# ROLE OF DIETARY CONSTITUENTS IN COGNITIVE DYSFUNCTION

#### Essay

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## علاقة خلل العناصر الغذائية باضطراب الوظائف المعرفية رسالة

توطئة للحصول على درجة الماجستير في الأمراض العصبية والنفسية

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#### **Summary**

Alzheimer's disease is a multifactorial disorder that has many physiological, biochemical and neurochemical facets. Aging is the major risk factor for AD that coexists with other causes of cognitive decline, particularly vascular dementia.

The processes underlying the pathology of AD involve several factors including mitochondrial dysfunction, abnormal protein aggregation, metal accumulation, inflammation and excitotoxicity. Although the relationship between these factors and the development of AD is multidirectional, oxidative damage is considered a common thread linking some of these factors is evident before cytopathologic hallmarks of the disorder.

A growing body of research indicates that nutritional deficiencies contribute to age-related cognitive decline including that which accompanies AD. But, it is important to mention that intervention trials have not been adequately designed to test whether these associations are causal.

Controlled studies in mice and patients with MCI and dementia have demonstrated that cognitive performance is subject to dietary compromise and that key dietary supplementation can alleviate and in some cases reverse the impact of dietary deficiencies on cognitive performance.

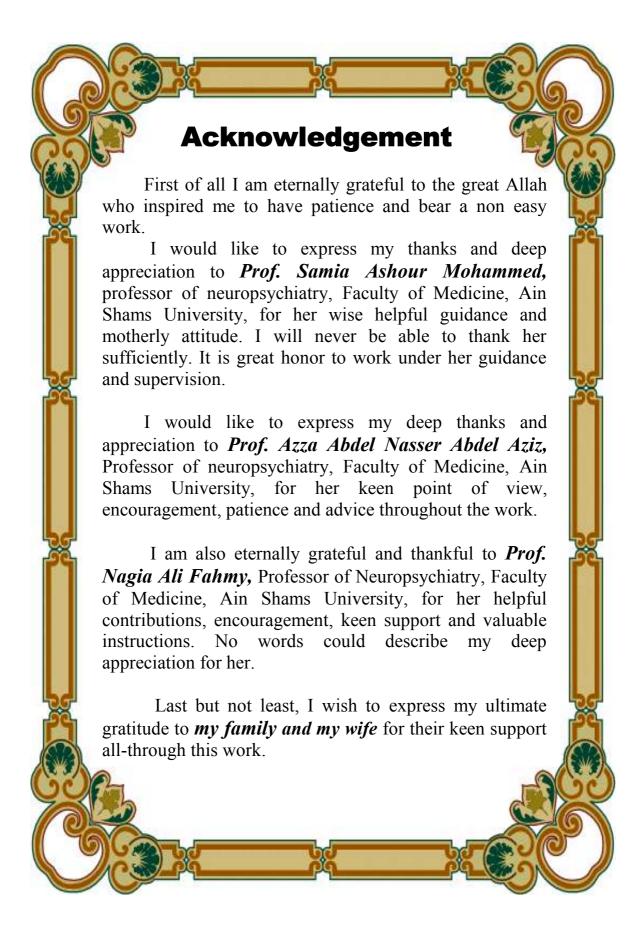


Table of contents	pages
List of abbreviations	
List of tables & figures	
Introduction	
Aim of the work	
Chapter one: Pharmacology of Dietary constituents	
Chapter two: Role of Dietary constituents in	53
cognitive functions	
Chapter three: Detection of dietary dysregulation in	
dementia	
Chapter four: Dietary treatment of cognitive	131
dysfunction and dementia	
Discussion	
Conclusion & recommendations	
Summary	170
References	173
Arabic summary	

#### **List of abbreviations**

(MeOPhSe)2 : p,p-methoxyldiphenyl diselenide

(PhSe)2 : Diphenyl diselenide

**1,25-OH)2D** : 1,25-dihydroxy-vitamin D

**250HD** : 25-hydroxyvitamin D

**5HTP** : 5-hydroxytryptamine

**8-OHdG** : 8-hydroxy-2-deoxyguanosine

**AA** : Arachidonic acid

**ACAT** : Acyl-coenzyme A cholesteryl acyltransferase

**AChE** : Acetylcholinesterase

**AD** : Alzheimer's disease

**ADAM10** : A Disintegrin And Metallopeptidase 10

**ADAScog** :Alzheimer's disease Assessment Scale

cognitive subscale

**AFR** : Ascorbate free radical

**ALCAR** : Acetyl-L-carnitine

Am80 : Tamibarotene

**AMP** : Adenosine monophaosphate

**APO** : Apolipoprotein

**APP** : Amyloid precursor protein

**ATP** : Adenosine triphosphate

**ATRA** : All-trans retinoic acid

**Aβ** : Amyloid  $\beta$ 

**BDNF** : Brain-derived neurotrophic factor

**BMAA** : Beta-N-methylamino-L-alanine

**XC-** : Cystine/glutamate antiporter

**CADPR** : Cyclic ADP-ribose

**CH** : Cognitive health

**ChAT** : Choline acetyltransferase

**CLN** : Colostrinin

**ALCAR** : Acetyl-Lcarnitine

**COX** : Cyclooxygenase

**CRABP** : Cellular retinoic acid-binding protein

**CRBP** : Cellular Retinoid Binding Proteins

**CSF** : Cerebrospinal fluid

**DHA** : Docosahexaenoic acid

**DHLA** : Dihydrolipoic acid

**DTI** : Diffusion tensor imaging

**DTMP** : Deoxythymidine monophosphate

**DUMP** : Deoxyuridine monophosphate

**EAE** : Experimental autoimmune encephalomyelitis

**EPA** : Eicosapentaenoic acid

**FA** : Friedreich's ataxia

**FAD** : Flavine adenine dinucleotide

fAβ : fibrils β-amyloid

**FDN** : Flavin dinucleotide

**FGAR** : Formylglycinamide ribonucleotide

**FMN** : Flavin mononucleotide

**GABA** : γ-amino butyric acid

**GAD** : Glutamic acid decarboxylase

**GAR** : Glycinamide ribonucleotide

**GDNF** : Glial cell line–derived neurotrophic factor

GLUT : Glucose transporter isoforms

**GPXs** : Glutathione peroxidases

**GSH** : Glutathione

**HAT** : Histone acetyltransferase

**HDAC** : Histone deacetylase

**HDL** : High-density lipoproteins

**HHcy** : Hyperhomocysteinemia

**HIF-1** : Hypoxia-inducible factor 1

**HNE** : 4-hydroxy-2-transnonenal

**HVD** : Hydroxy vitamin D

**ICAM-1** : Intracellular adhesion molecule-1

**IDIs** : Iodothyronine deiodinases

**IDL** : Intermediate-density lipoprotein

**IFN** : Interferon

Ig : Immunoglobulin

IL : Interleukin (e.g. IL-1)

**INOS** : Inducible nitric oxide synthetase

**IPF** : Isoprostane F

**JAK** : Janus kinase

**LDL** : Low-density lipoproteins

**LF** : Lactotransferrin

**LOAD** : Late onset Alzheimer disease

**LPL** : Lipoprotein lipase

**LPS** : Lipopolysaccharide

**LTDM** : Long-term declarative memory

MCI : Mild cognitive impairment

**MCO** : Metal-catalyzed oxidation

MDA : Malondehyde

**MHC** : Major histocompatibility complex

MS : Multiple sclerosis

MTA : Medial temporal lobe atrophy

**NAD** : Nicotinamide adenine dinucleotide

**NADH** : Nicotinamide adenine dinucleotide

hydrogenase

**NADP** : Nicotinamide adenine dinucleotide phosphate

**NADPH** : Nicotinamide adenine dinucleotide phosphate

**NBIA** : Neurodegeneration with brain iron

accumulation

NMDA : N-Methyl D-aspartate

NO : Nitric oxide

**NPD1** : NeuroprotectionD1

**PC**: Phosphatidylcholine

**PD** : Parkinson disease

**PE**: phosphatidylethanolamine

**PG** : Prostaglandin

**PL** : Total phospholipids

**PLP** : Pyridoxal phosphate

**PRP** : Proline-rich polypeptide

**PS** : Phosphatidyl serine

**PUFA** : Polyunsaturated fatty acids

**RA** : Retinoic acid

**RALDH** : Retinaldehyde dehydrogenase

**RAR** : Retinoic acid receptor

**RARE** : Retinoic acid response element

**RBP** : Retinol-binding protein

**ROS** : Reactive oxygen species

**RXR** : Retinoid X receptor

**SAM** : S-adenosylmethionine

**SCD** : Subacute combined degeneration

**SOCS** : Suppressors of cytokine signaling

**SOD** : Superoxide dismutase

**STAT** : Signal transducers and activators of

transcription

**STWM** : Short-term/working memory

**SVCT1** : Sodium-dependent vitamin C transporter

types 1 and 2

**T2DM** : Type 2 diabetes mellitus

**TBA-RS** : Thiobarbituric acid-reactive substances

**TH** : T helper

**TMP**: Thiamin monophosphate

**TNF** : Tumour necrosis factor

Trx : Thioredoxin

TT : Tetanus toxoid

**TTP** : Thiamin triphosphate

TTR : Transthyretin

**VDR** : Vitamin D receptor

**VLDL** : Very-low-density lipoproteins

	List of tables	No.
Table 1-1:	Reactions catalysed by nicotinamide nucleotide	18
	coenzymes within the glycolytic	
	pathway/tricarboxylic acid cycle and pentose	
	phosphate pathway of glucose metabolism.	
<b>Table 1-2:</b>	Biogenic amines derived from amino acids or	21
	amino acid derivatives by PLP-dependent	
	nonoxidative decarboxylation.	
<b>Table 1-3:</b>	Copper-Dependent Enzymes and Their Functions	33
	in Animals.	
<b>Table 2-1:</b>	Proposed metabolic mechanisms of vitamin-B12 or folate	70
	disorders	
<b>Table 2-2:</b>	<b>2-2</b> Epidemiological studies and n-3 PUFA.	93
<b>Table 3-1:</b>	Adjusted Risk of Low Cognitive Functioning	111
	(Score , 25 <sup>th</sup> percentile) Associated With Plasma	
	Carotenoids For Each Cognitive Test.	
<b>Table 3-2:</b>	Multivariate-adjusted hazard ratios (HRs) for any	114
	dementia by tertile (T) of plasma tocopherols, $\alpha$ -	
	tocopherylquinone, and 5-nitro-γ-tocopherol per	
	unit of cholesterol.	
Table 3-3:	Odds ratios for PBVL per year over 5 years for	

117

119

	B12 status and homocysteine, folate and white-	
	matter lesions.	
<b>Table 3-5:</b>	Logistic Regression Models Illustrating the Odds	120
	of Cognitive Impairment (95% Confidence	
	Intervals) by Quartiles of Serum [25(OH)D] for	
	1050 Older Noninstitutionalized Persons.	
<b>Table 3-6:</b>	Correlation of neuropsychological examination	123
	scores with copper, markers of neurodegeneration	
	in CSF, and biologic variables of copper	
	metbolism in the serum.	
<b>Table 3-7:</b>	L-Amino acids in the cerebrospinal fluid from the third	128
	ventricle of Alzheimer patients and of control patients.	
<b>Table 4-1:</b>		152
	regading its beneficial effect in AD.	

loss in the highest tertile vs the other two tertiles

by plasma vitamin B12, holoTC, and TC saturation

**Table 3-4:** Spearman correlation between markers of vitamin

levels.