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The effects of exercise programs on cognition, activities of daily living, and neuropsychiatric symptoms in community-dwelling people with dementia—a systematic review

Kathrin Steichele^{1*}, Anne Keefer¹, Nikolas Dietzel¹, Elmar Graessel², Hans-Ulrich Prokosch³ and Peter L. Kolominsky-Rabas¹

Abstract

Background: The prevalence of dementia is expected to increase dramatically. Due to a lack of pharmacological treatment options for people with dementia, non-pharmacological treatments such as exercise programs have been recommended to improve cognition, activities of daily living, and neuropsychiatric symptoms. However, inconsistent results have been reported across different trials, mainly because of the high heterogeneity of exercise modalities. Thus, this systematic review aims to answer the questions whether exercise programs improve cognition, activities of daily living as well as neuropsychiatric symptoms in community-dwelling people with dementia.

Methods: Eight databases were searched for articles published between 2016 and 2021 (ALOIS, CENTRAL, CINAHL, Embase, MEDLINE, PsycINFO, PubMed, Web of Science). Randomized controlled trials evaluating the effects of any type of physical activity on cognition, activities of daily living, or neuropsychiatric symptoms in community-dwelling people with a formal diagnosis of dementia were included in this systematic review. Two authors independently assessed eligibility and quality of the studies. The methodology was in line with the Preferred Reporting Items for Systematic Reviews and Meta-Analysis guidelines.

Results: Eight publications covering seven trials were included in this review with the majority investigating either a combination of strength and aerobic exercise or aerobic exercise alone. This review revealed that there is no clear evidence for the beneficial effects of exercise on cognition. None of the included trials found an impact on activities of daily living. Although different randomized controlled trials reported inconsistent results, one trial indicated that especially aerobic exercise may improve neuropsychiatric symptoms.

Conclusion: Our systematic review did not confirm the impact of exercise on cognition and activities of daily living in community-dwelling people with dementia. The results suggested that aerobic exercise might be effective to reduce neuropsychiatric symptoms. Well-designed trials including only community-dwelling people with a formal

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^{*}Correspondence: Kathrin.steichele@web.de

¹ Friedrich-Alexander-Universität Erlangen-Nürnberg, Interdisciplinary Center for Health Technology Assessment and Public Health (IZPH), Erlangen, Germany

diagnosis of dementia, large samples, long-term follow-ups, and detailed description of adherence to the intervention are needed to improve the scientific evidence on the best type of exercise modality.

Trial registration: PROSPERO, CRD42021246598.

Keywords: Non-pharmacological treatment, Physical activity, Cognitive function, BPSD, ADLs

Introduction

Improvements in health care in the past decades have contributed to an increase in life expectancy. Although dementia is not an inevitable part of normal aging, incidence increases with age. Currently, over 55 million people worldwide live with dementia, and prevalence of dementia is expected to increase dramatically as the population ages [1].

Due to the limited availability of pharmacological treatment, non-pharmacological interventions have been recommended as first-line approaches for over a decade to improve cognition, activities of daily living (ADLs), and neuropsychiatric symptoms (NPS) in people with dementia (pwd) [2, 3]. Among these, exercise has been recommended as an effective treatment for slowing down cognitive decline in pwd [4-7]. Furthermore, recent research partially shows physical activity to be a promising method to reduce NPS and improve ADLs [4, 5, 8]. However, these findings are not consistent, and conflicting results have been reported across different trials [9–11]. Although this is a field of high interest and many trials have been conducted, recent evidence seems controversial, as various systematic reviews reported conflicting results [6, 12-14]. As people with mild cognitive impairment (MCI), pwd living in a long-term care facility, and community-dwelling pwd have different needs and capabilities, it is necessary to distinguish between these groups, which has not been done in previous reviews. Thus, this systematic review aims to give a broad overview of the effects of exercise programs on cognition, ADLs, and NPS in community-dwelling people with a formal diagnosis of dementia. Moreover, frequency, intensity, duration, and setting of the interventions have been hardly discussed in previous reviews. Therefore, we aim to analyze how training modalities (type of training, frequency, duration, intensity, setting) influence the effects of exercise.

Methods

This systematic review was conducted and reported following the guidelines of Preferred Reporting Items for Systematic Reviews and Meta-Analysis (PRISMA) [15] and was registered at the national prospective register of systematic reviews (PROSPERO registration number: CRD42021246598).

Data sources and searches

PubMed, MEDLINE, Embase, ALOIS, Web of Science, PsychINFO, CINAHL, and CENTRAL were systematically searched using different terms for exercise and dementia. Full search strategy can be found in Additional file 2. Whenever possible, searches with additional filters such as randomized controlled trials as article type, English or German as language and publication date from 2016 until 2021, were conducted. Since a Cochrane Review on this topic has been conducted in 2015, we aim to focus on trials published after 2015.

Study selection

Randomized controlled trials (RCTs) over any length of time with the aim of improving cognition, ADLs, or NPS in pwd were eligible for this systematic review. As an intervention, we included RCTs providing any combination of resistance, endurance, or balance training. Multidomain interventions in which isolated effects of exercise cannot be measured (e.g., combination with cognitive training) had to be excluded. For control groups, usual care or social activities were included, while following regular exercise was used as exclusion criteria. Furthermore, we eliminated trials in which people with MCI or subjective cognitive impairment (SCI) or institutionalized pwd were involved. We considered people living in the community or assisted-living facilities at the time of intervention which is why acute hospitalized pwd or people living in long-term care facilities were excluded. Since most of the pwd living at home with community-dwelling people having different capabilities than institutionalized pwd, we excluded institutionalized pwd, where the disease is often more advanced [16, 17]. We had no restrictions regarding the type of dementia, as long as they had a formal medical diagnosis.

After merging search results and discarding duplicates, title, abstracts, and full texts were independently screened for inclusion by the first two authors (K.S. and A.K). In cases of disagreements, the last author was consulted for the final decision (P.K.-R.).

Data extraction and risk of bias assessment

Information from the articles was extracted by the first author, with the second author checking the collected data, which included study setting, publication year, country, way of recruitment, funding, inclusion and exclusion criteria, sample size, and drop-out rates.

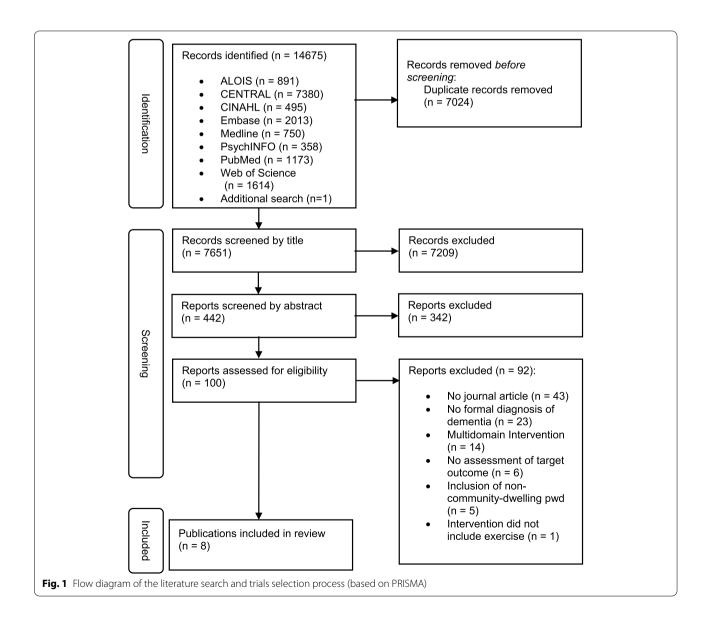
Extracted data also covered participants' baseline characteristics such as gender, age, diagnostic criteria, and diagnosis as well as Mini-Mental Status Examination (MMSE) score at baseline when available. Furthermore, a detailed description of the exercise modalities (e.g., type, frequency, duration, actual and planned intensity) and outcome data of the first follow-up was gathered. If a study combined multiple types of exercise, it was regarded as a multimodal intervention. Between-group differences in the mean changes from baseline to follow-up in domains of cognition, ADLs, and NPS were reported.

The quality of included studies was assessed by the first and the third authors using the "Revised Cochrane Risk-of-Bias Tool for Randomized Trials (RoB2)" [18].

Results

Included studies

In total, 14,675 studies were identified through the search. After removing duplicates, 7651 records were screened of which 7551 were excluded as they covered irrelevant topics (Fig. 1). Of the remaining 100 articles, 92 studies were further excluded mainly because of: (1) other publication formats such as conference papers, (2) participants without a formal diagnosis [8, 19–22], (3) a combination of different interventions [23–28], (4) no



assessment of target outcome [29–31], (5) or institutionalized pwd [32–36].

Consequently, eight publications of seven trials with a total of 1135 participants were included in this review. Detailed study characteristics are summarized in Table 1. The mean age ranged from 70.5 ± 7.4 [37] to 84.3 ± 7.7 years [38] and participants baseline MMSE varied between 14.7 ± 5.65 [38] and 23.94 ± 3.6 [37]. Different types of exercise were conducted, with the majority of included studies investigating either aerobic training [37, 39, 40] or a combination of aerobic, strength, and balance training (multimodal training) [41–43]. With the exception of Park et al. [38], all trials examined the effects of exercise on cognition. Effects on ADLs were investigated in two studies [37, 42] and NPS in four studies [37, 38, 42, 44]. Adherence to the protocol ranged from 65.63% [42] to 87% [26].

Effects of exercise on cognition

Of the seven included trials, six examined the effects of exercise on cognition in pwd, with five trials covering global cognition, measured by the Mini-Mental State Examination (MMSE) [37, 41, 43] or Alzheimer's Disease Assessment Scale (ADAS-Cog) [40, 42]. Although Yu et al. [40] concluded that aerobic exercise is effective to reduce global cognitive decline, none of the studies demonstrated a significant superiority of the intervention group performing either aerobic [37, 39, 40] or multimodal training [41–43] compared to the control group receiving usual care.

Assessment tools for cognitive subdomains varied between trials. Executive function was assessed in four studies [39–41, 43], either using the Clock Drawing test or composite scores of multiple tests, with only one trial finding mild positive effects after 52 weeks of multimodal training [43] (Table 2). None of the trials reported an impact on the domains verbal fluency and language [37, 40, 41, 43] as well as attention and processing speed [37, 40, 41]. While the effects of exercise through either aerobic [39] or multimodal training [37] on psychomotor speed were analyzed in two trials, only Karssemeijer et al. [39] demonstrated that twelve weeks of aerobic training leads to improvements in psychomotor speed.

Effects of exercise on activities of daily living (ADLs)

The effects of moderate-to-high intensity exercise trainings over 16 weeks on ADLs were investigated in two studies [37, 42] (Table 3). Although Lamb et al. [42] found an improvement in physical fitness, these effects could not be translated into improvements in ADLs measured by Alzheimer's Disease Cooperative Study ADL Scale (ADCS-ADL). These findings are in line with Hoffmann

et al., who found no improvement in ADLs assessed with the Bristol Activities of Daily Living Scale (BADLS) [34].

Effects of exercise on neuropsychiatric symptoms (NPS)

Four studies investigated the effects of aerobic training [37], multimodal training [42, 44], and chair-based strengthening or yoga exercises [38] on NPS, which were assessed by Neuropsychiatric Inventory (NPI) in three trials [37, 42, 44], while Park et al. [38] measured agitation, depression, and anxiety by Cohen-Mansfield Agitation Inventory-Short Form (CMAI) and Hospital Anxiety and Depression Scale (HADS), respectively (Table 4). Whereas Hoffmann et al. [37] described significant differences in change in total NPS, indicating less severe NPS in the intervention group after sixteen weeks of training, neither of the other interventions led to improvements in NPS after twelve [38], 16 [42], or 52 [44] weeks.

Risk of bias in included studies

Risk of bias varied between some to high concerns in the included studies (Table 5). For the detailed description, please refer to Additional file 1. Incompleteness of outcome data, selective reporting as well as measurement of the outcome were the predominant reasons for high concerns. Concerns in the measurement of the outcome mainly occurred because outcome assessors were aware of the intervention received. As participants were also aware of the assigned intervention, the possibility of bias due to deviations from the intended intervention did lead to some concerns in all included trials. Reporting of the trial of Karssemeijer et al. [39] and Lamb et al. [42] did not raise further concerns apart from blinding and thus should be considered as the included studies with the lowest risk of bias.

Discussion

Effects on cognition

This systematic review aimed to gather the current state of research on the effects of exercise on cognition, ADLs, and NPS in community-dwelling pwd. We found that pwd receiving exercise interventions did not yield additional benefits on global cognition in any of the included trials. In line with previous reviews, exercise could thus not be described as being effective for slowing down cognitive decline in pwd [12, 45]. Moreover, the results of Lamb et al. (2018) are of high interest, as they described a worsening of cognition in the intervention group after long-term follow-up and thus do not justify recommendation of physical exercise interventions as a treatment for cognitive decline in community-dwelling pwd.

Although our findings are consistent with Forbes et al. [12], they contrast with other studies and reviews [46,

 Table 1
 Study characteristics of included studies

Study	Country	Sam	ple	Sample Age [years] size		Gender [% female]	e e	MMSE (mean ± SD)	an ± SD)	Diagnosis criteria	Type of training	bu	Trial duration [weeks]	Relevant outcomes	Risk of bias
		<u>ច</u>	មួ	10	9)	<u> </u>	9	10	95		10	g			
De Oliveira et al. [41]	Brazil	13	4	13 14 77.54±8.05 81.22	± 8.88	78.6	38.5	20.9土4.34	20.66±5.19	DSM-IV	Multimodal training	Usual care	12	Cognition	High concerns
Hoffmann et al. [37]	Denmark	107	93	107 93 69.8 ± 7.4	71.3 ± 7.3	47.7	38.7	47.7 38.7 23.8±3.4	24.1 ± 3.8	NINDS- ADRDA	Aerobic training	Usual care	16	Cognition ADLs NPS	Some con- cerns
Karssemeijer et al. [39]	The Nether- lands	39ª	38	80.9±6.1	79.8±6.5	44.7	46.2	22.5 ± 3.1	21.9 ± 3.1	DSM-IV	Aerobic training	Stretching	12	Cognition	Some con- cerns
Lamb et al. [42]	N N	329	165	329 165 76.9±7.7	78.1 ± 7.7	40	37	22.1 ± 4.6	22.1 ± 4.6	DSM-IV	Multimodal training	Usual care	16	Cognition ADLs NPS	Some con- cerns
Öhman et al. [43, 44]	Finland	2 2	70	77.5 ± 5.4 78.3 ± 5.1	78.1 ± 5.3	42.9	37.1	17.8±6.6 18.5±6.3	17.7 ± 6.2	NINDS- ADRDA	Multimodal training	Usual care	52	Cognition NPS	High concerns
Park et al. [38]	USA	= ;	10	84.3±7.7		41.9		14.7±5.65		Z	Strength training	Music inter- vention	12	NPS	High concerns
Yu et al. [40] USA	USA	5 2	32	77.4±6.6	77.5±7.1	4	47	21±3.5	22.2 ± 2.7	NINDS- ADRDA	Cnair yoga Aerobic training	Stretching	24	Cognition	High concerns

ADLs Activities of daily living, NPS Behavioral and psychological symptoms of dementia, CG Control group, SM-IV Diagnostic and Statistical Manual of Mental Disorders—fourth edition, IG Intervention group, MMSE Mini-Mental State Examination, NINDS-ADRDA National Institute of Neurological and Communicative Diseases and Stroke/Alzheimer's Disease and Related Disorders Association

^a RCT with 2 intervention groups, but one did not meet inclusion criteria of this review

Study	Type of training	Trial duration	Frequency [sessions/	Intensity	Session duration	Setting	Adherence	Assessment tool	Domain	Outcome Baseline (mean ± SD)	eline	Outcome Follo	Outcome Follow-up (mean ±SD)	<i>p</i> value	Conclusion
		[weeks]	week		Ē					<u> </u>	ទូ	9	99		
De Oliveira Silva et al.	Multi-modal training	12	2	70–80% VO ₂ max	09	Super- vised	87%	MMSE	Global cognition	20.58 ± 4.91	21.43 ± 4.18	20.31 ±4.68	19.5 ± 5.37	11.	No effect
[41]								CDT	Executive function	1.5	_	-	_	.54	
								٧F	Verbal flu- ency	12	6	10	10	.16	
								ST3	Selective attention, inhibitory control, and processing speed	42	41.53	43.51	40.08	6.	
Hoffmann et al. [37]	Aerobic training	16	m	70–80% HRmax	09	Super- vised groups of	76%	SDMT	Mental speed and attention	27.1±14.7	25.4 ± 14.3	26.2 ± 15.6	24.1 ± 14.9	.179	No effect
						2–5		ADAS-Cog verbal memory	Verbal memory (immediate recall)	11.2±4.5	11.2 ± 4.5	112±4.6	11.4±4.4	.813	
								٧F	Verbal flu- ency	23.2±12	24.2 ± 12.4	23.1 ± 11.8	25.6 ± 14.9	.133	
								SCWT	Psychomo- tor speed	17.6±10.2	18±9.6	17.5 ± 10.4	18.3 ± 9.9	.441	
								MSSE	Global cognition	23.8 ± 3.4	24.1 ± 3.8	23.9 ± 3.4	23.9 ± 3.9	.244	
Karssemeijer et al. [39]	Aerobic training	12	m	50-75% HRR	30-50	Super- vised individu-	81.1%	Z-score (TMT-B short, SCWT, VF, and RS)	Executive function	0.05±0.72	-0.03 ±0.8	0.15±0.74	-0.12±0.87	.338	No effect
						ally		Z-score (LLT)	Episodic memory	-0.08 ± 1.05	-0.08±0.86	-0.11±1.15	-0.34±1.21	.184	
								Z-score (WAIS- III DST and WMS-III SS)	Working memory	0.02±0.73	0.15 ± 0.95	0.04±0.80	-0.12±1.02	.153	
								Z-score (TMT-A and SCWT)	Psychomo- tor speed	0.14±0.73	0.0 ± 0.81	0.32 ± 0.64	-0.25 ± 1.04	.004 ^a	Favors intervention
								Z-score (WMS-R LM and HVLT-R)	Episodic memory	Z				7447	No effect
[42]	Multi-modal training	91	2 (+1) ^b	Moderate to hard	06-09	Super- vised groups of 6–8 and unsu- pervised home-	65.63%	ADAS-Cog	Global cognition	21.4±9.6	218±7.7	22.9 ± 11.6	224±94	.24	No effect

Table 2 (continued)

Study	Type of training	Trial duration	Frequency Intensity Session [sessions/ duration	Intensity	Session	Setting	Adherence	Assessment Domain tool	Domain	Outcome Baseline (mean ± SD)	seline	Outcome F	Outcome Follow-up (mean ± SD)	p value	Conclusion
		[weeks]			m [ri					<u>9</u>	99	<u> </u>	95		
Öhman et al. [43]	Multi-modal training	52	2	Z	09	Super- vised	72.88%	CDT	Executive	2.32±2.04	2.45 ± 2.09	₹		.03	Favors inter- vention
						home				2.31±2.09				.07	No effect
								ΛF	Verbal flu-	8.34±4.75	7.89±4.25	Z		.93	
						Super-	72.12%		ency	8.05±4.3					
						vised		MMSE	Global	17.8±6.6	17.7 ± 6.2	₹			
						of 10			cognition	18.5±6.3					
Yu et al. [40]	Aerobic training	24	es .	50-75% HRR	40–60	Super- vised	> 70%	ADAS-Cog	Global cognition	19.3 ± 7.4	17.8 ± 6.5	Z		.386	No effect
						groups up to 3		Z-score (WMS-R LM and HVLT-R)	Episodic memory	Z				.447	
								Z-score (TMT- B, EXIT-25, and CDT)	Executive function	Z				.849	
								WAIS-R DST	Attention	Z				.539	
								Z-score (WAIS- III DST and TMT A)	Processing speed	Z				.778	
								Z-score (COWAT, VF, and BNT)	Language	Z				.925	

ADA5-Cog Alzheimer's Disease Assessment Scale – Cognitive Subscale, BNT Boston Naming Test, CDT Clock-Drawing Test, GG Control group, COWAT Controlled Oral Word Association Test, HRmax Maximal heart rate, EXTZ5 Executive Interview-25 Items, HVLT-R Hopkins Verbal Learning Test—Revised, HRR Heart rate reserve, IG Intervention group, LLT Location Learning Test, MMSE Mini-Mental State Examination, NI No information, RS Rule Shift Card Test, SCWT Stroop Color and Word Test, SDMT Symbol Digit Modalities Test, ST3 Stroop Test – third card, TMT-A Trail Making Test – Subtest A, TMT-B Trail Making Test – Subtest B, VF Verbal Fluency Test, VO₂max Maximal aerobic capacity, WMS-III DSTWechsler Adult Intelligence Scale – Third Edition Digit Span Test, WAIS-R DST Wechsler Adult Intelligence Scale – Revised Digit Symbol Test, WMS-R Wechsler Memory Scale – Third Edition Spatial Span

^a Group differences in the changes marked as significant in publication

 $^{^{\}mathrm{b}}$ Group-based training was offered 2 times per week, but participants were encouraged to perform one session at home

Table 3	Table 3 Included studies evaluating the effect of exercise programs on ADLs	s evaluating	יויר בווכבר כו											
Study	Type of training	Trial duration	Frequency [sessions/	Frequency Intensity [sessions/	Session	Setting	Adherence	Adherence Adherence Outcome Baseline (mean ± SD)	Outcome Bas (mean ± SD)	Baseline D)	Outcome foll (mean ± SD)	Outcome follow-up P-value Conclusion (mean ± SD)	P-value	Conclusion
		[weeks	week]		[min]				<u>ల</u>	CG	<u>9</u>	CG		
Hoffmann et al. [37]	Aerobic training	16	м	70–80% HRmax	09	Supervised 76% groups of 2–5	76%	ADCS-ADL 64.8±8.8 62.4±10.8 64.4±9.4 62.7±10.4 .868	64.8±8.8	62.4±10.8	64.4±9.4	62.7 ± 10.4	.868	No effect
Lamb et al. [42]	Multimodal 16 training	91	2 (+1) ^a	Moderate to 60–90 hard	06-09	Supervised groups of 6–8 and unsuper- vised home- based	65.63%	BADLS	01	1	14.6 ± 9.5	14.6±9.5 14.6±10.4 .15	.15	No effect

ADCS-ADL Alzheimer's Disease Cooperative Study ADL Scale, BADLS Bristol ADLs Scale, CG Control group, HRmax Maximal heart rate, IG Intervention group, NI No information

^a Group-based training was offered 2 times per week, but participants were encouraged to perform one session at home

 Table 4
 Included studies evaluating the effect of exercise programs on NPS

Study	Type of training	Trial		Intensity	Session	Setting	Adherence	Assessment tool	Outcome Baseline (mean ± SD)	seline	Outcome follow-up (mean±SD)	dn-w	P-value	P-value Conclusion
		[weeks]	week]		<u>u</u>				פ	CG	10	CG		
Hoffmann et al. [37]	Aerobic training	91	м	70–80% HRmax	09	Super- vised groups of 2-5	76%	NPI-12	10	9.4	8.8	4.1	.002 ^b	Favors interven- tion
Lamb et al. [42]	Multi- modal training	91	2 (+1)³	Moderate to hard	06-09	Super- vised groups of 6–8 and unsu- pervised home- based	65.63%	<u>a</u>	ω	0	12	8.5	4	No effect
Öhman et al. [44]	Multi- modal training	52	7	₹	09	Super- vised home based	72.88%	IdN	13.5±12.6 16.6±15.2		2.73 (1.08 to 5.05) ²	0.64 (– 2.23 to 3.46) ^c	4. L	No effect
						Super- vised groups of 10	72.12%		12.1 ± 9.8		0.88 (-1.30 to 2.84) ²			
Park et al. [38]	Strength training	12	2	₹	45	Super- vised groups	77.92%	CMAI	39.55±8 44.3±8.74 6.82±3.6	42.89±11.79 9.4±4.09	45.33±11.96 53.25±16.27 8.33±5	46.4±12.38 12.3±4.76	90. 74. 18:	No effect
	Chair yoga						76.25%	ssion anxiety	ω.		8.38±4.07 6.78±3.42 8.5±6.4	7.89±3.18	.89 .34	

CG Control group, CMAI Cohen-Mansfield Agitation Inventory-Short Form, HADS Hospital Anxiety and Depression Scale, HRmax Maximal heart rate, NPI Neuropsychiatric Inventory, NPI-12 Neuropsychiatric Inventory – 12 item version, IG Intervention group, NI No information

^a Group-based training was offered 2 times per week, but participants were encouraged to perform one session at home

 $^{^{\}rm b}$ Group differences in the changes marked as significant in publication

^c Change from baseline

47]. Differences in the study population might explain why our findings are inconsistent with other systematic reviews, which describe exercise as an effective treatment for cognitive decline in pwd without a formal diagnosis [46, 47]. De Oliveira et al. [41] did find significant improvements in cognition after a multimodal training intervention in people with MCI, but not in those with dementia and thus concluded that physical exercise should only be recommended in the early stages of neurocognitive disorders. Therefore, it seems useful to distinguish between people with MCI and pwd, since people affected experience different symptoms and biological adaptions so that possible mediators by which physical activity improves cognition may occur differently as the disease progresses [48]. In order to establish clear evidence and recommendations for physical activity in pwd, it is necessary to analyze a homogenous group in terms of diagnosis, as trainings recommendations might not be applicable for pwd as for people with MCI [49].

However, occasional significant superiorities for intervention groups in cognitive subdomains could be identified in two trials [39, 43]. Analyzing cognition, three trials used supervised sessions in groups [37, 40, 42], one did not report how the intervention was delivered [41], one compared two different settings [43], and one performed exercise individually guided [39]. Apart from Öhman et al. [43], none of the included trials reported beneficial effects for executive functions. These inconsistencies could have been arisen through the trainings content or the duration of the trial, as this was the only trial including training of executive function as part of the intervention, which lasted 52 weeks. Nevertheless, it needs to be stated that this multimodal training did lead to improvements in executive function for the homebased intervention group, while the same intervention did not affect executive function in a group-based setting. Deriving from this, there is an indication that training intervention for community-dwelling pwd are most beneficial if they are delivered individually guided and customized through a healthcare provider or the person's caregiver. This hypothesis may further explain deviating findings from Karssemeijer et al. [39], who described, in contrast to other trials [37, 40], a significant amelioration in psychomotor speed after 12 weeks of aerobic training in the intervention group. Since training modalities, adherence to the protocol, design of control groups, measurements of the outcome, and exercises did not differ widely between the three trials, individual training sessions seem to be preferable as they might not overwhelm pwd and allow individually adapted designs.

Effects on activities of daily living

In contrast to a previous Cochrane review [12], we could not find any beneficial effects on ADLs through physical activity in pwd. According to this review, there has been an unexplainable high heterogeneity between included studies, with only two trials [50, 51] out of six [52-55] favoring exercise over control. Furthermore, the sample size of studies included in the Cochrane review was comparatively small, ranging from six to 56 participants in the intervention group. Especially the power of the two studies observing beneficial effects on ADLs is limited, due to the sample size of eight [50] and eleven [51] participants in the intervention group. Therefore, Forbes et al. [12] suggested that these findings should be interpreted with caution and rated the quality of the evidence as low. Despite larger sample sizes in the included trials of this this review, ranging from 107 [37] to 329 [42], there were only two trials analyzing the effects of exercise on ADLs in pwd. We could not find an impact of adherence to the protocol.

Table 5 Risk of bias assessment of the included studies

Study	Domain 1: Risk of bias arising from the randomization process	Domain 2: Risk of bias due to deviations from the intended intervention	Domain 3: Risk of bias due to missing outcome data	Domain 4: Risk of bias in measurement of the outcome	Domain 5: Risk of bias in selection of the reported result	Overall risk of bias
De Oliveira et al. [26]	Low concerns	High concerns	High concerns	High concerns	Some concerns	High concerns
Hoffman et al. [37]	Some concerns	Some concerns	Low concerns	Low concerns	Low concerns	Some concerns
Karssemeijer et al. [39]	Low concerns	Some concerns	Low concerns	Low concerns	Low concerns	Some concerns
Lamb et al. [42]	Low concerns	Some concerns	Low concerns	Low concerns	Low concerns	Some concerns
Öhman et al. [43, 44]	Low concerns	Some concerns	Low concerns	High concerns	High concerns	High concerns
Park et al. [38]	Some concerns	Some concerns	High concerns	High concerns	Some concerns	High concerns
Yu et al. [40]	Low concerns	Low concerns	High concerns	Low concerns	High concerns	High concerns

Effects on neuropsychiatric symptoms

Trials investigating the effects of exercise on NPS showed inconsistent results. Since training modalities such as duration, intensity, and setting did not differ widely between trials, different types of training might explain divergences. While strength and multimodal trainings intervention showed no beneficial effects on NPS [38, 42, 44], Hoffmann et al. [37] described a reduction of NPS after 16 weeks of aerobic training. Since only one trial analyzed the impact of aerobic exercise on BSPD, it seems possible that aerobic exercise might be effective to reduce NPS in pwd and this is in line with a recently published review [56]. Even though this might seem plausible, we could not find evidence for differences between studies with active or passive control groups or deviations in exercise adherence.

Limitations

Although the search was conducted in eight different databases and 7651 trials were identified, we cannot rule out the possibility that we might have missed relevant trials due to limitations in language and year of publication. This might also be applicable for the requirement of a formal diagnosis of dementia. Some trials were excluded in this review because they included participants based on the results of screening instruments.

Conclusion

Implications for practice

There is little evidence that both strength and aerobic exercise or a combination of these cannot be recommended as a treatment option for cognitive impairment in community-dwelling pwd. Furthermore, moderate to high-intensity interventions might even worsen the cognitive decline in community-dwelling pwd after finishing the intervention [42]. In this context, it is mentionable that this was the study with the highest methodological quality and largest sample size. Furthermore, there is no evidence for the beneficial effects of exercise for ADLs. The effects on NPS are unclear, as one out of three studies found improvements after aerobic training. That is why healthcare providers and caregivers should be confident to promote the maintenance of an active and healthy lifestyle [57] among community-dwelling pwd instead, although recent recommendations [58] of moderate-intensity aerobic exercise for community-dwelling pwd are not underpinned by the results of this review. The development of best practice guidelines for healthcare providers is urgently needed. Exercise adherence does not seem to influence these outcomes.

Implications for research

As our review shows, there is a necessity for improvement in methodological approaches in the research of the effects of exercise on cognition, ADLs, and NPS in community-dwelling pwd. Due to its large sample size and high methodological quality, the trial of Lamb et al. [42] should be considered as a best practice example. Recent research recommends at least 150 min/week of moderate-intensity aerobic exercise for older people, but this might not be appropriate for community-dwelling pwd [58]. Following on from this, future RCTs should require a formal diagnosis of dementia and should distinguish between pwd and people with MCI, as the conditions lead to different capabilities and needs, so that effects of exercise could therefore result in different outcomes [41]. High methodological quality, large sample sizes and longterm follow-ups should be implemented in future trials. In respect to possible impacts of social stimulation and activities on cognition and NPS, control groups should be designed accordingly to the intervention group. Especially if supervised group sessions are analyzed in a trial, control group should receive comparable social stimulation. To give answer to the question which type of training is most beneficial for community-dwelling pwd, it would be necessary that training modalities are described in detail and to compare different exercise protocols within three-armed RCTs. To compare different exercise programs and to be able to transfer research results in practice, it is inescapable to give a detailed description of the content and exercises of the trials, as it was the case in most of the included studies.

Registration

This review was registered at the national prospective register of systematic reviews and no amendments were made (PROSPERO registration number: CRD42021246598). To view please visit https://www.crd.york.ac.uk/prospero/display_record.php?RecordID=246598.

No protocol was published in advance.

Abbreviations

ADAS-Cog: Alzheimer's Disease Assessment Scale; ADCS-ADL: Alzheimer's Disease Cooperative Study ADL Scale; ADLs: Activities of daily living; BADLS: Bristol Activities of Daily Living Scale; CMAI: Cohen-Mansfield Agitation Inventory-Short Form; HADS: Hospital Anxiety and Depression Scale; MCI: Mild cognitive impairment; MMSE: Mini-Mental Status Examination; NPI: Neuropsychiatric Inventory; NPS: Neuropsychiatric Symptoms; PRISMA: Preferred Reporting Items for Systematic Reviews and Meta-Analysis; PROSPERO: National Prospective Register of Systematic Reviews; pwd: People with dementia; RCTs: Randomized controlled trials; RoB2: Revised Cochrane Risk-of-Bias Tool for Randomized Trials; SCI: Subjective cognitive impairment.

Supplementary Information

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Additional file 1. Risk of bias assessment. This additional file provides the detailed description of risk of bias assessment after discussion of both authors (K.S. and N.D.).

Additional file 2. Search strategy. This file provides the full search strategy for each database with detailed description (e.g. additional filters and dates of search).

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Authors' contributions

The present manuscript was written by K.S. A.K. was involved in the screening process and in the review of the extracted data. Risk of bias assessment was performed by K.S. and N.D. P.K.-R. was a major contributor in writing the manuscript and in the screening process. All authors read and approved the final manuscript.

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Availability of data and materials

Data sharing is not applicable to this article as no datasets were generated or analyzed during the current study.

This paper has not been previously published and is not currently under consideration for publication elsewhere.

Declarations

Ethics approval and consent to participate

Not applicable.

Consent for publication

Not applicable.

Competing interests

The authors declare that they have no competing interests.

Author details

¹Friedrich-Alexander-Universität Erlangen-Nürnberg, Interdisciplinary Center for Health Technology Assessment and Public Health (IZPH), Erlangen, Germany. ²Department of Psychiatry and Psychotherapy, Center for Health Services Research in Medicine, University Hospital Erlangen, Erlangen, Germany. ³Friedrich-Alexander-Universitat Erlangen-Nurnberg, Department of Medical Informatics, Biometrics and Epidemiology, Erlangen, Germany.

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