

SUPPLEMENTARY TABLES

Supplementary Table 1A. Distribution of MCI accompanied by clinical data in each AT(N) biomarker category.

| AT(N) biomarker categories | Group | Normal | p value | AD continuum | p value | Non-AD pathologic change | p value | Number of missing data | | |
|----------------------------|-------|---------------|---------|---------------|---------|--------------------------|---------|------------------------|--------------|--------------------------|
| | | | | | | | | Normal | AD continuum | Non-AD pathologic change |
| Number in each group | HIGH | n = 30 | | n = 15 | | n = 4 | | | | |
| | LOW | n = 32 | | n = 4 | | n = 4 | | | | |
| Age at diagnosis | HIGH | 79.6 (± 3.8) | | 81.3 (± 5.4) | | 80.8 (± 3.8) | | 0 | 0 | 0 |
| | LOW | 64.7 (± 7.2) | | 64.3 (± 9.4) | | 67.0 (± 3.8) | | 0 | 0 | 0 |
| Disease duration (m) | HIGH | 21.1 (± 14.1) | | 15.2 (± 13.1) | | 6.5 (± 3.5) | | 0 | 0 | 0 |
| | LOW | 23.6 (± 14.8) | | 13.3 (± 13.2) | | 7.8 (± 6.0) | | 0 | 0 | 0 |
| Education (y) | HIGH | 11.6 (± 2.7) | 0.001** | 12.1 (± 2.7) | 0.48 | 11.5 (± 4.5) | 0.81 | 7 | 4 | 2 |
| | LOW | 14.5 (± 2.7) | | 13.3 (± 1.9) | | 13.0 (± 5.3) | | 11 | 1 | 0 |
| MMSE scores | HIGH | 26.8 (± 2.0) | 0.003** | 27.6 (± 1.9) | 0.86 | 27.7 (± 1.2) | 0.96 | 4 | 3 | 1 |
| | LOW | 28.4 (± 1.7) | | 27.3 (± 1.7) | | 27.8 (± 2.3) | | 7 | 1 | 0 |
| MCI, n (%) | HIGH | 3 (10.0%) | | 1 (6.7%) | | 0 | | 0 | 0 | 0 |
| | LOW | 1 (3.1%) | | 0 | | 1 (25.0%) | | 0 | 0 | 0 |
| ApoE ε4, n (%) | HIGH | 4 (13.8%) | | 6 (42.9%) | | 0 | | 1 | 1 | 1 |
| | LOW | 4 (14.2%) | | 1 (25.0%) | | 0 | | 4 | 0 | 1 |

Continuous variables are presented as the mean (± standard deviation). AD continuum = Alzheimer's continuum, Non-AD pathologic change = Non-Alzheimer's pathologic change, HIGH = a group of patients aged ≥ 73 at diagnosis, LOW = a group of patients aged < 73 at diagnosis, MMSE = mini-mental state examination, MCI = mild cognitive impairment, ApoE = Apolipoprotein E, Aβ42 = amyloid-beta 42, p-tau = phosphorylated tau, t-tau = total tau, ** = $p < 0.01$. *P* values represent the result of Welch's t test comparing the HIGH group vs. the LOW group within each AT(N) category.

Supplementary Table 1B. Contingency table for Fisher's exact test showing the numbers of MCI in each AT(N) category within the HIGH group.

| | Normal | AD continuum | Non-AD pathologic change | Total |
|------------------|--------|--------------|--------------------------|-------|
| MCI | 3 | 1 | 0 | 4 |
| Normal cognition | 27 | 14 | 4 | 45 |
| Total | 30 | 15 | 4 | 49 |

MCI = mild cognitive impairment, HIGH group = a group of patients aged ≥ 73 at diagnosis, AD continuum = Alzheimer's continuum, Non-AD pathologic change = Non-Alzheimer's pathologic change.

Supplementary Table 1C. Contingency table for Fisher's exact test showing the numbers of MCI in each AT(N) category within the LOW group.

| | Normal | AD continuum | Non-AD pathologic change | Total |
|------------------|--------|--------------|--------------------------|-------|
| MCI | 1 | 0 | 1 | 2 |
| Normal cognition | 31 | 4 | 3 | 38 |
| Total | 32 | 4 | 4 | 40 |

MCI = mild cognitive impairment, LOW group = a group of patients aged < 73 at diagnosis, AD continuum = Alzheimer's continuum, Non-AD pathologic change = Non-Alzheimer's pathologic change.

Supplementary Table 2. Cutoff levels of CSF amyloid-beta 42 in prior studies using INNOTEST® and our study.

| No. | CSF A β 42 cutoff (pg/mL) | Study | References |
|-----|---------------------------------|---------------|------------|
| 1 | 333 | J-ADNI | 26 |
| 2 | 380 | Krakow | 25 |
| 3 | 430 | Brussels | 25 |
| 4 | 445 | Lisbon | 25 |
| 5 | 450 | Bremen | 25 |
| 6 | 450 | Gothenburg | 25 |
| 7 | 450 | Ljubljana | 25 |
| 8 | 450 | Mannheim | 25 |
| 9 | 450 | Thessaloniki | 25 |
| 10 | 459 | St. Louis | 25 |
| 11 | 482 | Mattsson | 25 |
| 12 | 490 | Athens | 25 |
| 13 | 500 | Barcelona CUH | 25 |
| 14 | 500 | Brescia | 25 |
| 15 | 500 | Nijmegen | 25 |
| 16 | 500 | | This study |
| 17 | 542 | Coimbra | 25 |
| 18 | 550 | Amsterdam | 25 |
| 19 | 550 | Barcelona HSP | 25 |
| 20 | 550 | DESCRIPA | 25 |
| 21 | 550 | Lausanne | 25 |
| 22 | 550 | LeARN | 25 |
| 23 | 550 | Lorenskog | 25 |
| 24 | 550 | Madrid | 25 |
| 25 | 550 | Perugia | 25 |
| 26 | 550 | Stockholm | 25 |
| 27 | 600 | DCN | 25 |
| 28 | 600 | Scinawa | 25 |
| 29 | 610 | Warsaw | 25 |
| 30 | 638.5 | Antwerp | 25 |
| 31 | 662.65 | Chandigarh | 25 |

Median (IQR) cutoff level: 530.3 (452.3-550.0)

Of the 31 cutoff levels, 30 were taken from prior studies cited in Table 3 (29 from Reference [25] and one from Reference [26]), and the remaining one was our own. These levels ranged from 333 to 662.65 pg/mL. CSF = cerebrospinal fluid, No.= Number, A β 42 = amyloid-beta 42, J-ADNI = Japanese Alzheimer's Disease Neuroimaging Initiative, IQR = interquartile range. See the original paper about study names.

Supplementary Table 3A. Comparison of the prevalence of amyloid positivity between the two groups at a cutoff level of 400 pg/mL.

| | HIGH group (N = 49) | LOW group (N = 40) | Odds ratio | 95% CI | p value |
|---------------------------|-------------------------------|------------------------------|-------------------|---------------|----------------|
| Amyloid positivity, n (%) | 11 (22.4%) | 1 (2.5%) | 11.1 | 1.5–496.9 | 0.01** |

HIGH group = patients aged ≥ 73 at diagnosis, LOW group = patients aged < 73 at diagnosis, CI = confidence interval, ** = $p < 0.01$. *P* value represents the result of Fisher's exact test.

Supplementary Table 3B. Comparison of the prevalence of amyloid positivity between the two groups at a cutoff level of 600 pg/mL.

| | HIGH group (N = 49) | LOW group (N = 40) | Odds ratio | 95% CI | p value |
|---------------------------|-------------------------------|------------------------------|-------------------|---------------|----------------|
| Amyloid positivity, n (%) | 19 (38.8%) | 9 (22.5%) | 2.2 | 0.8–6.3 | 0.11 |

HIGH group = patients aged ≥ 73 at diagnosis, LOW group = patients aged < 73 at diagnosis, CI = confidence interval. *P* value represents the result of Fisher's exact test.