

*Supplementary Material*

SUPPLEMENTARY TABLE 1. *Demographic characteristics of included subjects*

| Group                    |                       | PIB+(n = 15)  | PIB-(n = 15)  |
|--------------------------|-----------------------|---------------|---------------|
| Sex                      | Male                  | 11            | 10            |
|                          | Female                | 4             | 5             |
| Diagnosis                | AD                    | 8             | 0             |
|                          | MCI with AD pathology | 7             | 0             |
| MCI without AD pathology | 0                     | 4             |               |
|                          | HC                    | 0             | 11            |
| Age(y)                   |                       | 68.07 ± 9.80  | 68.13 ± 3.90  |
| Weight (kg)              |                       | 77.67 ± 10.18 | 78.53 ± 13.43 |
| MMSE score               |                       | 25.53 ± 2.69  | 28.20 ± 2.43  |

SUPPLEMENTARY TABLE 2. *Description of the VOIs used for estimating  $k'_2$  values*

| Name                     | Origin   |
|--------------------------|--|
| Whole Brain              | All voxels on the image  |
| Cerebrum + Brainstem     | All voxels of cortical and white matter tissue from the cerebrum and the brainstem   |
| Gray Matter              | All cortical tissue from the cerebrum  |
| Frontal Lobe             | Superior frontal gyrus, middle frontal gyrus, inferior frontal gyrus, precentral gyrus, straight gyrus, anterior orbital gyrus, lateral orbital gyrus, medial orbital gyrus, posterior orbital gyrus, subcallosal area, subgenual frontal cortex, and pre-subgenual frontal cortex           |
| Parietal Lobe            | Postcentral gyrus, superior parietal gyrus, and inferolateral remainder of parietal lobe   |
| Temporal Lobe            | Hippocampus, amygdala, anterior temporal lobe lateral part, anterior temporal lobe medial part, parahippocampal and ambient gyri, superior temporal gyrus anterior part, superior temporal gyrus posterior part, middle and inferior temporal gyrus, fusiform gyrus, posterior temporal lobe |
| White Matter             | All white matter tissue from the cerebrum  |
| Brainstem                | VOI from the Hammer's atlas  |
| White Matter + Brainstem | Union of white matter tissue from the cerebrum and brainstem   |
| Threshold                | Drawn by fixing a lower limit on the $BP_{ND}^{SRTM}$ parametric map and taking all voxels that were above the limit (threshold chosen for this study: 0.01, 0.05, and 0.1)  |
| SPM                      | Voxels that presented statistically significant differences between PIB+ and PIB- groups using a two-sample t-test in SPM12  |

SUPPLEMENTARY TABLE 3. *Mean and standard deviation from estimated  $k'_2$  values per group, and p-values of difference between groups*

| <b>Method</b>            | <b>PIB+ (Mean ± SD)</b> | <b>PIB- (Mean ± SD)</b> | <b>p-value</b> |
|--------------------------|-------------------------|-------------------------|----------------|
| Whole Brain              | 0.05 ± 0.01             | 0.06 ± 0.01             | 0.03*          |
| Cerebrum + Brainstem     | 0.06 ± 0.01             | 0.07 ± 0.01             | 0.04*          |
| Gray Matter              | 0.05 ± 0.01             | 0.08 ± 0.02             | <0.01*         |
| Frontal Lobe             | 0.05 ± 0.01             | 0.07 ± 0.02             | <0.01*         |
| Parietal Lobe            | 0.05 ± 0.01             | 0.07 ± 0.02             | 0.01*          |
| Temporal Lobe            | 0.06 ± 0.01             | 0.10 ± 0.02             | <0.01*         |
| White Matter             | 0.07 ± 0.01             | 0.06 ± 0.01             | 0.16           |
| Brainstem                | 0.08 ± 0.01             | 0.08 ± 0.01             | 0.60           |
| White Matter + Brainstem | 0.07 ± 0.01             | 0.06 ± 0.01             | 0.20           |
| Threshold 0.01           | 0.05 ± 0.01             | 0.06 ± 0.01             | 0.11           |
| Threshold 0.05           | 0.05 ± 0.01             | 0.06 ± 0.01             | 0.27           |
| Threshold 0.1            | 0.05 ± 0.01             | 0.06 ± 0.01             | 0.71           |
| SPM                      | 0.05 ± 0.01             | 0.08 ± 0.02             | <0.01*         |

\* $p < 0.05$

SUPPLEMENTARY TABLE 4. *Results from the paired t-test comparing the differences between Whole Brain method and other methods for PIB+ and PIB- subjects on the  $k'_2$  estimation*

| <b>Method</b>            | <b>Group</b> | <b>Mean of the differences</b> | <b>Confidence Interval</b> | <b>p-value</b> |
|--------------------------|--------------|--------------------------------|----------------------------|----------------|
| Cerebrum + Brainstem     | PIB+         | -0.006                         | [-0.008, -0.004]           | <0.001*        |
|                          | PIB-         | -0.008                         | [-0.011, -0.005]           | <0.001*        |
| Gray Matter              | PIB+         | 0.001                          | [-0.001, 0.003]            | 0.643          |
|                          | PIB-         | -0.018                         | [-0.025, -0.011]           | <0.001*        |
| Frontal Lobe             | PIB+         | 0.006                          | [0.003, 0.010]             | 0.001*         |
|                          | PIB-         | -0.009                         | [-0.019, 0.000]            | 0.053          |
| Parietal Lobe            | PIB+         | 0.003                          | [-0.001, 0.007]            | 0.157          |
|                          | PIB-         | -0.009                         | [-0.017, 0.000]            | 0.049*         |
| Temporal Lobe            | PIB+         | -0.010                         | [-0.016, -0.005]           | 0.001*         |
|                          | PIB-         | -0.037                         | [-0.045, -0.029]           | <0.001*        |
| White Matter             | PIB+         | -0.015                         | [-0.018, -0.013]           | <0.001*        |
|                          | PIB-         | -0.003                         | [-0.006, -0.001]           | 0.013*         |
| Brainstem                | PIB+         | -0.025                         | [-0.028, -0.022]           | <0.001*        |
|                          | PIB-         | -0.019                         | [-0.023, -0.015]           | <0.001*        |
| White Matter + Brainstem | PIB+         | -0.016                         | [-0.018, -0.014]           | <0.001*        |
|                          | PIB-         | -0.003                         | [-0.007, 0.000]            | 0.045*         |
| Threshold 0.01           | PIB+         | -0.001                         | [-0.003, 0.001]            | 0.189          |
|                          | PIB-         | 0.001                          | [-0.002, 0.003]            | 0.622          |
| Threshold 0.05           | PIB+         | -0.001                         | [-0.003, 0.001]            | 0.153          |
|                          | PIB-         | 0.002                          | [0.000, 0.005]             | 0.071          |
| Threshold 0.1            | PIB+         | -0.001                         | [-0.003, 0.001]            | 0.148          |
|                          | PIB-         | 0.005                          | [0.001, 0.008]             | 0.013*         |
| SPM                      | PIB+         | 0.0014                         | [-0.001, 0.004]            | 0.286          |
|                          | PIB-         | -0.016                         | [-0.023, -0.009]           | <0.001*        |

\* $p < 0.05$

SUPPLEMENTARY FIGURE 1. Plot of the relative changes in BPND SRTM2 with relative increment of  $k_2'$ . In the x-axis is the fixed value of  $k_2'$  (which varied from 0.005 to 0.09 with a step of 0.005), and in the y-axis is the BPND SRTM2 of the fixed value of a specific brain region. The dashed line represents a possible lower threshold for  $k_2'$  estimation. Black dots represent data from the Superior Parietal Gyrus left region (a region with high PIB binding), in dark gray, the Inferior Frontal Gyrus right (a region with medium binding), and in light gray, the Lateral Remainder of Occipital Lobe right (a region with low binding of PIB).

