Emission of Mitochondrial Biophotons and their Effect on Electrical Activity of Membrane via Microtubules

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Abstract

In this paper we argue that, in addition to electrical and chemical signals propagating in the neurons of the brain, signal propagation takes place in the form of biophoton production. This statement is supported by recent experimental confirmation of photon guiding properties of a single neuron. We have investigated the interaction of mitochondrial biophotons with microtubules from a quantum mechanical point of view. Our theoretical analysis indicates that the interaction of biophotons and microtubules causes transitions/fluctuations of microtubules between coherent and incoherent states. A significant relationship between the fluctuation function of microtubules and alpha-EEG diagrams is elaborated on in this paper. We argue that the role of biophotons in the brain merits special attention.

Keywords: mitochondrial biophoton, microtubule (MT), coherence, fluctuation function

1. Introduction

All living cells of plants, animals and humans continuously emit ultraweak biophotons (ultraweak electromagnetic waves) in the optical range of the spectrum, which is associated with their physiological states and can be measured using special equipment¹. Neural cells

¹ Updated from: <u>http://www.transpersonal.de/mbischof/englisch/webbookeng.htm</u>, 23 November 2010.

also continuously emit biophotons. The intensity of biophotons is in direct correlation with neural activity, cerebral energy metabolism, EEG activity, cerebral blood flow and oxidative processes [37,50].

According to Van Wijk et al. [96], there are significant correlations between the fluctuations in biophoton emission and fluctuations in the strength of electrical alpha wave production in the brain. Some unpublished observations suggest that the state of the biophoton field of a person may be connected to the state of the brain as measured by the EEG (e.g., degree of synchronization and coherence) [9]. Certain meditative states characterized by a high degree of coherence in the EEG may well be accompanied by a high degree of coherence in the biophoton field [9], although measurements correlating the coherence of the biophoton field and the EEG readings have not been made yet.

Here, we investigate theoretically the interaction of biomolecules with biophotons taking place within the neurons of the brain. We have adopted a quantum mechanical formalism in an attempt to quantitatively investigate possible connections between the EEG and the biophoton production.

2. Biophoton Production Mechanism inside Neurons

During natural metabolic processes taking place in diverse living organism, permanent and spontaneous ultraweak biophoton emission has been observed without any external excitation [1,9,29,36,37,49,50,54,66,70,77,83,89,96,97,107]. The emergence of biophotons is due to the bioluminescent radical and non-radical reactions of Reactive Oxygen Species (ROS) and Reactive Nitrogen Species (RNS), and involves simple cessation of excited states [52,61,102]. The main source of biophotons derives from the oxidative metabolism of mitochondria [94].

Neurons also incessantly emit biophotons [37,50]. Biophoton emission from neural tissue depends on the neuronal membrane depolarization and Ca^{2+} entry into the cells [44]. This biophoton emission can be facilitated by the membrane depolarization of neurons by a high concentration of K⁺ and can be attenuated by application of tetrodotoxin or removal of extracellular Ca^{2+} [44]. Recently, Sun et al. [82] demonstrated that neurons can conduct photon signals. Moreover, Wang et al. [100] presented the first experimental proof of the existence of spontaneous and visible light induced biophoton emission form freshly isolated rat's whole eye, lens, vitreous humor and retina. They proposed that retinal phosphenes may originate from natural bioluminescent biophotons within the eyes [10,100]. However, the retina is part of the central nervous system. Recently, Bókkon suggested that biophysical pictures may emerge due to redox regulated biophotons in retinotopically organized cytochrome oxidase-rich neural networks during visual perception and imagery within early visual areas [12]. It seems that bioelectronic and biophotonic processes are not independent biological events in the nervous system. Therefore, we conclude that biophoton emission within neurons can be directly correlated with biochemical processes.

However, the term ultraweak bioluminescence (ultraweak biophoton emission) can be misleading, because it may suggest that biophotons are not important for cellular processes. Estimates indicate that for a measured intensity of biophotons, the corresponding intensity of the light field within the organism can be up to two orders of magnitude higher [15,80].

According to Bókkon et al. [11], the real biophoton intensity within cells and neurons can be considerably higher than one would expect from the measurement of ultraweak bioluminescence, which is generally carried out macroscopically several centimeters away from the tissue or cell culture [86]. Moreover, the most significant fraction of natural biophoton intensity cannot be accurately measured because it is absorbed during cellular processes.

Numerous findings have provided evidence of fundamental signal roles of ROS and RNS in cellular processes under physiological conditions [13,14,20,27,33,35,92,95]. There are experimental indications that ROS and RNS are also necessary for synaptic processes and normal brain functions. Free radicals and their derivatives act as signaling molecules in cerebral circulation and are necessary for molecular signaling processes in the brain such as synaptic plasticity, neurotransmitter release, hippocampal long-term potentiation, memory formation, etc. [47,48,85,87,88,99]. Because the generation of ROS and RNS is not a haphazard process, but rather a coordinated mechanism used in signaling pathways, biophoton emission may not be a byproduct of biochemical processes but it can be linked to precise signaling pathways of ROS and RNS. Consequently, regulated generation of ROS and RNS can also produce regulated biophotons within cells and neurons. This means that regulated electrical (redox) signals (*spike-related electrical signals along classical axonal-dendritic pathways*) of neurons can be converted into biophoton signals by various bioluminescent reactions.

Biophotons can be absorbed by natural chromophores such as porphyrin rings, flavinic, pyridinic rings, lipid chromophores, aromatic amino acids, etc. [43,45,57,86]. Mitochondrial electron transport chains contain several chromophores, among which cytochrome oxidase enzymes are most prominent [43,45]. The absorption of biophotons by a photosensitive molecule can produce an electronically excited state. As a result, molecules in electronically excited states often have very different chemical and physical properties compared to their electronic ground states. Regulated biophotons are not dissipated in random manner within cells and neurons, but are absorbed - close to the place where they originated - by chromophores and can excite nearby molecules and trigger/regulate complex signal processes [17]. Thus, absorbed biophotons could have effects on electrical activity of cells and neurons via signal processes.

3. Interaction of Mitochondrial Biophotons with Microtubules (MTs)

3.1 MTs and Mitochondria

Microtubular structures have been implicated as playing an important role in the signal and information processing taking place in the human (and possibly animal) brain [32,41,42,55,56]. Vertebrate neurons show typically filamentous mitochondria associated with the microtubules (MTs) of the cytoskeleton, forming together a continuous network (mitochondrial reticulum) [79]. The rapid movements of mitochondria are MT-based and the slower movements are actin-based [91]. MT formation can be regulated by redox-dependent phosphorylations and Ca^{2+} signals. Since the rapid movements of mitochondria are MT-based, mitochondrial trafficking can be organized by redox and Ca^{2+} -dependent MT regulation. Moreover, the refractive index of both mitochondria and MTs is higher than the surrounding cytoplasm [86]. Consequently, both the mitochondria and the MTs could act as optical waveguides, i.e. electromagnetic radiation can propagate within their networks [21,42,86]. Associated mitochondrial and MT networks may act as redox and Ca^{2+} regulated organic quantum optical-like fiber systems in neurons. MTs are composed of tubulin dimers.

Tubulin dimer is an intrinsically fluorescent molecule mainly due to 8 tryptophan residues it contains, as can be seen e.g. in the 1TUB structure from the Protein Data Bank [62]. It is well known that the absorption (ca. 280 nm) and fluorescence (ca. 335 nm) wavelength (and intensity) of tryptophan is dependent on the conformation of tubulin. Probing absorption and fluorescence of tubulin is a standard method to determine the polymerization state of MTs. This can be considered one of possible qualitative connections between the fluctuations of MT growth and its corresponding biophoton absorption and emission characteristics. Additionally, there exist other energy states (of both optical and vibrational nature [19,40,65]) which tubulin dimers and the whole MT can support. These states can be excited by energy supply provided by mitochondria [16]. Furthermore, MT polymerization is sensitive to UV [84] and blue light [59] and mitochondria are known to be sources of biophotons corresponding to the same wavelengths [8,34,98], which makes an immediate logical connection. Figure 1 illustrates how mitochondria emit light into MTs.

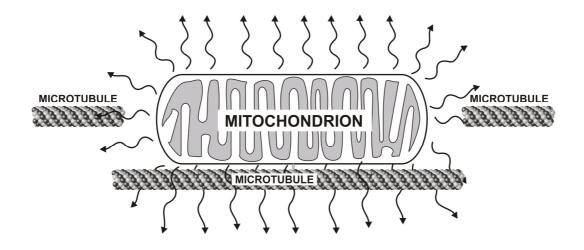


Figure 1 Representation of biophotons produced by mitochondria and the interaction of biophotons with microtubules.

3.2 Interaction of Biophotons with MTs

It is worth stressing here that centrioles and cilia, which are complex microtubular structures, are involved in photoreceptor functions in single cell organisms and primitive visual systems. Cilia are also found in all retinal rod and cone cells. The dimensions of centrioles and cilia are comparable to the wavelengths of visible and infrared light [31]. In a series of studies spanning a period of some 25 years G Albrecht-Buehler (AB) demonstrated that living cells possess a spatial orientation mechanism located in the centriole [2,3,4]. This is based on an intricate arrangement of MT filaments in two sets of nine triplets each of which is perpendicular to the other. This arrangement provides the cell with a primitive "eye" that allows it to locate the position of other cells within a two to three degree accuracy in the azimuthal plane and with respect to the axis perpendicular to it [2]. He further showed that electromagnetic signals are the triggers for the cells' repositioning. It is still largely a mystery how the reception of electromagnetic radiation is accomplished by the centriole. Another

mystery related to these observations is the original electromagnetic radiation emitted by a living cell [94].

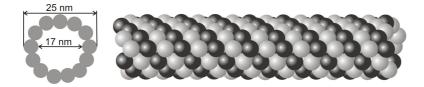


Figure 2 MTs are hollow cylinders composed of protein units called tubulin. The inner diameter of an MT is 17nm and the outer diameter is 25 nm. The lengths of MTs vary widely from nanometers to micrometers. MTs have been considered to act as QED-cavities [55, 56].

Earlier, MTs have been considered as optical cavities [55] with quantum properties [56], capable of supporting only a single mode [41] or perhaps a few widely spaced (in the frequency domain) modes. Our approach is based on a fully quantum mechanical formalism of the Jaynes-Cummings model [39]. MTs are biological hollow cylinders with a 17 nm inner diameter and a 25 nm outer diameter [16], composed of units called tubulin dimers, each of which has the dimensions $4nm \times 8nm \times 6nm$ [55]. Tubulin can be viewed as a typical *two-state* quantum mechanical system, where the dimers couple to conformational changes with $10^{-9} - 10^{-11}$ sec transitions due to electron transitions in hydrophobic pockets, corresponding to an angular frequency in the range $\omega_0 \sim O(10^{10}) - O(10^{12}) Hz$ [55]. Using a first-order-approximation estimate of the quality factor for the MT cavities (i.e. Q_{MT}), it has been found that $Q_{MT} \sim O(10^8)$ [55]. High-quality cavities encountered in Rydberg atom experiments dissipate energy on time scales of $O(10^{-3}) - O(10^{-4})$ sec and have Q's which are comparable to Q_{MT} [55]. We consider the tubulin dimer to represent a two-state system with ground $|g\rangle$ and excited $|e\rangle$ states, respectively. Now, we assume that tubulin interacts with a single-mode cavity field of biophotons (the coherent nature of biophotons will be discussed in section 3.4). Here, we introduce the tubulin transition operators $\hat{\sigma}_{_+}$, $\hat{\sigma}_{_-}$ and $\hat{\sigma}_F$, where $\hat{\sigma}_+ = |e\rangle\langle g|$ is an operator which takes tubulin into the excited state, the operator $\hat{\sigma}_{-} = |g\rangle\langle e| = \hat{\sigma}_{+}^{\dagger}$ takes tubulin into the ground state, and the fluctuation operator $\hat{\sigma}_F = |e\rangle \langle e| - |g\rangle \langle g|$ causes transitions between excited and ground states. We have

$$\hat{\sigma}_{+} |g\rangle = |e\rangle \qquad \hat{\sigma}_{+} |e\rangle = 0 \hat{\sigma}_{-} |g\rangle = 0 \qquad \hat{\sigma}_{-} |e\rangle = |g\rangle$$

$$(1)$$

Frequencies of visible light are on the order of THz, and as explained before, Wang et al, [100] have detected visible light in the brain as biophotons. Also, transition frequencies in tubulins are on the order of THz [55], so, the interaction between a two-state system (here represented by tubulin) and a single mode quantized field (here represented by biophotons) is given by the total Hamiltonian in the approximation $O(\omega_0) \sim O(\omega)$ [26] as:

$$\hat{H} = \hat{H}_{tubulin} + \hat{H}_{Biophotons} + \hat{H}_{Interation} = \frac{1}{2}\hbar\omega_0\hat{\sigma}_F + \hbar\omega\hat{a}^{\dagger}\hat{a} + \hbar\lambda(\hat{\sigma}_+\hat{a} + \hat{\sigma}_-\hat{a}^{\dagger})$$
(2)

where $\hat{H}_{tubulin}$ is the Hamiltonian operator for tubulin, $\hat{H}_{Biophotons}$ is the Hamiltonian operator for biophotons and $\hat{H}_{Interation}$ is the Hamiltonian operator representing the interaction between tubulin and biophotons. Here, ω_0 is the frequency of tubulin transitions and ω is the frequency of biophotons. *a* and a^{\dagger} are annihilation and creation operators, respectively.

$$\hat{a}|n\rangle = \sqrt{n}|n-1\rangle$$

$$\hat{a}^{\dagger}|n\rangle = \sqrt{n+1}|n+1\rangle$$
(3)

where ket $|n\rangle$ is the number state of photons. In Hamiltonian (2) the quantity λ is defined as $\lambda = \frac{dg}{\hbar}$ where *d* is the dipole moment of tubulin, $\hbar = \frac{h}{2\pi}$ where *h* is Planck's constant and $g \approx \frac{1}{2}\sqrt{\frac{\hbar\omega}{\varepsilon V}}$ where ε is the dielectric constant of the environment in an MT where $\frac{\varepsilon}{\varepsilon_0} \sim 80$, in which ε_0 is dielectric constant of vacuum [55], and *V* is the volume of an MT. Here, the state vector of tubulin is $|tubulin\rangle = c_g(t)|g\rangle + c_e(t)|e\rangle$ where $c_g(t)$ and $c_e(t)$ are time-dependent coefficients in which *t* represents time. The state of the field is $|Biophotons\rangle = \sum_{n=0}^{\infty} c_n |n\rangle$ where c_n is a constant coefficient and $|n\rangle$ is the number state of photons. The total state is the tensor product $|\psi(t)\rangle = |tubulin\rangle \otimes |Biophotons\rangle$. Inserting the total state into the Schrödinger's equation $i\hbar \frac{d}{dt}|\psi(t)\rangle = \hat{H}_{Interaction}|\psi(t)\rangle$, the result is readily obtained as [26]

$$\rightarrow |\psi(t)\rangle = \sum_{n=0}^{\infty} \frac{\left\{ \left[c_e c_n \cos(\lambda t \sqrt{n+1}) - i c_g c_{n+1} \sin(\lambda t \sqrt{n+1}] |e\rangle |n\rangle + \left[-i c_e c_{n-1} \sin(\lambda t \sqrt{n}) + i c_g c_n \cos(\lambda t \sqrt{n}) \right] |g\rangle |n\rangle \right\}$$

$$(4)$$

3.3 Coherence and Decoherence Problem for the system of MT

The Wu-Austin Hamiltonian has been used to describe the interaction of quantized electromagnetic field with MTs to yield a coherent Froehlich's state of the dipolar biological system. Wu and Austin [104,105,106] proposed a dynamical model containing a biological system composed of electric dipoles with N modes connected to harmonic heat baths representing a quantized electromagnetic source and the surrounding thermal-relaxation bath. The interaction of quantized field with the system of electric dipoles gives a Froehlich coherent state. We believe that the system of neuronal MTs is a good candidate for being properly described by the above Hamiltonian. MTs are composed of tubulin dimers which can be considered as biological electric dipoles.

Previously, one of the concerns of considering coherent states in the brain was due to the fact that the Bose-Einstein condensation happens only at low enough temperatures believed to be lower than body temperature. Recently, Reimers et al. [27] have argued that a very fragile Froehlich coherent state may paradoxically only emerge at very high temperatures and thus there is no possibility for the existence of Froehlich coherent states in biological systems, so quantum models based on the Froehlich coherence should be ruled out. However, it has been shown that there are serious problems in the calculations made by Reimers et al [27] and consequently their conclusions appear to be flawed [75].

Another important problem that remains when considering coherent states for MTs is the so-called *decoherence* problem. The question is "how is it possible for MTs to be in a coherent state while the environment surrounding them is relatively hot, wet and noisy?" Although evidence was found that quantum spin transfer between quantum dots connected by benzene rings (the same structures found in aromatic hydrophobic amino acids) is more efficient at relatively warm temperatures than at absolute zero [63], Tegmark [84] calculated decoherence times for MTs based on the collisions of ions with MTs leading to the decoherence times on the order of:

$$\tau = \frac{D^2 \sqrt{mkT}}{Ngq^2} \approx 10^{-13} s \tag{5}$$

where *D* is the tubulin diameter, *m* the mass of the ion, *k* Boltzmann's constant, *T* absolute temperature, *N* the number of elementary charges in the MT interacting system, $g = \frac{1}{4\pi\varepsilon_0}$

the Coulomb constant and q the charge of an electron. Hagan et al. showed that Tegmark used wrong assumptions for his investigation of MTs. Another main objection to the estimate in (5) is that Tegmark's formulation yields decoherence times that increase with temperature contrary to a well-established physical intuition and the observed behavior of quantum coherent states [74]). In view of these (and other) problems in Tegmark's estimates, Hagan et al. [30] asserted that the values of quantities in Tegmark's relation are not correct and thus the decoherence time should be approximately 10^{10} times greater. According to Hagan et al. [30], MTs in neurons of the brain can process information quantum mechanically and they could avoid decoherence via several mechanisms over sufficiently long times for quantum processing to occur. As a result, we conclude that coherent states in MTs are still theoretically possible. Below, we explore the consequences of this conclusion.

3.4 Fluctuation Function and Simulation

We begin by representing the state of the MTs as a superposition of the coherent and incoherent (ground) states, and the state of the biophotons as a field composed of n photons. In our approach, transitions involving the coherent-incoherent process determined by a function called the fluctuation function, F(t).

As we explained before, we assume two states for a MT: (a) the ground state (incoherent without energy pumping), and (b) an excited state (coherent with energy pumping). We reformulate our previous calculations for the system of MTs and investigate its interaction with biophotons. First, prior to the interaction, MTs may be in one of the two

states: the ground state or the coherent state.

The state of MTs can be written in the form of a superposition of the ground state and coherent state. MTs vibrate in different frequency modes before the energy pumping occurs. This state can be written in the form $|n_1\rangle|n_2\rangle|n_3\rangle....\equiv |n_1,n_2,n_3,...\rangle = |\{n_j\}\rangle$ in which each $|n_i\rangle$ is a special frequency mode state of a tubulin in an MT. When a quantized electromagnetic field is pumped into the MT, according to Froehlich's theory [22,23,24], this leads to the occupation of one frequency mode with a higher energy. This higher energy state is a coherent state. For simplicity in our calculations, we assume the ground state of the MT to be equivalent to the ket $|0\rangle$. Now, the state of MT can be written as the superposition of the ground state and coherent state,

$$\left|\psi(0)\right\rangle_{MT} = c_g \left|0\right\rangle + c_e \left|z\right\rangle \tag{6}$$

where $|z\rangle$ is our representation for the coherent state. It is written as $|z\rangle = e^{-\frac{|z|^2}{2}} \sum_{n=0}^{\infty} \frac{z^n}{\sqrt{n!}} |n\rangle$, where the coherent state $|z\rangle$ satisfies in $\hat{a}|z\rangle = z|z\rangle$ where \hat{a} is the annihilation operator.

We now investigate its interaction with N biophotons. The state of biophotons is considered as

$$\left|\phi(0)\right\rangle_{biophotons} = \sum_{n=0}^{\infty} c_n \left|n\right\rangle \tag{7}$$

and the total state is

$$|\psi(0)\rangle = |\psi(0)\rangle_{MT} \otimes |\varphi(0)\rangle_{biophotons}$$
(8)

After solving the Schrödinger's equation we have the time-dependent wave function as

$$|\psi(t)\rangle = \sum_{n=0}^{\infty} \left\{ \left\{ \left[c_e c_n \cos(\lambda t \sqrt{n+1}) - i c_g c_{n+1} \sin(\lambda t \sqrt{n+1}] | z \right] + \left[-i c_e c_{n-1} \sin(\lambda t \sqrt{n}) + i c_g c_n \cos(\lambda t \sqrt{n}) \right] | 0 \right\} \right\} | n\rangle$$

In this case, the MT system first occupies the ground state. It means $c_g = 1$, and $c_e = 0$. After the substitution of these coefficients, the state becomes

$$|\psi(t)\rangle = \sum_{n=0}^{\infty} \begin{cases} \left\{ \left[-ic_{n+1}\sin(\lambda t\sqrt{n+1}] | z \rangle \right] \\ + \left[ic_{n}\cos(\lambda t\sqrt{n}) \right] | 0 \rangle \right\} \end{cases} | n \rangle$$
(9)

Written in another form the state is

$$|\psi(t)\rangle = |\psi_g(t)\rangle|0\rangle + |\psi_e(t)\rangle|z\rangle$$
⁽¹⁰⁾

where

$$|\psi_{e}(t)\rangle = -i\sum_{n=0}^{\infty}c_{n}\sin(\lambda t\sqrt{n})|n-1\rangle$$
(11)

$$|\psi_{g}(t)\rangle = i\sum_{n=0}^{\infty} c_{n} \cos(\lambda t \sqrt{n}) |n\rangle$$
(12)

Now, we introduce the fluctuation operator σ_F as

$$\sigma_{F} = |e\rangle\langle e|-|g\rangle\langle g| = |z\rangle\langle z|-|0\rangle\langle 0|$$
(13)

To determine when the MT is in the coherent state and when in the ground state, we use the fluctuation function. We define the fluctuation function as

$$F(t) = \left\langle \psi \left| \sigma_F \right| \psi \right\rangle$$

This function determines the rate of transitions between the coherent state and the ground state. Eventually, the fluctuation function for this state is

$$F(T,z) = \sum_{n=0}^{\infty} c_n^2 \cos^2(\sqrt{n}T) \left[e^{-|z|^2} - 1 \right] + \sum_{n=0}^{\infty} c_n^2 \sin^2(\sqrt{n}T) \left[e^{-\frac{|z|^2}{2}} + e^{-|z|^2} \right]$$
(14)

where $T = \lambda t$. We refer to the parameter *T* as "scaled time". Here, c_n is the coefficient describing the biophoton field and thus $|c_n|^2$ is the probability of detection of biophotons. We may infer information about coherent light from photo-count statistics (PCS) [26]. One calculates the probability $p(n, \Delta t)$ of registering n photons in preset time interval Δt by recording the number of photons during the measurement. A fully coherent field satisfies a Poissonian distribution $p(n) = \frac{N^n e^{-N}}{n!}$ for all times $\Delta t > 0$, where *N* is the average number of detected photons per time interval Δt . The fact that a coherent field satisfies a Poissonian distribution is rooted in quantum theory. The amplitude and photon numbers are not simultaneously measurable with arbitrary certainty. However, definite and robust experimental proof is often complicated due to very small photon counts in Δt . Biophotons have been considered coherent by other researchers and there are some claims of experimental observations of coherent biophotons [6,7,9, 66, 67], but in general the claim of biophoton coherence still requires a concrete proof. However, here we hypothetically assume the coherence property of biophotons and investigate its consequences. Thus for biophotons we have

$$c_n = e^{-\frac{|\alpha|^2}{2}} \frac{\alpha^n}{\sqrt{n!}}$$

where the normalization implies that $|\alpha|^2 = N$ [26], keeping in mind that N is the average number of biophotons. Thus, $c_n^2 = e^{-N} \frac{N^n}{n!}$. Finally, the fluctuation function becomes

$$F(T, z) = \sum_{n=0}^{\infty} e^{-N} \frac{N^n}{n!} \left[1 - \cos^2(\sqrt{nT}) + \sin^2(e^{-|z|^2/2} \sqrt{nT}) \right]$$

The 3D diagrams of the fluctuation function are plotted in terms of the scaled time and coordinate z, for different numbers of biophotons N. According to earlier calculations [11], at least 100 biophotons can be produced within each human visual neuron per second during visual perception. So, a few biophotons per second are probably simultaneously present within the 1-10 micrometer length of an MT. Here, we have considered different values for the numbers of biophotons around an MT such as N=2, 5, 7 and 10 and plotted 3D and 2D diagrams of the fluctuation function in Figures 3 and 4, respectively.

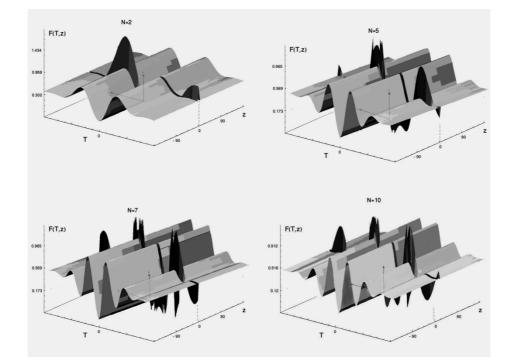


Fig. 3. 3D diagrams of the fluctuation function for different numbers of biophotons N. It is seen that the maximum of fluctuation is around z=1. MT is initially in the ground state

As introduced above, F(T,z) is the fluctuation function which determines the transition rate of biomolecules between the coherent states and the ground state. When in coherent states, the biophotons are absorbed via biomolecules and when in the ground state, the biophotons are absorbed via the vacuum. In this model, the information can be restored from the vacuum, and conscious states can be repeated as before. Then, the information can be sent back to the memory site again via emission of biophotons. This cycle can be repeated an arbitrary number of times. We have plotted the F(T,z) function for different values of its variables. According to 3D diagrams in Figure 3, we let the values near 1 for z, since the main fluctuations are in this area. In the next diagrams the scaled times are evaluated as $T=10^4$ and $T=10^5$.

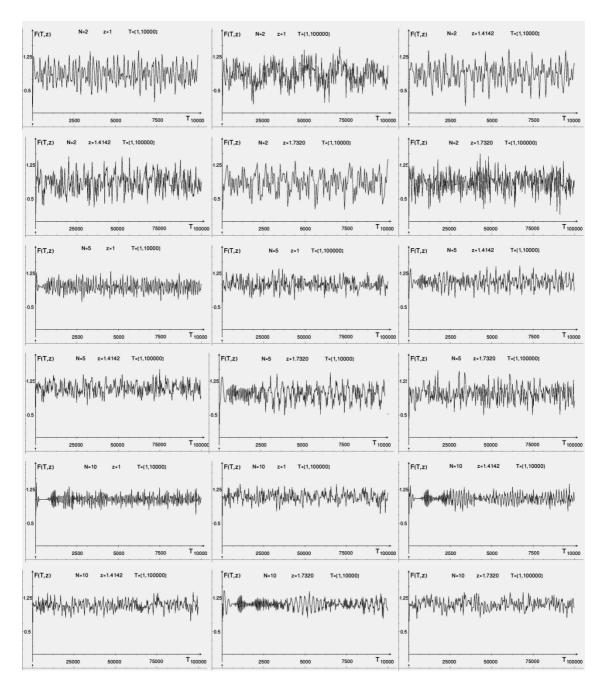


Fig. 4. 2D diagrams of the fluctuation function for different values of N and z in different scaled time intervals. MT are initially in the ground state.

Now, we investigate the behavior of the fluctuation function with the assumption that the MT is first in the excited state. Here $c_e(0) = 1$ and $c_g(0) = 0$. With the substitution of these coefficients we have

$$\left|\psi_{e}(t)\right\rangle = \sum_{n=0}^{\infty} c_{n} \cos(\lambda t \sqrt{n+1}) \left|n\right\rangle$$
(15)

$$\left|\psi_{g}(t)\right\rangle = -i\sum_{n=0}^{\infty}c_{n}\sin(\lambda t\sqrt{n+1})\left|n+1\right\rangle$$
(16)

Continuing our calculations for the fluctuation function we find that

$$F(T, z) = \sum_{n=0}^{\infty} e^{-N} \frac{N^n}{n!} \left[1 - \sin^2(\sqrt{n+1}T) + \cos^2(e^{-|z|^2/2}\sqrt{n+1}T) \right]$$

We have plotted 3D and 2D diagrams of the fluctuation function in Figures 5 and 6, respectively. As shown before, the maximum of fluctuations is found to be around z=1. Here, we see again these fluctuations around z=1.

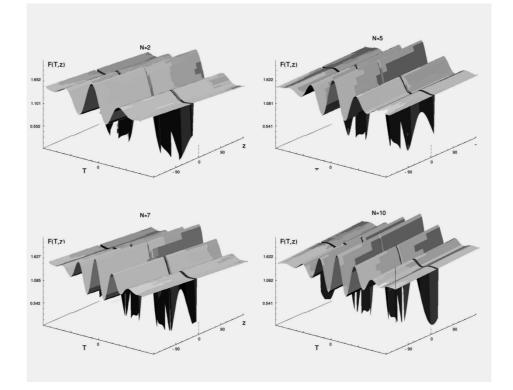


Fig. 5. 3D diagrams of the fluctuation function for different values of biophotons N. In this interaction, MT is initially in the coherent state

Next, we plot F(T,z) again for the above case at different values of its variables.

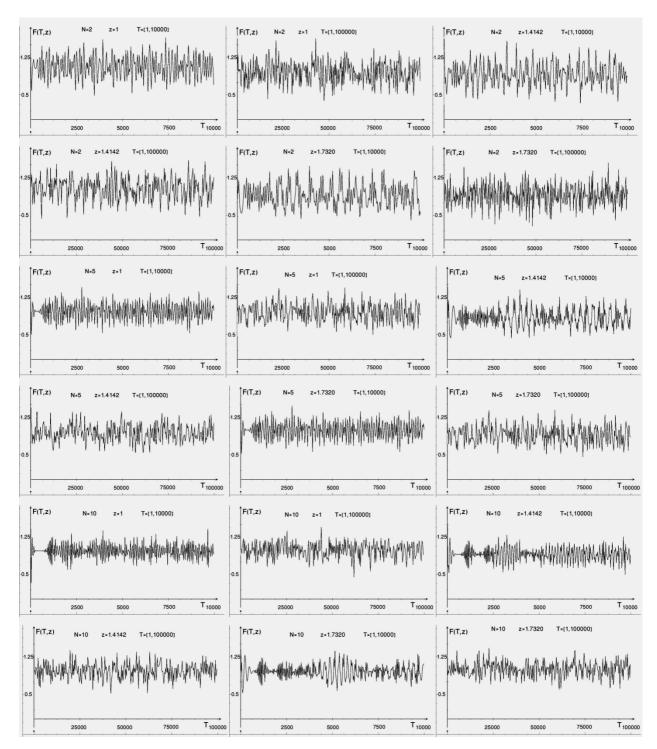


Fig. 6. 2D diagrams of the fluctuation function for different values of N and z. MT is initially in the coherent state.

In Figures 4 and 6 it is seen that the amplitudes of the fluctuation function are decreasing by increasing the number of biophotons which interact with an MT, and vice versa.

4. Electrical properties of MTs and their effects on the membrane activity

In previous sections we explained how biophotons can help MTs support coherent states. We have shown the existence of fluctuations between coherent states and normal states for MTs during the interaction with biophotons. Now the question arises how biophotons can affect

the electrical activity of membrane via MTs. To answer this question we should investigate how MT activity may affect the electrical activity of the membrane. It is well known that MTs play key roles in the trafficking of neurotransmitters to the synapse. According to Alvarez and Ramirez [5] action potential leads to a decrease of the disassembly rate of MTs. Very recently, Gardiner et al. [25] reviewed the evidence for neurotransmitters regulation (i.e. serotonin, melatonin, dopamine, glutamate, glycine, and acetylcholine) of the MT cytoskeleton. They postulated that MTs may play a direct role in propagating action potentials via conductance of electric current.

Both experimental and theoretical approaches have been used to study electrical signaling along MTs. For experimental investigations, the dual patch-clamp set up was used. In such experiments, electrical data were gathered and taxol-stabilized MTs were shown to behave as biomolecular transistors responding to brief pulses of electric current whose voltage amplitude was in the range of ±200mV [68]. Individual MTs were shown to amplify applied electrical current two-fold indicating a capability for ionic signal propagation that appeared to involve the condensed positive counterion clouds distributed along the length of the MT (where approximately 20 unit charges are present per tubulin monomer). This ionic cloud was attracted to the negative surface charge of the MT retaining its longitudinal mobility. Further measurements and theoretical modeling showed that MTs support nonlinear wave propagation [68]. It was also found that MTs exhibited transistor-like properties. It is worth noting that MTs were also shown to be conductive using an independent experiment involving an electroorientation approach [58]. Intact MTs were demonstrated to have conductance of 157 ± 7 mS/m, while MTs treated with subtilisin (which cleaves tubulin's Ctermini) lowered it to 96 \pm 6 mS/m. An argument was put forth that counterions on the surface (many of which interact with the negatively charged C-termini) are responsible for the observed conductance. Ionic conduction along MTs was modeled in terms of a nonlinear electrical circuit using cable equations [69]. Based on these results which indicate that MTs transmit electric signals between distant points within a neuron, the sources of potential information need to be identified in order to understand how processing sensory-based information occurs into a higher cognitive state. The most significant source of input is from the neuronal membrane which contains postsynaptic densities making contacts with other neurons.

In order for actin filaments (AFs) to conduct an electric signal to the MT network inside the neuron, there must be a functional link between these two types of cytoskeleton. There are at least three potential mechanisms through which AFs and MTs could interact: (1) direct physical contact, (2) via various types of linking proteins, and (3) indirectly through signal transduction. AFs are often found in cells forming direct association with MTs. MTs, in turn, frequently migrate correlated with actin bundles [76] as was demonstrated by bualwavelength fluorescent speckle microscopy analyses. AFs interact with MTs as part of neuronal migration, growth cone development, and neuronal receptor and ion channel transport [38]. While detailed mechanisms governing MT/AF interactions remain an open issue, numerous recently acquired insights reveal a growing number of cross-linker proteins playing important roles.

Cross-linker protein	References	
CLIP-115	[38]	
CLIP-170	[38],[101]	
CLASP1	[38],[93]	
CLASP2	[38],[93]	
Lis1	[38],[46]	
EB family	[38],[53]	
MAP2c	[51],[78]	
Tau	[78]	

Table 1. Major linker proteins of actin filaments with microtubules

Table 1 summarizes the known cross-linker proteins that bind MTs to AFs. Note that many of these proteins are MT plus-end tracking (so called +TIPs) [38]. Cytoplasmic linker proteins, CLIP-115 and CLIP-170, bind plus-ends of MTs and link them to cargo or to AFs though scaffolding protein intermediaries [101]. CLIP-associated proteins, CLASP1 and CLASP2, represent two additional examples of +TIPs. CLASP2 α contains a binding site for actin [93]. Lis1 tethers AF to MTs through interactions with scaffolding proteins [46]. Similarly, the EB family of +TIPs, which include EB1 and EB2, interact with other proteins (including CLASP1/2, CLIP-170) in order to bind MTs to AFs [53]. There are also the microtubule-associated proteins (MAPs): MAP2 and tau, which are known to bind AFs at relatively low affinities [51,78]. Signal transduction molecules, such as calmodulin, and Ca²⁺ and phosphorylation have been found to modulate the ability of MAP2 and tau to bind AFs and MTs. Consequently, in spite of numerous gaps in our understanding of these interactions, it is clear that electric signals arising from synaptic input can reach the internal cytoskeleton using various molecular pathways.

4.1 An Overview

We strongly believe that electrodynamic interactions between various cytoskeletal structures, with MTs playing a central role, and ion channels crucially regulate the neural informationprocessing mechanism. These interactions involve long-range ionic wave propagation along microtubule networks (MTNs) and AFs and exhibit subcellular control of ionic channel activity. Hence, they have an impact on the computational capabilities of the entire neural function. Cytoskeletal biopolymers, most importantly AFs and MTs, constitute the basis for wave propagation, and interact with membrane components leading to a modulation of synaptic connections and membrane ion channels. Association of MTs with AFs in neuronal filopodia guides MT growth and affects neurite initiation [18]. This is seen in neurons by the presence of proteins that interact with both MTs and AFs, as well as proteins that mediate interactions between both types of filaments. For example, MAP1B and MAP2 interact with actin in vitro [64,90] but cross-linking, MAP2 and/or MAP1B is associated with both types of filaments contributing to the guidance of MTs along AF bundles. Direct interaction between AFs and ion channels has been seen and a regulatory functional role has been associated with actin. Thus, it is clear that the cytoskeleton directly and indirectly affects membrane components, in particular ion channels and synapses.

MTs and AFs also interact during the migration of developing growth cones, a process that involves +TIPS, which tend to aggregate at the end of the axon shaft in the area where AFs are highly localized. Additionally, the neurotrophin NGF and the signal transduction molecule GSK-3 β act to assure a balance between stable and unstable MTs [28]. The dynamic relationships between AFs and MTs during growth cone migration could be partially controlled by electric signals transmitted between these networks, which is supported by experimental evidence. Applied electric fields were found to guide growth cones of *Xenopus* spinal neurons towards the positive current source and this effect depended on intact AFs and MTs [71].

While learning-related changes in MTs, AFs, MAPs and signal transduction molecules are well documented [103], much less is known about how electric signaling in the MT and AF networks might be involved memory formation processes. Electric signaling by AFs and MTs may play active roles in coincidence detection and storage of spatiotemporal patterns of inputs, and signaling within the cytoskeleton may be particularly critical to information storage over longer time scales than LTP times. The initial route to the MT network could be through the AFs concentrated in the spines. Inputs to arbitrary sites in the neuron can be transmitted from the neuronal membrane to AFs in spines via scaffolding proteins and signal transduction molecules. Electric signals can then be transmitted utilizing AF cross-linker proteins to MTs, and subsequently through MAPs and signal transduction molecules to other MTs in the network. A spatiotemporal pattern inherent to a complex stimulus (or cogneme) can therefore be readily envisaged and indeed mathematically modeled as a convergence of electrical signals to a particular stretch of an MT inside the dendrite.

5. Concluding Remarks

There is no doubt that EEG waves are deeply involved with the basic functioning of the brain but the origin and the exact function of EEG has remained a mystery. The EEG waves associated with two distant neurons are strongly correlated and this supports the view that EEG waves are related to the properties of the brain as a coherent quantum system. It is not possible for a scalp EEG to determine the activity within a single dendrite or neuron. Rather, a surface EEG reading is the summation of the synchronous activity of thousands of neurons that have similar spatial orientation, radial to the scalp².

Synaptic transmission and axonal transfer of nerve impulses are too slow to organize coordinated activity in large areas of the central nervous system. Numerous observations confirm this view [73]. The duration of a synaptic transmission is at least 0.5 ms, thus the transmission across thousands of synapses takes about hundreds or even thousands of milliseconds. The transmission speed of action potentials varies between 0.5 m/s and 120 m/s along an axon. More than 50% of the nerves fibers in the corpus callosum are without myelin, thus their speed is reduced to 0.5 m/s. How can these low velocities (i.e. classical signals) explain the fast processing in the nervous system? We believe that quantum theory is able to explain some of the above mysteries. As an example, recently it has been shown theoretically that the biological brain has the possibility to achieve large quantum bit computing at room temperature, superior when compared with the conventional processors [60].

² Updated from : <u>http://www.reference.com/browse/electro-encephalogram</u>, 13 November 2009.

Neuroscientists or brain specialists record the EEG diagrams of their patients when their eyes are closed, because when their eyes are open the amplitudes of diagrams dramatically would be reduced and they cannot diagnose changes in the amplitudes. In Figure 7 we represent the EEG diagrams of a person with different situations as explained in the caption to the figure. When the eyes are opened, the number of interacting photons with the biomolecules becomes high and the amplitudes of EEG become low. It is clear when the intensity of incident light is high; it must produce more action potentials than low intensity light. In classical physics, it is argued that this situation is because the superposition of several waves tends to decrease the amplitude of the total wave. This argument ignores the intensity of the incident light entering the eye. This classical argument also cannot explain when and how the synchronization happens in order to decrease the amplitude of EEG.

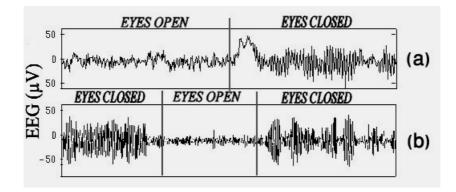


Fig. 7. A schematic alpha-EEG diagram of a person when (a) the eyes are opened and then he/she closes the eyes (b) the eyes are closed and then she/he opens his eyes and then closes them again.

Our argument is based on the interaction of (bio)photons and MTs inside the neurons. When the eyes start to be opened, the incident numbers of photons into the eyes increases and according to the diagrams in Figures 4 and 6 the amplitudes are reduced. Also, the amount of incident photons can increase the amount of biophoton production inside the neurons, since the visible light can be produced inside the brain in the form of biophotons [37,49,50,100] and this interaction is not only limited to the external incident photons with MTs. So far we have not been able to find an exact relation between the EEG diagrams and the fluctuation function, but the synchronous and coherent vibrations of billions electric dipoles of biomolecules cannot be ignored in the EEG diagrams. MTs are particularly numerous in the brain where they form highly ordered bundles and are the best candidate for long coherence and large synchrony [32]. The argument for connection between Alpha-EEG diagrams and MTs activity is their similar behavior in increasing and decreasing of amplitudes of fluctuation function for MTs and potential difference in EEG in response to the intensity of photons. This similarity during opening and shutting of the eyes indicates a significant relation between the EEG diagrams and the fluctuation function.

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References

- [1] Adamo AM, Llesuy SF, Pasquini JM, Boveris A, Brain chemiluminescence and oxidative stress in hyperthyroid rats, *Biochem J* **263**:273–277, 1989.
- [2] Albrecht-Buehler G, Autofluorescence of live purple bacteria in the near infrared, *Exp Cell Res* **236**:43–50, 1997.
- [3] Albrecht-Buehler G, Cellular infrared detector appears to be contained in the centrosome, *Cell Motil Cytoskeleton* **27**:262–271, 1994.
- [4] Albrecht-Buehler G, Changes of cell behavior by near-infrared signals, *Cell Motil Cytoskeleton* **32**:299–304, 1995.
- [5] Alvarez J, Ramirez BU, Axonal microtubules: their regulation by the electrical activity of the nerve, *Neurosci Lett* **15**:19–22, 1979.
- [6] Bajpai RP, Coherent nature of biophotons: Experimental evidence and phenomenological model, pp. 323–339, in Chang J, Fisch J, Popp FA (eds.), *Biophotons*, Kluwer Academic Publishers, Dordrecht, 1998.
- [7] Bajpai RP, Coherent Nature of the Radiation Emitted in Delayed Luminescence of Leaves, *J Theor Biol* **198**:287–299, 1999.
- [8] Batyanov AP, Distant optical interaction of mitochondria. *Bull Exp Biol Med* **97**:740–742, 1984.
- [9] Bischof M, Biophotons The light in our cells, *J Optom Photother*, March, 1–5, 2005.
- [10] Bókkon I, Phosphene phenomenon: A new concept, *BioSystems* **92**:168–174, 2008.
- [11] Bókkon I, Salari V, Tuszynski J, Antal I, Estimation of the number of biophotons involved in the visual perception of a single object image: Biophoton intensity can be considerably higher inside cells than outside. *J Photochem Photobiol B* **100**:160–166, 2010.
- [12] Bókkon I, Visual perception and imagery: a new hypothesis, *BioSystems* **96**:178–184, 2009.
- [13] Chiarugi P, Reactive oxygen species as mediators of cell adhesion, *Ital J Biochem* **52**: 28–32, 2003.
- [14] Chiarugi P, Src redox regulation: there is more than meets the eye, *Mol Cells* **26**:329–337, 2008.
- [15] Chwirot BW, Ultraweak luminescence studies of microsporogenesis in Larch, pp. 259–285, in Popp FA, Li KH, Gu Q (eds.), *Recent advances in biophoton research and its applications*, World Scientific, Singapore, River Edge, NJ, 1992.
- [16] Cifra M, Pokorný J, Havelka D, Kučera O, Electric field generated by axial longitudinal vibration modes of microtubule, *BioSystems* **100**:122 131, 2010.
- [17] Cilento G, Photobiochemistry without light, *Experientia* **44**:572–576, 1988.
- [18] Dehmelt L, Halpain S, Actin and microtubules in neurite initiation: Are MAPs the missing link? *J Neurobiol* **58**:18–33, 2004.
- [19] Deriu MA, Soncini M, Orsi M, Patel M, Essex JW, Montevecchi FM, Redaelli A, Anisotropic Elastic Network Modeling of Entire Microtubules, *Biophys J* 99:2190 2199, 2010.
- [20] Dröge W, Free Radicals in the Physiological Control of Cell Function, *Physiol Rev* 82:47–95, 2002.
- [21] Faber J, Portugal R, Rosa LP, Information processing in brain microtubules, *BioSystems* 83:1–9, 2006.
- [22] Fröhlich H, Long range coherence and energy storage in biological systems, *Int J Quantum Chem* **2**:641–649, 1968.
- [23] Fröhlich H, Long range coherence and the action of enzymes, *Nature* 228:1093, 1970.
- [24] Fröhlich H, The extraordinary dielectric properties of biological materials and the action of enzymes, *PNAS* **72**:4211–4215, 1975.
- [25] Gardiner J, Overall R, Marc J, The microtubule cytoskeleton acts as a key downstream effector of neurotransmitter signaling, *Synapse* **65**:249–256, 2011.

- [26] Gerry C, Knight P, *Introductory Quantum Optics*, Cambridge University Press, Cambridge, 2005. And also to see the criteria of coherent state of the light see: Walls DF, Millburn GJ, *Quantum Optics*. Springer-Verlag New York, 1995.
- [27] Gordeeva AV, Zvyagilskaya RA, Labas YA, Cross-talk between reactive oxygen species and calcium in living cells, *Biochemistry (Mosc)* **68**:1077–1080, 2003.
- [28] Gordon-Weeks PR, Microtubules and growth cone function, J Neurobiol 58:70–83, 2004.
- [29] Grass F, Kasper S, Humoral phototransduction: light transportation in the blood, and possible biological effects, *Med Hypotheses* **71**:314–317, 2008.
- [30] Hagan S, Hameroff SR, Tuszynski JA, Quantum computation in brain microtubules: Decoherence and biological feasibility, *Phys Rev E* **65**:061901, 2002.
- [31] Hameroff S, Tuszynski J, Quantum states in proteins and protein assemblies: The essence of life? *Proceedings of SPIE Conference on Fluctuationa and Noise*, Canary Islands, 2004.
- [32] Hameroff SR, Penrose R, Conscious events as orchestrated space-time selections, J Conscious Stud 3:36–53, 1996.
- [33] Hidalgo C, Carrasco MA, Muñoz P, Núñez MT, A role for reactive oxygen/nitrogen species and iron on neuronal synaptic plasticity, *Antioxid Redox Signal* **9**:245–255, 2007.
- [34] Hideg E, Kobayashi M, Inaba H, Spontaneous ultraweak light emission from respiring spinach leaf mitochondria, *Biochim Biophys Acta* **1098**:27–31, 1991.
- [35] Hoidal JR, Reactive Oxygen Species and Cell Signaling, *Am J Respir Cell Mol Biol* **25**:661–663, 2001.
- [36] Imaizumi S, Kayama T, Suzuki J, Chemiluminescence in hypoxic brain--the first report. Correlation between energy metabolism and free radical reaction, *Stroke* **15**:1061–1065, 1984.
- [37] Isojima Y, Isoshima T, Nagai K, Kikuchi K, Nakagawa H, Ultraweak biochemiluminescence detected from rat hippocampal slices, *NeuroReport* **6**:658–660, 1995.
- [38] Jaworski J, Hoogenraad CC, Akhmanova A, Microtubule plus-end tracking proteins in differentiated mammalian cells, *Int J Biochem Cell Biol* **40**:619–637, 2008.
- [39] Jaynes ET, Cummings FW, Comparison of quantum and semiclassical radiation theories with application to the beam maser, *Proc IEEE* **51**:89–109, 1963.
- [40] Jelínek F, Pokorný J, Microtubules in biological cells as circular waveguides and resonators, *Electro- magnetobiol* **20**:75–80, 2001.
- [41] Jibu M, Brinbram KH, Yasue K, From conscious Experience to Memory Storage and Retrieval: The role of quantum brain dynamics and boson condensation of evanescent photons, *Inter J Mod Physics B* 10:1735–1754, 1996.
- [42] Jibu M, Hagan S, Hameroff SR, Pribram KH, Yasue K, Quantum optical coherence in cytoskeletal microtubules: implications for brain function, *Biosystems* **32**:195–209, 1994.
- [43] Karu T, Primary and secondary mechanisms of action of visible to near-IR radiation on cells, *J Photochem Photobiol B* **49**:1–17, 1999.
- [44] Kataoka Y, Cui Y, Yamagata A, Niigaki M, Hirohata T, Oishi N, Watanabe Y, Activity-Dependent Neural Tissue Oxidation Emits Intrinsic Ultraweak Photons, *Biochem Biophys Res Commun* **285**:1007–1011, 2001.
- [45] Kato M, Shinzawa K, Yoshikawa S, Cytochrome oxidase is a possible photoreceptor in mitochondria, *Photobiochem Photobiophys* **2**:263–269, 1981.
- [46] Kholmanskikh SS, Koeller HB, Wynshaw-Boris A, Gomez T, Letourneau PC, Ross ME, Calcium-dependent interaction of Lis1 with IQGAP1 and Cdc42 promotes neuronal motility, *Nat Neurosci* **9**:50–57, 2006.
- [47] Kishida KT, Klann E, Sources and targets of reactive oxygen species in synaptic plasticity and memory, *Antioxid Redox Signal* **9**:233–244, 2007.
- [48] Knapp LT, Klann E, Role of reactive oxygen species in hippocampal long-term potentiation: contributory or inhibitory? *J Neurosci Res* **70**:1–7, 2002.
- [49] Kobayashi M, Takeda M, Ito K, Kato H, Inaba H, Two-dimensional photon counting imaging and spatiotemporal characterization of ultraweak photon emission from a rat's brain in vivo, *J Neurosci Methods* **93**:163–168, 1999.

- [50] Kobayashi M, Takeda M, Sato T, Yamazaki Y, Kaneko K, Ito K, Kato H, Inaba H, In vivo imaging of spontaneous ultraweak photon emission from a rat's brain correlated with cerebral energy metabolism and oxidative stress, *Neurosci Res* **34**:103–113, 1999.
- [51] Kotani S, Nishida E, Kumagai H, Sakai H, Calmodulin inhibits interaction of actin with MAP2 and Tau, two major microtubule-associated proteins, *J Biol Chem* **260**:10779–10783, 1985.
- [52] Kruk I, Lichszteld K, Michalska T, Wronska J, Bounias M, The formation of singlet oxygen during oxidation of catechol amines as detected by infrared chemiluminescence and spectrophotometric method, *Z Naturforsch [C]* **44**:895–900, 1989.
- [53] Lansbergen G, Akhmanova A, Microtubule plus end: a hub of cellular activities, *Traffic* 7:499–507, 2006.
- [54] Mansfield JW, Biophoton distress flares signal the onset of the hypersensitive reaction, *Trends Plant Sci* **10**:307–309, 2005.
- [55] Mavromatos NE, Mershin A, Nanopoulos DV, QED-cavity model of microtubules implies dissipationless energy transfer and biological quantum teleportation, *Inter J Mod Physics B* **16**:3623–3642, 2002.
- [56] Mavromatos NE, Nanopoulos DV, On Quantum Mechanical Aspects of Microtubules, *Inter J Mod Physics B* 2:517 –542, 1998.
- [57] Mazhul' VM, Shcherbin DG, Phosphorescent analysis of lipid peroxidation products in liposomes, *Biofizika* 44:676–681, 1999.
- [58] Minoura I, Muto E, Dielectric measurement of individual microtubules using the electroorientation method, *Biophys J* **90**:3739–3748, 2006.
- [59] Murata T, Kadota A, Wada M, Effects of Blue Light on Cell Elongation and Microtubule Orientation in Dark-Grown Gametophytes of Ceratopteris richardii, *Plant Cell Physiol* 38:201–209, 1997.
- [60] Musha T, Possibility of High Performance Quantum Computation by Superluminal Evanescent Photons in Living Systems, *BioSystems* **96**:242–245, 2009.
- [61] Nakano M, Low-level chemiluminescence during lipid peroxidations and enzymatic reactions, *J Biolumin Chemilum* **4**:231–240, 1989.
- [62] Nogales E, Wolf SG, Downing K, Structure of the alpha beta tubulin dimer by electron crystallography, *Nature* **391**:199–203, 1998.
- [63] Ouyang M, Awschalom DD, coherent spin transfer between moleculary bridged quantum dots, *Science* **301**:1074–1078, 2003.
- [64] Pedrotti B, Colombo R, Islam K, Interactions of microtubule associated protein MAP2 with unpolymerized and polymerized tubulin and actin using a 96-well microtiter plate solid phase immunoassay, *Biochemistry* **33**:8798–8806, 1994.
- [65] Pokorný J, Jelínek F, Trkal V, Lamprecht I, Hölzel R, Vibrations in Microtubules, *J Biol Phys* 23:171–179, 1997.
- [66] Popp FA, Nagl W, Li KH, Scholz W, Weingartner O, Wolf R, Biophoton emission. New evidence for coherence and DNA as source, *Cell Biophys* **6**:33–52, 1984.
- [67] Popp FA, Yan Y, Delayed Luminescence of Biological Systems in Terms of Coherent States, *Phys Lett A* **293**:93–97, 2002.
- [68] Priel A, Ramos AJ, Tuszynski JA, Cantiello HF, A biopolymer transistor: electrical amplification by microtubules, *Biophys J* **90**:4639–4643, 2006.
- [69] Priel A, Tuszynski JA, A nonlinear cable-like model of amplified ionic wave propagation along microtubules, *Europhysics Lett* **83**:68004, 2008.
- [70] Quickenden TI., Que Hee SS, Weak luminescence from the yeast Sachharomyces-Cervisiae, *Biochem Biophys Res Commun* **60**:764–770, 1974.
- [71] Rajnicek AM, Foubister LE, McCaig CD, Growth cone steering by a physiological electric field requires dynamic microtubules, microfilaments and Rac-mediated filopodial asymmetry, *J Cell Sci* 119:1736–1745, 2006.
- [72] Reimers JR, McKemmish LK, McKenzie RH, Mark AE, Hush NS, Weak, strong, and coherent regimes of Fröhlich condensation and their applications to terahertz medicine and quantum consciousness, *PNAS* **106**:4219–4224, 2009.

- [73] Reinis S, Holub RF, Smrz P, A quantum hypothesis of brain function and consciousness, Translated version of Ceskoslovenska fyziologie 54:26–31, 2005.
- [74] Salari V, Rahnama M, Tuszynski J 2011. Toward Quantum Visual Information Transfer in the Human Brain, arXiv.org/quant-phys:0809.0008, to appear in Foundations of Physics.
- [75] Salari V, Tuszynski J, Rahnama M, Bernroider G, On the Possibility of Quantum Coherence in Biological Systems 2010, arXiv.org/quant-phys:1012.3879, to appear in Journal of Physics, Conf Ser.
- [76] Salmon WC, Adams MC, Waterman-Storer CM, Dual-wavelength fluorescent speckle microscopy reveals coupling of microtubule and actin movements in migrating cells, *J Cell Biol* **158**:31–7, 2002
- [77] Scott RQ, Roschger P, Devaraj B, Inaba H, Monitoring a mammalian nuclear membrane phase transition by intrinsic ultraweak light emission, *FEBS Lett* **285**:97–98, 1991.
- [78] Selden SC, Pollard TD, Interaction of actin filaments with microtubules is mediated by microtubule-associated proteins and regulated by phosphorylation, *Ann N Y Acad Sci* **466**:803–812, 1986.
- [79] Skulachev VP, Mitochondrial filaments and clusters as intracellular power-transmitting cables, *Trends Biochem Sci* **26**:23–29, 2001.
- [80] Slawinski J, Luminescence research and its relation to ultraweak cell radiation, *Experientia* 44:559–571, 1988.
- [81] Staxén I, Bergounioux C, Bornman J, Effect of ultraviolet radiation on cell division and microtubule organization in Petunia hybrida protoplasts. *Protoplasma* **173**:70–76, 1993.
- [82] Sun Y, Wang C, Dai J, Biophotons as neural communication signals demonstrated by in situ biophoton autography, *Photochem Photobiol Sci* **9**:315–322, 2010.
- [83] Takeda M, Tanno Y, Kobayashi M, Usa M, Ohuchi N, Satomi S, Inaba H, A novel method of assessing carcinoma cell proliferation by biophoton emission, *Cancer Lett* **127**:155–160, 1998.
- [84] Tegmark M, Importance of quantum decoherence in brain processes, *Phys Rev E* **61**: 4194–4206, 2000.
- [85] Tejada-Simon MV, Serrano F, Villasana LE, Kanterewicz BI, Wu GY, Quinn MT, Klann E, Synaptic localization of a functional NADPH oxidase in the mouse hippocampus, *Mol Cell Neurosci* 29:97–106, 2005.
- [86] Thar R, Kühl M, Propagation of electromagnetic radiation in mitochondria? J Theor Biol **230**:261–270, 2004.
- [87] Thiels E, Klann E, Hippocampal memory and plasticity in superoxide dismutase mutant mice, *Physiol Behav* **77**:601–605, 2002.
- [88] Thiels E, Urban NN, Gonzalez-Burgos GR, Kanterewicz BI, Barrionuevo G, Chu CT, Oury TD, Klann E, Impairment of long-term potentiation and associative memory in mice that overexpress extracellular superoxide dismutase, *J Neurosci* **20**:7631–7639, 2000.
- [89] Tilbury RN, Cluickenden TI, Spectral and time dependence studies of the ultraweak bioluminescence emitted by the bacterium Escherichia coli, *Photobiochem Photobiophys* **47**:145–150, 1988.
- [90] Togel M, Wiche G, Propst F, Novel features of the light chain of microtubule associated protein MAP1B: Microtubule stabilization, self interaction, actin filament binding, and regulation by the heavy chain, *J Cell Biol* **143**:695–707, 1998.
- [91] Tong JJ, Mitochondrial delivery is essential for synaptic potentiation, *Biol Bull* **212**:169–175, 2007.
- [92] Touyz RM, Reactive oxygen species as mediators of calcium signaling by angiotensin II: implications in vascular physiology and pathophysiology, *Antioxid Redox Signal* **7**:1302–1314, 2005.
- [93] Tsvetkov AS, Samsonov A, Akhmanova A, Galjart N, Popov SV, Microtubule-binding proteins CLASP1 and CLASP2 interact with actin filaments, *Cell Motil* Cytoskeleton 64:519– 530, 2007
- [94] Tuszynski JA, Dixon JM, Quantitative analysis of the frequency spectrum of the radiation emitted by cytochrome oxidase enzymes, *Phys Rev E* 64: 051915, 2001.

- [95] Valko M, Leibfritz D, Moncol J, Cronin MT, Mazur M, Telser J, Free radicals and antioxidants in normal physiological functions and human disease, Int J Biochem Cell Biol **39**:44–84, 2007.
- [96] Van Wijk R, Bosman S, Ackerman J, Van Wijk E PA, *NeuroQuantology* **16**:452–463, 2008.
- [97] Van Wijk R, Schamhart DH, Regulatory aspects of low intensity photon emission, *Experientia* **44**:586–593, 1988.
- [98] Vladimirov YA, Proskurnina EV, Free radicals and cell chemiluminescence, Biochem (Moscow) 74:1545-1566, 2009. original txt in Uspekhi Biologicheskoi Khimii 49: 341–388, 2009.
- [99] Volterra A, Trotti D, Tromba C, Floridi S, Racagni G, Glutamate uptake inhibition by oxygen free radicals in rat cortical astrocytes, *J Neurosci* 14:2924–2932, 1994.
- [100] Wang Ch, Bókkon I, Dai J, Antal I. Spontaneous and visible light-induced ultraweak photon emission from rat eyes, *Brain Res* **1369**:1–9, 2011.
- [101] Watanabe T, Wang S, Noritake J, Sato K, Fukata M, Takefuji M, Nakagawa M, Izumi N, Akiyama T, Kaibuchi K, Interaction with IQGAP1 links APC to Rac1, Cdc42, and actin filaments during cell polarization and migration, *Dev Cell* 7:871–883, 2004
- [102] Watts BP, Barnard M, Turrens JF, Peroxynitrite-dependent chemiluminescence of amino acids, proteins, and intact cells, *Arch Biochem Biophys* **317**:324–330, 1995.
- [103] Woolf NJ, Priel A, Tuszynski JA, Nanoneuroscience: Structural and Functional Roles of the Neuronal Cytoskeleton in Health and Disease, Springer Verlag, Heidelberg, 2010.
- [104] Wu TM, Austin S, Bose condensation in biosystems, *Phys Lett A* 64:151–152, 1977.
- [105] Wu TM, Austin S, Cooperative behavior in biological systems, *Phys Lett A* **65**:74–76, 1978.
- [106] Wu TM, Austin SJ, Froehlich's model of bose condensation in biological systems, *J Biol Phys* **9**:97–107, 1981.
- [107] Yoon YZ, Kim J, Lee BC, Kim YU, Lee SK, Soh KS, Changes in ultraweak photon emission and heart rate variability of epinephrine-injected rats, *Gen Physiol Biophys* **24**:147–159, 2005.