

DeepPsy Biomarker's Report Guide

April 2024 - v1.0.0

Table of Contents

- 1. Introduction
- 2. Overview of the report sections:
 - a. Analysis Metadata
 - b. Biomarker Plots
 - c. Biomarker Values & Interpretations
 - d. Biomarker Descriptions
- 3. Frequently Asked Questions (FAQs)

Introduction

The DeepPsy Biomarkers Report is generated from an analysis of patient EEG and ECG data. It provides a concise summary of the analysis performed, detailing the biomarkers identified from the physiological signals. The report includes charts that offer a comprehensive view of the mainly predictive markers for optimization of treatment and the patient's condition, alongside interpretations of the data, which are based on current scientific literature.

This guide will provide a description of the different sections in the report and how they can be interpreted and used.

Evidence level

Each "Interpretation" in the 3rd page (interpretations section) has a label with the "evidence level". This is a statement about the strength of the evidence in this finding.

To rank the evidence level, we are using the "The Oxford 2011 Levels of Evidence" from the Oxford Centre for Evidence-Based Medicine.

The system categorizes evidence from Level 1 (high-quality evidence) to Level 5 (lowest evidence level). In the case of our predictive biomarkers, we follow:

Level 1	Level 2	Level 3	Level 4	Level 5
Systematic review of inception cohort studies	Inception cohort studies	Cohort study or control arm of randomized trial*	Case-series or case-control studies, or poor quality prognostic cohort study**	n/a

The header and footer have basic identifying information that appears in every page. The patient information is printed in every page to avoid the mixing of reports.

This section gives general information about the EEG or ECG itself date, length etc) and the analysis performed by the DeepPsy specialist.

If there were any problems during the analysis, they're shown here. Additionally, if there's any information from the analysis that is important for the interpretation of the results, it'll be shown here as well.



Name:

Patient ID: Age: 29 Sex: Female Case ID:

Report ID: 1723-163-9625832 Analysis Date: 25.04.2024 Creation Date: 25.04.2024

Biomarkers Report

This report CAN be used in the following circumstances:

- · When requested and interpreted by a physician or psychiatric/medical institutions.
- When requested and interpreted following a verified diagnosis or suspected diagnosis by a physician.

tion, and the relevance of the report is weighed in the context of the entire clinical picture.

- · When used as an additional source of information alongside other clinical and paraclinical sources of informa-
- · When the final decision and responsibility regarding the use of the report's information rests with the requesting doctor or health institution.
- · When the EEG and ECG data from which this report is generated come from certified medical device ampli-
- · When the EEG and ECG data have been reviewed by the requesting doctor for further neurological or cardiac pathologies or the influence of medications.

This report CANNOT be used under the following circumstances:

- · Not if it is intended for diagnostic purposes. The report is only intended for use with patients who have already been given a (suspected) diagnosis by a trained professional.
- · Not if it is to decide whether to undergo psychiatric treatment.
- Not if the report is to be considered as a binding guideline.
- · Not if it is intended for self-diagnosis or self-treatment.
- · Not in the presence of neurological pathologies (in the EEG) or cardiac pathologies (in the ECG).

Analysis Characteristics

- · EEG analysis completed successfully. ECG analysis completed successfully.
- · All biomarkers were computed successfully.

Recording Date: 26.02.2024 Sampling Frequency: 1000 Hz

Total Recording Duration: 1289s (21.5m) Analysis Interval: 201s - 787s (9.8m) Number of Channels: EEG(21) EOG(1) ECG(1) Channel Types:

EOG (POL PG1 - POL PG2) Bad Channels Interpolated: F7, Fp2 EOG Channel:

Number of Epochs: Epochs with Artefacts: 8 (1.4%) FCG Peaks: 119 Peaks Corrected: None

The Vigilance section (EEG) has a plot with the level of vigilance of the patient throughout the whole recording (left panel) and the slope of the vigilance level over the first two minutes (right panel).

The slope is the value that has predictive clinical implications.

The plot shows an example of segments with the Alpha Peak Frequency (APF) for the patient (EEG). From the APF as the basic rhythm of the brain, several predictive aspects can be concluded.

This plot shows the distribution of Heart Rate Variability, derived from the ECG.

High values of Low Frequency (LF) power reflect the adaptive capabilities of the autonomic nervous system. Low LF values are usually associated with the potential for good relaxation.

High values of High Frequency (HF) power are associated with a relaxed state, where low HF power values can be associated with stress or anxiety

Case ID: Patient ID: Report ID: 1723-163-9625832 Analysis Date: 25.04.2024 Sex: Female Creation Date: 25.04.2024 Vigilance Vigilance Slope 0.00Time [sec] Alpha Peak Frequency Frontal Alpha Asymmetry Cha Time [sec] Heartrate Variability Low Frequency (LF) SD 10000 -0.05Hz 0.10Hz 0.15Hz 0.20Hz 0.25Hz 0.30Hz High Frequency (HF) Histogram of **Heart Rate** Heart Rate Slope beat per minute -0.75 (BPM) for the ECG. This marker has predictive value for treatment options. v0.4.32MD 2 of 6 For any questions, please contact us at: +44 555 111 4444 | support@deeppsy.io

The hemisphere with greater frontal alpha power was highlighted for the frontal alpha asymmetry.

This marker s has predictive clinical implications specific in female patients.

Slope of the beat per minute (BPM) for the ECG. This marker has predictive value for treatment options.

The individual values or each of the biomarkers can be found here. For eligible biomarkers, the normal range in the healthy population is presented. Values outside the boundary of two standard deviations are presented in bold type.

Report interpretations concisely summarize the associations found in the scientific literature between EEG and ECG biomarkers and treatment effectiveness. All references to the corresponding studies are given alongside with a rating of the clinical evidence (level 1 -highest level of evidence- to level 4 -lowest level of evidence)

Note: Since there exist several different markers for specific treatments (e.g. for SSRIs), the results sometimes can be contradictory between biomarkers. The results need to be put into the clinical context by a physician and to be discussed with the patient.



Name: Patient ID: Age: 29 Sex: Female Case ID:

Report ID: 1723-163-9625832 Analysis Date: 25.04.2024 Creation Date: 25.04.2024

Biomarker Values

	Value (first 2min)	Normal Interval (2 SD)
EEG:	, ,	, ,
Alpha Frontal Asymmetry	-0.0865	(-0.3 - 0.3)
Alpha Peak Frequency (Hz)	12.0*	(8.0 - 11.7)
qEEG Alpha (µV²)	45.4	(0.0 - 144.0)
qEEG Beta (μV^2)	17.55*	(0.0 - 16.0)
qEEG Delta (μV^2)	6.02	(0.0 - 16.0)
qEEG Gamma1 (µV2)	1.51	(0.0 - 2.0)
qEEG Gamma2 (μV^2)	0.48*	(0.0 - 0.2)
qEEG Theta (μV^2)	5.37	(0.0 - 32.0)
Vigilance Regulation A1 Stages (%)		_
Vigilance Level (Level)	3.0	(2.2 - 6.0)
Vigilance Regulation (Level/min)	-0.0	(-0.5 - 0.4)
Vigilance Regulation 9.8min (Level/min)	-0.0	-
ECG:		
Heart Rate (BPM) (beats/min)	60.0	(53.0 - 76.0)
Heart Rate Trend (BPM Slope) (beats/min ²)	-0.75	(-2.91 - 2.73)
Parasympathetic Activity (HF) (ms ²)	644.73	(0.0 - 4320.0)
Sympathetic/Parasympathetic Ratio	65.49*	(0.0 - 8.0)
Sympathetic Activity (LF) (ms ²)	1223.39	(0.0 - 4242.0)

Interpretations

EEG

- Frontal Alpha Asymmetry (FAA) Left dominant frontal alpha activity (-). Female patients with depression show a
 decreased response to SSRIs (Evidence level: 2)*
- Alpha Peak Frequency (APF) High Alpha Peak Frequency (+). For depression, 1Hz TMS protocols over the rDLPFC may be more effective than 10Hz protocols over the left DLPFC (Evidence level: 2). There is also a negative correlation with the response to Sertraline (Evidence level: 2). In the case of ADHD, medication with Methylphenidate might be more effective than biofeedback treatment (Evidence level: 4).
- Vigilance Level Low vigilance level (-). Low vigilance levels can be associated with increased fatigue, drowsiness, medication or manic syndroms.
- Vigilance Regulation 2min Increase or no initial decrease of vigilance during the first 2 minutes (+). Literature shows lower response rates in depression to SSRIs. SNRIs can be more effective (Evidence levels: 2)*. Increasing Vigilance levels were found more often in patients after a recent suicide attempt in comparison to healthy controls (Evidenzyrad: 4).
- Vigilance Regulation No decrease of vigilance (+) in the analysis interval of (9.8min). Commonly observed in patients diagnosed with depression or OCD (Evidence Level 2). Response to medication in case of ADHD less probable (Evidence Level 4).

Interpretation for ECG (if present) data follow the same format as for EEG. The associations of biomarker values found for the patient and treatment effectiveness are reported. All references to the corresponding studies are given alongside with a rating of the clinical evidence (level 1 -highest level of evidence- to level 4 -lowest level of evidence)

The descriptions offer more comprehensive information about each biomarker. This can be useful in cases where one wishes to explain the meaning of the biomarkers to the patient, or simply to obtain more information in order to better support clinical decisions.



Name: Patient ID: Age: 29 Sex: Female Case ID:

Report ID: 1723-163-9625832 Analysis Date: 25.04.2024 Creation Date: 25.04.2024

ECG

- Heart Frequency (BPM) Normal heart rate (0). Normal heart frequency is not associated with special treatment recommandations.
- Heart Frequency Regulation Decrease or no evidence of substantial rising of BPM (0). In depression, there is a
 correlation with lower response rates for venlafaxine (SNRI). SSRIs can be more effective. (Evidence level: 2)*
- Absolute parasympathetic Activation Normal parasympathetic activation (0). Compared to the average population, parasympathetic activity is normal.
- Relative Sympathetic/Parasymathetic Activation
 Balanced Sympathetic and Parasympathetic activity (0). A
 balanced sympathetic-parasympathetic activity is not associated with a special treatment indication.
- Absolute Sympathetic Activation High sympathetic activation (+). Sympathetic activity is increased compared to the average population.

Biomarker Descriptions

Frontal Alpha Asymmetry (FAA) Female patients with left frontal alpha asymmetry and depressive syndromes responsed to SNRIs (venlafaxine or duloxetine) more likely than to SSRIs (citalopram or sertraline).

Alpha Peak Frequency (APF) The APF is 12.0Hz and higher than the average of a healthy control population. There is evidence that treatment using a TMS protocol with 1 Hz stimulation over the rDLPFC can be more effective than treatment using a 10 Hz protocol over the IDLPFC in case of depression. If the APF is high and ADHD is present, a better response to medication with methylphenidate can be expected than to biofeedback.

Vigilance Regulation 2min No substantial drop found in the vigilance of the first two minutes of the resting EEG. Stable vigilance regulation correlates with larger non-response to a SSRI (Sertraline or Citalopram) in MDD patients in comparison to SNRIs.

Vigilance Regulation The vigilance in the present EEG does not drop over the course of 9.8min, and is very stable. Stable vigilance regulation is found more frequently in patients with a depressive syndrome or with obsessive-compulsive disorders. Rapidly falling vigilance levels, on the other hand, are more often associated with manic syndromes, emotionally unstable personality disorders, ADHD and MS- or cancer related Fatigue.

Heart Frequency (BPM) A normal heart rate during resting state is a physiological profile and shows a good balance of the autoinomic nervous system.

Heart Frequency Regulation A decrease or a slow increase of the heart rate during 2min of rest is associated with a good relaxation during rest. In depression, it has been associated with less likely response to ventafaxine (SNRI) and a higher response rate for SSRIs.

Absolute parasympathetic Activation In the ECG, the absolute parasympathetic activity is in a normal range compared to a healthy control population. This finding is correlated with a good balance between relaxation and drive.

4 of 6

v0.4.32MD For any questions, please contact us at: +44 555 111 4444 | support@deeppsy.io

Literature references of all cited papers in the interpretation section.

The "interpretations" section in page 3 has a number of reference number next to each title. You can find further information about each interpretation in the literature listed here.



Name: Patient ID: Age: Unknown Sex: Unknown Case ID:

Report ID: 49OTDQD6IL Analysis Date: 29.04.2024 Creation Date: 29.04.2024

References

- Dohrmann A-L, Stengler K, Jahn I, Olbrich S (2017) EEG-arousal regulation as predictor of treatment response in patients suffering from obsessive compulsive disorder. Clin Neurophysiol 128:1906–1914. https://doi.org/10.1016/j.clinob.2017.07.406
- [2] Hegerl U, Wilk K, Olbrich S, et al (2011) Hyperstable regulation of vigilance in patients with major depressive disorder. The World Journal of Biological Psychiatry: The Official Journal of the World Federation of Societies of Biological Psychiatry. https://doi.org/10.3109/15622975.2011.579164
- [3] Hegerl U, Stein M, Mulert C, et al (2008) EEG-vigilance differences between patients with borderline personality disorder, patients with obsessive-compulsive disorder and healthy controls. Eur Arch Psychiatry Clin Neurosci 258:137–143. https://doi.org/10.1007/s00406-007-0765-8
- [4] Olbrich S, Sander C, Jahn I, et al (2011) Unstable EEG-vigilance in patients with cancer-related fatigue (CRF) in comparison to healthy controls. The World Journal of Biological Psychiatry. The Official Journal of the World Federation of Societies of Biological Psychiatry. https://doi.org/10.3109/15622975.2010.545434
- [5] Small JG, Milstein V, Medlock CE (1997) Clinical EEG findings in mania. Clin Electroencephalogr 28:229–235.
- [6] Olbrich S, Sander C, Minkwitz J, et al (2012) EEG Vigilance Regulation Patterns and Their Discriminative Power to Separate Patients with Major Depression from Healthy Controls. Neuropsychobiology 65:188–194. https://doi.org/10.1159/000337000
- [7] Sander C, Arns M, Olbrich S, Hegerl U (2010) EEG-vigilance and response to stimulants in paediatric patients with attention deficit/hyperactivity disorder. Clin Neurophysiol 121:1511–1518. https://doi.org/10.1016/j.clinph.2010.03.021
- [8] Olbrich S, Tränkner A, Surova G, et al (2016) CNS- and ANS-arousal predict response to antidepressant medication: Findings from the randomized iSPOT-D study. Journal of Psychiatric Research 73:108–115. https://doi.org/10.1016/j.jpsychires.2015.12.001
- [9] Ip C-T, Ganz M, Dam VH, et al (2021) NeuroPharm study: EEG wakefulness regulation as a biomarker in MDD. Journal of Psychiatric Research 141:57–65. https://doi.org/10.1016/j.jpsychires.2021.06.021
- [10] Strauß M, Reif A, Ulke C, et al (2020) Is brain arousal regulation a predictor of response to psychostimulant therapy in adult ADHD patients? Eur Arch Psychiatry Clin Neurosci 270:1073–1076. https://doi.org/10.1007/s00406-019-01085-y
- [11] Roelofs CL, Krepel N, Corlier J, et al (2021) Individual alpha frequency proximity associated with repetitive transcranial magnetic stimulation outcome: An independent replication study from the ICON-DB consortium. Clin Neurophysiol 132:643-649. https://doi.org/10.1016/j.clinph.2020.10.017
- [12] Corlier J, Carpenter LL, Wilson AC, et al (2019) The relationship between individual alpha peak frequency and clinical outcome with repetitive Transcranial Magnetic Stimulation (rTMS) treatment of Major Depressive Disorder (MDD). Brain Stimulation 12:1572–1578. https://doi.org/10.116/j.br.2019.07.018
- [13] Voetterf H, van Wingen G, Michelini G, et al (2022) Brainmarker-I Differentially Predicts Remission to Various Attention-Delicit/Hyperactivity Disorder Treatments: A Discovery, Transfer, and Blinded Validation Study. Biological Psychiatry: Cognitive Neuroscience and Neurolimaging. https://doi.org/10.1016/j.bpse.2022.02.007
- [14] Arns M, Bruder G, Hegerl U, et al (2016) EEG alpha asymmetry as a gender-specific predictor of outcome to acute treatment with different antidepressant medications in the randomized iSPOT-D study. Clin Neurophysiol 127:509–519. https://doi.org/10.1016/j.clinph.2015.05.032
- [15] Ip C-T, Olbrich S, Ganz M, et al (2020) Pretreatment qEEG biomarkers for predicting pharmacological treatment outcome in Major Depressive Disorder: Independent validation from the NeuroPharm study, submitted
- [16] UK Biobank, https://www.ukbiobank.ac.uk/.
- [17] Annia Rüesch, Tania Villar de Araujo, Anna Bankwitz, Christoph Hörmann, Atalia Adank, Cheng-Teng Ip, Georgios Schoretsanitis, Birgit Kleim, Sebastian Olbrich, A recent suicide attempt and the heartheat: Electrophysiological findings from a trans-diagnostic cohort of patients and healthy controls, Journal of Psychiatric Research, Volume 157 2023, Pages 257-263.
- [18] Annia Rüesch EEG wakefulness regulation in transdiagnostic patients after a recent suicide attempt ScienceDirect

v0.4.32MD For any questions, please contact us at: +44 555 111 4444 | support@deeppsy.io 6 of 6

Frequently Asked Answers (FAQs)

- Why are some of the recommendations contradictory? It's possible that some biomarkers give contradictory information. This is the nature of assessing different sources of information. It's important to weigh the merits of the interpretations against each other and against the patient's medical history.