

## CASE REVIEW

### PNES – Case Review in Neuroimaging Context

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#### Abstract

Psychogenic nonepileptic seizures (PNES) – episodic failure of behavioural control – create a heterogeneous group of diseases. There is no common approach in the diagnostics of mentioned diseases. The absence of the common diagnostic approach contributes to a significant delay in the diagnostic process, often up to 7 years. Very important part of this process is neuroimaging. There are quite inhomogeneous results in up to date literature review whether in context of morphology or function. The main aim of the presented paper is a review of specific cases in neuroimaging context to collect data for further research. For the review the authors introduce 2 case studies of patients with PNES on different psychopathological background and their representation via EEG – current density maps using eLORETA software. On the basis of the studied literature and analyzed cases, conclusions are drawn and suggestions for further investigation are proposed.

#### Keywords

PNES, EEG, eLORETA, current density maps, neuronal networks

#### 1. Introduction

Current bio-psycho-social model of PNES depicts the multifactoriality of the disease. Based on previous research (Hingray et al., 2011; Brown & Reuber, 2016; Green, Norman & Reuber, 2017; Dworetzky & Baslet, 2017) certain life events can be considered as causal in the formation of psychogenic nonepileptic seizures. These events include severe trauma in childhood (24 – 58 % sexual abuse, child abuse & neglect (CAN) syndrome); adult life trauma (rape, physical abuse, severe accidents); relevant emotional loss (decease of a close person); acute or situational stress (dissonance in the family, developmental unreadiness, insufficient maturity). Motoric and behavioural expressions of PNES are different in different patients (remarkable motoric activity, wilt or falling, vocalisation or ictal injury). Seizures appear typical in home setting when there are other people present, mostly during the day, typically lasting longer than 2 minutes. Psychogenic nonepileptic seizure is described by patients as out of their wilful control. (Brown & Reuber, 2016). Prevalence of PNES is approximately 33 cases on 100 000 people per year. (Green et al., 2017). Manifestation of PNES occurs mostly in the 2nd or 3rd decennium (Asadi-Pooya et al., 2014), predominantly by females (approx. ¾ of all patients) and people with lower education (Brown & Reuber, 2016) or mental issues (Kanemoto et al., 2017). As a comorbidity with epilepsy, a PNES is reported in 10 to 30% of patients. The most common psychiatric conditions in these patients are depressive and anxious disorder, personality disorders (mostly borderline personality disorder) and other somatoform and dissociative disorders. (Dworetzky & Baslet, 2017). Neurologic comorbidities in PNES patients are migraine, other types of headaches, chronic pain, weakness, lowered sensitivity (Mintzer, 2015). Psychopharmacologic treatment is indicated when the patient is diagnosed along with

panic disorder, depression, anxious-depressive disorder or psychosis. Most commonly used drugs are antidepressants of SSRI group. Benzodiazepines can be used temporarily. (Dworetzky & Baslet, 2017). The mere fact, that from 20 % to 25 % of pharmacoresistent epilepsy patients suffer from PNES or combination of both, suggests that the process of PNES diagnosis requires improvement. (Schachter & LaFrance, 2010). Proper and early diagnosis can prevent significant damage to iatrogen, the inadequate treatment of which can lead to the death of the patient (Asadi-Pooya et al., 2014). Foreign research includes the financial burden of the inadequate therapy. (Magee et al., 2014). Absolute specificity in PNES diagnostic is fundamental taking into account the diagnostic delay reaching up to 5 to 10 years (Alessi & Valente, 2014). „Golden standard“ for the definitive diagnosis of PNES is video EEG monitoring. Psychologic examination consists of neuropsychologic assessment, projective methods, personality questionnaires and tests for simulation and effort. Basics of a neuropsychological assessment are detailed diagnostic interview, personal history mapping overloading moments and traumatic life experience. It can be stated that in PNES patient group there is maladaptive emotional regulation in their perception (Novakova et al., 2015). Psychogenic nonepileptic seizures can reflect a deficit in recognition, thinking and emotion description. From this perspective can be PNES seen as symptom of emotional discomfort or arousal which person incorrectly attaches to physical cause or as a valve for releasing unrecognised emotional stress. (Brown & Reuber, 2016). Patients with PNES represent a heterogeneous group with various manifestations of psychiatric symptoms and somatic discomfort. Therefore it is less probable that this disorder would be associated with one model of emotional regulation and procession. This is very important for the need of individual approach in psychotherapy with these patients. . Therapy of this

patient group demands a multidisciplinary approach beginning with a sensitive way of communicating the diagnose, terminating the anticonvulsive medication, treating the comorbid psychiatric symptoms and early beginning of psychotherapy as the main therapy approach. (Dworetzky & Baklet, 2017). Initial psychoeducation of patients and their relatives is a very important step in the therapy process. (Chen et al., 2014; Mayor et al., 2013). Retrospective studies (for example Duncan et al., 2016) show that approximately one third of patients after appropriate diagnose communication were within 3-6 months afterwards without other intervention seizure free. Positive prognostic factors that affect the overall condition are: lower age, shorter duration of the disease, identification of the causal trauma, absence of epileptic seizures, absence of psychiatric comorbid condition, higher socioeconomic state, IQ at least in norm range. Predictors of seizure persistence are: depression, personality disorders, drug abuse. (Dworetzky & Basket, 2017). Understanding of the conditions, especially by the patient, is inevitable for the efficiency of the following therapy process. The anticipation of the fear from the seizure and the internalisation of the suffering identity can cause more harm than the seizure itself. Understanding the diagnosed disease from the patient's point of view is essential for the seizure frequency reduction and regaining control over patients own life. (LaFrance & Wincze, 2015).

## 2. Neuroimaging of PNES

In summary the most neuroimaging studies finds in PNES patients multifocal submacroscopic abnormalities associated with specific neuropsychiatric traits. The common denominator in this disease is a neuronal network dysfunction (Szaflarski 2018).

Despite of the fact that the results of recent neuroimaging studies impose inconclusive they are an existence proof of connection between structural and functional brain abnormality which corresponds with the bio-psycho-social model of PNES formation. Findings on brain imaging – neurobiological correlates of this disease – contradict their purely psychogenic aetiology. Further important fact is heterogeneity of PNES patient group and therefore essential subpopulation selection which need to be seen separately. (Mcsweeney 2017)

Approximately 30% patients seen nowadays by neurologists are concluded as idiopathic, roughly 18% get diagnosed with dissociative =functional movement disorder (FMD). In the epilepsy patient group are at the end of the day around 7 to 20% diagnosed with PNES (Perez, 2015).

### 2.1 Neuronal circuits

The key structures in neuroimaging of functional movement disorder – PNES included – are the prefrontal, insular, amygdale parts, posterior parietal cortex/temporoparietal junction (PPC/TPJ), premotor parts and their connections. Medial prefrontal cortex, especially anterior cingulate cortex (ACC) shows functional and structural changes in PNES patients. ACC can be divided into ventral – affective and dorsal – cognitive part. Subgenual and perigenual areas take part in emotion regulation and fear extinction. The dorsal part of ACC mediates emotion expression and their cognitive control. There is a connection between ACC and n. accumbens/ventral caudate, ventral globus pallidum and mediodorsal and ventral thalamic nuclei. Subgenual and perigenual ACC are interconnected to amygdale whereas the

dorsal ACC is in connection with lateral prefrontal parts including supplementary motor area (SMA) and dorsolateral prefrontal cortex (dlPFC). Hypoactivation of subgenual and perigenual ACC imposes reduction of top-down activity regulation of amygdale. This finding corresponds to its hyperactivity in patients with fibromuscular dysplasia (FMD) in affective and motoric tasks.

Recent neuroimaging results show a connection between dissociation and increased resting-state perigenual ACC activity. This seems important taking into account that psychotraumatization in childhood, PTSD, dissociation are very common in patients with PNES.

From the structural point of view – volumetric findings in patients with PNES show reduction of grey matter in dorsal ACC and SMA areas, reduced cortical thickness in paracentral and precentral gyrus. Described structural alteration in dorsal ACC-SMA parts can lead to abnormal behavioural expression. Reciprocal ventral and dorsal ACC connection can mediate interactions between impaired emotional regulation and expression (Perez, 2015).

Recent studies (Szaflarski, 2018) imply causal relationship between altered brain connectivity and formation and maintenance of PNES (Figure 1).

Figure 1 Neuronal network scheme by PNES formation. TPJ, PI, dAI vAI P/ACC LN VMPFC VLPFC (Szaflarski, 2018)

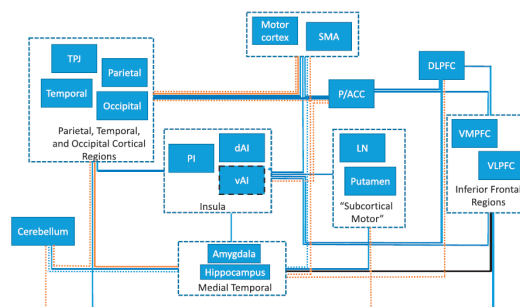


FIGURE. Schematics of possible nodes of the network underlying development and maintenance of psychogenic nonepileptic seizures (PNES) derived from structural neuroimaging studies of patients with PNES and other functional neurological disorders/conversion disorders (FNDs/CDs) and possible connections between those nodes derived from functional and structural connectivity studies. Colors indicate from which studies the connections were derived: BLUE = resting state connectivity; BLACK = structural findings/structural connectivity (dashed line around vAI is derived from the meta-analysis of some of the neuroimaging studies of PNES 6); ORANGE = task-related functional connectivity; DASHED BLUE = connectivity noted in other FNDs/CDs, in addition to the connectivity in PNES; DASHED ORANGE = task-related functional connectivity (various tasks).

## 3. Material and methods

In this chapter, study material collected from 2 cases of patients diagnosed per exclusionem with PNES is presented, as well as the description of the methods for the processing of obtained data.

### Case No.1.

Patient – female, 21yrs.

**Personal History:** Seizures in diff. Dg. Process – PNES, Canabinoid Misuse, St.p. TS with medication 02/20, St.p. Nephrostomy

**Family History:** no neuropsychiatric burden found

**Social History:** parents have lived separated since she was 3 years old, father pays alimony, communication with father via grandfather, patient studies 3<sup>rd</sup> year of opera singing, acting and piano at musical academy, after the 1<sup>st</sup> year of studies she changed the school for bullying; for the last couple of months she has lived in a rented flat, after school classes she works as a student job receptionist in a fitness centre, she lives off the alimony and money she earns, after paying her bills she is left over with ca 60 euro for a



month, as 16 yrs old she had a relationship, which ended with an unpleasant experience in a park where her drunk boyfriend sexually harassed her and gossiped on her afterwards, at the time being she has had a 19yrs old boyfriend for 4 months, she is happy in this relationship.

**Seizure semiology:** Seizure starts with a sudden fall, the patient imposes atonic, lays without moving with closed eyes and calm breathing. Ca after 2.5 min she suddenly sits up, begins to hyperventilate and cough, cries for the next 3-4 minutes until she calms down.

#### **Diff. Dg. Procedure:**

**EEG**—background activity occipital, less regular with frequency 8-9 Hz, amplitude up to do 50uV, less modulated, Berger reaction is present. In frontal electrodes beta activity, which does not change by hyperventilation, rarely electrode and muscle artefacts. By the end of recording - signs of vigilance oscillation. In ECG is regular rhythm. EEG finding - without specific abnormality, without signs of focal injury, vigilance oscillations. Recommendation: Control according to clinical course, long term EEG.

**Long-term video-EEG monitoring** - Day: EEG basic activity occipital less regular, with frequency 9-10 Hz, amplitude up to do 50 uV, less modulated. In frontal electrodes beta activity, frequent muscle and ocular artifacts, patient lays herself to sleep stands afterward up, sits and plays with her mobile phone. Often IRDA discharges, bilaterally FT. At 14:16 she falls asleep until 14:51 with appearance of K-complexes, after waking up she repeatedly falls asleep until the end of recording at 19:31. In video recording IRDA without ocular artefacts. Night: awake state until falling asleep at 21:47 with typical course of nonREM and REM sleep. Conclusion: EEG without specific abnormality.

**Brain MRI:** no significant morphologic abnormality

**Psychodiagnostic Examination:** ROR, Figure Drawing Test, Baumtest, Association experiment, WAIS-R; Personality in maturation process, at the moment immature with average intellect. Autonomy tendency dominates by difficult self-enforcement. Orientation in own experiences imposes limited. As problematic seems the acceptance of the female role.

**Conclusion:** PNES based on PTSD

#### **Case No.2**

Patient – female, 41 yrs.

**Personal history:** Brain MRI 12/2017 - P1 part ACP bilat. Hypoplasia, since 2017 diagnosed with hyperventilation tetany – seizure regulation by breathing, subsequently no effect of Diazepam per Rectum, EEG after sleep deprivation - negative result, 03/2020 neurological hospitalisation with record of tetanic seizure with atonoid movements of upper and lower extremities, opisthotonus – concluded as susp. PNES; Vertebroalgalic syndrome of cervical spine, Micronodular Struma, St.p. Appendectomy, St.p. Tonsilectomy, St.p. Curretage for a Myoma, St.p. H.pylori Eradication

**Family History:** Father committed a suicide by strangulation in his 47 yrs. "he felt like he cannot earn for the living of the family", brother of the father with posttraumatic epilepsy, otherwise no neuropsychiatric burden found.

**Social History:** she considers her childhood as very good, her parents didn't argue, she has 2yrs younger sister, high school – hairdresser, she didn't finish the school because she got a job, 1 marriage which lasted 2 yrs, her husband didn't have much interest in the family, he does not have interest on their daughter (16yrs), after divorce she returned back to her mother where she opened her own hairdresser studio. She met her 2<sup>nd</sup> husband 8 years ago, have been married for 5 years, her 2<sup>nd</sup> husband has 2 children from an earlier marriage, problematic relationship with his ex-wife. These children are staying with the patient and her 2<sup>nd</sup> husband on weekends and all holidays. They live in a house of her husband parents. They have altogether good relationships. As an investment she bought her own flat.

**Seizure semiology:** series of habitual movements (confirmed by the patient who communicates easily during the seizure), semiology of incongruent and inconsistent movements – atetoid movements in quadrudistribution of crescendo-decrescendo type, turning over, interponed voluntary movements – patient makes up her pillow, recorded opisthotonus, duration several tens of minutes.

#### **Diff.-dg.Procedure:**

**Long-term video-EEG monitoring** – nocturnal recording – modulated basic activity with amplitude up to 9 Hz, in sleep stages I-III, 4 cycles of REM and nonREM, no epileptic activity in EEG or in video recording. Regular rhythm found in ECG. Daily record: during recording at 6:53 a.m. a seizure – movement in lying with opisthotonus indicated, movements of upper and lower limbs, in EEG muscle and electrode artifacts, before or after seizure no epileptic activity. The seizure was terminated by conversation. Seizure manifestation imposes as psychogenic. No epileptic process in EEG or clinically proven. Recommendation: Control according to clinical course, psychological treatment.

**Psychodiagnostic examination. – Personality, Intellect** - ROR, AE, PSSI, WAIS-R; Personality with dominant histrionic traits. The actual intellectual performance is in lower average range. The performance appears underneath the ability level. A good readiness and concentration ability were observed by easy tasks. Subjectivistic, intuitive thinking. Critical-analytical thinking is reduced. Contact with reality is preserved, however, there were some distortions based on emotional repression observed. Tendency to negative emotion attenuation and self-focusing affections are present.

**Conclusion:** Dissociative seizures based on personality disorder.

### **3.1 EEG Data Acquisition and Postprocessing**

EEG Data recording was performed on a standard 19-channel digital EEG amplifier BrainScope EADS with 19 Ag/AgCl surface electrodes placed according to the international 10/20 system. The data-sampling rate was 1000 Hz and the acquired signals were filtered with digital high- and low-pass filters at 0.15 and 70 Hz, respectively. Our selected data were a part of the video-EEG examination in duration of several hours.

Prior to data analysis artifact detection was visually performed to exclude all epochs containing eye blink, eye-rolling artifact, head movements, muscle artifacts. Thus, from each EEG at least 60 sec of resting state, artifact-free, and highly reliable data were subjected to further analysis. The number of epochs, as well as the length of the samples processed, did not differ between the patients and the healthy control.

Data analysis was performed using the exact low-resolution electromagnetic tomography – eLORETA an inverse solution technique that estimates the intracranial distribution of electrical activity (current density) in the cortex based on a three shell spherical head model co-registered with Talairach coordinates (Pascual-Marqui, 2002). We used the eLORETA-Key software (Key Institute for Brain-Mind Research, Zurich, Switzerland), available at <http://www.uzh.ch/keyinst/loreta.htm>. Using the eLORETA transformation matrix, cross spectra of each subject and for each frequency band were transformed to eLORETA files. This resulted in a corresponding 3D cortical distribution of the electrical neuronal generators for each subject. The computed eLORETA images reflect the cortical current density distribution in 6,239 voxels with a spatial resolution of  $5 \times 5 \times 5$  mm. The eLORETA algorithm has no localization of bias even in the presence of structured noise, which allows us to increase the localization accuracy, compared to the previous version of sLORETA (54). The advantage of eLORETA is that it belongs to a reference-free method of EEG analysis, therefore, determining the source distribution for EEG data is not affected by the selected electrode reference. Current density values were computed in eight frequency bands delta (0.5–3.5 Hz), theta (4–8 Hz), alpha-1 (8.5–10 Hz), alpha-2 (10.5–12 Hz), beta-1 (12.5–18 Hz), beta-2 (18.5–21 Hz), beta-3 (21.5–30 Hz), and omega (0.5–30 Hz).

### 3.2 Statistical Analysis

For the statistical analysis eLoreta data were used. We performed no normalisation of the data. Our data were approached as paired groups. No baseline correction was computed. Tests were performed for all timeframes/frequencies. For the analysis the F-Test – Log of F-Ratio was chosen. In the eLORETA analyses, the localization of the differences in baseline activity between the group of responders and non-responders was assessed using a voxel-by-voxel unpaired t-test of the eLORETA images, based on the power of estimated electric current density. In the resulting statistical three-dimensional images, cortical voxels showing significant differences were identified using a nonparametric approach (statistical nonparametric mapping or SnPM) via randomizations. This randomization strategy determined the critical probability threshold values for the actual observed t-values, with correction for multiple comparisons across all voxels and all frequencies. A total of 5,000 permutations were used to determine significance for each randomization test.

### 5. Discussion

In our patient with PNES based on PTSD we found significant baseline activity difference - decrease in primary somatosensory cortex (postcentral gyrus) which plays important role in each stage of emotional processing, including identification of emotional significance in a stimulus, generation of emotional states, and regulation of emotion. Furthermore there was decrease of baseline activity found in area involved in the analysis and integration of higher order visual, auditory and somesthetic information, area in parietal cortex where translation and interpretations of visual impressions take place. Increased baseline activity was found in subgenual area, part of the ventromedial prefrontal cortex implicated in a variety of social, cognitive, and affective functions that are commonly disrupted in mental illness.

Patient with PNES based on personality disorder had their baseline activity significantly decreased in postcentral gyrus of primary

somatosensory cortex taking part in emotional processing, in primary motoric cortex, in area in aprietal cortex playing role in visuo-motor coordination and in secondary visual cortex. Baseline activity increase was seen in associative visual cortex.

As shown by our results it is inevitable to take into account the psychopathological background of PNES formation. It can be stated that precisely this aspect is a condition by which the patient population can be subdivided where certain similarity of results can be expected.

Further moment would be targeting of neuroimaging - neuronal connectivity for instance – according to up to date neuroimaging results.

One quite remarkable design for subsequent research would be comparison of neuroimaging of PNES patient with a group of patients with the same psychiatric condition without seizure manifestation. Perhaps, it would be this approach which will bring us closer to the neurophysiological correlate of PNES.

## 4. Results

### Case No.1

Baseline activity decrease in Alpha 1 frequency band in BA 2 – postcentral gyrus, parietal lobe (Fig.1), alpha 2 in BA - 7 – superior parietal lobule, parietal lobe (Fig.2), beta 2 in BA 19 – precuneus (Fig. 3), parietal lobe, beta 3 in BA 19 – precuneus, parietal lobe (Fig 4.). Baseline activity increase in delta frequency band in BA 25 – anterior cingulate, limbic lobe (Fig.5).

Figure 2 Alpha 1

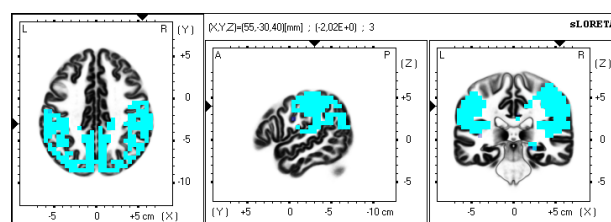


Figure 3 Alpha 2

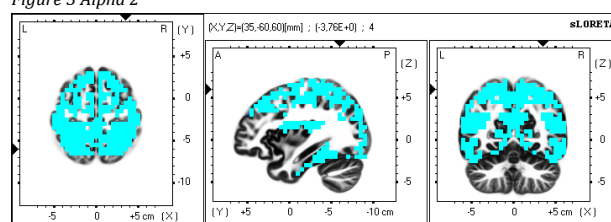


Figure 1 Beta 2

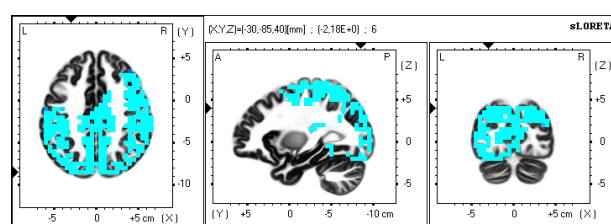




Figure 5 Beta 3

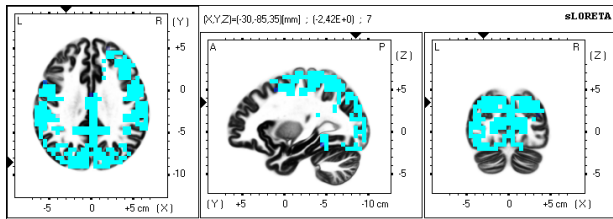
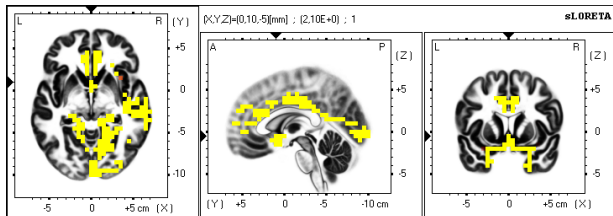


Figure 6 Delta



### Case No.2

Baseline activity decrease in delta frequency band in BA 7 – precuneus, parietal lobe (Fig.6), theta in BA 3 – postcentral gyrus, parietal lobe (Fig.7), alpha 2 frequency band in BA 4 – precentral gyrus, frontal lobe (Fig.8), beta 1 in BA 4 – precentral gyrus, frontal lobe (Fig. 9), beta 2 in BA 4 – precentral gyrus, frontal lobe (Fig. 10), beta 3 in BA 18 – cuneus, occipital lobe (Fig 11.).

Baseline activity increase in alpha 1 frequency band in BA 19 – lingual gyrus, occipital lobe (Fig. 12).

Figure 7 Delta

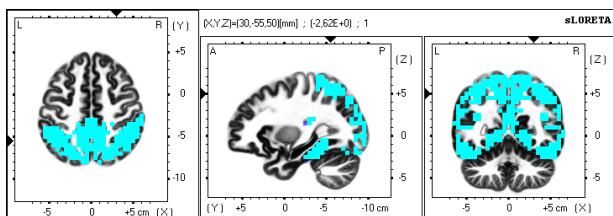


Figure 2 Theta

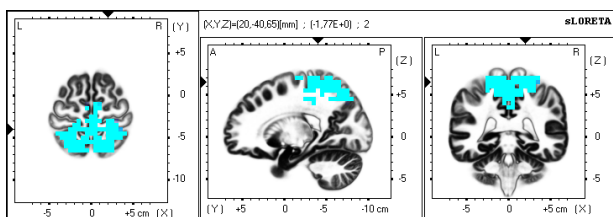


Figure 3 Alpha 2

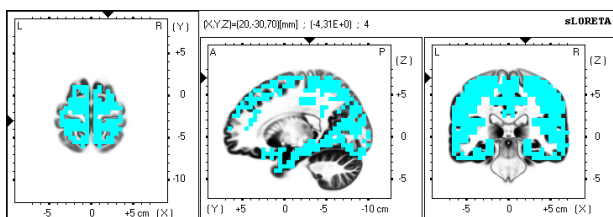


Figure 4 Beta 1

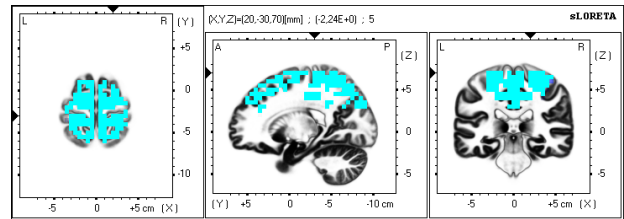


Figure 5 Beta 2

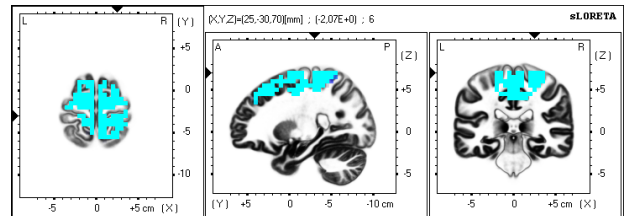


Figure 6 Beta 3

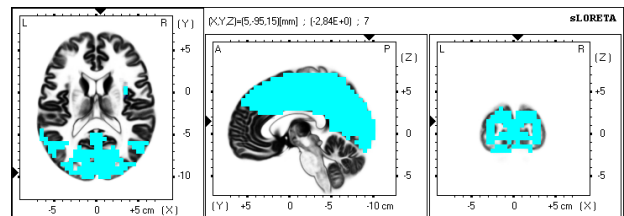
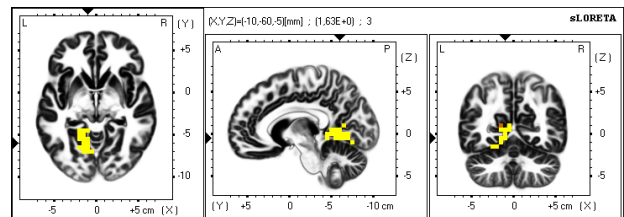


Figure 7 Alpha 1



## 6. Conclusion

Neuroimaging studies present the next step in understanding of formation and maintenance processes of PNES. Considering the multifactoriality of this disease the subdivision of patient group into subpopulations depending on psychopathological background appears inevitable. Some other diagnostic tools like psycho diagnostics seem quite helpful. However, PNES cannot be seen as a homogenous disease which corresponds with the heterogeneity of results found in recent neuroimaging studies. In this context, the authors of the presented paper agreed that further comparison of neuroimaging of PNES patients with a group of patients with the same psychiatric condition without seizure manifestation and targeting of neuroimaging are crucial for better understanding of the PNES disease and optimization of the treatment options for the affected group of patients.

## References

- Hingray, C., Maillard, L., Hubsch, C., Vignal, J. P., Bourgoignon, F., Laprevote, V., Schwan, R. (2011). Psychogenic nonepileptic seizures: Characterization of two distinct patient profiles on the basis of trauma history. *Epilepsy & Behavior*, 22, 532–536.



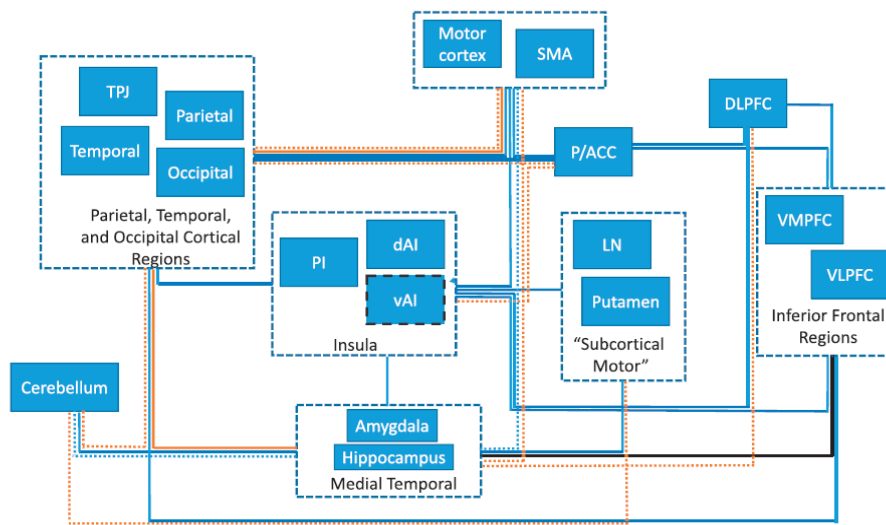
- Mayor, R., Brown, R. J., Cock, H., House, A., Howlett, S., Smith, P., . . . Reuber, M. (2013). A feasibility study of a brief psycho-educational intervention for psychogenic nonepileptic seizures. *Seizure*, 22, 760–765.
- Green, B., Norman, P., & Reuber, M. (2017). Attachment style, relationship quality, and psychological distress in patients with psychogenic nonepileptic seizures versus epilepsy. *Epilepsy & Behavior*, 66, 120–126.
- Dworetzky, B. A., & Baslet, G. C. (2017). Psychogenic nonepileptic seizures. Toward the integration of care. New York, The United States of America: Oxford University Press.
- Asadi-Pooya, A. A., Emami, Y., Emami, M., & Sperling, M. R. (2014). Prolonged psychogenic nonepileptic seizures or pseudostatus. *Epilepsy & Behavior*, 31, 304–306.
- Kanemoto, K., LaFrance Jr., W. C., Duncan, R., Gigineishvili, D., Park, S. P., & Tandokoro, Y. (2017). PNES around the world: Where we are now and how we can close the diagnosis and treatment gaps—an ILAE PNES Task Force report. *Epilepsia Open*, 2(3), 307–316.
- Mintzer, S. (2015). Driven to tears: epilepsy specialists and the automobile. *Epilepsy Currents*, 15(5), 279–282.
- Schachter SC, LaFrance WC Jr (Eds). *Gates and Rowan's Nonepileptic Seizures*. 3rd Ed. Cambridge; New York: Cambridge University Press, 2010
- Magee, J. A., Burke, T., Delanty, N., Pender, N., & Fortune, G. M. (2014). The economic cost of nonepileptic attack disorder in Ireland. *Epilepsy & Behavior*, 33, 45–48.
- Mayor, R., Brown, R. J., Cock, H., House, A., Howlett, S., Smith, P., . . . Reuber, M. (2013). A feasibility study of a brief psycho-educational intervention for psychogenic nonepileptic seizures. *Seizure*, 22, 760–765.
- Alessi R, Valente KD. Psychogenic nonepileptic seizures: should we use response to AEDS as a red flag for the diagnosis?. *Seizure*. 2014;23(10):906-908. doi:10.1016/j.seizure.2014.07.016
- Novakova, B., Howlett, S., Baker, R., & Reuber, M. (2015). Emotion processing and psychogenic non-epileptic seizures: A cross-sectional comparison of patients and healthy controls. *Seizure*, 29, 4–10.
- Chen, H., Ding, J., An, D., Liao, W., Wu, G., Xu, Q., Zhou, D., (2014). Abnormal functional connectivity density in psychogenic non-epileptic seizures. *Epilepsy Res*. 108, 7, 1184-1194.
- LaFrance, W. C., & Wincze J. P. (2015). *Treating Nonepileptic Seizures: Therapist Guide (Treatments That work)*. New York, The United States of America: Oxford University Press.
- Szaflarski J.P., LaFrance C.W., (2018), Psychogenic Nonepileptic Seizures (PNES) as a Network Disorder – Evidence From Neuroimaging of Functional (Psychogenic) Neurological Disorders, *Epilepsy Currents*, Vol. 18, No. 4, pp. 211–216
- Mcsweeney, M., Reuber, M., Hoggard, N., Levita, L., (2018), Cortical thickness and gyrification patterns in patients with psychogenic non-epileptic seizures, *Neuroscience Letters* 678 (2018) 124–130
- Perez, D.L., Dworetzky, B.A., Dickerson, B.C., Leung, L., Cohn, R., Baslet, G., Silbersweig, D.A., (2015), An Integrative Neurocircuit Perspective on Psychogenic NonEpileptic Seizures and Functional Movement Disorders: Neural Functional Unawareness, *Clin EEG Neurosci*. 46(1): 4–15
- Pascual-Marqui RD, Esslen M, Kochi K, Lehmann D. Functional imaging with low-resolution brain electromagnetic tomography (LORETA): a review. *Methods Find Exp Clin Pharmacol* (2002) 24 Suppl C:91–5.
- Pascual-Marqui RD. Standardized low-resolution brain electromagnetic tomography (sLORETA): technical details. *Methods Find Exp Clin Pharmacol* (2002) 24 Suppl D:5–12

**Received: 12.1. 2021**

**Accepted after review: 2.2. 2021**

**Published on-line: 9.2. 2021**

Figure 1 Neuronal network scheme by PNES formation. TPJ, PI, dAI vAI  
P/ACC LN VMPFC VLPFC (Szaflarski, 2018)



**FIGURE.** Schematics of possible nodes of the network underlying development and maintenance of psychogenic nonepileptic seizures (PNES) derived from structural neuroimaging studies of patients with PNES and other functional neurological disorders/conversion disorders (FND/CDs) and possible connections between those nodes derived from functional and structural connectivity studies. Colors indicate from which studies the connections were derived: BLUE = resting state connectivity; BLACK = structural findings/structural connectivity (dashed line around vAI is derived from the meta-analysis of some of the neuroimaging studies of PNES; 6); ORANGE = task-related functional connectivity; DASHED BLUE = connectivity noted in other FNDs/CDs, in addition to the connectivity in PNES; DASHED ORANGE = task-related functional connectivity (various tasks).

Figure 8 Alpha 1

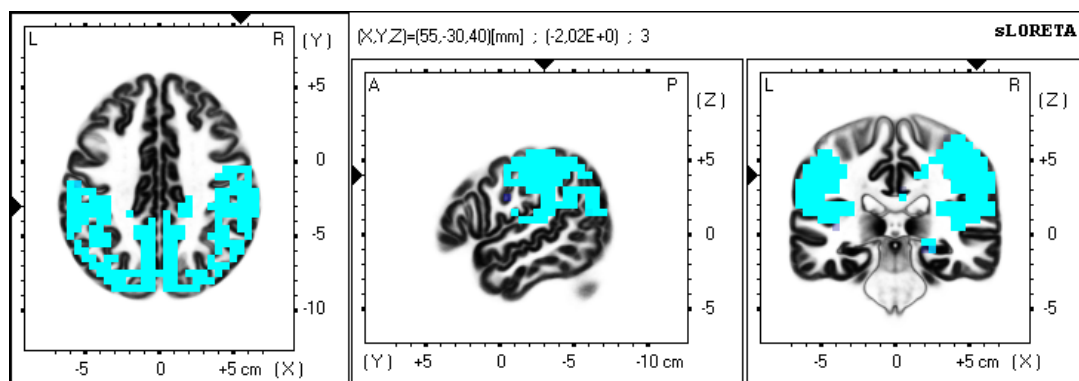


Figure 9 Alpha 2

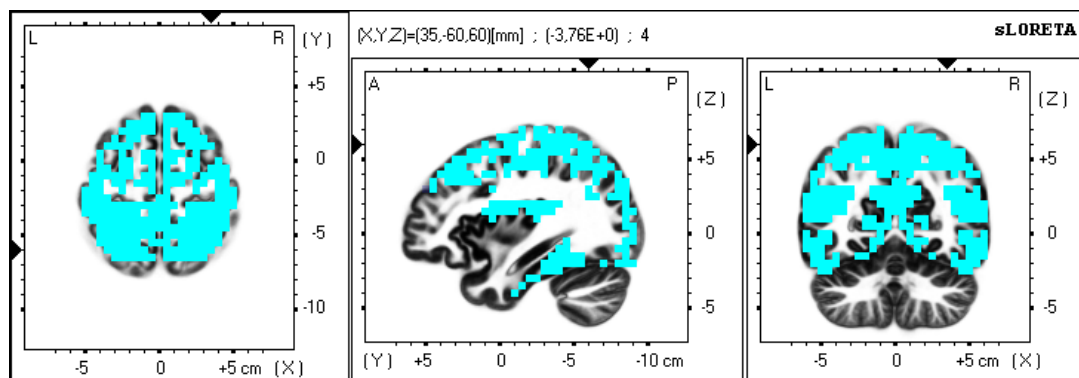


Figure 10 Beta 2

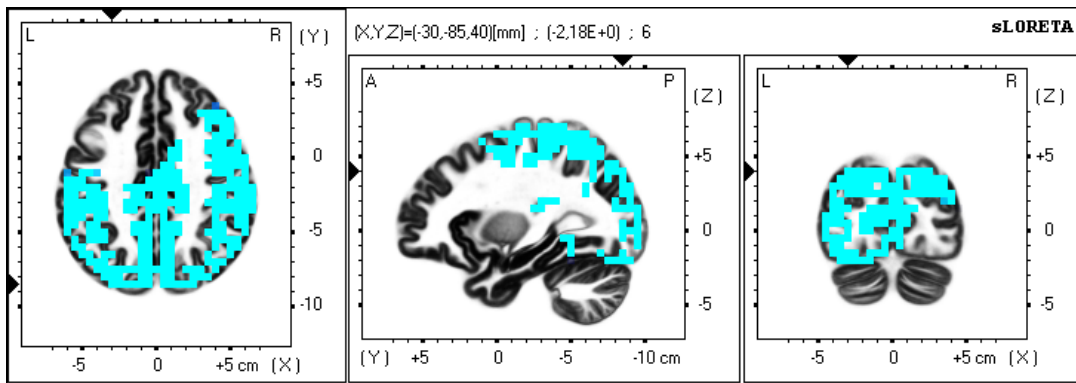


Figure 5 Beta 3

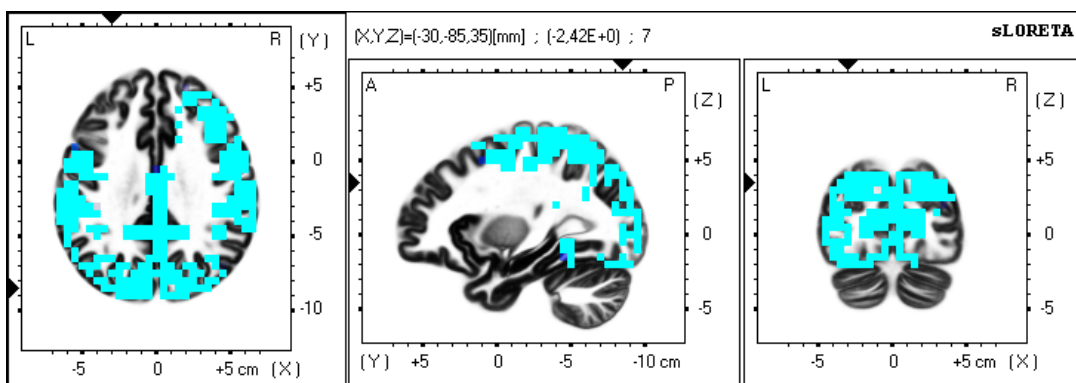


Figure 11 Delta

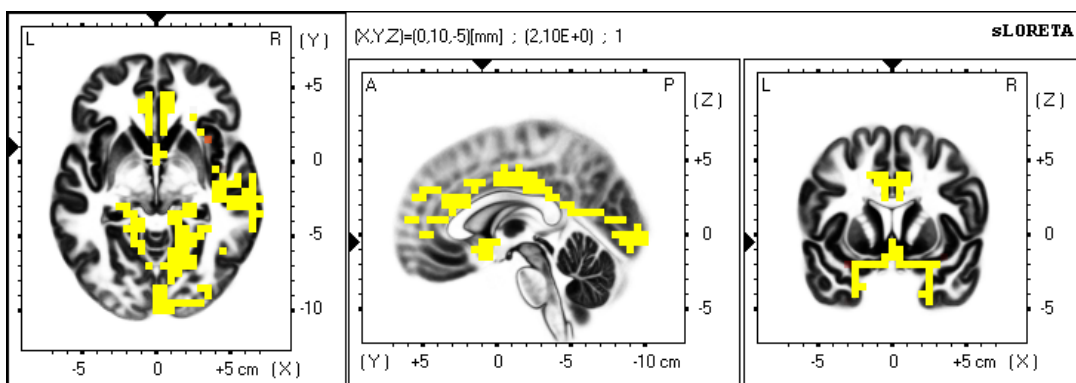


Figure 7 Delta

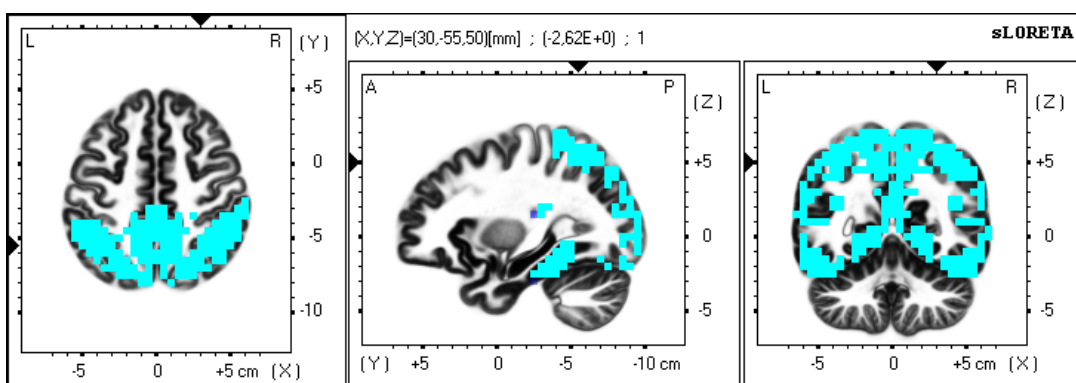




Figure 12 Theta

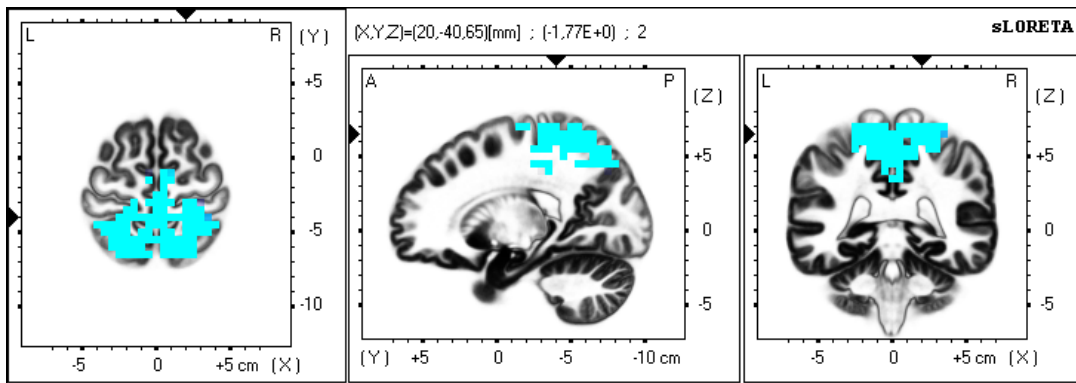


Figure 13 Alpha 2

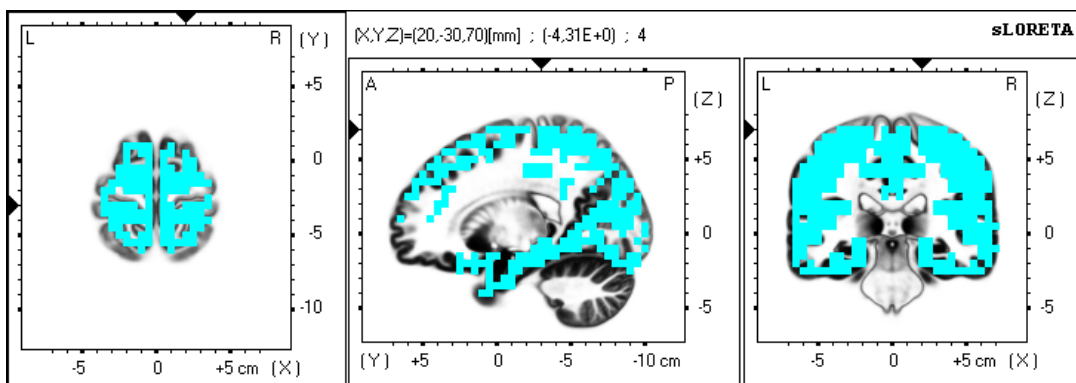


Figure 14 Beta 1

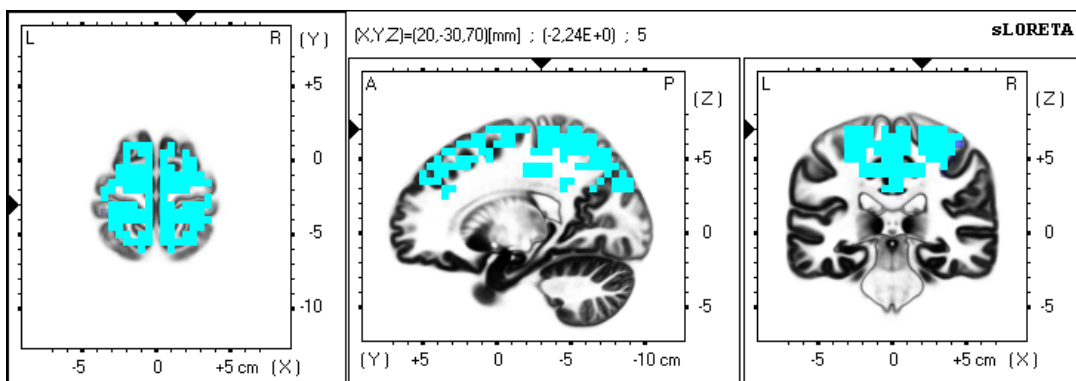


Figure 15 Beta 2

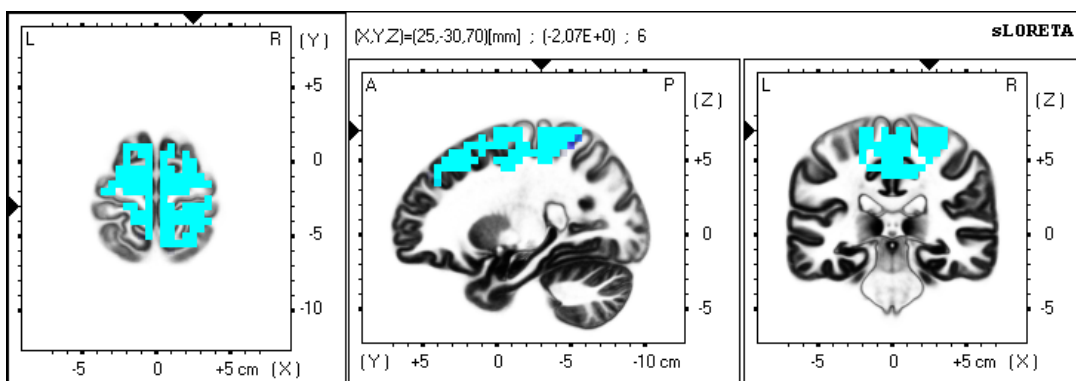


Figure 16 Beta 3

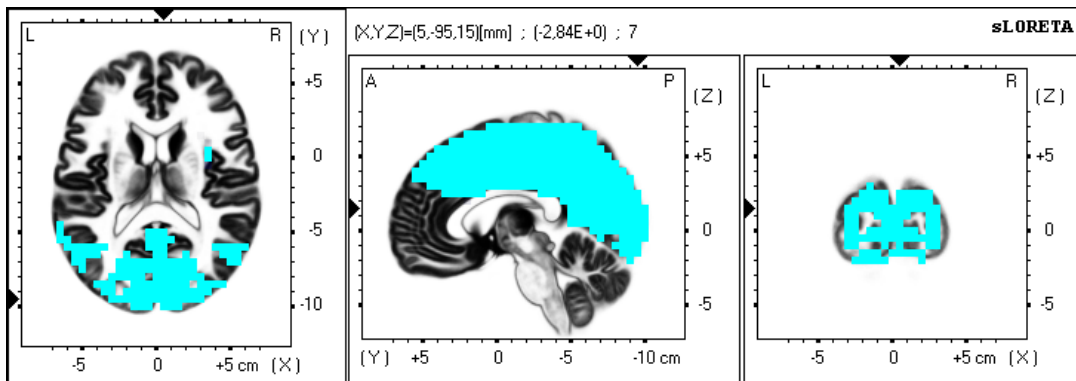
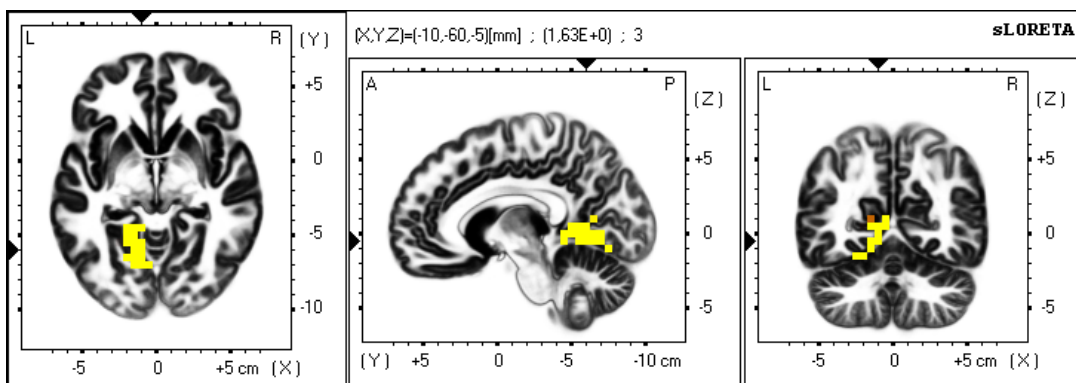


Figure 17 Alpha 1



## CITATION:

Bruncvik, L., Rakús, T., Hubčíková, K., Pechanová, Z., Brunovsky, M. (2021). PNES – Case Review in Neuroimaging Context. *Cognitive Remediation Journal* [online], 10(1), 1-10 [put a date of citation]. Available on WWW: [https://cognitive-remediation-journal.com/artkey/crj-202101-0002\\_pnes-8211-case-review-in-neuroimaging-context.php](https://cognitive-remediation-journal.com/artkey/crj-202101-0002_pnes-8211-case-review-in-neuroimaging-context.php). ISSN 1805-7225.