Cureus

Review began 05/30/2022 Review ended 06/07/2022 Published 06/08/2022

© Copyright 2022

Fontani et al. This is an open access article distributed under the terms of the Creative Commons Attribution License CC-BY 4.0., which permits unrestricted use, distribution, and reproduction in any medium, provided the original author and source are credited.

Long-Lasting Efficacy of Radio Electric Asymmetric Conveyer Neuromodulation Treatment on Functional Dysmetria, an Adaptive Motor Behavior

Vania Fontani ^{1, 2}, Arianna Rinaldi ³, Chiara Rinaldi ⁴, Laura Araldi ⁵, Alida Azzarà ⁶, Antonio M. Carta ⁵, Nicoletta Casale ⁵, Alessandro Castagna ², Maurizio Del Medico ⁷, Maurizio Di Stasio ⁵, Marina Facchini ⁸, Monica Greco ⁹, Savino LaMarca ¹⁰, Giovanni Loro ¹¹, Anna Marrone ⁵, Alessandra Palattella ¹², Giulio Pellegata ¹³, Daniele Ruini ¹⁴, Corrado Schmitt ⁵, Franco Vianini ⁵, Margherita Maioli ³, Carlo Ventura ¹⁵, Franco Caltabiano ¹⁶, Adriano J. Bueno ¹⁷, Amélia Fugino Matuoka ¹⁸, Edison Massahiro Nabechima ¹⁸, Fabio A. Bechelli ¹⁸, Fabricio da Silveira Bossi ¹⁹, Greice C. Nitschke Fontana ¹⁸, Jaques Finkielsztejn ¹⁸, José Alfredo Coelho Pereira ¹, Juarez Nunes Callegaro ²⁰, Kleiner Vasconcelos Pinheiro ²¹, Lara R. Ferreira Alves ¹⁸, Marcelo Kodja Daguer ¹⁷, Márcia C. Marins Martins ²², Mauricio Bezerra Uliana ¹⁷, Nelson Knop Zisman ¹⁸, Paulo Cezar Schütz ¹⁷, Paulo R. Fochesato ¹⁸, Pollyanna Celso Felipe de Castro ²³, Rosa M. Tanaka Nabechima ¹⁸, Roseli B. Randon ²², Salvatore Rinaldi ^{1, 2}

1. Research, Rinaldi Fontani Foundation, Florence, ITA 2. Regenerative Medicine, Rinaldi Fontani Institute, Florence, ITA 3. Biomedical Sciences, University of Sassari, Sassari, ITA 4, Neuroscience, Psychology, Drug Area and Child Health (NEUROFARBA), University of Florence, Florence, ITA 5. Internal Medicine, International Scientific Society of Neuro Psycho Physical Optimization with REAC Technology, Florence, ITA 6, Obstetrics and Gynecology, International Scientific Society of Neuro Psycho Physical Optimization with REAC Technology, Florence, ITA 7. Medical Physics, International Scientific Society of Neuro Psycho Physical Optimization with REAC Technology, Florence, ITA 8. Neurology, International Scientific Society of Neuro Psycho Physical Optimization with REAC Technology, Florence, ITA 9. Geriatrics, International Scientific Society of Neuro Psycho Physical Optimization with REAC Technology, Florence, ITA 10, Oral Medicine, International Scientific Society of Neuro Psycho Physical Optimization with REAC Technology, Florence, ITA 11. Diabetes and Endocrinology, International Scientific Society of Neuro Psycho Physical Optimization with REAC Technology, Florence, ITA 12. Psychiatry, International Scientific Society of Neuro Psycho Physical Optimization with REAC Technology, Florence, ITA 13. Anesthesiology, International Scientific Society of Neuro Psycho Physical Optimization with REAC Technology, Florence, ITA 14. Family Medicine, International Scientific Society of Neuro Psycho Physical Optimization with REAC Technology, Florence, ITA 15. Cardiology and Microbiology, National Laboratory of Molecular Biology and Stem Cell Engineering, National Institute of Biostructures and Biosystems, Bologna, ITA 16. Surgery and Urology, International Scientific Society of Neuro Psycho Physical Optimization with REAC Technology, Florence, ITA 17. Orthopaedics, International Scientific Society of Neuro Psycho Physical Optimization with REAC Technology, São Paulo, BRA 18. Internal Medicine, International Scientific Society of Neuro Psycho Physical Optimization with REAC Technology, São Paulo, BRA 19. Surgery, Endoscopy and Gastroenterology, International Scientific Society of Neuro Psycho Physical Optimization with REAC Technology, São Paulo, BRA 20. Psychiatry, International Scientific Society of Neuro Psycho Physical Optimization with REAC Technology, São Paulo, BRA 21. Integrative/Complementary Medicine, International Scientific Society of Neuro Psycho Physical Optimization with REAC Technology, São Paulo, BRA 22. Pediatrics, International Scientific Society of Neuro Psycho Physical Optimization with REAC Technology, São Paulo, BRA 23. Neurology, International Scientific Society of Neuro Psycho Physical Optimization with REAC Technology, São Paulo, BRA

Corresponding author: Salvatore Rinaldi, srinaldi@irf.it

Abstract

Background

Fluctuating asymmetry (FA) is widely defined as the deviation from perfect bilateral symmetry and is considered an epigenetic measure of environmental stress. Rinaldi and Fontani hypothesized that the FA morpho-functional changes originate from an adaptive motor behavior determined by functional alterations in the cerebellum and neural circuits, not caused by a lesion, but induced by environmental stress. They called this phenomenon functional dysmetria (FD). On this premise, they developed the radio electric asymmetric conveyer (REAC) technology, a neuromodulation technology aimed at optimizing the best neuro-psycho-motor strategies in relation to environmental interaction.

Aims

Previous studies showed that specific REAC neuro postural optimization (NPO) treatment can induce stable FD recovery. This study aimed to verify the duration of the NPO effect in inducing the stable FD recovery over time.

Materials and methods

Data were retrospectively collected from a population of 29,794 subjects who underwent a specific semiological FD assessment and received the NPO treatment, regardless of the pathology referred.

How to cite this article

Results

The analysis of the data collected by the various participants in the study led us to ascertain the disappearance of FD in 100% of the cases treated, with a stability of the result detected up to 18 years after the single administration of the REAC NPO treatment.

Conclusions

The REAC NPO neurobiological modulation treatment consisting of a single administration surprisingly maintains a very long efficacy in the correction of FD. This effect can be explained as the long-lasting capacity of the NPO treatment to induce greater functional efficiency of the brain dynamics as proven in previous studies.

Categories: Internal Medicine, Public Health

Keywords: neurobiological modulation, neurostimulation, neuromodulation, functional dysmetria, radio electric asymmetric conveyor, epigenetic, neuroplasticity, neurotransmission, endogenous bioelectric activity

Introduction

In living organisms, the loss of perfect somatic symmetry is constantly observed. This phenomenon called fluctuating asymmetry (FA) is considered an epigenetic measure of environmental stress [1]. Fluctuating asymmetry is defined as the deviation from perfect bilateral symmetry. This deviation is caused by environmental stressors, developmental instability, and genetic/epigenetic problems during development [1-4]. Therefore, both genomic and environmental or epigenetic changes can increase FA. Consequently, FA is a useful phenomenon for stress monitoring both in nature and in the laboratory.

Fluctuating asymmetry is the manifestation of a deterioration of the allostatic mechanisms of development, which can be highlighted through changes in the individual morphological symmetry [2-4]. Since FA is an omnipresent phenomenon in all living beings, its presence is not considered a pathology, but an indicator of environmental stress [1-4]. Although there is almost unanimous agreement on its meaning, the mechanisms by which FA occurs or how it can be countered have not yet been identified.

Rinaldi and Fontani have focused on the processes that result in FA. They started from the observation that to induce a morphological asymmetry such as FA, the affected structures must be subjected to a different mechanical load. In living organisms, muscles are the structures that contract and generate mechanical force that is transmitted to their insertion points [5]. If the forces generated by the muscles, particularly by the hetero-lateral ones, are asymmetrical, a difference in both pressure and traction is created. This determines the so-called piezoelectric effect [6], a phenomenon that induces variations of the endogenous bioelectric activity (EBA) [7], i.e., electro negativity/electro positivity both at the bone level and in the organic component [8,9]. A morpho-functional remodeling follows in the structures affected by the piezoelectric phenomenon [10]. In summary, the authors have hypothesized that the asymmetrical activation (dysmetria) and the consequent piezoelectric effect are the cause of those morpho-functional changes that are manifested in the FA.

The lack of motor coordination and executive precision is already defined in the literature as dysmetria and is typically identified as a manifestation of cerebellar dysfunction or lesion [11]. Dysmetria can be defined as an error in trajectory due to an abnormal range, rate, and/or force of motion as in Holmes's [12] definition, or a disturbance of the trajectory or placement of a body part during active movement, both in range and direction, as in Textbook of Clinical Neurology (third edition) [13].

Thanks to the new knowledge on cerebellar functions, today the term dysmetria can also indicate dysmetria of thought or the inability to adapt i.e., modulate precise and adequate behaviors around an allostatic-homeostatic baseline [14]. The cerebellum plays numerous key roles in the emotional [15], cognitive, and behavioral spheres [16], and in motor control, coordination, sense of balance, learning, and some cognitive functions related to language and attention. Considering the variety and complexity of the functions it performs, it follows that the cerebellum is widely involved in adaptive phenomena [17-19].

When dysmetria can be observed in healthy subjects [20,21], it has no lesional or pathological origin. Therefore, Rinaldi and Fontani deduced that it could be the result of an adaptive dysfunction at the level of the neural circuits, such as the cerebellar ones [22], that control the "symmetry and adequacy" of movement, and the emotional, cognitive, and behavioral sphere.

Proceeding in this research, Rinaldi and Fontani developed the term and the concept of functional dysmetria (FD) [20,21,23], to describe an adaptive motor behavior [24] determined by functional alterations in the cerebellum [18,25] and neural circuits [26], induced by the experienced environmental stress or allostatic overload [19]. Similarly, the FA is the morphological manifestation of an epigenetic phenomenon and an indicator of environmental stress [1]. While in the specialized orthopedic literature, the term dysmetria is sometimes erroneously used to indicate a heteromeric measurement of the limbs or bone segments [27],

Rinaldi and Fontani correlated FD to the phenomena of dysfunctional adaptive motor behavioral response to environmental stressors.

Over the years, Rinaldi and Fontani have investigated how to recognize and differentiate the presence of FD and they identified a specific motor task to assess the asymmetrical activation of symmetrical muscle groups [20,21]. This semiological procedure highlights the asymmetrical activation of both the quadriceps, during a motor task [20,21]. Once they had standardized the FD assessment, Rinaldi and Fontani focused on how to recover the dysfunctional neuro-psycho-physical adaptive mechanisms underlying FD. For this purpose, they developed the radio electric asymmetric conveyer (REAC) technology, a neurobiological modulation technology aimed at recovering the best motor and behavioral strategies in response to environmental stressors [28-35].

Since previous studies using a specific treatment of REAC technology, the neuro postural optimization (NPO), has been shown to induce the stable recovery of FD [20,21], the purpose of this study was to verify the duration of the NPO effect over time and formulate the most likely hypothesis about the verified correlation between the REAC NPO treatment and the stable duration of the recovery of the FD. This FD recovery could be explained as the REAC NPO efficacy in inducing greater functional efficiency of the brain dynamics, previously observed in functional magnetic resonance imaging (fMRI) studies [20,21,23].

Materials And Methods

Ethics

This research was conducted according to the guidelines of the Declaration of Helsinki. The Italian branch of this research was approved by the Institutional Review Board of the Centre for Developmental Biology and Reprogramming (CEDEBIOR) of the University of Sassari, Italy (approval no. 03/2021), while the Brazilian branch was approved by the Ethical Committee of Federal University of Amapá (UNIFAP) (approval no. 3.238.424 02). Informed consent was obtained from all subjects that received the NPO Treatment, involved in the research.

Study design

In this research, we retrospectively analyzed the effects of the REAC NPO treatment over time. In the practice of the clinical centers affiliated with the Society of Neuro-Psycho-Physical Optimization with REAC technology (SONC), the stability of the disappearance of the FD after the REAC NPO administration is checked every time the patient comes back for control without time programming.

The coordinator of the research launched an initial invitation to the members of SONC, to participate in this research project. Following this invitation, a webinar was organized to illustrate the objectives and how to collect the data.

Clinicians who agreed to participate were asked to provide an anonymized grid, where the following data were reported: gender and age of the patient at the date of the FD assessment, date of the NPO administration, and verification of its effectiveness. Each operator analyzed their database of the patients treated with NPO, and each patient was investigated to verify whether over time the functional dysmetria had recurred in the checks following the administration of the NPO treatment. For the data collection, each patient was entered only once, considering only the control date furthest from the administration of the NPO treatment. This method allows highlighting for each patient the longest stability over time of the effects of NPO on functional dysmetria. The datasheet was collected and sent to the coordinator, then aggregated and subsequently processed.

Population

The data presented in this research were retrospectively collected from a population comprising patients who showed up at the SONC clinical centers involved in the project, over 18 years starting from September 2003.

The data collection concerned a population of 29,794 subjects, 11,981 males and 17,813 females. The average age was 45.07 for males and 50.33 for females. The overall average age was 47.70 (Table 1).

Cureus

POPULATION = 29,794	
Males	Females
11,981 (40.21%)	17,813 (59.79%)
AVERAGE AGE OF THE POPULATION = 47.70	
Males	Females
45.07	50.33

TABLE 1: Population demographics

In the clinical practice of the SONC centers, all patients undergo the assessment for FD. All the patients showed FD and received the REAC NPO treatment, regardless of the symptomatology, and pathology initially referred.

Functional dysmetria assessment

Functional dysmetria is a phenomenon not perceptible to the subject, but evident to the examiner during the motor task of the assessment maneuver [20,21]. The assessment of FD is an evaluation of neurological semiology aimed at observing the presence of dysfunctional adaptive motor behavior and not a morphological modification.

To assess the presence of FD, the subjects were asked to lie down on the medical examination table and move from the supine to the sitting position [20,21]. During the execution of this motor task, the examinator places his hands lightly on the femoral quadriceps of the subject, and in such a way as to perceive muscle contraction and movement, without opposing and taking care that the nails of his left and right thumbs are perfectly aligned.

In the transition of the subjects from the supine to the sitting position, Rinaldi and Fontani observed a progressive misalignment of the two thumbs. This motor task allows the operator to perceive and highlight the asymmetric activation (dysmetria) of symmetrical muscle groups, such as the quadriceps muscles.

REAC neuro postural optimization treatment

The REAC NPO neuromodulation is a preprogrammed single session treatment. It is administered by applying the tip of the metallic REAC asymmetric conveyer probe (ACP) to a specific area of the ear located at the superior edge of the lower third of the scapha of the ear pavilion, according to the NPO protocol [20,21,23].

Study replicability

The REAC device's parameter for administering the NPO treatment is set by the manufacturer of the device and cannot be modified by the operators. The methods of administration have been extensively described in previous papers [15,20,21,23].

The models of the equipment used were the CRM, BENE 101, and BENE 110 models, which are equivalent to each other for the administration of REAC NPO. All the devices were produced by ASMED S.r.l, Florence, Italy.

Results

The data collected on the outcomes of the verification of the stable recovery of the FD are distributed over 18 years and show a higher concentration of cases in the first 36 months after the administration of the NPO treatment, while the cases tend to rarefy as we move away from the date of administration (Table 2).

Months											
Time slot	1	2	3	4	5	6	7	8	9	10	11
Number of patients	2410	2508	976	832	261	1455	272	222	256	153	108
% Stable effects	100%	100%	100%	100%	100%	100%	100%	100%	100%	100%	100%
Years											
Time slot	1	2	3	4	5	6	7	8	9	10	11
Number of patients	3809	3662	2680	1972	1829	1329	732	790	654	637	438
% Stable effects	100%	100%	100%	100%	100%	100%	100%	100%	100%	100%	100%
Time slot	12	13	14	15	16	17	18				
Number of patients	424	382	366	277	250	217	189				
% Stable effects	100%	100%	100%	100%	100%	100%	100%				

TABLE 2: Distribution of patients by time slots and percentage of patients with stable disappearance of FD

This phenomenon can be explained by the physiological tendency to abandonment by patients, who gradually thin out follow-up visits until they don't show up anymore. This explains why out of a total population of 29,794 subjects, the data collected 18 years after the administration of the NPO relate only to 189 subjects (as seen above in Table 2). However, the percentage of patients who presented the stable disappearance of FD is equal to 100% in all the time slots of the data collection (Table 2).

Discussion

Any environmental interaction, even unconscious, can only take place through the nervous system, regardless of its evolutionary level [36,37]. For this reason, the changes produced by exposome [17], environmental stress, or allostatic overload are present in all living beings [19,36] and can be observed even as morphological asymmetry phenomena [38,39].

This phenomenon is known as FA [1], and in scientific literature, it is described as a morphological asymmetry resulting from errors in the development of normally symmetrical bilateral traits under stressful conditions [2-4].

Based on the phenomenon of FA, which is already considered an epigenetic measure of environmental stress, Rinaldi and Fontani developed the concept of FD, a phenomenon arising upstream of the FA. The FD is an adaptive motor behavior [24] and is detectable with a specific semiological assessment as an asymmetrical activation of symmetrical muscle groups during a motor task. This dysfunctional motor behavior does not originate from lesions or pathologies of the cerebellum, but from a dysfunctional response to environmental stressors by the cerebellum and neural circuits.

From these premises, the authors studied how to act on the nervous system to allow the recovery of the correct adaptive mechanisms and they conceived the radio electric asymmetric conveyer (REAC) technology for neurobiological stimulation treatments, aimed at counteracting the dysfunctional effects induced by the adaptive phenomenon [21,29-35].

Among the REAC neurobiological stimulation protocols, the treatment called neuro postural optimization (NPO) was devised.

In previous studies conducted with fMRI, NPO treatment was able to re-modulate the activation of the brain areas involved in a motor task, increasing the efficiency and competence of the activation pattern, as highlighted by the blood-oxygen-level-dependent (BOLD) signal level modification [20,21,23]. A higher BOLD signal was observed in the brain areas where the greater neural activity requires an increase in oxygen consumption, which in turn is detectable through an increase of deoxyhemoglobin in brain blood flow.

After the NPO treatment, the greater functional efficiency of the brain dynamics is detectable during fMRI through a reduction of BOLD signal in areas not competent for the motor task performed and a reduction in the extension and degree of their activation since the activation is concentrated in specifically competent areas [20,21].

In other previous studies, the NPO treatment has been shown to induce a long-lasting disappearance of FD through the induction of greater functional efficiency in the execution of motor tasks [20,21], even in subjects suffering from neurodegenerative diseases such as Alzheimer's [40-42].

In the present research, a single REAC NPO treatment session showed to be effective in 100% of the treated subjects and the disappearance of FD lasted up to 18 years.

Retrospective studies have important limitations in the quality and quantity of data available for analysis since data was rarely collected per the needs of the study. But in this study the assessment of FD was performed as it would have been done in a prospective study as the characteristics of the subjects included, the data to be collected, and the outcomes to be measured were defined before the study was carried out.

Conclusions

According to the conceptuality formulated by Rinaldi and Fontani, the verified correlation between the REAC NPO treatment and the stable duration of the disappearance of the FD can be explained as optimization of some functional components of the adaptation processes. The authors correlated the concept of FD as a phenomenon of dysfunctional adaptive origin to the role of the cerebellar functions and other neurodynamics, involved not only in motor control but also in emotional, affective, cognitive, relational control, and stress response.

Based on the hypothesis of the mechanism of action of REAC technology, the NPO treatment is supposed to induce a stable improvement of the neurodynamics and greater efficiency and competence of the brain areas activated in the motor task. This effect would find its clinical demonstration in the long-lasting recovery of FD, likely resulting from a better neurobiological response capacity of the subject to the exposome, and therefore from better control of the symmetry of the adaptive motor behavior.

Additional Information

Disclosures

Human subjects: Consent was obtained or waived by all participants in this study. Institutional Review Board of the Centre for developmental biology and reprogramming (CEDEBIOR) of University of Sassari, Italy. issued approval 03/2021. This research was conducted according to the guidelines of the Declaration of Helsinki. The Italian branch of this research was approved by the Institutional Review Board of the Centre for developmental biology and reprogramming (CEDEBIOR) of the University of Sassari, Italy, with the number 03/2021, while the Brazilian branch was approved by the Ethical Committee of Federal University of Amapá (UNIFAP) with the number 3.238.424 02. Informed consent was obtained from all subjects that received the NPO Treatment, involved in the research. Animal subjects: All authors have confirmed that this study did not involve animal subjects or tissue. Conflicts of interest: In compliance with the ICMJE uniform disclosure form, all authors declare the following: Payment/services info: All authors have declared that no financial support was received from any organization for the submitted work. Financial relationships: All authors have declared that they have no financial relationships at present or within the previous three years with any organizations that might have an interest in the submitted work. Intellectual property info: SR and VF are the authors of the REAC patent. Other relationships: AR and CR are daughters of SR and VF.

Acknowledgements

We thank Dr. Alessandra Cappelli and Dr. Eng. Matteo Lotti Margotti for their precious cooperation.

References

- Parsons PA: Fluctuating asymmetry: an epigenetic measure of stress. Biol Rev Camb Philos Soc. 1990, 65:131-145. 10.1111/j.1469-185x.1990.tb01186.x
- Deleon VB: Fluctuating asymmetry and stress in a medieval Nubian population . Am J Phys Anthropol. 2007, 132:520-534. 10.1002/ajpa.20549
- Furlow FB, Armijo-Prewitt T, Gangestad SW, Thornhill R: Fluctuating asymmetry and psychometric intelligence. Proc Biol Sci. 1997, 264:823-829. 10.1098/rspb.1997.0115
- 4. Ozener B: Does urban poverty increase body fluctuating asymmetry? . Coll Antropol. 2011, 35:1001-1005.
- Lemke SB, Schnorrer F: Mechanical forces during muscle development. Mech Dev. 2017, 144:92-101. 10.1016/j.mod.2016.11.003
- Steinem C, Janshoff A: SENSORS | Piezoelectric Resonators. Encyclopedia of analytical science (second edition). Worsfold P, Townshend A, Poole C (ed): Elsevier, Oxford; 2005. 269-276. 10.1016/B0-12-369397-7/00556-2
- Rajabi AH, Jaffe M, Arinzeh TL: Piezoelectric materials for tissue regeneration: a review. Acta Biomater. 2015, 24:12-23. 10.1016/j.actbio.2015.07.010
- Wojnar R: Piezoelectric nanomaterials for biomedical applications. Piezoelectric phenomena in biological tissues. Ciofani G, Menciassi A (ed): Springer, Berlin, Heidelberg, New York; 2012. 173-185. 10.1007/978-3-642-28044-3_6
- 9. Kwon J, Cho H: Piezoelectric heterogeneity in collagen type I fibrils quantitatively characterized by

piezoresponse force microscopy. ACS Biomater Sci Eng. 2020, 6:6680-6689. 10.1021/acsbiomaterials.0c01314

- Cerrolaza M, Duarte V, Garzón-Alvarado D: Analysis of bone remodeling under piezoelectricity effects using boundary elements. Journal of Bionic Engineering. 2017, 14:659-671.
- 11. Manto M: Mechanisms of human cerebellar dysmetria: experimental evidence and current conceptual bases. J Neuroeng Rehabil. 2009, 6:10. 10.1186/1743-0003-6-10
- 12. Holmes G: The symptoms of acute cerebellar injuries due to gunshot injuries . Brain. 1917, 40:461-535. 10.1093/brain/40.4.461
- Timmann D, Diener HC: Coordination and ataxia. Textbook of clinical neurology (third edition). Goetz CG (ed): 2007. 307-325. 10.1016/B978-141603618-0.10017-7
- 14. Schmahmann JD: Disorders of the cerebellum: ataxia, dysmetria of thought, and the cerebellar cognitive affective syndrome. J Neuropsychiatry Clin Neurosci. 2004, 16:367-378. 10.1176/jnp.16.3.367
- 15. Adamaszek M, D'Agata F, Ferrucci R, et al.: Consensus paper: cerebellum and emotion. Cerebellum. 2017, 16:552-576. 10.1007/s12311-016-0815-8
- Argyropoulos GP, van Dun K, Adamaszek M, et al.: The cerebellar cognitive affective/Schmahmann syndrome: a task force paper. Cerebellum. 2020, 19:102-125. 10.1007/s12311-019-01068-8
- 17. Erzin G, Gülöksüz, S: The exposome paradigm to understand the environmental origins of mental disorders . Alpha Psychiatry. 2021, 22:171-176. 10.5152/alphapsychiatry.2021.21307
- Gheorghe D: Investigating the effects of psychosocial stress on cerebellar function. School of Psychology, University of East Anglia, 2018.
- McEwen BS, Wingfield JC: The concept of allostasis in biology and biomedicine. Hormones and behavior . Horm Behav. 2003, 43:2-15. 10.1016/s0018-506x(02)00024-7
- Rinaldi S, Mura M, Castagna A, Fontani V: Long-lasting changes in brain activation induced by a single REAC technology pulse in Wi-Fi bands. Randomized double-blind fMRI qualitative study. Sci Rep. 2014, 4:5668. 10.1038/srep05668
- Mura M, Castagna A, Fontani V, Rinaldi S: Preliminary pilot fMRI study of neuropostural optimization with a noninvasive asymmetric radioelectric brain stimulation protocol in functional dysmetria. Neuropsychiatr Dis Treat. 2012, 8:149-154. 10.2147/NDT.S29971
- Barmack NH, Pettorossi VE: Adaptive balance in posterior cerebellum. Front Neurol. 2021, 12:635259. 10.3389/fneur.2021.635259
- Rinaldi S, Fontani V, Castagna A: Brain activity modification produced by a single radioelectric asymmetric brain stimulation pulse: a new tool for neuropsychiatric treatments. Preliminary fMRI study. Neuropsychiatr Dis Treat. 2011, 7:649-654. 10.2147/NDT.S26123
- Crivelli D, Peviani V, Salvato G, Bottini G: Exploring the interaction between handedness and body parts ownership by means of the implicit association test. Front Hum Neurosci. 2021, 15:681904. 10.3389/fnhum.2021.681904
- Gheorghe DA, Panouillères MT, Walsh ND: Psychosocial stress affects the acquisition of cerebellardependent sensorimotor adaptation. Psychoneuroendocrinology. 2018, 92:41-49. 10.1016/j.psyneuen.2018.03.013
- Beauchaine TP, Neuhaus E, Zalewski M, Crowell SE, Potapova N: The effects of allostatic load on neural systems subserving motivation, mood regulation, and social affiliation. Dev Psychopathol. 2011, 23:975-999. 10.1017/S0954579411000459
- 27. Canuti M, Giorgi B, Valenti C: The Ilizarov apparatus in the treatment of congenital dysmetria of the leg . Ital J Orthop Traumatol. 1988, 14:483-491.
- 28. Collodel G, Moretti E, Fontani V, et al.: Effect of emotional stress on sperm quality. Indian J Med Res. 2008, 128:254-261.
- 29. Rinaldi S, Fontani V, Aravagli L, Margotti ML: Psychological and symptomatic stress-related disorders with radio-electric treatment: psychometric evaluation. Stress Health. 2010, 26:350-358. 10.1002/smi.1298
- Fontani V, Rinaldi S, Aravagli L, Mannu P, Castagna A, Margotti ML: Noninvasive radioelectric asymmetric brain stimulation in the treatment of stress-related pain and physical problems: psychometric evaluation in a randomized, single-blind placebo-controlled, naturalistic study. Int J Gen Med. 2011, 4:681-686. 10.2147/IJGM.S24628
- Olivieri EB, Vecchiato C, Ignaccolo N, et al.: Radioelectric brain stimulation in the treatment of generalized anxiety disorder with comorbid major depression in a psychiatric hospital: a pilot study. Neuropsychiatr Dis Treat. 2011, 7:449-455. 10.2147/NDT.S23420
- Rinaldi S, Fontani V, Aravagli L, Mannu P, Castagna A, Margotti ML, Rosettani B: Stress-related psychophysiological disorders: randomized single blind placebo controlled naturalistic study of psychometric evaluation using a radio electric asymmetric treatment. Health Qual Life Outcomes. 2011, 9:54. 10.1186/1477-7525-9-54
- 33. Fontani V, Aravagli L, Margotti ML, Castagna A, Mannu P, Rinaldi S: Neuropsychophysical optimization by REAC technology in the treatment of: sense of stress and confusion. Psychometric evaluation in a randomized, single blind, sham-controlled naturalistic study. Patient Prefer Adherence. 2012, 6:195-199. 10.2147/PPA.S29734
- 34. Rinaldi A, Rinaldi C, Coelho Pereira JA, et al.: Radio electric asymmetric conveyer neuromodulation in depression, anxiety, and stress. Neuropsychiatr Dis Treat. 2019, 15:469-480. 10.2147/NDT.S195466
- Rinaldi A, Maioli M, Marins Martins MC, et al.: REAC Non-invasive neurobiological stimulation for mitigating the impact of internalizing disorders in autism spectrum disorder. Adv Neurodev Disord. 2021, 5:446-456. 10.1007/s41252-021-00217-7
- 36. Torrey EF: Brain environment interactions. Psychiatry. 2004, 67:412-415. 10.1521/psyc.67.4.412.56571
- Hari R: From brain-environment connections to temporal dynamics and social interaction: principles of human brain function. Neuron. 2017, 94:1033-1039. 10.1016/j.neuron.2017.04.007
- Breed MD, Moore J: Mating systems. Animal behavior (second edition). Breed MD, Moore J (ed): Academic Press, San Diego; 2016. 357-393. 10.1016/B978-0-12-801532-2.00011-8
- 39. Møller AP: Developmental instability as a general measure of stress . Advances in the study of behavior.

Møller AP, Milinski M, Slater PJB (ed): Academic Press, 1998. 27:181-213. 10.1016/S0065-3454(08)60365-4

- Fontani V, Rinaldi S, Castagna A, Margotti ML: Noninvasive radioelectric asymmetric conveyor brain stimulation treatment improves balance in individuals over 65 suffering from neurological diseases: pilot study. Ther Clin Risk Manag. 2012, 8:73-78. 10.2147/TCRM.S28812
- 41. Olazarán J, González B, López-Álvarez J, et al.: Motor effects of REAC in advanced Alzheimer's disease: results from a pilot trial. J Alzheimers Dis. 2013, 36:297-302. 10.3233/JAD-130077
- Olazarán J, González B, Osa-Ruiz E, et al.: Motor effects of radio electric asymmetric conveyer in Alzheimer's disease: results from a cross-over trial. J Alzheimers Dis. 2014, 42:325-332. 10.3233/JAD-140417