

Reply to Reviewer's Comments on "EEG characterization and classification based on image gradient histograms" (now "Histogram of Gradient Orientations of Signal Plots applied to P300 Detection")

We are grateful to the reviewer for pointing out relevant issues in our manuscript.
 In the following, we discuss how we dealt with each raised issue.

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REVIEWER #1 TRANSCRIPT:

I have read the manuscript "EEG characterization and classification based on Histogram of Gradients of Signal Plots" with great interest since its title promised a solution to an important problem. In spite of presenting an interesting methodology and some interesting issues related to EEG characterization, I consider the manuscript is not ready for publication.

First of all, the title is misleading since a first, and naive, read suggest being able to classify and characterize any EEG with their different components (i.e., P200, P300, P600, N100, N200, N400 to name a few). Clearly, the title does not match the content since the authors focus on the P300; of course, there is evidence of the benefits of working with these signals, but, is the proposed method extensible to other signals?, this is not clear from the manuscript.

Another issue with the manuscript is its mathematical vagueness. For example, the variable T is used as a vector or two-tuple but its notation is different from the notation the authors use for vectors. Moreover, there is something wrong in equation (4) and something inconsistent between equations (7) and (8). Also, the authors go back and forth the digital space and the continuous space and although there is an equivalence, the manuscript could benefit from a clearer explanation.

Another issue that I find worrisome is the design of experiments. First of all, it seems that the number of subjects is rather low, it would be interesting to know why the authors decided to use such a subset. Also, the authors speculate that their results are not that good because they did not take into account the latency and magnitude of the signals, there is previous evidence in the literature that the P300 suffers from these, and other, problems and, thus, one wonders why the authors did not anticipate such problems.

A couple of finale observations. First, it seems the authors assume that the readers have some previous deep knowledge of some topics such as that some channels should carry more or better information without a previous reference. Another issue has to do with the way the manuscript is written, sometimes is very difficult to follow the ideas the authors want to convey, an example can be found in the paragraph containing equation (5), the manuscript surely benefit from a careful review before submitting.

General Comments

I have read the manuscript "EEG characterization and classification based on Histogram of Gradients of Signal Plots" with great interest since its title promised a solution to an important problem. In spite of presenting an interesting methodology and some interesting issues related to EEG characterization, I consider the manuscript is not ready for publication.

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Thank you for your comments and observations. We have modified several points according to your recommendations, we hope the article is now better suited for publication.

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2.1) the title is misleading since a first, and naive, read suggest being able to classify and characterize any EEG with their different components (i.e., P200, P300, P600, N100, N200, N400 to name a few). Clearly, the title does not match the content since the authors focus on the P300

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Thank you for the observation. The title of the article was changed accordingly to your suggestion.

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2.2) there is evidence of the benefits of working with these signals P300, but, is the proposed method extensible to other signals?, this is not clear from the manuscript.

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Thank you very much for your comment. We published in (Ramele *et al.*, 2016) the application of the same method to identify rhythmic EEG events like Visual Occipital Alpha Waves and to classify Motor Imagery. We are also working on unpublished material where we are analyzing the same approach for the detection of K-Complexes and classification of SSVEP patterns but the results are in progress. We added this information on the article (Introduction and Conclusion, section 3). We hope this new information enriches the article and clarifies this point.

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Third Paragraph

3.1) Another issue with the manuscript is its mathematical vagueness. For example, the variable T is used as a vector or two-tuple but its notation is different from the notation the authors use for vectors

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Thank you for the observation. We worked hard to improve all the mathematical description of the method. The issue with variable T was fixed and we improved on the font styling to differentiate vectors and scalars. Section 1.1 was modified as well as section 1.2.3. The mathematical terminology were corrected all along the article. We hope we have full addressed this point.

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3.2) Moreover, there is something wrong in equation (4)

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We appreciate a lot your truthful comment. Equation (4) is very important for us because it is the essence of the method. We reviewed the equation, corrected their terms and we hope it is now more clear.

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3.3) and something inconsistent between equations (7) and (8)

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We have revised the manuscript and modified equations (7) and (8) (now (8) and (9)) which are located on Section 1.2.3 (Parameters). We added more information on the text and specified subscript indexes to avoid the inconsistency. We thank you for noticing this.

3.4) Also, the authors go back and forth the digital space and the continuous space and although there is an equivalence, the manuscript could benefit from a clearer explanation.

Thank you for the observation. We modified all equations to emphasize the point that they are calculated in digital space, particularly on equation (4). We also added that image gradients are also calculated using finite differences, and added information about the support of each equation to explain that they are actually calculated in digital space. With this, we hope we have full addressed your point.

Fourth Paragraph

4.1) Another issue that I find worrisome is the design of experiments. It seems that the number of subjects is rather low, it would be interesting to know why the authors decided to use such a subset.

Thank you for your comment. The original dataset was published as the 008-2014 dataset at the BNCI-Horizon website, which is very important in BCI Research. As far as we are aware of, experiments on this dataset of ALS patients has not been published by any independent group. In order to address this issue, we included a new own dataset with the same experimental conditions that we generated ourselves, on healthy subjects. We published this dataset on the kaggle platform for replication.

4.3) Also, the authors speculate that their results are not that good because they did not take into account the latency and magnitude of the signals, there is previous evidence in the literature that the P300 suffers from these, and other, problems and, thus, one wonders why the authors did not anticipate such problems.

Thank you for your comments. We added more detailed information that address this issue. First in Section 1.1.4 we included information about the signal averaging procedure. In Section 1.2.3 we added the parameters that were used. More importantly, in Section 2 we added information and references about the tests that we did to address latency and amplitude issues. We included in Section 3 information about future and current works where we are also dealing with these problems. We hope we full addressed your remarks.

Fifth Paragraph

5.1) it seems the authors assume that the readers have some previous deep knowledge of some topics such as that some channels should carry more or better information without a previous reference.

Thank you for your suggestion. The Results section was modified to clarify the expectation about the EEG channels where a visual P300 oddball paradigm generates stronger signals. We also added references. We hope we have addressed your point with this.

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5.2) issue has to do with the way the manuscript is written

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Thank you a lot for this important comment. We re-read all the manuscript and asked colleagues to proofread the article. They gave us interesting suggestions that helped us to improve different sections. We would be very pleased if you find it now easier to read it.

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5.3) sometimes is very difficult to follow the ideas the authors want to convey, an example can be found in the paragraph containing equation (5)

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Thank you very much for this information. We reviewed completely that equation (it is now equation (6) and equation (7)). We included a step-by-step procedure that describes the proposed algorithm and we added a schematic chart in Figure 3.

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REFERENCES

- Schomer, D.L.; Silva, F.L.D. *Niedermeyer's Electroencephalography: Basic Principles, Clinical Applications, and Related Fields*; Walters Klutter -Lippincott Williams & Wilkins, 2010.
- De Vos, M.; Debener, S. Mobile EEG: Towards brain activity monitoring during natural action and cognition. *International Journal of Psychophysiology* **2014**, *91*, 1–2.
- Wolpaw, J.; E., W. *Brain-Computer Interfaces: Principles and Practice*; Oxford University Press, 2012.
- Carlson, T.; del R. Millan, J. Brain-Controlled Wheelchairs: A Robotic Architecture. *IEEE Robotics & Automation Magazine* **2013**, *20*, 65–73.
- Guger, C.; Daban, S.; Sellers, E.; Holzner, C.; Krausz, G.; Carabalona, R.; Gramatica, F.; Edlinger, G. How many people are able to control a P300-based brain-computer interface (BCI)? *Neuroscience Letters* **2009**, *462*, 94–98.
- Lotte, F.; Faller, J.; Guger, C.; Renard, Y.; Pfurtscheller, G.; Lécuyer, A.; Leeb, R., Combining BCI with Virtual Reality: Towards New Applications and Improved BCI. In *Towards Practical Brain-Computer Interfaces: Bridging the Gap from Research to Real-World Applications*; Springer Berlin Heidelberg: Berlin, Heidelberg, 2013; pp. 197–220.
- Jure, F.; Carrere, L.; Gentiletti, G.; Tabernig, C. BCI-FES system for neuro-rehabilitation of stroke patients. *Journal of Physics: Conference Series* **2016**, *705*, 1–8.
- Clerc, M.; Bougrain, L.; Lotte, F. *Brain-computer interfaces, Technology and applications 2(Cognitive Science)*; ISTE Ltd. and Wiley, 2016.
- Riener, R.; Seward, L.J. Cybathlon 2016. *2014 IEEE International Conference on Systems, Man, and Cybernetics (SMC)* **2014**, pp. 2792–2794.
- Guger, C.; Allison, B.Z.; Lebedev, M.A. Introduction. In *Brain Computer Interface Research: A State of the Art Summary 6*; Springer, Cham, 2017; pp. 1–8.
- Huggins, J.E.; Alcaide-Aguirre, R.E.; Hill, K. Effects of text generation on P300 brain-computer interface performance. *Brain-Computer Interfaces* **2016**, *3*, 112–120.
- Alvarado-González, M.; Garduño, E.; Bribiesca, E.; Yáñez-Suárez, O.; Medina-Bañuelos, V. P300 Detection Based on EEG Shape Features. *Computational and Mathematical Methods in Medicine* **2016**, pp. 1–14.

- Yamaguchi, T.; Fujio, M.; Inoue, K.; Pfurtscheller, G. Design Method of Morphological Structural Function for Pattern Recognition of EEG Signals During Motor Imagery and Cognition. Fourth International Conference on Innovative Computing, Information and Control (ICICIC), 2009, pp. 1558–1561.
- Berger, S.; Schneider, G.; Kochs, E.; Jordan, D. Permutation Entropy: Too Complex a Measure for EEG Time Series? *Entropy* 2017, Vol. 19, Page 692 **2017**, 19, 692.
- Ramele, R.; Villar, A.J.; Santos, J.M. BCI classification based on signal plots and SIFT descriptors. 4th International Winter Conference on Brain-Computer Interface, BCI 2016; IEEE: Yongpyong, 2016; pp. 1–4.
- Hartman, a.L. *Atlas of EEG Patterns*; Vol. 65, Lippincott Williams & Wilkins, 2005.
- Boiman, O.; Shechtman, E.; Irani, M. In defense of nearest-neighbor based image classification. *26th IEEE Conference on Computer Vision and Pattern Recognition, CVPR* **2008**.
- Farwell, L.A.; Donchin, E. Talking off the top of your head: toward a mental prosthesis utilizing event-related brain potentials. *Electroencephalography and clinical neurophysiology* **1988**, 70, 510–23.
- Knuth, K.H.; Shah, A.S.; Truccolo, W.A.; Ding, M.; Bressler, S.L.; Schroeder, C.E. Differentially Variable Component Analysis: Identifying Multiple Evoked Components Using Trial-to-Trial Variability. *Journal of Neurophysiology* **2006**, 95, 3257–3276.
- Hu, L.; Mouraux, A.; Hu, Y.; Iannetti, G.D. A novel approach for enhancing the signal-to-noise ratio and detecting automatically event-related potentials (ERPs) in single trials. *NeuroImage* **2010**, 50, 99–111.
- Renard, Y.; Lotte, F.; Gibert, G.; Congedo, M.; Maby, E.; Delannoy, V.; Bertrand, O.; Lécuyer, A. OpenViBE: An Open-Source Software Platform to Design, Test, and Use BrainComputer Interfaces in Real and Virtual Environments. *Presence: Teleoperators and Virtual Environments* **2010**, 19, 35–53.
- Krusienski, D.J.; Sellers, E.W.; Cabestaing, F.; Bayoukh, S.; McFarland, D.J.; Vaughan, T.M.; Wolpaw, J.R. A comparison of classification techniques for the P300 Speller. *Journal of Neural Engineering* **2006**, 3, 299–305.
- Liang, N.; Bougrain, L. Averaging techniques for single-trial analysis of oddball event-related potentials. *4th International Brain-Computer* **2008**, pp. 1–6.
- Bresenham, J.E. Algorithm for computer control of a digital plotter. *IBM Systems Journal* **1965**, 4, 25–30.
- Lowe, G. SIFT - The Scale Invariant Feature Transform. *International Journal* **2004**, 2, 91–110.
- Vedaldi, A.; Fulkerson, B. VLFeat - An open and portable library of computer vision algorithms. *Design* **2010**, 3, 1–4.
- Riccio, A.; Simione, L.; Schettini, F.; Pizzimenti, A.; Inghilleri, M.; Belardinelli, M.O.; Mattia, D.; Cincotti, F. Attention and P300-based BCI performance in people with amyotrophic lateral sclerosis. *Frontiers in Human Neuroscience* **2013**, 7, 732.
- Brunner, C.; Blankertz, B.; Cincotti, F.; Kübler, A.; Mattia, D.; Miralles, F.; Nijholt, A.; Otal, B. BNCI Horizon 2020 Towards a Roadmap for Brain / Neural Computer Interaction. *Lecture Notes in Computer Science* **2014**, 8513, 475–486.
- Schalk, G.; McFarland, D.J.; Hinterberger, T.; Birbaumer, N.; Wolpaw, J.R. BCI2000: a general-purpose brain-computer interface (BCI) system. *IEEE transactions on bio-medical engineering* **2004**, 51, 1034–43.
- Rao, R.P.N. *Brain-Computer Interfacing: An Introduction*; Cambridge University Press: New York, NY, USA, 2013.
- Polich, J. Updating P300: An integrative theory of P3a and P3b, 2007.
- Casarotto, S.; Bianchi, A.; Cerutti, S.; Chiarenza, G. Dynamic time warping in the analysis of event-related potentials. *IEEE Engineering in Medicine and Biology Magazine* **2005**, 24, 68–77.
- Ouyang, G.; Hildebrandt, A.; Sommer, W.; Zhou, C. Exploiting the intra-subject latency variability from single-trial event-related potentials in the P3 time range: A review and comparative evaluation of methods. *Neuroscience and Biobehavioral Reviews* **2017**, 75, 1–21.
- Van Drongelen, W. *Signal processing for neuroscientists: an introduction to the analysis of physiological signals*; Academic press, 2006.
- Nijboer, F.; Broermann, U. Brain Computer Interfaces for Communication and Control in Locked-in

Patients. In *Graimann B., Pfurtscheller G., Allison B. (eds) Brain-Computer Interfaces. The Frontiers Collection.*; Springer Berlin Heidelberg, 2009; pp. 185–201.