Filming Session 3	
Topics Functional Biochemistry  Toxicity Chemicals	
Principles of Functional Biochemistry	



Enzymes are protein catalysts that regulate the rates at which physiological processes take place. They are encoded by specific genes which in turn are stimulated by hormones.

There are 3870 enzymes catalogued in the ENZYME DATABASE.

## There are two types

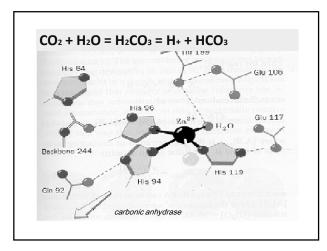
- those that require a coenzyme such as the oxido-reductases.
   of known enzymes require coenzymes to function.
- 2) those that do not require a coenzyme such as the digestive enzymes.

# Four parts

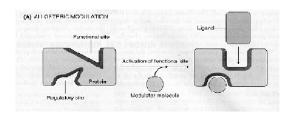
- 1. The apoenzyme is the protein part of an enzyme.
- 2. The coenzyme is required for the activation of an enzyme.

# 3.Metal ion catalysts

- a) Metalloenzymes contain tightly bound metal ions most commonly transition metal ions such as Fe2+, Fe3+, Cu2+, Zn2+,Mn2+ or Co3+.
- b) Metal activated enzymes loosely bind metal ions from solution, usually alkaline earth metal ions Na+, K+, Mg2+ or Ca2+

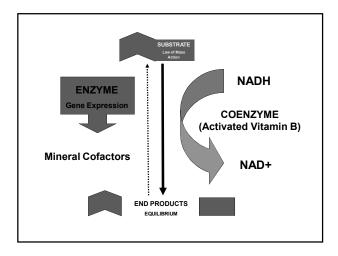


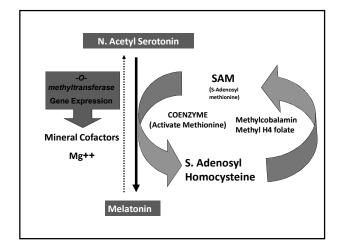
4. Low molecular weight allosteric effectors modulate the catalytic activity of certain regulatory enzymes.



# **Factors affecting enzyme function**

- 1. Temperature
- 2. Enzyme concentrations
- 3. Substrate concentration
- 4. pH
- 5. Inhibitors can poison enzymes e.g. certain chemicals e.g. toiletries, cosmetics, toxic metals and mycotoxins.

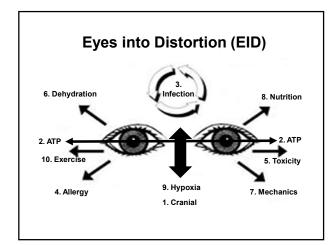




# CHALLENGES FOR ENZYME PATHWAY INHIBITION

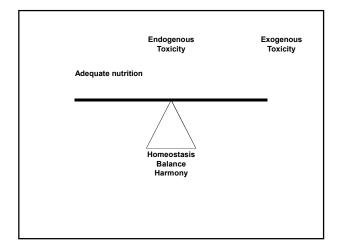
- 1. A WEAK ASSOCIATED MUSCLE WILL STRENGTHEN TO THE REQUIRED END PRODUCT.
- 2. A STRONG INDICATOR MUSCLE WILL WEAKEN WHEN CHALLENGED TO THE SUBSTRATE.
- 3. THIS WEAKNESS WILL BE NEGATED BY THE MINERAL COFACTORS AND / OR THE COENZYME (USUALLY AN ACTIVATED VITAMIN B)

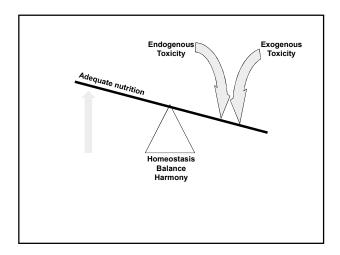
Toxicity





Toxicity at the root of many disorders

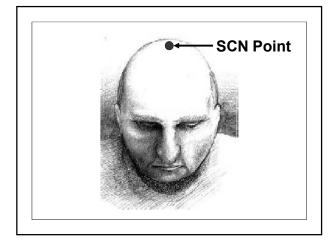




# **Toxicity**

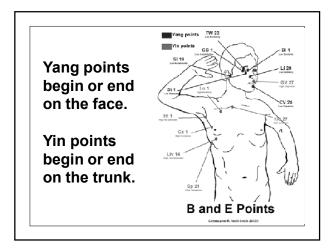
- 1. Internal Chemicals e.g. neurotransmitters, hormones generated by the mind External Chemicals xenobiotics
- 2. Toxic metals
- 3. Radiation e.g. radioactive
- . isotopes, electromagnetic stress

Endogenogous toxins are generated from the mind are mediated by the hypothalamus hormones – Thyrotrophic releasing hormone Corticotrophic releasing hormone Gonadotrophic releasing hormones Growth hormone releasing hormone Somatostatin Prolactin inhibiting hormone 300 Neuropeptides	
External toxins (Xenobiotics) come from  What we eat What we drink What we breath What we put on our skin Electromagnetic pollution Harmful solar wavelengths	
Identifying the priority meridian imbalance	



**Identifying the Definitive Meridian** 

- 1. Therapy localise the SCN point (should remain strong)
- 2. Cross Therapy localise to each meridian B&E point
- 3. Only one will weaken. This is the *definitive* meridian
- 4. You can confirm with the respective neurotransmitter vial
- 5. Identify weak associated muscle



Yang points indicate neurotransmitter deficiencies.  Yin points indicate neurotransmitter excesses  B and E Points  Chicaparell' And I and B 130	
Detoxification	
Detoxification is a generic term for the metabolism (catabolism) of both endogenous and exogenous chemicals.	

The main endogenous chemicals to be metabolised are	
<ul> <li>Neurotransmitters</li> </ul>	
• Hormones	
• Eicosonoids	
Certain Fatty acids	
Retinoids	
	1
The main everences chemicals	
The main exogenous chemicals (xenobiotics) to be metabolised	
are either	
Water soluble	
• Lipid soluble	
	1
l inid caluble abomicale are	
Lipid soluble chemicals are generally metabolised by	
1. Hydroxylation to make them	
more water soluble	
ОН	
OH	
	1

	_
Lipid soluble chemicals are generally metabolised by	
Hydroxylation to make them more water soluble	
2. Conjugation to aid their elimination through the kidneys	
or biliary system.	
Manage harmon and an anadem data d	
Many hormones are methylated after hydroxylation before they are conjugated.	
are conjugated.	
СH <sub>3</sub>	
СНз	
СНз	
СНз	
СH <sub>3</sub>	
CH <sub>3</sub>	
Probably 90% of detoxification involves the metabolism of the	
Probably 90% of detoxification	
Probably 90% of detoxification involves the metabolism of the endogenously produced	

Like Neurotransmitters and Hormones	
However to fully understand endogenous detoxification its easier to start by learning about exogenous detoxification.	
Those of medical relevance are	

	-
1. Drugs 2. Chemical carcinogens 3. Pesticides and other various compounds.	
More than 75,000 synthetic chemicals now exist.  Most will require detoxification, with the liver being the main organ involved.  Occasionally a xenobiotic maybe excreted unchanged.	
It is convenient to consider the metabolism of xenobiotics in two phases.  1. Phase 1 hydroxylation catalyzed by the mono-oxygenases cytochrome P450's.	
2. Phase 2 Methylation or Conjugation.	

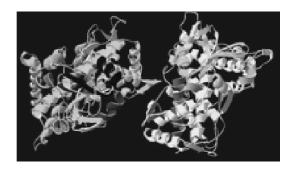
The overall purpose of the two
phases is to increase their water
solubility (polarity) and thus
facilitate their excretion from the
body.

Very hydrophobic xenobiotics would persist in adipose tissue indefinitely if they were not converted to more polar forms.

In certain cases, Phase 1 metabolic reactions convert xenobiotics from inactive to biologically active compounds.

In some instances the original xenobiotics are pro-carcinogens which then become converted to carcinogens by the phase 1 hydroxylation.

## **Phase 1 Hydroxylation**



There are 14 families of the Cytochrome P450 enzyme	
encoding for between 35-60 distinct P450 enzymes.	
They all use the abbreviated root	
symbol CYP.	
This is followed by a number	
designating the family having similar sequence identity.	
This is followed by a capital letter	
indicating the subfamily.	
Lastly this is followed by another number indicating the individual	
P450's in the family.	
	1
Examples:-	
CYP1A1 metabolises PAH's 2-estrogens	
CYPIA2 metabolises 16-estrogens	
CYPIB1 metabolises 4-estrogens and synthetic estrogens	
CYP2A6 metabolises nicotine CYP2B4 metabolises	
phenobarbital	

CYP2C9 metabolises warfarin

CYP2C19 and CYP2D6 most antidepressants and antipsychotics

CYP2E1 metabolises ethanol, solvents and components in tobacco smoke.

CYP3A4 50% pharmaceutical drugs. Induced by St John's Wort. Inhibited by grapefruit.

# PHASE 1 (HYDROXYLATION)

(Cytochrome P450)

NADPH + H+

NADP+ Fe++

Fe+++

 $XH + 2O_2 \longrightarrow XOH + H_2O + O_2$ 

## Cytochrome P450 enzymes are

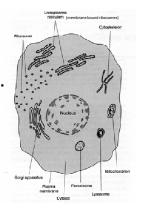
- 1. Hemoproteins (like hemoglobin).
- 2. Widely distributed across species especially in bacteria.
- 3. Present in the endoplasmic reticulum of all cells but greatest in the liver, small intestine, lung and glial cells.

Endoplasmic reticulum.

Major site of protein synthesis.

Synthesis of various lipids.

Oxidation of many xenobiotics.



# **Cytochrome P450 enzymes**

- 4. Require NADPH not NADH in their activation.
- 5. Require adequate levels of phosphatidyl choline rich in the membranes of the endoplasmic reticulum for optimal function.

# Cytochrome P450 enzymes

6. Are inducible and is therefore the mechanism of drug interaction.





Cytochrome P450 enzymes	
6. Are inducible and is therefore the mechanism of drug interaction.	
7. Can have polymorphisms (individual genetic isoforms)	
which can exhibit low catalytic activity.	
50% of all drugs prescribed to humans are metabolised by the	

# **Inability to Phase 1 detoxify**

various P450 enzymes.

However many P450 enzymes are inhibited by various drugs or their metabolic products, producing another cause of drug interaction.

- 1. Leads to either the absorption and displacement in phospholipid cell membranes,
- 2. Inactivation of specific enzymes
- 3. The toxin binding with serum albumin, which is antigenic leading to the production of antibodies against it.

I	P	450	IN	IDI	ICI	ΓΙΟΝ	ΝI	ITR	IFN	ITS
1	_	+	114	u	,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,		INL	, , ,		

Phosphatidylcholine, NADPH, (thus Mg), Fe, FMN, FAD, Thiolate (α-lipoic for Sulfur), Broccoli (IA2), Brussel sprouts (1A2), St John's Wort (3A4,5,7), Licorice, Black Walnut

High protein, Low carbohydrate, Ethanol (2E1), Zn, Cu, Cr, Ca, Mol, Se, Vit E, Vit C, Bioflavonoids, Beta Carotene, NAC, (NUTRIENT PHASE 1 & 2 contains all the above nutrients)

SUPPLEMENT ACCORDINGLY

	Phase 1	Biomarkers	Phase 2	Biomarkers	Key nutrients
	CYP 1A1	Estradiol Estrone Ethoxyresorufin, PAH, Ranaparene	Glutathione (finland Glutathione)	Napthalene Aspatame MSG	Cyskine, NAC, Glycine, Glatamic acid, PSF, Vit C, Pantethine, EFAs, Brassien
	10%	Caffeine Estradiol Estrone Methosyronerafia	Glucuronid- ation (CDP- Glucuronic orid)	Benesic seid Bilirobin Phenylltarbitol Vasillin	Glemenic acid NAD Mg, Fish oil
	CYP 1B1	Estradiol Estrone Ethocyresorufin PAH	Sulfation (PAPS Methionius SAM Transine Gire sulphore MSM Sulfar or Alsonici	Acetominophes (Paracetomol) Acetone DDT/DDE Ethylene glycol Toluene TRIC	PAPs Methionior, SAM Cystaine, PSP Gletathiose, Taurine Mal Mg Vit B2, Mn Glerosamine sultate, MSM, Suffor, av-Lionic neid
20dy Typoe	CYP 284	BHT Phonobarbital	Selfinidation Cellity solders	Sedium perobiosition	Cysteine Mal, Mg, BZ, Mn
Body Types and	CYP 2C9 10%	ibuprolen Warfarin	Acetylation (Acctyl Circle	boniaride Newsprint Petroleum SLS	Acetsl CoA factor Panenthine Vir B5 P-5-P Cyntaine Mr Vir C
anu	* CYP 2C19	Phenobarbitel Diazepum	Methylation (SAM)	Caffeine Phenol Nicotine	
Toxicity	CYP 206 30%	Codelac	Glycine	Aspirin Cholyl-CoA Sedium benzoate	Glycine, Co-factors (Felic Acid, PSP, Mg, Zn, 102, 103)
	CYF2EI	Acetominophen (Pursuetamol) Benzene Carbon tetrackloride Ethanal Juministal	Taurine	Carbon totrackluride Cholys-CaA	Tourine Cystoline Viz ID Iron P.S.P Mg Zn Cu Viz C
	CYP 3A4 50%	Acetaminophen Aflassin BI, GI Cudeine Nicotine Testasterone Warfarin	Cysteine		Cystaine Methionine Mg Zu PSP VitC
			Glatamine		Glatarnine Mg
			Threetine	Facest tesin	Thromise
		_	Ornithing	Ammonia	Тгурозрама
			Actinias	NAME OF TAXABLE PARTY.	

Phase 2
Conjugation

Phase 2 reactions conjugate the derivatives from Phase 1, where applicable, with molecules such as Glutathione, Glucuronic acid, Sulfate, Acetyl CoA, SAM, Taurine, Cysteine, Glycine and Threonine. This makes the derivatives even more water soluble for excretion through the urine or bile. Phase 1 toxic intermediate glutathione-s-transferase Reduced Glutathione Zn ++ Spinach Onion Garlic \* glutathione peroxidase Selenium

Broccoli \* Lemongrass Celery \* Watercress

Glutathione (P-5-P) conjugate

Oxidised Glutathione

A failure in the glutathione conjugation would lead to covalent combination to DNA and RNA and other cell proteins creating serious cell damage. Natural inducers of glutathione-stransferase are Spinach, Onion, Garlic, Broccoli, Lemon grass, Celery, Rosemary and Watercress.	
Glutathione conjugates are further metabolised before excretion. The glutamic and glycine groups are removed and an acetyl group donated by Acetyl CoA is added to the cysteine moiety.	
The resulting compound is a mercapturic acid, a conjugate of N. Acetyl Cysteine (NAC), which is then excreted in the urine.  N.Acetyl Cysteine (NAC) is thus an excellent supplement to use to activate this pathway.	

<ul><li>N. Acetyl Cysteine may activate detoxification via -</li><li>1. Glutathione</li><li>2. Acetylation</li><li>3. Sulfation</li></ul>	
4. Cysteine	
Glucuronidation conjugation is catalyzed by a variety of glucuronosyl-transferases with UDP-glucuronic acid as the glucuronyl donor. Glucuronidation conjugation is the favoured pathway for the metabolism of many neurotransmitters, hormones, phenol and benzoic acid.	
Natural Glucuronates Artichokes Cashew Soy Licorice Flax Alfalfa	

Sulfation conjugation uses 3-phosphoadenosine-5-phosphosulfate (PAPs), or sulfates or most commonly elemental sulfur or MSM or cysteine or  $\alpha$ -lipoic acid as the sulfur donor.

Many neurotransmitters and hormones are conjugated via this pathway.

Natural Sulfate donors

**Broccoli** 

**Asparagus** 

**Garlic** 

Mustard

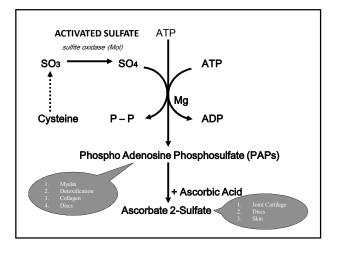
Dill

**Parsnip** 

Horseradish

Cabbage

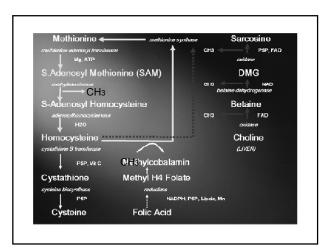
Stinging nettle



	-
Chemicals conjugated by Sulfation	
1. Acetone	
2. DDT / DDE	
3. Ethylene glycol	
4. Fluorine	
5. Toluene	
6. TRIC	
Acetylation conjugation uses Acetyl CoA as the acetyl donor.	
The reactions are catalyzed by	
acetyltransferases. Natural acetylaters - Endive, Pea,	
Cucumber, Watercress, Tomato	
The drug isoniazid used in the	
treatment of tuberculosis is conjugated by acetylation.	
conjugated by dectylation.	
	]
Chemicals conjugated by	
Acetylation	
1. Petroleum	
2. Newsprint	
3. Hypochlorite	

Methylation conjugation is catalyzed by the various *methyltransferases* employing S-Adenosyl methionine as the methyl donor (SAM).

Many hormones are initially hydroxylated, then methylated and lastly conjugated usually by glucuronidation or sulfation.



Amino acid conjugation can use either
Taurine,
Glycine,
Cysteine or
Threonine as conjugating donors.

Sodium benzoate is conjugated

with glycine.

CONJUGATE	BIOMARKER	NUTRIENTS
XOH + GLUTATHIONE	GLUTATHIONE-S- TRANSFERASE	GLUTATHIONE (NAC, Glutamate, Glycine) B6, Zn
XOH+ GLUCURONIDATION	GLUCURONIC ACID	GLUCURONIC ACID
1. XOH + SULFATION	1. PAPs	PAPs, S, MSM
2. SULFITE OXIDASE	2. SULFITE OXIDASE	Mol, Fe.
XOH + ACETYLATION	ACETYL CoA	Acetyl CoA (B5, Mg, Acetic acid)
XOH + METHYLATION	SAM	Methionine, MgATP, B12, Folic, Betaine, DMG
XOH + TAURINE	TAURINE	Taurine, NAD, Vit C, Vit A
XOH + THREONINE	THREONINE	Threonine
XOH + GLYCINE	GLYCINE	Glycine, B6, B2, Mg, Folic.
XOH + CYSTEINE	CYSTEINE	Cysteine, NAC, Methionine B6



Nightshades

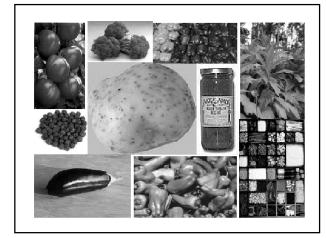


Solanidine
An alkaloid produced by the
decomposition of solanine, as
white crystalline substance
having a harsh bitter taste.

It is a fat soluble	neurotoxin
OH OH OH	
Solatrines modely	- 300modine - 100e07

Other acetylcholine blockers are prevalent in Solanine containing foods.

Egg plant (Aubergine)
Green peppers Paprika
Potato Potato starch
Tomato Tomato paste
Tobacco Relish
Spices Broccoli
Black pepper Chili



They may cause a variety of symptoms Joint pains **Abdominal bloating** Urinary bladder weakness **Pupillary dilation Dry mouth** Skin rashes Weight gain **Tiredness** 

#### Short Term:

Snort term:

Effects on the nervous system included increased heart, pulse, and respiratory rates, sedation, and coma.

Effects resulting from cell membrane disruption included internal hemorrhaging, edema, diarrhea, constriction of the abdominal muscles, and lesions of the stomach and duodenum Haemorrhagic damage to the gastrointestinal tract as well as to the

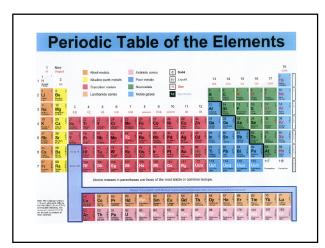
retina
Higher concentrations inhibited fibroblast cell growth
Abdominal cramps, gas, Diarrhea
PMS

Depressed central nervous system

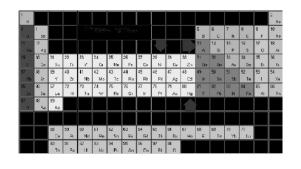
Depressed central nervous system kidney inflammation reduced iron uptake Dizziness sleepiness Vision problems including ocular pressure Mental confusion Flu like illness in higher dosages Heart Attack Death (rare)

Long Term: Congenital spina bifida Low thyroid and other endocrine dysfunctions Vision problems, due to wide iris allowing too much light, that can cause damage Loss of night vision Cancers of the brain, breast, endometrium, lung, and thyroid Loss of memory and thinking ability Overeating and malnourish due to diarrhea Birth defects General weakness Depression Immune system dysfunctions Osteoporosis Arthritis Joint Problems Loss of Concentration Sexual dysfunctions Aphrodisiac symptoms over stimulation	
Challenging for toxicity From weakness with the definitive meridian.	
Chemicals vial Toxic metals vial Radiation vial	
Toxins	
Chemicals - Black walnut Coriander spice NAC Lemon balm Rosemary Yarrow Other spices Charcoal	
Charcoai	

# **Toxic Metals**



# **The Periodic Table**



	•
Toxic Metals	
Toxic metals inhibit various enzyme pathways.	
Many active sites on enzymes are sulphur amino acids.	
Toxic metals such as mercury always attack these SH bonds.	
	1
Sulfur bearing Amino Acids	
1. Methionine	
2. Cysteine	
3. Cystine	
4. Glutathione	
5. Taurine	
	1
Madala that have a transfer	
Metals that have a transient charge (Fe++ and Fe+++) are	
capable of inducing oxidative	
stress directly. Other metals may induce	
oxidative stress by their reactions with low molecular thiols	
(glutathione and others) and	
therefore reduce the antioxidant potential.	
potential.	

	_
Fe++ Fe+++	
reduced oxidised	
Cu+ Cu++	
Colours and Toxic Metals	
The cone colour that a person	
weakens to gives some indication	
of potential toxic metal affinity.	
Aluminium	
Absorption	
619 nm	
Emission	

#### **Aluminium**

Salt in tap water as a deflocculant and softener. Antacid, Anti-inflammatory, antidiarrhoeal medication. Aluminium silicates in medications.

All foods wrapped with aluminium foil. Oxo cubes.

Insides of milk and fruit juice cartons.

Aluminium take-away cartons.

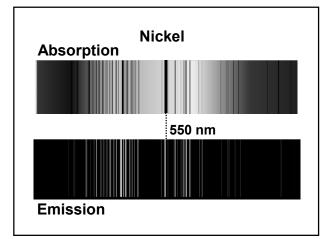
Aluminium food, soft drinks and beer cans.

Squeezy tubes such as tomato paste.

Baking powder, Self raising flour, Salt and certain food additives. Naturally high levels in Tea, spearmint and peppermint teas, tea bags, instant coffee, Spinach and Potatoes. Processed cheese.

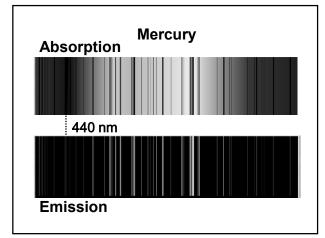
Deodorants, antiperspirants, skin lotions, make-ups, douches, toothpaste.

Saucepans, frying pans, kettles, baking sheets.



#### Nickel

Sunflower seeds, Licorice, Hydrogenated oils, Peanut butter, Vegetable shortenings. Rolled oats. 7% Stainless steel. Watch straps and glasses frames. Non silver or gold jewellery such as earrings. Dental fillings and retainers. Cooking utensils and cappucino machines. Nickel / cadmium batteries. Cosmetics and permanent waves. Tobacco smoke, industrial exposure and ceramics. Superphosphate fertilizers.



#### Mercury

Dental amalgams. High fructose corn syrup.
Sanitary towels, Cotton balls and buds, Dental floss, Toothpicks, Paints, Explosives, Batteries, Mercurial diuretics, Fungicides, Laxatives containing camomel, Hemorroid suppositories, Fluorescent lamps, Cosmetics, Hair dyes. Fibreglass, Manufacture and delivery of petroleum. Sewage sludge.
Methylmercury chlorine bleaches. Fabric softeners, Polishes, wood preservers, Latex, Solvents, Plastics, Inks used by printers and tatooists, some Paints.
Salt, Fish from contaminated water such as tuna. Vaccines (thermerasol)

## Potentially toxic elements

**RED CONE** 

**BLUE CONE** 

Aluminium, Lead, Fluoride, Indium, Palladium, Copper, Manganese, Vanadium Mercury Gallium Thallium

**GREEN CONE** 

Arsenic Nickel

Cobalt



	1
Diagnosis	
Any enzymatic pathway, which exhibits acquired inhibition.	
•	
Most common positive meridians are	
Kidney and Bladder	
	1
Procedure	
Positive meridian will negate against the TOXIC METAL nosode.	
Patient will weaken to a specific toxic metal(s).	
Best to test with full strength toxic metal solution from strength	
to weakness.	
Treatment	
Often the best chelating agent is adjacent the toxic metal in the	
periodic table.	

Toxins Toxic metals - Black walnut Coriander herb Coriander spice Lemon balm Lipoic acid Yarrow Glutathione Vitamin C for nickel NAC	
Toxic metals in the water soluble components of the body will chelate easily with any of the previous nutrients.  Toxic metals in the fat soluble components of the body may need a solvent such as alkylglycerol from Chlorella from algae.	
Amalgam extraction  Chlorella 2 caps immediately before extraction. 2 caps immediately after extraction. 2 caps one hour later. Use Liquid Selenium as a mouthwash during amalgam extractions as it bonds with mercury to form selenates. Spit out after rinsing the mouth each time.	

# RADIATION

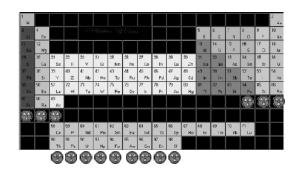
# **Ionising Radiation**



lonising radiations come from alpha and gamma particles from the radioactive isotopes of certain elements.

All the elements past number 83 are unstable and radioactive.

# The Periodic Table



Ionising Radiation	
May also come from exposure to UV light and cosmic radiations in high altitude jet travel	
	I
DIAGNOSIS Positive meridian will negate with the RADIATION nosode. Identify causative element.  Spleen 4	
Toxins  Radiation - Chlorella Coriander spice Smart Vitamin C (Rutin) Turmeric Yarrow	