
- Wannamethee SG, Shaper AG and Walker M (1998) Changes in physical activity, mortality, and incidence of coronary heart disease in older men. *The Lancet* **351**, 1603–1608.
- Young J, Angevaren M, Rusted J and Tabet N (2015) Aerobic exercise to improve cognitive function in older people without known cognitive impairment. The Cochrane Database of Systematic Reviews 4, CD005381