### S3 Table. Risk of Parkinson’s disease in individuals with PPI exposure stratified by various clinical variables

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| --- | --- | --- | --- | --- |
| **Variable** | **N** | **Including a lag window of 1 year** | **Including a lag window of 2 years** | **Including a lag window of 3 years** |
| **Adjusted OR (95% CI)†** |  | **Adjusted OR (95% CI)†** |  | **Adjusted OR (95% CI)†** |
| Age group | 　 | 　 | 　 | 　 | 　 | 　 |
| <50 years | 5,550 | 1.11 (0.93–1.31) | 　 | 1.04 (0.87–1.25) | 　 | 0.98 (0.79–1.20) |
| ≥50 years | 23,955 | 1.10 (1.07–1.13)\*\*\* | 　 | 1.11 (1.07–1.14)\*\*\* | 　 | 1.10 (1.06–1.13)\*\*\* |
| Sex | 　 | 　 | 　 | 　 | 　 | 　 |
| Male | 69,799 | 1.06 (1.01–1.11)\* | 　 | 1.07 (1.02–1.12)\*\* | 　 | 1.04 (0.99–1.10) |
| Female | 86,830 | 1.13 (1.09–1.18)\*\*\* | 　 | 1.14 (1.09–1.18)\*\*\* | 　 | 1.13 (1.09–1.18)\*\*\* |
| BMI group (kg/m2) | 　 | 　 | 　 | 　 | 　 | 　 |
| Underweight (<18.5) | 4,150 | 1.31 (1.05–1.63)\* | 　 | 1.29 (1.03–1.62)\* | 　 | 1.30 (1.02–1.67)\* |
| Normal (18.5–<25.0) | 9,569 | 1.06 (1.01–1.11)\* | 　 | 1.08 (1.03–1.13)\*\* | 　 | 1.07 (1.02–1.12)\*\* |
| Overweight (25.0–<30.0) | 50,090 | 1.16 (1.09–1.23)\*\*\* | 　 | 1.15 (1.09–1.23)\*\*\* | 　 | 1.13 (1.06–1.21)\*\*\* |
| Obesity (≥30.0) | 6,720 | 1.49 (1.25–1.79)\*\*\* | 　 | 1.36 (1.14–1.63)\*\* | 　 | 1.46 (1.20–1.77)\*\*\* |
| HTN | 　 | 　 | 　 | 　 | 　 | 　 |
| No | 155,630 | 1.09 (1.06–1.13)\*\*\* | 　 | 1.10 (1.07–1.14)\*\*\* | 　 | 1.09 (1.05–1.13)\*\*\* |
| Yes | 1,000 | 16.97 (7.76–37.11)\*\*\* | 　 | 5.02 (2.62–9.62)\*\*\* | 　 | 7.68 (3.54–16.65)\*\*\* |
| DM | 　 | 　 | 　 | 　 | 　 | 　 |
| No | 156,290 | 1.10 (1.07–1.13)\*\*\* | 　 | 1.10 (1.07–1.14)\*\*\* | 　 | 1.09 (1.06–1.13)\*\*\* |
| Yes | 340 | 3.78531e+19 (0.00–Inf) | 　 | 3.08 (0.25–38.34) | 　 | 5.11 (0.32–82.72) |
| CCI score group | 　 | 　 | 　 | 　 | 　 | 　 |
| 0 | 65,350 | 1.05 (0.98–1.13) | 　 | 1.05 (0.97–1.13) | 　 | 1.03 (0.95–1.13) |
| 1 | 46,386 | 1.03 (0.95–1.12) | 　 | 1.03 (0.94–1.12) | 　 | 1.01 (0.92–1.10) |
| 2 | 24,364 | 1.06 (0.92–1.21) | 　 | 1.05 (0.91–1.20) | 　 | 1.14 (0.99–1.33) |
| ≥3 | 20,529 | 1.29 (1.10–1.52)\*\* | 　 | 1.31 (1.12–1.53)\*\* | 　 | 1.38 (1.17–1.61)\*\*\* |
| Smoking status | 　 | 　 | 　 | 　 | 　 | 　 |
| Never | 111,611 | 1.11 (1.07–1.15)\*\*\* | 　 | 1.11 (1.07–1.16)\*\*\* | 　 | 1.12 (1.07–1.16)\*\*\* |
| Ex/Current | 62,869 | 1.22 (1.04–1.44)\*\*\* | 　 | 1.15 (0.97–1.36) | 　 | 1.08 (0.91–1.30) |
| Alcohol consumption (times/week) | 　 | 　 | 　 | 　 | 　 |
| None | 114,274 | 1.12 (1.08–1.17)\*\*\* | 　 | 1.14 (1.10–1.18)\*\*\* | 　 | 1.13 (1.08–1.17)\*\*\* |
| 1–2 | 25,554 | 1.01 (0.87–1.17) | 　 | 0.97 (0.84–1.13) | 　 | 0.93 (0.79–1.09) |
| ≥3 | 16,801 | 1.17 (0.93–1.46) | 　 | 1.04 (0.83–1.30) | 　 | 0.94 (0.73–1.20) |

Abbreviations: BMI, body mass index; CCI, Charlson Comorbidity Index; CI, confidence interval; DM, diabetes mellitus; HTN, hypertension; OR, odds ratio; PPI, proton pump inhibitor
The odds ratio (OR) and 95% confidence intervals (CIs) were computed from the conditional logistic regression model.
† Adjusted for the calendar year of the index date, CCI score, smoking status, and alcohol consumption.
\*p<0.05; \*\*p<0.01; \*\*\*p<0.001