

Investigation of Coenzyme Q10 As An Anti-Aging Supplement

Sarah Dunn

Beloit College, Beloit, Wisconsin

Abstract

Coenzyme Q10 (CoQ10) is recognized as an important coenzyme, or semi-vitamin, that functions in many capacities in aerobic organisms. Of specific importance is CoQ10's function as an electron carrier in the electron transport chain in mitochondria, which is responsible for the production of ATP. In that process, CoQ10 contributes to the translocation of protons from the mitochondrial matrix to the inter-membrane space, thereby creating a proton gradient, which is the crux of ATP generation. Consumption of CoQ10 in supplements has become a growing trend meant to enhance bioenergetic capacity and mitigate adverse effects of aging related to certain pathological conditions. I hypothesize that exogenous intake of CoQ10 through a CoQ10 supplement will be ineffective against aging and may have hazardous side effects. Preliminary results suggest that CoQ10 does not mitigate aging, and is damaging in high dosages.

Introduction

CoQ is a lipophilic, redox molecule located within the phospholipid bilayer of cellular membranes (4). In mitochondria, CoQ10 is located within the inner membrane and functions to drive the creation of the transmembrane proton gradient, which is the crux of aerobic respiration and drives ATP generation (4,6,8) CoQ10 can exist in three stable oxidation states, which is important for its biosynthesis and its role in aerobic respiration. These states include the fully reduced ubiquinol form (CoQ10H₂), the radical semiquinone intermediate (CoQ10H[•]), and the fully oxidized ubiquinone form (CoQ10) (3,6,7). While the CoQ intermediate (CoQ10H[•]) is a requisite for aerobic respiration, it is also the primary source of mitochondrial superoxide anion radical production, which promulgates oxidative damage and thus aging (4,6,7,8). Because CoQ10 functions as both an essential component of the electron transport chain and as the primary source of mitochondrial radical production, it is unclear whether increased exogenous intake will prolong or curtail lifespan (4).

References

- (1) Marieb E, Katja H. *Human Anatomy & Physiology*. 7th ed. Benjamin Cummings, 2006.
- (2) Crane FL, Hafezi Y, Lester RI, Widmer C. Isolation of a quinone from beef heart mitochondria. *Biochimica et Biophys. Acta* 1957;25:220-221.
- (3) Micronutrient Information Center for Coenzyme Q10. Linus Pauling Institute, Micronutrient Research for Optimum Health, 2003 [cited 2009 Oct 20]. Available from: <http://lpi.oregonstate.edu/info/center/othernuts/coq10/#deficiency>.
- (4) Sumien N, Heinrich KR, Shetty R, Sohal RS, Forster MJ. Prolonged intake of coenzyme Q10 impairs cognitive functions in mice. *J. Nutr.* 2009;139:1926-1932
- (5) Guidance for Industry Estimating the maximum safe starting dose in initial clinical trials for therapeutics in adult healthy volunteers. U.S. Department of Health and Human Services FDA, Center for Drug Evaluation and Research, 2005 [cited 2009 Oct 10]. Available from: <http://www.fda.gov/downloads/Drug/GuidanceComplianceRegulatoryInformation/Guidances/UCM078932.pdf>.
- (6) Chance B, Sies H, Boveris A. Hydroperoxide metabolism in mammalian organs. *Physiol Rev.* 1979;59:527-605.
- (7) Turrens JF, Alexandre A, Lehninger AL. Ubisemiquinone is the electron donor for superoxide formation by complex III of heart mitochondria. *Arch Biochem Biophys.* 1985;237:408-414.
- (8) Ernster L, Dallner G. Biochemical, physiological and medical aspects of ubiquinone function. *Biochim. Biophys. Acta.* 1995;1271:195-204.
- (9) Matthews R, Yang L, Browne S, Baik M, Beal MF. Coenzyme Q10 administration increases brain mitochondrial concentrations and exerts neuroprotective effects. *PNAS.* 1998;95:8892-8897.
- (10) Zhang Y, Aberg F, Appelkvist E, Dallner G, Ernster L. Uptake of dietary coenzyme Q supplement is limited in rats. *J. Nutr.* 1995;125:446-453.
- (11) Niklowitz P, Sonnenschein A, Janetzky B, Andler W, Menke T. Enrichment of coenzyme Q10 in plasma and blood cells: defense against oxidative damage. *Int. J. Biol. Sci.* 2007;3:257-262.
- (12) Yan J, Fujie Y, Kishida H, Hosoe K, Sawashita J, Takeda T, Mori M, Higuchi, K. Reduced coenzyme Q10 supplementation decelerates senescence in SAMP1 mice. *Exp. Gerontology.* 2006;41:130-140.
- (13) Morisco C, Trimarco B, Condonelli M. Effect of coenzyme Q10 therapy in patients with congestive heart failure: a long-term, multicenter randomised study. *Clin Investig.* 1993;71:134-136.
- (14) Mueller T, Buettner T, Gholipour AF, Kuhn W. Coenzyme Q10 supplementation provides mild symptomatic benefit in patients with Parkinson's disease. *Neurosci Lett.* 2003;341:201-204.
- (15) Shults CW, Haas RH, Beal MF. A possible role of coenzyme Q10 in the etiology and treatment of Parkinson's disease. *BioFactors.* 1999;9:267-272.
- (16) Hofman-Bang C, Rehnquist N, Swedberg K, Wiklund I, Astrom H. Coenzyme Q10 as an adjunctive in the treatment of chronic congestive heart failure. The Q10 study group. *J Card Fail.* 1995;1:101-107.
- (17) Ferrante RJ, Andressen OA, Dedeoglu A, et al. Therapeutic effects of coenzyme Q10 and remacemide in transgenic mouse models of Huntington's disease. *J Neurosci.* 2002;22:1592-1599.

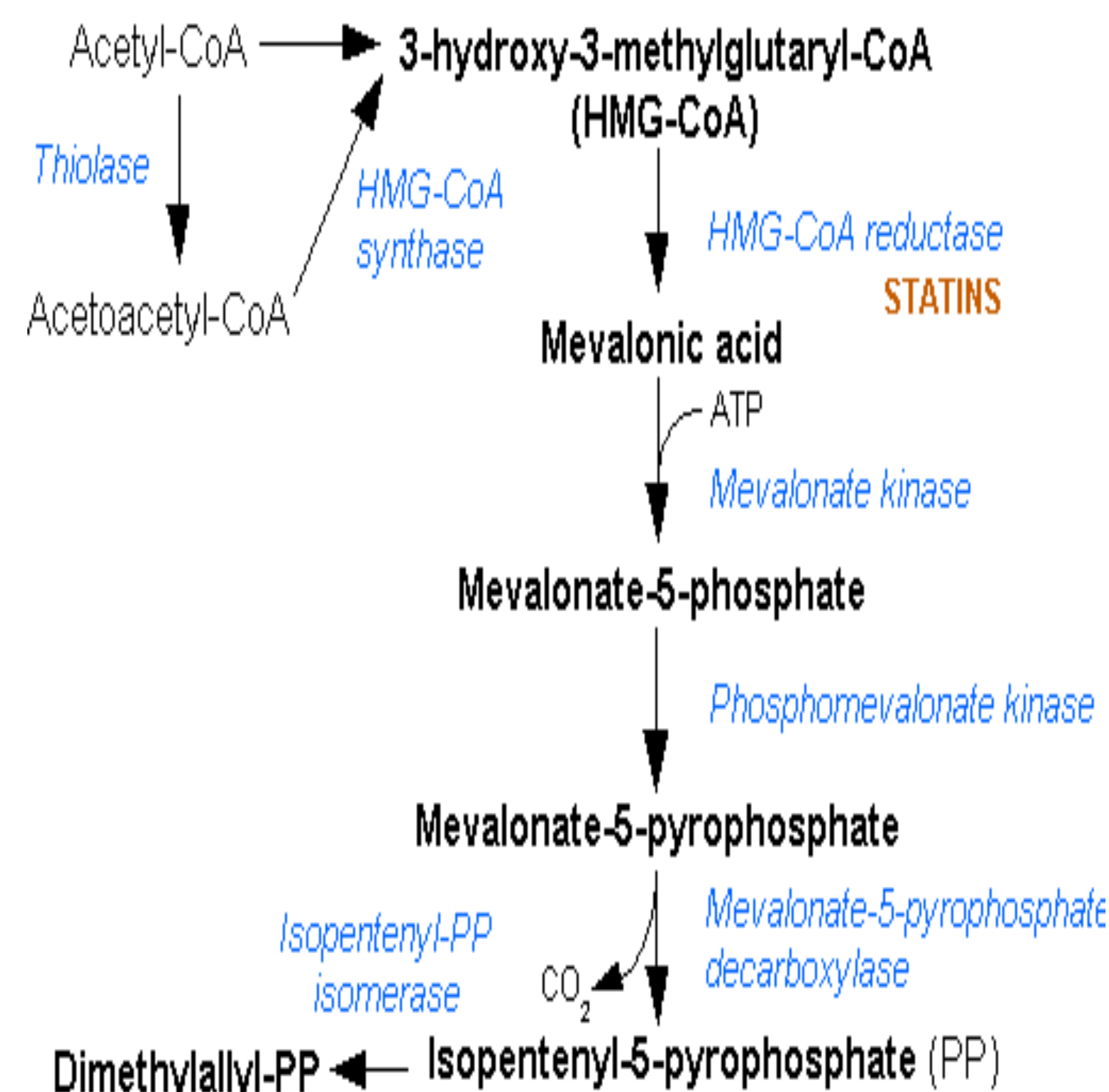


Figure 1: Mevalonate Pathway; Biosynthesis of CoQ. Mevalonate is a precursor to isopentenyl pyrophosphate, which combines with its isomer, dimethylallyl pyrophosphate, in repeating alternations to form isoprene (or polyprenyl) chains.

http://en.wikipedia.org/wiki/File:Mevalonate_pathway.png

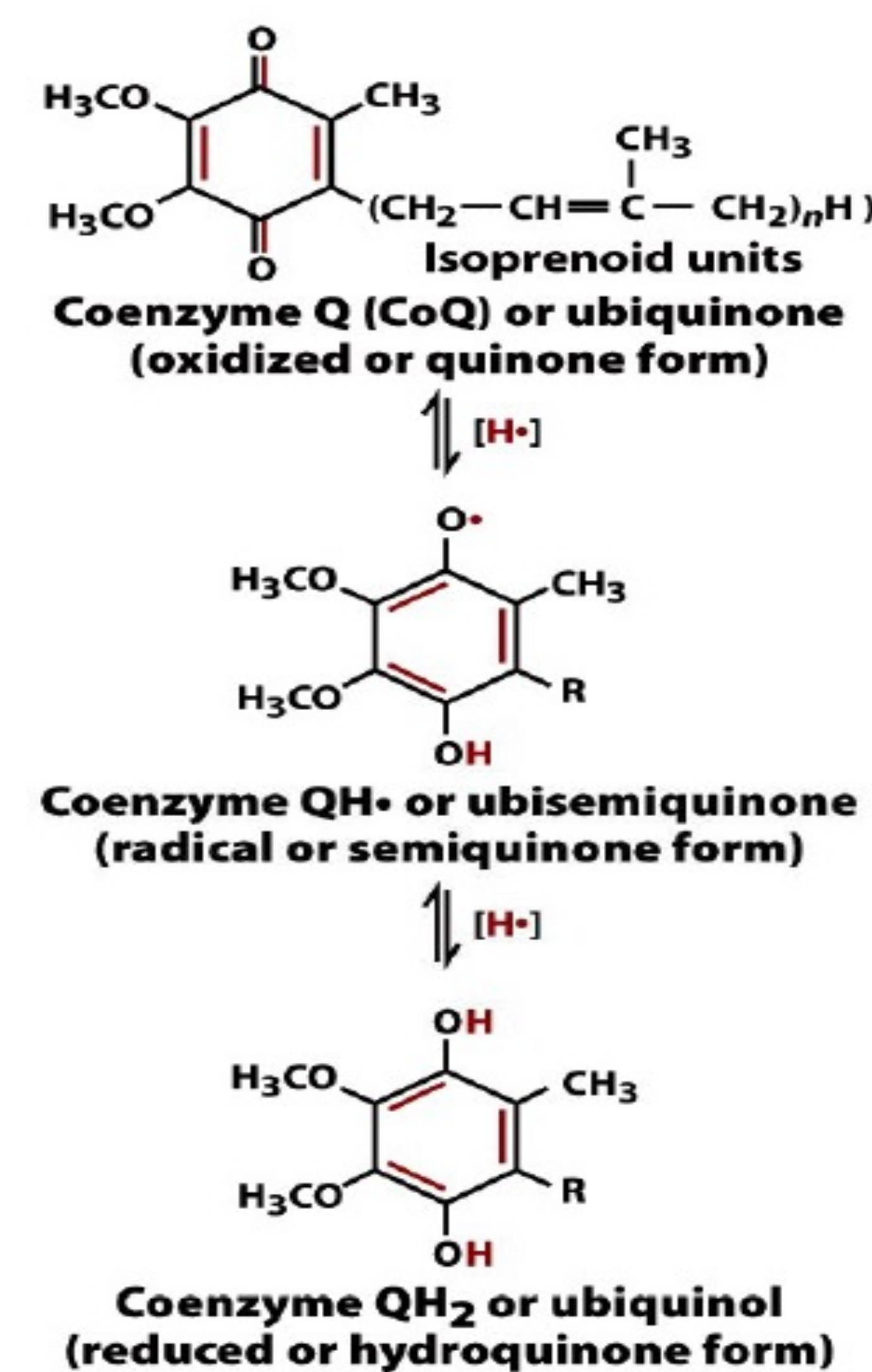


Figure 2: CoQ10 oxidation states. This is how CoQ acts as an electron carrier in the ETC transferring a high energy electron.

<http://biology.ucsd.edu/classes/bibc102.SU2.08/objects/ch05-OxPhos.pdf>

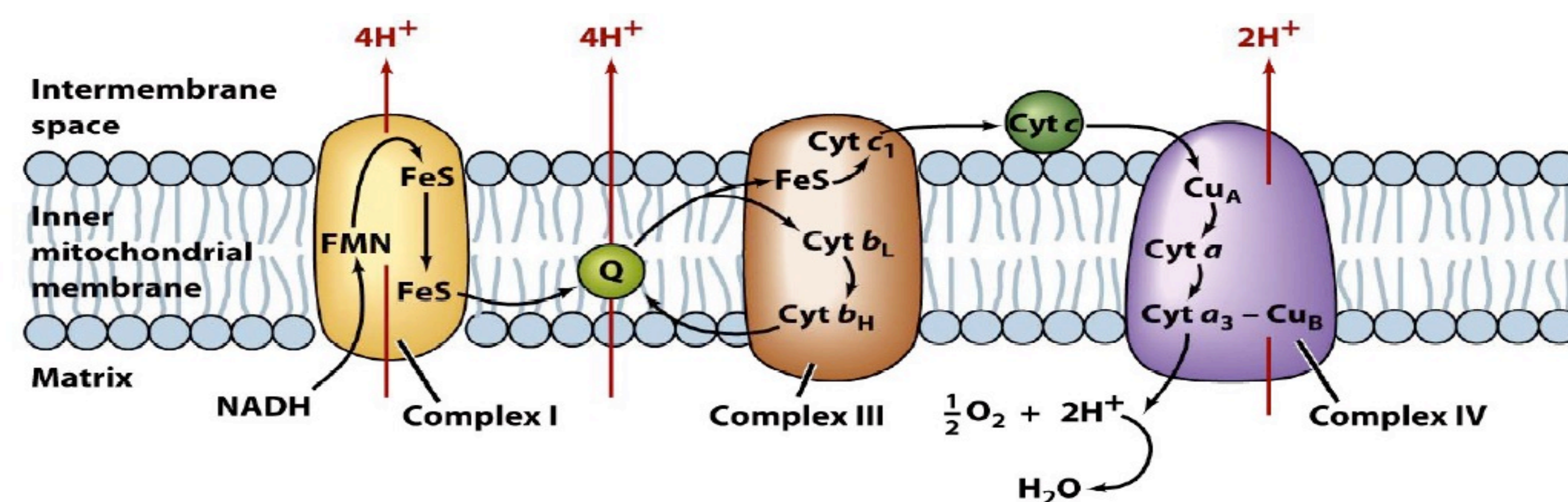


Figure 3: Electron Transport Chain (ETC)

The ETC is a series of linked electron carriers in mitochondria, leading to the eventual reduction of O₂ to H₂O and the creation of an electrochemical proton gradient, which is utilized in oxidative phosphorylation to synthesize stored energy as ATP. Complex I passes electrons from NADH to CoQ, Complex II passes electrons from succinate to CoQ, CoQ passes its electrons to cytochrome c in complex III and then water is reduced in Complex IV.

<http://biology.ucsd.edu/classes/bibc102.SU2.08/objects/ch05-OxPhos.pdf>

Methods

A literature review of peer-reviewed journal articles was conducted.

Results

Early evidence suggests that there are possible benefits of CoQ10 in ameliorating some disease states. Various diseases including Huntington's and Parkinson's involve decreased concentrations of CoQ10, and early clinical trials in humans with these diseases suggest that a low dosage CoQ supplementation may be beneficial (3, 4, 11,13-17). It is not known, however, if the decreased concentrations are a cause or effect of the disease (11). Furthermore, a mouse model was used to determine adverse effects of CoQ10 supplementation and found that at high dosage, 2.6 mg/g, cognitive and sensory impairments encountered in old mice were exacerbated (4). Additionally, supplements of CoQ10 in both high and low dosages, 0.68 mg/g and 2.0 mg/g, had no significant effect on survival (p>.565) (4). There is no evidence to suggest a direct correlation between CoQ intake and prolonged life span (4,8,9,11,12).

Discussion

From the preliminary data I would suggest that a healthy person should not take CoQ10 supplements. The assertion that the supplements delay aging is unsubstantiated and safe and effective dosages have not been established in a human model. Furthermore, the adverse effects of a high dosage outweigh the possible benefits.

Conclusion

CoQ10 supplements are new and have had little clinical evaluation. Research conducted thus far has had small sample sizes or were based on mouse models. Therefore, CoQ10 supplements have not been shown to be safe nor proven beneficial. A health provider should be advised when an individual is considering taking a CoQ10 supplement, and a risk benefit analysis should be undertaken.