

A case report of fatal familial insomnia with cerebrospinal fluid leukocytosis during the COVID-19 epidemic and review of the literature

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ABSTRACT

Fatal familial insomnia (FFI) is a rare autosomal dominant genetic neurodegenerative disease. Generally, FFI patients will develop rapidly progressive dementia, sleep disturbance, autonomic dysfunction, and so on. Cerebrospinal fluid examination of FFI patients normally shows no obvious abnormalities. Here, we report a young male patient who was diagnosed with FFI during the COVID-19 epidemic. Clinical symptoms include psychobehavioral abnormality, cognitive decline, sleep disturbance, and autonomic dysfunction. No abnormalities were found in routine examinations after admission. However, the number of white blood cells in the cerebrospinal fluid increased. Though the patient was treated with anti-infection and immunotherapy, the symptoms were not relieved. A lumbar puncture was performed again, and it was found that the total Tau protein in the cerebrospinal fluid was elevated, and PET results showed that brain metabolism decreased. Finally, a genetic test was used to confirm the diagnosis of FFI. This case suggests that patients with FFI may also have elevated white blood cells in cerebrospinal fluid and timely detection of Tau protein in cerebrospinal fluid is helpful for early identification of FFI. And precise diagnosis relies on genetic testing.

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KEYWORDS

Clinical features; CSF analysis; fatal family insomnia; PET-CT; PRNP; Tau protein

Introduction

FFI, a rare autosomal dominant disease, is associated with the D178N-129M missense mutations of the prion protein gene (PRNP) on chromosome 20, and the main pathological feature is scrapie prion protein (PrP^{Sc}) deposition in the brain [1,2]. The average onset age of FFI is about 51 years old, and the course of the disease ranges from 8 to 72 months [3]. Clinical symptoms of FFI patients are variable. Typically, FFI patients are accompanied with rapidly progressive dementia, sleep disturbance, autonomic dysfunction, and so on. Eventually, it will lead to death [4].

In this article, we report one case of a young male patient diagnosed with FFI who had been vaccinated against COVID-19 during the COVID-19 pandemic. This patient had no positive family history but with severe sleep disturbances, progressive cognitive decline, significant changes in autonomic nervous function, and psychobehavioral abnormality. Early detection of cerebrospinal fluid showed increased white blood cells. Multiple magnetic resonance imaging (MRI) and electroencephalography (EEG) examination showed no abnormalities, and the effects of anti-infection and immunotherapy treatment were poor. Finally, the patient was diagnosed with FFI by genetic testing.

Case presentation

The patient, a 28-year-old courier, was admitted to the hospital in July 2021, because of 'Memory impairment and psychotic behaviour disorder for more than one month'. The patient suffered from memory loss one month before admission without obvious incentives. He often made repeated mistakes at work and forgot whether he had eaten. There are often involuntary movements of limbs and frequent changes in his body position during sleep. Later, hallucinations and delusion gradually appeared. When his brother was not at home, he claimed to have seen him. He also said that his child was adopted, and he had another one. The patient had seen a doctor in another hospital before coming to our department. He was diagnosed with 'anxiety disorder' and was treated with 'Olanzapine Tablets 2.5mg/day'. After treatment, the patient's sleep quality has slightly improved, but he often moves his hands and feet around during sleep and tears the quilt with his hands and teeth.

When admitted to the hospital, the patient's body temperature was normal, his consciousness was clear, his facial expression was stolid, and his feeling of time orientation was not good. His short-term memory was impaired. He could not recall three things (ball, flag,

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Scientific Report On Fatal Familial Insomnia

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Neuroimmune Pharmacology and Therapeutics Howard E. Gendelman, Tsuneya Ikezu, 2024-12-26 Bridging neuroscience immunology and pharmacology and bringing together the foremost authorities Neuroimmune Pharmacology and Therapeutics 3rd edition is an invaluable reference and textbook The text discusses the immunology of the nervous system New chapters are offered on innate humoral and cellular immune responses innate and adaptive immunity The additions join each component of the immune response in descriptions for how each affects nervous system function in both health and disease Next discussions of neuropharmacology now include both drug development and delivery into brain subregions to optimize clinical responses This edition features greatly expanded coverage of therapeutics The new extensions have blossomed into focused therapies engaging the immune system directly deploying it for drug delivery attracting the newly evolving field of genetics vaccinations and bioengineering ultimately leading to improved therapeutic disease outcomes All of the revisions in this edition are designed to bring an early trainee together with a skilled clinical and translational scientist to discuss the state of the art in each part of the emerging field of immunity as it affects the nervous system during steady state and disease and how it can be harnessed for therapeutics and clinical benefit **The Age of Scientific Wellness** Leroy Hood, Nathan Price, 2023-04-04 Taking us to the cutting edge of the new frontier of medicine a visionary biotechnologist and a pathbreaking researcher show how we can optimize our health in ways that were previously unimaginable We are on the cusp of a major transformation in healthcare yet few people know it At top hospitals and a few innovative health tech startups scientists are working closely with patients to dramatically extend their healthspan the number of healthy years before disease sets in In The Age of Scientific Wellness two visionary leaders of this revolution in health take us on a thrilling journey to this new frontier of medicine Today most doctors wait for clinical symptoms to appear before they act and the ten most commonly prescribed medications confer little or no benefit to most people taking them Leroy Hood and Nathan Price argue that we must move beyond this reactive hit or miss approach to usher in real precision health a form of highly personalized care they call scientific wellness Using information gleaned from our blood and genes and tapping into the data revolution made possible by AI doctors can catch the onset of disease years before symptoms arise revolutionizing prevention Current applications have shown startling results diabetes reversed cancers eliminated Alzheimer's avoided autoimmune conditions kept at bay This is not a future fantasy it is already happening but only for a few patients and at high cost It's time to make this gold standard of care more widely available Inspiring in its possibilities radical in its conclusions The Age of Scientific Wellness shares actionable insights to help you chart a course to a longer healthier and more fulfilling life

Human Prion Diseases, 2018-06-07 Human Prion Diseases Volume 153 is designed to update the reader on the latest advances and clinical aspects of prion diseases The book is organized into five sections including the pathophysiology of prions and a description of animal and human diseases This is followed by detailed reports on recent advances in diagnosis

strategies for the development of novel anti prion molecules and possible designs of clinical trials in such a rare disease An introductory chapter gives an extensive historical background of prion research with a final chapter highlighting recent progress and more importantly unsolved problems Offers an authoritative overview of prion diseases in humans detailing the pathogenesis of the disease clinical investigations and the diagnosis of both the genetic and acquired forms Provides clarity and context by presenting prion diseases in relation to other neurodegenerative diseases in humans Emphasizes the unique properties of prion diseases and consequent problems they can cause both clinically and in public health terms

Advancing Prion Science Institute of Medicine, Medical Follow-up Agency, Committee on Transmissible Spongiform Encephalopathies: Assessment of Relevant Science, 2003-04-20 In Advancing Prion Science the Institute of Medicine's Committee on Transmissible Spongiform Encephalopathies Assessment of Relevant Science recommends priorities for research and investment to the Department of Defense's National Prion Research Program NPRP Transmissible spongiform encephalopathies TSEs also called prion diseases are invariably fatal neurodegenerative infectious diseases that include bovine spongiform encephalopathy commonly called mad cow disease chronic wasting disease scrapie and Creutzfeldt Jakob disease To develop antemortem diagnostics or therapies for TSEs the committee concludes that NPRP should invest in basic research specifically to elucidate the structural features of prions the molecular mechanisms of prion replication the mechanisms of TSE pathogenesis and the physiological function of prions normal cellular isoform Advancing Prion Science provides the first comprehensive reference on present knowledge about all aspects of TSEs from basic science to the U S research infrastructure from diagnostics to surveillance and from prevention to treatment This report summarizes the progress thus far **The Best American Science and Nature Writing 2020** Michio Kaku, Jaime Green, 2020 Presents an anthology of the best science and nature writing published in the previous year selected from American periodicals

Rosenberg's Molecular and Genetic Basis of Neurological and Psychiatric Disease, Seventh Edition Roger N. Rosenberg, Juan M. Pascual, 2024-10-04 Rosenberg's Molecular and Genetic Basis of Neurologic and Psychiatric Disease Seventh Edition provides a comprehensive introduction and reference to the foundations and key practical aspects relevant to neurologic and psychiatric disease This volume has been thoroughly revised and includes newly commissioned chapters on ethics genetic counselling and gene therapy for the central nervous system disorders A favorite of over four generations of students clinicians and scholars this new edition retains and expands the informative concise and critical tone of the previous edition This is an essential reference for general medical practitioners neurologists psychiatrists geneticists and related professionals and for the neuroscience and neurology research community at large Both volumes combined provide a comprehensive coverage on the neurogenetic foundation of neurological and psychiatric disease This volume provides a detailed introduction on both the clinical and basic research implications of molecular and genetics surrounding the brain Includes new chapters on genomics of human neurological disorders CRISPR and genome engineering *EBOOK:*

Psychology: The Science of Mind and Behaviour, 4e Nigel Holt, Andy Bremner, Ed Sutherland, Michael Vliek, Michael Passer, Ronald Smith, 2019-03-01 EBOOK Psychology The Science of Mind and Behaviour 4e

Dementia 3Ed David Ames, Alistair Burns, John O'Brien, 2005-08-26 The rising prevalence of dementia in the population continues to pose a serious public health challenge in both the developed and the developing world Previous editions of Dementia have become acknowledged as a key gold standard work in this field and have had a genuinely international approach The third edition has been fully revised and updated

Encyclopedia of Neuroscience, Volume 1 Larry R. Squire, 2009-06-12 The Encyclopedia of the Neuroscience explores all areas of the discipline in its focused entries on a wide variety of topics in neurology neurosurgery psychiatry and other related areas of neuroscience Each article is written by an expert in that specific domain and peer reviewed by the advisory board before acceptance into the encyclopedia Each article contains a glossary introduction a reference section and cross references to other related encyclopedia articles Written at a level suitable for university undergraduates the breadth and depth of coverage will appeal beyond undergraduates to professionals and academics in related fields

The Year's Best Science Fiction: Thirty-First Annual Collection Gardner Dozois, 2014-07-15 In the new millennium what secrets lay beyond the far reaches of the universe What mysteries belie the truths we once held to be self evident The world of science fiction has long been a porthole into the realities of tomorrow blurring the line between life and art Now in The Year's Best Science Fiction Thirty First Annual Collection the very best SF authors explore ideas of a new world in the year's best short stories This venerable collection brings together award winning authors and masters of the field such as Robert Reed Alastair Reynolds Damien Broderick Elizabeth Bear Paul McAuley and John Barnes And with an extensive recommended reading guide and a summation of the year in science fiction this annual compilation has become the definitive must read anthology for all science fiction fans and readers interested in breaking into the genre

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The ^AOxford Handbook of Adult Cognitive Disorders Michael L. Alosco, Robert A. Stern, 2019-05-07 The Oxford Handbook of Adult Cognitive Disorders is an up to date scholarly and comprehensive volume that covers most diseases conditions and injuries resulting in impairments in cognitive function in adults The Handbook is appropriate for both clinicians and clinical researchers from advanced trainees to seasoned professionals

Pathogenicity and Drug Resistance of Human Pathogens Saif Hameed, Zeeshan Fatima, 2020-01-23 The book comprehensively discusses the mechanisms of pathogenesis and drug resistance current diagnostics landscape of four key human pathogens bacterial fungal protozoans and viral which are the causes of major infectious diseases It also assesses the emerging technologies for the detection and quantification of these pathogens Further it discusses the novel opportunities to fight against these infectious diseases and to identify pertinent drug targets with novel methodologies It also reviews the current and future insights into the control elimination and eradication of these infectious

diseases Importantly the book discusses the epidemiological characteristics and various challenges in combating Ebola and Influenza diseases Finally the book highlights the growing role of nanotechnology and bioinformatics resources for combating the infectious diseases In summary the book provides the mechanistic insight of the pathogenicity drug resistance therapeutic strategies and identification of the novel drug targets of Mycobacterium tuberculosis Plasmodium Candida Hepatitis C and emerging viral infections Incredible Consequences of Brain Injury Alexander R. Toftness,2022-11-11

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Oxford Textbook of Neuroimaging Massimo Filippi,2015 Part of the Oxford Textbooks in Clinical Neurology series this

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