

EEG findings in MCI. Reduced delta power during resting state EEG has been identified in patients with MCI (Liddell et al., 2007). Furthermore, in the study, individuals with MCI demonstrated a significant positive correlation between delta power and immediate memory recall. Liddell et al. (2007) proposed that these findings suggest that delta power may be linked to memory decline in MCI, indicating that it could serve as a sensitive indicator of prodromal or early cognitive decline. However, other studies have shown increased delta power in MCI patients compared to healthy controls, particularly in frontal and centroparietal regions (Adler, Bramesfeld & Jajcevic, 1999; Moretti, Zanetti, Binetti & Frisoni, 2012). A decrease in beta power has been found in individuals with mild AD (Hogan, Swanwick, Kaiser, Rowan & Lawlor, 2003).

EEG in MCI and AD

D'Atri et al. (2021): <https://doi.org/10.1016/j.isci.2021.102386>

The hallmark of the resting state EEG in patients with AD is the slowing of cortical rhythms, consisting of increased low-frequency (0.5-7.0 Hz) and decreased high-frequency activity (Babiloni et al., 2015; Jeong, 2004). Similar EEG features affect mild cognitive impairment (MCI) subjects, a condition being prodromal to AD in more than half of cases (Babiloni et al., 2006; Galluzzi et al., 2001; Petersen et al., 2001; Scheltens et al., 2002). The EEG slowing correlates with the functional, structural, and cognitive changes in the disease progression (Babiloni et al., 2006; Claus et al., 2000; Jelic et al., 1996) and has been considered an EEG expression of the neurodegenerative process (Dringenberg, 2000).

Results: The waking EEG activity recorded in the evening hours displayed significant differences at the prefrontal and right frontotemporal sites for the **delta** band and at the right occipital derivation only for the **alpha** band. Prefrontal delta power was significantly higher in the AD compared to the HC group, while the right frontotemporal **delta** activity increased in the AD compared to both the HC and MCI groups. As expected, the occipital **alpha** power was reduced in the AD and MCI groups compared to the HC group. In the morning EEG, the three groups showed differences only in the **delta** band with a prevalence of the **delta** activity in AD compared to both HC and MCI groups. On the other hand, the EEG activity of the MCI group differed from that of HC only at the frontal sites. Greater EEG slowing was associated with worse cognitive impairment, as indicated by lower MMSE scores.

Liu, Wang, Xin, Wang, Jiang & Meng, 2024: <https://doi.org/10.1186/s12877-024-05041-x>

Commented [JH1]: EEG in AD and MCI

Commented [JH2]: The EEG index that showed the strongest correlation with cognitive deterioration is the synthetic index of the EEG slowing during REM sleep. This result confirms previous findings (Montplaisir et al., 1996), and it also suggests that this composite index may be better suited as a disease marker than others based on cortical activity in a single frequency band measured during REM and NREM sleep or resting state, as well as than the same index evaluated during wakefulness.

Commented [JH3]: @article{d2021eeg, title={EEG alterations during wake and sleep in mild cognitive impairment and Alzheimer's disease}, author={D'Atri, Aurora and Scarpelli, Serena and Gorgoni, Maurizio and Truglia, Ilaria and Lauri, Giulia and Cordone, Susanna and Ferrara, Michele and Marra, Camillo and Rossini, Paolo Maria and De Gennaro, Luigi}, journal={Iscience}, volume={24}, number={4}, year={2021}, publisher={Elsevier} }

Commented [JH4]: EEG in cognitive impairment (MCI)

Commented [JH5]: @article{liu2024relationship, title={The relationship and pathways between resting-state EEG, physical function, and cognitive function in older adults}, author={Liu, Hairong and Wang, Jing and Xin, Xin and Wang, Peng and Jiang, Wanting and Meng, Tao}, journal={BMC geriatrics}, volume={24}, number={1}, pages={463}, year={2024}, publisher={Springer} }

Studies have found that EEG signals in older adults with cognitive impairment exhibit specific characteristics, such as a basic rhythm slowdown, manifested by an increase in low-frequency band (delta, theta) power and a decrease in high-frequency band (alpha, beta) power [8, 9]. Slowing of alpha power may be an early sensitive indicator of the brain transitioning from normal physiological function to aging or its pathological state [10], while increased theta power may have a good predictive effect on cognitive decline [11].

In individuals with cognitive impairment, acute exercise effects include decreased delta and theta power and increased beta power [18, 20]; longterm exercise appears to result in decreased delta [18, 20, 21] and theta rhythm power [19, 21], and increased alpha and beta rhythm power [18, 20, 22].

The left frontal area, located at the front of the brain, is associated with advanced cognitive functions, decision-making, problem-solving, and personality traits. It plays a role in executive functions, emotional regulation, and modulation of social behavior.

The severity of cognitive impairment is positively correlated with increased theta activity, and an increase in theta waves in EEG serves as a good predictor of cognitive decline [36].

As the activation level of the frontal lobe cortex increases, theta wave power decreases, leading to improved cognitive function. A decrease in theta wave power was observed after both single and prolonged exercise [17].

Finnigan & Robertson, 2011: <https://doi.org/10.1111/j.1469-8986.2010.01173.x>

We address the degree to which resting EEG bandpower is associated with cognitive performance in 73 healthy older adults (aged 56–70). Relative theta (4–6.5 Hz) power was significantly correlated with immediate and delayed verbal recall, attention, and executive function measures. Relative delta and alpha power and peak alpha frequency did not correlate with any cognitive measures. These data indicate that high resting theta power in healthy older adults is associated with better cognitive function and may be a marker of healthy neurocognitive aging.

In summary, these outcomes indicate that high resting-state theta power in older adults is associated with relatively greater cognitive impairment; whereby such impairment either may already exist, be developing, and/or may (be predestined to) subsequently manifest or increase (this is from introduction).

Across 73 healthy older adults, relative resting theta power was significantly correlated with performance on numerous cognitive tests, which assess verbal memory, attention, or executive function.

Commented [JH6]: 8. Lejko N, Larabi D I, Herrmann C S, et al. Alpha Power and Functional Connectivity in Cognitive decline: a systematic review and Meta-analysis [J]. J Alzheimers Dis. 2020;78:1047–88.

9. Garrido-Chaves R Perezv, Zapater-Fajari M, et al. EEG markers and subjective memory complaints in young and older people [J]. Int J Psychophysiol. 2022;182:23–31.

Commented [JH7]: 10. Babiloni C, Ferri R, Noce G et al. Abnormalities of Cortical Sources of Resting State Alpha Electroencephalographic Rhythms are Related to Education Attainment in Cognitively Unimpaired Seniors and Patients with Alzheimer's Disease and Amnesic Mild Cognitive Impairment [J]. Cerebral cortex (New York, NY: 1991), 2021;31(4):2220–37

Commented [JH8]: 11. Musaeus C S, Engedal K, Høgh P, et al. EEG Theta Power is an early marker of Cognitive decline in Dementia due to Alzheimer's disease [J]. J Alzheimer's Disease: JAD. 2018;64(4):1359–71.

Commented [JH9]: 18. Amjad I, Toor H, Niazi I K, et al. Therapeutic effects of aerobic exercise on EEG parameters and higher cognitive functions in mild cognitive impairment

Commented [JH10]: 20. Amjad I, Toor H, Niazi I K, et al. Xbox 360 Kinect Cognitive games improve

Commented [JH11]: 18. Amjad I, Toor H, Niazi I K, et al. Therapeutic effects of aerobic exercise on EEG

Commented [JH12]: 20. Amjad I, Toor H, Niazi I K, et al. Xbox 360 Kinect Cognitive games improve

Commented [JH13]: 21. Styliadis C, Kartsidis P, Paraskevopoulos E, et al. Neuroplastic effects of com-

Commented [JH14]: 19. Hong S G, Kim J H, Jun T W. Effects of 12-Week Resistance Exercise on Elec-

Commented [JH15]: 21. Styliadis C, Kartsidis P, Paraskevopoulos E, et al. Neuroplastic effects of com-

Commented [JH16]: 18. Amjad I, Toor H, Niazi I K, et al. Therapeutic effects of aerobic exercise on EEG

Commented [JH17]: 20. Amjad I, Toor H, Niazi I K, et al. Xbox 360 Kinect Cognitive games improve

Commented [JH18]: 22. Jiang H, Chen S, Wang L et al. An investigation of limbs Exercise as a treat-

Commented [JH19]: 36. Babiloni C, Visser P J, Frisoni G, et al. Cortical sources of resting EEG rhythms in

Commented [JH20]: 17. Pedroso R V, Lima-Silva A E, Tarachuque P E, et al. Efficacy of Physical Exercise

Commented [JH21]: EEG in cognitive impairment (older adults)

Commented [JH22]: @article{finnigan2011resting, title={Resting EEG theta power correlates with cognitive performance in healthy older adults}, author={Finnigan,

Lejko, Larabi, Herrmann & Aleman, 2020: <https://doi.org/10.3233/JAD-200962>

We have therefore decided to take the first step by focusing on the alpha band for three main reasons.

First, alpha is the most prominent rhythm during quiet wakefulness [33]. Second, it has large amplitudes ranging from 10 to 50 V [34]. This is in contrast to theta, beta, and gamma waves, which have smaller amplitudes at rest [34–36], and can be best identified using tasks with many repetitions. This makes measuring alpha activity in research and clinical practice highly feasible, as it does not require people with cognitive decline to perform challenging and tire-some tasks.

Lastly, lower power and synchronization of alpha oscillations have consistently been associated with neurodegenerative dementias, especially AD [29, 37–39]. These decreases were found to correlate with lower cognitive scores [40, 41], higher atrophy of the hippocampus [42] and higher amyloid burden [43], as well as with genetic susceptibility for AD [44, 45]. Though alpha activity in other neurodegenerative dementias is less well-researched, there is evidence of lower alpha power in people with dementia due to Lewy body disorders [46–49]. Alpha power and synchronization are therefore an important part of the changes in the EEG spectrum associated with cognitive decline.

Babiloni et al. (2011): <https://doi.org/10.3233/JAD-2011-0051>

It has been reported that a positive ERP peaking 600 ms after the zerotime of stimuli to be encoded (P600) was reduced in patients with AD and mild cognitive impairment (MCI), particularly in those MCI patients who subsequently converted to AD [7, 8]. Furthermore, a positive ERP peaking 300 ms after the zerotime of oddball stimuli (P300) was reduced in amplitude in AD patients [5, 9], even during its early stages [10].

When compared to the resting state EEG rhythms of healthy normal elderly (Nold) subjects, AD patients showed an amplitude increase of widespread delta and theta sources and an amplitude decrease of posterior alpha (8–13 Hz) and/or beta (13–30 Hz) sources [45, 55, 56, 76–78]. The observation of these abnormalities of the EEG rhythms could allow discrimination among different dementia diagnoses, for instance a marked decline of posterior slow-frequency alpha power shows peculiar features in mild AD subjects when compared to cerebrovascular dementia, fronto-temporal dementia and normal elderly subjects with similar cognitive impairment.

Furthermore, pathological increased amplitude of the theta sources characterized cerebro-vascular dementia patients [56].

Commented [JH23]: EEG in AD (alpha)

Commented [JH24]: @article{lejko2020alpha, title={Alpha power and functional connectivity in cognitive decline: a systematic review and meta-analysis}, author={Lejko, Nena and Larabi, Daouia I and Herrmann, Christoph S and Aleman, Andre and {\c}ur{\v{c}}-Blake, Branislava}, journal={Journal of Alzheimer's disease}, volume={78}, ...

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Commented [JH30]: [42] Babiloni C, Frisoni GB, Pievani M, Vecchio F, Lizio R, Buttiglione M, Geroldi C, Fracassi C, Eusebi F, Ferri ...

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Commented [JH32]: [44] Babiloni C, Benussi L, Binetti G, Bosco P, Busonero G, Cesaretti S, Dal Forno G, Del Percio C, Ferri R, Frisoni G, ...

Commented [JH33]: [46] Andersson M, Hansson O, Minthon L, Ros'en I, Londos E (2008) Electroencephalogram variability in dementia with ...

Commented [JH34]: EEG in MCI and AD

Commented [JH35]: @article{babiloni2011resting, title={Resting state cortical rhythms in mild cognitive impairment and Alzheimer's disease: ...

Commented [JH36]: [7] Olichney JM, Morris SK, Ochoa C, Salmon DP, Thal LJ, Kutas M, Irarui VJ (2002) Abnormal verbal event ...

Commented [JH37]: 5] Rossini PM, Rossi S, Babiloni C, Polich J (2007) Clinical neurophysiology of aging brain: from normal aging to neu- ...

Commented [JH38]: Polich J, Corey-Bloom J (2005) Alzheimer's disease and P300: review and evaluation of task and modality. Curr ...

Commented [JH39]: [45] Dierks T, Jelic V, Pascual-Marqui RD, Wahlund L, Julin P, Linden DE, Maurer K, Winblad B, Nordberg A (2000) ...

Commented [JH40]: 56] Babiloni C, Binetti G, Cassetta E, Cerboneschi D, Dal Forno G, Del Percio C, Ferreri F, Ferri R, Lanuzza B, Miniussi C, ...

Despite the evidence of abnormal cortical rhythms in MCI and AD subjects, EEG analysis alone is unable to allow a diagnosis of disease.

The hypothesis of some strict relationships between brain activity in MCI and AD subjects implies the prediction of similar features of resting state EEG rhythm in MCI and AD subjects as a function of genetic risk factors.

Farina et al., 2020: <https://doi.org/10.1016/j.neuroimage.2020.116795>

Among the most promising EEG markers are reduced **alpha** power and increased **theta** power, as well as increased **theta** band functional connectivity (Cassani et al., 2017; Hatz et al., 2015; Musaeus et al., 2018). EEG ratios, such as **theta**/gamma and high **alpha**/low **alpha**, have also been suggested as promising markers (Moretti et al., 2012, 2009).

Similar to AD models, the best features for distinguishing aMCI from healthy ageing were increased **theta** and **delta** power in left temporo-parietal electrodes, while the best predictor of control status was increased frontal **beta2** power. **Theta** and **beta1** power also discriminated aMCI from AD participants; that is, higher **theta** in left frontal and right parietal electrodes was associated with AD status, and higher temporo-parietal **beta1** power was associated with aMCI status.

The EEG markers highlighted here are consistent with the neurophysiological changes typically associated with AD; specifically, increases in slow wave activity (i.e., **delta** and **theta**) and decreases in fast wave activity (i.e., **alpha** and **beta**), which are thought to reflect loss of cholinergic innervations as the disease progresses (Cassani et al., 2018; Musaeus et al., 2018). **Theta** power was the best overall predictor of patient status, in line with the suggestion that increased **theta** is one of the first changes to occur in AD (Musaeus et al., 2018).

Differences in **theta** were most pronounced at temporo-parietal sites, where connectivity between electrodes was increased in AD and aMCI. AD participants were distinguished from aMCI participants by increased **theta** in frontal and parietal electrodes, likely reflecting widespread changes that occur at advanced stages (Fraga et al., 2013). Higher **delta** power in left temporo-parietal areas was also indicative of patient status, though to a lesser degree, consistent with evidence that **delta** changes occur later (Roh et al., 2011). As expected, healthy ageing was associated with higher **alpha** power, both in amplitude (temporo-parietal areas) and connectivity (with frontal electrodes), and **beta** power, which was increased in controls relative to patients, and in aMCI relative to AD.

Commented [JH41]: EEG in MCI and AD

Commented [JH42]: @article{farina2020comparison, title={A comparison of resting state EEG and structural MRI for classifying Alzheimer's disease and mild cognitive impairment}, author={Farina, Francesca R and Emek-Sava{\c{s}}, DD and Rueda-Delgado, L and Boyle, Rory and Kiiski, Hanni and Yener, G{\o}rsev and Whelan, Robert}, journal={Neuroimage}, volume={215}, pages={116795}, year={2020}, publisher={Elsevier} }

Fröhlich, Kutz, Müller & Claudia Voelcker-Rehagen (2021):

<https://doi.org/10.3389/fnagi.2021.675689>

Compared with healthy older adults, patients with Alzheimer's disease show decreased **alpha** and **beta** power as well as increased **delta** and **theta** power during resting state electroencephalography (rsEEG). Findings for mild cognitive impairment (MCI), a stage of increased risk of conversion to dementia, are less conclusive. Results indicate no rsEEG power differences between healthy individuals and those with MCI.

Thus, the hypotheses that MCI is characterized by lower **alpha** and **beta** power as well as stronger **delta** and **theta** power during EC could not be confirmed in our sample. This is not in complete agreement with prior findings of changes in the rsEEG in patients with MCI. For the rest with EC, it was shown that **alpha** and **beta** powers were reduced and **theta** and **delta** powers were either elevated or reduced in MCI compared with healthy OA (Koenig et al., 2005; Babiloni et al., 2006b, 2010; Kwak, 2006; Ya et al., 2015). In fact, when specifying former studies, each study only showed some of the listed changes, but the overlap between results was often not great even though similar parameters were studied.

Caravaglios et al., 2023: <https://doi.org/10.1177/15500594221110036>

Amnesic-MCI had: i) increased **delta/beta** activity in the superior frontal gyrus and decreased **alpha1** activity in the paracentral lobule (ie, default mode network); ii) greater **delta/theta/alpha/beta** in the superior frontal gyrus (i.e., attention network); iii) lower **alpha** in the left superior parietal lobe, as well as a lower **delta/theta** and **beta**, respectively in post-central, and in superior frontal gyrus(ie, attention network).

➔ Not that much about power, but other things

Özbek, Fide & Yener, 2021: <https://doi.org/10.1016/j.clinph.2021.05.012>

Compared to healthy controls individuals with early-onset Alzheimer's disease (EOAD) showed an increase in slow frequency bands and a decrease in fast frequency bands. Frontal **alpha/theta** power ratio is the best discriminating value between EOAD and young HC with the sensitivity and specificity greater than 80% with area under the curve (AUC) 0.881.

This study is the first to report that resting-state EEG power can be a promising marker for diagnostic accuracy between EOAD and healthy controls.

Additionally, our study showed that resting-state **alpha/theta** power ratio can accurately discriminate between individuals with EOAD and healthy controls.

Commented [JH43]: EEG in MCI and AD

Commented [JH44]: @article{froehlich2021characteristics, title={Characteristics of resting state EEG power in 80+-year-olds of different cognitive status}, author={Fr{o}hlich, Stephanie and Kutz, Dieter F and M{u}ller, Katrin and Voelcker-Rehage, Claudia}, journal={Frontiers in aging neuroscience}, volume={13}, pages={675689}, year={2021}, publisher={Frontiers Media SA} }

Commented [JH45]: EEG in MCI

Commented [JH46]: @article{caravaglios2023eeg, title={EEG resting-state functional networks in amnesic mild cognitive impairment}, author={Caravaglios, G and Muscoso, EG and Blandino, V and Di Maria, G and Gangitano, M and Graziano, F and Guajana, F and Piccoli, T}, journal={Clinical EEG and neuroscience}, volume={54}, number={1}, pages={36-50}, year={2023}, publisher={SAGE Publications Sage CA: Los Angeles, CA} }

Commented [JH47]: EEG in AD (early-onset AD)

Commented [JH48]: @article{ozbek2021resting, title={Resting-state EEG alpha/theta power ratio discriminates early-onset Alzheimer's disease from healthy controls}, author={Ozbek, Ya{u}gur and Fide, Ezgi and Yener, G{o}rsev G}, journal={Clinical Neurophysiology}, volume={132}, number={9}, pages={2019-2031}, year={2021}, publisher={Elsevier} }

Perez, Duque, Hidalgo & Salvador, 2024:

<https://doi.org/10.1016/j.biopsycho.2024.108823>

Over the years, a substantial body of research has amassed compelling evidence pointing to a progressive alteration in brain electrical activity in neurodegenerative disorders such as MCI or AD. This alteration is characterized by an increase in **theta** power and a decrease in **beta** power, followed in subsequent stages by a decrease in **alpha** power and an increase in **delta** power (Gouw et al., 2017; Jeong et al., 2021; Prichep et al., 2006).

In Iliadou's study, the MCI group exhibited a significant increase in overall **delta** power compared to the SCD group, highlighting distinctive spectral alterations. Moreover, in their spectral analysis comparing SCD and MCI, Sibilano et al. (2023) identified higher spectral power in the **delta** band that was associated with the clinical progression from SCD to MCI. Iliadou et al. (2021) observed lower spectral power in **theta** in SCD compared to individuals with MCI. Sibilano et al. (2023) utilized rsEEG to discriminate between SCD and MCI, identifying the **delta** and **theta** bands as discriminating the most between SCD and MCI. We identified six articles that reported alterations in **alpha** power in individuals with SCD compared to those with MCI and healthy control groups. Iliadou et al. (2021) observed a decrease in **alpha** spectral power in individuals with SCD compared to those with MCI. Individuals with SCD exhibited decreased **alpha**₁, compared to the healthy control group, and increased **alpha**₂ in the left temporal, central, and parietal cortex compared to those with MCI, as reported in the study by Mazzon et al. (2018). Iliadou et al. (2021) observed a decrease in **beta** spectral power in individuals with SCD compared to those with MCI. Additionally, Mazzon et al. (2018) identified a decrease in **beta** power in the left frontal regions when comparing the SCD group to the MCI group.

Overall, individuals with SCD exhibited a pronounced increase in **theta** spectral power compared to healthy controls, whereas those with MCI showed a further increment compared to individuals with SCD, indicating alterations across the MCI continuum. Similar alterations across the MCI continuum were observed in **alpha** spectral power.

Specifically, individuals with SCD displayed decreased **alpha** spectral power compared to healthy controls, but higher levels than in individuals with MCI. However, this trend is inconsistent because two studies reported increases, rather than decreases, in this band in both SCD and MCI. Additionally, findings for the **delta** and **beta** frequency bands are rather inconclusive, given that half of the studies did not identify significant effects within these bands. In the studies that did observe effects, an increase in **delta** was noted in individuals with SCD compared to healthy controls, as well as in individuals with MCI compared to

Commented [JH49]: EEG in SCD

Commented [JH50]: @article{perez2024eeg, title={EEG Frequency Bands in Subjective Cognitive Decline: A Systematic Review of Resting State Studies}, author={Perez, Vanesa and Duque, Ar{\`a}nazu and Hidalgo, Vanesa and Salvador, Alicia}, journal={Biological Psychology}, pages={108823}, year={2024}, publisher={Elsevier} }

individuals with SCD. The findings related to **beta** band activity are also unclear. Whereas one study reported an increase in **beta** band activity when comparing SCD and healthy controls, an examination of the MCI continuum reveals a tendency towards decreased **beta** band activity in individuals with MCI compared to those with SCD.

Furthermore, studies contrasting SCD and MCI reveal that rsEEG abnormalities persist and intensify as cognitive decline progresses. Our review revealed the following evidence: 1) increased **delta** power in individuals with SCD compared to both healthy controls and people with MCI, although these findings were not consistently reported across all the studies; 2) a progressive increase in **theta** frequency bands in individuals with SCD compared to healthy controls, which intensified when comparing MCI to SCD; 3) a decrease in the **alpha** frequency band in individuals with SCD compared to healthy controls, with this decrease being more pronounced in those diagnosed with MCI compared to SCD. However, this trend was not observed in two studies that compared individuals with SCD to healthy controls and individuals with MCI, respectively. In these studies, unexpected increases, rather than decreases, in **alpha** and **beta** frequency band spectral power were reported in the SCD groups compared to healthy controls, and in the MCI groups compared to the SCD groups; 4) a decrease in **beta** band activity was only noted in studies that compared MCI with SCD. Conversely, the other two studies reported increases in the frequency of this band.

Jelic, Shigeta, Julin, Almkvist, Winblad & Wahlund (1996):

<https://doi.org/10.1159/000106897>

A significant difference between the AD group and groups with objective and subjective memory disturbances was found for **delta**, **theta**, and **alpha** relative power. AD group significantly higher **theta** relative power in all investigated regions, and significantly lower **alpha** relative power in all investigated regions, in relation to the rest of the study population. For **delta** relative power, a lower level of significance ($p < 0.05$) was observed in the left and right temporal and parieto-occipital, and left frontal regions. No significant difference was found for **beta** relative power in the groups studied, but a tendency towards higher values was observed in frontal regions in groups with objective memory disturbance and subjective memory complaints, as well as an increase in the left and right temporal and parieto-occipital regions in the AD group. A significant difference was found between the AD group and the other groups for mean frequency in the 4- to 20- Hz range in all regions, except for the left temporal region where the AD group was significantly different compared to controls and the

Commented [JH51]: EEG in AD and MCI / objective and subjective decline

Commented [JH52]: @article{jelic1996quantitative, title={Quantitative electroencephalography power and coherence in Alzheimer's disease and mild cognitive impairment}, author={Jelic, Vesna and Shigeta, Masahiro and Julin, Per and Almkvist, Ove and Winblad, Bengt and Wahlund, Lars-Olof}, journal={Dementia and Geriatric Cognitive Disorders}, volume={7}, number={6}, pages={314--323}, year={1996}, publisher={S. Karger AG Basel, Switzerland} }

group with subjective memory disturbances. There was no significant difference in the mean frequency between AD patients and the group with objective memory disturbances in that region. All 4 spectral ratios were significantly lower in all investigated regions in the AD group. Significantly lower temporoparietal **alpha** band coherence was found in the AD group. A tendency towards a decrease in temporofrontal coherence was observed in the AD group. No significant differences between the groups with subjective or objective memory impairment when compared to the controls. The lack of clear qEEG changes in the present study in subjects with MCI can have at least two explanations. 1. Group is heterogeneous and, according to some follow-up studies, only a proportion of them will show further cognitive decline and develop manifest disease (4,9). 2. Some of these subjects have preclinical AD with pathology still restricted to medial temporal lobes, which cannot be detected as changes in EEG power.

Claus et al. (2000): <https://doi.org/10.1159/000017219>

Slowing on the electroencephalogram (EEG) in patients with Alzheimer's disease (AD), compared to normal control subjects, evidenced by increase of **theta** activity and decrease of **beta** or **alpha** power, is a uniform finding in previous studies [1–5].

More impairment in overall cognitive function was most strongly reflected in loss of parieto-occipital and fronto-central **alpha** activity.

Detailed analysis of cognitive domains in relation to localized EEG values also revealed most consistently associations with decrease in **alpha** activity. Lower temporal and parietal rCBF were significantly associated with lower parieto-occipital **alpha** activity, while presence of leukoaraiosis was significantly associated with lower relative **beta** activity and higher absolute **delta** and **theta** activity.

General level of cognitive function, assessed in several previous studies with the MMSE, is most consistently related to **alpha** activity on EEG in AD patients [8, 13, 16 17, 46], in agreement with our findings. However, also correlations between overall cognitive function and **delta** and **theta** activity were observed in our study and in previous reports [13–16].

Alpha power as strongest correlate of cognitive domains finds support in the study of Jelic et al. [19], where visuospatial functions were strongly related to **alpha** power in left parieto-occipital and right temporal regions. In the study by Jelic et al. [19] measures of frontal lobe function, including attention and abstraction, were significantly related to fronto-central **theta** activity. We also found that, in addition to **alpha** activity, fronto-central **theta** activity was selected as a significant predictor of performance in attention and abstraction.

Commented [JH53]: EEG in cognitive impairment in AD

Commented [JH54]: @article{claus2000determinants, title={Determinants of quantitative spectral electroencephalography in early Alzheimer's disease: cognitive function, regional cerebral blood flow, and computed tomography}, author={Claus, JJ and Ongerboer de Visser, BW and Bour, LJ and Walstra, GJM and Hijdra, A and Verbeeten Jr, B and Van Royen, EA and Kwa, VIH and Van Gool, WA}, journal={Dementia and geriatric cognitive disorders}, volume={11}, number={2}, pages={81–89}, year={2000}, publisher={S. Karger AG Basel, Switzerland} }

Commented [JH55]: 88 Dement Geriatr Cogn Disord 2000;11:81–89 Claus/Ongerboer de Visser/Bour/Walstra/Hijdra/Verbeeten/Van Royen/Kwa/van Gool

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Commented [JH57]: 13 Dierks T, Frolich L, Ihl R, Maurer K: Correlation between cognitive brain function and electrical brain activity in dementia of Alzheimer type. *J Neural Transm* 1995;99:55–62.

14 Elmstahl S, Rosen I, Gullberg B: Quantitative EEG in elderly patients with Alzheimer's disease and healthy controls. *Dementia* 1994;5:

Commented [JH58]: Jelic V, Shigeta M, Julin P, Almkvist O, Winblad B, Wahlund LO: Quantitative electroencephalography power and coherence in Alzheimer's disease and mild cognitive impairment. *Dementia* 1996;7:314–323

Commented [JH59]: Jelic V, Shigeta M, Julin P, Almkvist O, Winblad B, Wahlund LO: Quantitative electroencephalography power and coherence in Alzheimer's disease and mild cognitive impairment. *Dementia* 1996;7:314–323

Interestingly, our relative **beta** activity, but also absolute **theta** and **delta** activity, were significantly related to leukoaraiosis on CT. This analysis may demonstrate that increase of absolute **theta** and **delta** power is sometimes less clearly reflected in the relative power values.

This is probably due to the fact that **theta** and also **delta** power determine the main part of the total power. As, for instance, **theta** or **delta** increases, the total power more or less increases proportionally. An increase in **theta** or **delta** may then be reflected by the absolute power, rather than by the relative EEG values. The preclinical finding that **beta** activity is found in subcortical or lower cortical structures [51–53] may either suggest that neuronal function of these brain structures is compromised by the presence of leukoaraiosis or that leukoaraiosis results in disconnection of subcortical and cortical structures.

Thus, the results suggest that leukoaraiosis in AD patients is related to slowing of the EEG, evidenced mainly by increase of **theta** and loss of **beta** activity.

In conclusion, **alpha** activity may be closely associated with cognitive function and rCBF, while **beta** and **theta** activity are related to lower cortical or subcortical changes. Our study therefore suggests that the EEG bands reflect differential pathophysiologic changes in AD.

Duffy, McAnulty & Albert (1995): <https://doi.org/10.1093/cercor/5.3.215>

These results demonstrate that there are significant and topographically consistent alterations in qEEG in mild to moderately impaired AD patients. The qEEG differences between patients and controls were best reflected by increased **theta**, decreased **beta**, and reduced amplitude in the long-latency VER and AER. Spectral findings were basically the same for both absolute and relative measures.

By EEG spectral analysis, **theta** was increased and **beta** decreased for the AD patients (abstract).

Wada, Nanbu, Jiang, Koshino, Yamaguchi & Hashimoto (1997):

[https://doi.org/10.1016/0006-3223\(95\)00651-6](https://doi.org/10.1016/0006-3223(95)00651-6)

Compared with the normal controls, the AD patients had a significantly lower **alpha**-2 and **beta** band power in the resting EEG as well as a significant increase in **delta** and **theta** band power.

In addition, our patients were found to have a significantly lower EEG power for the **alpha**-2, **beta**-1 and **beta**-2 bands in the resting condition. These findings are in general agreement

Commented [JH60]: 51 Leung LS: Fast (beta) rhythms in the hippocampus: A review. *Hippocampus* 1992;2:93–98.
52 Riekkinen PJ, Riekkinen M, Sirvio J, Miettinen R, Koudstaal PJ: Loss of cholinergic neurons in the nucleus basalis induces neocortical electroencephalographic and passive avoidance deficits. *Neuroscience* 1992;47:823–831.
53 Boeijinga PH, Lopez da Silva FH: Differential distribution of beta and theta EEG activity in the entorhinal cortex of the cat. *Brain Res* 1988;448:272–286

Commented [JH61]: EEG in AD

Commented [JH62]: @article{duffy1995temporoparietal, title={Temporoparietal electrophysiological differences characterize patients with Alzheimer's disease: a split-half replication study}, author={Duffy, Frank H and McAnulty, Gloria B and Albert, Marilyn S}, journal={Cerebral Cortex}, volume={5}, number={3}, pages={215--221}, year={1995}, publisher={Oxford University Press} }

Commented [JH63]: EEG in AD

Commented [JH64]: @article{wada1997electroencephalographic, title={Electroencephalographic abnormalities in patients with presenile dementia of the Alzheimer type: quantitative analysis at rest and during photic stimulation}, author={Wada, Yuji and Nanbu, Yuko and Jiang, Zheng-Yan and Koshino, Yoshifumi and Yamaguchi, Nariyoshi and Hashimoto, Takuma}, journal={Biological psychiatry}, volume={41}, number={2}, pages={217--225}, year={1997}, publisher={Elsevier} }

with those of earlier studies showing that AD patients had background EEG slowing with a reduction in **alpha** and fast activity (Liddell 1958; Swain 1959; Horie et al 1990; Miyauchi et al 1989, 1994).

Topographic analyses of the resting EEG showed a significant increase in **delta** band power at the frontal regions. This finding is consistent with that of previous EEG studies both with visual inspection (Liddell 1958; Swain 1959) and with significant probability mapping (Miyauchi et al 1989, 1994), in which **delta** activity was observed predominantly at the frontal areas.

Previous studies showed the relationships between cognitive impairment and EEG abnormalities in AD patients (Johannesson et al 1979; Sulkava 1982) although these studies evaluated EEG by visual inspection. Although in our AD patients the MMS score was positively correlated with **alpha** and **beta** band power of the resting EEG, no significant correlations were found with **delta** or **theta** band power.

Schreiter-Gasser, Gasser & Ziegler (1993) : [https://doi.org/10.1016/0013-4694\(94\)90144-9](https://doi.org/10.1016/0013-4694(94)90144-9)

The degree of dementia is strongly reflected by an increase of power in the **delta** frequency band, accentuated on the left hemisphere, as well as a decrease of **alpha** activity. Longer duration of disease is associated with a decrease of power in the **alpha** frequency band, earlier age at onset with an additional increase of power in the **theta** frequency band. Visual EEG evaluation correlates highly with the degree of dementia, in contrast to visually assessed **CCT**.

The quantitative EEG shows a surprising power in reflecting stages of Alzheimer's disease and indeed the most striking associations occurred between the degree of dementia and **delta** power. Correlations of absolute **delta** power with MMSE scores ranged typically from 0.6 to 0.9 at different locations. The **delta** band is by far the best indicator for the degree of dementia. The **theta** band, which best separated patients from controls (Schreiter-Gasser et al. 1993), separates poorly different stages of dementia. This can be explained by the fact that **theta** activity is greatly increased already in mild to moderate cases. In the **delta** band, on the other hand, mild to moderate cases are intermediate between severe cases and controls. Thus, progression of Alzheimer's disease goes along with gradual increase of **delta** power. This is in general agreement with the literature (Cohen et al. 1985; Penttilfi et al. 1985; Soininen and Partanen 1988; Soininen et al. 1991), but the relationships are more clear-cut in our Study. This may be attributed to using absolute rather than relative power and also to LOG artifact correction.

Commented [JH65]: EEG in dementia and AD (what's the difference?)

Commented [JH66]: @article{schreiter1994quantitative, title={Quantitative EEG analysis in early onset Alzheimer's disease: correlations with severity, clinical characteristics, visual EEG and CCT}, author={Schreiter-Gasser, Ursula and Gasser, Theo and Ziegler, Peter}, journal={Electroencephalography and clinical Neurophysiology}, volume={90}, number={4}, pages={267--272}, year={1994}, publisher={Elsevier} }

Commented [JH67]: Craniale Computertomographie

Elmståhl, Rosen & Gullberg (1994): <https://doi.org/10.1159/000106706>

The control subjects showed increasing **theta** activity with age but the EEG changes did not correlate significantly with psychometric features. The AD patients showed highly significant increases in **delta** and **theta** activity and decreases in **beta** activity compared with controls. Our finding of a steadily increasing relative **theta** power with age is in accordance with earlier studies [20, 21], but at variance with other studies which indicate a continuous decline in **delta** and **theta** activity with age [10, 22].

Not only **theta** power but also **delta** power is significantly increased. This is most likely due to the fact that our patient sample is one of late-onset AD in a rather advanced stage. In studies of the progress of the EEG with increasing severity of AD **theta** power is increased, followed by a decrease in **beta** power and later by an increase in **delta** power and a decrease in **alpha** power [3, 25]. Our group of elderly institutionalized patients had obviously reached a stage of marked **delta** power increase.

The topographical analysis showed a widespread increase in **delta** and **theta** power over most cortical areas, whereas the decrease in **beta** power was more restricted to posterior temporoparietal areas. **Delta** waves are considered to reflect primarily abnormalities of connections between subcortical and cortical areas whereas the **beta** power decrease is considered to reflect cortical degenerative changes [26, 27]. The results of our study would therefore indicate a profound derangement of subcortical function in combination with degeneration of posterior cortical areas.

Dierks, Frölich, Ihl & Maurer (1994): <https://doi.org/10.1007/BF01271469>

Summarizing the present investigation, we demonstrated a) an increase of dipole strength in the slow frequency bands, b) a more anterior equivalent dipole of **alpha**- and **beta**-activity, and c) a slowing of the EEG with increasing cognitive deterioration. The results support the assumption that cognitive decline in dementia can be assessed by measuring the electrical activity of the brain.

Kuskowski, Mortimer, Morley, Malone & Okaya (1993): [https://doi.org/10.1016/0006-3223\(93\)90108-P](https://doi.org/10.1016/0006-3223(93)90108-P)

Log-absolute EEG power in the **alpha** bandwidth (8–12 Hz) was found to be correlated with the computed rate of **MMSE decline**. This association was present for electrode sites across all regions of the scalp and remained significant when the effects of current cognitive severity

Commented [JH68]: akzent

Commented [JH69]: EEG in AD

Commented [JH70]: @article{elmstaahl1994quantitative, title={Quantitative EEG in elderly patients with Alzheimer's disease and healthy controls}, author={Elmståhl, Sören and Rosen, Ingemar and Gullberg, Bo}, journal={Dementia and Geriatric Cognitive Disorders}, volume={5}, number={2}, pages={119--124}, year={1994}, publisher={S. Karger AG Basel, Switzerland} }

Commented [JH71]: 20 Obrist WD: Problems of aging; in Rasmussen A (ed): Handbook of Electroencephalography and Clinical Neurophysiology. Amsterdam. Elsevier. 1976. vol 6A. pp 275-292.
21 Bussc EW; Electroencephalography: in Reisberg B (ed): Alzheimer's Disease. New York. The Free Press, 1983, pp 231-236

Commented [JH72]: 10 Williamson PC, Merskey H, Morrisott S, Rabbitt K, Fox H, Wands K, Wong C, Hachinski V: Quantitative electroencephalographic correlates of cognitive decline in normal elderly subjects. Arch Neurol 1990;47: 1185-1188
22 Duffy FH, Albert MS, McNulty G, Garvey AJ: Age-related differences in brain electrical activity of healthy subjects. Ann Neurol 1984; 16:430-438.

Commented [JH73]: 3 Coblen LA, Danziger W, Storandt M: A longitudinal EEG study of mild senile dementia of Alzheimer type: Changes at 1 year and at 2.5 years. J Electroencephalogr Clin Neurophysiol 1985;61:101-112.
25 Soininen H, Partanen J, Laulumaa V, Paakkonen A, Helkala EL, Riekkinen P: Serial EEG in Alzheimer's disease: 3 year follow-up and clinical outcome. J Electroencephalogr Clin Neurophysiol 1991;79:342-348.

Commented [JH74]: 26 Kellaway P: An orderly approach to visual analysis: Parameters of the normal EEG in adults and children; in Klass DW, Daly DD (eds): Current practice of Clinical Electroencephalography. New York, Raven Press. 1979. pp 69-148.

27 Gloor P, Ball G, Schaul N: Brain lesions that

Commented [JH75]: EEG in dementia

Commented [JH76]: @article{dierks1995correlation, title={Correlation between cognitive brain function and electrical brain activity in dementia of Alzheimer type}, author={Dierks, T and Frölich, L and Ihl, R and Maurer, R}}

Commented [JH77]: EEG in AD

Commented [JH78]: @article{kuskowski1993rate, title={Rate of cognitive decline in Alzheimer's disease is associated with EEG alpha power}, author={Kuskowski, Michael A and Mortimer, James A and Morley, Gerald K and}}

Commented [JH79]: Mini Mental State Examination

were partialled out. These data suggest that a quantitative EEG measure (absolute **alpha** power) is related to the rate of cognitive decline in patients with Alzheimer's disease.

The data presented here suggest that a quantitative EEG measure (absolute **alpha** power) is correlated with the rate of cognitive decline in patients with AD. This association was present for electrode sites across all regions of the scalp and remained significant when the effects of current cognitive severity were partialled out.

Dringenberg (2000): [https://doi.org/10.1016/S0166-4328\(00\)00261-8](https://doi.org/10.1016/S0166-4328(00)00261-8)

The generalized slowing of the neocortical EEG is a characteristic symptom in AD and refers to a reduction in desynchronized, activated EEG patterns that are replaced by deactivated, synchronized activity. Typical EEG changes in AD include a loss of **beta** (13–30 Hz) activity, a decrease in power and mean frequency of **alpha** activity (8–12 Hz), and increased power in the **theta** (4–7 Hz) and **delta** (B4 Hz) bands [21,39,71,74].

Also, **alpha** EEG coherence decreases while **delta** coherence increases in patients with clinically probable AD [56]. These quantitative EEG changes provide a sensitive index of the cognitive status of AD patients; McAdam and Robinson [59] reported a positive correlation of 0.79 between EEG abnormalities and the severity of dementia in a population of demented patients, and Penttilä et al. [71] noted a significant positive correlation between occipital peak (**alpha**) frequency and neuropsychological test scores in AD patients (i.e. lower frequency was associated with lower test scores).

Some EEG changes, such as the increase in **theta** power, occur together with the earliest signs of cognitive deterioration, while others are associated with more advanced cognitive decline (e.g. increased **delta** power [71,74]). The close relation between EEG slowing and the severity of cognitive symptoms suggests that a disruption of processing in cortical networks contributes importantly to the behavioral disorganization present in AD.

Babiloni et al. (2006): <https://doi.org/10.1016/j.clinph.2005.09.019>

In the present study, low-band (8–10.5 Hz) **alpha** sources in parietal, occipital, temporal, and limbic areas had an intermediate magnitude in MCI subjects when compared to mild AD and Nold subjects. Furthermore, magnitude of these five EEG sources showed positive linear and non-linear (i.e. correlations with MMSE score (global cognitive level) across all Nold, MCI, and mild AD subjects as a single group. These results suggest that the global neurophysiological variables (posterior cortical rhythmicity) were linearly and not linearly

Commented [JH80]: EEG in AD

Commented [JH81]: @article{dringenberg2000alzheimer, title={Alzheimer's disease: more than a 'cholinergic disorder'—evidence that cholinergic–monoaminergic interactions contribute to EEG slowing and dementia}, author={Dringenberg, Hans C}, journal={Behavioural brain research}, volume={115}, number={2}, pages={235–249}, year={2000}, publisher={Elsevier} }

Commented [JH82]: [21] Coben LA, Danziger W, Storandt M. A longitudinal EEG study of mild senile dementia of Alzheimer's type: Changes at 1 year and at 2.5 years. *Electroencephalogr Clin Neurophysiol* 1985;61:101–12.

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Commented [JH83]: [56] Locatelli T, Cursi M, Liberati D, Franceschi M, Comi G. EEG coherence in Alzheimer's disease. *Electroencephalogr Clin Neurophysiol* 1998;106:229–37.

Commented [JH84]: [59] McAdam W, Robinson RA. Senile intellectual deterioration and the electroencephalogram: a quantitative correlation. *J Ment Sci* 1956;102:438–45.

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Commented [JH86]: [71] Penttilä M, Partanen JV, Soininen H, Riekkinen PJ. Quantitative analysis of occipital EEG in different stages of Alzheimer's disease. *Electroencephalogr Clin Neurophysiol* 1985;60:1–6. [74] Pritchep LS, John ER, Ferds SH, Reisberg B, Almas M, Alper K, Cancro R. Quantitative EEG correlates of cognitive deterioration in the elderly. *Neurobiol Aging* 1994;15:85–90.

Commented [JH87]: EEG in MCI AD and normal old (Nold)

Commented [JH88]: @article{babiloni2006sources, title={Sources of cortical rhythms change as a function of cognitive impairment in pathological aging: a multicenter study}, author={Babiloni, Claudio and Binetti, Giuliano and Cassetta, Emanuele and Dal Forno, Gloria and Del Percio, Claudio and Ferreri, Florinda and Ferri, Raffaele and Frisoni, Giovanni and Hirata, Koichi and Lanuzza, Bartolo and others}, journal={Clinical neurophysiology}, volume={117}, number={2}, pages={252–268}, year={2006}, publisher={Elsevier} }

correlated with global clinical and cognitive status (MMSE score) across the shadow region between physiological and pathological aging.

The present results extend in spatial detail previous EEG evidence showing a decrease of **alpha** power in MCI compared to normal subjects (Frodl et al., 2002; Grunwald et al., 2001; Huang et al., 2000; Jelic et al., 1996; 2000).

Furthermore, they complement previous evidence of early atrophy signs in limbic, precuneus, and posterior cingulate areas of MCI subjects (Baron et al., 2001; Callen et al., 2001).

In our MCI group, the **alpha** findings paralleled those in occipital **delta** (2–4 Hz), which had an intermediate magnitude compared to mild AD and Nold subjects. Furthermore, magnitude of these EEG sources showed negative linear and non-linear correlations with MMSE score (global cognition) across all subjects. These results are compatible with previous EEG evidence showing increased slow rhythms in MCI compared to normal controls (Grunwald et al., 2001; Jelic et al., 2000; Prichep et al., 1994; Wolf et al., 2003). Furthermore, previous evidence have shown that the increase of slow EEG rhythms in AD is secondary to progressive cortical hypoperfusion (Brenner et al., 1986; Dossi et al., 1992; Kwa et al., 1993; Niedermeyer, 1997; Nobili et al., 1998; Passero et al., 1995; Rae-Grant et al., 1987; Stigsby et al., 1981; Steriade et al., 1994; Rodriguez et al., 1999a; Young, 1987). From a physiological viewpoint, **delta** rhythms have been intensively studied during slow wave sleep. These rhythms are then replaced by fast (**beta** and gamma) cortical oscillations induced by the depolarizing effects of meso-pontine cholinergic neurons acting on thalamocortical neurons and by the depolarizing effects of nucleus basalis cholinergic neurons acting on cortical neurons (Steriade, 2003). Therefore, it can be speculated that the increment of **delta** oscillations in MCI and AD subjects might be related to loss of hippocampal and posterior cortical neurons, which are impinged by cholinergic inputs. Indeed, it has been demonstrated that early degeneration in mesial temporal cortex of MCI and AD subjects can affect functional connectivity between hippocampal formation and temporoparietal cortex (Killiany et al., 1993). Furthermore, a bilateral reduction of gray matter volume in the hippocampal formation and entorhinal cortex of AD subjects was correlated with an increment of **delta** rhythms in posterior cortex (Fernandez et al., 2003).

In the present study, the **theta** sources in parietal, occipital, temporal and limbic areas had a stronger magnitude in mild AD subjects than MCI and Nold subjects. These results extend in spatial detail previous EEG evidence showing an increase of **theta** power in mild AD compared to normal subjects (Coben et al., 1983; Huang et al., 2000; Mattia et al., 2003; Ponomareva et al., 2003).

The results of the present study showed that cortical sources of EEG rhythms changed across Nold, MCI, and mild AD subjects, as a function of the global cognitive level.

This was true for occipital **delta** and **alpha** 1 sources in parietal, occipital, temporal, and limbic areas, which had an intermediate magnitude in MCI subjects compared to mild AD and Nold subjects and were correlated with MMSE score across all subjects.

Jeong (2004): <https://doi.org/10.1016/j.clinph.2004.01.001>

Since Hans Berger, the discoverer of the electroencephalogram (EEG), first observed pathological EEG sequences in a historically verified AD patient (Berger, 1931, Berger, 1932), a large number of studies about the EEG of AD have been performed. The hallmark of EEG abnormalities in AD patients is slowing of the rhythms and a decrease in coherence among different brain regions. An increase in **theta** and **delta** activities and a decrease in **alpha** and **beta** activities are repeatedly observed (Brenner et al., 1986, Coben et al., 1983, Coben et al., 1985, Giaquinto and Nolfé, 1986), and a reduced coherence of the **alpha** and **beta** bands is frequently found (Dunkin et al., 1994, Leuchter et al., 1987, Locatelli et al., 1998).

Furthermore, these abnormalities are correlated with the severity of the disease (Hughes et al., 1989, Kowalski et al., 2001). For the last 2 decades, the EEG has been utilized as a useful tool for diagnosing dementias.

There is a good correlation between the degree of the EEG abnormality and cognitive impairment (Brenner et al., 1988; Erkinjuntti et al., 1988; Johannesson et al., 1979; Kaszniak et al., 1979; Liddle, 1958; Merskey et al., 1980; Obrist et al., 1962; Rae-Grant et al., 1987; Roberts et al., 1978; Soininen et al., 1982; Wiener and Schuster, 1956).

It is of clinical interest to find that the EEG abnormality is associated with cognitive deficits. A good correlation is found between EEG spectral measures and cognitive deterioration scores, such as the Folstein (Mini-mental) score (Brenner et al., 1986; Elmsa^{hl} et al., 1994; Filipovitch et al., 1989; Leuchter et al., 1987; Leuchter et al., 1993; Schreiter-Gasser et al., 1994; Strijers et al., 1997), the global deterioration score (Helkala et al., 1991; Passero et al., 1995; Prichep et al., 1994), and a composite neuropsychological test score (Penttilä et al., 1985). There are, however, some studies reporting only a weak correlation or no correlation between EEG changes and the cognitive decline in AD (Hughes et al., 1989; Prinz and Vitiello, 1989).

Babiloni (2015): <https://doi.org/10.1016/j.iopsycho.2015.02.008>

Commented [JH89]: EEG in AD and cognitive impairment

Commented [JH90]: Here more information about delta, theta, beta and alpha waves in AD

Commented [JH91]: @article{jeong2004eeg, title={EEG dynamics in patients with Alzheimer's disease}, author={Jeong, Jaeseung}, journal={Clinical neurophysiology}, volume={115}, number={7}, pages={1490--1505}, year={2004}, publisher={Elsevier} }

Commented [JH92]: EEG in MCI and AD
EEG introduction, why it's great to use

Commented [JH93]: Good paper!

Commented [JH94]: @article{babiloni2016brain, title={Brain neural synchronization and functional coupling in Alzheimer's disease as revealed by resting state EEG rhythms}, author={Babiloni, Claudio and Lizio, Roberta and Marzano, Nicola and Capotosto, Paolo and Soricelli, Andrea and Triggiani, Antonio Ivano and Cordone, Susanna and Gesualdo, Loreto and Del Percio, Claudio}, journal={International Journal of Psychophysiology}, volume={103}, pages={88--102}, year={2016}, publisher={Elsevier} }

Resting state EEG power density differed between AD patients and amnesic MCI subjects, who were considered to be at high risk of suffering from prodromal AD. There was an “intermediate” power density of low-frequency **alpha** rhythms (8–10.5 Hz) in the parietal and occipital regions in MCI compared to mild AD and Nold subjects (Babiloni et al., 2006b). Furthermore, maximum **alpha** and **beta** power density shifted more anteriorly in AD patients compared to Nold and MCI subjects (Huang et al., 2000). Moreover, longitudinal studies have shown that increased **delta** or **theta** power density, decreased **alpha** and **beta** power density, and slowing of mean EEG frequency were in some way predictors of the progression from MCI to dementia at about 1-year follow-up (Huang et al., 2000; Jelic et al., 1996, 2000; Grunwald et al., 2001; Kwak, 2006; Rossini et al., 2006). High power density of the posterior **alpha** rhythms also predicted a stable global cognitive function in MCI subjects at 1-year follow-up (Babiloni et al., 2010a).

In the MCI subjects, the EEG markers of disease progression included an increase of the power density at the **theta** and **delta** rhythms in the temporal and occipital regions as well as a decrease of the power density at **beta** rhythms in temporal and occipital regions (Jelic et al., 2000).

In the AD patients with global cognitive impairment, hippocampal atrophy was associated with increased power density at **delta** and **theta** rhythms in the temporal and parietal regions (Helkala et al., 1996), in line with recent magnetoencephalographic (MEG) evidence (Fernandez et al., 2003).

Furthermore, a volume decrement of hippocampus was related to the decreased power density at **alpha** rhythms in the temporal, parietal, and occipital regions in MCI and AD subjects (Babiloni et al., 2009b).

Furthermore, the global **delta** and **alpha** power density was related to the total amount of atrophy of cortical gray matter in the amnesic MCI and in the AD subjects, as revealed by MRI voxel-to-voxel volumetry of lobar brain volume; the higher the total gray matter volume, the lower the global **delta** power density and the higher the global **alpha** power density (Babiloni et al., 2012).

The power density of the resting state eyes-closed EEG rhythms was repeatedly found to be correlated to cognitive status in MCI and AD subjects. It has been shown that the posterior **alpha** power density was positively correlated to the subjects' global cognitive status, as measured by ADAS-cog in the MCI or AD subjects; namely, the lower the **alpha** power density, the lower the cognitive status (Luckhaus et al., 2008). This relationship can be extended to the cognitive health condition. Furthermore, the posterior **delta** and **alpha** power

density was correlated to the MMSE score in the Nold, MCI and AD subjects; namely, the lower the **alpha** power, the higher the **delta** power and the lower the cognitive status (Babiloni et al., 2006b).

It has been shown that in the MCI subjects, the markers of disease progression included an increase of the power density at **theta** and **delta** rhythms in the temporal and occipital lobes as well as the reduction of the **beta** power density in the temporal and occipital lobes (Jelic et al., 2000).

AD patients were characterized by an increase of **theta** and **delta** power density and by a reduction of the **alpha** and **beta** power density in the parieto-occipital lobes (Coben et al., 1985). Furthermore, half of the AD patients showed an increase of the **theta** and **delta** power density in the temporal-occipital lobe (Soininen et al., 1989).