Autism, Environment, Subtypes, and Remissions

Webinar Febuary 25, 2016



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"No nation is any healthier than its children."

Harry Truman (1884-1972) 33rd President of USA



- 54% of Kids Chronic Illness
- Type 2 Diabetes- 12 Million
- Allergies- 7.3 Million
- Asthma- 6.7 million 300%
- ADHD- 5 Million
- Obesity 17% 2-19 yo
- 1 in 6 Neurodeveloptmental disorder
- 1 in 6 public school special ed
- 1 in 7 kids antipsychotic med
- Maternal Mortality 33rd;
 Maternal Health 61st in World
- Infant Mortality 28^{th:} Children's well being 42nd
- Autism 1 in 45

You are Your Child's Hero

Steps You Can Take to Improve Your Autistic Child's Neurological Health and Quality of Life



Norman Schwartz, M.D.

With Nancy Hovan Carpenter

Chapter 1: UNDERSTANDING AUTISM

What is Autism?....

Rare to Epidemic in a few decades.....

What is Different about Today?

The Evolution of Our Understanding of Autism Spectrum D Why is my Child Autistic?.....

Why Doesn't My Doctor know this if it is Science-Based?....

Chapter 2: STEPS TO TAKE: Where Are My Tights and C

Remember Your Child's Strengths and Intelligence.....

Reduce Toxic Exposure

Improve Diet.....

Support Detoxification

Simple Steps to Enhance Drainage for Your Child and fo Biomedical Treatments to Support Detoxification

GI Support and Gut/Digestive Health.....

Support Sleep Healing Modalities.....

Chapter 3: PREVENTING AUTISM SPECTRUM DISORDER Preconception and Prenatal Steps for Mothers.....

Preventing Autism in Babies.....

Chapter 4: SELF CARE: Be Your Own Hero

Parental Stress.....

Stress Management Techniques.....

Find Your Tribe.....

Chapter 5: GLOBAL CRISIS: The Clarion Call.....

Join your voice to others within the autism community Join your voice to others in the environmental movement....

You Come First.....

Suggested Books.....

Chapter 1: UNDERSTANDING AUTISM

"These children are trying so hard to show us how to live in a cleaner world. I believe these kids are here for a reason: to teach us to eat better, clean up the air, and get rid of toxins because they can't survive."

Jenny McCarthy

Autism What is Going On??? Rare to Pandemic Cause, prevention, cure?

- 1943- psych Leo Kanner first report-multiple med problems
- 2015- 72 yrs & > 18,000 peer-reviewed papers later
- Why? Initial focus-social, behavior, communication
- 3 generations of clinicians, siloed thinking
- No clear biological markers or medical criteria
- Science set back 1950-1970's Autism due to BAD PARENTING;
- Bernie Rimland- 1969. evidence of biological causes
- 1980-2016 primacy of genetics
- THINGS ARE CHANGING: Importance Environment, disruption fundamental control symptoms

Multi-system from the start? Kanner 1943 on <u>body</u> symptoms

- Case 1: "Eating has always been a problem" for him. He has never shown a normal appetite."
- Case 2: "...large and ragged tonsils."
- Case 3: diarrhea and fever following smallpox vaccination healthy except for large tonsils and adenoids.
- Case 4: vomited a great deal during his first year... feeding formulas were changed frequently ... tonsils were removed...
- Case 5: nursed very poorly ... quit taking any kind of nourishment at three months... tube-fed five times daily up to one year of age...At camp she slid into avitaminosis and malnutrition but offered almost no verbal complaints."
- Case 7: vomited all food from birth through the third month....
- Case 8: feeding formula caused ...concern. ... colds, bronchitis, streptococcus infection, impetigo...
- Case 9: none of the usual children's diseases." [? Overactive immune system?]
- Case 10: frequent hospitalizations because the feeding problem ... repeated colds and otitis media
- Case 11: was given anterior pituitary and thyroid preparations for 18 months

 Kanner's original paper, discussed in Jepson 2007

Autism: A <u>Behaviorally Defined</u> Syndrome Biology is not part of the definition (and neither is prognosis)

DSM-IV Criteria for Autistic Disorder (299.0)

- 1. Impaired social interaction
- 2. Impaired social communication
- 3. Markedly restricted repertoire of activities and interests

Secondary Features of Autism

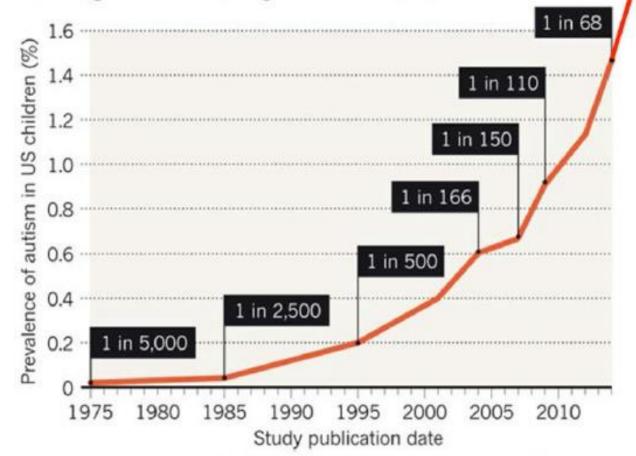
Seizures (~30%+), cognitive deficits, sensorimotor abnormalities, savant skills, inflammation immune impairments, GI distress(50-75%), food allergies (~50+%)

No biological markers exist to identify autism at this time Autism is presumably <u>Heterogeneous</u> biologically But autism is <u>biological</u>

Background: The Frightening Reality

AUTISM DIAGNOSES RISING

Almost 1.5% of US children are now diagnosed with autism, according to data from 11 regions in the United States.



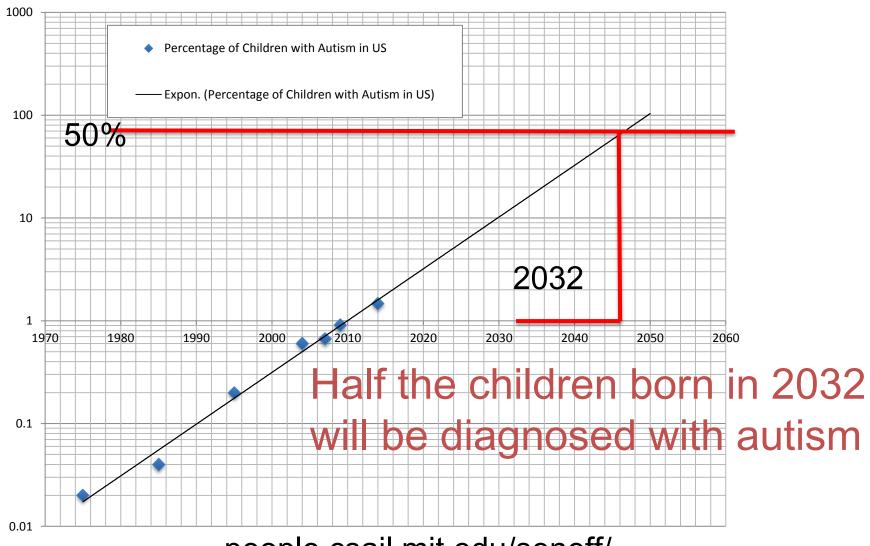
1 in 42 boys¹

2016/1 in 45

- 1 in 189 girls¹
- If one child in a family has an ASD, odds are others will

PURDUE

Percentage of children with Autism in the US



people.csail.mit.edu/seneff/

Environmental Health Perspectives 108, Supplement 3 June 2000

The Environment as an Etiologic Factor in Autism: A New Direction for Research

Eric London^{1,2} and Ruth A. Etzel³

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(ap

¹The National Alliance for Autism Research, Princeton, New Jersey, USA; ²University of Medicine and Dentistry of New Jersey, New Brunswick, New Jersey, USA; 3Division of Epidemiology and Risk Assessment, Food Safety and Inspection Service, Washington, D.C., USA

Autism is one of a group of developmental disorders that have devastating lifelong effects on its victims. Despite the severity of the disease and the fact that it is relatively common (15 in 10,000), there is still little understanding of its etiology. Although believed to be highly genetic, no abnormal

genes have been found. Recent findings in autism and in related disorders point to the possibility that the disease is caused by a gene-environment interaction. Epidemiologic studies indicate that

the number of cases of autism is increasing dramatically each year. It is not clear whether this is

due to a real increase in the disease or whether this is an artifact of ascertainment. A new theory

research on pesticides. We describe the current state of environmental research in autism to illustrate the issues faced by those interested in pursuing the

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the developmental disorders rather than just

etiology of developmental brain disorders. Because this work is in its infancy, we include

Believed to be highly genetic, no abnormal genes have been found. Despite the severity of the disease and the fact that it is relatively common:15 in 10,000 [2016: 222 in 10,000]

The autism spectrum disorders (otherwise known as the pervasive developmental disor-

ders) include autistic disorder as well as pervasive developmental disorder not otherwise specified and Asperger's syndrome. These would bring the total population of people with developmental brain disorders in the United States to over 20 million individuals. Except for a few types of mental retardation

studies. There is a concordance rate of approximately 2-6% in dizygotic twins as opposed to the 66% concordance rate in monozygotic twins. This 2-6% rate is also the rate estimated for family recurrence, which should be

autism spectrum disorders (9,10). This has

been established through twin and family

such as Down's syndrome and Fragile X synthree are all widely believed to be developdrome, the etiology of the developmental brain mental brain disorders. It is quite likely that disorders is unknown. The most common eti-

PERSPECTIVE

nature neuroscience

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meeting

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000 twin

ses?

Time to give up on a single explanation for autism

Francesca Happé, Angelica Ronald & Robert Plomin

We argue that there will be no single (genetic or cognitive) cause for the diverse symptoms defining autism. We present recent evidence of behavioral fractionation of social impairment, communication difficulties and

rigid and re nonoverlap cognitive le symptoms and treatm

Autism, and syndrome, a a triad of bel We argue that there will be no single cause for the diverse symptoms defining autism.

conversation going or likes to do things over and over again in the same

communication and restricted and repetitive interests and activities1. These core diagnostic features, which reflect Kanner's first reports of

pairs assessed between the ages of 7 and 9, we have found modest-tolow correlations between autistic-like behavioral traits in the three

Recent work suggests that autistic-like traits can be measured in

the general population⁷⁻⁹. We can ask, for example, whether a child

joins in playing games with other children easily, can keep a two-way

NUMBER 10 | OCTOBER 2006 NATURE NEUROSCIENCE

SCIENTIFIC AMERICAN

SEARCH

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New Study: Autism Linked to Environment

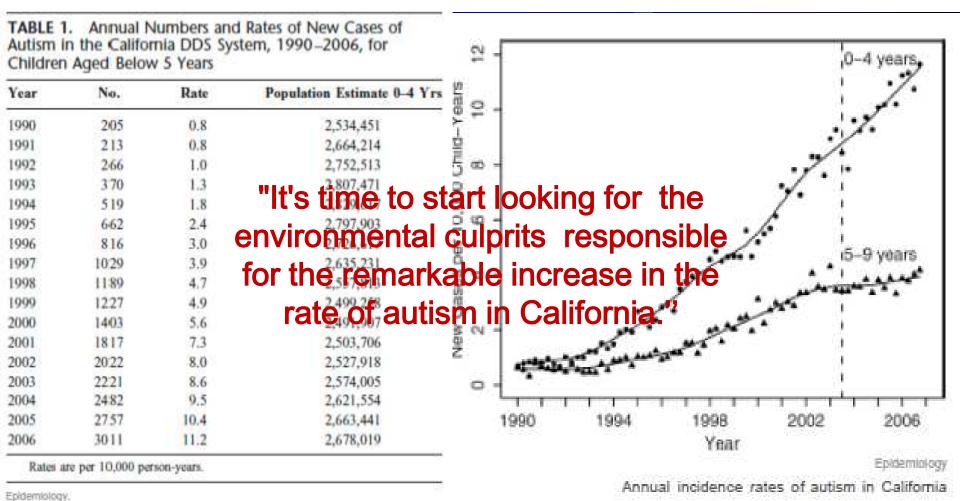
Research links soaring incidence of the mysterious neurological disorder to fetal and infant exposure to pesticides, viruses, household chemicals

By Marla Cone | January 9, 2009 | ₹76

Epidemiology • Volume 20, Number 1, January 2009

The Rise in Autism and the Role of Age at Diagnosis

Irva Hertz-Picciotto*, and Lora Delwiche*



nature

3 NOVEMBER 2011 | VOL 479 | NATURE



THISWEEK

EDITORIALS

we fail to find the Higgs boson p.8 effect of contraceptives and HIV p.7 John Mc(dies p

The mind's tangled web

Efforts to elucidate how genes and the environment shape the development of autism, although making progress, still fall far short of their goal.

a mong psychiatric disorders, autism has received particularly strong support from government and philanthropic funders in recent years. And that investment has paid scientific dividends, above all the uncovering of genetic clues to underlying mechanisms for the disorder. But, as discussed in this special issue and in a web collection of content published this week in other Nature journals (see nature.com/autism), those developments, although pointing a way forward, have themselves revealed just how small a distance we have travelled towards a fuller mechanistic understanding.

"The growth in the prevalence of autism can be explained only partly by changes in diagnostic practice."

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Autism counts

"Idea of whether the prevalence is increasing is so contentious for autism, but not for asthma, type 1 diabetes, food allergies. It is clear there is a real increase in autism, researchers need more funding and encouragement to look at environmental causes."

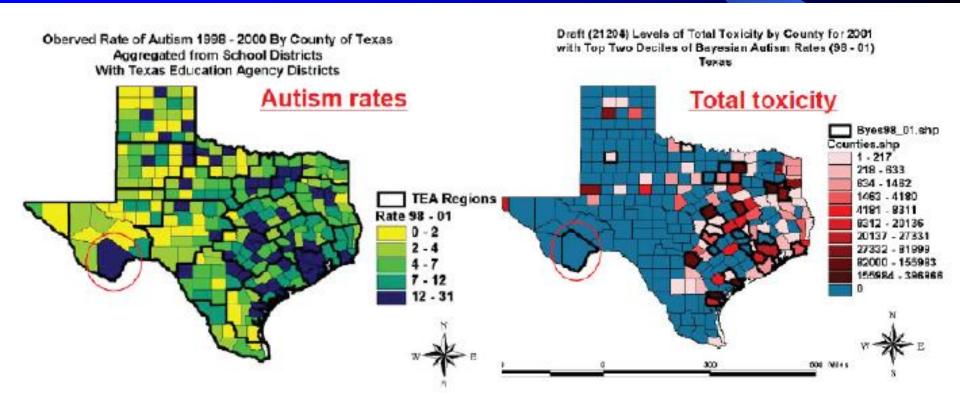
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physician and psychiatrist at Johns Hopk Thomas Insel, Director NIMH
University in Baltimore, Maryland, predictor and psychiatrist at Johns Hopk Thomas Insel, Director NIMH

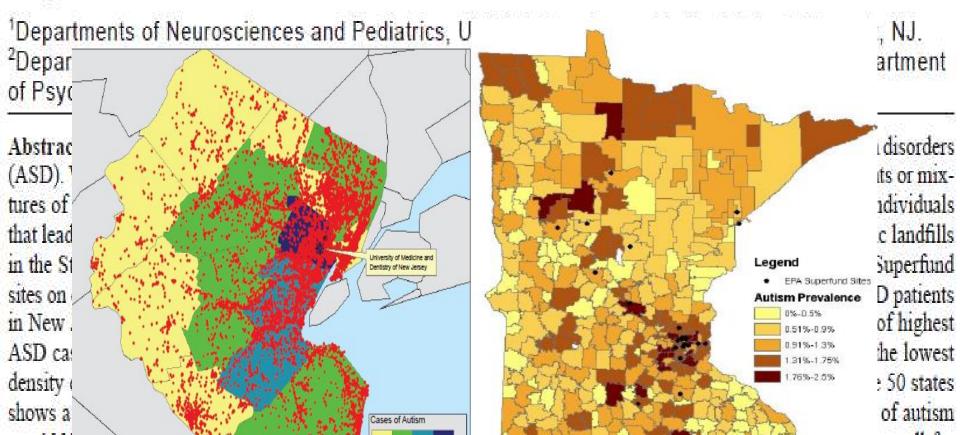
Association between autism rates, environmental mercury, other toxins in Texas Palmer, et al., Health and Place 2006 Jun;12(2):203-9

For each 1000 lb of environmentally released mercury, there was a 43% increase in the rate of special education services and a 61% increase in the rate of autism.



Autism Spectrum Disorders and Identified Toxic Land Fills: Co-Occurrence Across States

Xue Ming¹, Michael Brimacombe², Joanne H. Malek³, Nisha Jani² and George C. Wagner³



Maternal Residence Near Agricultural Pesticide Applications and Autism Spectrum Disorders among Children in the California Central Valley





https://imfar.confex.com/imfar/2010/webprogram/Paper6362.html

International Society for Autism Research



Start | Browse by Day or Format | Author Index

Breastfeeding and Autism

Saturday, May 22, 2010

Franklin Hall B Level 4 (Philadelphia Marriott Downtown)

10:00 AM

P. G. Williams , Pediatrics, University of Louisville, Louisville, KY L. L. Sears , Pediatrics, University of Louisville, Louisville, KY

Background:

Breas prote Conclu

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Exposure to PCB's in breast milk, environmental trigger, disrupting brain development - ASD in genetically predisposed.

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months was similar to the general rate for Kentucky, but the rate of breastfeeding in children with autism was nearly 3 times higher. The Kenet study pointed to possible concerns associated with substances in breast milk. While PCB's were banned in the late 1970's, these substances are very slowly degraded and maintain a si of plasma PCB concentrations compared to those who were not breastfed. The authors hypothesized that exposure to PCB's in breast milk might act as an environmental trigger, disrupting brain development and resulting in autism for those already genetically predisposed. While the known breastfeeding benefits still.

Curr Opin Pediatr. 2010 Apr;22(2):219-25. doi: 10.1097/MOP.0b013e328338eb9a.

What causes autism? Exploring the environmental contribution.

Landrigan PJ.

Author information

Abstract

PURPOSE OF REVIEW: Autism is a biologically based disorder of brain development. Genetic factors--mutations, deletions, and copy number variants--are clearly implicated in causation of autism. However, they account for only a small fraction of cases, and do not easily explain key clinical and epidemiological features. This suggests that early environmental exposures also contribute. This review explores this hypothesis.

RECENT FINDINGS: Indirect evidence for an environmental contribution to autism comes from

Genetic factors account for only a small fraction of casesearly environmental exposures contributes

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valproic acid; maternal rubella infection; and the organophosphate insecticide, chlorpyrifos. There is no credible evidence that vaccines cause autism.

Exposures in early pregnancy - thalidomide, misoprostol, and valproic acid; maternal rubella infection; and the organophosphate insecticide, chlorpyrifos.

oday adult have on

suspect chemicals, which combines expanded toxicological screening, neurobiological research and prospective epidemiological studies.



NIH Public Access

Author Manuscript

Epidemiology. Author manuscript; available in PMC 2012 July 1.

Published in final edited form as:

Epidemiology. 2011 July; 22(4): 476–485. doi:10.1097/EDE.0b013e31821d0e30.

Prenatal vitamins, one-carbon metabolism gene variants, and risk for autism

Periconceptional use of prenatal vitamins

may reduce the risk of having children with

autism, especially for genetically susceptible

mothers and children

³ Department of Pediatrics, University of California Davis School of Medicine, Davis, CA

⁴ Department of Preventive Medicine, Institute for Genetic Medicine, University of Southern California Keck School of Medicine, Los Angeles, CA

⁵ Department of Biochemistry and Molecular Medicine, University of California Davis School of



Short Communication Open Acces

Vitamin/Mineral Supplements for Children and Adults with Autism

James B Adams*

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Director, Autism/Asperger's Research Program, President's Professor, Arizona State University, PO Box: 876106, Temple, AZ, USA

Abstract

Vitamins and minerals are the most widely used medical treatment for autism. Many research studies have demonstrated that children and adults with autism often have nutritional and metabolic problems, including problems with methylation, glutathione, oxidative stress, sulfation, lithium, and more. This review summarizes the results of several vitamin/mineral treatment studies conducted by our group, which demonstrate that vitamin/mineral supplements are highly effective in improving many nutritional and metabolic problems, and result in significant improvements in symptoms based on a large, randomized, double-blind, placebo-controlled study. We recommend that all children and adults with autism consider a 2-3 month trial of a vitamin/mineral supplement designed for individuals with autism that is similar to the one used in our studies. By starting at a low dose, and gradually increasing it, there is minimal risk of adverse effects, and many children and adults are likely to benefit, sometimes substantially.

Vitamin/mineral supplements highly effective in improving nutritional and metabolic problems, in a large, randomized, double-blind, placebo-controlled study

widely used treatments for autism.

The definition of a "vitamin" or "essential mineral" is that the

supplementation - with vitamin methyl-B12, found acid (a special type of folic acid), and (in one of the studies) trimethylglycine resulted in significant improvements in oxidative stress, methylation and plutathions [3,4]. A 30 resolv double blind placebe controlled

erals and/o nat childre methylatio smpared to nutritions What's really behind the increase in autism rates. May 7, 2012

THE COLLABORATIVE ON HEALTH AND THE ENVIRONMENT



Face the reality- our children are sick from preventable causes NOT better diagnosis
1980- 200 billion pounds of chemicals/year
Now- 27 trillion pounds of chemicals/yr
1960- 400 million pounds of pesticides/yr
Now 4.5 billion pounds pesticides/yr
National Academy of Sciences 50% of pregnanciesfetal death or less than healthy child

CHEMICAL FALLOUT | UPDATE

EPA a failure on chemicals, audit finds

Assessment of toxic risks inadequate, says new chief

FDA relied heavily on BPA lobby

Regulators actively reached out to industry, e-mails show

By Susanne Rust and Meg Kissinger of the Journal Sentinel

Posted: May. 16, 2009



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Mobile/PD

EPA To Drop 'E,' 'P' From Name

"We're not really environmental anymore, and we certainly aren't protecting anything"



THE MICROBIOME IN AUTISM SPECTRUM DISORDER

A model for the induction of autism in the ecosystem of the human body: the anatomy of a modern pandemic?

Staci D. Bilbo1, Cynthia D. Nevison2 and William Parker3*

¹Department of Psychology & Neuroscience, Systems & Integrative Neuroscience Group, Duke University, Durham, NC, USA; ²Institute for Arctic and Alpine Research, University of Colorado Boulder, Boulder, CO, USA; ³Department of Surgery, Duke University Medical Center, Durham, NC, USA

Background: The t

Two lines of reasoning are examined which suggest that autism is indeed a pandemic of modern culture

uestion regarding

culture.

ence is

the nature of autis pandemic of modern culture with environmental factors at the roots. Others are convinced that the disease is not pandemic in nature, but rather that it has been with humanity for millennia, with its biological and neurological underpinnings just now being understood Unequivocal way to resolve the debate-perform an objective.

experiment: monitor the prevalence of autism after normalizing immune function in a Western population

reviewed reviewed culture. In light of this reasoning, current epidemiological evidence regarding the incidence of autism,

VOLUME 123 | NUMBER 3 | March 2015 · Environmental Health Perspectives

Air Pollution and ASDs

Homing In on an Environmental Risk Factor

Although researchers have begun making significant inroads into understanding the genetic and biological basis for autism spectrum disorders (ASDs) and other neurodevelopmental disorders, it's been estimated that environmental factors could account for just over half the risk of developing ASDs.\(^1\) In this issue of EHP, researchers have used data from 1,767 women in the Nurses' Health Study (NHS) II to study exposure to particulate matter (PM) as one potential environmental risk factor for ASDs.\(^2\)

In one of the first reports of a relationship between air pollution and ASDs, investigators in California found associations between estimated exposures to airborne heavy metals and other pollutants and risk of ASDs.³ Later studies using the same exposure

models as the California study reported links between multiple hazardous air pollutants and ASDs in North Carolina and West

50% higher odds ASD in women with PM2.5 exposure during pregnancy In this study, higher exposures to PM2s air pollution during the third trimester of pregnancy were associated with increased risk of having a child with an autism spectrum disorder. KldStock/Blend images/Corbis

> exposed to concentrated ambient ultrafine PM showed ventriculomegaly, altered neurochemistry, and activation of glial cells in the

What Do All These Have In Common

- Vinyl flooring
- Pesticides flea/tick sprays
- Living near power plants, landfills, incinerators

ssriuse in pregnancy. rpussociated rewirth Autism

- Antibiotics
- Increased rain fall
- Northern latitude
- Gut Ecology
- Living near freeway
- Increased cable usage
- Obesity/DM mom

- Shorter birth interval
- Breast feeding
- June birth,
- First born
- Infection, meds in pregnancy
- Fetal distress, low apgar
- In vitro fertilization
- Unplanned C-section
- Vaccinations
- Older parents
- Higher Socioeconomic

THEMATIC CLUSTER: FOCUS ON AUTISM SPECTRUM DISORDERS

Microbial Ecology in Health & Disease 2012, 23:

Should autism be considered a canary bird telling that *Homo sapiens* may be on its way to extinction?

Olav Albert Christophersen, Cand. mag.*

Pensioned Norwegian state stipendiate

There has been a dramatic enhancement of the reported incidence of autism in different parts of the world over the last 30 years. This can apparently not be explained only as a result of improved diagnosis and reporting, but may also reflect a real change. The causes of this change are unknown, but if we shall follow T.C. Chamberlin's principle of multiple working hypotheses, we need to take into consideration the possibility that it partly may reflect an enhancement of the average frequency of responsible alleles in large populations. If this hypothesis is correct, it means that the average germline mutation rate must now be much higher in the populations concerned, compared with the natural mutation rate in hominid ancestors before the agricultural and industrial revolutions. This is compatible with the high prevalence of impaired human semen quality in several countries and also with what is known about high levels of total exposure to several different unnatural chemical mutagens, plus some natural ones at unnaturally high levels. Moreover, dietary deficiency

Chapter 2: STEPS TO TAKE: Where Are My Tights and Cape?

"...That a disease is complex or multifactorial does not imply that simple solutions cannot be found or that clinical advances following insight cannot be swift."

JA Rees (Science, 2002;296: 698-701)

Action Steps

- Remember your child's strengths and intelligence
- Reduce toxic exposures
- Improve diet
- Support detoxification
- Heal gut/digestion
- Support sleep
- Employ healing modalities

Steps to Take

- Toxins in our bodies can be cumulative.
 Every single thing you do counts.
- Eat Clean Food
- Avoid processed foods
- Avoid artificial sweeteners, coloring, additives, and flavorings
- Drink Pure Water
- Breathe Clean Air
- Use Clean Personal Care Products

Steps to Take

- Reduce Hormone/Endocrine Disruptors
- Clean Clothing/Laundry Products
- Eliminate Plastics
- Eliminate Toxic Metals
- Avoid Electromagnetic Radiation
- Avoid Excitotoxins
- Eat lots of Healthy Fats

Steps to Take

- Biomedical Treatments to Support Detoxification
- GI Support and Gut/Digestive Health
- Support Sleep
- Healing Modalities
- B12 Injections

JOURNAL OF CHILD AND ADOLESCENT PSYCHOPHARMACOLOGY Volume X, Number x, 2016

Mary Ann Liebert, Inc.

Original Article

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Pp. 1-10

DOI: 10.1089/cap.2015.0159

Randomized, Placebo-Controlled Trial of Methyl B12 for Children with Autism

Robert L. Hendren, DO, S. Jill James, PhD, Felicia Widjaja, MPH, Brittany Lawton, BS, Abram Rosenblatt, PhD, and Stephen Bent, MD

Abstract

Objective: Children with autism spectrum disorder (ASD) have been reported to have reduced ability to methylate DNA and elevated markers of oxidative stress. We sought to determine if methyl B12, a key metabolic cofactor for cellular methylation reactions and antioxidant defense, could improve symptoms of ASD.

Symptom improvement correlated with \tau methionine and \tau S-adenosylmethionine (SAM)/SAH

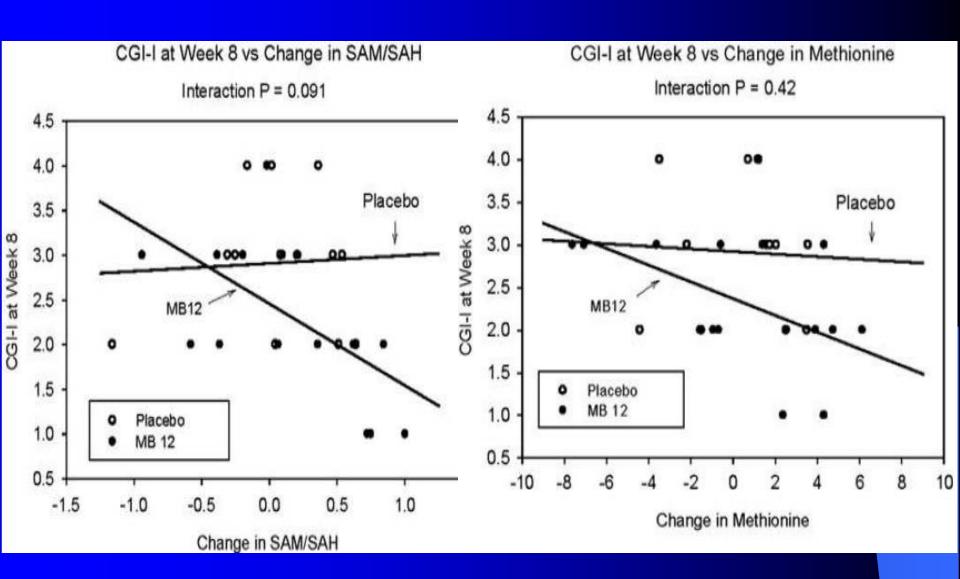
measures of meanonine meanytation and antioxidant gratautione metabolism were assessed at baseline and o weeks.

Results: A total of 50 children (mean age 5.3 years, 79% male) completed the study. The primary outcome measure - the

Improved cellular methylation capacity

cating an improvement in cellular methylation capacity. No improvements were observed in the parent-rated ABC or SRS.

Conclusions: Methyl B12 treatment improved clinician-rated symptoms of ASD that were correlated with improvements in



Chapter 4: SELF CARE: Be Your Own Hero

• "Do not let either the medical authorities or the politicians mislead you. Find out what the facts are, and make your own decisions about how to live a happy life and how to work for a better world."

Linus Pauling, PhD, 1901-1994 Two- time Nobel Laureate

Self Care

- You Come First
- Parental Stress- helpful articles:
 - Family Autism Self Care Strategies
 - Article Self Care is not Selfish
 - Article Caring for the Caregiver
 - 15 Tips for Your Family
 - Sibling Resources
 - 10 Things Parents of ASD Would Love To Hear

Self Care

- Stress Management Techniques
- Find Your Tribe
 - Documenting Hope/Epidemic Answers
 - Generation Rescue
 - Talk About Curing Autism
 - Safe Minds
 - National Autism Association
 - Fearless Parent
 - Thinking Moms Revolution
 - The Canary Party
 - BioMed Heals
 - Moms in Charge
 - Parents Helping Parents: The Autism Education
 Network

Almost Autism: Recovering Children from Sensory
Processing Disorder
by Maria Rickert Hong

A Compromised Generation: The Epidemic of Chronic Illness in America's Children by Beth Lambert and Victoria Kobliner

Healing Without Hurting by Jennifer Kozek, LPC, NCC

Kids Beyond Limits by Anat Baniel

The Hidden Connection by Kathleen DiChiara, FDN

Nourishing Hope for Autism, Autism Diet & Nutrition Book

Healing the New Childhood Epidemics Kenneth Bock, M.D. Cameron Stauth

Autism Revolution by Martha Herbert, M.D. with Karen Weintraub

Nutritional Supplement Use for Autistic Spectrum Disorder

by Jon Pangborn

Children with Starving Brains by Jaquelyn McCandless, M.D.

Autism: Effective Biomedical Treatments by Jon Pangborn, PhD and Sidney MacDonald Baker, M.D. The Dirt Cure by Maya Shetreat-Klein, MD Changing the Course of Autism by Bryan Jepson Detoxification & Healing: the Key to Optimal Health by Sid Baker, MD

The Autism Book by Robert W Sears, MD, FAAP Infantile Autism by Bernard Rimland, MD

Treating Autism: Parent Stories of Hope & Success Edited by Stephen Edelson, PhD & Bernard Rimland, PhD

The Road to Immunity by Kenneth Bock, MD & Nellie Sabin

Healing Our Autistic Children by Julie A Buckley, MD

Born on a Blue Day by Daniel Tammet

Understanding Autism for Dummies by Steven M Shore,

Chapter 5: GLOBAL CRISIS: The Clarion Call

"A new consciousness is developing that sees the earth as a single organism, and recognizes that an organism at war with itself is doomed."

Carl Sagan, 1934-1996

"Worldwide grassroots movement of hope and humanity. Immune System of our planet"



"I observe the physician with the same diligence as the disease." John Donne, 1572-1631 The specific disease doctrine is the grand refuge of weak, uncultured, unstable minds such as now rule in the medical profession.

Florence Nightingale 1820-1910



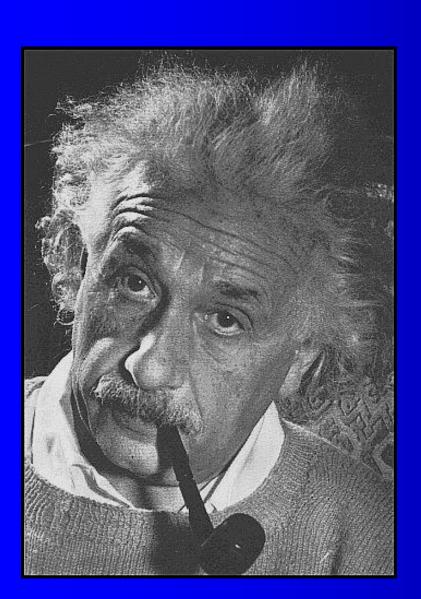


»YOU CAN KNOW the name of a bird in all the languages of the world, but when you're finished, you'll know absolutely nothing whatever about the bird. So let's look at the bird and see what it's doing — that's what counts. I learned very early the difference between knowing the name of something and knowing something.«

RICHARD P. FEYNMAN

Ranked as one of the ten greatest physicists of all time

Opinions about obviousness are to a certain extent a function of time.



Albert Einstein

- The earth is flat.
- Children should be seen and not heard
- Airplanes are impossible
- Women are not logical enough to vote
- Autism caused by bad parenting
- Autism is genetic.

ASD Complex Whole Body

- Oxidative Stress
- Inflammation
- Mitochondrial Disorders
- Seizure Disorder
- CalciumDysregulation
- Gastrointestinal
- Fatty AcidDeficiencies
- Toxic Metals Burden

- Decrease Brain Blood Flow
- Brainstem Abn
- Glutathione Decrease
- Melatonin Decrease
- Oxytocin/Serotonin Abnormalities
- Low and High Cholesterol
- Nutrient, Mineral Deficiencies
- Multiple Immune Abnormalities

Subtypes

- 1. Redox balance- cell vitality, ability to prevent oxidativem stress and free radical damage
- 2. Digestive competence- break down, assimilate, and eliminate, entire digestive system
- 3. Anabolic/catabolic balance- repair competences or their deficit, including unresolved inflammation.
- 4. **Detoxification-** ability to safely convert toxins into more easily excreted and less harmful
- 5. Somatosenory- information processing, limited reserves can lead to system overload and dysfunction
- 6. Neuroimmunohormonal integration- immune neurotransmitter, and hormonal systems function as a single, integrated control system.

Autism: Thinking the unthinkable

To cling to a purely genetic explanation of autism is a desperate attempt to maintain the illusion that one lives in a comfortable and rational world where new chemicals and technologies always mean progress, experts are always objective and thorough, corporations are honest, and authorities can be trusted. That human actions, rather than genetics, might be responsible for compromising the health of a significant proportion of a whole generation is so painful as to be, for many unthinkable.

Martha Herbert MD, PhD Pediatric Neurologist. Harvard Medical School
48

Autisms are Treatable

Observed Improvements:

- Cognitive deficits
- Sensorimotor abnormalities
- Disordered sleep
- Immune impairments
- -GI distress
- Food allergies
- Systemic metabolic disturbances
- Seizures

My Approach to Treatment

- Currently most behavioral/developmental disorders are MEDICAL not MENTAL Preventable &Treatable
- Comprehensive, functional, integrative approach
- Rx based on predictive, sensitive, specific lab testing ; clinical observations for individualized therapy
- Understand biochemical imbalances to prioritize low risk, high gain treatments.
- Engage family, "Doctor Mom & Dad" as colleagues to determine what works best
- Assess and treat environmental toxicity
- Not a single cause,-complex, need to think deeply about causes and consequences in neurodevelopment.

The Autism Intensive















Stephanie Seneff, PhD Jeffrey Bland, PhD









Paul Thomas, MD



Anat Baniel



Nancy H. O'Hara, MD Julie Matthews, CNC

Erica Peirson, ND

Elizabeth Mumper, MI













Jim Adams, PhD





Russ Jaffe, MD, PhD



Donna Gates





Stu Freedenfeld, MD

Derrick MacFabe, MD

Jolene Brighten, ND

Geri Brewster RD, MPH Dan Rossignol, MD

Tom Malterre, MS

Kim Stagliano

Those who have the privilege to know have the duty to act." Albert Einstein

Together, we know enough...to mitigate, remediate, and prevent the epidemics of autism, chronic illness and lack of wellness. If not now, when?

Thank you!!!

Norm Schwartz MD 10602 N Pt Washington Rd Suite 101 Mequon WI 53092 drnormmd@aol.com 262 240-0133

Closely Spaced Pregnancies Are Associated With Increased Odds of Autism in California Sibling Births

Pediatrics 2011;127:246-253



WHAT'S KNOWN ON THIS SUBJECT: Autism has been associated with pregnancy and birth complications that may indicate a suboptimal prenatal environment. Although the interpregnancy interval (IPI) may affect the prenatal environment, the association between the IPI and risk for autism is not known.



what this study adds: Using full-sibling pairs from a large population, the authors examined the association between autism and IPIs. Second-born children conceived after an IPI of <12 months had more than threefold increased odds of autism relative to those with IPIs of ≥36 months.

AUTHORS: Keely Cheslack-Postava, PhD, MSPH, Kayuet Liu, DPhil, and Peter S. Bearman, PhD^b

Robert Wood Johnson Foundation Health and Society Scholars, Columbia University, New York, New York, and Paul F. Lazarsfeld Center for the Social Sciences, Columbia University, New York, New York

KEY WORDS

autism, IPI, birth intervals, siblings, California

ARREST/IA TIONS

IPI-interpregnancy interval

DDS-Department of Developmental Services

OR-odds ratio

abs'

Children born after shorter intervals between pregnancies increased risk of developing ASD; the highest risk was associated with pregnancies spaced < 1year apart.

METHODS. Pairs of first- and second-born singleton run storings were identified from all California births that occurred from 1992 to 2002 using birth records, and autism diagnoses were identified by using linked records of the California Department of Developmental Services.

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PEDIATRICS (ISSN Numbers: Print, 0031-4005; Online, 1098-4275).

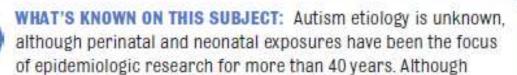
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PEDIATRICS®

OFFICIAL JOURNAL OF THE AMERICAN ACADEMY OF PEDIATRICS

Perinatal and Neonatal Risk Factors for Autism: A Comprehensive Meta-analysis Pediatrics 2011;128:344–355



AUTHORS: Hannah Gardener, ScD,^a Donna Spiegelmar ScD,^{a,b} and Stephen L. Buka, ScD^c

^aDepartment of Epidemiology and ^aDepartment of Biostatisti

Meta analysis of 60 factor s,17 significant: abnormal presentation, umbilical-cord compl, fetal distress, birth trauma, multiple birth, maternal hemorrhage, summer birth, low birth weight, small for gestational age, congenital malformation, low 5-minute Apgar score, feeding difficulties, meconium aspiration, anemia, ABO or Rh incompatibility, and hyperbilirubinemia.

BACKGROUND: The etiology of autism is unknown, although perinatal and neonatal exposures have been the focus of epidemiologic research

Spiegelman contributed to the study concept and design, analysis and interpretation of data, critical revision of the manuscript for important intellectual content, statistical J Am Coll Nutr. 2011 Oct;30(5):348-53.

Autism rates associated with nutrition and the WIC program.

Shamberger RJ.

Author information

Abstract

OBJECTIVES: Autism rates in the United States are increasing at a rate of 15% per year. Autistic children are diagnosed by age 3 when they have problems communicating and interacting socially. This study uses nutritional epidemiology and an ecologic study design to link the possible cause of autism to nutrition by creating autism rates for the 50 states of America and comparing them with published measures of infant nutrition such as duration of exclusive breast-feeding and participation in the Women, Infants, and Children (WIC) program. The percentage of infants with measles, mumps, and rubella (MMR) inoculations was also compared with the autism rates. Study

DESIGN: Autism rates for each state were established. The percentage of infants who participate in

the WIC pro New Jersey design with

The states with the highest WIC participation have significantly lower ASD

0.02). A similar pattern was observed in 21 New Jersey counties (p < 0.02) and 30 Oregon counties (p < 0.05). In contrast, there was a direct correlation with the increasing percentage of women exclusively breast-feeding from 2000-2004 (p < 0.001). Infants who were solely breast-fed had diets that contained less thiamine, riboflavin, and vitamin D than the minimal daily requirements (MDR). There was no correlation of MMR inoculations with the autism rate.

CONCLUSION: The mothers who are exclusively breast-feeding should also continue their prenatal vitamins or their equivalent and make better dietary choices. These results suggest that autism may be nutritionally related to a possible deficiency of riboflavin or the cognitive vitamins such as thiamine or vitamin D. However, due to an ecologic study design there is a potential for fallacy because individuals were not examined. The results suggest the need for a robust observational

ONLINE FIRST

Arch Gen Psychiatry. 2011;68(11):1104-1112.

Antidepressant Use During Pregnancy and Childhood Autism Spectrum Disorders

Lisa A. Croen, PhD; Judith K. Grether, PhD; Cathleen K. Yoshida, MS; Roxana Odouli, MSPH; Victoria Hendrick, MD

Context: The prevalence of autism spectrum disorders (ASDs) has increased over recent years. Use of antidepressant medications during pregnancy also shows a secular increase in recent decades, prompting concerns that prenatal exposure may contribute to increased risk of ASD.

Objective: To systematically evaluate whether prenatal exposure to antidepressant medications is associated with increased risk of ASD. **Results:** Prenatal exposure to antidepressant medications was reported for 20 case children (6.7%) and 50 control children (3.3%). In adjusted logistic regression models, we found a 2-fold increased risk of ASD associated with treatment with selective serotonin reuptake inhibitors by the mother during the year before delivery (adjusted odds ratio, 2.2 [95% confidence interval, 1.2-4.3]), with the strongest effect associated with treatment during the first trimester (adjusted odds ratio, 3.8

SSRI 2-fold increased risk ASD during the year before delivery strongest during the first trimester

cal covariates.

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Setting: The Kaiser Permanente Medical Care Program in Northern California.

Participants: A total of 298 case children with ASD (and their mothers) and 1507 randomly selected control children (and their mothers) drawn from the membership of the Kaiser Permanente Medical Care Program in North-

Conclusion: Although the number of children exposed prenatally to selective serotonin reuptake inhibitors in this population was low, results suggest that exposure, especially during the first trimester, may modestly increase the risk of ASD. The potential risk associated with exposure must be balanced with the risk to the mother or fetus of untreated mental health disorders. Further studies are needed to replicate and extend these findings.

Anal. Com Donald store 2011-60(11)-1104-1112

ANN NEUROL 2013;

Association of Gestational Maternal Hypothyroxinemia and Increased Autism Risk

Gustavo C. Román, MD,^{1,2} Akhgar Ghassabian, MD, PhD,^{3,4}
Jacoba J. Bongers-Schokking, MD, PhD,^{5,6} Vincent W. V. Jaddoe, MD, PhD,^{3,5,7}
Albert Hofman, MD, PhD,^{3,5} Yolanda B. de Rijke, PhD,^{8,9}

Association between severe, early gestation maternal hypothyroxinemia and autistic symptoms in offspring

ı lesions resemnal weeks 6–18)

antibodies) were assessed in 5,100 women. We defined severe maternal hypothyroxinemia as fT_4 < 5th percentile with normal TSH. Six years later, parents reported behavioral and emotional symptoms in 4,039 children (79%) using the Pervasive Developmental Problems (PDP) subscale of the Child Behavior Checklist and/or the Social Responsive-

Objective: bling those and autistic

Evidence that Increased Acetaminophen use in Genetically Vulnerable Children Appears to be a Major Cause of the Epidemics of Autism, Attention Deficit with Hyperactivity, and Asthma

William Shaw, PhDa

©2013, William Shaw, PhD Journal Compilation ©2013, AARM DOI 10.14200/jrm.2013.2.0101

Journal of Restorative Medicine 2013; 2:

ABSTRACT

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It appears that the marked dedcit with hyperactivity by the marked increase in metabolically susceptible

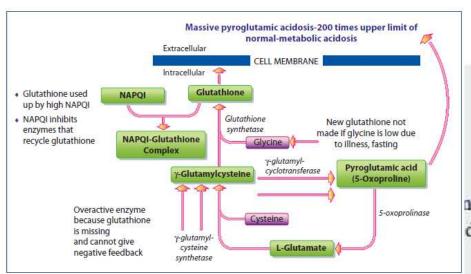


Figure 3b: Metabolism of GSH after exposure to high doses of acetaminophen.

women. Toxicity of acetaminophen may cause autism by overloading the

Acetaminophen may cause autism by overloading defective sulfation leading to overproduction of the toxic metabolite N-acetylp- benzoquinone imine (NAPQI)

| Harmful effects of acetaminophen or its metabolites | References |
|--|------------|
| Neurotoxic effects on rat brain neurons both in vitro and in vivo | 13, 15 |
| Maternal use during pregnancy is associated with teratogenic defects in testicular function and gastrointestinal tract | 30, 31 |
| Very toxic metabolites formed from or induced by acetaminophen (NAPQI and peroxynitrite) that can cause oxidative damage to proteins, nucleic acids, amino acids, and lipids, causes increased mitochondrial and cellular damage and death | 15, 32–35 |
| Severe immune abnormalities at doses that do not damage the liver, including depression of speciac antibody production and reduction of speciac B-cells | 36 |
| Depresses the immune response to vaccination in humans and animals | 37, 38 |
| Depletion of glutathione by acetaminophen causes severe metabolic acidosis | 39, 40 |
| The leading cause of liver failure in the United States | 14, 41,42 |
| A major cause of side effects that lead to emergency room visits in the United States (56,000 per year in the United States) | 41, 42 |
| Associated with increased rates of certain blood cancers | 43 |
| Acetaminophen prenatally or postnatally is associated with increased asthma | 14, 44-46 |

NAPQI, N-acetyl-p-benzoquinone imine.



- 1. 1977 Warning labels recommended for acetaminophen products
- 2. 1980 Warning of Reye's syndrome risk with children's aspirin use children's acetaminophen products increase in sales
- 3. 1982 First acetaminophen scare: seven murders in Chicago using cyanide-laced acetaminophen
- 4. 1986 Second acetaminophen scare: woman murdered in New York using cyanide-laced acetaminophen

Editorial

Autism spectrum disorder: prevalence and cause may be bound together[†]

Emily Simonoff



Summary

Autism has been in the forefront of science and public concern because of reported increases in its prevalence. Changing diagnostic practice and improved identification explain some of this rise, but there may include environmental factors to understand the causes of autism.

Declaration of interest

Research needs to include environmental factors to understand the causes of autism.

Emily Simonoff is Professor of Child and Adolescent Psychiatry, Department of Child and Adolescent Psychiatry, King's College London, Institute of Psychiatry, and National Institute for Health Research Biomedical Research Centre for Mental Health, London.

example, a UK study showed that 42% of 11- to 12-year-olds meeting consensus criteria for autism spectrum disorder (ASD) had not been so identified by services – but virtually all had at least one other diagnosis. However, the DSM-5 Working Group has been concerned about the current diagnostic criteria and is proposing significant changes. The DSM-IV criteria allow an

JAMA. 2013;309(16):1696-1703. doi:10.1001/jama.2013.2270.

Prenatal Valproate Exposure and Risk of Autism Spectrum Disorders and Childhood Autism

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Therese Koops Grønborg, MSc

Merete Juul Sørensen, PhD

Diana Schendel, PhD

Erik Thorlund Parner, PhD

Lars Henning Pedersen, PhD

Mogens Vestergaard, PhD

Importance Valproate is used for the treatment of epilepsy and other neuropsychological disorders and may be the only treatment option for women of childbearing potential. However, prenatal exposure to valproate may increase the risk of autism.

Objective To determine whether prenatal exposure to valproate is associated with an increased risk of autism in offspring.

Design, Setting, and Participants Population-based study of all children born alive in Denmark from 1996 to 2006. National registers were used to identify children exposed to valproate during pregnancy and diagnosed with autism spectrum disorders (childhood autism [autistic disorder], Asperger syndrome, atypical autism, and other

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Valproate during pregnancy associated with increased risk of autism spectrum disorder in the offspring

malformations and delayed cognitive development in the offspring, 1-3 but little is known about the risk of other serious neuropsychiatric disorders. Autistic symptoms have been described in case series of children exposed in utero

til the day of autism spectrum disorder diagnosis, death, emigration, or December 31, 2010, whichever came first.

Main Outcomes and Measures Absolute risk (cumulative incidence) and the hazard ratio (HR) of autism spectrum disorder and childhood autism in children after exposure to valproate in pregnancy.

Soy Formula Linked to Seizures in Autism

"There was a 2.6-fold higher rate of febrile seizures, a 2.1-fold higher rate of epilepsy comorbidity and a 4-fold higher rate of simple partial seizures in the autistic children fed soy-based formula"



Traffic-Related Air Pollution, Particulate Matter, and Autism

Heather E. Volk, PhD, MPH; Fred Lurmann; Bryan Penfold; Irva Hertz-Picciotto, PhD; Rob McConnell, MD

Context: Autism is a heterogeneous disorder with genetic and environmental factors likely contributing to its origins. Examination of hazardous pollutants has suggested the importance of air toxics in the etiology of autism, yet little research has examined its association with local levels of air pollution using residence-specific exposure assignments.

Main Outcome Measures: Crude and multivariable adjusted odds ratios (AORs) for autism.

Results: Children with autism were more likely to live at residences that had the highest quartile of exposure to traffic-related air pollution, during gestation (AOR, 1.98 [95% CI, 1.20-3.31]) and during the first year of life (AOR,

th control chil-

gen dioxide and

um in diameter ith autism dur-

ide: AOR, 1.81

OR, 2.08 195%

2.17 [95% CL

le (exposure to

37-3.091; expo-

Objective: I related air po

Design: Thi cludes data of trol children rolled in the (the Environn Exposure to traffic-related air pollution, nitrogen dioxide, during pregnancy first year of life associated ASD.

dress from the pirth ceruncate and addresses reported from a residential history questionnaire were used to estimate exposure for each trimester of pregnancy and first year of life. Traffic-related air pollution was assigned to each location using a line-source air-quality dispersion model. Regional air pollutant measures were based on the Environmental Protection Agency's Air Quality System data. Logistic regression models compared estimated and measured pollutant levels for children with autism and for control children with typical development.

Setting: Case-control study from California.

Participants: A total of 279 children with autism and a total of 245 control children with typical development.

sure to PM₂₃: AOR, 2.12 [95% CI, 1.45-3.10]; exposure to PM₁₀: AOR, 2.14 [95% CI, 1.46-3.12]). All regional pollutant estimates were scaled to twice the standard deviation of the distribution for all pregnancy estimates.

Conclusions: Exposure to traffic-related air pollution, nitrogen dioxide, PM₂₃, and PM₁₀ during pregnancy and during the first year of life was associated with autism. Further epidemiological and toxicological examinations of likely biological pathways will help determine whether these associations are causal.

JAMA Psychiatry. 2013;70(1):71-77. Published online November 26, 2012. doi:10.1001/jamapsychiatry.2013.266

Environmental Health Perspective April 2011

RESEARCH ARTICLE

OPEN ACCESS

Prenatal Exposure to Organophosphate Pesticides and IQ in 7-Year Old Children

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7-Year Neurodevelopmental Scores and Prenatal Exposure to Chlorpyrifos, a Common Agricultural

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Prenatal Exposure to Organophosphates, Paraoxonase
 1, and Cognitive Development in Childhood

RESEARCH ARTICLE

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Determinants of Agricultural Pesticide Concentrations in Carpet Dust

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HTML version of this article is available at http://dx.doi.org/10.1289/ehp.1206187.

Environ Health Perspect 121:978–984 (2013). doi.org/10.1289/ehp.1206187

Perinatal Air Pollutant Exposures and Autism Spectrum Disorder in the Children of Nurses' Health Study II Participants

Andrea L. Roberts, Kristen Lyall, Jaime E. Hart, Francine Laden, Allan C. Just, Jennifer F. Bobb, Karestan C. Koenen, Alberto Ascherio, And Marc G. Weisskopf, Allan C. Just, Jennifer F. Bobb, Karestan C. Koenen, Alberto Ascherio, And Marc G. Weisskopf, Allan C. Just, Jennifer F. Bobb, The Karestan C. Koenen, Alberto Ascherio, Allan C. Just, Jennifer F. Bobb, The Karestan C. Koenen, Alberto Ascherio, Allan C. Just, Jennifer F. Bobb, The Karestan C. Koenen, Alberto Ascherio, Allan C. Just, Allan C. Just, Jennifer F. Bobb, The Karestan C. Koenen, Allan C. Just, Allan C. Ju

¹Department of Social and Behavioral Sciences, and ²Department of Nutrition, Harvard School of Public Health, Boston, Massachusetts, USA; ³University of California, MIND Institute, Davis, California, USA; ⁴Channing Division of Network Medicine, Brigham and Women's Hospital and Harvard Medical School, Boston, Massachusetts, USA; ⁵Department of Environmental Health, ⁶Department of Epidemiology, and ⁷Department of Biostatistics, Harvard School of Public Health, Boston, Massachusetts, USA; ⁸Department of Epidemiology, Mailman School of Public Health, Columbia University, New York, New York, USA

OBJECTIVE: Air pollution contains many toxicants known to affect neurological function and to have effects on the fetus in utero. Recent studies have reported associations between perinatal exposure to air pollutants and autism spectrum disorder (ASD) in children. We tested the hypothesis that perinatal exposure to air pollutants is associated with ASD, focusing on pollutants associated

mutagens (Agency for Toxic Substances and Disease Registry 2011), and *de novo* DNA mutations have been implicated in ASD etiology (Kinney et al. 2010; Sebat et al. 2007; Smith et al. 2009). Therefore, we focused on

2X

Associated ASD: diesel, lead, manganese, mercury, methylene chloride- highest to lowest risk

and census tract-level socioeconomic measures (e.g., tract median income and percent college educated), as well as maternal age at birth and year of birth. We also examined possible differences in the relationship between ASD and pollutant exposures by child's sex.

RESULTS: Perinatal exposures to the highest versus lowest quintile of diesel, lead, manganese, mercury, methylene chloride, and an overall measure of metals were significantly associated with ASD, with odds ratios ranging from 1.5 (for overall metals measure) to 2.0 (for diesel and mercury). In addition, linear trends were positive and statistically significant for these exposures (p < .05 for each). For most pollutants, associations were stronger for boys (279 cases) than for girls (46 cases) and significantly different according to sex.

CONCLUSIONS: Perinatal exposure to air pollutants may increase risk for ASD. Additionally, future studies should consider sex-specific biological pathways connecting perinatal exposure to pollutants with ASD.

in the association of environmental toxicants with executive function (Braun et al. 2011) have been suggested by prior research. The one previous study that reported associations between exposure to pollutants and ASD according to sex did not find statistically significant differences (Kalkbrenner et al. 2010).

Methods

Selection of cases and controls, We used data from the Nurses' Health Study II, a cohort of 116,430 female nurses from 14 U.S. states that was ostablished in 1989 and hes been fol

KEY WORDS air pollution, autism, diesel, heavy metals, prenatal exposure

Tylenol and Autism

- Tylenol is detoxified/eliminated via sulfation
- Autistic kids are severely impaired in this process
 - Reflects reduced supply of sulfate
- Tylenol therapy further depletes sulfate stores
- This same pathway is necessary to eliminate excess adrenalin and dopamine from the brain
 - Impairment could lead to the formation of neurotoxic substances with psychedelic effects

* Alberti et al., Biol Psychiatry 46, 420–424, 1999.

Autism Epidemic

Awareness of importance of individual susceptibility to environmental pollutants.

Understanding that safe levels for an "average person" leave many individuals at risk.

Understanding biochemical individuality and susceptibility will provide the basis of stronger health policy to protect the most vulnerable.

Sulfur, Asthma and Tylenol

- Autistic kids are especially susceptible to asthma
- There has been a sharp increase in childhood asthma over the past 30 years
- Timing corresponds with aspirin scare in 1980's
- Bronchial tube epithelial cells produce cholesterol sulfate
- Cholesterol sulfate protects from asthma through its effect on profilaggrin
- Tylenol causes increased bronchial constriction and wheezing
- Tylenol depletes sulfur stores through its elimination process

C. Aschwanden, "Studies suggest an acetaminophen-asthma link," Dec. 19, 2011, NY Times

Autism spectrum disorder: interaction of air pollution with the MET receptor tyrosine kinase gene.

Volk HE¹, Kerin T, Lurmann F, Hertz-Picciotto I, McConnell R, Campbell DB.

Author information

Abstract

BACKGROUND: Independent studies report association of autism spectrum disorder with air pollution exposure and a functional promoter variant (rs1858830) in the MET receptor tyrosine kinase (MET) gene. Toxicological data find altered brain Met expression in mice after prenatal exposure to a model air pollutant. Our objective was to investigate whether air pollution exposure

MET CC genotype and high air pollutantsincreased risk ASD compared to heterogeneous
and lower air pollutant exposures.

RESULTS: Subjects with both MET rs1858830 CC genotype and high air pollutant exposures were at increased risk of autism spectrum disorder compared with subjects who had both the CG/GG genotypes and lower air pollutant exposures. There was evidence of multiplicative interaction between NO2 and MET CC genotype (P= 0.03).

CONCLUSIONS: MET rs1858830 CC genotype and air pollutant exposure may interact to increase the risk of autism spectrum disorder.

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Open Access Scientific Reports

Review Artilce Open Access

Prevalence of Autism is Positively Associated with the Incidence of Type 1 Diabetes, but Negatively Associated with the Incidence of Type 2 Diabetes, Implication for the Etiology of the Autism Epidemic

John B Classen*

Classen Immunotherapies, Inc., 6517 Montrose Avenue, Baltimore, MD 21212, USA

Abstract

Background: Epidemics of type 1 diabetes had been linked to inflammation. Previous reports have suggested the prevalence of autism is increased in patients with type 1 diabetes.

Epidemic of autoimmune/inflammation mediated ASD, likely related to simultaneous epidemic of type 1 diabetes in children

Conclusion: This suggests that patients with autoimmune autism likely represent a large subset of patients with autism and that the etiology of the epidemic of autoimmune/inflammation mediated autism in children is likely to be related to the etiology of the simultaneous epidemic of type 1 diabetes in children.

Keywords: Autism; Diabetes; Cortisol

Introduction

Type 1 diabetes has been linked to an increased risk of autism [1]. Furthermore patients with autism have an increased family history of

Results

Data from two published studies of US children allowed direct correlation between the incidence of type 1 diabetes, type 2 diabetes [11] and the prevalence of autism [5] in multiple different races (Whites, Blacks, Hispanics, Asians, Native Americans). Data on the prevalence

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Citation: Transl Psychiatry (2014) 4, e360; doi:10.1038/tp.2014.4 © 2014 Macmillan Publishers Limited All rights reserved 2158-3188/14

udies)

ASD, a

www.nature.com/tp

ORIGINAL ARTICLE

Environmental toxicants and autism spectrum disorders: a systematic review

DA Rossignol¹, SJ Genuis² and RE Frye³

hav een Toxicants implicated: pesticides, phthalates, PCBs, env dies esti solvents, toxic waste sites, air pollutants, heavy metals h of inve nine electronic scientific databases through November 2013 was performed. In the first category examining ASD risk and estimated toxicant exposures in the environment, the majority of studies (34/37; 92%) reported an association. Most of these studies were retr e, case Genes implicated paraoxonase PON1, glutathione S-transferase oxic rep (GSTM1 and GSTP1, t, δ-aminolevulinic acid dehydratase, SLC11A3,

Although the involvement of genetic abnormalities in autism spectrum disorders (ASD) is well-accepted, recent studies point to an equal contribution by environmental factors, particularly environmental toxicants. However, these toxicant-related studies in ASD

large number was dedicated to examining heavy metals. Such studies demonstrated mixed findings, with only 19 of 40 (47%) case–control studies reporting higher concentrations of heavy metals in blood, urine, hair, brain or teeth of children with ASD

and the metal regulatory transcription factor 1

Environmental and State-Level Regulatory Factors Affect the Incidence of Autism and Intellectual Disability

Andrey Rzhetsky^{1,2,3}*, Steven C. Bagley⁴, Kanix Wang^{2,5}, Christopher S. Lyttle⁶, Edwin H. Cook, Jr.⁷, Russ B. Altman⁸, Robert D. Gibbons⁹

- 1 Department of Medicine, the University of Chicago, Chicago, Illinois, United States of America, 2 Institute for Genomics and Systems Biology, the University of Chicago, Chicago, Illinois, United States of America, 3 Computation Institute, Department of Human Genetics, the University of Chicago, Chicago, Chicago, United States of America,
- 4 Department of Genetics, Stanford University, Stanford, California, United States of America, 5 Committee on Genetics, Genomics, and Systems Biology, the University of Chicago, Chicago, Illinois, United States of America, 6 The Center for Health and the Social Sciences, the University of Chicago, Chicago, Illinois, United States of America,
- 7 Department of Psychiatry, Institute for Juvenile Research, University of Illinois at Chicago, Chicago, United States of America, 8 Departments of Bioengineering, Genetics, Medicine, and Computer Science, Stanford University, Stanford, California, United States of America, 9 Departments of Medicine and Health Studies, the University of Chicago, Center for Health Statistics, Chicago, Illinois, United States of America

ASD incidence rates linked to rates of congenital malformations of the reproductive system in males-

283% increase for every percent increase in incidence of malformations

congenital malformations in males (excluding those affecting the reproductive system) appeared to significantly affect both phenotypes: 31.8% ASD rate increase (CI: [12%, 52%], $p < 6 \times 10^{-5}$), and 43% ID rate increase (CI: [23%, 67%], $p < 6 \times 10^{-5}$). Furthermore, the state-mandated rigor of diagnosis of ASD by a pediatrician or clinician for consideration in the special education system was predictive of a considerable decrease in ASD and ID incidence rates (98.6%, CI: [28%, 99.99%], p = 0.02475 and 99% CI; [68%, 99.99%], p = 0.00637 respectively). Thus, the observed spatial variability of both ID and ASD rates is associated with environmental and state-level regulatory factors; the magnitude of influence of compound environmental predictors was approximately three times greater than that of state-level incentives. The estimated countylevel random effects exhibited marked spatial clustering, strongly indicating existence of as yet unidentified localized factors driving apparent disease incidence. Finally, we found that the rates of ASD and ID at the county level were weakly but American Journal of Epidemiology

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DOI: 10.1093/aje/kwn250 Advance Access publication October 21, 2008

Original Contribution

Advanced Parental Age and the Risk of Autism Spectrum Disorder

Maureen S. Durkin, Matthew J. Maenner, Craig J. Newschaffer, Li-Ching Lee, Christopher M. Cunniff, Julie L. Daniels, Russell S. Kirby, Lewis Leavitt, Lisa Miller, Walter Zahorodny, and Laura A. Schieve

Initially submitted March 21, 2008; accepted for publication July 16, 2008.

This study evaluated independent effects of maternal and paternal age on risk of autism spectrum disorder. A case-cohort design was implemented using data from 10 US study sites participating in the Centers for Disease Control and Pre 994 birth cohort in-Maternal and paternal age were cluded 253,347 ,251 children aged 8 years with co s having an autism independently associated with autism spectrum disord ition, Text Revision criteria. After adjustment for the other parent's age high order maternal education, and other covariates, both maternal an nal age >35 Firstborn offspring older parents 3x times vs. 25-29 ye 's vs. 25-29 years = 1.4ore likely to more likely to develop ASD develop auti < 40 years

(odds ratio = 3.1, 95% confidence interval: 2.0, 4.7). The increase in autism risk with both maternal and paternal age has potential implications for public health planning and investigations of autism etiology.

autietic disorder: hirth order: maternal age: naternal age

Air pollution and brain inflammation

Authorized Charles (2014) No. 2014 - Discharge (2014) No.

Urban air pollutants may damage IQs before baby's first breath, scientists say

Epidemiology. 2010 Jun 17. [Epub ahead of print]

Perinatal Exposure to Hazardous Air Pollutants and Autism Spectrum Disorders at Age 8.

Kalkbrenner AE, Daniels JL, Chen JC, Poole C, Emch M, Morrissey J.

Scientists find 'baffling' link between autism and vinyl flooring

Children who live in homes with vinyl floors, which can emit phthalates, are twice as likely to have autism, according a Swedish and U.S. researchers. Scientists call the discovery "intriguing and baffling." Experts suspect that genetic and

Prenatal exposure to \(\beta^2\)-adrenergic receptor agonists and risk of autism spectrum disorders

Lisa A. Croen · Susan L. Connors · Marilyn Matevia ·

Yinge Qian · Craig Newschaffer ·

Andrew W. Zimmerman

Terbutaline exposure for >2 days during the third trimester was associated with more than **4X increased** risk ASDs

© Springer Science+Business Media, LLC 2011

Abstract This study aims to investigate the association between prenatal exposure to terbutaline and other β2 adrenergic receptor (B2AR) agonists and autism spectrum disorders (ASDs). The methodology used is a case–control study among children born from 1995 to 1999 at Kaiser Permanente Northern California hospitals. Cases (n=291) were children with an ASD diagnosis; controls (n=284) were children without ASDs, randomly sampled and frequency-matched to cases on sex, birth year, and delivery hospital. Exposure to B2AR agonists during 30 days prior to conception and each trimester of pregnancy was ascertained from prenatal medical records and health plan

databases. The frequency of exposure to any B2AR agonist during pregnancy was similar for mothers of children with ASD and mothers of controls (18.9% vs. 14.8%, P=0.19). Exposure to B2AR agonists other than terbutaline was not associated with an increased risk for ASDs. However, terbutaline exposure for >2 days during the third trimester was associated with more than a fourfold increased risk for ASDs independent of indication although the limited sample size resulted in an imprecise and nonsignificant effect estimate (OR_{adj}=4.4; 95% confidence interval, 0.8–24.6). This analysis does not offer evidence linking B2AR exposure in pregnancy with autism risk. However, exposure

Deficits in Bioenergetics and Impaired Immune Response in Granulocytes From Children With Autism.

Napoli E1, Wong S, Hertz-Picciotto I, Giulivi C.

Author information

Abstract

Despite the emerging role of mitochondria in immunity, a link between bioenergetics and the immune response in autism has not been explored. Mitochondrial outcomes and phorbol 12-myristate 13-acetate (PMA)-induced oxidative burst were evaluated in granulocytes from age-, race-, and

mitochon

Majority of granulocytes exhibited defects
in oxidative phosphorylation, immune
race-, and typically granulocytes exhibited defects
in oxidative phosphorylation, immune
response, and antioxidant defense

dysfunction in children with autism was accompanied by a lower (26% of TD children) oxidative burst by PMA-stimulated reduced nicotinamide-adenine dinucleotide phosphate (NADPH) oxidase and by a lower gene expression (45% of TD children's mean values) of the nuclear factor erythroid 2-related factor 2 transcription factor involved in the antioxidant response. Given that

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Health & Place ■ (■■■) ■■■



www.elsevier.com/locate/healthplace

Proximity to point sources of environmental mercury release as a predictor of autism prevalence

Raymond F. Palmer^{a,*}, Stephen Blanchard^b, Robert Wood^a

^aUniversity of Texas Health Science Center, San Antonio Department of Family and Community Medicine, 7703 Floyd Curl Drive, San Antonio Texas, Mail Code 7794, TX 78229-3900, USA



Autism Spectrum Disorders in Relation to Distribution of Hazardous Air Pollutants in the San Francisco Bay Area

Gayle Divisi Healtl

Toxins with greatest Risk: Mercury, Cadmium, Nickel, Trichloroethylene, Vinyl Chloride



NIH Public Access

Author Manuscript

Neurotoxicology. Author manuscript; available in PMC 2013 October 01.

Published in final edited form as:

Neurotoxicology. 2009 September; 30(5): 822-831. doi:10.1016/j.neuro.2009.01.011.

Associations between indoor environmental factors and parental-reported autistic spectrum disorders in children 6-8 years of age

Malin Larsson^a, Bernard Weiss^b, Staffan Janson^a, Jan Sundell^{c,d}, and Carl-Gustav Bornehag^{a,c,f}

Abatuant

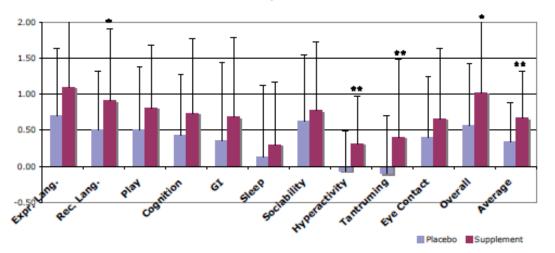
ASD significant correlations 1) maternal smoking 2) male sex 3) family financial problems 4) window condensation, proxy for low ventilation rate 5) PVC flooring, especially in the parents' bedroom. Physician-diagnosed asthma at baseline were associated with ASD five years later.

smoking, allergic symptoms, type of residence, moisture-related problems, and type of flooring material, which included polyvinyl chloride (PVC). The 2005 survey, based on the same children, now 6.8 years of any also asked if during the intervening period, the shild had been discussed.

Oxidative Stress, Methylation, and Glutathione:

| | units | Neuro-typicals (n = 44) | Arizona Treatment Group (n = 18) | | | |
|--|-----------|----------------------------|-------------------------------------|---------------------|----------|-----------|
| | | | Pre | Post | % change | P - value |
| Piasma nitro-tyrosine | Ug/I | 7.4 +/-5.1 | 14.1 +/-6.5 | 9.9 +/-5.4 | -29% | 0.004 |
| Reduced plasma glutathione (GSH) | nmoi / mi | 4.09 +/-0.79 | 3.27 +/-0.59 | 3.84 + / - 0.61 | + 17% | 0.0008 |
| Oxidized glutathione (GSSG) | nmoi / mi | 0.362 +/-0.10 | 0.467 +/-0.12 | 0.403 + / - 0.09 | - 14% | 0.02 |
| Ratio of oxidized to reduced plasma glutathione | | 0.093 +/-0.04 | 0.150 +/-0.05 | 0.109 +/-0.03 | -27% | 0.002 |
| SAM (RBC) | umol /dl | 228.4 +/-12 | 218 +/-17 | 230 +/-16 | + 6% | 0.003 |
| Uridine (plasma) | 10 - 6 M | 7.9 +/-2.7 | 16.2 +/-9.5 | 10.4 +/-4.7 | -36% | 0.008 |

Parental Global Impressions - Revised



| ANRC Essentials Formulation (recommended dosage for a 60 pound ch |
|---|
| 8000 IU Vitamin A (as mixed carotenoids from algae) |
| 500 mg Vitamin C (as calcium ascorbate) |
| 1500 IU Vitamin D3 |
| 100 mg Vitamin E (as mixed tocopherols, including gamma tocopherol) |
| 50 mcg K1, 50 mcg K2 |
| 30 mg Vitamin B1 (as thiamin hydrochloride) |
| 40 mg Vitamin B2 (Riboflavin) |
| 50 mg Vitamin B3 (as inositol hexanicotinate) |
| 20 mg Pyridoxine hydrochloride, 20 mg pyridoxal 5 phosphate (P5P) |
| 600 mcg MTHF (methyl-tetra-hydrofolate) |
| 600 mcg for Vitamin B12 as methylcobalamin |
| 500 mcg Biotin |
| 30 mg Pantothenic acid (as calcium d-pantothenate) |
| 100 mcg lodine (as potassium iodide) |
| 350 mcg Lithium (as lithium orotate) |
| 250 mg Choline (as choline bitartrate) |
| 100 mg Inositol |
| 100 mg Calcium (calcium ascorbate, calcium pantothenate, calcium citrate) |
| 250 mg Magnesium (as magnesium citrate) |
| 15 mg Zinc (as zinc sulfate) |
| 40 mcg Selenium (80% as selenomethionine, 20% as sodium selenite) |
| 1 mg Manganese (as manganese aspartate) |
| 70 mcg Chromium (as chromium picolinate) |
| 100 mcg Molybdenum (as molybdenum nicotinate glycinate) |
| 50 mg Potassium (potassium chloride) |
| 500 mg MSM (methylsulfonylmethane) |
| 50 mg Co-Enzyme Q10 |
| 45 mg N-acetyl-cysteine |
| 200 mg Acetyl-L-carnitine |
| |



200 mg L-carnitine





For Release: April 25, 2011 12:01 am (ET)

AAP SAYS U.S. FAILS TO PROTECT CHILDREN FROM HAZARDOUS CHEMICALS

PEDIATRICS

OFFICIAL JOURNAL OF THE AMERICAN ACADEMY OF PEDIATRICS

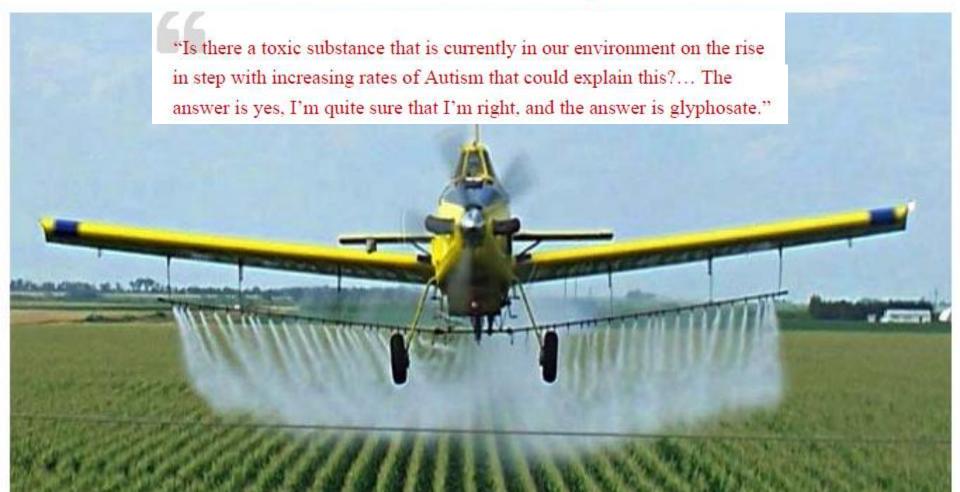
Attention-Deficit/Hyperactivity Disorder and Urinary Metabolites of Organophosphate Pesticides

13

The findings showed that children with higher urinary levels of organophosphate metabolites were more likely to meet the diagnostic criteria for ADHD.

2015

Autism Will Afflict Half Of The American Children By 2025, And Glyphosate Is To Blame, MIT Doctor Says





OA Autism 2014 Jun 10;2(2):11.

Revi

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Etiology of autism spectrum disorders: Genes, environment, or both?

C A Shaw¹, S Sheth¹, D Li¹, L Tomljenovic¹

Abstract

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Introduction

Thus far, most of the research on both neurodevelopmental and neurodegenerative disorders has been may provide important clues of Al's putative role in autism. Because of the

connection tight between development of the immune and the central nervous system, the possibility several decades) rules out a purely genetic origin.

In ALS, for example, much of the literature of the last 20 years has ch for motor

focus under Heritability factors cannot adequately explain all reported less poten cases nor their drastic increase over the last few decades

some forms of autism are clearly the fact remains that genetic,

periou. These include a number of There is now sufficient evidence from variations on the so-called "toxic gain of oding

Evidence shows autism may in part result from early-life decades immune insults induced by environmental xenobiotics.

twins have now shown that common environmental factors account for 55% of their risk for developing autism while genetic susceptibility explains only 37% of cases. Because

only medical intervention that we attempt to deliver to every living human on earth and that by far the largest target population for vaccination are healthy

protein TDP-43, FUS, or VEGF, or the newest player, C9orf72, and all have added to the complexity of the polygenic picture without necessarily increasing the total percentage of all devi

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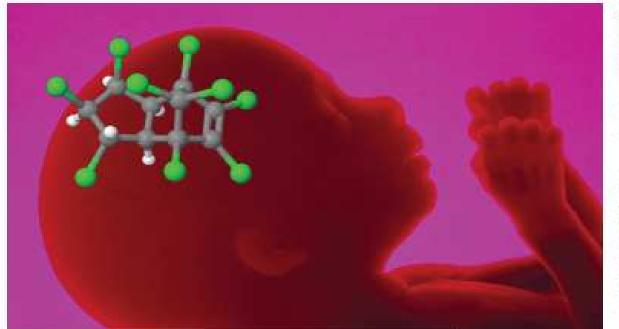
Clues to Autistic Behaviors

Exploring the Role of Endocrine Disruptors

Two lines of evidence suggest that endocrine disruption may be a factor in autism spectrum disorders (ASDs). First, the observation that males may be four times as likely to be diagnosed with ASDs as females suggests hormonal involvement. Second, adrenal, gonadal, and thyroid hormones play an important role in fetal Examination Survey. After the researchers adjusted for potential confounding variables, they found that higher maternal exposures to trans-nonachlor and PBDE-28 were associated with higher average SRS scores. Trans-nonachlor is a component of the highly persistent banned pesticide chlordane, and PBDE-28 is one of the polybrominated diphenyl ether compounds used as flame retardants in commercial goods containing polyurethane foam (including furniture and mattresses) made before 2005. The study also found negative associations between four

Endocrine disrupters (ED) factor in ASD: 1) males 4 X as likely to be diagnosed 2) ED impact adrenal, gonadal, and thyroid hormones role in fetal neurodevelopment causing abnormal brain development

mers. ated opliults hors s in each.



is innovative and mirrors the real world, where we are all exposed to a mixture of chemicals, and where the neurotoxicants may have different effects that may even depend on the time of exposure," says Philippe Grandjean, an adjunct professor of environmental health at the Harvard School of Public Health, who was not involved with the study.

But Grandjean also cautions against comparing apples and oranges. "Those substances that have been measured with large imprecision, e.g., because of common short-term variability, are likely to be underestimated as possible contributors to autismlike behaviors," he explains. On the other hand, "persistent substances will have less imprecision and could therefore erroneously appear as if they are more important."

Although the study measured only

Biomarkers for Autism Stephanie Seneff PhD people.csail.mit.edu/seneff/

- Disrupted gut bacteria; inflammatory bowel
- Low serum sulfate
- Methionine deficiency
- Serotonin and melatonin deficiency
- Defective aromatase
- Zinc and cobalamin deficiency
- Urinary p-cresol
- Mitochondrial disorder
- Seizures; Glutamate toxicity in the brain
 These can all be explained as potential
 effects of glyphosate on biological systems





Review

Empirical Data Confirm Autism Symptoms Related to Aluminum and Acetaminophen Exposure

Stephanie Seneff 1,*, Robert M. Davidson 2 and Jingjing Liu 1

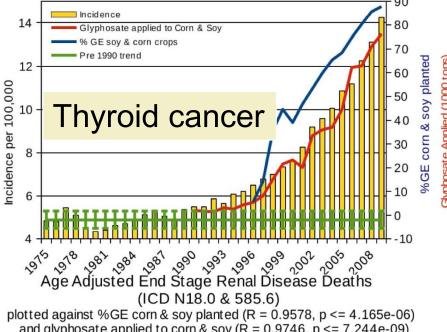
Glyphosate enhances aluminum toxicity

Glyphosate interferes with acetaminophen metabolism

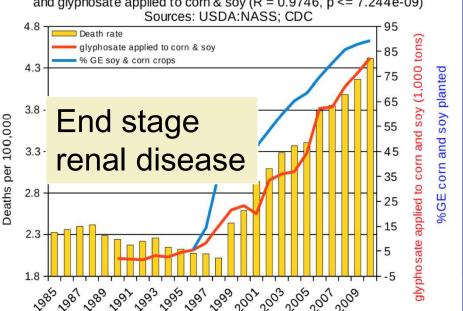


Thyroid Cancer Incidence Rate (age adjusted)

plotted against glyphosate applied to U.S. corn & soy (R = 0.988, p <= 7.612e-09) along with %GE corn & soy crops R = 0.9377, p <= 2.152e-05 sources: USDA:NASS: SEER

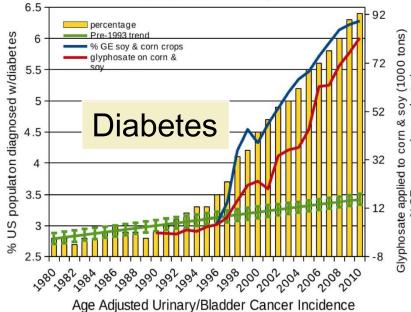


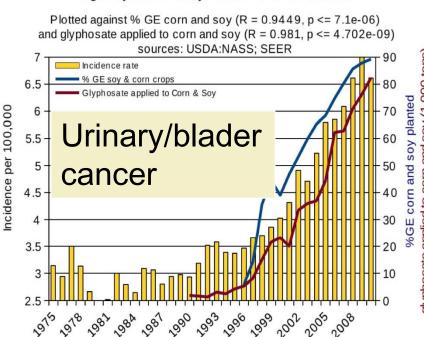
and glyphosate applied to corn & soy (R = 0.9746, $p \le 7.244e-09$)



Prevalence of Diabetes in US (age adjusted)

plotted against glyphosate applied to corn & soy (R = 0.971, p <= 9.24e-09) along with %GE corn & soy grown in US (R=0.9826, $p \le 5.169e-07$) sources: USDA:NASS; CDC





THE MICROBIOME IN AUTISM SPECTRUM DISORDER

Gut bacteria in children with autism spectrum disorders: challenges and promise of studying how a complex community influences a complex disease

Rosa Krajmalnik-Brown^{1,2*}, Catherine Lozupone³, Dae-Wook Kang¹ and James B. Adams⁴

¹Swette Center for Environmental Biotechnology, Biodesign Institute, Arizona State University, Tempe, AZ, USA: 2

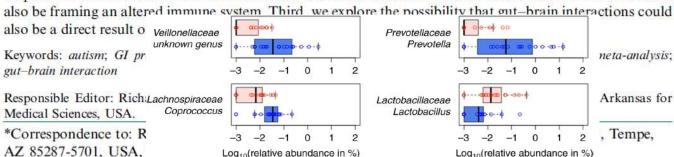
USA; of Mat

Shifted microbiota from westernization could be framing an altered immune system

Recent studies suggest a role for the microbiota in autism spectrum disorders (ASD), potentially arising from

their ro Gut brain interactions could be a direct depende are com result of microbially produced metabolites. summai the role

we provide some evidence that the shifted microbiota can be a result of westernization and that this shift could also be framing an altered immune system. Third, we explore the possibility that gut-brain interactions could

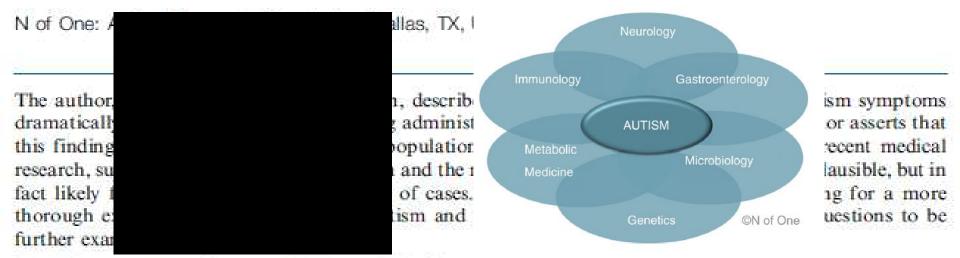




THE MICROBIOME IN AUTISM SPECTRUM DISORDER

An n=1 case report of a child with autism improving on antibiotics and a father's quest to understand what it may mean

John Rodakis*



Keywords: autism; microbiome; antibiotic; n = 1; N of One; vancomycin; amoxicillin

Responsible Editor: Richard E, Frye, Arkansas Children's Hospital Research Institute and University of Arkansas for Medical Sciences, USA.

*Correspondence Microbial Ecology in Health & Disease 2015, 26: 26382

Commentary

Life Imprints: Living in a Contaminated World

David Crews^{1,2} and Andrea C. Gore^{2,3}

¹Section of Integrative Biology, ²Institute for Cellular and Molecular Biology, and ³Division of Foundation of Texas at Austin, Austin, Texas, USA

BACKGROUND: The links between nature and nurture need to be redefined to accommodate anthropogenic chemical contamination. Although some local remediation of contamination has occurred toon, by never expenses all levels of biological organization. Where the change, we must move on and consider the types of adaptations that could occur as a consequence.

OBJECTIVES: We propose a paradigm shift in the field that integrates various disciplines involved in

To understand the future, we must study what is likely to happen and less to what has happened..

illuminate how the causal mechanisms and functional outcomes of related processes operate at each level of biological organization while at the same time revealing the relations among the levels.

Mary Claire King and BRCA 1 and 2





"The lifetime risk of breast cancer among female mutation carriers is presently 82%. Risks appear to be increasing with time. Before 1940 it was 24%. Lack of physical exercise and obesity in adolescence may be important modulating factors for risk in carriers". Science 2003; 302: 643-50.

Original Investigation

The Familial Risk of Autism

```
RESULTS In the sample, 14 516 children were diagnosed with ASD, of whom 5689 had autistic
disorder. The RRR and rate per 100 000 person-years for ASD among monozygotic twins was
      Environmental factors as important as genes in
estir
                                                                           or
dizy
                         understanding ASD
sibli
                                                                          alf
siblings, 3.3 (95% CI, 2.6-4.2; rate, 492 for exposed vs 94 for unexposed); for paternal half
sil
       We were surprised by our findings as we did not expect
(9
    importance of environmental factors in ASD to be so strong
aι
including only additive genetic and nonshared environmental effects. The ASD heritability
wa
    Factors which are unique to the individual, or 'non-shared
est
   environments' were the major source of environmental risk.
CONCLUSIONS AND RELEVANCE Among children born in Sweden, the individual risk of ASD
```

and autistic disorder increased with increasing genetic relatedness. Heritability of ASD and autistic disorder were estimated to be approximately 50%. These findings may inform the



RESEARCH Open Access

A comparison of temporal trends in United States autism prevalence to trends in suspected environmental factors

Cynthia D Nevison

- Data suggest ~75-80% of the tracked increase in autism since 1988 is due to an actual increase in the disorder rather than to changing diagnostic criteria
- Polybrominated diphenyl ethers (fire retardants), *aluminum* adjuvants, and the herbicide *glyphosate* have increasing trends that correlate positively to the rise in autism.

C. Nevison Environmental Health 2014;13:73.



Autism and the Environment Workshop Agenda

http://www.iom.edu/CMS/3740/35684/39826/42257.aspx

CANDIDATE FACTORS I

- Childhood vaccines
- Thimerosal
- Prenatal ultrasound, Pasko Rakic Antenatal steroids
- Epidural anesthesia
- Magnesium sulfate
- Maternal/paternal age

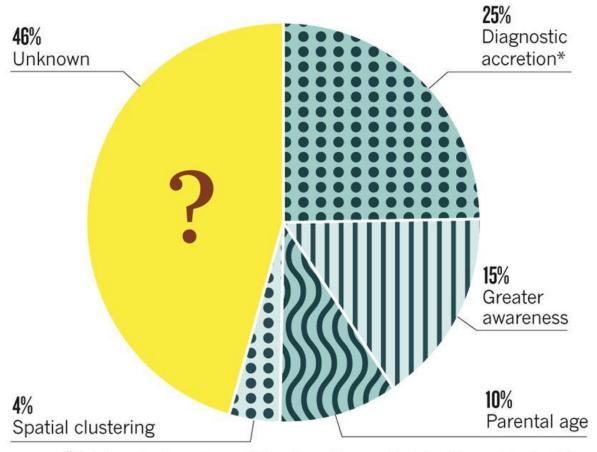
CANDIDATE FACTORS II

- Tocolysis for preterm labor
 - Terbutaline toxicity in rats
- ART/ICSI & cryopreservation of embryos; ICSI and imprinting defects causing AS and BWS
- Chemical exposures
- Folate, vitamins, diet

94

"If it is an environmental cause contributing to an increase, we certainly want to find it."

Reasons: unclear



^{*}Children who formerly would have been diagnosed solely with mental retardation

K. Weintraub, Nature 479, Nov. 3 2011, 22-24.

Children's Environmental Mount Sinai Medical Center Health Center



"It is increasingly clear that environmental exposures also contribute to autism"

Biomarkers, Immune-Mediated Disorders and Autism June 1, 2013

_{By} Judy Van de Water, Ph.D.

Environmental Health



ASDs: Need New Model

- Looks to molecular causes:
 - Complexity theory
 - Systems biology
 - Network theory
- Defines subtypes
- Family as colleagues
- Biology before pharmacology