## Table A: Summary Comparison of Characteristics of Autism & Mercury Poisoning

Mercury Poisoning	Autism
Psychiatric Disturbances	
Social deficits, shyness, social withdrawal	Social deficits, social withdrawal, shyness
Depression, mood swings; mask face	Depressive traits, mood swings; flat affect
Anxiety	Anxiety
Schizoid tendencies, OCD traits	Schizophrenic & OCD traits; repetitiveness
Lacks eye contact, hesitant to engage others	Lack of eye contact, avoids conversation
Irrational fears	Irrational fears
Irritability, aggression, temper tantrums	Irritability, aggression, temper tantrums
Impaired face recognition	Impaired face recognition
Speech, Language & Hearing Deficits	
Loss of speech, failure to develop speech	Delayed language, failure to develop speech
Dysarthria; articulation problems	Dysarthria; articulation problems
Speech comprehension deficits	Speech comprehension deficits
Verbalizing & word retrieval problems	Echolalia; word use & pragmatic errors
Sound sensitivity	Sound sensitivity
Hearing loss; deafness in very high doses	Mild to profound hearing loss
Poor performance on language IQ tests	Poor performance on verbal IQ tests
Sensory Abnormalities	
Abnormal sensation in mouth & extremities	Abnormal sensation in mouth & extremities
Sound sensitivity	Sound sensitivity
Abnormal touch sensations; touch aversion	Abnormal touch sensations; touch aversion
Vestibular abnormalities	Vestibular abnormalities
Motor Disorders	
Involuntary jerking movements - arm flapping, ankle jerks, myoclonal jerks, choreiform movements, circling, rocking	Stereotyped movements - arm flapping, jumping, circling, spinning, rocking; myoclonal jerks; choreiform movements
Deficits in eye-hand coordination; limb apraxia; intention tremors	Poor eye-hand coordination; limb apraxia; problems with intentional movements

Gait impairment; ataxia - from incoordination & clumsiness to inability to walk, stand, or sit; loss of motor control	Abnormal gait and posture, clumsiness and incoordination; difficulties sitting, lying, crawling, and walking
Difficulty in chewing or swallowing	Difficulty chewing or swallowing
Unusual postures; toe walking	Unusual postures; toe walking
Cognitive Impairments	
Borderline intelligence, mental retardation - some cases reversible	Borderline intelligence, mental retardation - sometimes "recovered"
Poor concentration, attention, response inhibition	Poor concentration, attention, shifting attention
Uneven performance on IQ subtests	Uneven performance on IQ subtests
Verbal IQ higher than performance IQ	Verbal IQ higher than performance IQ
Poor short term, verbal, & auditory memory	Poor short term, auditory & verbal memory
Poor visual and perceptual motor skills, impairment in simple reaction time	Poor visual and perceptual motor skills, lower performance on timed tests
Difficulty carrying out complex commands	Difficulty carrying out multiple commands
Word-comprehension difficulties	Word-comprehension difficulties
Deficits in understanding abstract ideas & symbolism; degeneration of higher mental powers	Deficits in abstract thinking & symbolism, understanding other's mental states, sequencing, planning & organizing
Unusual Behaviors	
Stereotyped sniffing (rats)	Stereotyped, repetitive behaviors
ADHD traits	ADHD traits
Agitation, unprovoked crying, grimacing, staring spells	Agitation, unprovoked crying, grimacing, staring spells
Sleep difficulties	Sleep difficulties
Eating disorders, feeding problems	Eating disorders, feeding problems
Self injurious behavior, e.g. head banging	Self injurious behavior, e.g. head banging
Visual Impairments	
Poor eye contact, impaired visual fixation	Poor eye contact, problems in joint attention
"Visual impairments," blindness, near-sightedness, decreased visual acuity	"Visual impairments"; inaccurate/slow saccades; decreased rod functioning
Light sensitivity, photophobia	Over-sensitivity to light
Blurred or hazy vision	Blurred vision
Constricted visual fields	Not described
Physical Disturbances	

Increase in cerebral palsy; hyper- or hypo-tonia; abnormal reflexes; decreased muscle strength, especially upper body; incontinence; problems chewing, swallowing, salivating	Increase in cerebral palsy; hyper- or hypotonia; decreased muscle strength, especially upper body; incontinence; problems chewing and swallowing
Rashes, dermatitis/dry skin, itching; burning	Rashes, dermatitis, eczema, itching
Autonomic disturbance: excessive sweating, poor circulation, elevated heart rate	Autonomic disturbance: unusual sweating, poor circulation, elevated heart rate
Gastro-intestinal Disturbances	
Gastroenteritis, diarrhea; abdominal pain, constipation, "colitis"	Diarrhea, constipation, gaseousness, abdominal discomfort, colitis
Anorexia, weight loss, nausea, poor appetite	Anorexia; feeding problems/vomiting
Lesions of ileum & colon; increased gut permeability	Leaky gut syndrome
Inhibits dipeptidyl peptidase IV, which cleaves casomorphin	Inadequate endopeptidase enzymes needed for breakdown of casein & gluten
Abnormal Biochemistry	
Binds -SH groups; blocks sulfate transporter in intestines, kidneys	Low sulfate levels
Has special affinity for purines & pyrimidines	Purine & pyrimidine metabolism errors lead to autistic features
Reduces availability of glutathione, needed in neurons, cells & liver to detoxify heavy metals	Low levels of glutathione; decreased ability of liver to detoxify heavy metals
Causes significant reduction in glutathione peroxidase and glutathione reductase	Abnormal glutathione peroxidase activities in erythrocytes
Disrupts mitochondrial activities, especially in brain	Mitochondrial dysfunction, especially in brain
Immune Dysfunction	
Sensitivity due to allergic or autoimmune reactions; sensitive individuals more likely to have allergies, asthma, autoimmune-like symptoms, especially rheumatoid-like ones	More likely to have allergies and asthma; familial presence of autoimmune diseases, especially rheumatoid arthritis; IgA deficiencies
Can produce an immune response in CNS	On-going immune response in CNS
Causes brain/MBP autoantibodies	Brain/MBP autoantibodies present
Causes overproduction of Th2 subset; kills/inhibits lymphocytes, T-cells, and monocytes; decreases NK T-cell activity; induces or suppresses IFNg & IL-2	Skewed immune-cell subset in the Th2 direction; decreased responses to T-cell mitogens; reduced NK T-cell function; increased IFNg & IL-12
CNS Structural Pathology	

Selectively targets brain areas unable to detoxify or reduce Hg-induced oxidative stress	Specific areas of brain pathology; many functions spared	
Damage to Purkinje and granular cells	Damage to Purkinje and granular cells	
Accummulates in amygdala and hippocampus	Pathology in amygdala and hippocampus	
Causes abnormal neuronal cytoarchitecture; disrupts neuronal migration & cell division; reduces NCAMs	Neuronal disorganization; increased neuronal cell replication, increased glial cells; depressed expression of NCAMs	
Progressive microcephaly	Progressive microcephaly and macrocephaly	
Brain stem defects in some cases	Brain stem defects in some cases	
Abnormalities in Neuro-chemistry		
Prevents presynaptic serotonin release & inhibits serotonin transport; causes calcium disruptions	Decreased serotonin synthesis in children; abnormal calcium metabolism	
Alters dopamine systems; peroxidine deficiency in rats resembles mercurialism in humans	Possibly high or low dopamine levels; positive response to peroxidine (lowers dopamine levels)	
Elevates epinephrine & norepinephrine levels by blocking enzyme that degrades epinephrine	Elevated norepinephrine and epinephrine	
Elevates glutamate	Elevated glutamate and aspartate	
Leads to cortical acetylcholine deficiency; increases muscarinic receptor density in hippocampus & cerebellum	Cortical acetylcholine deficiency; reduced muscarinic receptor binding in hippocampus	
Causes demyelinating neuropathy	Demyelination in brain	
EEG Abnormalities / Epilepsy		
Causes abnormal EEGs, epileptiform activity	Abnormal EEGs, epileptiform activity	
Causes seizures, convulsions	Seizures; epilepsy	
Causes subtle, low amplitude seizure activity	Subtle, low amplitude seizure activities	
Population Characteristics		
Effects more males than females	Male:female ratio estimated at 4:1	
At low doses, only affects those geneticially susceptible	High heritability - concordance for MZ twins is 90%	
First added to childhood vaccines in 1930s	First "discovered" among children born in 1930s	
Exposure levels steadily increased since 1930s with rate of vaccination, number of vaccines	Prevalence of autism has steadily increased from 1 in 2000 (pre1970) to 1 in 500 (early 1990s), higher in 2000.	
Exposure occurs at 0 - 15 months; clinical silent stage means symptom emergence delayed;	Symptoms emerge from 4 months to 2 years old; symptoms emerge gradually, starting with movement & sensation	

symptoms emerge gradually, starting with	
movement & sensation	

## Table II: Summary of Psychiatric Disturbances Found in Autism & Mercury Poisoning

Mercury Poisoning	Autism
Extreme shyness, social withdrawal, feeling overly sensitive, introversion	Social deficits, social withdrawal, self reports of extreme shyness, aloofness
Mood swings; flat affect; mask face; laughing or crying without provocation; episodes of hysteria	Mood swings; flat affect in some; no facial expression; laughing or crying without reason
Anxiety; nervousness; tremulousness; somatization of anxious feelings	Anxiety, nervousness; anxiety disorder
Schizoid tendencies, neurosis, obsessive-compulsive traits, repetitive dreams	Schizophrenic traits; OCD traits; repetitive behaviors and thoughts
Lack of eye contact; being less talkative; hesitancy to engage others	Lack of eye contact, gaze avoidance; avoids conversation
Depression, lack of interest in life, lassitude, fatigue, apathy; feelings of hopelessness; melancholy	Association with depression; lack of initiative, diminished outward emotions
On the one hand, less overtly active, unwilling to go outside or be with others; on the other hand, increased restlessness	Tendency to withdraw, especially to own rooms, prefer to be alone; hyperactivity
Irrational fears	Irrational fears
Irritability, anger, and aggression; in children this may manifest as frequent and severe temper tantrums	Irritability and aggression; severe temper tantrums in children
Psychotic episodes; hallucinations, hearing voices; paranoid thoughts	Psychotic talk, paranoid thoughts
Impaired face recognition	Impaired face recognition

# Table III: Summary of Speech, Language & Hearing Deficits in Autism & Mercury Poisoning Mercury Poisoning Autism

Complete loss of speech in adults or children; failure to develop speech in infants	Delayed language onset; failure to develop speech
Dysarthria; speech difficulties from intention tremor; slow and slurred speech	Dysarthria; dyspraxia and oral-motor planning difficulties; unintelligible speech

Aphasia, the inability to use or understand words, inability to comprehend speech although ability to hear sound is intact

Difficulties verbalizing; word retrieval problems

Auditory disturbance; difficulties differentiating voices in a crowd
Sound sensitivity

Hearing loss; deafness in very high doses

Poor performance on standardized language tests

Speech comprehension deficits, although ability to hear sound is intact

Echolalia; pronoun reversals, word meaning and pragmatic errors; limited speech production

Difficulties following conversational speech with background noise

Sound sensitivity

Mild to profound hearing loss

Poor performance on verbal IQ tests

Table IV: Summary of Sensory Abnormalities in Mercury Poisoning & Autism

Mercury Poisoning	Autism
Abnormal sensation or numbness around mouth and extremities (paresthesia); burning feet	Abnormal sensation in mouth and extremities; excessive mouthing of objects (infants); toe walking; difficulty grasping objects
Sound sensitivity	Sound sensitivity
Excessive pain when bumping; abnormal touch sensations; touch aversion	Insensitivity or overreaction to pain and touch; touch aversion; stiff to hold
Loss of position in space	Vestibular system abnormalities; difficulty orienting self in space
Normal pinprick tests	Normal pinprick tests

## Table V: Summary of Motor Disorder Behaviors in Mercury Poisoning & Autism

### **Mercury Poisoning**

Involuntary jerking movements, e.g., arm flapping, ankle jerks, myoclonal jerks; choreiform movements; circling (cats); rocking; purposeless movement of extremities; twitching, shaking; muscular spasticity

Unsteadiness in handwriting or an inability to hold a pen; deficits in eye-hand coordination; limb apraxia; intention tremors; loss of fine motor skills

Ataxia: gait impairment; severity ranging from mild incoordination, clumsiness to complete inability to walk, stand, or sit; staggering, stumbling; loss of motor control

#### Autism

Stereotyped movements such as arm flapping, jumping, circling, spinning, rocking; myoclonal jerks; choreiform movements

Difficulty in writing with or holding a pen; poor eye-hand coordination; limb apraxia; problems carrying out intentional movements (praxia)

Abnormal gait and posture, clumsiness and incoordination; difficulties sitting, lying, crawling, and walking in infants and toddlers Toe walking

Difficulty in chewing or swallowing

Unusual postures

Areflexia

Tremors in general, tremors of the face and tongue,

hand tremors

Toe walking

Difficulty chewing or swallowing

Unusual postures

None described

None described

## Table VI: Summary of Areas of Mental Impairment in Mercury Poisoning & Autism

#### **Mercury Poisoning**

Some aspect of mental impairment in all symptomatic cases

Borderline intelligence on testing among previously normal individuals; mental retardation occurring in severe cases of pre-/postnatal exposure; some cases of MR reversible; primate studies indicate core intelligence spared with low exposures

Uneven performance on subtests of intelligence

Verbal IQ higher than performance IQ; compromised language/verbal expression and comprehension

Poor concentration, shortened attention span, general lack of attention; poor response inhibition

Forgetfulness, loss of memory, particularly short term, verbal and auditory memory; mental confusion

Poor visual and perceptual motor skills, poor eye-hand coordination; impairment in simple reaction time

Not reported as being tested

Difficulty carrying out complex commands

Alexia (inability to comprehend the meaning of written words)

Deficits in constructional skills, conceptual abstraction, understanding abstract ideas and symbolism; degeneration of higher mental powers

Lack of understanding of object permanence (primates)

**Autism** 

Some aspect of mental impairment in all cases

Borderline intelligence or mental retardation on standard tests among previously normally appearing infants; some cases of MR "reversible"; indications that normal IQ might be present in MR-labeled individuals

Uneven performance on subtests of intelligence

Verbal IQ higher than performance IQ; compromised language/verbal expression and comprehension

Lack of concentration, short attention span, lack of attention, difficulty shifting attention

Poor short term/working memory; poor auditory and verbal memory; lower verbal encoding abilities

Poor visual and perceptual motor skills, poor eye-hand coordination; lowered performance on timed tests

Difficulty processing multiple stimuli

Difficulty carrying out multiple commands

Hyperlexia (ability to decode words while lacking word comprehension)

Deficits in abstract/conceptual thinking, symbolism, understanding other's mental states; impairment in sequencing, planning, organizing

Deficient understanding of object permanence (children)

## **Table VII: Summary of Unusual Behaviors** in Mercury-Poisoned Animals and Humans & in Autism

**Mercury Poisoning** 

Autism

Stereotyped sniffing (rats) Stereotyped, repetitive behaviors

Hyperactivity (rats); poor response inhibition Hyperactivity; ADHD-traits

(humans), restlessness

Agitation (humans) Agitation

Insomnia; difficulty falling or staying Insomnia; difficulty falling asleep (humans) asleep

Eating disorders: anorexia, poor appetite, food Eating disorders: anorexia; restricted aversion, narrow food preferences, decided food diet/narrow food preferences; feeding and

preferences (salty food) (humans) suckling problems Masturbation, priapism (children) Masturbatory tendencies

Unintelligible cries; continuous crying; unprovoked Unprovoked crying

crying (infants and children)

Self injurious behavior, including head banging and Self injurious behavior, including head

hitting the head (toddlers and children) banging and hitting the head

Grimacing Grimacing (children) Staring spells (infants and children) Staring spells

## **Table VIII: Summary of Visual Impairments** Seen in Mercury Poisoning & Autism

#### **Mercury Poisoning**

**Autism** 

Lack of eye contact; difficulties with Lack of eye contact; gaze abnormalities; problems in visual fixation

joint attention "Visual impairments," blindness, near-"Visual impairments"; inaccurate or slow saccades;

sightedness, decreased visual acuity decreased functioning of the rods; retinal sheen

Light sensitivity, photophobia Over-sensitivity to light

Blurred or hazy vision Blurred vision Constricted visual fields Not described

## **Table IX: Physical Disturbances** in Mercury Poisoning & Autism

### **Mercury Poisoning**

#### **Autism**

Increase in cerebral palsy; hyper- or hypotonia; Increase in cerebral palsy; hyper- or paralysis, abnormal reflexes; spasticity; hypotonia; decreased muscle strength, decreased muscle strength and motor power, especially in the upper body; incontinence; problems chewing, swallowing, and salivating Rashes, dermatitis, dry skin, itching; burning sensation

especially in the upper body; incontinence/toilet training difficulties; problems chewing and swallowing

Rashes, dermatitis, eczema; itching

Autonomic disturbances: excessive sweating; poor circulation; elevated heart rate

Autonomic disturbances: sweating abnormalities; poor circulation; elevated heart

## **Table X: Summary of Gastrointestinal Problems** in Mercury Poisoning & Autism

#### **Mercury Poisoning**

Gastroenteritis, diarrhea; abdominal pain, rectal itching, constipation, "colitis"

Anorexia, weight loss, nausea, poor appetite

Lesions of the ileum and colon; increased intestinal permeability

Inhibits dipeptidyl peptidase IV, which cleaves casomorphin

#### **Autism**

Diarrhea, constipation, gaseousness, abdominal discomfort, colitis

Anorexia; feeding difficulties, vomiting as

infants

Leaky gut syndrome from sulfur deficiency

Inadequate endopeptidase enzymes responsible for breakdown of casein and gluten

## Table XI: Abnormalities in Biochemistry **Arising from Hg Exposure & Present in Autism**

#### **Mercury Poisoning**

Ties up sulfur groups; prevents sulfate absorption

Has special affinity for purines and pyrimidines

Depletes cellular tyrosine in yeast

Reduces bioavailability of glutathione, necessary in Low levels of glutathione; decreased cells and liver for heavy metal detoxification Can cause significant reduction in glutathione peroxidase and glutathione reductase

Disrupts mitochondrial activities, especially in brain

### Low sulfate levels

Errors in purine and pyrimidine metabolism can lead to autistic features PKU, arising from disruption in tyrosine production, results in autism

**Autism** 

ability of liver to detoxify heavy metals Abnormal glutathione peroxidase activities in erythrocytes

Mitochondrial dysfunction, especially in

## Table XII: Summary of Immune System Abnormalities in Mercury Exposure & Autism

## **Mercury Poisoning**

### Autism

Individual sensitivity due to allergic or autoimmune More likely to have allergies and asthma; reactions; sensitive individuals more likely to have allergies and asthma, autoimmune-like symptoms, especially rheumatoid-like ones

Can produce an immune response, even at low levels; can remain in CNS for years

Presence of autoantibodies (IgG) to neuronal cytoskeletal proteins, neurofilaments, and myelin basic protein; astrogliosis; transient ANA and AnolA

familial presence of autoimmune diseases, especially rheumatoid arthritis; IgA deficiencies

Indications of on-going immune response in CNS

Presence of autoantibodies (IgG and IgM) to cerebellar cells, myelin basis protein

Causes overproduction of Th2 subset; diminishes capacity to produce TNF(alpha) and IL-1; kills lymphocytes, T-cells, and monocytes; inhibits lymphocyte production; decreases NK T-cell activity; may induce or suppress IFN(gamma) and IL-2 production

Skewed immune-cell subset in the Th2 direction and abnormal CD4/CD8 ratios; decreased responses to T-cell mitogens; increased neopterin; reduced NK T-cell function; increased IFN(gamma) and IL-12

## Table XIII: CNS Lesions in Mercury Poisoning & Autism

#### **Mercury Poisoning**

## Primarily impacts CNS

Selectively targets brain areas - those unable to detoxify heavy metals or reduce Hg-induced oxidative stress

Damage to Purkinje and granular cells

Accumulates in amygdala and hippocampus

Causes abnormal neuronal cytoarchitecture; interferes with neuronal migration and depresses cell division in developing brains; reduces NCAMs

Head size differences: progressive microcephaly

Brain stem defects in some cases

#### Autism

Neurological impairments primary

Specific areas of brain pathology; many functions spared

Damage to Purkinje and granular cells Pathology in amygdala and hippocampus Neuronal disorganization; increased neuronal cell replication, small glia to neuron ration, increased glial cells; depressed expression of NCAMs

Head size differences: progressive microcephaly and macrocephaly Brain stem defects in some cases

## Table XIV: Abnormalities in Neurons & Neurochemicals from Mercury & in Autism

#### **Mercury Poisoning**

Can increase tissue concentration of serotonin in newborn rats; causes calcium disruptions in neurons, preventing presynaptic serotonin release and inhibiting serotonin transport activities

Alters dopamine systems; disrupts calcium and increases synaptosome membrane permeability, which affect dopamine activities; peroxidine deficiency in rats results in acrodynia

Increases epinephrine and norepinephrine levels by blocking the enzyme which degrades epinephrine

Elevates glutamate; decreases glutamate uptake; reduces functional activity of glutamatergic system Alters choline acetyltransferase, leading to

Alters choline acetyltransferase, leading to acetylcholine deficiency; inhibits acetylcholine neurotransmitter release via impact on calcium

#### Autism

Serotonin abnormalities: decreased serotonin synthesis in children; oversynthesis in adults; elevated serotonin in platelets; positive response to SSRIs; calcium metabolism abnormalities present Indications of either high or low dopamine levels; positive response to peroxidine by lowering dopamine levels; positive response to dopamine antagonists Elevated norepinephrine and epinephrine; positive response to norepinephrine reuptake inhibitors

Elevated glutamate and aspartate

Abnormalities in cholinergic neurotransmitter system: cortical acetylcholine deficiency and reduced homeostasis; causes cortical acetylcholine

deficiency; increases muscarinic receptor density in

hippocampus and cerebellum

Causes demyelating neuropathy

muscarinic receptor binding in

hippocampus

Demyelation in brain

## **Table XV: EEG Activity & Epilepsy** in Mercury Poisoning & Autism

## **Mercury Poisoning**

Causes abnormal EEGs and unusual epileptiform

activity

Causes seizures, convulsions

Causes subtle, low amplitude seizure activity

Abnormal EEG activity; epileptiform

Autism

activity

Seizures; epilepsy

Subtle, low amplitude seizure activities