

Article

K-Nearest Neighbor (KNN) Entropy Estimates of Complexity and Integration from Non-Stationary Electroencephalographic (EEG) Recordings of the Human Brain

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Supplementary Material

1.1 Categorization Task Stimuli Spatial Frequencies and Orientations

Table S1. Spatial frequencies and orientations for 1-Exemplar Task Gabor stimuli

Category Stimulus	Category 1		Category 2	
	Frequency	Orientation	Frequency	Orientation
<i>Task Order 1</i>	3.86	57.50	3.24	68.30
<i>Task Order 2</i>	1.62	57.50	2.23	68.30
<i>Task Order 3</i>	3.24	21.70	3.86	32.43
<i>Task Order 4</i>	2.24	21.70	1.62	32.43
<i>Task Order 5</i>	3.24	57.50	3.86	68.30
<i>Task Order 6</i>	2.24	57.50	1.62	68.30
<i>Task Order 7</i>	3.86	21.70	3.24	32.43
<i>Task Order 8</i>	1.62	21.70	2.24	32.43

Spatial frequency values are in cycles/° and orientation values are in degrees from vertical.

Table S2. Spatial frequencies and orientations for 2-Exemplar Task Gabor stimuli

Category Stimulus		Category 1		Category 2	
		Frequency	Orientation	Frequency	Orientation
<i>Task Order 1</i>	1	1.49	18.07	2.33	18.07
	2	2.33	36.01	1.49	36.01
<i>Task Order 2</i>	1	3.14	18.07	3.97	18.07
	2	3.97	36.01	3.14	36.01
<i>Task Order 3</i>	1	1.49	53.95	2.33	53.95
	2	2.33	71.89	1.49	71.89
<i>Task Order 4</i>	1	3.14	53.95	3.97	53.95
	2	3.97	71.89	3.14	71.89

Spatial frequency values are in cycles/° and orientation values are in degrees from vertical.

1.2 Distributional Testing of EEG Property Measures

Parametric statistical analyses such as ANOVA typically assume that data is normally distributed. In order to check this assumption, Jarque-Bera tests of univariate normality [67] were performed for each EEG and behavioral measure across subjects. Given the large number of tests, Type-I error was minimized by correcting the p-values for multiple comparisons using the Holm-Bonferroni procedure [49]. None of the EEG measures were significantly different from a

normal distribution ($p_s > 0.14$). (It should be noted, however, that even though the final EEG integration and complexity values entering into the ANOVA were normally-distributed across subjects, the EEG data from which they were computed were non-normal; see Supplementary Materials Section 1.3., below. This suggests that the operations involved in the computation of $I(X)$ and $C_i(X)$ smooth the data in accordance with the Central Limit Theorem). Any concern that this approach was overly conservative and thus might lead to Type-II error in this analysis (i.e. some of the measures were non-normal, but did not depart from normality enough to yield a significant test) may be mitigated by the fact that ANOVAs and GEEs are fairly robust to minor violations of distributional assumptions [45-47].

The distributions of the behavioral reaction time data were also not significantly different from normal for both the 1-Exemplar and 2-Exemplar Categorization tasks. However, while the 2-Exemplar task accuracy rates were normally-distributed, the 1-Exemplar accuracy rates were not; this is because accuracy in this task was near ceiling (see Section 3.3 of the main text). This was accounted for in two ways: first, a nonparametric Wilcoxon signed rank test was used to assess accuracy differences between the two categorization tasks. Second, for GEE-based regressions relating accuracy to EEG integration and complexity, accuracy was treated as the independent variable and the EEG measures as the dependent variable; dependent variables are model-dependent in GEE analyses, whereas independent variables are model-independent [52]. Moreover, the GEE analyses used a robust covariance estimator, which allowed for a model-free estimate of the data covariance structure [53].

1.3 Distributional Testing and Gaussian-Transformation of EEG Data

In order to examine how KNN-based entropy estimation of $I(X)$ and $C_i(X)$ compared to the Gaussian-based estimation used by previous studies (Section 3.5 of the main text) it was first necessary to assess to what degree the statistical distribution of the dimension-reduced EEG data deviated from normality, then perform a Gaussian-transformation of the data, and finally assess whether or not the data transformation was successful. Following [11], the univariate and multivariate normality of the EEG signals was assessed via Jarque-Bera tests [67] and Royston's Test of multivariate normality [68], respectively, for each trial, condition, and participant. Royston's Test was computed via a publically available MATLAB script [69]. All tests were conducted at the $p < 0.05$, two-tailed, corrected-level. Gaussian transformation of the data was achieved using a previously established method that has been successfully used before with EEG data [11, 66]. The EEG data was transformed on a trial-by-trial basis for each separate condition, and participant. The distributional testing and Gaussian transformation were performed on the observed and simulated EEG data.

Although the distributions of less than 1% of EEG signals on average violated the univariate normality assumption on any given trial prior to Gaussian-transformation, the multivariate normality assumption was violated on 80% of trials on average. After the Gaussian transformation, none of the EEG trials violated univariate or multivariate normality (all $p_s = 1$).

1.4 Gaussian-Based Entropy Estimation

Marginal and joint entropies of the EEG signals were computed via explicit analytic expressions based on the assumption that a set X of N_s EEG signals realize continuous univariate and multivariate Gaussian processes with variances σ_{ii}^2 and covariance matrix K [6,17,70,71]:

$$H(X_i) = \frac{1}{2\ln(2)} \cdot \ln(2\pi e \sigma_{ii}^2) , \quad (S1)$$

$$H(X) = \frac{1}{2\ln(2)} \cdot \ln\left\{(2\pi e)^{N_s} |K|\right\} . \quad (S2)$$

Conditional entropy was computed according to Equation 3 of the main text. All entropy functions were computed with a correction for any bias that may arise due to the estimation of the covariance

matrices from limited data [70, 72-74]. All entropies were computed in terms of binary units (bits) of information.

Tables S3 – S7 display the quantitative and statistical results of the Gaussian-based computation of $I(X)$ and $C_I(X)$. The general pattern and interpretation of these findings are discussed in the main text Sections 3.5, 4.2, and 4.3.

Table S3. Mean KNN estimator-based EEG integration and complexity of Gaussian-transformed observed data

Induced EEG		Observed Data		Surrogate Data	
Task	Condition	$I(X)$	$C_I(X)$	$I(X)$	$C_I(X)$
1-Exemplar Task	Prestimulus	69.44 (0.36)	82.83 (0.08)	57.82 (57.18, 58.46)	86.69 (86.30, 87.08)
	Poststimulus	68.98 (0.42)	82.97 (0.05)	57.30 (56.71, 57.88)	86.57 (86.15, 86.98)
2-Exemplar Task	Prestimulus	69.43 (0.42)	82.88 (0.07)	57.88 (57.03, 58.72)	86.65 (86.22, 87.08)
	Poststimulus	68.71 (0.41)	82.89 (0.06)	57.12 (56.48, 57.76)	86.46 (86.06, 87.86)
Resting Task	Eyes Open	70.70 (0.41)	82.10 (0.04)	58.31 (57.65, 58.97)	86.50 (85.98, 87.02)
	Eyes Closed	73.02 (0.46)	82.21 (0.05)	60.19 (59.32, 61.05)	86.40 (85.97, 86.83)
Evoked EEG					
Task	Condition				
1-Exemplar Task	Prestimulus	66.13 (0.69)	79.45 (0.86)	57.81 (57.20, 58.42)	86.73 (86.37, 87.09)
	Poststimulus	79.20 (1.35)	82.72 (0.84)	57.13 (56.59, 57.68)	86.43 (86.13, 86.73)
2-Exemplar Task	Prestimulus	65.58 (0.64)	80.50 (0.85)	57.88 (57.04, 58.72)	86.74 (86.30, 87.19)
	Poststimulus	81.61 (1.05)	84.84 (0.60)	57.16 (56.56, 57.76)	86.59 (86.01, 87.17)

Note: All values are in bits; SE in parentheses for observed data, 95% CIs in parentheses for surrogate data.

Table S4. Analysis of variance (ANOVA) results for KNN estimator-based EEG integration and complexity of Gaussian-transformed observed data

Induced EEG					
Task	EEG Measure	Effect	F	P	η^2_P
<i>Categorization</i>	I(X)	Task	0.57	0.461	0.04
		TI	21.87	0.001	0.59
		Task x TI	6.27	0.024	0.30
	C _I (X)	Task	0.06	0.815	0.01
		TI	1.15	0.301	0.07
		Task x TI	1.57	0.230	0.10
<i>Resting State</i>	I(X)	RS	35.50	0.001	0.70
	C _I (X)	RS	7.15	0.017	0.32
Evoked EEG					
<i>Categorization</i>	I(X)	Task	1.42	0.252	0.09
		TI	193.36	0.001	0.93
		Task x TI	10.56	0.005	0.41
	C _I (X)	Task	3.83	0.069	0.20
		TI	17.95	0.001	0.55
		Task x TI	0.81	0.382	0.05

ANOVA factor labels: Task, Behavioral Task; TI, Time Interval; RS, Resting State. All dfs = 1, 15.

Table S5. Mean Gaussian estimator-based EEG integration and complexity of Gaussian-transformed observed data

Induced EEG		Observed Data		Surrogate Data	
Task	Condition	I(X)	C_i(X)	I(X)	C_i(X)
<i>1-Exemplar Task</i>	<i>Prestimulus</i>	72.96 (0.31)	40.46 (0.22)	56.70 (55.35, 58.06)	47.95 (46.87, 49.03)
	<i>Poststimulus</i>	72.11 (0.29)	40.85 (0.18)	55.90 (54.63, 57.17)	47.60 (46.50, 48.70)
<i>2-Exemplar Task</i>	<i>Prestimulus</i>	73.00 (0.34)	40.46 (0.24)	56.85 (55.24, 58.48)	47.98 (46.85, 49.11)
	<i>Poststimulus</i>	72.01 (0.31)	40.87 (0.18)	55.62 (54.30, 56.93)	47.44 (46.34, 48.53)
<i>Resting Task</i>	<i>Eyes Open</i>	76.93 (0.26)	39.20 (0.17)	58.29 (57.07, 59.50)	48.54 (47.28, 49.80)
	<i>Eyes Closed</i>	78.23 (0.34)	38.36 (0.21)	60.68 (59.24, 62.11)	48.29 (47.33, 49.25)
Evoked EEG					
Task	Condition				
<i>1-Exemplar Task</i>	<i>Prestimulus</i>	70.72 (0.81)	40.15 (0.65)	57.65 (56.42, 58.89)	45.45 (44.64, 46.26)
	<i>Poststimulus</i>	93.32 (2.46)	46.10 (0.81)	72.79 (68.00, 77.58)	49.41 (48.06, 50.76)
<i>2-Exemplar Task</i>	<i>Prestimulus</i>	70.51 (1.04)	40.04 (0.69)	57.46 (56.03, 58.90)	45.66 (44.65, 46.67)
	<i>Poststimulus</i>	96.53 (1.98)	47.65 (0.59)	75.10 (71.00, 79.19)	50.28 (49.15, 51.41)

Note: All values are in bits; SE in parentheses for observed data, 95% CIs in parentheses for surrogate data.

Table S6. Analysis of variance (ANOVA) results for Gaussian estimator-based EEG integration and complexity of Gaussian-transformed observed data

Induced EEG					
Task	EEG Measure	Effect	F	P	η^2_P
<i>Categorization</i>	I(X)	Task	0.07	0.795	0.01
		TI	70.34	0.001	0.82
		Task x TI	2.32	0.149	0.13
	C _I (X)	Task	0.01	0.953	0.0
		TI	17.43	0.001	0.54
		Task x TI	0.03	0.859	0.01
<i>Resting State</i>	I(X)	RS	27.33	0.001	0.65
	C _I (X)	RS	45.10	0.001	0.75
Evoked EEG					
<i>Categorization</i>	I(X)	Task	1.62	0.222	0.10
		TI	138.25	0.001	0.90
		Task x TI	3.29	0.090	0.18
	C _I (X)	Task	3.51	0.081	0.19
		TI	156.86	0.001	0.91
		Task x TI	9.77	0.007	0.40

ANOVA factor labels: Task, Behavioral Task; TI, Time Interval; RS, Resting State. All dfs = 1, 15.

Table S7. Mean KNN estimator-based and Gaussian estimator-based EEG integration and complexity of Gaussian-transformed dipole simulation data

Data Condition	Prestimulus	Poststimulus	F	p	η^2_P
KNN-Based					
<i>I(X)</i> _{Induced}	68.48 (0.02)	69.19 (0.06)	161.3	0.001	0.92
<i>C_I(X)</i> _{Induced}	82.04 (0.08)	82.72 (0.10)	19.72	0.001	0.57
<i>I(X)</i> _{Evoked}	64.25 (0.62)	73.59 (0.46)	119.11	0.001	0.89
<i>C_I(X)</i> _{Evoked}	80.58 (0.60)	83.48 (0.82)	10.19	0.006	0.40
Gaussian-Based					
<i>I(X)</i> _{Induced}	73.31 (0.03)	71.81 (0.04)	1461.14	0.001	0.99
<i>C_I(X)</i> _{Induced}	40.28 (0.01)	40.87 (0.03)	438.93	0.001	0.97
<i>I(X)</i> _{Evoked}	69.33 (1.14)	81.50 (0.71)	76.17	0.001	0.84
<i>C_I(X)</i> _{Evoked}	37.97 (0.35)	40.60 (0.31)	54.07	0.001	0.78

Complexity and integration values are in bits, GFP values are in μV , order parameters are dimensionless; SE in parentheses. ANOVA parameters describe the significance of prestimulus versus poststimulus differences.

1.5 Basic EEG Dipole Simulations

The development of the dipole simulations of the empirically-observed categorization task data involved the initial exploration of EEG signals that in terms of dipole source amplitude, phase, and changes in synchronization of dipole oscillations and amplitude over time. Each simulation consisted of the creation of one hundred 2 second trials of simple, wide-range (4 – 13 Hz) oscillatory waveforms that varied in terms of initial amplitude, initial synchronization of the waveform phase, and changes in amplitude and phase coupling. General simulation procedures are described in Section 2.9 of the main text, with manipulation of the following parameters: initial amplitude (high: 60 μ A-cm, low: 30 μ A-cm), starting phase (random: 2π , synched: $\pi/50$), Kuramoto parameter K (0, 5, 10), amplitude change across time (constant; variable: $\sigma = 250$ ms Gaussian envelopes with ± 1000 ms peak latency; synchronized: $\sigma = 250$ ms Gaussian envelopes with ± 39 ms peak latency), and the dependency of variable/synchronized amplitude changes (full independence or full dependence). In addition, high and low amplitude versions of two special waveforms were simulated, one with a nonstationary discontinuous phase created via the method of Theiler et al. [58] and a second created as a multivariate normal process with $\mu = 0$ and $\sigma = 1$.

Tables S8 – S11 display the quantitative results of these basic EEG dipole simulations. In general, these simulations showed qualitatively that EEG integration increased with increasing oscillatory synchronization and synchronized amplitude changes, but decreased with overall reductions in dipole amplitude. EEG complexity followed the theoretically predicted relationship with integration as given in Figure 1, but was mainly affected by changes in dipole amplitude when the dipole oscillations and/or amplitude changes were unsynchronized. Induced and evoked EEG

Table S8. Basic EEG dipole source simulations: High amplitude, random start phase

Data Condition	I(X)	C_i(X)	GFP^{Induced}	GFP^{Evoked}	Λ^{Induced}	Λ^{Evoked}
<i>K_{param} = 0, Constant Amplitude</i>	126.71	111.78	819.69	86.94	0.192	0.220
<i>K_{param} = 5, Constant Amplitude</i>	132.33	116.86	1271.70	100.91	0.296	0.279
<i>K_{param} = 10, Constant Amplitude</i>	136.90	117.38	1593.98	156.00	0.331	0.319
<i>K_{param} = 0, Variable Independent Amplitude</i>	116.18	117.70	877.85	80.35	0.213	0.201
<i>K_{param} = 5, Variable Independent Amplitude</i>	121.63	120.40	1234.61	100.43	0.277	0.259
<i>K_{param} = 10, Variable Independent Amplitude</i>	130.78	118.98	1604.06	136.03	0.331	0.294
<i>K_{param} = 0, Variable Dependent Amplitude</i>	131.25	109.75	894.95	85.45	0.221	0.238
<i>K_{param} = 5, Variable Dependent Amplitude</i>	132.99	114.98	1300.91	124.26	0.299	0.289
<i>K_{param} = 10, Variable Dependent Amplitude</i>	139.87	116.54	1709.29	152.93	0.333	0.323
<i>K_{param} = 0, Synched Dependent Amplitude</i>	130.59	111.26	804.79	78.96	0.217	0.203
<i>K_{param} = 5, Synched Dependent Amplitude</i>	135.41	114.62	1213.14	85.85	0.291	0.317
<i>K_{param} = 10, Synched Dependent Amplitude</i>	142.83	115.93	1632.33	115.78	0.33	0.327
<i>K_{param} = 0, Variable Amplitude, Nonstationary phase</i>	116.74	117.91	910.25	90.30	0.219	0.217
<i>Multivariate Normal Process ($\mu = 0, \sigma = 1$)</i>	67.85	102.70	647.87	67.29	0.252	0.263

All values are averages of electrodes, time, and trials. Complexity and integration values are in bits, power values are in μ V, order parameter is dimensionless; SE in parentheses.

power were generally larger for synchronized versus unsynchronized dipole oscillations and amplitude changes, which reflects the constructive versus destructive summation of EEG signals as they are volume-conducted through the scalp. However, as for the empirical data, induced EEG power was greater than evoked power. Finally, induced and evoked EEG power were generally larger for synchronized versus unsynchronized dipole oscillations and amplitude changes, but like EEG complexity was mainly affected by changes in dipole amplitude when the dipole oscillations and/or amplitude changes were unsynchronized.

Table S9. Basic EEG dipole source simulations: Low amplitude, random start phase

Data Condition	I(X)	C_i(X)	GFP^{Induced}	GFP^{Evoked}	Λ^{Induced}	Λ^{Evoked}
<i>K_{param} = 0, Constant Amplitude</i>	126.71	111.78	416.23	31.81	0.220	0.167
<i>K_{param} = 5, Constant Amplitude</i>	132.33	116.86	614.35	70.41	0.304	0.318
<i>K_{param} = 10, Constant Amplitude</i>	136.90	117.38	812.27	49.84	0.328	0.334
<i>K_{param} = 0, Variable Independent Amplitude</i>	116.18	117.70	448.79	44.51	0.218	0.184
<i>K_{param} = 5, Variable Independent Amplitude</i>	121.63	120.40	624.51	58.63	0.286	0.253
<i>K_{param} = 10, Variable Independent Amplitude</i>	130.78	118.98	812.15	64.55	0.334	0.318
<i>K_{param} = 0, Variable Dependent Amplitude</i>	131.25	109.75	450.85	40.32	0.221	0.186
<i>K_{param} = 5, Variable Dependent Amplitude</i>	132.99	114.98	644.77	47.72	0.294	0.261
<i>K_{param} = 10, Variable Dependent Amplitude</i>	139.87	116.54	857.65	71.65	0.330	0.333
<i>K_{param} = 0, Synched Dependent Amplitude</i>	130.59	111.26	417.81	35.05	0.216	0.222
<i>K_{param} = 5, Synched Dependent Amplitude</i>	135.41	114.62	603.40	51.09	0.290	0.304
<i>K_{param} = 10, Synched Dependent Amplitude</i>	142.83	115.93	816.21	62.57	0.328	0.309
<i>K_{param} = 0, Variable Amplitude, Nonstationary phase Multivariate Normal Process ($\mu = 0, \sigma = 1$)</i>	116.74	117.91	449.22	47.41	0.215	0.148
	67.85	102.70	325.85	31.12	0.247	0.244

All values are averages of electrodes, time, and trials. Complexity and integration values are in bits, power values are in μV , order parameter is dimensionless; SE in parentheses.

Table S10. Basic EEG dipole source simulations: High amplitude, synchronized start phase

Data Condition	I(X)	C _i (X)	GFP ^{Induced}	GFP ^{Evoked}	Λ ^{Induced}	Λ ^{Evoked}
<i>K_{param} = 0, Constant Amplitude</i>	126.71	111.78	1948.82	476.19	0.372	0.374
<i>K_{param} = 5, Constant Amplitude</i>	132.33	116.86	1948.40	378.26	0.372	0.373
<i>K_{param} = 10, Constant Amplitude</i>	136.90	117.38	1948.62	427.74	0.372	0.374
<i>K_{param} = 0, Variable Independent Amplitude</i>	116.18	117.70	1910.16	406.21	0.367	0.369
<i>K_{param} = 5, Variable Independent Amplitude</i>	121.63	120.40	1907.12	427.04	0.368	0.372
<i>K_{param} = 10, Variable Independent Amplitude</i>	130.78	118.98	1914.63	397.18	0.368	0.366
<i>K_{param} = 0, Variable Dependent Amplitude</i>	131.25	109.75	2107.94	418.37	0.374	0.374
<i>K_{param} = 5, Variable Dependent Amplitude</i>	132.99	114.98	2115.96	454.54	0.374	0.374
<i>K_{param} = 10, Variable Dependent Amplitude</i>	139.87	116.54	2104.34	465.12	0.374	0.375
<i>K_{param} = 0, Synched Dependent Amplitude</i>	130.59	111.26	1958.32	281.76	0.370	0.374
<i>K_{param} = 5, Synched Dependent Amplitude</i>	135.41	114.62	1952.69	302.75	0.371	0.374
<i>K_{param} = 10, Synched Dependent Amplitude</i>	142.83	115.93	1962.71	225.23	0.371	0.374
<i>K_{param} = 0, Variable Amplitude, Nonstationary phase</i>	116.74	117.91	890.42	80.13	0.227	0.240
<i>Multivariate Normal Process ($\mu = 0, \sigma = 1$)</i>	67.85	102.70	650.18	62.33	0.250	0.255

All values are averages of electrodes, time, and trials. Complexity and integration values are in bits, power values are in μV , order parameter is dimensionless; SE in parentheses.

Table S11. Basic EEG dipole source simulations: Low amplitude, synchronized start phase

Data Condition	I(X)	C _i (X)	GFP ^{Induced}	GFP ^{Evoked}	Λ ^{Induced}	Λ ^{Evoked}
<i>K_{param} = 0, Constant Amplitude</i>	126.71	111.78	973.45	237.56	0.370	0.375
<i>K_{param} = 5, Constant Amplitude</i>	132.33	116.86	969.83	256.34	0.371	0.374
<i>K_{param} = 10, Constant Amplitude</i>	136.90	117.38	976.09	220.55	0.371	0.376
<i>K_{param} = 0, Variable Independent Amplitude</i>	116.18	117.70	953.97	218.06	0.367	0.365
<i>K_{param} = 5, Variable Independent Amplitude</i>	121.63	120.40	954.19	223.16	0.367	0.37
<i>K_{param} = 10, Variable Independent Amplitude</i>	130.78	118.98	955.17	188.01	0.366	0.369
<i>K_{param} = 0, Variable Dependent Amplitude</i>	131.25	109.75	1055.30	205.92	0.372	0.374
<i>K_{param} = 5, Variable Dependent Amplitude</i>	132.99	114.98	1054.52	215.45	0.372	0.371
<i>K_{param} = 10, Variable Dependent Amplitude</i>	139.87	116.54	1054.28	224.26	0.372	0.374
<i>K_{param} = 0, Synched Dependent Amplitude</i>	130.59	111.26	976.55	158.35	0.367	0.373
<i>K_{param} = 5, Synched Dependent Amplitude</i>	135.41	114.62	979.88	137.57	0.366	0.374
<i>K_{param} = 10, Synched Dependent Amplitude</i>	142.83	115.93	977.67	152.26	0.367	0.372
<i>K_{param} = 0, Variable Amplitude, Nonstationary phase Multivariate Normal Process ($\mu = 0, \sigma = 1$)</i>	116.74	117.91	450.49	41.58	0.208	0.200
	67.85	102.70	324.42	30.68	0.247	0.252

All values are averages of electrodes, time, and trials. Complexity and integration values are in bits, power values are in μV , order parameter is dimensionless; SE in parentheses.

References

All references cited in the Supplementary Materials are listed in the References section of the main text.