

Simulation and Modelling of near Infrared Spectroscopy (NIRS) as Brain Monitor

Mohamed Shaaban Ali*

Anesthesia, Faculty of Medicine, Assiut University, Assiut, Egypt Email: <u>msali58@hotmail.com</u>

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Near infrared spectroscopy (NIRS) is a method for non-invasive monitoring of cerebral oxygenation and haemodynamics. Different devices provide information on changes of oxygenated (HbO₂) and deoxygenated haemoglobin (HHb), oxidized cytochrome aa3 (CytOx) or regional oxygen saturation (rSO₂). NIRS has been used during adult and paediatric cardiac surgery.

1. Physical Principles of NIRS

The unique ability of light in the near-infrared (NIR) range (700 - 1300 nm) to detect the oxygenation-state of living tissue was first realized by Jobsis in 1977 [1]. Since then, NIRS has emerged as a technique for *in vivo* and continuous monitoring of cerebral oxygenation.

This technique is based on several physical characteristics of NIR light. First, the skin, scalp and skull are relatively transparent to NIR light, allowing transmission of NIR photons into the brain. As these photons pass through the brain, they undergo attenuation as a result of scattering and absorption by specific chromophores. These chromophores are intravascular haemoglobin and intracellular cytochrome aa3. Measurement of changes in cerebral chromophore concentration is related to the overall optical attenuation of NIRS light by a modified Beer Lambert Law [2]:

$$\mathbf{A} = (\mathbf{a} [\mathbf{c}] \mathbf{L} \mathbf{B}) + \mathbf{G}$$

where A = the attenuation of light in optical densities, a = the extinction coefficient of the chromophore $(mM^{-1} \cdot cm^{-1})$, [c] = the concentration of chromophore (mM), L = the distance between light entry and exit (cm), B = a path length factor that takes into of the increased optical path length due to scattering of light in the tissue, and G = a factor related to tissue and optode geometry.

The attenuation due to scattering and the geometric factor is not known and thus the concentration of chromophore cannot be quantified absolutely. Nonetheless, in a given tissue with a fixed optical geometry, it is likely to remain constant. Changes in attenuation, therefore, can be attributed solely to changes in the concentration of chromophore

$$\Delta[\mathbf{c}] = \Delta \mathbf{A} / (\mathbf{a} \times \mathbf{L} \times \mathbf{B})$$

where ΔC is the change in chromophore concentration; ΔA is the change in optical density.

^{*}PhD, MD, Associate professor of Anesthesia.

Optical densities are measured at several wavelengths selected for their proximity to absorption peaks of chromophores of interest. NIRS measures changes in concentration of cerebral oxyhaemoglobin (HbO₂), deoxy-genated haemoglobin (HHb), and, from their sum changes of total haemoglobin (THb), as well as changes in the cerebral concentration of oxidized cytochrome aa3 (CytOx). In addition, claims are made to measure the ratio of HbO₂/THb and this is reported as "regional cerebral saturation" (rSO₂) or as "tissue oxygen index" (TOI).

Cytochrome oxidase is the terminal member of the mitochondrial respiratory chain, producing the energy necessary for conversion of adenosine diphosphate (ADP) to adenosine triphosphate (ATP). If oxygen is unavailable, electron transfer cannot take place and electrons accumulate on the haem and copper atoms of cytochrome aa3, causing it to become reduced. When oxygen becomes available, electrons are transferred to oxygen and cytochrome aa3 becomes oxidised. Thus, monitoring changes in oxidised form of the enzyme cytochrome aa3 can provide valuable information on oxidation-reduction state of the enzyme and hence on oxygenation at the cellular level [3] [4].

2. Limitations of NIRS

NIRS has the following limitations [5]-[7]:

- 1) It detects regional cerebral oxygenation (small area under the optodes).
- 2) Optical path length is difficult to identify.
- 3) Arterial versus venous changes are not distinguished.
- 4) The effect of extracranial tissue on NIRS signal is unknown.
- 5) Absolute quantification is lacking.
- 6) Vulnerable to light and movement artifacts.
- 7) Dissolved oxygen not measured.

3. NIRS Simulation and Modelling

Several methods of simulation and modelling were used to overcome such limitations [8]-[11]. Chang *et al.* [9] used a simulation method for understanding photon migration dynamics in human brain based on Monte Carlo algorithm by using three-dimensional magnetic resonance imaging (MRI) data with multi-wavelength illumination. MRI imaged the human brain with five layers; scalp, skull, cerebrospinal fluid (CSF), gray matter and white matter. They found that such patient-oriented simulation model can provide significant characteristics on the optimal choice of source-detector separation within 3.3 cm for individualized case. Also, significant distortions were observed around cerebral cortex folding. Spatial sensitivity penetrated deeper to the brain in the case of expanded CSF. Such finding suggests that the optical method may provide not only functional signal from the brain activation but also structural information of brain atrophy with expanded CSF. It appeared that the three-dimensional time-resolved brain modelling method approached the realistic human brain and provides useful information for NIRS systematic design and calibration for individualized case with prior MRI.

Others [11] described a four-layered for NIR light propagation in a human head based on Mont Carlo method. They used wave lengths of 690 nm, 800 nm and 1300 nm. The four layers are scalp and skull (one layer), CSF, grey matter and white matter. They found that the wave length of 1300 nm is more appropriate for monitoring of human brain activity. Also, grey matter absorption was more sensitive at 1300 nm, and the spatial resolution was improved at 1300 nm.

It appears that NIRS is a promising brain oxygenation monitor. It has all the theoretical advantages that could make it a suitable device for real-time brain monitoring during cardiac surgery. It is to be hoped that increasingly sophisticated computer modelling will permit a better understanding of its limitations and that modern equipment will make possible the real-time correction for the scattering effects of extracerebral tissue.

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