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Micronutrients: Vitamin B1 - Thiamin

Dr. Ritamarie Loscalzo


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B Vitamins At a Glance

Letter	Names	Notes/Actions
B1	Thiamin, Benfotiamine	Energy, heart, muscle, and nerve function
B2	Riboflavin, R 5'-Phosphate	Energy, red blood cells, vision
B3	Niacin, Nicotinic Acid, Niacinamide	Energy, nerve function, circulation and heart
B4	Choline, Adenine, Carnitine	Loosely considered as B vitamins - cell membranes, memory, neuromuscular
B5	Pantothenic Acid	Coenzyme A, adrenals, skin
B6	Pyridoxine, Pyridoxal 5'-Phosphate	Brain and nerve, hormones, protein synthesis
B7	Biotin	Hair, metabolism
B8	Inositol	Loosely considered a B vitamin
B9	Folate, Methylfolate, Folinic Acid	Red blood cell production, DNA repair, brain
B10	Pteroylmonoglutamic Acid (PABA – Para-aminobenzoic Acid)	Really a form of folate, skin protector
B11	Salicylic Acid	Not technically a vitamin, loosely categorized
B12	Cobalamin	Red blood cells, DNA repair, nervous system

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General Info - Thiamin

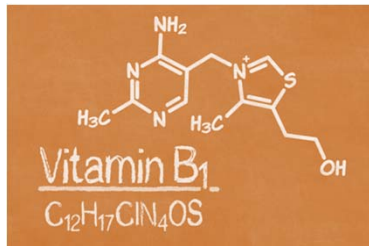
- ✓ The first vitamin isolated in pure form
- ✓ Water soluble
- ✓ Free form in plants
- ✓ Phosphorylated form in animal sources: thiamin diphosphate (TDP) and thiamin pyrophosphate (TPP)
- ✓ Thiamin hydrochloride (HCl) and mononitrate in supplements
- ✓ Benfotiamine is a fat-soluble form which lasts longer, yielding potentially therapeutic benefits over ordinary thiamin



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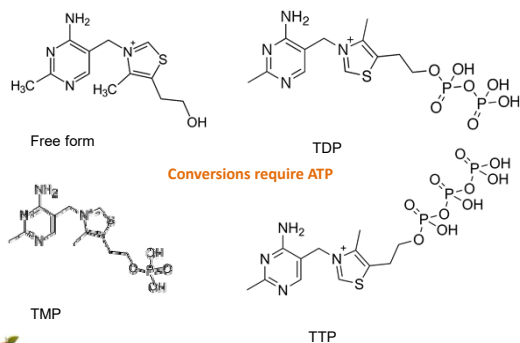
Thiamin Chemistry

- ✓ Oxidative phosphorylation of keto acids and 2-keto sugars
- ✓ Pyrimidine ring plus thiazole ring linked by a methylene (CH₂) bridge



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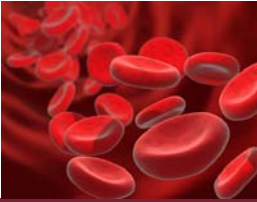
Thiamin Forms



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Thiamin Transport

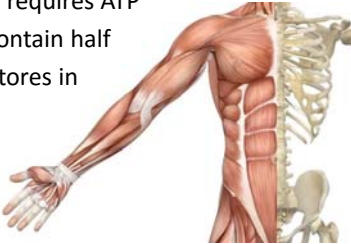
- ✓ Transported in blood **mostly as free form** or bound to albumin
- ✓ Some transported as **thiamine monophosphate**
- ✓ **90% is in the blood cells**, not the plasma
- ✓ In red blood cells, **most thiamin is as TDP** with smaller amounts as TMP or free



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Thiamin Storage

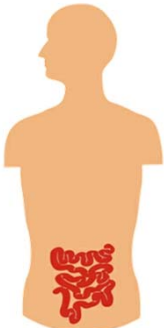
- ✓ Humans store approximately 30 mg
- ✓ 80% of all thiamin in body is thiamin diphosphate (TDP)
- ✓ Conversion to TDP requires ATP
- ✓ Skeletal muscles contain half
- ✓ Other significant stores in
 - Liver
 - Brain
 - Heart
 - kidneys



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Thiamin Absorption

- ✓ Free form absorbed
- ✓ Mainly in jejunum, somewhat in ileum and duodenum
- ✓ Absorbed via active transport (sodium dependent) at low intakes
- ✓ Absorbed via diffusion at higher levels



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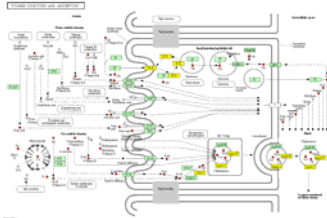
Influences on Thiamin Absorption

Decreases

- ✓ Ethanol
- ✓ Thiaminases in raw fish and raw horsetail
- ✓ Polyhydroxyphenols
 - Tannic acid
 - Caffeic acid
 - Tea
 - Coffee
 - Blueberries
 - Brussels sprouts
 - Red cabbage
- ✓ Calcium and magnesium with polyhydroxyphenols

Increases

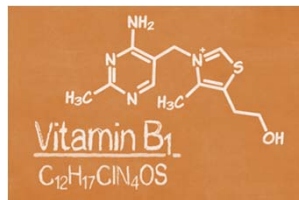
- ✓ Vitamin C: deactivates effect of polyhydroxyphenols
- ✓ Citric acid: deactivates effect of polyhydroxyphenols



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Main Thiamin Functions

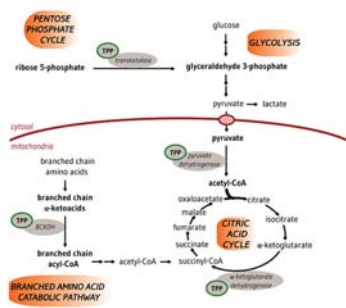
- ✓ Coenzyme for energy transformation
- ✓ As a coenzyme for synthesis of pentoses and nicotinamide adenine dinucleotide (NADPH)
- ✓ Membrane and nerve conduction in a non-coenzyme capacity



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Thiamine Biochemistry

Figure 1. Metabolic Pathways Requiring Thiamin Pyrophosphate




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Thiamin Functions

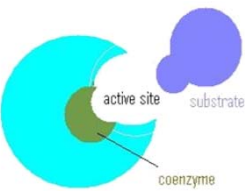
- ✓ Enhances circulation
- ✓ Assists in blood formation
- ✓ Important in carbohydrate metabolism
- ✓ Role in the production of hydrochloric acid
- ✓ Optimizes cognitive activity and brain function
- ✓ Important co-factor in the Kreb's cycle (energy)
- ✓ Needed for proper muscle tone of the intestines, stomach, and heart
- ✓ Acts as an antioxidant
- ✓ Role in appetite regulation
- ✓ Helps as part of a leaky gut protocol to rebuild mucosa
- ✓ Role in immune function and protection from stress response



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Thiamin as a Coenzyme in Kreb's Cycle


- ✓ Pyruvate decarboxylase complex
- ✓ Alpha ketoglutarate dehydrogenase
- ✓ Branched chain alpha keto acid dehydrogenase
- ✓ Transketolase needed for pentoses and NADPH



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Thiamin and Blood Sugar Balance

- ✓ Coenzyme for transketolase (Tk), pyruvate dehydrogenase and α -ketoglutarate dehydrogenase complexes - intracellular glucose metabolism
- ✓ Thiamine levels and thiamine-dependent enzyme activities are reduced in diabetics
- ✓ Genetics link thiamine and diabetes: Tk, *SLC19A2* gene, transcription factor *Sp1*, α -1-antitrypsin, and *p53*
- ✓ Deficiency has a definitive role in diabetic complications: endothelial vascular diseases, lipid profile, retinopathy, nephropathy, cardiopathy, and neuropathy

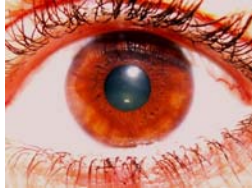


The Impact of Thiamine Treatment in the Diabetes Mellitus, Khanh Vinh quoc Luong, b and Lan Thi Hoang Nguyen, J Clin Med Res. 2012 Jun; 4(3): 153-160. May 2012.

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Cataracts and Thiamin

- ✓ Thiamine, along with other nutrients, may lower risk of cataracts
- ✓ People with plenty of protein and vitamins A, B1, B2, and B3 (or niacin) in their diet are less likely to develop cataracts
- ✓ Getting enough vitamins C, E, and B complex vitamins -- particularly B1, B2, folate, and B12 -- may further protect the lens



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Alzheimer's Disease and Thiamin

- ✓ Lack of thiamine can cause dementia
- ✓ Oral thiamine has been shown to improve cognitive function of patients with Alzheimer's
- ✓ Absorption of thiamine is poor in elderly individuals



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Thiamin Effects on Heart and Brain Health

- ✓ Diuretic use in people with CHF may deplete thiamin; taking thiamine supplements may help
- ✓ Low levels of thiamine are associated with depression




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Things That Deplete Thiamin


- ✓ Gluten
- ✓ Processed and overly cooked foods
 - they strip minerals from the body
- ✓ Alcohol
- ✓ Herbicides and pesticides
- ✓ Refined sugar, corn syrup and artificial sweeteners
- ✓ Prolonged stress
- ✓ Table salt
- ✓ Tap water - sodium fluoride
- ✓ Unfermented soy products
- ✓ Regular and decaffeinated coffee or black tea
- ✓ Excess zinc and vitamin D



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Causes of Thiamin Deficiency


- ✓ Health Conditions:
 - Alcoholism
 - Crohn's disease
 - Anorexia
 - Kidney dialysis
- ✓ Deficient soil
- ✓ Processed foods
- ✓ Fluoride in water
- ✓ Drugs: antibiotics, Dilantin, phenytoin, sulfa drugs, oral contraceptives
- ✓ Heavy caffeine consumption
- ✓ High carbohydrate diet



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Impact of Thiamin Deficiency

- ✓ Headache
- ✓ Nausea
- ✓ Fatigue
- ✓ Irritability
- ✓ Depression
- ✓ Abdominal discomfort
- ✓ Difficulty digesting carbohydrates
 - leading to pyruvic acid build up resulting in:
 - Loss of mental alertness
 - Difficulty breathing
 - Heart damage
- ✓ Extreme deficiency is beriberi



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Thiamin Deficiency: Beriberi

✓ Wet (edema) and dry (emaciation/body wasting)

- ❑ Confusion
- ❑ Trouble breathing because of fluid in the lungs
- ❑ Uncontrolled eye movements (nystagmus)
- ❑ Anorexia and weight loss
- ❑ Neurological: apathy, confusion, short-term memory loss, irritability
- ❑ Cardiac: hypertrophy and altered heart rate - may lead to death from heart failure
- ❑ Swelling, tingling, or burning sensation in the hands and feet
- ❑ Rare in developed world



❑ Occurs in alcoholics

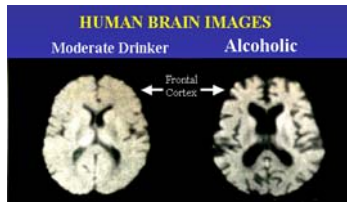


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Thiamin Deficiency: Wernicke-Korsakoff Syndrome

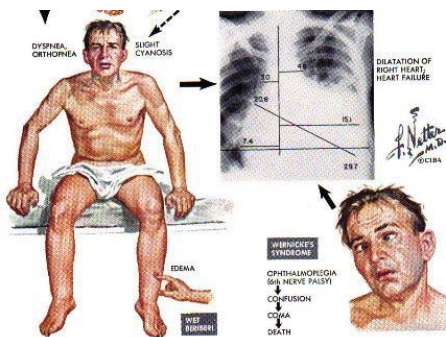
✓ Common in alcohol abusers

- ❑ Short-term memory loss
- ❑ Jerky eye movement
- ❑ Staggering gait
- ❑ Encephalopathy
- ❑ Nystagmus
- ❑ Ataxis
- ❑ Opthamoplegia



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Thiamin Deficiency Diseases



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Deficiency Signs From Nutrient Assessment Chart

Vitamin B1	<ul style="list-style-type: none"><input type="checkbox"/> Anxiety<input type="checkbox"/> Depression<input type="checkbox"/> Hysteria<input type="checkbox"/> Loss of appetite; in extreme cases beriberi (mostly in alcoholics).<input type="checkbox"/> Muscle cramps
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Impact of Thiamin Excess


- ✓ Little risk of toxicity
- ✓ Studies done with 500 mg per day or more
- ✓ 100x RDA parenterally have induced:
 - Headache
 - Convulsions
 - Arrhythmia
 - Anaphylactic shock



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Thiamin Drug/Herb Interactions

- ✓ **Horsetail (*Equisetum arvense* L.):** Contains thiaminase, an enzyme which inhibits thiamine
- ✓ **Digoxin:** May reduce the ability of heart cells to absorb and use vitamin B1, particularly when digoxin is combined with Lasix, aka furosemide, a diuretic
- ✓ **Diuretics, particularly Lasix:** Reduce levels of vitamin B1
- ✓ **Phenytoin (Dilantin):** Some people have lower levels of thiamine in their blood, which may contribute to the side effects of the drug



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Assessing Thiamin Status

- ✓ Serum is not accurate
- ✓ Spectracell
- ✓ Erythrocyte transketolase
- ✓ NutrEval by Genova / Metamatrix
- ✓ Questionnaires and good history taking for signs and symptoms



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Thiamin RDA

Table 1. Recommended Dietary Allowance (RDA) for Thiamin			
Life Stage	Age	Males (mg/day)	Females (mg/day)
Infants	0-6 months	0.2 (AI)	0.2 (AI)
Infants	7-12 months	0.3 (AI)	0.3 (AI)
Children	1-3 years	0.5	0.5
Children	4-8 years	0.6	0.6
Children	9-13 years	0.9	0.9
Adolescents	14-18 years	1.2	1.0
Adults	19 years and older	1.2	1.1
Pregnancy	all ages	-	1.4
Breast-feeding	all ages	-	1.4



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Methods of Repletion

- ✓ Food and herbs
- ✓ Oral supplements
- ✓ Intravenous (Myer's cocktail)
- ✓ Intramuscular




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Thiamin Food Sources

- ✓ Brown rice
- ✓ Egg yolks
- ✓ Fish
- ✓ Legumes
- ✓ Liver
- ✓ Peanuts
- ✓ Peas
- ✓ Pork
- ✓ Poultry
- ✓ Whole grains
- ✓ Sunflower seeds
- ✓ Macadamia nuts
- ✓ Most nuts

- ✓ Asparagus
- ✓ Brewer's yeast
- ✓ Broccoli
- ✓ Dulse
- ✓ Kelp
- ✓ Squash
- ✓ Brussels sprouts
- ✓ Oatmeal
- ✓ Plums
- ✓ Dried prunes
- ✓ Raisins
- ✓ Spirulina
- ✓ Watercress




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Herbs High In Thiamin

- ✓ Alfalfa
- ✓ Bladderwrack
- ✓ Burdock root
- ✓ Catnip
- ✓ Cayenne
- ✓ Chamomile
- ✓ Chickweed
- ✓ Dandelion
- ✓ Eyebright
- ✓ Fennel seed
- ✓ Fenugreek
- ✓ Hops
- ✓ Horsetail
- ✓ Lemongrass

- ✓ Licorice
- ✓ Mullein
- ✓ Nettle
- ✓ Oat straw
- ✓ Paprika
- ✓ Parsley
- ✓ Peppermint
- ✓ Raspberry leaf
- ✓ Red clover
- ✓ Rose hips
- ✓ Sage
- ✓ Shepherd's purse
- ✓ Yarrow
- ✓ Yellow dock



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WHFoods.com Thiamin Chart

World's Healthiest Foods ranked as quality sources of vitamin B1

Food	Serving Size	Cals	Amount (mg)	DRI/DV (%)	Nutrient Density	World's Healthiest Foods Rating
Asparagus	1 cup	39.6	0.29	24.17	11.0	excellent
Sunflower Seeds	0.25 cup	204.4	0.52	43.33	3.8	very good
Green Peas	1 cup	115.7	0.36	30.00	4.7	very good
Flax Seeds	2 TBS	74.8	0.23	19.17	4.6	very good
Spinach	1 cup	41.4	0.17	14.17	6.2	very good
Brussels Sprouts	1 cup	56.2	0.17	14.17	4.5	very good
Cabbage	1 cup	43.5	0.11	9.17	3.8	very good
Eggplant	1 cup	34.6	0.08	6.67	3.5	very good
Romaine Lettuce	2 cups	16.0	0.07	5.83	6.6	very good
Mushrooms, Crimini	1 cup	15.8	0.07	5.83	6.6	very good

<http://www.drRitamarie.com/go/ThiaminFoods>


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Forms of Supplementation

Absorption of oral magnesium supplements vary from 4% absorbed to about 50%.

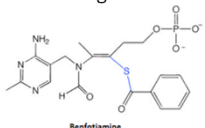
Thiamine HCl	Thiamin Pyrophosphate (aka cocarboxylase)	Benfotiamine
<ul style="list-style-type: none">✓ Most common form✓ Least expensive form	<ul style="list-style-type: none">✓ <u>Active form</u>➢ Pyruvate dehydrogenase➢ Pyruvate decarboxylase in ethanol fermentation➢ Alpha-ketoglutarate dehydrogenase➢ Branched-chain amino acid dehydrogenase➢ 2-hydroxyphytanoyl-CoA lyase➢ Transketolase	<ul style="list-style-type: none">✓ Lipid soluble form of thiamine✓ Helpful in diabetic neuropathy✓ Reduces microvascular damage from high blood sugar✓ Transketolase activator – reduces AGEs by directing to the pentose phosphate pathway



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Benfotiamine: Absorption and Structure

- ✓ Absorption approximately 5-fold Thiamin and 3.7-fold Thiamin Hydrochloride
- ✓ Serum increases detected in serum within one hour
- ✓ S-{{(Z)-2-{{(4-amino-2-methylpyrimidin-5-yl) methyl-formylamino}-5-phosphonooxypent-2-en-3-yl} benzenecarbothioate
- ✓ Common dose: 300-600 mg over the course of the day




Benfotiamine

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Benfotiamine Research

- ✓ Alzheimer's
- ✓ Pain
- ✓ Cardiac tissue
- ✓ Glucose metabolism
- ✓ Skeletal muscle and bone
- ✓ Inflammation and immunology
- ✓ Oxidative stress protection
- ✓ Lungs
- ✓ Kidneys
- ✓ Eyes



http://examine.com/supplements/benfotiamine/

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Benfotiamine Studies

- ✓ Alzheimer's: Attenuates glucose-induced increase in β -amyloid protein synthesis and normalize the glutathione:glutathione disulfide (GSSG) ratio in the cerebral cortex
Jun Xi, et al. Benfotiamine prevents increased β -amyloid production in HEK cells induced by high glucose. *Neurosci Bull*. (2012)
- ✓ Weakly attenuates Advanced Glycemic End product (AGE) bound to serum albumin (a complex known as AGE-albumin) from activating macrophages and inducing oxidative stress
- ✓ Attenuates LPS induced mitochondrial membrane potential loss in the macrophages



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Thiamin Resources

- ✓ *Advanced Nutrition and Human Metabolism – Gropper, Smith and Groff*
- ✓ Linus Pauling Institute:
<http://www.drRitamarie.com/go/LPIThiamin>
- ✓ <http://www.drRitamarie.com/go/UMMCThiamin>
- ✓ <http://www.drRitamarie.com/go/ExamineBenfotiamine>



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