Vitamins B6 and B12		
Water-Soluble	B vitamins	
Vitamins	Thiamin (B1), riboflavin (B2), niacin (B3), pantothenic acid (B5), pyridoxine (B6), biotin (B7), cobalamin (B12), folate	
	Characteristics:	
	1-Not significantly stored in the body	
	2-Must be supplied regularly in the diet	
	3-Excess excreted	
Vitamin B Complex	1-Present in small quantities in different types of food	
	2-Important for growth and good health	
	3-Help in various biochemical processes in cell	
	4-Function as coenzymes	
	Vitamin R <sub>c</sub>	

#### VILUIIIIII D6

#### Has three forms:

1-Pyridoxine, 2-Pyridoxal, 3-Pyridoxamine

All 3 are converted to \*\* active form\*\*: pyridoxal phosphate (PLP)

**Functions of Vitamin B6:** 

- 1-Transamination
- 2-Deamination
- 3-Decarboxylation
- **4-Condensation reactions**

#### **Transamination**



Convert alanine to pyruvate by: Alanine Aminotransferase (ALT) enzyme and converting alpha-KG to glutamate by pyridoxal phosphate (PLP)

## **Decarboxylation**

- 1-Formation of Chatecholamines: Dopamine, norepinephrine and epinephrine
- 2- Formation of histamine
- 3- Formation of Serotonin

#### **Condensation reactions**

Formation of ALA by ALA synthase, The regulatory step in hemoglobin synthesis

(ALA) = Aminolevulinic acid **Disorders of Vitamin** 

**B6 Deficiency** 

A-Dietary deficiency: (rare)

1-Newborn infants fed on formulas low in B6, 2-Women on oral contraceptives, 3-Alcoholics.

\* Isoniazid treatment for tuberculosis can lead to vitamin B<sub>6</sub> deficiency by forming inactive derivative with PLP

B- Deficiency leads to poor activity of PLP-dependent enzymes causing:

1-Deficient amino acid metabolism,

	2-Deficient lipid metabolism, 3-Deficient neurotransmitter synthesis:
	Serotonin, epinephrine, norepinephrine and gamma amino butyric acid (GABA)
	C- PLP is involved in the synthesis of sphingolipids so Its deficiency leads to
	demyelination of nerves and consequent peripheral neuritis.
Symptoms of	A-Mild deficiency leads to:
deficiency	1-Irritability 2-Nervousness 3-Depression
	B-Severe deficiency leads to:
	1-Peripheral neuropathy 2-Convulsions
	Vitamin D

### Vitamin B<sub>12</sub>

# Forms of Vitamin $B_{12}$ :

- 1-Cyanocobalamin 2-Hydroxycobalamin 3-Adenosylcobalamin (major storage form in the liver)
- 4-Methylcobalamin (mostly found in blood circulation)

Coenzyme forms of  $B_{12}$ :

Adenosylcobalamin and Methylcobalamin

- -Coenzymes for metabolic reactions
- \* Body can convert other cobalamins into active coenzymes

### Vitamin B12 (cobalamin):

- Mainly found in animal liver
- Bound to protein as Methylcobalamin or 5'-deoxyadenosylcobalamin

#### Characteristics:

- Essential for normal nervous system function and red blood cell maturation
- Not synthesized in the body and must be supplied in the diet
- Binds to intrinsic factor and absorbed by the ileum
- Intrinsic factor is a protein secreted by cells in the stomach

# Vitamin B<sub>12</sub> Storage

- Liver stores vitamin B12 (4-5 mg)
- Other B vitamins are not stored in the body
- Vitamin B12 deficiency is observed in patients with IF deficiency due to autoimmunity or by partial or total gastrectomy
- \* Clinical deficiency symptoms develop in several years

Functions of Vitamin	Two reactions require B12:
B <sub>12</sub>	1. Conversion of methylmalonyl-CoA to succinyl-CoA.
	The enzyme in this pathway, methyl-malonyl-CoA mutase, requires B12
	2. Conversion of homocysteine to methionine.
	Methionine synthase requires B12 in converting homocysteine to methionine
B <sub>12</sub> Deficiency and	■ Homocysteine re-methylation reaction is the only pathway where N⁵-methyl
Folate Trap	TH4 can be returned back to tetrahydrofolate pool
	Hence folate is trapped as
	N⁵-methyltetrahydrofolate (folate trap)
	■ This leads to folate deficiency and deficiency of other TH4 derivatives (N⁵-N¹0
	methylene TH4 and N <sup>10</sup> formyl TH4) required for purine and pyrimidine
	syntheses
	TH4: Tetrahydrofolate
Disorders of Vitamin	Pernicious anemia

B <sub>12</sub> Deficiency	1-Megaloblastic anemia
	2-Vitamin B <sub>12</sub> deficiency is mainly due to the deficiency of intrinsic factor
	Demyelination
	Myelin sheath of neurons is chemically unstable and damaged
	Neuropathy
	■ Peripheral nerve damage
Causes of neuropathy	
	tamin B <sub>12</sub> leads to accumulation of methylmalonyl CoA
<ul><li>Deficiency of vi</li></ul>	tamin B <sub>12</sub> leads to accumulation of methylmalonyl CoA ethylomalonyl CoA are used instead of malonyl CoA for fatty acid synthesis
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<ul> <li>Deficiency of vi</li> <li>High levels of m</li> <li>Myelin synthes</li> </ul> Neuropsychiatric	ethylomalonyl CoA are used instead of malonyl CoA for fatty acid synthesis zed with these abnormal fatty acids is unstable and degraded causing neuropathy  Neurological symptoms
■ Deficiency of vi ■ High levels of m ■ Myelin synthes  Neuropsychiatric symptoms of Vitamin	ethylomalonyl CoA are used instead of malonyl CoA for fatty acid synthesis zed with these abnormal fatty acids is unstable and degraded causing neuropathy  Neurological symptoms  Paraesthesia (abnormal sensation) of hands and feet

**Psychiatric symptoms:** 

DepressionUnstable mood

■ Confusion and memory loss