Experimental Physiology - Symposium Report

Pharmacogenomics of neuroimmune interactions in human psychiatric disorders

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There is bidirectional communication between the brain and the immune system. Overproduction of interleukin-L3 (IL-L3) leads to systemic inflammatory response syndrome (SIRS). The crucial role of IL-1\(\beta\) in inflammation has been highlighted by studies performed in caspase-1 knockout mice (casp1"/"), transgenic mice that lack mature IL-1/3 and survive lethal doses of lypopolysaccharide (LPS). We have previously shown that IL-1/3, its receptor IL-1 receptor I (IL-1RI) and caspase-1 are expressed within the brain. Moreover, we documented that peripherally injected LPS triggers a specific spatiotemporal pattern of expression of IL-1/3 mRNA within the brain, suggesting that IL-1/3 could be a major regulator of the central inflammatory cascade. Therefore, we studied brain transcriptional patterns that occur during LPS-induced SIRS in wild-type and casp1-/- mice. We showed patterns of gene expression in wild-type and casp1"/" mice that included differential expression of several genes, such as those for cytokines, chemokines, nitric oxide synthase 2 and cyclo-oygenase 2. A key component of the neuroimmune-endocrine axis that is increased by IL-1 β is corticotrophin releasing hormone (CRH). We found increased response to antidepressants in patients homozygous for the GAG haplotype of CRH receptor-1. Our results support the hypotheses that the CRH receptor-1 gene and possibly other genes in stress-inflammatory pathways are involved in the response to antidepressant treatment. Since dysregulation of the neuroimmune-endocrine axis appears to be one of the fundamental biological mechanisms that underlie psychiatric disorders, our findings might contribute to increase the understanding of the molecular pathways that are altered in these diseases.

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The field of neuroimmunomodulation has evolved dramatically during the last 20 years. It is now very clear that peripheral cytokines modulate brain functions such as regulation of the hypothalamic-pituitary-adrenal (HPA) axis, fever, feeding, sickness behaviour and sleep (Vitkovic et al. 2000). Peripheral cytokines influence brain activity through at least four different routes: peripheral organs synthesize and release cytokines that, by paracrine action, act on their receptors present in nerve fibres of the autonomic nervous system to modulate brain function: (2) circulating cytokines act within the brain vasculature to increase the expression of prostanoids and nitric oxide, which in turn diffuse through the brain parenchyma to trigger the inflammatory cascade; (3) cytokines might signal into the brain through specific areas that lack the blood-brain barrier (BBB), such as

the circunventricular organs; or (4) alternatively, cytokines might enter the brain through a saturable transport mechanism. One of the most potent and well-described pro-inflammatory cytokines that modulates physiological and pathophysiological processes within the brain is interleukin-1\beta (IL-1\beta; Licinio & Wong, 1997\alpha; Vitkovic er \alpha L 2000; Allan & Rothwell, 2003; Elenkov \u00e4 al. 2005).

Systemic administration of IL-1β produces several behavioural changes that include decreased locomotor activity, exploration and feeding (Swiergiel & Dunn, 2006). The crucial role exerted by IL-1β during lypopolysaccharide (LPS)-induced inflammation was previously demonstrated in knockout (KO) and transgenic mice with altered components of the IL-1β network (Li er al. 1995; Hirsch et al. 1996). Mice with a KO of the caspase-1 (casp1) gene, a cysteine-protease that cleaves

Neuroimmune Interactions In Neurologic And Psychiatric Disorders

Valentin A. Pavlov, Niccolo Terrando

Neuroimmune Interactions In Neurologic And Psychiatric Disorders:

Neuro-Immune Interactions in Neurologic and Psychiatric Disorders P. Patterson, C. Kordon, Y. Christen, 2012-12-06 The idea that the brain is an immune privileged site has perhaps served to slow our realization that the intact brain can generate its own inflammatory reactions. These responses can be to peripheral infection or they can arise from local internal causes for instance as a response to stress or to the se vere changes in neuronal activity in seizure or the loss of oxygen in stroke We are also becoming increasingly aware of the contribution of local inflam matory reactions to certain neurodegenerative diseases such as Alzheimer's In fact evidence is accumulating that inflammatory processes disease AD contribute to the progression of AD suggesting the possibility of using cur rently available or novel anti inflammatory agents to interfere with this terri ble disease Correlations are also being made between inflammatory signs and mental illness which is a new frontier of research This book presents the current state of knowledge in a variety of areas relevant to neuro immune interactions Neuroglia in Neurologic and Psychiatric Disorders, Part II ,2025-03-28 Neuroglia in with particular attention to AD Neurologic and Psychiatric Disorders Part II discusses glia relative to neurologic and psychiatric disorders emphasizing cross disease glial cell dysfunctions Driven by advances in genetics transcriptomics electrophysiologic and imaging techniques as well as by the increase in knowledge on neuropathologic features and underlying disease mechanisms this book bridges the gap between neuroglia basic science and neuroglia clinical research Provides an overview of neuroglia biology Identifies neuroglia dysfunction in neurologic and psychiatric disorders Covers ALS dementia Alzheimer Parkinson Huntington diseases and more Includes TBI Stroke COVID Encompasses disorders of mood eating sleep anxiety schizophrenia autism and more

Neuro-immune interaction in CNS injury and disease Bo Li, Guangzhi Ning, Ruiping Wang, 2023-07-24 Rosenberg's Molecular and Genetic Basis of Neurological and Psychiatric Disease Roger N. Rosenberg, Juan M. Pascual, 2014-10-28 Rosenberg s Molecular and Genetic Basis of Neurologic and Psychiatric Disease Fifth Edition provides a comprehensive introduction and reference to the foundations and key practical aspects relevant to the majority of neurologic and psychiatric disease A favorite of over three generations of students clinicians and scholars this new edition retains and expands the informative concise and critical tone of the first edition This is an essential reference for general medical practitioners neurologists psychiatrists geneticists and related professionals and for the neuroscience and neurology research community. The content covers all aspects essential to the practice of neurogenetics to inform clinical diagnosis treatment and genetic counseling Every chapter has been thoroughly revised or newly commissioned to reflect the latest scientific and medical advances by an international team of leading scientists and clinicians. The contents have been expanded to include disorders for which a genetic basis has been recently identified together with abundant original illustrations that convey and clarify the key points of the text in an attractive didactic format Previous editions have established this book as the leading tutorial reference on neurogenetics Researchers will find great value in the coverage of genomics animal models and diagnostic

methods along with a better understanding of the clinical implications Clinicians will rely on the coverage of the basic science of neurogenetics and the methods for evaluating patients with biochemical abnormalities or gene mutations including links to genetic testing for specific diseases Comprehensive coverage of the neurogenetic foundation of neurological and psychiatric disease Detailed introduction to both clinical and basic research implications of molecular and genetic understanding of the brain Detailed coverage of genomics animal models and diagnostic methods with new coverage of evaluating patients with biochemical abnormalities or gene mutations Controlling Biochemical Weapons A. Kelle, K. Nixdorff, M. Dando, 2006-05-26 Scientific and technological is currently transforming the problem of preventing biological warfare and biological terrorism Examples from the areas of immunology the neurosciences and the neuroendocrine immune system are used to show the magnitude of the problem The book outlines the measures required to control biochemical Neuro-Immune Interactions in Inflammation and Autoimmunity Valentin A. Pavlov, Niccolo weapons today Terrando, 2018-07-24 The nervous system plays an important role in the regulation of immunity and inflammation On the other hand unbalanced immune responses in inflammatory and autoimmune conditions may have a deleterious impact on neuronal integrity and brain function Recent studies have characterized neural pathways communicating peripheral inflammatory signals to the CNS and brain and spinal cord derived circuitries controlling various innate and adaptive immune responses and inflammation A prototypical neural reflex circuit that regulates immunity and inflammation is the vagus nerve based inflammatory reflex Ongoing research has revealed cellular and molecular mechanisms underlying these neural circuits and indicated new therapeutic approaches in inflammatory and autoimmune disorders Pharmacological and bioelectronic modulation of neural circuitry has been successfully explored in preclinical settings of sepsis arthritis inflammatory bowel disease obesity driven disorders diabetes and other diseases These studies paved the way to successful clinical trials with bioelectronic neuronal modulation in rheumatoid arthritis and inflammatory bowel disease Dysregulated release of cytokines and other inflammatory molecules may have a severe impact on brain function Brain inflammation neuroinflammation imbalances in brain neuronal integrity and neurotransmitter systems and cognitive impairment are characteristic features of post operative conditions sepsis liver diseases diabetes and other disorders characterized by immune and metabolic dysregulation Derangements in cytokine release also play a pivotal role in depression Characteristic brain reactive antibodies in autoimmune conditions including systemic lupus erythematosus and neuromyelitis optica significantly contribute to brain pathology and cognitive impairment These studies and the simultaneous characterization of neuro protective cytokines identified new therapeutic approaches for treating neurological complications in inflammatory and autoimmune disorders This Frontiers Research Topic is a forum for publishing research findings and methodological and conceptual advances at the intersection of immunology and neuroscience We hope that presenting new insight into bi directional neuro immune communication in inflammation and autoimmunity will foster further collaborations and facilitate

the development of new efficient therapeutic strategies *Neuro-Immune Connections to Enable Repair in CNS Disorders* Tim Vanmierlo, Bieke Broux, Jack Van Horssen, Niels Hellings, 2020-10-28 This eBook is a collection of articles from a Frontiers Research Topic Frontiers Research Topics are very popular trademarks of the Frontiers Journals Series they are collections of at least ten articles all centered on a particular subject With their unique mix of varied contributions from Original Research to Review Articles Frontiers Research Topics unify the most influential researchers the latest key findings and historical advances in a hot research area Find out more on how to host your own Frontiers Research Topic or contribute to one as an author by contacting the Frontiers Editorial Office frontiers in org about contact **Biomarkers of Neurological** and Psychiatric Disease Sabine Bahn, Paul Guest, 2011-11-09 Published since 1959 International Review of Neurobiology is a well known series appealing to neuroscientists clinicians psychologists physiologists and pharmacologists Led by an internationally renowned editorial board this important serial publishes both eclectic volumes made up of timely reviews and thematic volumes that focus on recent progress in a specific area of neurobiology research This volume reviews existing theories and current research surrounding the movement disorder Dyskinesia Leading authors review state of the art in their field of investigation and provide their views and perspectives for future research Chapters are extensively referenced to provide readers with a comprehensive list of resources on the topics covered All chapters include comprehensive background information and are written in a clear form that is also accessible to the non specialist Neuroimmune Diseases Hiroshi Mitoma, Mario Manto, 2024-11-18 This Springer Major Reference Work provides a detailed translational overview of neuroimmune diseases for neuroscientists and clinicians clarifying the pathological mechanisms underlying neuroimmune diseases and building a comprehensive bridge between the latest research findings and their clinical implications in daily practice The material is presented in two steps The first section comprises a review of the pathogenic actions of immune cells in brain diseases Here the authors discuss the mechanisms through which immune cells disrupt the functions of nerve cells The second section explores the ways in which the brain becomes dysfunctional due to impaired nerve cell function Based on pathogenesis diagnostic and therapeutic strategies are discussed for each clinical category. The new edition features updated chapters throughout including coverage of new findings new diagnostic criteria and therapeutic algorithms Neuroimmune Diseases 2nd edition also includes new chapters with a translational approach The book will be invaluable for use in clinical practice of neuroimmune diseases Advances in Nervous System Research and Application: 2012 Edition ,2012-12-26 Advances in Nervous System Research and Application 2012 Edition is a ScholarlyBrief that delivers timely authoritative comprehensive and specialized information about Nervous System in a concise format The editors have built Advances in Nervous System Research and Application 2012 Edition on the vast information databases of ScholarlyNews You can expect the information about Nervous System in this eBook to be deeper than what you can access anywhere else as well as consistently reliable authoritative informed and relevant The content of Advances in Nervous System Research and

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explores the medical and philosophical implications of neuroscience in the mental health field Frontiers in Synaptic Plasticity: Dendritic Spines, Circuitries and Behavior Alberto A. Rasia-Filho, Rochelle S. Cohen, Oliver von Bohlen und Halbach, 2016-09-22 The term synaptic plasticity is a broad concept which is studied with a variety of experimental approaches One focus is the impact of changes in synaptic neuronal and glial morphology on brain circuitry and behavior In this regard unique animal models have been key to the study of affective and social behaviors and neurological and psychiatric diseases However there is a paucity of compilations directed toward the correlation of alterations in synaptic structure with various physiological and behavioral paradigms This Frontiers Research Topic will therefore serve as an exciting forum for the exchange of novel hypotheses and data and an important resource and reference for investigators studying synaptic and brain plasticity as well as those in related fields Immune Ethics Walter Glannon, 2023-08-03 The immune system maintains homeostasis within human organisms and protects them from pathogenic threats But sometimes it cannot provide this protection on its own and vaccines may be necessary to ensure our health and survival Immune functions can become dysregulated and result in autoimmune disease or multi system damage Pharmacological and genomic interventions may activate or modulate immune functions to prevent these outcomes This Element is an analysis and discussion of some of the ethical implications of these interventions After describing the main components of innate and adaptive immunity and how it might be enhanced it considers the potential benefit and harm from vaccines against addiction and viruses immunotherapy for cancer neuroimmunomodulating agents to prevent or treat neurodevelopmental and neurodegenerative diseases and gene editing of immunity to enable xenotransplantation and prevent infectious disease The Element concludes with an exploration of a possible outcome of natural competition between humans and microbes

Neuroinflammatory and Oxidative/Nitrosative Pathways in Neuropsychiatric and Neurological Diseases and Their Possible Neuropharmacological Regulation, Volume I Javier R. Caso, Marta P. Pereira, Borja Garcia-Bueno, 2022-11-03 Postdoctoral Research Fellowship Opportunities National Institutes of Health (U.S.), 1995

Pharma-Nutrition Gert Folkerts, Johan Garssen, 2014-07-08 Natural products and functional medical foods are now widely acknowledged as having an effect on the microbiome of the intestine which in turn influences the outcome of certain disease This book reviews the impact and effects of natural products and functional medical foods nutritional programming on disease management specifically focusing on diseases related to 1 Inflammation and Immunity 2 Cancer COPD and Cachexia 3 Allergy and 4 Brain Neuro Immune Hippocrates said let medicine be thy food and food be thy medicine While most of us are familiar with Hippocrates famous words we admit that in recent times the disciplines of pharma and nutrition have evolved separately Today with the ever growing burden of diseases in modern society we see a convergence of the two in relation to specific disease prevention and treatment This re discovered common ground between the complementary values of pharma and nutrition can be conceptualized in the term pharma nutrition Various chapters in the book review the



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