

## The Roles of Ascorbic Acid in the Nervous System

Soleimanitadi S<sup>1</sup>, Saeediniya E<sup>2</sup>, Reaei P<sup>1</sup>, Dalirian A<sup>3</sup>, Dabbagh S<sup>1</sup>, Partoandaz H<sup>1</sup>, Sheikhi B<sup>4</sup>, Nasiri S<sup>5</sup>, Amini P<sup>6</sup>, Naderi M<sup>6</sup> and Esmaeili H<sup>7\*</sup>

<sup>1</sup>Department of Medicine, Iran University of Medical Science, Iran

<sup>2</sup>School of Medicine International Campus Iran University of Medical Science, Iran

<sup>3</sup>Medical Student, Department of Medicine, Iran University of Medical Science, Tehran, Iran

<sup>4</sup>School of Medicine, Iran University of Medical Sciences, Iran

<sup>5</sup>Bachelor of Science in Nursing, Islamic Azad University Semnan Branch, Iran

<sup>6</sup>Medical Student, Tehran Medical Sciences, Islamic Azad University, Iran

<sup>7</sup>Young Researcher and Elite Club, Medical Department, Tehran Medical Sciences, Islamic Azad University, Iran

ISSN: 2637-8078



**\*Corresponding author:** Hossein Esmaeili, Young Researchers and Elite Club, Medical Department, Tehran Medical Sciences, Islamic Azad University, Iran

**Submission:**  July 09, 2021

**Published:**  August 06, 2021

Volume 5 - Issue 2

**How to cite this article:** Soleimanitadi S, Saeediniya E, Reaei P, Dalirian A, Dabbagh S, Partoandaz H, Sheikhi B, Nasiri S, Amini P, Naderi M, Esmaeili H. The Roles of Ascorbic Acid in the Nervous System. Significances of Bioengineering & Biosciences. 5(2). SBB. 000606. 2021. DOI: [10.31031/SBB.2021.05.000606](https://doi.org/10.31031/SBB.2021.05.000606)

**Copyright@** Esmaeili H, This article is distributed under the terms of the Creative Commons Attribution 4.0 International License, which permits unrestricted use and redistribution provided that the original author and source are credited.

### Abstract

However different aspect of vitamin c's affect is not cleared till now, but it is proved that Vitamin C plays several key roles in our body specially during early postnatal period and in natural developing brain specially hippocampus and spatial memory neurons . It must be noticed that it's deficiency also effects memory involved neurons in early life. Some other researches [1] show that Vitamin C regulates transcription of many vital proteins such as collagen notably in sciatic nerves which prevents some sensorimotor disorders.

**Keywords:** Vitamin C; Nervous system; Ascorbic acid

**Abbreviations:** DHA: Dehydroascorbic Acid; AA: Ascorbic Acid; SNS: Sympathetic Nervous System; PTZ: Pentylentetrazol; CA<sub>3</sub>: Cornu Ammonis 3; BH<sub>4</sub>: Tetrahydrobiopterin; BH<sub>2</sub>: Dihydrobiopterin; CA<sub>1</sub>: Cornu Ammonis 1; A<sub>3</sub>R: A<sub>3</sub> Receptors; ROS: Reactive Oxygen Species; SVCT: Sodium±Vitamin C Co-Transporters; AD: Alzheimer Disease; VDR: Vitamin D Receptor

### Vitamin C and Nervous System

Vitamin C (vit C) in body transports by 2 mechanisms: 1) Efficient transport of dehydroascorbic acid (DHA) through glucose transporters. 2) Sodium-dependent co-transport of ascorbic acid (AA) [2]. Vitamin C plays many roles in our body. It increases neurogenesis and can be helpful in prenatal noise stress condition. Prenatal stress increases free radicals [3]. As Vit C is an electron donor, it can protect neurons from free radicals [1,2]. Oxidative stress influence the sympathetic hyperactivity characterizing essential hypertension and contributes to the development of hypertension through activation of the Sympathetic Nervous System (SNS). Vit C also modulates the effects of ethanol and PTZ, which reduce BCL2 protein expression and increase bax and cytochrome c levels and caspase-9 protein and caspase-3 protein (due to increased cytochrome C level) expression, thereby increasing apoptosis is occurred [4]. This condition decreases the number of neurons in all subdivisions of hippocampus . Therefore, the number of axons in CA3 area that is efficient in spatial

memory is reduced. This means Vit C deficiency in early postnatal life impairs spatial memory. It seems that Vit C deficiency in high-risk individuals is harmful for natural developing of brain [5]. Vit C deficiency decreases tetrahydrobiopterin ( $BH_4$ ) and increases dihydrobiopterin ( $BH_2$ )/ $BH_4$  in plasma, but Vit C deficiency has shown no changes in morphology of pyramidal neurons in Cornu Ammonis 1 ( $CA_1$ ) area of hippocamp [6]. Activation of  $A_3R$ , not  $A_1R$  and  $A_2AR$ , activates adenosine receptor and can modulate ascorbate transportation by increasing its diffusion, so intracellular ascorbate will decrease and Reactive Oxygen Species (ROS) will be accumulated in cells [7].  $A_3R$  activation also alters the oxidation balance through SVCT<sub>2</sub> without any change in the production or consumption of ATP neurons [6]. Morphometry significantly reduced myelin thickness but unaffected axon diameters and hydroxyproline levels are unchanged. Collagen transcription is regulated by ascorbic acid in sciatic nerves. the results show that a decrease in the concentration of AA, hypomyelination and a defect in collagen leads to sensorimotor disorders [8].

Vit D protects the brain against the adverse effects Alzheimer Disease (AD). Behavioural effects of adult vit D deficiency are not associated with proliferation or survival of neurons in the adult hippocampus [9]. But Administration of high doses of Vit D without knowing its serum level can cause Hypovitaminosis D and neurological complications. The difference is in the expression of polymorphic variants of VDR gene. Cooperating between astrocyte & neuron make vitamin c being recycled while AA plays a pivotal role in defending which is not used so much by tumor cell contrariwise hippocampus [10]. Healthy people have the ability to modulate sympathetic- mediated peripheral vascular resistance through endothelial synthesis of nitric oxide despite the people who have CHF [11]. The key rule of Vitamin C in protection of DNA, proteins and lipids from oxidative damage [12] Based on some assays it's found that vitamin C has effects on the sensorimotor [13]. It's examined and cleared that Vitamin C by presenting anti-inflammatory mechanism down-regulates the levels of pro-inflammatory mediators [14] Antioxidants such as Vitamin C and N-Acetyl-L- cysteine play a key role in deface damages which are caused by treatment of ethanol [15].

deficiency could be an effective factor which cause embryonic midbrain deficiency of VC and later reduced midbrain-type dopamine [16]. VC deficiency in brain with redox imbalance together impose negative effects on neuronal morphology [17].

## Conclusion

In this paper we have studied on the Vit C transportation, its effect in hypertension and some proteins expression and its deficiency affects on human body. We understood that accumulation of reactive oxygen species could be occurred in cell during diffusion of Vitamin C. Vitamin C prevents hypertension and sensorimotor disorders; Vitamin C also could regulates apoptosis by having effect on protein transcription. Its deficiency also attracted our attention that could be so harmful for natural brain developing in high-risk groups specially in postnatal period. During our study about the

vitamin c roles in CNS, Vitamin D protects the brain against the adverse effects Alzheimer Disease; however must pay attention to its consumption dose in order to prevent Hypovitaminosis.

## References

- Huff TC, Sant DW, Camarena V, Booven VD, Andrade NS, et al. (2021) Vitamin C regulates Schwann cell myelination by promoting DNA demethylation of pro-myelinating genes. *Journal of neurochemistry* 157(6): 1759-1773.
- Evayanti LG, Viskasari PK, Abdurachman (2018) The effect of Vitamin C on the cerebral cortex neurons of rats exposed by prenatal noise stress. *Asian Journal of Microbiology, Biotechnology & Environmental Sciences Paper* 20: 34-37.
- Shankardass K, McConnell R, Jerrett M, Milam J, Richardson J, et al. (2009) Parental stress increases the effect of traffic-related air pollution on childhood asthma incidence. *Proceedings of the National Academy of Sciences USA* 106(30): 12406-12411.
- Naseer MI, Ullah N, Ullah I, Koh PO, Lee HY, et al. (2011) Vitamin C protects against ethanol and PTZ-Induced apoptotic neurodegeneration in prenatal rat hippocampal neurons. *Synapse* 65(7): 562-571.
- Tveden-Nyborg P, Johansen LK, Raida Z, Villumsen CK, Larsen JO, et al. (2009) Vitamin C deficiency in early postnatal life impairs spatial memory and reduces the number of hippocampal neurons in guinea pigs. *The American journal of clinical nutrition* 90(3): 540-546.
- Camila CP, Encarnação T, Sagrillo MA, Pereira MR, Relvas JB, et al. (2021) Activation of adenosine  $A_3$  receptors regulates Vitamin C transport and redox balance in neurons. *Free Radical Biology and Medicine* 163: 43-55.
- Parsons HT, Fry SC (2010) Reactive oxygen species-induced release of intracellular ascorbate in plant cell-suspension cultures and evidence for pulsing of net release rate. *New Phytologist* 187(2): 332-342.
- Schultheis PJ, Fleming SM, Clippinger AK, Lewis J, Tsunemi T, et al. (2013) Atp13a2-deficient mice exhibit neuronal ceroid lipofuscinosis, limited  $\alpha$ -synuclein accumulation and age-dependent sensorimotor deficits. *Human Molecular Genetics* 22(10): 2067-2082.
- Groves NJ, Bradford D, Sullivan RKP, Kyna-Anne C, Aljelaify RF, et al. (2016) Behavioural effects of adult Vitamin D deficiency in balb/c mice are not associated with proliferation or survival of neurons in the adult hippocampus. *PLoS One* 11(4).
- Astuya Allisson, Caprile T, Castro M, Salazar K, García MA, et al. (2005) Vitamin C uptake and recycling among normal and tumor cells from the central nervous system. *Journal of neuroscience research* 79(1-2): 146-156.
- Gianfranco P, Raffaele Q, Fimognari F, Moisè A, Mario M, et al. (2004) Influence of L-arginine and vitamin C on the autonomic nervous system in chronic heart failure secondary to ischemic cardiomyopathy. *The American journal of cardiology* 93(5): 650-654.
- EFSA Panel on Dietetic Products, Nutrition and Allergies (NDA) (2009) Scientific opinion on the substantiation of health claims related to vitamin C and protection of DNA, proteins and lipids from oxidative damage (ID 129, 138, 143, 148), antioxidant function of lutein (ID 146), maintenance of vision (ID 141, 142), collagen formation (ID 130, 131, 136, 137, 149), function of the nervous system (ID 133), function of the immune system (ID 134), function of the immune system during and after extreme physical exercise (ID 144), non-haem iron absorption (ID 132, 147), energy -yielding metabolism (ID 135), and relief in case of irritation in the upper respiratory tract (ID 1714, 1715) pursuant to Article 13(1) of Regulation (EC) No 1924/2006. *EFSA Journal* 7(10): 1226.
- Burkhard G, Röhr D, Fledrich R, Sereda MW, Kleffner I, et al. (2011) Sodium-dependent vitamin C transporter 2 deficiency causes hypomyelination and extracellular matrix defects in the peripheral nervous system. *Journal of Neuroscience* 31(47): 17180-17192.

14. Wei-Guo W, Xiu RJ, Xu Z, Yin Y, Feng Y, et al. (2015) Protective effects of Vitamin C against spinal cord injury- induced renal damage through suppression of NF- $\kappa$ B and proinflammatory cytokines. *Neurological Sciences* 36(4): 521-526.
15. Yang Su, Jian Mao, Runhong Lei, Rui Wang, Fankai Lin, et al. (2013) Vitamin C and N-Acetyl-L-cysteine prevent ethanol induced cultured cerebellar granule neurons apoptosis through nuclear factor-kappa B pathway. 2013 ICME International Conference on Complex Medical Engineering.
16. Xi-Biao H, Mirang Kim, Seon-Young K, Sang-Hoon Y, Yong-Hee R, et al. (2015) Vitamin C facilitates dopamine neuron differentiation in fetal midbrain through TET 1- and JMJD 3-dependent epigenetic control manner. *Stem cells* 33(4): 1320-1332.
17. Hansen SN, Jørgensen JMB, Nyengaard JR, Lykkesfeldt J, Tveden-Nyborg P (2018) Early life Vitamin C deficiency does not alter morphology of hippocampal CA<sub>1</sub> pyramidal neurons or markers of synaptic plasticity in a guinea pig model. *Nutrients* 10(6): 749.

For possible submissions Click below:

[Submit Article](#)