Supplementary Materials for

Broadband slow-wave modulation in posterior and anterior cortex tracks distinct states of propofol-induced unconsciousness Emily P. Stephen, Gladia C. Hotan, Eric T. Pierce, P. Grace Harrell, John L. Walsh, Emery N. Brown, Patrick L. Purdon Correspondence to: patrick.purdon@mgh.harvard.edu

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Figure S1: Phase-amplitude coupling between the slow phase (0.1-4 Hz) and the alpha amplitude (8-16 Hz) using mean vector analysis for each subject. For each subject, the polar plot on the left shows the mean vectors for all electrodes for 100 30-second intervals spread over the entire session. The polar plot on the right shows the 2.5%, 50% (median), and 97.5% percentiles of the mean vectors, computed for 16 equally-spaced phase bins. Unlike the correlation-based cross-frequency coupling metric used in the rest of the paper, the mean vector estimate of phase is sensitive to coupling at 90 degrees and 270 degrees, corresponding to the rising phase and the falling phase of the slow oscillation. The results show that the amplitude of the high frequency signals tends to couple to either the peak or trough of the slow oscillation, and helps justify the use of the correlation between slow phase and fast amplitude as a means of quantifying phase-amplitude coupling.



Figure S2: Sensor-level summary for Subject 2 (20 years old). Layout as in Figure 1. Note that this subject entered burst suppression (Burst Supp) at the highest dose of propofol, so the previous level was used for the Unconscious High Dose condition.



Figure S3: Sensor-level summary for Subject 3 (22 years old). Layout as in Figure 1. A large number of posterior electrodes were excluded for this subject due to electrical bridging (seee Methods), so the posterior cross-frequency coupling has been omitted.



Figure S4: Sensor-level summary for Subject 4 (20 years old). Layout as in Figure 1. Note that this subject entered burst suppression (Burst Supp) at the highest dose of propofol, so the previous level was used for the Unconscious High Dose condition.



Figure S5: Sensor-level summary for Subject 5 (22 years old). Layout as in Figure 1.



Figure S6: Sensor-level summary for Subject 8 (23 years old). Layout as in Figure 1.



Figure S7: Sensor-level summary for Subject 9 (25 years old). Layout as in Figure 1.



Figure S8: Sensor-level summary for Subject 10 (25 years old). Layout as in Figure 1. Note that the Unconscious High Dose condition for this subject does not exhibit broadband peakmax dynamics over frontal electrodes: we interpret this to indicate that the subject was in a lighter state of unconsciousness than other subjects at the highest propofol dose.



Figure S9: Sensor-level summary for Subject 13 (23 years old). Layout as in Figure 1. Note that this subject entered burst suppression (Burst Supp) at the highest dose of propofol, so the previous level was used for the Unconscious High Dose condition.



Figure S10: Sensor-level summary for Subject 15 (31 years old). Layout as in Figure 1.



Figure S11: Cross-frequency coupling results for two electrodes (indicated in the insets, right), across all subjects (traces), for the four levels of interest.



Figure S12: Cross-frequency coupling patterns estimated by lobe in source space, projected onto the first principal mode. Each marker represents one subject. The markers are colored based on the value of the projection (also shown on the y-axis). Because the first mode is positive for all frequencies, positive values (red) correspond to broadband peakmax and negative values (blue) correspond to broadband troughmax.



Figure S13 Example EEG traces from two electrodes of Subject 7 (see Figure 1). Ten seconds of EEG data were selected from the four levels of interest (Baseline, Sedation, Unconscious Low Dose, and Unconscious High Dose), and the EEG data are displayed for a frontal and a posterior electrode (electrode locations shown in the insets to right). While Baseline and Sedation have little coupling of activity to the slow-wave, Unconscious Low Dose and Unconscious High Dose reflect the cross-frequency coupling evident in Figure 1: at Low Dose, the posterior electrode shows broadband coupling to the peak of the slow-wave; at High Dose, both posterior and frontal electrodes show broadband coupling to the peak of the slow-wave.