The Association between Activity of Daily Living and the Combination of Alzheimer’s Disease and Cataract in Elderly Requiring Nursing Care

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Abstract

Some clinical and epidemiological studies have suggested that Alzheimer’s Disease (AD) and cataract, may share common pathogenetic mechanisms, subsequently a positive association between the prevalence of AD and cataract, although other studies found no significant relationship between dementia and visual impairment including cataract in the elderly. Little is known about the association between Activity of Daily Living (ADL) and the combination of AD and cataract. To examine the association between ADL and the combination of AD and cataract, we performed a national survey in nursing care institutions in Japan, examining the decreased ADL in elderly with and without AD and cataract for 453 elderly aged 85.0 ± 8.13 years. The proportion, 43.5% of AD in subjects without cataract was significantly higher than that, 23.5% with cataract. Almost all ADL in AD with cataract was significantly lower than that without cataract, although all ADL in cataract

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patients receiving surgery in AD was significantly (all p < 0.002) higher than that in cataract patients without surgery in AD, as was confirmed by the multiple regression analysis incorporating into the relevant factors as independent variables. These results showed that ADL scores including cognitive functions decreased by cataract were increased by the surgery in the patients with AD. A significantly negative association between AD and cataract seen in our data, which was inconsistent with the previous result, might lead to the necessities of the clinical diagnosis for slight severity of cataract for patients with AD.

**Keywords**

Alzheimer’s Disease, Cataract, Elderly, Activity of Daily Living, Cognitive Function

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### 1. 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 the cognitive impairment of elderly in need of care, consequently interfering with daily tasks and decreasing Activities of Daily Living (ADL) in many ways [2]. Although epidemiologic studies have shown many risk factors such as occupational exposures (exposure to pesticides, electromagnetic fields, organic solvents and volatile anesthetics), pre-existing medical conditions (cerebrovascular disease, hypertension, diabetes, dyslipidemia, traumatic brain injury, depression and cancer) and lifestyle factors (smoking, consumptions of alcohol and coffee, body mass index, physical activity and cognitive activity) as non-genetic etiology of AD [3] [4], the accurate involvements of the occupational, pre-existing medical conditions and lifestyle factors in the development of AD remain to be elucidated.

It is estimated that there are 39 million blind persons and that cataract is the first cause of blindness worldwide [5]. Epidemiological studies have shown that smoking, diabetes mellitus, asthma, chronic bronchitis, cardiovascular disease, exposure to UVB light and corticosteroids use would increase cataract risk [6] [7]. Furthermore, over the last few years, clinical and epidemiological studies have suggested that dementia and visual impairment, in particular AD and cataract, may share common pathogenetic mechanisms [8]-[10]. Age-related vision impairment has been found to be closely associated with cognitive and behavioral manifestations [11]. Some epidemiologic studies also have demonstrated a positive association between AD and cataract [12]-[14]. Nevertheless, other studies found no significant relationship between cognitive impairment and vision loss including cataract in the elderly [15] [16]. In addition to the inconsistent studies on the association between AD and cataract, little is known about the association between Activity of Daily Living (ADL) and the combination of AD and cataract [17].

Both AD and cataract which increase with the aging deteriorate ADL and subsequently require nursing care [18]-[20]. To examine the association between ADL and the combination of AD and cataract, we performed a national survey in nursing care institutions in Japan, examining the decreased ADL in elderly with and without AD and cataract.

### 2. Methods

Fifty facilities were randomly selected out of 3410 nursing care institutions in Japan. Five resident patients and 5 daycare patients were sampled at random from each facility. A total of 500 patients without cerebral infarction, cerebral and subarachnoid hemorrhage were prospective subjects in this study. We mailed a questionnaire to a chief medical doctor in charge of each facility and subjects inquiring about ADL and medical conditions, including AD and cataract, as described below and obtained 453 responses (90.6%, 273 resident patients and 180 daycare patients). This survey was conducted from 2010 to 2012. The average age and SD of the 453 subjects were 85.0 and 8.13 years old. This research was conducted after obtaining approval from the ethical committee of the Japan Association of Geriatric Health Service Facilities.

The questionnaires regarding medical conditions in subjects were related to AD and cataract with and without surgery. To examine AD, the questions concerned neuropsychological examinations including the patients’
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medical history, neurological testing, Mini-Mental State Examination [21], and standard clinical evaluation including brain scanning. In the diagnostic process, we used DSM-IV-TR and ruled out other factors. The National Institute of Neurological and Communicative Disorders and Stroke-Alzheimer’s Disease and Related Disorders Association guidelines [22] were used for detection and diagnosis of either possible or probable AD, and stage of dementia according to the Clinical Dementia Rating (CDR) Scale [23]. We categorized AD all those with CDR scale scores of 0.5 or higher also meeting the criteria for probable or possible AD. Cataract was diagnosed according to the International Classification of Diseases (ICD), 10th Revision (H25) by family practice medical doctor. Diabetes mellitus was diagnosis according to Japan Diabetes Society clinical diagnosis guideline by family practice medical doctor [24]. The diagnoses of AD and cataract were performed within one year prior to the contact with nursing care institutions for elderly in this study. Our subjects included 125 patients with possible or probable AD, 253 and 108 patients with cataract treated with surgery and conservatively, respectively, and 52 elderly without AD or cataract.

Regarding ADL, the questions were developed on the basis of a standard described previously [19] [25] and [26]. They dealt with bed mobility, transfer and locomotion, cognitive function (orientation), cognitive function (communication), cognitive function (mental activity), eating (swallowing), eating (feeding), toilet use, and bathing for elderly requiring both residential nursing care systems and those for 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.

Table 1 shows ADL adjusted by age and sex according to AD and cataract. The two-way ANCOVA score by analysis of covariance (ANCOVA) with age and sex as covariants, in which age was assigned as a continuous variable and sex were dummy-coded with follows: male 0, female 1. Mean 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 cataract treated with or without surgery, we used two-way ANCOVA with age and sex as covariants. To confirm the results of ANCOVA, we performed multiple regression analysis for each ADL score incorporating into facility, sex, age, DM, cataract surgery, and Clinical Dementia Rating (CDR) scale as independent variables. The statistical software SPSS ver. 21 was used. P-values less than 5% with two tails were considered statistically significant.

3. Results

Table 1 shows numbers (proportion) and age of residential or day care users with and/or without cataract and AD in this study. The proportion, 43.5% of AD in subjects without cataract were significantly higher than that, 23.5% with cataract (Chi square test, p < 0.000). The proportions of woman using residential was significantly higher than that using day care. The proportions, 85.5% of women in subjects with AD were significantly higher than that, 66.1% without AD (p = 0.000). The proportions, 69.6% of women without cataract was significant lower than that, 79.5% with cataract, respectively (p = 0.042) (Table 1).

Table 2 shows that ADL adjusted by age and sex according to AD and cataract. The two-way ANCOVA showed statistically significant main effects of AD on ADL scores of cognitive functions including orientation,
The proportions of AD in subjects without and with cataract showed 43.5% and 23.5%, respectively (p < 0.000). The proportion of women using residential and day use showed 85.5% and 66.1%, respectively (p < 0.000). The proportion of women without and with cataract were 69.6% and 79.5%, respectively (p = 0.042).

Age and sex-adjusted ADL in 361 cataract subjects with and without AD and surgery against cataract are shown in Table 3. There were statistically significant main effects of AD on ADL scores of cognitive functions including orientation, communication and mental activity in addition to significant main effects of surgery on all ADL scores. Furthermore, there were significant interactive effects of AD and surgery on ADL of transfer and locomotion, cognitive functions including orientation and communication, and toilet use (Table 2).

Table 4 shows results on multiple regression analysis for each ADL score using facility, sex, age, DM, cataract surgery, and CDR score as independent variables in 356 subjects with cataract. Cataract surgery was statistically positively associated with ADL score of bed mobility, cognitive function (orientation), cognitive function (communication), cognitive function (mental activity) and toilet use. CDR was statistically negatively associated with all ADL scores but transfer and locomotion. There was no association of DM with any ADL score examined in this study (Table 4).

To summarize these results, almost all ADL in AD with cataract was significantly lower than those without cataract, although all ADL in cataract patients receiving surgery in AD was significantly higher than those in cataract patients without surgery in AD, as was confirmed by the multiple regression analysis incorporating into the relevant factors as independent variables.

4. Discussion

The present study showed a higher prevalence of AD in subjects without cataract than those with cataract, demonstrating a significant negative association between AD and cataract. On the contrary, recent population studies have demonstrated a significant positive relationship between AD and cataract. Mandas et al. [27], who analyzed data from 1168 subjects aged 65 years or older admitted between 2006 and 2013 to the outpatients clinic of the geriatric care unit at the University of Cagliari and to the one at Santissima Trinità Hospital in Cagliari, showed a relationships between mild cognitive impairment, various forms of age-related dementia (Alzheimer’s
Table 2. Activity of daily living adjusted by age and sex according to cataract and AD.

| AD | CA | Number | Proportion of woman (%) | Age (years; mean ± SD) | Activity of daily living (age-and sex adjusted score; means ± SD) |
|----|----|--------|-------------------------|-----------------------|---------------------------------------------------------------|
|    |    |        |                         |                       | Bed mobility | Transfer and locomotion | Cognitive function (orientation) | Cognitive function (communication) | Cognitive function (mental activity) | Eating (dysphagia) | Eating (feeding) | Toilet use | Bathing |
| (-) | 52 | 32 (61.5) | 81.94 ± 10.61 | 4.07 ± 1.33 | 2.33 ± 0.90 | 4.25 ± 1.06 | 4.23 ± 1.13 | 4.29 ± 1.13 | 4.45 ± 1.08 | 4.47 ± 1.06 | 3.94 ± 1.26 | 3.27 ± 0.87 |
| (+) | 276 | 216 (78.3) | 85.38 ± 7.93 | 3.70 ± 1.32 | 2.37 ± 0.89 | 3.98 ± 1.05 | 3.88 ± 1.12 | 3.94 ± 1.12 | 4.29 ± 1.07 | 4.35 ± 1.05 | 3.73 ± 1.25 | 3.12 ± 0.86 |
| (+) | 40 | 32 (80.0) | 84.05 ± 6.27 | 4.24 ± 1.32 | 2.73 ± 0.90 | 3.48 ± 1.05 | 3.58 ± 1.12 | 3.30 ± 1.12 | 4.46 ± 1.07 | 4.59 ± 1.05 | 3.94 ± 1.25 | 3.05 ± 0.86 |
| (+) | 85 | 71 (83.5) | 86.12 ± 7.42 | 3.37 ± 1.32 | 2.13 ± 0.90 | 2.41 ± 1.05 | 2.46 ± 1.12 | 2.45 ± 1.13 | 3.74 ± 1.07 | 3.92 ± 1.05 | 3.13 ± 1.25 | 2.83 ± 0.86 |

Statistics (p value)

- Sex: 0.412, 0.015, 0.065, 0.400, 0.497, 0.776, 0.968, 0.379, 0.084
- Age: 0.019, 0.000, 0.001, 0.012, 0.001, 0.004, 0.125, 0.012, 0.177
- AD: 0.612, 0.450, 0.000, 0.000, 0.000, 0.005, 0.025, 0.048, 0.017
- Cataract: 0.000, 0.012, 0.000, 0.000, 0.000, 0.000, 0.000, 0.000, 0.001, 0.084
- AD × cataract: 0.132, 0.003, 0.002, 0.005, 0.073, 0.163, 0.269, 0.049, 0.745

Statistics was performed by two-way (cataract and AD) ANCOVA with age and sex as covariants.

Table 3. Age and sex-adjusted activities of daily living in 361 cataract subjects with and/or AD and surgery against cataract.

| AD | Surgery | Number | Proportion of woman (%) | Age (years; mean ± SD) | Activity of daily living (age-and sex adjusted score; means ± SD) |
|----|---------|--------|-------------------------|-----------------------|---------------------------------------------------------------|
|    |         |        |                         |                       | Bed mobility | Transfer and locomotion | Cognitive function (orientation) | Cognitive function (communication) | Cognitive function (mental activity) | Eating (dysphagia) | Eating (feeding) | Toilet use | Bathing |
| (-) | 48 | 36 (75.0) | 82.41 ± 10.37 | 3.13 ± 1.36 | 2.18 ± 0.89 | 3.29 ± 1.04 | 3.16 ± 1.12 | 3.28 ± 1.12 | 3.74 ± 1.09 | 3.82 ± 1.11 | 3.16 ± 1.27 | 2.85 ± 0.86 |
| (+) | 228 | 180 (78.4) | 85.91 ± 7.28 | 3.80 ± 1.35 | 2.39 ± 0.88 | 4.10 ± 1.04 | 4.01 ± 1.11 | 4.06 ± 1.11 | 4.40 ± 1.08 | 4.45 ± 1.11 | 3.83 ± 1.26 | 3.16 ± 0.85 |
| (-) | 52 | 52 (80.7) | 85.29 ± 7.25 | 3.15 ± 1.35 | 1.89 ± 0.88 | 1.99 ± 1.04 | 2.07 ± 1.11 | 2.10 ± 1.11 | 3.56 ± 1.09 | 3.81 ± 1.11 | 2.86 ± 1.27 | 2.68 ± 0.86 |
| (+) | 25 | 19 (76.0) | 86.12 ± 6.64 | 3.85 ± 1.35 | 2.65 ± 0.88 | 3.37 ± 1.04 | 3.37 ± 1.11 | 3.25 ± 1.11 | 4.14 ± 1.08 | 4.17 ± 1.11 | 3.73 ± 1.26 | 3.17 ± 0.86 |

Statistics (p value)

- Sex: 0.833, 0.114, 0.325, 0.325, 0.325, 0.906, 0.856, 0.856, 0.341
- Age: 0.005, 0.000, 0.001, 0.001, 0.001, 0.000, 0.066, 0.066, 0.009
- AD: 0.853, 0.890, 0.000, 0.000, 0.000, 0.000, 0.162, 0.353, 0.353, 0.263
- Cataract surgery: 0.000, 0.000, 0.000, 0.000, 0.000, 0.000, 0.002, 0.002, 0.000
- AD × surgery: 0.925, 0.034, 0.056, 0.056, 0.056, 0.775, 0.387, 0.387, 0.591

Statistics was performed by two-way (AD and cataract surgery) ANCOVA with age and sex as covariants.
| Bed mobility   | Transfer and locomotion | Cognitive function (orientation) | Cognitive function (communication) | Cognitive function (mental activity) | Eating (dysphagia) | Eating (feeding) | Toilet use | Bathing   |
|---------------|-------------------------|---------------------------------|------------------------------------|--------------------------------------|--------------------|-----------------|-------------|-----------|
| **Status**    |                         |                                 |                                    |                                      |                    |                 |             |           |
| **0.284***    | 0.277***                | 0.063                           | 0.071                              | 0.091*                               | 0.294***           | 0.166**         | 0.292***    | 0.252***   |
| **0.513 to 1.103** | **0.33 to 0.719**     | **-0.49 to 0.38**              | **-0.038 to 0.429**               | **0.015 to 0.483**                   | **0.458 to 0.925** | **0.142 to 0.632** | **0.523 to 1.066** | **0.265 to 0.644** |
| **0.048**     | **-0.034**              | **-0.029**                      | **-0.006**                         | **0.015**                            | **0.0068**         | **0.049**        | **0.008**   | **-0.007** |
| **-0.194 to 0.521** | **-0.312 to 0.159**   | **-0.360 to 0.168**             | **-0.303 to 0.263**               | **-0.234 to 0.330**                  | **-0.091 to 0.473** | **-0.158 to 0.434** | **-0.301 to 0.356** | **-0.244 to 0.213** |
| **-0.119**    | **-0.15**               | **-0.116**                      | **-0.078**                         | **-0.106**                           | **-0.138**         | **-0.057**       | **-0.0093** | **-0.075** |
| **-0.040 to 0.002** | **-0.030 to -0.050** | **-0.033 to -0.006**             | **-0.028 to 0.001**               | **-0.330 to -0.003**                 | **-0.350 to -0.005** | **-0.024 to 0.007** | **-0.035 to 0.002** | **-0.020 to 0.004** |
| **Diabetes mellitus** | **-0.061**               | **0.010**                       | **0.021**                          | **0.039**                            | **0.035**          | **0.14**         | **-0.011** | **0.013**  |
| **-0.529 to 0.126** | **-0.008 to 0.045**    | **-0.177 to 0.308**             | **-0.136 to 0.383**               | **-0.148 to 0.369**                  | **-0.222 to 0.297** | **-0.300 to 0.544** | **-0.356 to 0.246** | **-0.183 to 0.238** |
| **0.133**     | **0.109**               | **0.233***                      | **0.215***                         | **0.189***                           | **0.159**          | **0.104**        | **0.152**   | **0.11**   |
| **0.055 to 0.746** | **0.008 to 0.448**    | **0.410 to 0.923**              | **0.350 to 0.900**                | **0.276 to 0.824**                   | **0.122 to 0.668** | **-0.030 to 0.544** | **0.121 to 0.763** | **-0.13 to 0.434** |
| **-0.096**    | **-0.088**              | **-0.504***                     | **-0.475***                        | **-0.494***                          | **-0.169**         | **-0.228***      | **-0.178**  | **-0.121** |
| **-0.322 to 0.031** | **-0.207 to 0.028**   | **-0.854 to -0.592**            | **-0.831 to -0.551**              | **-0.861 to -0.582**                 | **-0.352 to -0.073** | **-0.431 to -0.138** | **-0.423 to -0.095** | **-0.231 to -0.002** |
| **R²**        | **0.157**               | **0.177**                       | **0.489**                          | **0.427**                            | **0.439**          | **0.231**        | **0.141**   | **0.222**  |
| **Adjusted R²** | **0.142**               | **0.163**                       | **0.48**                           | **0.418**                            | **0.429**          | **0.218**        | **0.126**   | **0.209**  |
| **0.130**     |                         |                                 |                                    |                                      |                    |                 |             |           |

Statistical significant: *p < 0.05, **p < 0.01, ***p < 0.001. Regarding the effect of surgery against cataract on the ADL of several cognitive functions on, which effect is considered to be most typically seen in this study, cognitive function scores in the AD patients with cataract were decreased as compared to the scores in those without cataract, whereas the scores in the AD patients with cataract surgery were increased as compared to the scores in those with surgery.
Our findings also do not support the theory of a combined neurodegenerative mechanism in cataract and cognitive impairment, as we found a negative association between cataract (including treated and untreated) and AD. Thus, our results agree well with Newcastle 85+ Cohort study [12], which was drawn from elderly subjects with almost the similar age as the mean age of 85.0 years in this study. Our study relied on family practice medical doctor for diagnoses of cataract in case of subjects without surgery. The association between AD and cataract for elderly subjects with very higher age might be different from that for relatively younger elderly. Very old elderly with AD could not be diagnosed to be cataract because they would not complain of visual loss even if they had cataract actually. This seems to be the reason why we had a negative association between AD and cataract in this study. As the diagnosis for cataract was performed based on examination for apparent changes of lens in this study, the cataract might be diagnosed only in case of cataract with the severe degree. In fact, the overall prevalence of cataract in this study showed 79.7%, which is considered to be lower than 90% - 95% for subjects with the range of almost the same age [14]. Such a diagnostic bias accompanying AD, which has been pointed out by some studies [12] [14], might account for the negative association between AD and cataract. In fact, we recognized a very lower ratio of receiving cataract surgery in patients with AD as compared to those without AD. The lower ratio of receiving surgery also seems to be very closely associated with lower prevalence of cataract in case of patients with AD. Epidemiology by Rogers and Langa [28] demonstrates that 77.9% of cataract patients with normal cognition had received at least one previous eye procedure as compared with 51.7% of cataract actually. This seems to be the reason why we had a negative association between AD and cataract surgery in AD patients, our results are significantly lower rate of dementia in the cataract surgery group (hazard ratio 0.77%, 95% confidence interval 0.75 - 0.79, p < 0.001), concluded that patients undergoing cataract surgery were associated with a reduced risk of subsequent dementia compared with those without cataract surgery. A retrospective cohort study by using the database of the Taiwan National Health Insurance Program from 1999 to 2004 including 19,954 subjects aged 65 - 84 with newly diagnosed cataract showed that the adjusted HR of Alzheimer’s disease was 1.43 (95% CI 1.13, 1.82) for the cataract group, compared to the non- cataract group after adjustment for potential confounders, sex, age, diabetes mellitus, head injury and hypertension [14]. In contrast with these reports, cross-sectional analysis of data from the Newcastle 85+ Cohort study [12], comparing among no cataract, cataract diagnosis with and without surgery, found a lower degree of cognitive impairment in patients with cataracts, but no difference in cognitive impairment between cataract patients with and without surgery. The inverse results might be considered to be related to an enhanced help-seeking behavior in people with diagnosed cataract.

It is noteworthy that ADL regarding cognitive functions with cataract surgery in AD patients was significantly higher than those without surgery, as was shown in the Table 3. Simultaneously, our results suggests that that cataract surgery improves cognitive function scores in the AD patients decreased by cataract to almost the same level as those in the subjects without AD. A number of studies examining the effect of cataract surgery on cognitive performance suggest that cognitive performance can be improved with cataract surgery [29] [30]. A prospective observational study performed by Tamura et al. for 20 patients with cognitive impairment showed that the grade of cognitive impairment in 12 patients (60%) were made better after cataract surgery, concluding cataract surgery improved cognitive impairment in elderly Japanese patients [29]. However, a randomized study using a waiting list control group did not confirm this [31] [32]. Anstey et al. [31], who examined cataract surgery improve neuropsychological performance in healthy elderly using a randomized clinical trial of cataract surgery, showed no cognitive benefits of cataract surgery in cognitively normal adults, concluding that visual improvement following cataract surgery is not strongly associated with an improvement in neuropsychological test performance in elderly without cognitive impairment. Although there seems to be inconsistent results for the association between ADL regarding cognitive functions and cataract surgery in AD patients, our results are much different from other studies including Anstey et al. [31] [32] in the lights of age of the subjects and severity of AD. Furthermore, taken together with our results showing the multiple regression analysis controlling other relevant factors including AD severity confirmed our results, cataract surgery for highly aged elderly with AD seems to improve ADL regarding cognitive functions. The causal relationship between cataract surgery and
the improvement of ADL in highly aged elderly with AD will be confirmed by the future follow-up study. In additions, we have some limitations regarding the outcome of cataract surgery depending on various factors; not assessing the association of cataract type with incidence of cognitive, not focusing on the first or second surgery and sequential surgery, not referring to one or both eyes, not examining analyzing the postoperative duration for each individual patient [30] [33].

5. Conclusion
In conclusion, our results showed that ADL scores including cognitive functions decreased by cataract were increased by the surgery in the patients with AD. Therefore, cataract surgery seems to improve ADL including cognitive functions for highly aged elderly with AD. A significantly negative association between AD and cataract seen in our data, which was inconsistent with results from other studies, might lead to the necessities of the clinical diagnosis for slight severity of cataract for patients for AD.

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Conflict of Interest Statement
None of the authors have any proprietary interests or conflicts of interest related to this submission.

Limitations
There are some limitations in our study. For example, our study population is small. Furthermore, we must assess the association of ADL by combination AD and cataract in long time. Also, we have conducted a survey using a questionnaire by mail. So, there may be individual differences by investigated staff. We cannot assess the association of cataract types with incidence of pathological change, not focusing on the detail of surgery. Further studies are needed to clarify the improvement of cognitive functions by cataract surgery in the patients with AD.

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