

Innovations in Inorganic and Materials Chemistry

Ageing, Aluminium and Silicon

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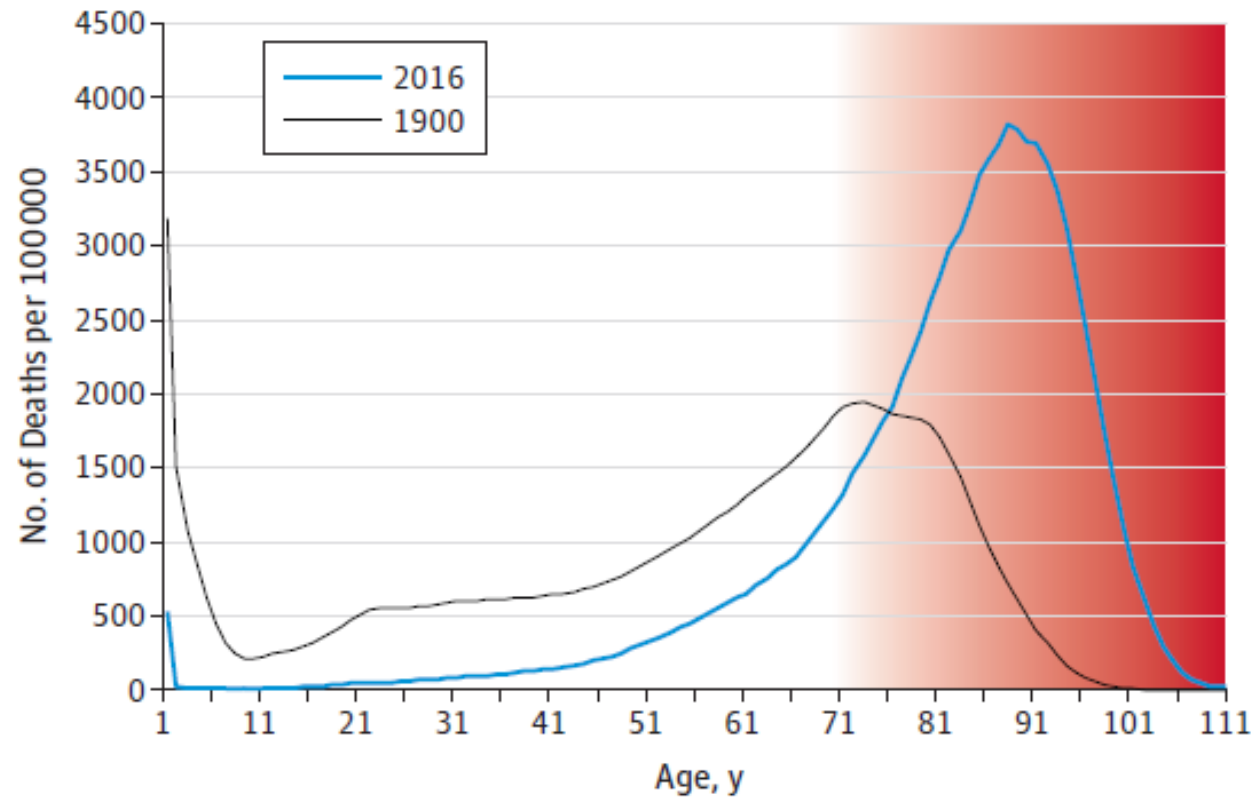
c.exley@keele.ac.uk

<http://www.keele.ac.uk/aluminium/>

<https://www.hippocraticpost.com/?s=Exley>



Figure. Age Distribution of Life Table Deaths for Women in the United States, per 100 000 People, 1900 and 2016



The red zone represents a period in life when the risk of frailty and disability begins to increase rapidly. The goal of aging science is to delay and compress the red zone, which may extend healthy life. Sources: 1900 data from Bell and Miller¹; 2016 data from Human Mortality Database.²

Neurones are the longest-lived cells of the human body and survive aging processes that ravage the remainder of the human body.

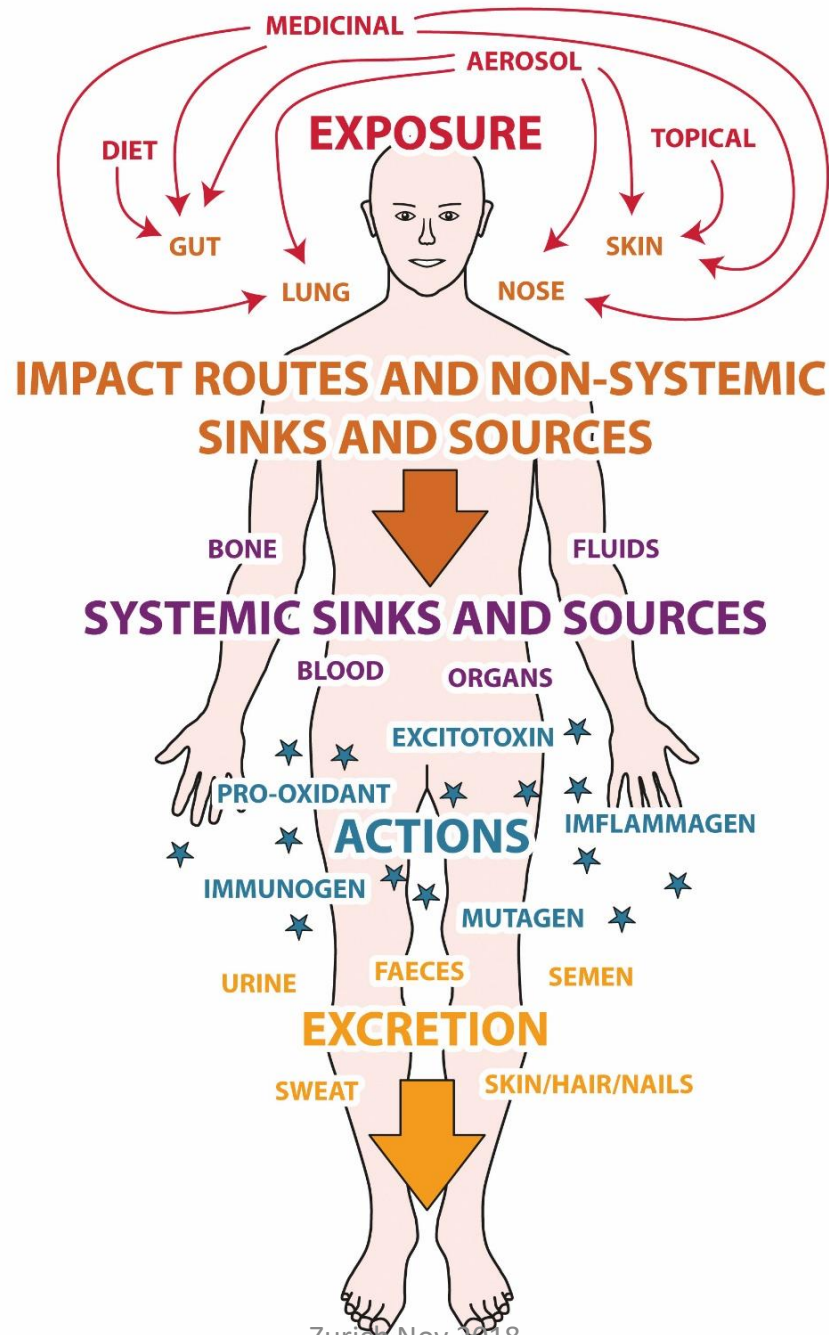
Evolution through natural selection has conferred biochemical advantages upon neurones and the neuronal microenvironment that have in turn enabled human beings to live for longer.

I would contend that the evolution of what is an ostensibly immortal cell line would not have occurred in the presence of biologically available aluminum.

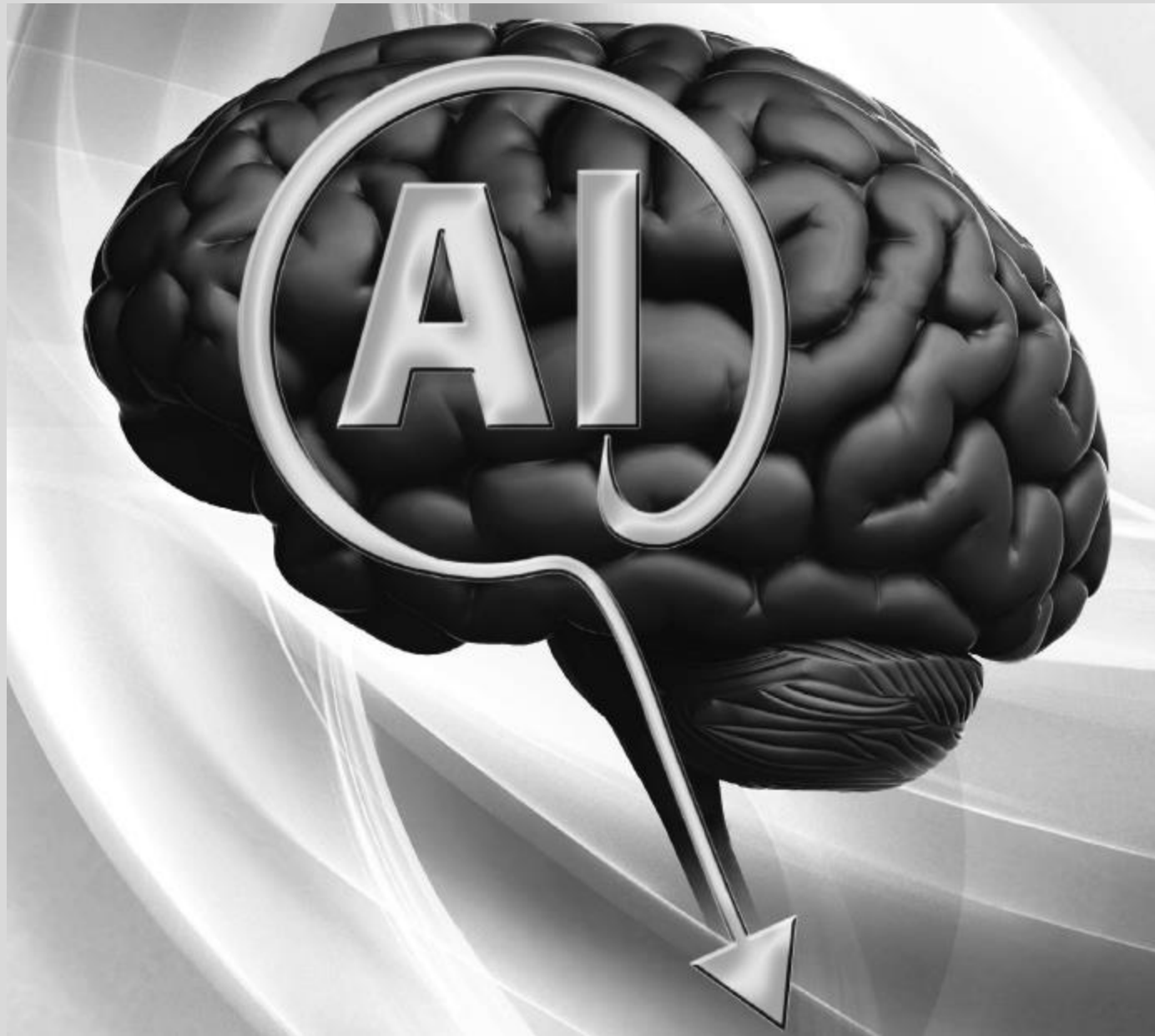
Indeed, the advent of the aluminum age must now have serious consequences for the health and longevity of such a cell line.

The lifespan of neurones predisposes them to a lifetime accumulation of aluminium.

<https://www.frontiersin.org/articles/10.3389/fneur.2014.00212/full>



Zurich Nov. 2018



Zurich Nov 2018

Cite this: *Metallomics*, 2012, **4**, 56–65

www.rsc.org/metallomics

PAPER

Aluminium, iron and copper in human brain tissues donated to the medical research council's cognitive function and ageing study

Emily House,^a Margaret Esiri,^b Gill Forster,^c Paul G Ince^c and Christopher Exley^{*a}

The median Al content of tissues from all **60 brains** (n=713) is 1 $\mu\text{g/g}$ dry wt.

In 52 out of 60 individuals at least one tissue sample exceeded 2 $\mu\text{g Al/g}$ dry wt.

In 41 out of 60 individuals at least one tissue sample exceeded 3.5 $\mu\text{g Al/g}$ dry wt.

Approximately 70% of individuals aged 70 – 103 years had at least one tissue Al content which should be considered as pathological

The Identification of Aluminum in Human Brain Tissue Using Lumogallion and Fluorescence Microscopy

Ambreen Mirza^a, Andrew King^{b,c}, Claire Troakes^c and Christopher Exley^{a,*}

^a*The Birchall Centre, Lennard-Jones Laboratories, Keele University, Staffordshire, UK*

^b*Department of Clinical Neuropathology, King's College Hospital, London, UK*

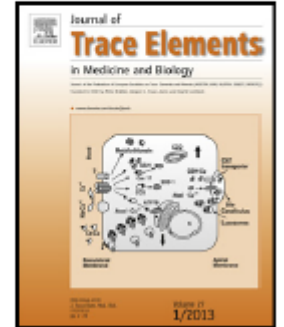
^c*MRC London Neurodegenerative Diseases Brain Bank, Institute of Psychiatry, Psychology and Neuroscience, King's College, London, UK*



Contents lists available at ScienceDirect

Journal of Trace Elements in Medicine and Biology

journal homepage: www.elsevier.com/locate/jtemb



Toxicology

Aluminium in brain tissue in familial Alzheimer's disease

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^b Department Of Clinical Neuropathology, King's College Hospital, London, SE5 9RS, United Kingdom

^c MRC London Neurodegenerative Diseases Brain Bank, Institute of Psychiatry, Psychology and Neuroscience, King's College, London, SE5 8AF, United Kingdom



<https://www.sciencedirect.com/science/article/pii/S0946672X16303777?via%3Dihub>

45000 Views on the Publisher's Website

Zurich, Nov 2018

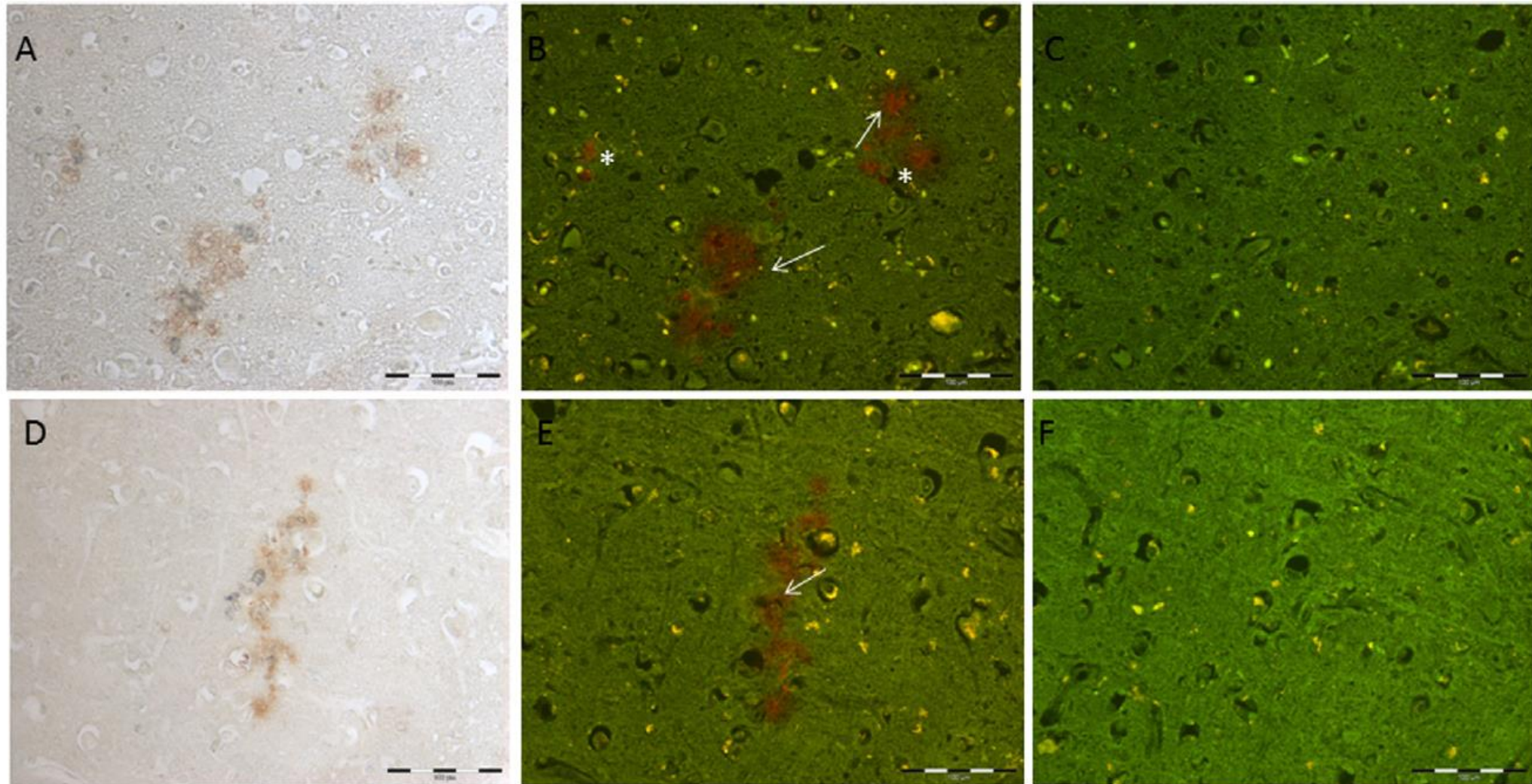


Fig. 1. Representative images of aluminium in frontal cortex. Light (A&D) and fluorescence (B&E) microscopy images of lumogallion-stained sections of frontal cortex. Asterisk label suggested intracellular deposits while arrows show diffuse deposits. Fluorescence microscopy of un-stained adjacent tissue sections (C&F) show autofluorescence. Scale bars are all 100 μm .

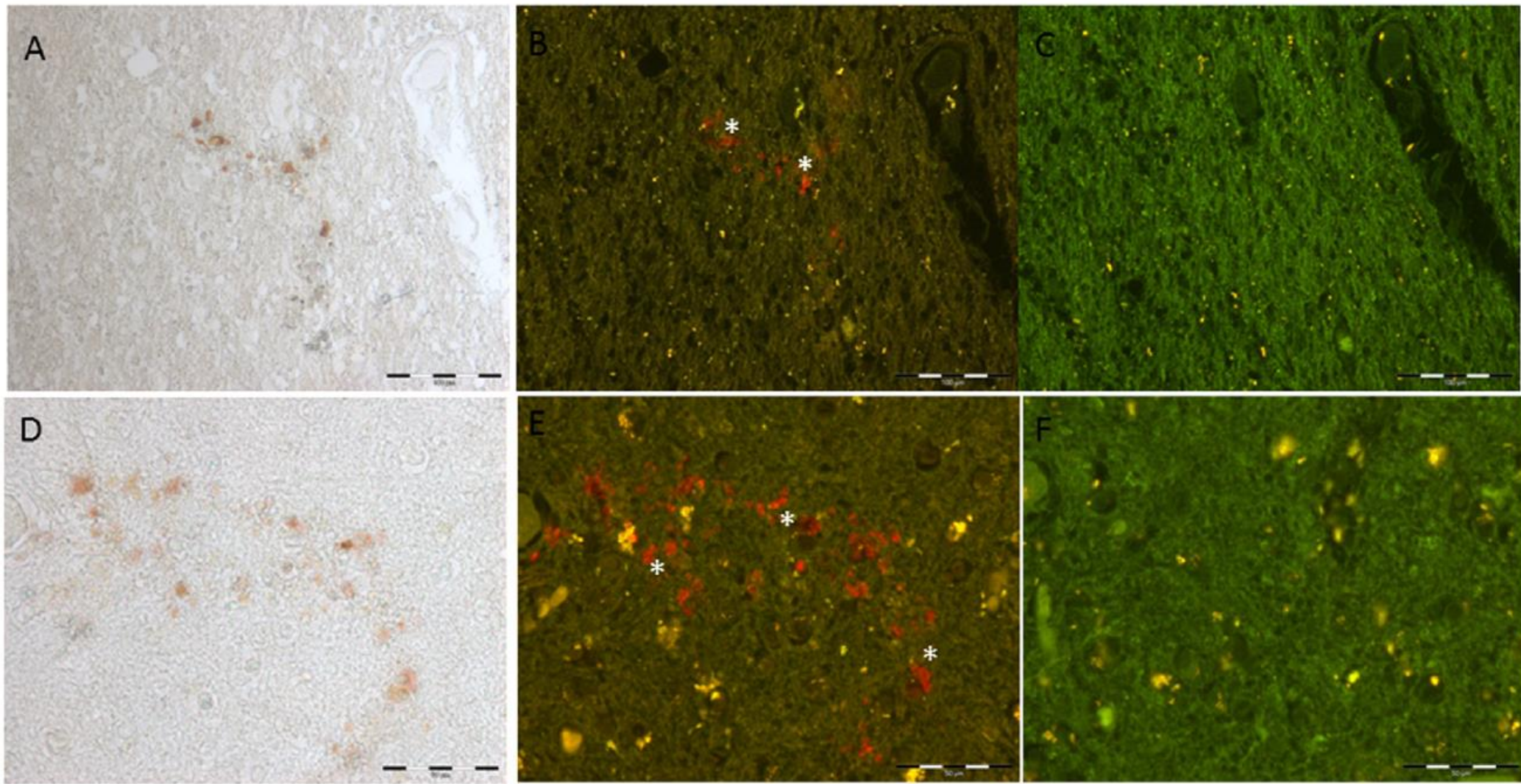


Fig. 2. Representative images of aluminium in parietal cortex. Light (A&D) and fluorescence (B&E) microscopy images of lumogallion-stained sections of frontal cortex. Asterisk label suggested intracellular deposits associated with both living and dead cells. Fluorescence microscopy of un-stained adjacent tissue sections (C&F) show autofluorescence. Scale bars are 100 μm (A–C) and 50 μm (D–F).

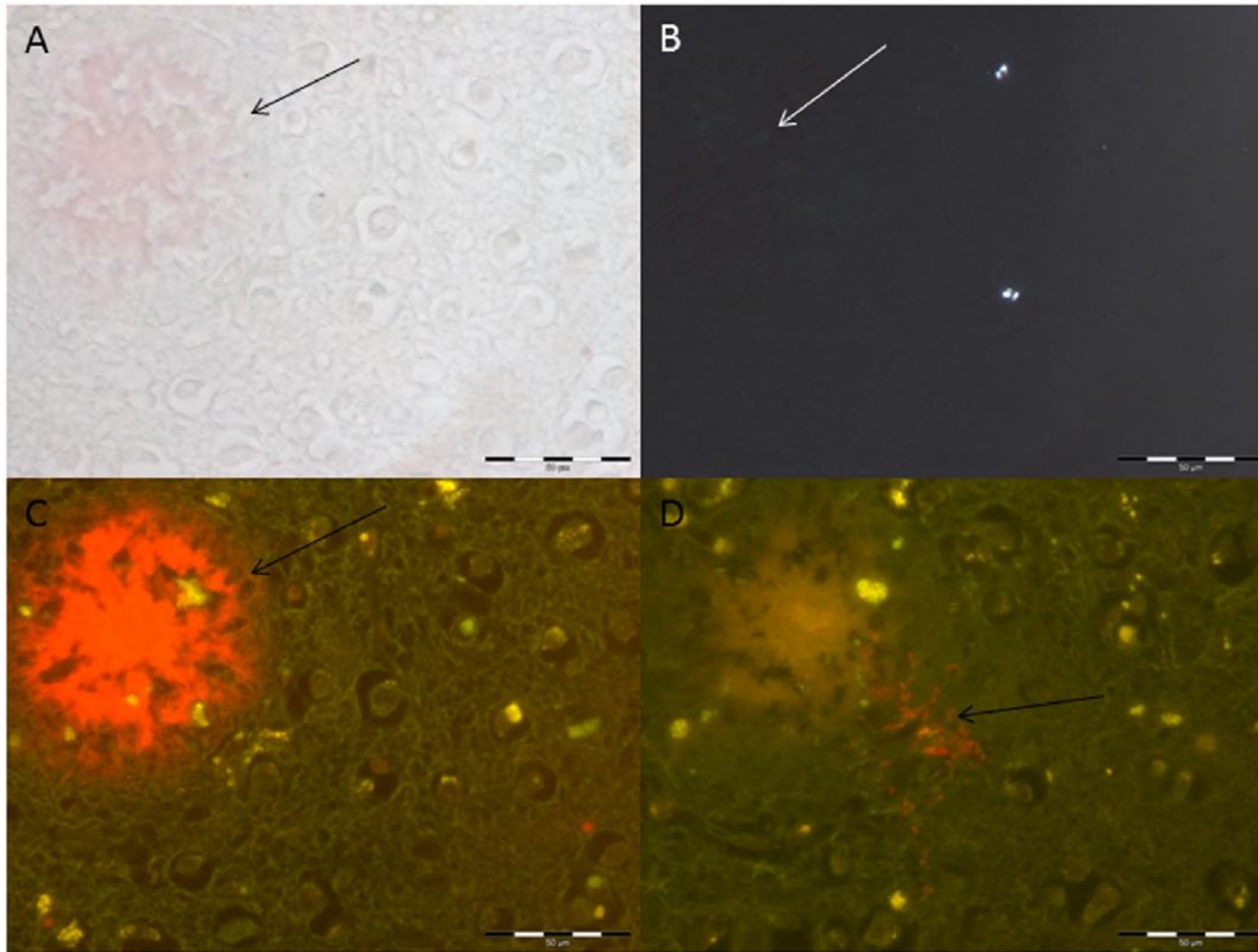


Fig. 4. Co-localisation of amyloid and aluminium in occipital cortex. (A) Light microscopy image of Congo red-stained tissue showing (arrow) senile plaque-like amyloid deposit. (B) Polarising microscopy image of Congo red-stained image showing (arrow) apple-green birefringence characteristic of amyloid in β sheet conformation. (C) Fluorescence microscopy image of Congo red-stained tissue showing (arrow) senile plaque-like amyloid deposit. (D) Fluorescence microscopy image of adjacent section of tissue stained with lumogallion and showing (arrow) significant deposits of aluminium. Scale bars are all 50 μm .

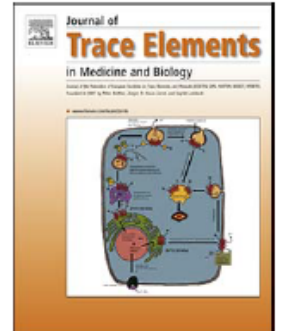


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journal homepage: www.elsevier.com/locate/jtemb



Aluminium in brain tissue in autism

Matthew Mold^a, Dorcas Umar^b, Andrew King^c, Christopher Exley^{a,*}

^a *The Birchall Centre, Lennard-Jones Laboratories, Keele University, Staffordshire, ST5 5BG, United Kingdom*

^b *Life Sciences, Keele University, Staffordshire, ST5 5BG, United Kingdom*

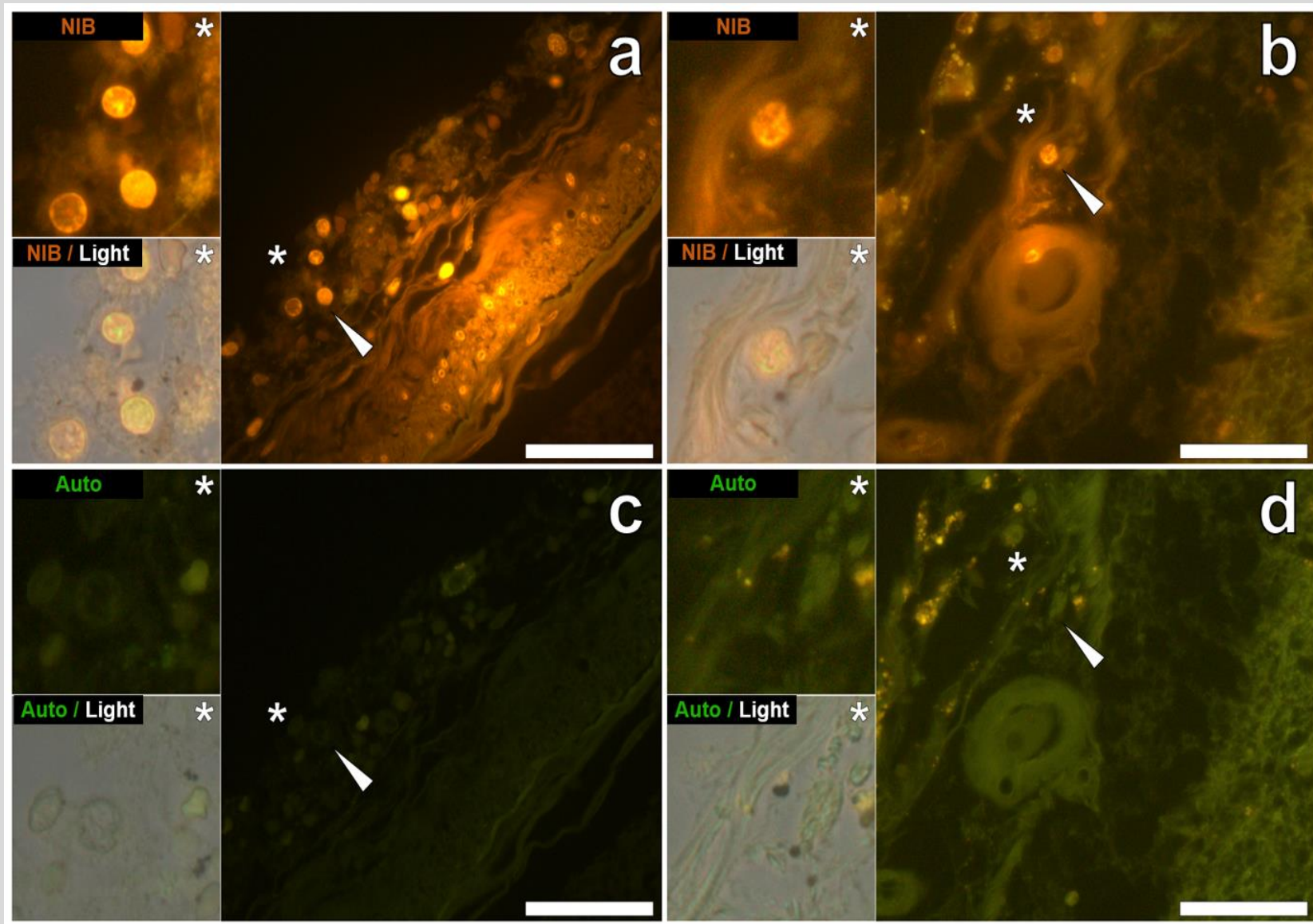
^c *Department of Clinical Neuropathology, Kings College Hospital, London, SE5 9RS, United Kingdom*



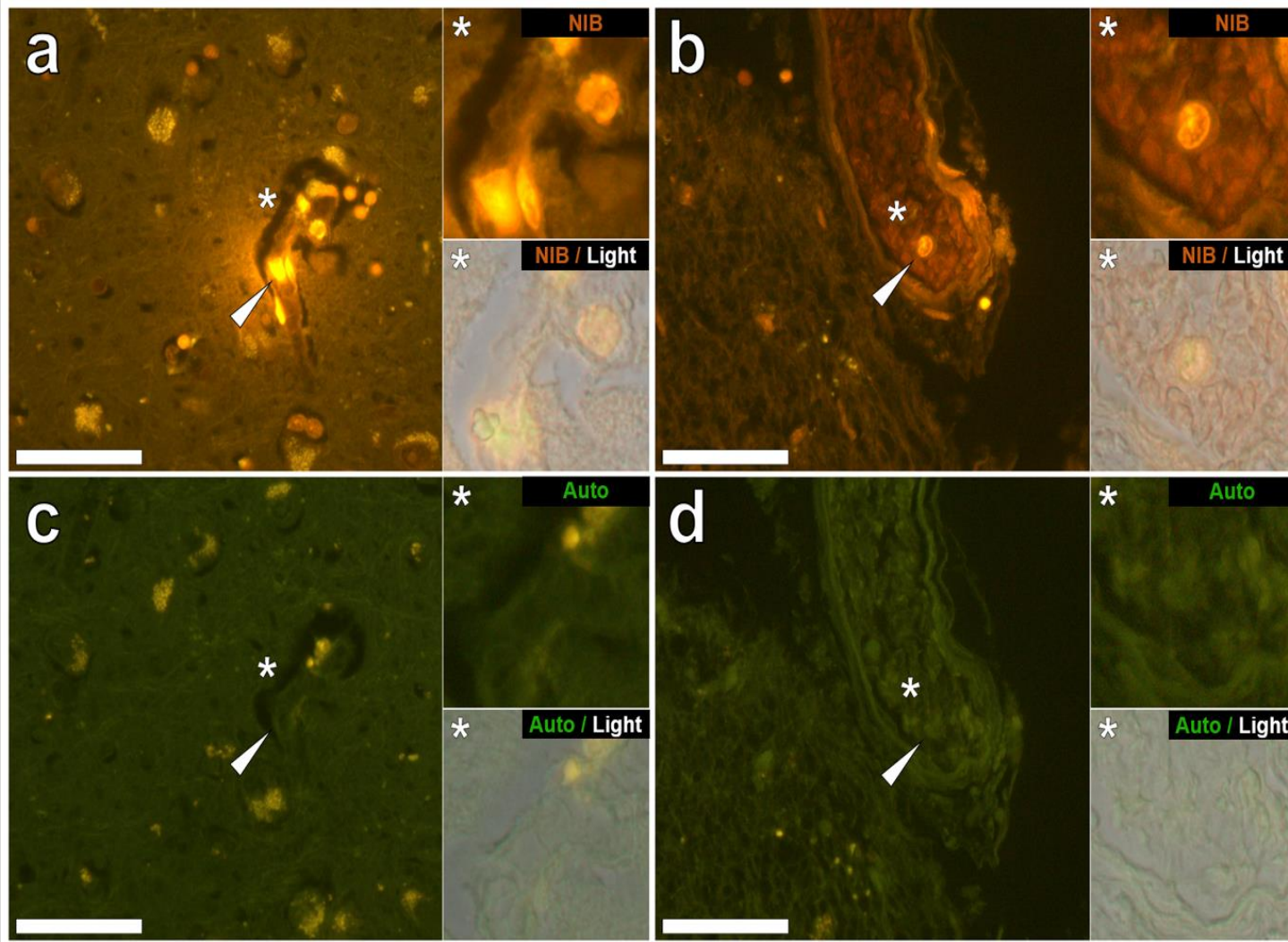
<https://www.sciencedirect.com/science/article/pii/S0946672X17308763>

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Zurich, Nov 2018

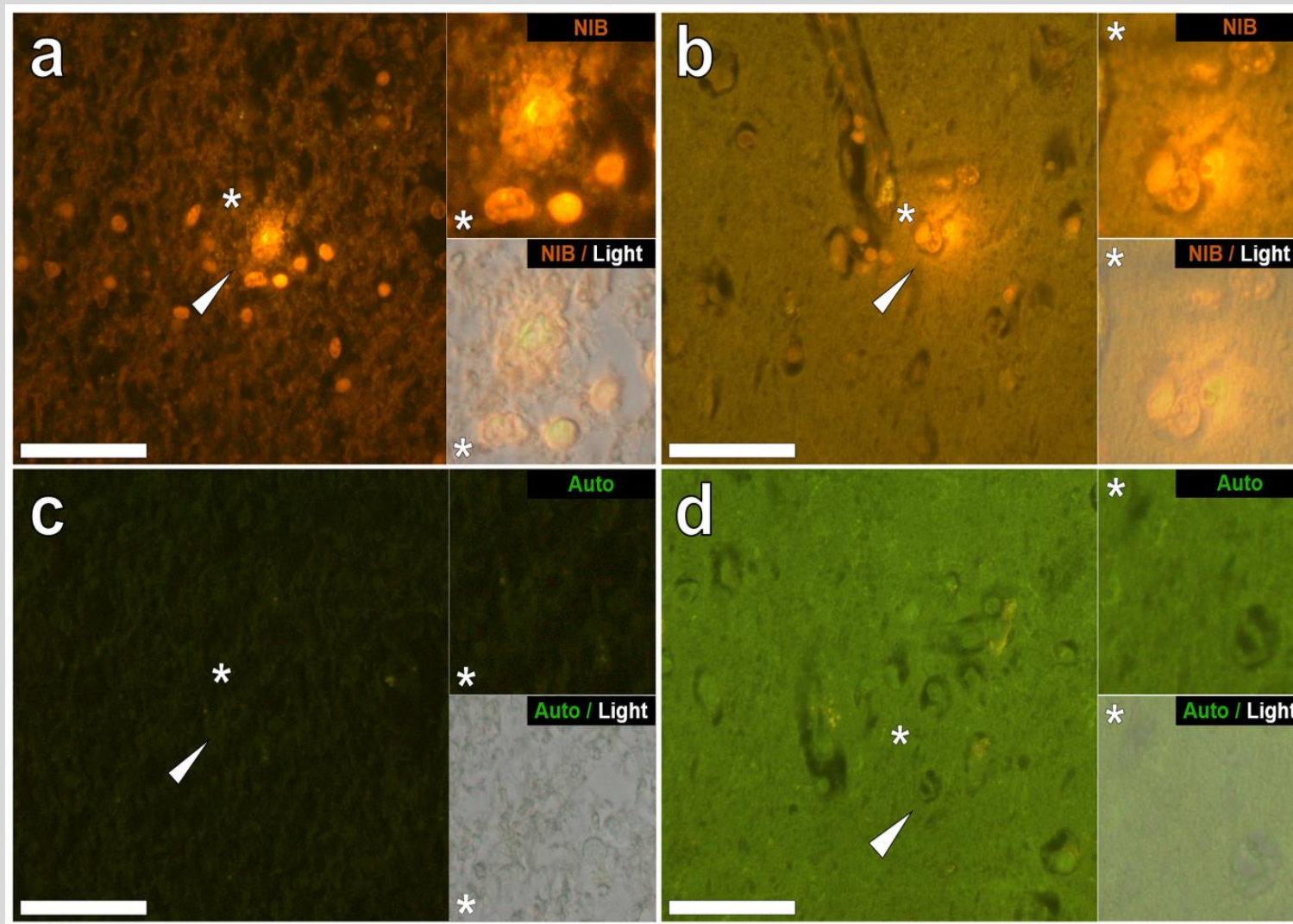


Intrameningeal lumogallion-reactive aluminium identified in the hippocampus (**a & c**) and frontal lobe (**b & d**) of a 50-year-old male donor diagnosed with autism.



Intravasculature lumogallion-reactive aluminium identified in the hippocampus (**a – d**) of a 50-year-old male donor diagnosed with autism.

<https://www.sciencedirect.com/science/article/pii/S0946672X17308763>



Lumogallion-reactive aluminium identified in the hippocampus (**a & c**) and parietal (**b & d**) lobe of a 15-year-old male donor diagnosed with autism.





International Journal of
*Environmental Research
and Public Health*



Article

Aluminium in Brain Tissue in Multiple Sclerosis

Matthew Mold ¹ , Agata Chmielecka ², Maria Raquel Ramirez Rodriguez ¹, Femia Thom ²,
Caroline Linhart ³, Andrew King ⁴ and Christopher Exley ^{1,*} 

<https://www.mdpi.com/1660-4601/15/8/1777>

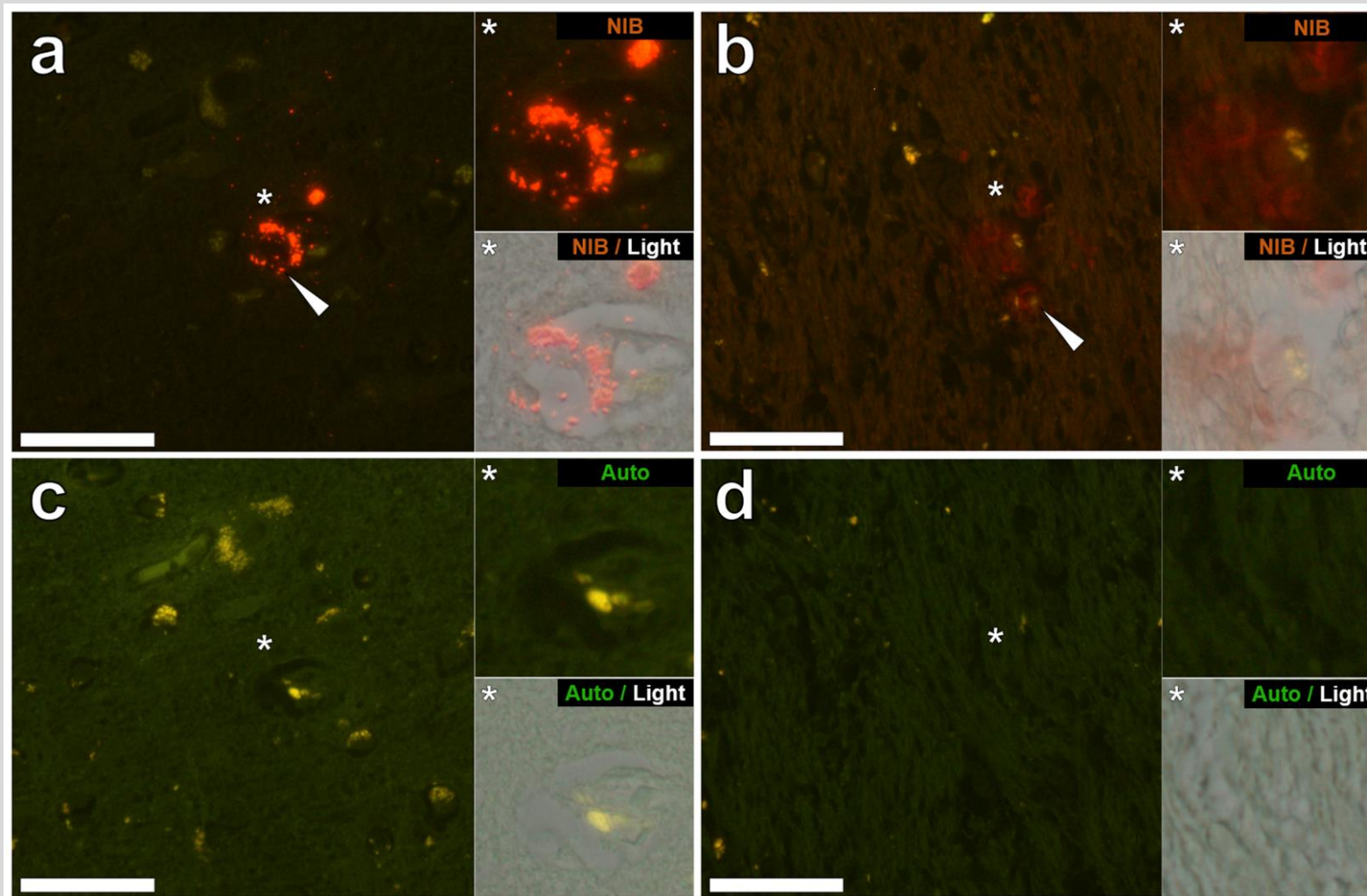


Figure 1. Extracellular aluminium in the frontal lobe and hippocampus of a 56-year-old male donor (MS274), diagnosed with RRMS. (a) Intense orange fluorescence (white arrow) indicating punctate deposits of aluminium was observed in the perivascular region of a small blood vessel in the white matter of the frontal lobe, in close proximity to lipofuscin, identified by yellow fluorescence. (b) Extracellular deposits of aluminium, identified as diffuse orange-red fluorescence, appear co-deposited with lipofuscin (white arrow) in white matter adjacent to the parahippocampal gyrus. (c,d) Autofluorescence of serial sections confirms the identity of aluminium in (a,b) respectively. Upper and lower panels depict magnified inserts of the fluorescence channel and bright field overlay. Magnification x400, scale bar 50 μ m.

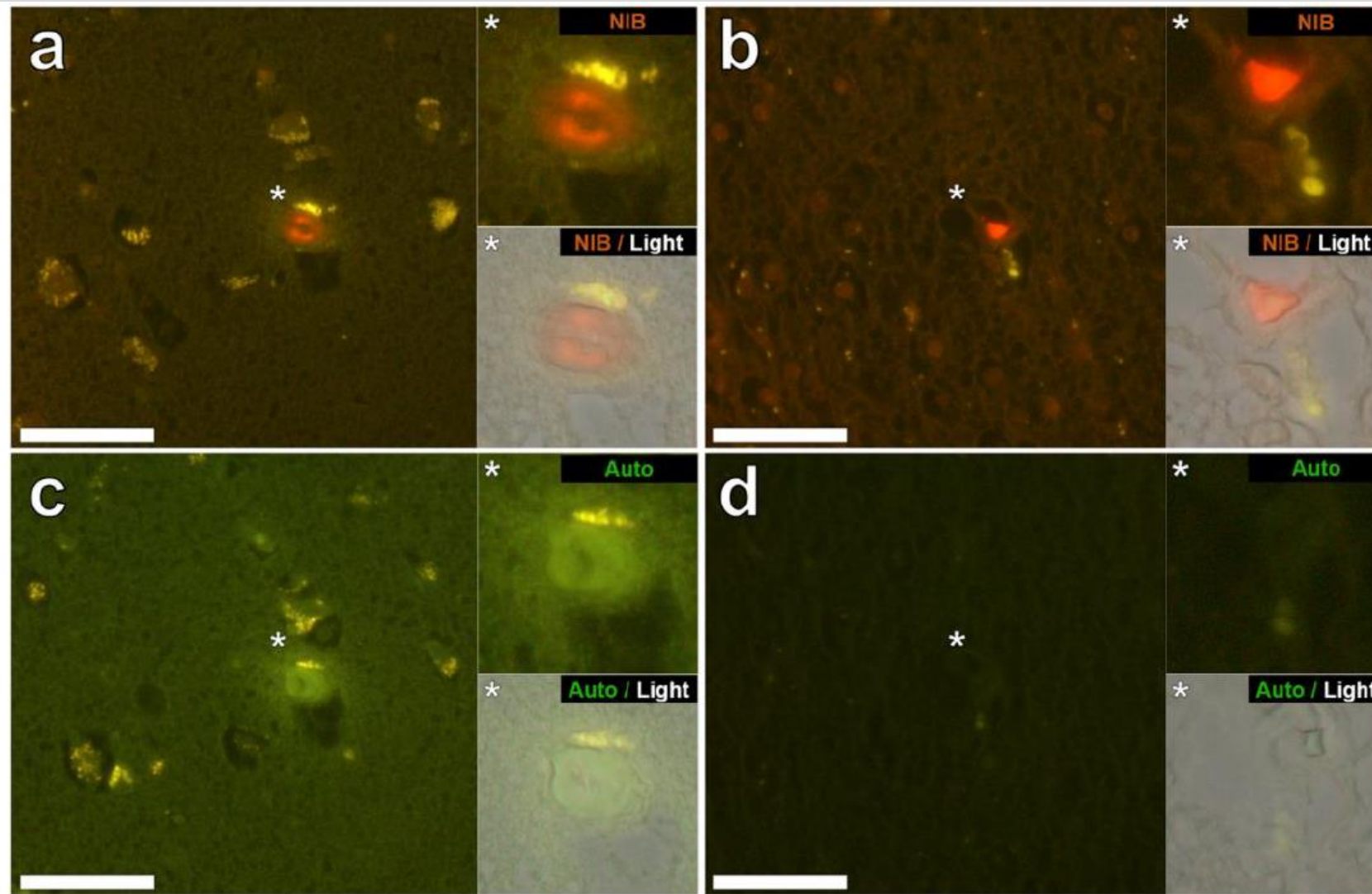
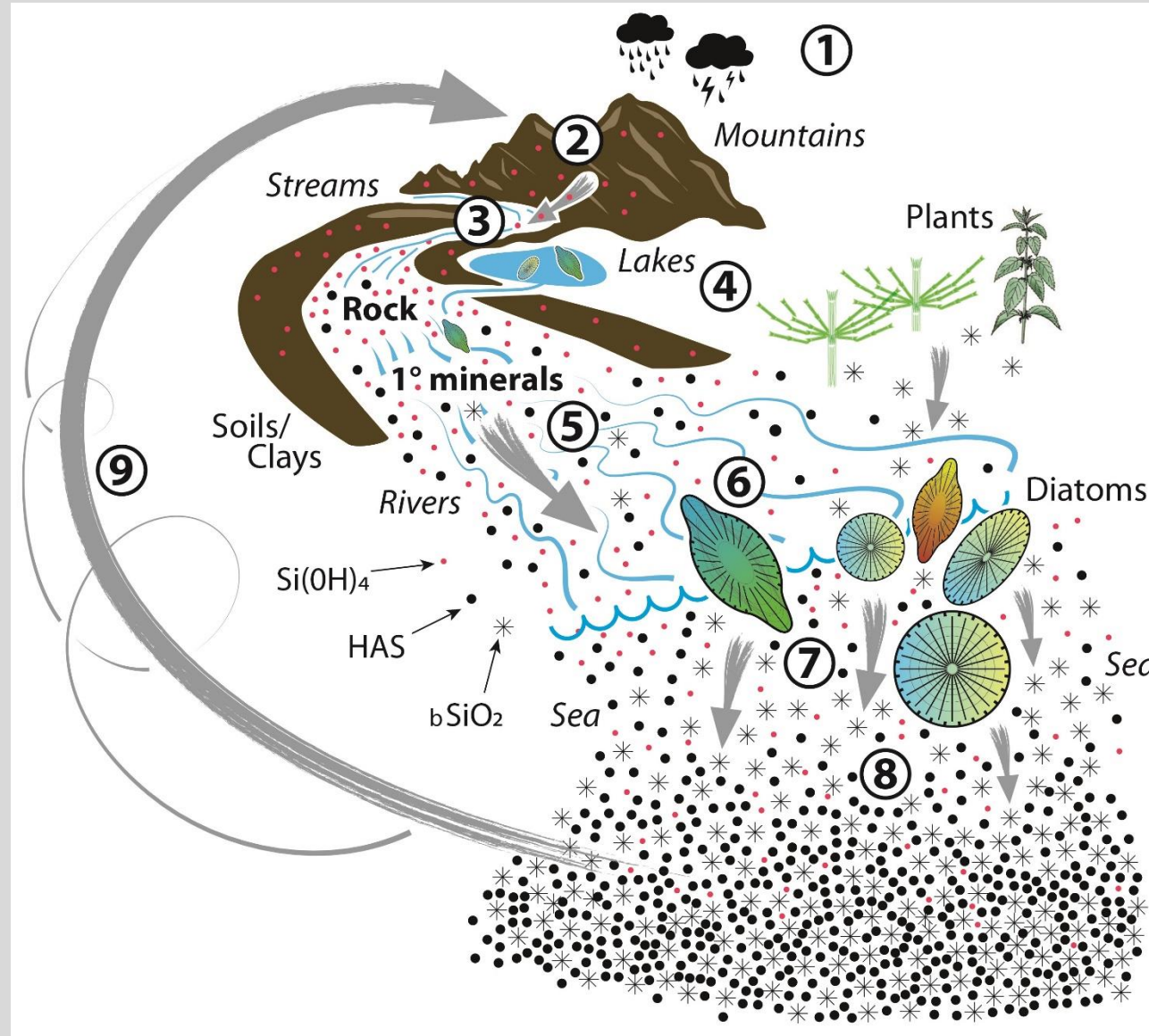


Figure 3. Aluminium in the frontal lobe and hippocampus of a 48-year-old female donor (MS317), diagnosed with SPMS. **(a)** Intense orange aluminium fluorescence was identified in refractile corpora amylacea (or mineralised deposits) in the frontal cortex grey matter). **(b)** Intracellular aluminium was also observed in occasional glial-like cells in the parahippocampal gyrus (white matter). Autofluorescence of serial sections **(c,d)** confirms the identity of aluminium in **(a,b)** respectively. Upper and lower panels depict magnified inserts of the fluorescence channel and bright field overlay. Magnification x400, scale bars: 50 μm .

THE SILICIC ACID CYCLE

The clue to Earth's (healthy) ageing strategy?

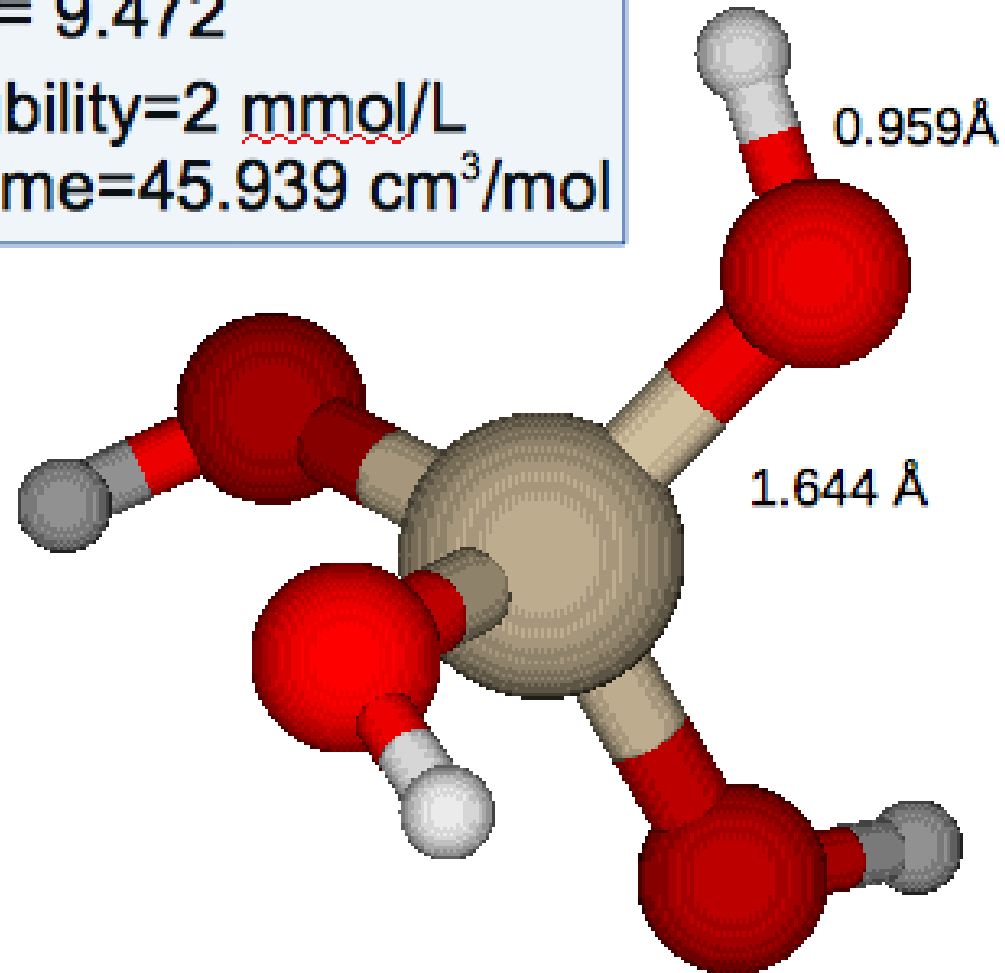


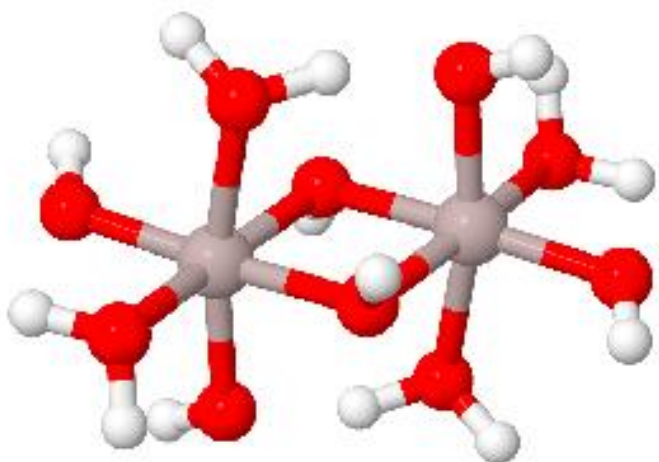
Silicic Acid

$pK_a = 9.472$

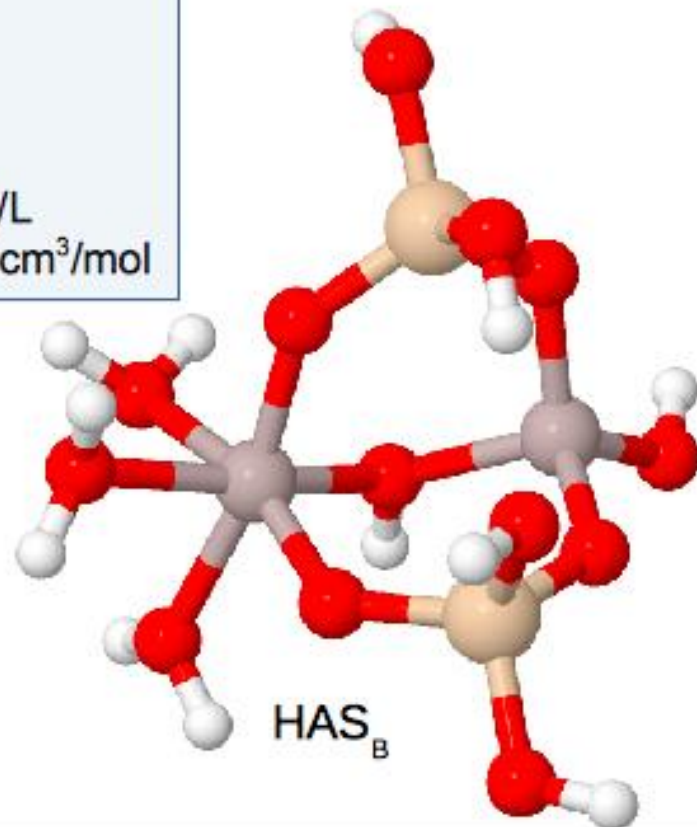
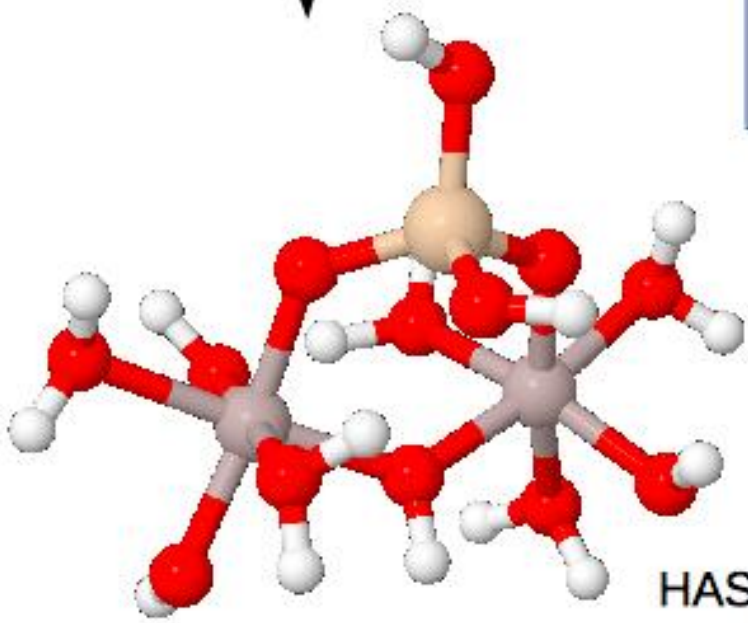
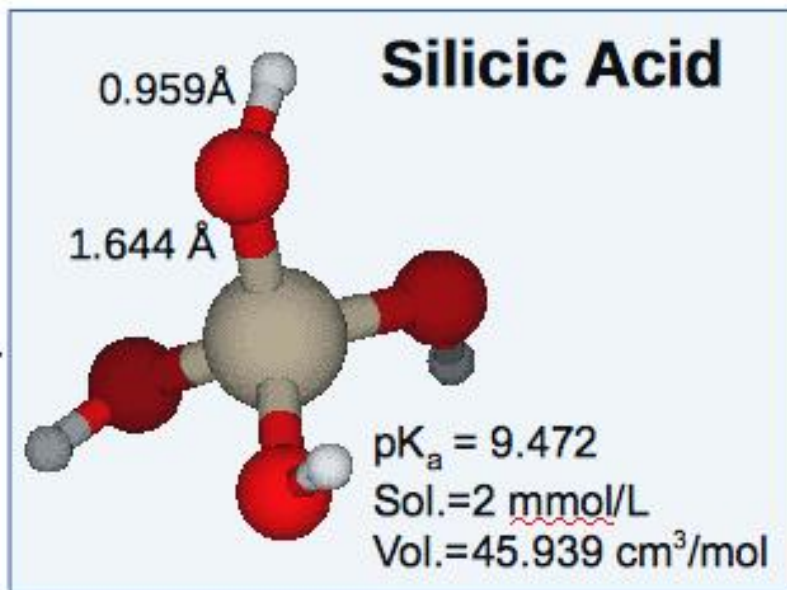
Solubility=2 mmol/L

Volume=45.939 cm^3/mol

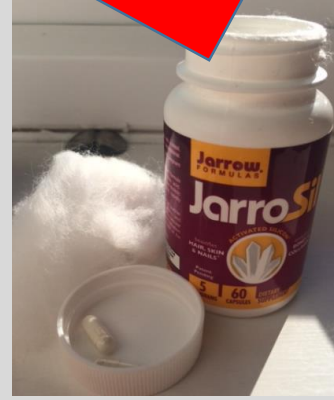




Aluminium Hydroxide



SILICON/SILICA SUPPLEMENTS



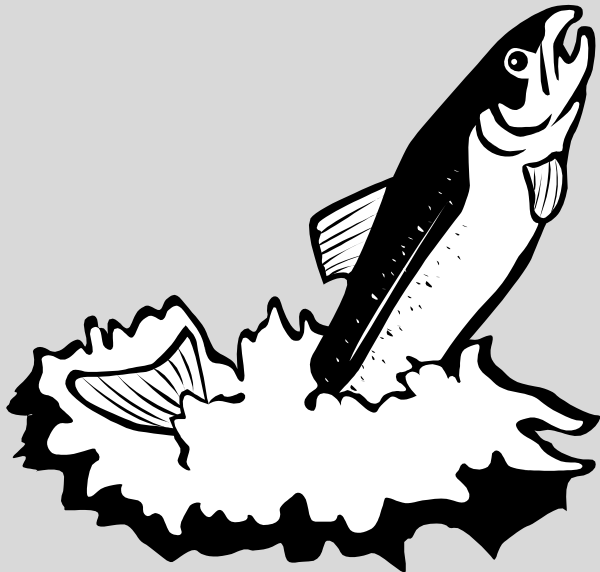
A Bioinorganic Solution to Aluminium-Related Disease?

1989

**Acute toxicity of aluminium to fish
eliminated in silicon-rich acid waters**

J. D. BIRCHALL, C. EXLEY,
J.S. CHAPPELL & M. J. PHILLIPS

***Nature* 338, 146 - 148 (09 March 1989);
doi:10.1038/338146a0**



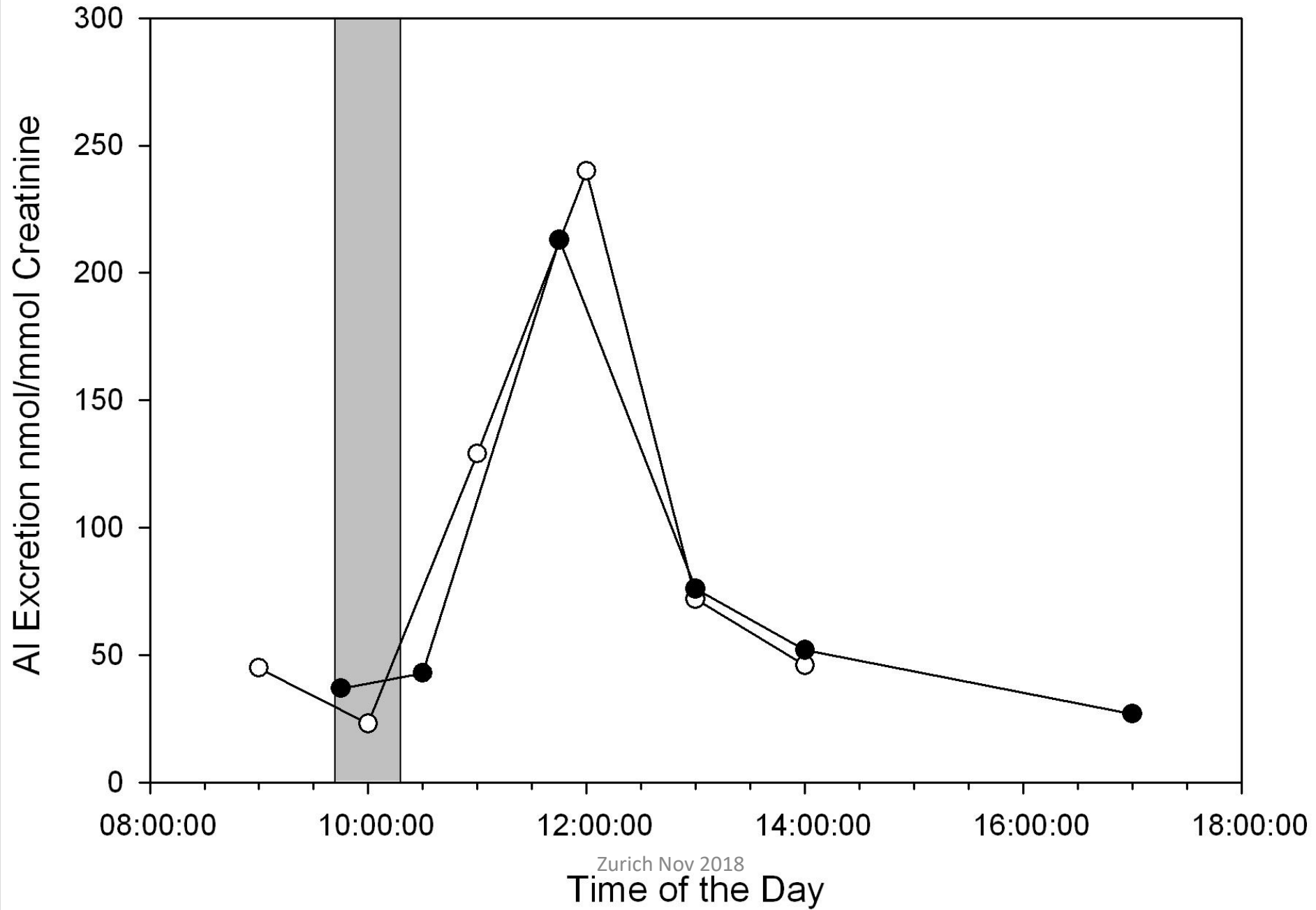
2006

**Non-invasive therapy to reduce the body
burden of aluminium in Alzheimer's
disease**

Christopher Exley, Olga Korchazhkina,
Deborah Job, Stanislav Strekopytov, Anthony
Polwart and Peter Crome

***Journal of Alzheimer's Disease* 10 (2006)
17-24**





Non-invasive therapy to reduce the body burden of aluminium in Alzheimer's disease

Christopher Exley^{a,*}, Olga Korchazhkina^b, Deborah Job^c, Stanislav Strekopytov^a,
Anthony Polwart^d and Peter Crome^{c,e}

^a*Birchall Centre for Inorganic Chemistry and Materials Science, Keele University, Staffordshire, UK*

^b*Institute for Science and Technology in Medicine, Keele University, Staffordshire, UK*

^c*Department of Gerontology, University Hospital of North Staffordshire, Staffordshire, UK*

^d*Life Sciences, Keele University, Staffordshire, UK*

^e*School of Medicine, Keele University, Staffordshire, UK*

The first 'test' (over only 5 days) of an 'aluminium hypothesis of Alzheimer's disease with a silicon-rich mineral water showed that silicon-rich mineral waters could be an effective and non-invasive method to lower the body burden of aluminium.

The Second Test!

Silicon-Rich Mineral Water as a Non-Invasive Test of the 'Aluminum Hypothesis' in Alzheimer's Disease

Samantha Davenward^a, Peter Bentham^b, Jan Wright^b, Peter Crome^c, Deborah Job^c,
Anthony Polwart^d and Christopher Exley^{a,*}

^a*The Birchall Centre, Lennard-Jones Laboratories, Keele University, Stoke-on-Trent, Staffordshire, UK*

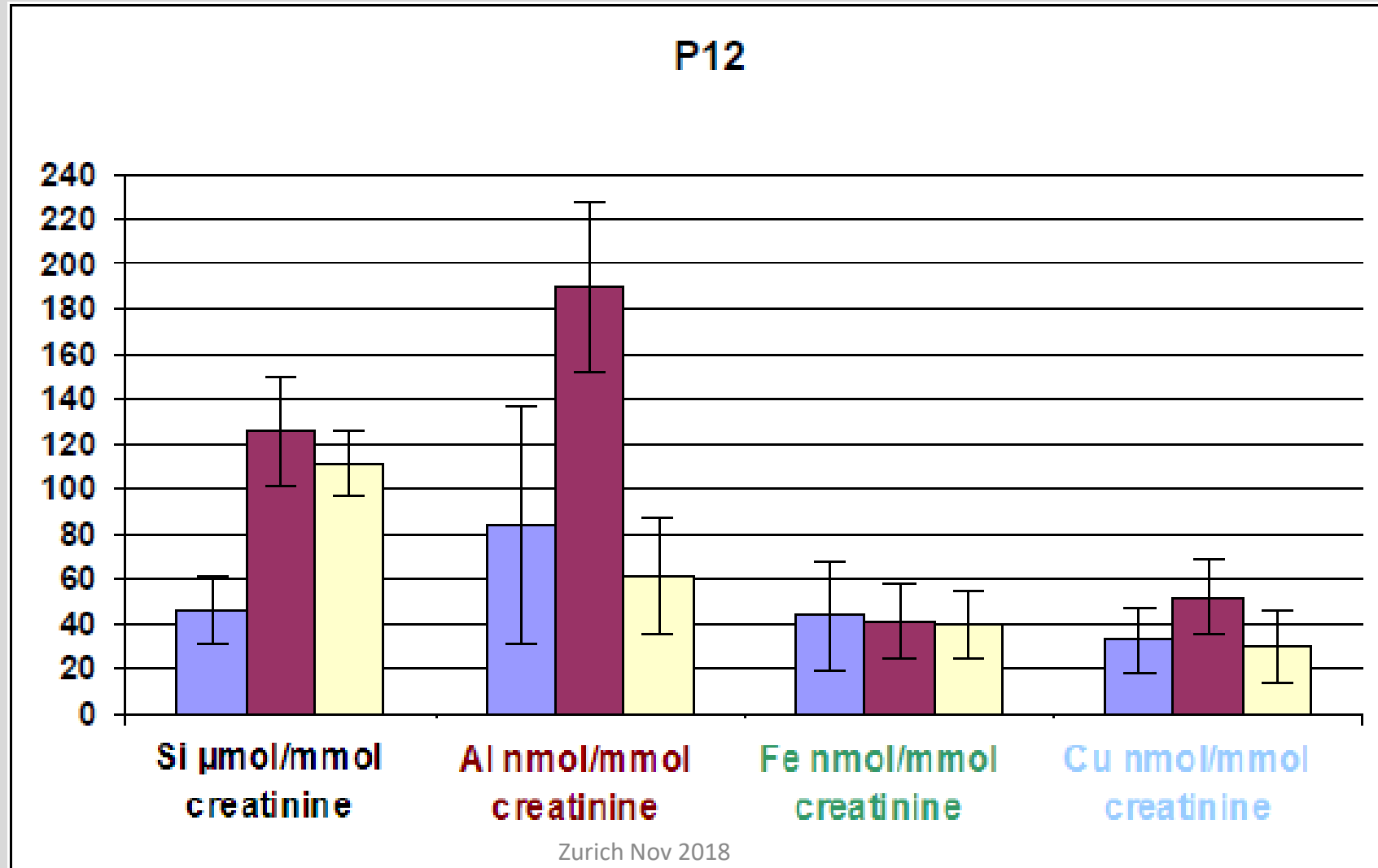
^b*Birmingham and Solihull Mental Health NHS Foundation Trust, The Barberry Centre, Birmingham, UK*

^c*North Staffordshire Combined Healthcare NHS Trust, Harplands Hospital, Stoke-on-Trent, UK*

^d*Life Sciences, Keele University, Stoke-on-Trent, Staffordshire, UK*

We have provided preliminary evidence that over 12 weeks of silicon-rich mineral water therapy the body burden of aluminium fell significantly in individuals with Alzheimer's disease and, concomitantly, cognitive performance showed clinically relevant improvements in at least 3 out of 15 individuals.

Alzheimer's Disease



Healthy Volunteers

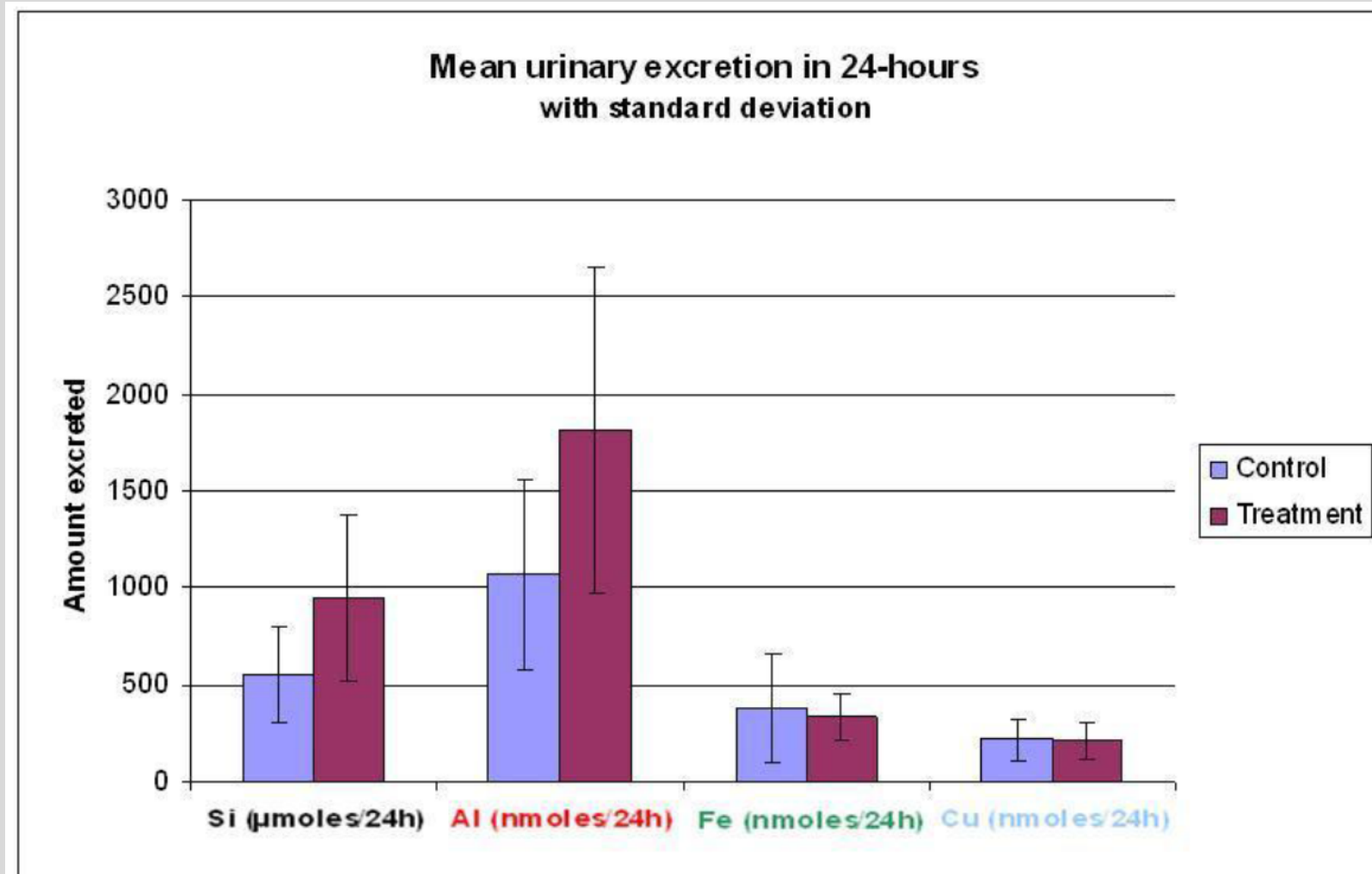


Figure 3.1.4.2: **Mean** amounts and standard deviation bars of excreted Si ($\mu\text{moles}/24\text{h}$), Al, Fe and Cu ($\text{nmoles}/24\text{h}$) in the control and treatment samples.



Contents lists available at ScienceDirect

EBioMedicine

journal homepage: www.ebiomedicine.com



Research Paper

Urinary Excretion of Aluminium and Silicon in Secondary Progressive Multiple Sclerosis



Krista Jones^a, Caroline Linhart^b, Clive Hawkins^c, Christopher Exley^{a,*}

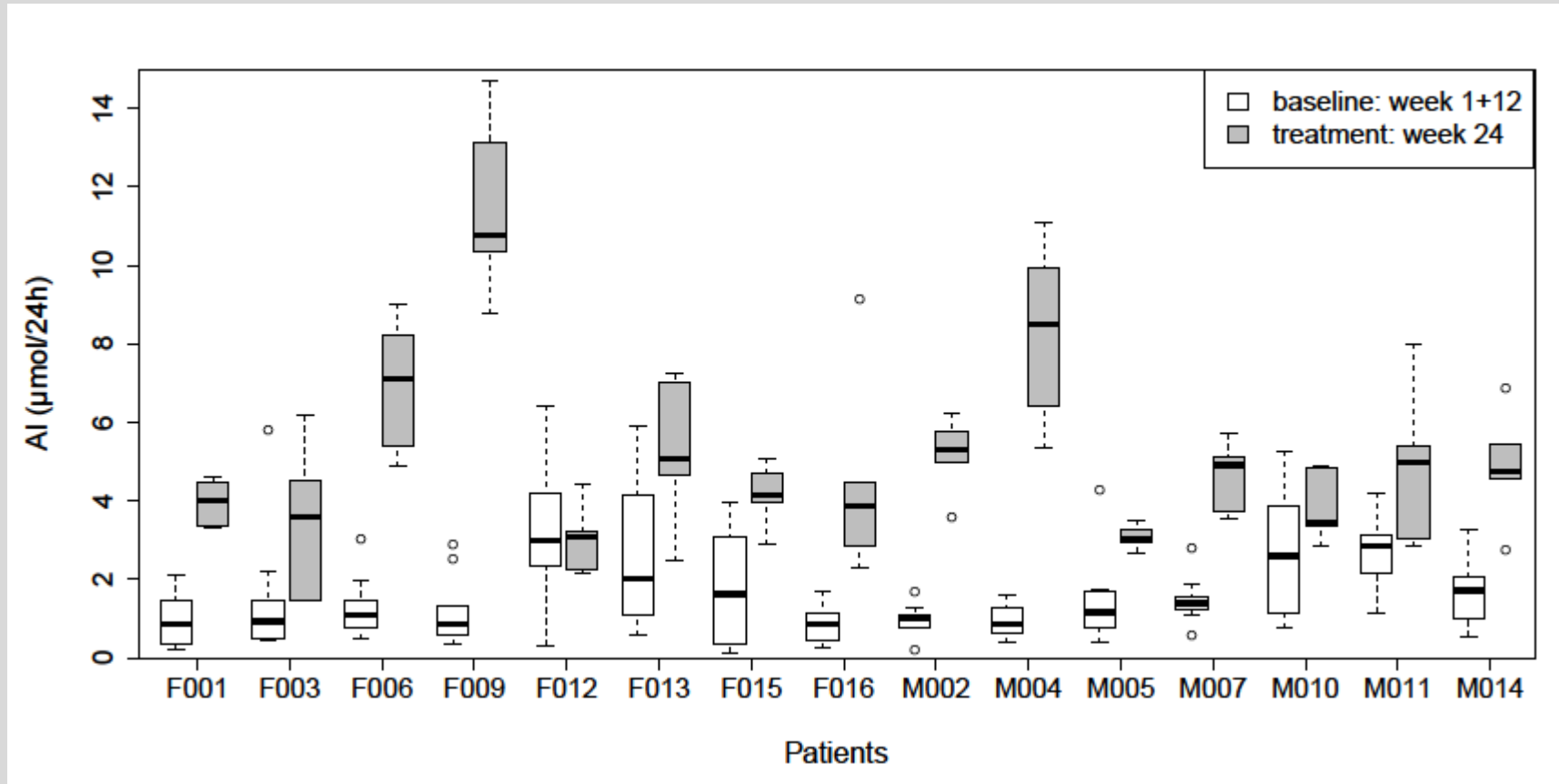
^a *The Birchall Centre, Lennard-Jones Laboratories, Keele University, United Kingdom*

^b *Department of Medical Statistics, Informatics and Health Economics, Medical University of Innsbruck, Austria*

^c *Institute of Science and Technology in Medicine, Keele University, United Kingdom*

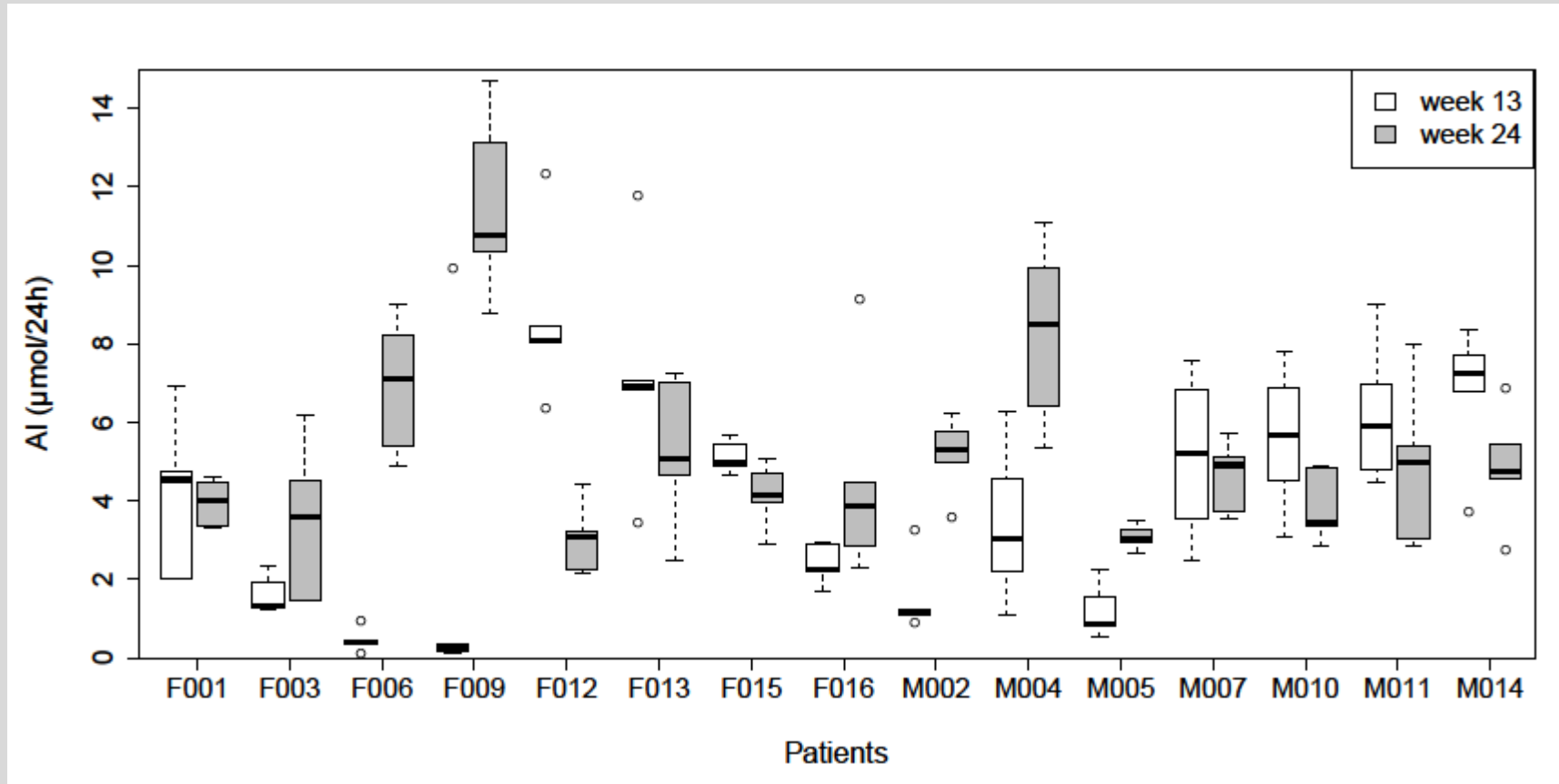
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Multiple Sclerosis



<https://www.sciencedirect.com/science/article/pii/S2352396417304280?via%3Dihub>

Multiple Sclerosis



<https://www.sciencedirect.com/science/article/pii/S2352396417304280?via%3Dihub>

*I think I have the
solution to healthy
ageing...
Have a good day!*



The Birchall Centre, Keele University and
Centro de Investigación Científica de Yucatán



THE THIRTEENTH
KEELE MEETING ON
Aluminium

**Future Challenges in
the Aluminium Age**

23rd
27th
March
2019

Hotel Uxmal Resort Maya
Yucatán, México



Keele
UNIVERSITY



CONACYT



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Zurich Nov 2018