## COMT Diplotype Amplifies Effect of Stress on Risk of Temporomandibular Pain

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## **Appendix**

**Appendix Table.** Cox Regression Models of Time-Constant and Time-Varying Influences of Psychological Stress on Hazard of First-Onset TMD: OPPERA Prospective Cohort Study.

| Model | Description  | No. of<br>Subjects | df    | $\chi^2$ | Interaction<br>P Value | Parameter  | Hazard Ratio<br>(95% CL) |
|-------|--|--------------------|-------|----------|------------------------|--|--------------------------|
| AI    | Model 6 from Table 2,                                      | 2,186              | 103.5 | 19       | 0.24                   | Time-constant baseline PSS score (per 6.4 units)   | 1.70 (1.44, 2.02)        |
|       | with interaction of<br>time-varying stress<br>and COMT LPS |                    |       |          |                        | Stratum with 0 copies of LPS: Time-varying average quarterly PSS score minus baseline (per 6.4 units)      | 2.38 (1.59, 3.58)        |
|       |  |                    |       |          |                        | Stratum with 1 copy of LPS: Time-varying average quarterly PSS score minus baseline (per 6.4 units)        | 1.54 (1.07, 2.22)        |
|       |  |                    |       |          |                        | Stratum with 2 copies of LPS: Time-varying average quarterly PSS score minus baseline (per 6.4 units)      | 1.80 (0.91, 3.56)        |
| A2    | Model 6 from Table 2,                                      | 2186               | 99.8  | 13       | 0.22                   | Time-constant baseline PSS score (per 6.4 units)   | 1.72 (1.46, 2.04)        |
|       | with interaction of<br>time-varying stress<br>and COMT HPS |                    |       |          |                        | Stratum with 0 copies of HPS: Time-varying average quarterly PSS score minus baseline (per 6.4 units)      | 1.70 (1.24, 2.34)        |
|       |  |                    |       |          |                        | Stratum with I to 2 copies of HPS: Time-varying average quarterly PSS score minus baseline (per 6.4 units) | 2.36 (1.47, 3.77)        |
| A3    | Model I from Table 2, with interaction of                  | 2186               | 102.2 | 13       | 0.19                   | Low-activity COMT stratum: Baseline PSS score (per 6.4 units)  | 1.57 (1.26, 1.95)        |
|       | baseline stress and COMT diplotype                         |                    |       |          |                        | High-activity COMT stratum: Baseline PSS score (per 6.4 units)   | 1.89 (1.52, 2.36)        |

All models include time-constant covariates of study site (4 categories), age (continuous measure), sex (2 categories), and are adjusted for population stratification using the first three eigenvectors from multidimensional scaling analysis. COMT, catechol-O-methyltransferase; HPS, high-pain sensitivity; LPS, low-pain sensitivity; OPPERA, Orofacial Pain: Prospective Evaluation and Risk Assessment; PSS, Perceived Stress Scale; TMD, temporomandibular disorder; 95% CL, 95% confidence limits for the estimated hazard ratio.

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