







Physical Signs Long Forgotten:

- :
- .
- Case 1: "Eating has always been a problem for him. He has never shown a normal appetite."
 Case 2: "Large and ragged tonsits."
 Case 3: darker and fever following smallpox vaccination ... healthy except for large tonsits and advisords.
 Case 5: Annual and fever following smallpox vaccination healthy except for large tonsits and advisords.
 Case 5: Annual every normality any kind of nourishment at three months. Tuber for live tree for large tonsits."
 Case 7: Jonne and Fever following smallpox vaccination healthy compliants.
 Case 7: Annual every normality of the series of age. At camp he slid into avitaminosis and mainutrition but offered almost to verbal compliants.
 Case 7: vormited all food from birth through the third month.... Gase 10: request hospitalizations because the feeding problem ... repeated cobs and offits media.
 Case 9: none of the usual children's diseases."
 Case 9: none of the usual children's diseases. The series of the series .
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Physiology: is the study of the mechanical, physical, and biochemical functions of living organisms.















Laboratory Investigations Can Help Clinical Decision Making Dys DysDysBytestighet Acid
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What is Autism: Parents Perspective

- Mismanagement of incoming stimuli:
 External environment
 - The five senses which bring information to the brain – Internal environment
 - Response to the senses
 - Regulation of the response to the stimuli
 - Inappropriate response to internal cues
 Responses are all biochemically mediated!

What is Autism: A Physician's Perspective

Genetically influenced



What is Autism: A Physician's Perspective

Genetically influencedEnvironmentally triggered



What is Autism: A Physician's Perspective

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- Environmentally
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- Involves the BODY and the BRAIN- attached to the body, downstream, and all that is "wet"

What is Autism: A Physician's Perspective

- Genetically influenced
- Environmentally triggered
 Involves the BODY and the BRAIN- attached to the body, downstream, and all that is
- "wet" Metabolic, immunologic, gastro, neuro and endocrine abnormalities



What is Autism: A Physician's Perspective

- Genetically influenced
- Environmentally triggered
- Involves the BODY and the BRAIN- attached to the body, downstream, and all that is "wet"
- Metabolic, immunologic, gastroenterological abnormalities
- Is treatable and recoverable
- ed he boody, it is c, c,



Parent's Concerns

- Foul smelling loose Coversears yellow green stools
 - Strange "eeeee" sounds
- · Bloated belly No language
- NOT SLEEPING: hard to get down, mom has to sleep with him
- Does not do well with running errands as he tantrums with each transition
- Tantrums severely, sometimes triggered, sometimes no clue · Fevers can come and go
- Not potty trained, no interest

Commonly Seen During an Office Visit

- Poor eye contact
- Non-stop moving
- · Climbing on everything
- Language deficit
- · Repetitive play on office Brio train
- Restricted diet: loves
- bread, cheese, juice, milk, chips (grazing in
- office)

• Physical exam: - Dark circles below eyes Dry, pale, translucent skin

- Dull hair, straw-like - White tongue
- Dilated pupils
- Low muscle tone - Tender "mass" in LLQ
- Enlarged lymph nodes

Considerations: Wrap child in blanket (the baby burrito) Recline in chair with child in lap 22 gauge intra cath needle

Blood Draws:

Sedation if necessary



Listing What's Biologically Different

- Gut Problems
- Inflammatory Bow Reflux Esophagitis
- Gastritis

- Dysbiosis
 Leaky gut
 Malabsorption
- Persistent Viral Illness
- Persistent Viral Illness Food allergy Recurring illnesses Auto immune/auto inflammatory processes-especially brain

Neurologic

Opjate effect Metaboli/Detoalfication Pathway Disruption Purine disorders Elevated ammonia Omega 3 deficits Suffation defect Methylation disorders Methylation disorders Methylation disorders Dopamie defect Moay metal burden Chromosomal issues

Seizures/Sensory Issues Low muscle tone- especially trunk Perfusion Defects- SPECT

Opiate effect

RESTORATION OF CHILDREN'S HEALTH

BREAK THE GRIDLOCK, INTERRUPT THE VICIOUS CYCLES, RESTORE NORMAL FUNCTION:

- Removing dietary/environmental stressors 1.
- 2. Bolstering nutrition
- Relieving inflammation 3.
- 4. Improving broken biochemistry
- 5. Improving immune function

6. Dr Pangborn

Detoxification



























Abnormal Bowels Soiling

Constipation • Diarrhea

- Anorexia • Nocturnal awakenings
- Reflux
- · Foul Smelling Stools
- · Foul Smelling Breath
- · Protuberant Belly

Laboratory Evaluation

- Urine Organic Acids Yeast
- Urine Opiates/Peptides
- Comprehensive Stool
- Cultures and Parasitology
- Food Sensitivity Profile Inflammatory Markers, ie IBD profile, CRP, ESR,
- Overall digestion

 - Endoscopy
- Calprotectin
- Dysbiotic Bacteria Parasite
- X ray/KUB

20				
:5				

What is Stool?

- 10 parts water
- 1 part bacteria
- 1 part fiber
- 1 part mucous, fat, dead cells, protein



Undigested Food in Stool

- Gastrointestinal Abnormalities in Children with Autistic Disorder
- Autistic Disorder RESULTS: Histologic examination in these 36 children revealed grade 1 or II reflux esophagitis in 25 (69.4%), chronic gastritis in 15, and chronic duodenitis in 24. The number of Paneth's cells in the duodenal crypts was significantly elevated in autistic children compared with non-autistic control subjects. Low intestinal carbohydrate digestive enzyme activity was reported in 21 children (58.3%) Horath K Bandentinicu, IC. Rebotm & Discherger C. Tildren (1.
- Linute of Control (J) Horvath K, Papadimitriou JC, Rabsztyn A, Drachenberg C, Tildon JT. J Pediatri 1999 Nov:135(5):559-63. Department of Pediatrics, University of Maryland School of Medicine, Baltimore, USA. PMID: 10547242 [PubMed indexed for MEDLINE]

Very Large Stools

 Constipation With Acquired Megarectum in **Children With Autism** Conclusions: Constipation is a frequent finding in children with gastrointestinal symptoms and autism, particularly in the rectosigmoid colon, often with acquired megarectum. PEDIATRICS Vol. 112 No. 4 October 2003 Nadeem Afzal, MRCPCH*; Simon Murch, PhD*; Kumran Thirrupathy, MBBS*; Leslie Berger, FRCR‡; Andrew Fagbemi, MRCPCH*; and Robert Heuschkel, FRCPCH

Large Amount of Flatulence Soft Serve **Snake Stools** · Fermenting · Poor absorption and • Baring down against a partially closed organisms digestion sphincter Incomplete • Straining • Fructose breakdown of • Lactose polysaccharides • Gluten/sprue Air swallowing · Old stool

Pebble Stool

- · Small pebbles often result due to the lack of fiber.
- · Decreased fluid intake

Rainbow Stools

- · Yellow: fat in stool
- · Red: blood, beets
- Green: GI infection
- Black: blood, Bismuth, iron
- · White/grey: lack of bile flow
- Reddish-black: intussusception (currant jelly)

Gut Disorders in Autism

Historical Clues: Difficulty breastfeeding Persistent Colic Gastro-esophageal reflux Infantile eczema Infantile eczema Food sensitivities Failure to thrive Frequent antibiotics (abnormal flora) Abnormal posturing Self injurious behavior Poor sleep

Physical/Lab Clues: Abnormal stools Abnormal cytokine profile Lymphonodular Hyperplasia of ileum Esophagitis Gastritis

When bowels improve.....

- Stools improve, still soft-serve like
- · Eye contact improves
- Word approximations may start
- Tantrums are remarkably improve
- Appetite improves; trying new things
 Just seems "happier in the skin he/she is in"
- · Health improves

Diarrhea

- 1-6 (or more) liquid movements per day
- Some can be explosive
- Can be very foul
- · Can have varied colors

Management of Diarrhea

- Diet: remove gluten, dairy, consider removing fruits and fruit juices
- Digestive Enzymes
- Antibiotics
- Antifungals
- Antiparisitics
- Zinc: adding 20-60mg per day
- Fiber: soluble 1 to 4 tsp per day

Management of Diarrhea

- Monolaurin
- · Ionic silver
- Transfer factor, such as colostrum
- Oral immunoglobulins (off label)
- Consider immunodeficiency work-up (ie, IgA deficiency)
- Secretin IV (off label)





Constipation

- · Can be daily to once weekly
- · Can be small balls, pellets, to soft ball sized
- · Very foul smelling • Don't forget about
- Hirshsprung's Disease



Treatments for Constipation

- Enemas daily for up to a Vitamin C 2000week if necessary
- · Glycerin suppository Polyethylene glycol
- Aloe
- Fiber (may require larger amounts)
- 10,000mg daily as tolerated (buffered) Magnesium 200-600mg as tolerated
- Magnesium citrate Water soluble contrast enema (in radiology suite)
- · Admit, NG tube, etc

Fiber

• Soluble Fiber helps prevent constipation, which in turn, diminishes the potential for dysbiosis.

Dysbiosis

- Stools studies can give information of many aerobic overgrowths as well as current probiotic bacterial counts
- Urine organic acid studies can give information on anaerobic bacterial growth
- · Fungal cultures with sensitivities will point to antifungal medication preferences
- Parasitology studies, when positive, can help with therapeutic interventions





Yeast Treatments

- Fluconazole 5mg/kg/day x 14-30 day
- Ketoconazole 5mg/kg/day x 14-30 days
 Itraconazole 5mg/kg/day x 14-30 days
- Nystatin compounded 1-2 million unit 2-3 times daily
- Amphotericin B 250mg 500mg four times daily
 Check Liver Function Tests every 8 weeks if on "azoles" for longer periods
- Bicarbonate/alkalinized water may play a role
- · Assure daily stooling

Yeast Treatments

- Monolaurin ¼ to ½ tsp twice daily
- Oregano 0.2 ml twice daily
- Olive leaf extract varies with preparation
- · Caprylic acid 500-1000mg with meals
- · Berberine (very bitter)
- Garlic 1-2 pills per day
- Nano particle silver ½ tsp twice daily

Treatment for Bacteria: Gram Positive Bacteria

- Azithromycin 5mg/kg/day once daily 5-30 days
- Cephalexin 25-50mg/kg/day divided twice daily
- Vancomycin 5-10mg/kg/day divide three times daily ORALLY (not absorbed)
- Clindamycin 5-10mg/kg/day divided three times daily orally



Treatment for Bacteria: Gram Negative Bacteria and Some Parasites

- Metronidazole and Metronidazole Benzoate (suspension form of metronidazole)
 30mg/kg/d divided three times daily for 7-10 days
- Nitaoxanide suspension: use published tables for 3 days...or longer

Small Bowel *Erosive* Ulcerations (with enteric protein loss)

Inflammatory Bowel Disease

- Mesalamine: individual dosing, starting with 250mg twice daily.
- Budesonide 3mg twice daily for up to 8 weeks
 Oral steroids
- Dietary changes
- Regular bowel movements
- Zinc 40-60 mg per day
- Purified distilled Aloe
- Fiber
- Probiotics

Inflammatory Bowel Disease

- Antiviral therapies
- Trichuris Suis Ova therapy
- IV Immunoglobulin therapy
- Curcumin
- Omega 3's
- Pain/discomfort management such as compounded ibuprofen



Reflux

- This is a common allergic/noxious stimuli response
- Associated with otitis media, sinusitis, asthma
- Dietary changes help
- Antacids: promote healing
- Gastric motility/constipation

Reflux Treatments

- Remove exogenous allergens
- Famotidine 20 to 40 mg once daily
- Ranitidine 5-10mg/kg/day divided twice daily
- Omeprazole 5-10kg give 5mg per day, 10-20kg give 10mg per day, over 20 kg, give 20mg per day
- Cimetadine 20-25mg/kg divided in 4 dosesEsomeprazole : 1-10yrs 10mg once daily, 11yrs
- and older, 20-40mg once daily

Biofilms



Current Therapeutic Practices

 Phase 1: Digestive enzymes that break down mucin/biofilm include Nattokinase, Pectinase, Amylase. Included are proteolytic blend composed of enzymes derived from the edible fungi, Aspergillus oryzae and Aspergillus melleus. There are several manufacturers that have combination products.

Current Therapeutic Practices

- Phase 2: NaEDTA (ethylenediaminetetraacetic acid) which is purported to breakdown the ionic matrix that holds the biofilm together.
- · Phase 3: the antibiotics, antiparasitics, antifungals.
- · Timing: may take several months

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- Appetite improves; trying new things
- · Just seems "happier in the skin he/she is in"
- Health improves · Sleep improves

Sile Melatonin: 1-3mg before bedtime (Start with Iower dose, also useful in chronic pain) 5-HTP: 100mg 5- burtcovtrontonhan before	ep • Vitamin D3 1000 IU and up • GABA 125mg or Gabapentin • Cortisol replacement • Vitamin B6/niacinamide: 50mg B6 and 500 mg niacinamide
har soys group and the special y useful when taken with vitamin B6 and magnesium Calcium/magnesium: can really be helpful	 Pantothenic acid (Vitamin B5): 50 mg daily. (good for relieving stress) Inositol: 100mg daily (Enhances REM sleep) Herbs: milk thistle, valerian, chamomile tea

Decreased Melatonin Activity

- Subgroup of children with very poor sleep cycles in autism
- Biol Psychiatry 1990 Nov 1;28(9):773-93. A novel biochemical model linking dysfunctions in brain melatonin, proopiomelanocortin peptides, and serotonin in autism. Chamberlain RS, Herman BH

Sleep

- Benadryl dye free/compounded 1-2mg/kg
 Ibuprofen 10mg/kg
 Clonidine 0.1mg: ¼ -2 tabs at bedtime
- Trazadone: 0.75mg-1mg/kg at bedtime
- Risperidone: 0.25mg-1mg at bedtime
- Buspirone: 2.5mg -5 mg divided twice daily
 Synthesized Tetrahydrocannibinol 2.5mg 1-3 times
- per day
- Clonazepam: 0.01-0.03 mg/kg/d divided three times dailyHBOT
- Neurontin

Dietary Recommendations

- Removal of dairy: that's anything that comes from a cow's udder! No casein, whey, etc
- · Removal of Gluten is next tacanow.org gfcfdiet.com
- Selective removal of specific foods your child may be sensitive to
- True addictions may make these transitions difficult!

Diets

- There are many, many different diets
- The best diet: the one that works for your child
- Laboratory investigations can point direction
- Trial and error may be the only way to find out what
- really works for your child As organic as possible, cook from "scratch"
- Medically prescribed diets are a tax deduction!!





Eicosanoids as Inflammatory Mediators the Sugar Link to Mood and Bad Behaviors.

- Generally underemphasized influence, over the inflammatory response in your child, is the eicosanoid pathway.
- the consumption of certain fats and/or sugar (starch/carbohydrates) influences the production of these inflammatory mediators (Diet Link) Vascular effects of eicosanoids, are likely part of the
- observed perfusion (blood flow) abnormalities in autism

Starting with Nutritional Supplements

- Vitamins and Minerals: need replenishment due to usual self limited diet
- Digestive Enzymes: well known and well described deficiency
- Omega 3's: crucial to start these
- Soluble Fiber: very important for bowel health

Vitamins

 Vitamins: Vitamins are micronutrients that are critical for normal cellular metabolism and function. They must be consumed regularly as our bodies are not able to synthesize these vital nutrients in sufficient amounts. Vitamins are required for proper cell signaling, mediators of cell growth, as antioxidants, and even as hormones

Minerals

 Minerals: The main electrolytes are Sodium, Potassium, Calcium, Magnesium and Chloride. Proper levels of these minerals in the body allow the body to have energy, better muscle and nerve function (including faster stress recovery) and improved function in many other areas. The so called trace minerals (such as selenium, iodine, chromium, magnesium, molybdenum, just to name a few) need to be replenished daily, ideally through the diet.

We have to address the Vitamins and
Minerals

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Poracolam	n	43- 95					
Phosphores	595	482- 745			-		
Copper	0.7	0.52- 0.69				-	
Zinc	3.3	8- 14-5	_		_	•	
irce	754	745- 1050		-			
Maganese	0.015	0.837- 0.03		1995	-		
Ororian	0.021	0.812- 0.87		-			
Selenium	0.15	0.19- 0.38		-	_		
Boros	0.09	0.885- 0.21		1255	-	_	
Vanadiam	0.0611	.0001-0.002			-		
Molabdenan	0.0009	.6035-0.002					

Digestive Enzymes

 Digestive Enzymes break down the foods that we eat into smaller compounds that can be readily absorbed and put into the bloodstream. The enzyme class of proteases breaks down proteins to amino acids. The enzyme class of lipase is responsible of breaking down fats to fatty acids, and the carbohydrases break down carbohydrates into simple sugars.

Omega 3's

- Omega 3 : the fatty acids, maintain the fluidity and stability of cell membranes and protect the body from harmful substances such as bacteria and viruses. They must be obtained through diet or supplementation with EFA-rich food oils.
- They have an amazing affect on behaviors, mood, focus and concentration.



Ca/Mg deficiency:

 Typically recommend supplementing 500-1000 mg of Calcium in Citrate, glycinate/malate complexes with at least 200-400mg of Magnesium (fully reactive chelate or ionic) daily. Calcium and magnesium blood levels should be checked regularly.

Zinc

- Consider supplementing with 20 mg (or more) of zinc daily. Some children require much higher doses.
- Some sources of Zinc may be better absorbed than others
- · Zinc Picolinate and Zinc Monomethionine are very good sources

Selenium

• Many children with ASD require additional selenium. We will start with 25-100 mcg daily. Packed red blood cell element evaluations may reveal higher levels of supplementation are necessary.

Chromium

• If this is revealed to be low on blood level tests, we recommend about 50 to 100 mcg daily.

Iron Deficiency

This can be easily evaluated by simple blood tests: a CBC, RETICULOCYTE COUNT, SERUM FERRITIN, AND A TOTAL IRON BINDING CAPACITY

lodine

- There is a move away form iodized salt in fast food and natural products.
- · Natural products often use sea salt (relatively
- low in iodine).
- Necessary for Thyroid function
- Lugol's 5% solution has 6.26mg per drop
- There is a lot of controversy about the optimal amount of lodine to recommend



Essential Fatty Acid Deficiency

- Omega 3 Fatty Acids in Particular (EPA) Eicosapentaenoic Acid and (DHA) Docosahexanoic Acid

 - (Cr.) Ecosperation Crick and Criny Docosahexanoic Acid Docosahexanoic Acid Prostaglandins Leukot Essent Fatty Acids 2001 Jul:65(1):-7. Plasma fatty acid levels in autistic children. Vancassel S, et al Prostaglandins Leukot Essent Fatty Acids 2000 Jul-Augr/3(1-2):21-5. Red blood cell fatty acid compositions in a patient with autistic spectrum disorder: a characteristic abnormality in neurodevelopmental disorders? Bell JG,

EFA deficiency:

- Just about all the children will require omega 3 fatty acids
- Start with Cod Liver Oil
- Children ages 2-5: ½ tsp of cod liver oil per day
- Older children: 1 tsp daily
- · A blood test, called an Essential Fatty Acid
- Profile will reveal other possible fatty acid deficits that can be supplemented.

Failure to Thrive



- Same weight for a long growth period Age of bone well behind
- chronological age • Thin, pale, irritable
- Height is well below 5th percentile or is falling of . graph

Causes of Failure to Thrive

- Stress
- · Poor caloric intake
- Poor digestion
- Poor absorption
- · Gut motility issues
- Chronic inflammation
- Many other issues

Stress and Growth Hormone

- Excessive Corticotropin-Releasing Hormone (CRH) from chronic stress.
- · Possible excess cortisol.
- Also reduces the body's secretion of Growth Hormone

Protein deficiency:

- RICE PROTEIN POWDER:
- 1 tablespoon is equal to 14 gms of protein. If under 3 years, supplement with 1 gm per 2 pounds of body weight.
- Ages 4-6 years, 1 tablespoon daily.
- 7-10 years of age supplement with 1 ¾ tablespoons daily.
- Older boys and girls: 2 tablespoon daily.
- Pea Protein is another great option

Consequences of Abnormal Physiologic Functions:

 Endocrine: abnormal feedback controls of the thyroid, adrenal, pituitary glands, and many other glandular functions

Thyroid Symptoms

- Lethargy
- Abnormal weight gain
- Constipation
- Mental slowness
- Dry skin
- Thin hair
- · Poor growth parameters

Thyroid

- · Labs: T3 Free, T4 Free, TSH, reverse T3
- Many aberrations found with feedback control
 Ideally, TSH is considered normal if falls within 1.0 and
- 2.0
- Insure daily amount of lodine: Children Tolerable Upper Intake Levels (UL): 200mcg/day for ages 1-3 years; 300mcg/day for 4-8 years; 600mcg/day for 9-13 years; 900mcg/day for 14-18 years
- Use compounded porcine thyroid for replacement/augmentation as indicated Remove sources of fluoride, chlorine



Methylation plays a role in:

- Alcoholism
- Bipolar disease
- Schizophrenia
- Depression
- Attention deficit disorder Autism
- And even constipation!

Methylation

- Methyl B 12 appears to work best when given sub-cutaneously, by the parents (this is a prescription). We tend to add folic acid and N- acetyl cysteine to the injection.
- TMG or DMG
- Folic/folinic acid
- Glutathione transdermally twice daily is prescribed as well





Back to the LAB: Pediatric Auto-immune Neuropsychiatric Disorder Associated with Streptococcus

PANDAS profile: . Anti-streptolysin O titers (ASO)
 Anti-DNAse antibodies

Keep in mind that PANDAS is a Clinical Diagnosis

Lab Results

- Elevated ASO titers and Anti DNAse antibodies
- Antibiotics for PANDAS:
 - Augmentin Zithromax
 - Keflex
 - Omnicef

Inflammation Disease States Laboratory Findings Allergies Inflammatory Bowel Inflammatory markers: elevated platelets, elevated erythrocyte sedimentation rate, elevated C-reactive Claritin Redi-Tabs Antacids Allergies environmental sources • Zyrtec Nexium Prilosec food sources - Pepcid Reflux disease

- protein Endoscopy/pathology

.

Small bowel disease

Large bowel disease

Low grade bacterial

infections

Chronic viral activation

- findings
- Viral titers: HHV 1,2,6, and CMV, EBV
- Stool cultures, anaerobic markers

Inflammation

Anti-inflammatories

Colazal
Pentasa

- Singulair
- Nasonex
- Gastrocrom
- Quercetin
- Provocation-Neutralization-
- Desensitization LDA
- Rowasa
 Steroids 6 mercapto purine

Inflammation

- Infections Anti-inflammatory Antibiotics Motrin/Advil Antifungals
 - Celebrex Naprosyn
- Antiparasitics Antivirals
- Steroids
- LongVida/Neuroprotek Natural agents
 - Curcumin
 Omega oils
 - Pycnogenols

Actos/Avandia

Language

- L Carnosine
- L Carnotine
- Namenda
- GABA or Gabapentin
- Steroids
- Gingko Biloba
- Anti-Seizure Medication
- HBOT
- IV Glutathione

Cognitive Function

- L-Carnosine 400 mg
- Coenzyme Q-10 up to 600 mg
 Gingko biloba leaf extract: 50 200mg

- Glingko biloba teal extract: 30 200mg
 American ginseng root extract (*Panax quinquefolium*): 200 mg q day
 DMAE (as dimethylaminoethanol bitartrate) 200 500 mg
- L-Theanine 25 200 mg
- Acetyl L-Carnitine (from acetyl L-carnitine HCI) 300 1200 mg

The Recurrently III Child

- Frequent ear infections
- Frequent sinus infections
- Frequent upper and lower respiratory infections
- Frequent fevers

The Recurrently III Child Laboratory Approach

- CBC with Diff
- Comprehensive Metabolic Profile
- Erythrocyte Sedimentation Rate
- C-Reactive Protein
- Quantitative Immunoglobulins: IgG, IgM, IgE, IgA
- Lymphocyte Enumeration with Natural Killer Cell activity

The Recurrently III Child The Treatments



- Isoprinosine (not available in US) which is a TH1 stimulator
- Oral Immunoglobulins
- Syndromes such as Kartagener's/immotile celia syndrome
- Bone Scan
- Anti-Virals



