REGULAR ARTICLE

Decreased activity of daily living produced by the combination of Alzheimer's disease and lower limb fracture in elderly requiring nursing care

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Received: 11 March 2012/Accepted: 18 April 2012/Published online: 11 May 2012 © The Japanese Society for Hygiene 2012

Abstract

Objectives Alzheimer's disease (AD) impairs cognitive functions, subsequently decreasing activity of daily living (ADL), and is frequently accompanied by lower limb fracture including hip fracture in the elderly. However, there have been few studies on what kinds of physical functions are affected or what degrees of dysfunction are produced by this combination. This study aims to clarify the relationship between decreased ADL and the combination of AD and lower limb fracture.

Methods We examined present illness and ADL in 4340 elderly aged 82.8 \pm 9.36 years [average \pm standard deviation (SD)] requiring nursing care and compared ADL between elderly with and without AD or lower limb fracture treated with surgery or conservatively using analysis of covariance (ANCOVA), with age and sex as covariants. *Results* We recognized that activities of cognitive function (p < 0.001), eating (dysphagia) (p < 0.001), eating (feeding) (p < 0.001), and toilet use (p < 0.001) in the elderly with AD were significantly lower than in those without the disease, even after adjusting for sex and age. Activities of bed mobility (p < 0.05), transfer and locomotion (p < 0.001), and bathing (p < 0.05) in the elderly with a fracture treated with surgery were significantly lower, which differed from the results of AD. Significant interactions of AD and fracture treated with surgery on the ADL scores for bed mobility (p < 0.001), dysphagia (p < 0.01), feeding (p < 0.001), and

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toilet use (p < 0.05) show that the combination had a much more profound influence on the ADL scores than AD or fracture alone. We obtained almost the same results for fractures treated conservatively as for fractures treated with surgery.

Conclusions These results demonstrated that the combined effects of AD and lower limb fracture were significantly greater than expected additive effects of AD and fracture, suggesting that the combination of AD and lower limb fracture has synergistic effects on almost all types of ADL except cognitive functions.

Keywords Alzheimer's disease · Hip fracture · Elderly · Activity of daily living · Physical function

Introduction

Alzheimer's disease (AD) is the most common form of dementia. The number of individuals with AD has been increasing considerably in recent years, accounting for more than 50 % of dementia cases [1]. AD causes cognitive impairment of elderly in need of care, consequently interfering with daily tasks and decreasing activities of daily living (ADL) in many ways [2]. It is known that patients with AD suffer from various kinds of complications [3]. The increased risk of falling associated with AD leads to lower limb fracture including hip fracture [4]. In addition, taken together with the fact that AD and osteoporosis have common risk factors [5, 6], the incidence of fracture is higher in elderly with AD. Thus, lower limb fractures could accompany AD as one of its complications.

To make matters worse, elderly made bedridden by the aftereffects of lower limb fracture could develop cognitive impairment as its complication [7]. Even if an elderly person

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is not made bedridden, the aftereffects of fracture sometimes impair cognitive functions, subsequently decreasing basic ADL [8–10]. AD and fracture, which are commonly encountered in the elderly, aggravate each other, being the two major factors subsequently deteriorating ADL in bedridden elderly. Therefore, AD and fracture produce a vicious spiral, resulting in not only cognitive impairments but also deterioration in ADL [3, 5]. However, it remains to be elucidated exactly what kinds of physical functions are affected and to what degree ADL is changed by the combination of AD and lower limb fracture. To clarify this, we performed a national survey in nursing care institutions in Japan, examining the decreased ADL in elderly with and without AD and lower limb fracture.

Subjects and methods

Subjects

Three hundred and ninety facilities were randomly selected out of 3410 nursing care institutions in Japan. Ten resident patients and 10 daycare patients were sampled at random from each facility. A total of 7800 patients were prospective subjects in this study. We mailed a questionnaire to caregivers in each facility inquiring about ADL and medical conditions, including AD and fracture, as described below and obtained 4340 responses (55.6 %, 2132 resident patients and 2208 daycare patients). This survey was conducted from December 2009 to February 2010. The average age and SD of the 4340 subjects were 82.8 and 9.36 years. This research was conducted after obtaining approval from the ethical committee of the Japan National Conference of Geriatric Health Care Facility.

Questionnaire

The questionnaires regarding medical conditions in subjects were related to AD and lower limb fracture (hip fracture and lower leg fracture) requiring surgery, and fractures treated without surgery. To examine AD, the questions concerned neuropsychological examinations including the patients' medical history, neurological testing, Mini-Mental State Examination [11], and standard clinical evaluation including brain scanning. In the diagnostic process, DSM-III [12] was used for diagnosis and to rule out other factors. The National Institute of Neurological and Communicative Disorders and Stroke-Alzheimer's Disease and Related Disorders Association guidelines [13] were used for detection and diagnosis of either possible or probable AD. Lower limb fracture was diagnosed after an appropriate procedure including X-ray roentgenogram. The diagnoses of AD and fracture were performed by a physician of the facility within 1 year prior to the contact with nursing care institutions for elderly in this study. Our subjects included 491 patients with possible or probable AD, 405 and 196 patients with fracture treated with surgery and conservatively, respectively, and 3248 elderly without AD or fracture.

Regarding ADL, the questionnaire was developed on the basis of a standard described previously [14, 15]. They dealt with bed mobility, transfer and locomotion, cognitive function (orientation), cognitive function (communication), cognitive function (mental activity), eating (dysphagia), eating (feeding), toilet use, and bathing for elderly requiring both residential nursing care and daycare. Each ADL was categorized into a five-point scale. In the case of bed mobility, for example, a score of 5 represents being able to stand on one's feet and maintain this posture; score 4: having difficulty maintaining a standing posture, but being able to transfer from one place to another in a sitting position; score 3: being unable to move in a sitting position, but being able to sit in a proper posture without support; score 2: having difficulty sitting in an upright posture, but being able to turn over on a bed; and score 1: not being able to turn over on a bed. In the case of eating (feeding), a score of 5 represents being able to eat well without any support of others; score 4: spilling food during eating; score 3: having difficulty eating by themselves, but being able to eat with support for their posture and position of the dish; score 2: being unable to eat without complete support; and score 1: being unable to eat with any support (receiving tube feeding). Thus, the requirement for more concentrated nursing care during eating for the elderly decreased the ADL score. A lower score for each ADL implies worse ADL. In Table 1, we show ADL and prevalence of AD, fracture treated with surgery, and fracture treated conservatively according to the status of nursing care (residential and daycare).

Statistics

Pearson's correlation coefficients of age showed weak but significant correlations with bed mobility (-0.118), transfer and locomotion (-0.145), cognitive function (orientation) (-0.262), cognitive function (communication) (-0.199), cognitive function (mental activity) (-0.232), eating (dysphagia) (-0.142), eating (feeding) (-0.088), toilet use (-0.126), and bathing (-0.077) (all p < 0.001). Because of significant differences in proportions in terms of sex and age between elderly with and without diseases in addition to significant correlations between age and ADL scores, we compared each ADL score by analysis of covariance (ANCOVA) with age and sex as covariants, in which age was assigned as a continuous variable and sex was dummy-coded as follows: male 0, female 1. Mean

Status	Number	Proportion	Age (years;	Activities of	daily living (ag	e-adjusted scor	e; mean \pm SD)					
		of women (%)	mean ± SD)	Bed mobility	Transfer and locomotion	Cognitive function (orientation)	Cognitive function (communication)	Cognitive function (mental activity)	Eating (dysphagia)	Eating (feeding)	Toilet use	Bathing
Residential Daycare	2132 2208	79.7 57.3	85.2 ± 8.34 80.5 ± 9.20	3.27 ± 1.45 4.24 ± 1.05	2.07 ± 0.72 2.74 ± 0.99	3.31 ± 1.33 4.25 ± 1.03	3.24 ± 1.36 4.19 ± 1.12	3.25 ± 1.34 4.22 ± 1.09	3.84 ± 1.28 4.66 ± 0.80	$\begin{array}{c} 4.01 \pm 1.25 \\ 4.64 \pm 0.74 \end{array}$	3.22 ± 1.26 4.23 ± 1.06	2.83 ± 0.69 3.35 ± 0.91
Status		Number	Proporti	on of women (r (%)	Age (years; me	an \pm SD)	Prevalence (ag	ge-adjusted scor	.e)		
								AD	Fracture (su	rgery)	Fracture (co	nservatively)
Residential		2132	79.7		~	35.2 ± 8.34		17.0	12.2		5.53	
Daycare		2208	57.3		~	80.5 ± 9.20		5.80	6.57		3.53	

ADL score and standard deviation adjusted by age and sex were calculated using regression coefficients corresponding to age, sex, and disease obtained by ANCOVA and raw mean values of age and sex in total subjects. To assess the interaction for ADL score between the combination of AD and fracture treated with or without surgery, we used two-way ANCOVA with age and sex as covariants. The statistical software SPSS version 17 was used. Twotailed *p*-values less than 5 % were considered statistically significant.

Results

Changes in ADL with AD and fracture

The proportion of women and age of subjects with AD were significantly higher than those without AD. The activities of cognitive function (orientation), cognitive function (communication), cognitive function (mental activity), eating (dysphagia), eating (feeding), and toilet use in the elderly with AD were significantly lower than those without the disease, even after controlling for sex and age (Table 2). Activities of bed mobility, transfer and locomotion, and bathing in elderly with a fracture requiring surgical operation were significantly lower, which were quite different results from AD (Table 3). Table 4 also shows lower scores on bed mobility, transfer and locomotion, and bathing of the patients with a fracture treated without surgery.

The interaction between AD and fracture

The two-way ANCOVA demonstrated a significant main effect of AD on ADL scores of all cognitive functions and eating functions, as well as toilet use, and a significant main effect of fracture on all scores of ADL except cognitive functions. Furthermore, we recognized significant interactions between AD and fracture on the activities of bed mobility, eating (dysphagia), eating (feeding), and toilet use after adjustment for age and sex (Table 5). Table 6 shows ADL scores in the cases of patients with complications of AD and fracture treated conservatively. ADL scores of all cognitive functions and eating functions, as well as toilet use, in the elderly with AD were significantly decreased compared with those without AD. The scores of all ADL except cognitive functions and eating functions were significantly decreased compared with those without fracture treated conservatively. Interactions between ADL and fracture treated conservatively were recognized for the scores of bed mobility, transfer and locomotion, and toilet use even after adjusting for sex and age. There were decreases in the scores of eating (dysphagia), eating

Table 2	Comparisons of act	tivities of daily	living in patien	its with and wi	thout AD						
AD	Number	Age (years;	Activities of (daily living (a£	ge- and sex-adjust	ted score; mean \pm S	D)				
	(proportion of women, %)	mean ± SU)	Bed mobility	Transfer and locomotion	Cognitive function (orientation)	Cognitive function (communication)	Cognitive function (mental activity)	Eating (dysphagia)	Eating (feeding)	Toilet use	Bathing
(-)(+)Statistics	3849 (67) 491 (82) ***	82.5 ± 9.56 85.3 ± 7.07 ***	3.76 ± 1.30 3.78 ± 1.35	2.41 ± 0.87 2.44 ± 0.91	3.92 ± 1.18 2.77 ± 1.17 ***	3.83 ± 1.24 2.85 ± 1.26 ***	3.86 ± 1.18 2.84 ± 1.24 ***	4.28 ± 1.12 4.07 ± 1.13 ***	4.36 ± 1.05 4.09 ± 1.06 ***	3.77 ± 1.24 3.46 ± 1.26 ***	3.10 ± 0.81 3.03 ± 0.84
ANCOV/ In the ant	A was used to detect dysis of the differe	t significant diffence in age and p	erence in the sco proportion betw	ore in elderly v veen sexes, Stu	with AD compared ident's t test and	d with the correspond χ^2 test were used, re	the function of the second se	ithout AD after 0.001	adjustment for	age and sex, *	$^{**}p < 0.05.$
Table 3	Comparisons of act	ivities of daily l	living in patien	ts with and wit	thout fracture trea	ated with surgery					
Fracture	Number	Age (years;	Activities of 6	daily living (ag	ge- and sex-adjust	ted score; mean \pm S	(D)				
	(proportion of women, %)	mean ± SD)	Bed mobility	Transfer and locomotion	Cognitive function (orientation)	Cognitive function (communication)	Cognitive function (mental activity)	Eating (dysphagia)	Eating (feeding)	Toilet use	Bathing
(-)	3935 (66)	82.3 ± 9.40	3.78 ± 1.32	2.43 ± 0.88	3.79 ± 1.19	3.73 ± 1.25	3.75 ± 1.25	4.26 ± 1.13	4.33 ± 1.07	3.74 ± 1.25	3.10 ± 0.82
(+)	405 (89)	87.4 ± 7.45	3.60 ± 1.35	2.24 ± 0.93	3.77 ± 1.25	3.66 ± 1.33	3.74 ± 1.29	4.26 ± 1.15	4.32 ± 1.09	3.67 ± 1.27	3.00 ± 0.85
Statistics	***	***	*	***							*
ANCOV ^{β} after adju: p < 0.001	A was used to detect stment for age and	t significant diffe sex, * $p < 0.05$	erence in the scc $; *** p < 0.00$	ore in elderly w 1. In the analy:	ith fracture treatesis of the differer	d with surgery components in age and propo	ared with the corresp ortion between sexes	onding one in el , Student's <i>t</i> tes	Iderly without f it and χ^2 test w	fracture treated /ere used, resp	with surgery ectively; ***

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Table 4	Comparisons of acti	ivities of daily l	living in patien	ts with and wit	thout fracture tre	ated conservatively					
Fracture	Number	Age (years;	Activities of d	laily living (ag	e- and sex-adjus	ted score; mean \pm S	D)				
	(proportion of women, %)	mean ± SD)	Bed mobility	Transfer and locomotion	Cognitive function (orientation)	Cognitive function (communication)	Cognitive function (mental activity)	Eating (dysphagia)	Eating (feeding)	Toilet use	Bathing
(+)	4144 (68) 196 (83) ***	82.7 ± 9.41 85.3 ± 7.79 ***	3.77 ± 1.29 3.56 ± 1.33 *	2.42 ± 0.90 2.25 ± 0.91 *	3.79 ± 1.22 3.87 ± 1.23	3.72 ± 1.29 3.70 ± 1.30	3.74 ± 1.22 3.80 ± 1.27	4.26 ± 1.09 4.18 ± 1.12	4.33 ± 1.03 4.35 ± 1.06	3.74 ± 1.22 3.63 ± 1.25	3.10 ± 0.84 2.96 ± 0.84 *
ANCOVA conservati p < 0.001	A was used to detec ively after adjustme	ct significant di nt for age and	fference in the sex, $* p < 0.05$	score in elder . In the analys	rly with fracture is of the differen	treated conservative nce in age and propo	ly compared with t rtion between sexes	the correspondi , Student's t te:	ng one in elde st and χ^2 test w	rly without fr vere used, resp	cture treated ectively; ***
Table 5	Comparisons of acti	vities of daily 1	iving in patient	ts with and wit	thout AD and fra	icture treated with su	ırgery				
Disease	Number	Age (years	; Activities of	of daily living	(age- and sex-ad	justed score; mean =	E SD)				
	(proportion of women, %)	IIICAII II 21	Ded mobility	Transfer and locomotion	Cognitive function (orientation)	Cognitive function (communication	Cognitive function (mental activity)	Eating (dysphagia)	Eating (feeding)	Toilet use	Bathing
(-)	3495 (64)	82.0 ± 9.6	1 3.77 ± 1.3	$0 2.43 \pm 0.8$	9 3.93 ± 1.18	3.84 ± 1.24	3.86 ± 1.24	4.28 ± 1.12	4.35 ± 1.06	3.77 ± 1.24	3.11 ± 0.83
AD	440 (81)	84.9 ± 7.0	9 3.86 ± 1.3	4 2.48 \pm 0.9	$0 2.77 \pm 1.17$	2.88 ± 1.26	2.85 ± 1.24	4.12 ± 1.11	4.15 ± 1.07	3.51 ± 1.26	3.06 ± 0.84
Fracture	354 (88)	87.2 ± 7.6	$3 3.69 \pm 1.3$	5 2.27 \pm 0.9	$2 3.89 \pm 1.19$	3.79 ± 1.28	3.85 ± 1.24	4.34 ± 1.13	4.42 ± 1.07	3.75 ± 1.26	3.02 ± 0.85
Combinat	ion 51 (94)	88.5 ± 5.9	$3 3.01 \pm 1.3$	4 2.05 ± 0.9	$1 2.74 \pm 1.18$	2.65 ± 1.26	2.78 ± 1.24	3.63 ± 1.12	3.60 ± 1.06	3.05 ± 1.25	2.81 ± 0.84
Main effe of AD	ct				* *	* *	***	* * *	* *		
Main effe of fractu	ct Ire		* * *	* * *				*	* *	*	*
Interaction	u		***					*	***	*	
Two-way four grout	ANCOVA with age	and sex as covito to be significant	ariants was use thy different (bo	d in the analysist $n < 0.001$	is of the different by v^2 test and c	ce in ADL score, $* p$ me-way analysis of y	< 0.05, ** p < 0.0	1, *** $p < 0.00$	1. The sex and	age distributio	ns among the

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Disease	Number	Age (years;	Activities of c	laily living (ag	ge- and sex-adju	sted score; mean \pm	SD)				
	(proportion of women, %)	mean ± SD)	Bed mobility	Transfer and locomotion	Cognitive function (orientation)	Cognitive function (communication)	Cognitive function (mental activity)	Eating (dysphagia)	Eating (feeding)	Toilet use	Bathing
(-	3676 (68)	82.34 ± 9.61	3.77 ± 1.33	2.42 ± 0.91	3.92 ± 1.15	3.83 ± 1.21	3.86 ± 1.21	4.29 ± 1.09	4.36 ± 1.03	3.77 ± 1.21	3.11 ± 0.85
AD	468 (82)	85.15 ± 7.10	3.82 ± 1.34	2.46 ± 0.91	2.77 ± 1.17	2.86 ± 1.28	2.84 ± 1.23	4.09 ± 1.12	4.11 ± 1.06	3.50 ± 1.25	3.05 ± 0.84
Fracture	173 (83)	85.05 ± 7.96	3.66 ± 1.33	2.30 ± 0.91	3.99 ± 1.17	3.83 ± 1.26	3.90 ± 1.24	4.25 ± 1.12	4.41 ± 1.07	3.74 ± 1.25	2.99 ± 0.84
Combination	23 (78)	87.22 ± 6.16	2.86 ± 1.34	1.89 ± 0.91	2.80 ± 1.17	2.64 ± 1.27	2.87 ± 1.23	3.64 ± 1.12	3.82 ± 1.06	2.73 ± 1.25	2.68 ± 0.84
Main effect of AD					* * *	***	***	* *	* *	***	
Main effect of fracture			* *	* * *						* *	*
Interaction			*	*						*	
Two-way ANG four groups w	OVA with age an	nd sex as covaria be significantly	unts was used in different (both	the analysis c $p < 0.001$) by	of the difference χ^2 test and one	in ADL score, * $p <$ -way ANOVA, resp	<pre>< 0.05, ** p < 0.01 ectively</pre>	, *** p < 0.00	1. The sex and	age distribution	is among the

(feeding), and bathing in the patients with the combination, but these were not statistically significant (Table 6).

Discussion

Decreased ADL in eating (dysphagia), eating (feeding), and toilet use as well as cognitive functions, but not in bed mobility, or transfer and locomotion, were recognized in AD patients in this study. Although AD in our subjects was not considered to produce symptoms severe enough to affect motor functions, the influences of AD on eating (dysphagia), eating (feeding) [16, 17], and toilet use [18] seemed to be associated with cognitive impairments due to AD. These findings are supported by epidemiological studies on the natural course of the progress of AD [19]. In contrast to the results for AD, the patients with fracture treated either with or without surgery showed decreased ADL scores in bed mobility, transfer and locomotion, and bathing. Motor dysfunctions including difficulties in bathing are common following a hip fracture or a lower limb fracture [20, 21]. Thus, many epidemiological studies have evaluated effects of AD or lower limb fracture on individual ADL in the elderly, albeit separately.

The kinds of function affected and the degrees of change in ADL in patients suffering from the combination of AD and lower limb fracture are clarified by the observation that the kinds of dysfunction for AD and fracture were quite different. Our results showed significant interactions on the ADL scores of bed mobility, eating (dysphagia), eating (feeding), and toilet use, demonstrating that the combination of AD and fracture treated with surgery has more profound influences on the ADL scores than AD or fracture alone. This implies that the combined effect was significantly greater than the expected additive effect of AD and fracture. The decreased ADL score of transfer and locomotion with this combination was not recognized as involving a significant interaction between AD and fracture with surgery, but it was in cases without surgery. Thus, the combined effects of AD and lower limb fracture on ADL excluding cognitive functions and bathing were much larger than the sum of the sole effects of AD and fracture. The lack of an interaction for cognitive functions seems to be due to severe influences of AD on cognitive functions, for which complication with fracture did not bring any further deterioration [2, 3].

A fracture in a lower limb has a much greater influence on toilet use because this ability largely depends upon the functions of the lower limbs [18]. However, there are few studies demonstrating how the complication of lower limb fracture with AD affects dysphagia or eating activity, which is unaffected by the fracture alone [22]. Our findings showing that fracture in a lower limb aggravates dysfunctions in eating (dysphagia) and eating (feeding) produced by AD suggest that the fracture makes AD worse and more progressive. This finding is supported by many studies [23– 25], which have shown that a lower limb fracture affects the central nervous system through motor dysfunction, resulting in cognitive impairment.

On the other hand, our finding showing that the activities of bed mobility, and transfer and locomotion that were decreased by the fracture were further deteriorated by its complication with AD supports the idea that AD has greater influence on motor functions than expected. It is well known that osteoporosis, which often accompanies AD, leads to motor dysfunctions [5]. Some studies have demonstrated that problems in abnormal behaviors in AD patients may produce muscular dystrophy, which further damages physical functions [18, 25]. Taking these findings together, motor dysfunctions such as in bed mobility, and transfer and locomotion are considered to be accelerated by the complication of fracture with AD.

Women are at higher risk of developing AD and fracture [26, 27]. Our results showed that women account for 94 % of patients with the complications of AD and fracture. Therefore, the synergistic effects of AD and lower limb fracture on ADL seen in this study explain well that ADL in all fields in elderly women are lower than those in elderly men of the same age.

There were few differences in decreased ADL between cases of fracture treated with and without surgery. Our data regarding fracture treated with and without surgery do not coincide well with the findings that patients with fracture treated with surgery show better prognosis in terms of ADL than those treated without surgery [7, 10]. We did not obtain any data regarding social circumstances and physical background under which the patients with fracture did not receive surgery. Analysis of the data might confirm effects of surgery on the recovery from fracture, which was not seen in this study. Since this study was performed with a cross-sectional design, the causal effects of AD and fracture on ADL must be demonstrated by further follow-up study.

In conclusion, our results showed that the combination of AD and lower limb fracture has more profound influence on ADL scores in terms of bed mobility, transfer and locomotion, eating (dysphagia), eating (feeding), and toilet use, demonstrating that the combined effects were significantly greater than expected additive effects of AD and fracture, suggesting that the complication of AD with lower limb fracture has synergistic effects on almost all types of ADL besides cognitive functions. The interaction between AD and fracture on ADL seemed to be due to effects of both AD on motor function and fracture on central nervous functions. Acknowledgments The work was funded by a Grant-in-Aid for Scientific Research (B) from the Ministry of Health, Labor, and Welfare (2009–2010), Japan.

Conflict of interest None.

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