A Three-Wavelength Pulse Oximeter for Carboxyhemoglobin Determination

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Abstract:

Pulse oximetry measures the relative concentration of arterial oxygenated hemoglobin and reduced hemoglobin. This paper presents the design of a pulse oximeter, which measures the presence of a third substance, carboxyhemoglobin (HbCO), by unfiltered LEDs at 660 nm, 810 nm, and 940 nm. Background information is given on the theory of pulse oximetry. Theoretical approximations and calculations which are implemented in our instrument are presented. Sensor design and a flowchart, showing the signal processing routine that accomplishes the required analysis is included. The problem of noise occurrence during the signal processing routine is also discussed.

Keywords: Oximeter, pulse oximeter, hemoglobin, oxygenated hemoglobin, carboxyhemoglobin, sensor design, signal processing

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1. Oximeters

The first oximeters used spectrophotometry at two wavelengths on withdrawn blood. Upon withdrawal, the blood sample was placed in a cuvette and exposed to two or more monochromatic beams of light. The transmitted light was detected by a photocell whereupon the signal was amplified and put through calculations to estimate the saturation of hemoglobin. In order to avoid withdrawal of blood, this technique was extended by using fiber optics to transmit the wavelengths into and back from the blood [1]. This technique can measure $\%O_2$ saturation (SaO_2) noninvasively by transmitting light through the pinna of the ear or by reflecting light through the finger pad.

2. Pulse Oximeters

An improved method, different from the above, is pulse oximetry. It relies on the arterial pulsations of a given appendage. The change in light transmission due to the arterial inflow corresponds to the same transmission as the spectrophotometric case except that the dc level of light transmission must be eliminated. The dc portion is made up of nonchanging entities such as tissue, venous blood, bone, etc. [2–4]. Pulse oximetry provides a noninvasive method of determining arterial oxygen saturation.

3. Reflectance/Transmittance Oximeters

Reflectance oximeters are more complex because they increase the scattering of light [5–8]. Reflectance oximeters have not gained wide popularity in pulse oximetry because the sensor and emitter are on the same side of the finger, which increases scattering. The reflectance type of sensor is also more difficult to construct due to the need to control the angles of light emission. For this reason, we chose to construct a transmittance type of sensor where the light and sensing element are on opposite sides of the finger. This type of arrangement provides the most stable and reliable output. It is also the most commonly used arrangement in commercial devices.

Both transmittance and reflectance oximeters share LED technologies and absorption coefficient data. The absorption coefficients used by both techniques differ only because of each instrument's inherent geometry and circuit components. The source of light (nonfocussed LEDs), the angle of light emission, and the intensity all play large roles in determining the absorption coefficients. Data for monochromatic light have been reported throughout the literature. It would be desirable to achieve standard absorption coefficients for the broader bandwidth LEDs as light sources.

4. Three-Component Oximeters

Oximeters have distinguished only two components: hemoglobin and oxygenated hemoglobin. Also, present systems yield inaccurate results when substantial amounts of carbon monoxide are bound to blood [9]. Clinical testing has shown that most oximeters suffer from nonlinearity below 70% SaO_2 and that carboxyhemoglobin, methemoglobin, and sulfhemoglobin are the cause of some of these errors [10]. The ability to measure a third component, carboxyhemoglobin, can offer a significant improvement over present pulse oximeters.

This paper presents a system to measure three components of hemoglobin bonding based on the presence of one of those components, carboxyhemoglobin. A general background on oximeter theory is presented, which concludes with the equations to be solved. Presented are theoretical curves which simulate the anticipated results and the resulting calculations. An outline of the oximeter's hardware, software, and sensor are covered. Further, the problems encountered with noise in the system are examined.

5. Beer's Law

Traditional oximeters have solved Beer's law for two components: oxygenated hemoglobin (HbO_2) and reduced hemoglobin (Hb). Beer's law is:

$$I = I_0\, 10^{-ecd} \tag{1}$$

where I = transmitted light, I_0 = incident light, e = absorption coefficient of the substance at some wavelength, c = concentration of the absorbing substance, and d = optical path length through the substance. Transmission T is related to absorption A by:

$$A = -\log T \tag{2}$$

where

$$T = I/I_0 \tag{3}$$

thus yielding

$$A = -\log ecd \tag{4}$$

Equation (4) can be expanded to include the effects of multiple absorbing materials.

$$A = e_1c_1d + e_2c_2d + e_3c_3d \ldots \tag{5}$$

We substitute the absorption coefficients for three substances and measure the absorbance at three wavelengths to obtain three equations:

$$A_1 = e_{11}c_1d + e_{12}c_2d + e_{13}c_3d \tag{6}$$

$$A_2 = e_{21}c_1d + e_{22}c_2d + e_{23}c_3d \tag{7}$$

$$A_3 = e_{31}c_1d + e_{32}c_2d + e_{33}c_3d \tag{8}$$

If we assume that only three components exist, namely Hb, HbO_2, and HbCO, we can make the following substitutions: A_1, A_2, and A_3 correspond to the absorbances at 660 nm, 805 nm, and 940 nm, respectively, which were chosen due to the commercial availability of LEDs at these wavelengths and their fairly wide spectral separation from each other; c_1, c_2,and c_3 correspond to the concentrations of Hb, HbO_2, and HbCO, respectively; e_{11}, e_{21}, and e_{31} correspond to the absorption coefficients of Hb at 660 nm, 805 nm, and 940 nm, respectively; e_{12}, e_{22}, e_{32} correspond to the absorption coefficients of HbO_2 and e_{13}, e_{23}, and e_{33} correspond to the absorption coefficients of HbCO.

Dividing (6) by (7) and (8) by (6) we can eliminate the distance component d. Note that the arterial inflow of blood can cause a time dependent variation of d, even though this value would prove to be slight. Nevertheless, whether d remains constant, assuming rigidity of the finger artery, or varies to a certain extent, its variability can be eliminated by dividing the elements of equation (6) by (7):

$$\frac{A_1}{A_2} = \frac{e_{11}c_1 + e_{12}c_2 + e_{13}c_3}{e_{21}c_1 + e_{22}c_2 + e_{23}c_3} \tag{9}$$

and (8) by (6):

$$\frac{A_3}{A_1} = \frac{e_{31}c_1 + e_{32}c_2 + e_{33}c_3}{e_{11}c_1 + e_{12}c_2 + e_{13}c_3} \tag{10}$$

We also assume that only three products are attached to hemoglobin (Hb, HbO_2, HbCO) and that these make up 100% of the hematocrit, thus a third equation is $c_1 + c_2 + c_3 = 100\%$, where c_1, c_2, and c_3 are expressed as percentages.

For case 1, where we calculate A_{660}/A_{805} we can substitute the value of c_1 with $c_1 = 100 - c_2 - c_3$. At 805 nm $e_{21} = e_{12}$, thus

$$\frac{A_{660}}{A_{805}} = \frac{100e_{11} + c_2(e_{12} - e_{11}) + c_3(e_{13} - e_{11})}{100e_{21} + c_3(e_{23} - e_{21})} \tag{11}$$

This reduces to a set of linear equations where the slope m_1 and the intercept y_1 are functions of c_3.

$$\frac{A_{660}}{A_{805}} = m_1c_2 + y_1 \tag{12}$$

$$m_1 = \frac{(e_{12} - e_{11})}{100e_{21} + c_3(e_{23} - e_{21})} \tag{13}$$

and

$$y_1 = \frac{c_3(e_{13} - e_{11}) + 100e_{11}}{100e_{21} + c_3(e_{23} - e_{21})} \tag{14}$$

A similar formulation can be made for A_{940}/A_{660}, however the slope and intercept, will vary as a function of c_2, making the ratio nonlinear.

$$\frac{A_{940}}{A_{660}} = m_2 + y_2 \tag{15}$$

$$m_2 = \frac{(e_{32} - e_{31})}{100e_{11} + c_2(e_{12} - e_{11}) + c_3(e_{13} - e_{11})} \tag{16}$$

$$y_2 = \frac{100e_{31} + c_3(e_{33} - e_{31})}{100e_{11} + c_2(e_{12} - e_{11}) + c_3(e_{13} - e_{11})} \tag{17}$$

6. Pulse Oximetry

Pulse oximetry assumes that the pulsatile arterial inflow causes a modulation in transmittance [2],[11]. If the sensor measures a dc level (E_{dc}) during diastole, arterial pulsations cause an increased attenuation in the amount of light that is transmitted during systole. This variation can be termed the ac response (E_{ac}). The dc level is entirely due to nonchanging entities such as venous blood, tissue, bone, skin, etc. Transmittance is given by:

$$T = \frac{E_{dc} + E_{ac}}{E_{dc}} \tag{18}$$

By taking the log of T, we obtain the absorbance A and can then substitute into (6), (7), and (8).

7. Absorbance Ratios

Fig. 1 shows a 3-D representation of the two different ratios of absorbances plotted on the x and y axes versus the hemoglobin saturation on the z axis. Isolateral lines represent variation of HbCO. Thus all five varying factors are plotted on one graph. The surface therefore represents the complete solution to Beer's law with our three components, but presents difficulty in estimating the actual saturation since there may be multiple solutions to concurrent ratios.

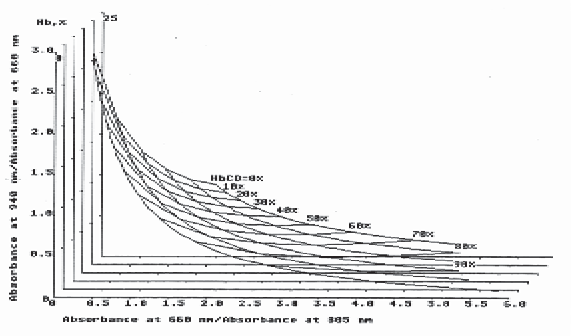

Fig. 1: A three-dimensional representation of carboxyhemoglobin plotted as a function of the hemoglobin saturation and absorbance ratios at 940 nm/660 nm and 660 nm/805 nm.

The 2-D graphs of Fig. 2, Fig. 3, and Fig. 4 are more informative. The A_{660}/A_{805} ratio and A_{940}/A_{660} yield two very different plots. The A_{660}/A_{805} ratio yields a linear plot whereas the A_{940}/A_{660} yields a plot with varying intercept and slope as predicted by (16) and (17). This is due to the isosbestic point (where the absorbances are equal) in ratio 1 versus the lack of one in ratio 2.

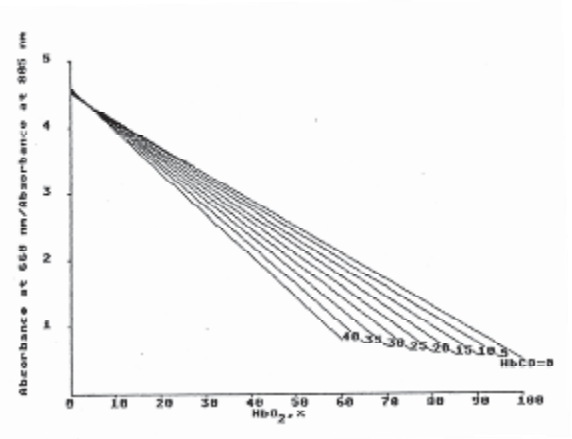

Fig. 2: The plot of absorbance at 660 nm/ absorbance at 805 nm versus the percentage of oxyhemoglobin (HbO_2). The different lines represent stepwise variations in carboxyhemoglobin (HbCO).

8. Sensor

It was important to design the shape of the sensor to avoid transmission of stray light and entrance of ambient light. Ambient light entered the sensor and caused variability in dc output levels. Inclusion of a back panel blocked the ambient light and minimized this problem. Without finger pads, the nonfocussed LEDs permitted light to scatter around and from the finger to the sensor, saturating it and making it highly dependent on finger motion. Slight movement of the finger caused changes in the dc baseline setting of the sensor and errors in the results. Black silicone rubber pads which absorb the infrared as well as the red light minimized this problem. The pads contained 13 × 13 mm square holes which allowed some finger movement without causing much variation in displayed output. As long as the finger covered the square, little variation occurred. However, allowing light to pass upward and around the finger once again caused error.

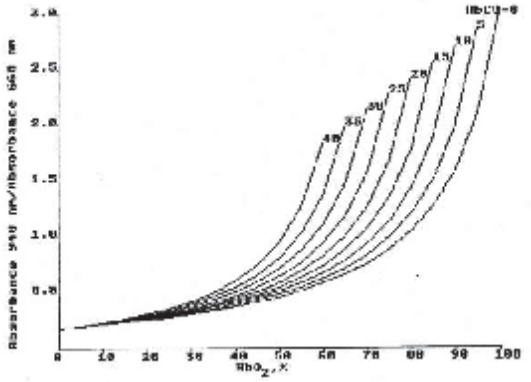

Fig. 3: The plot of absorbance at 940 nm/absorbance at 660 nm versus the percentage of oxyhemoglobin. The different lines represent stepwise changes in carboxyhemoglobin.

Spring pressure proved to be a critical aspect of the sensor. The pressure exerted on the finger by a spring force of 0.5 N seemed to hold the sensor adequately without causing loss of circulation to the finger. Forces greater than 0.5 N caused discomfort and loss of pulse when applied for great lengths of time.

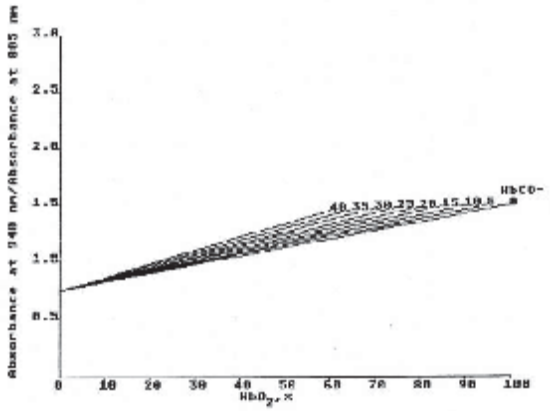

Fig. 4: The plot of absorbance of 940 nm/absorbance at 805 nm versus the percentage of oxyhemoglobin. The narrow separation between varying carboxyhemoglobin levels indicates difficulty in separating out carboxyhemoglobin at this set of wavelengths.

Besides the finger sensor, nose sensors and ear sensors are also used to measure oxygen saturation. These other sensors have the advantage of not being subjected to motion as are the finger sensors. The finger sensor can easily be implemented into an ear probe simply by changing the hardware assembly. A nose sensor would require a different design. Neither of these two sensors has yet been designed, but our system includes two connector control pins and software routines which will recognize the presence of these other two sensors.

The peak wavelengths of the LEDs are 660 nm, 810 nm, and 940 nm with bandwidths of 20 nm, 50 nm, and 50 nm, respectively. The LED dies are available from Silicon Sensors. The detector is a Siemens Corp. BPX-79 photovoltaic cell. Its rise time is 6 μs and fall time is 10 μs for 10% to 90% of minimal and maximal output. Maximal wavelength sensitivity is 800 nm. Sensitivity is 0.1 μA/lx minimum. The sensor assembly utilizes surface mount technology for its operational amplifier, resistors, and capacitors, all of which are mounted within the sensor compartment. Including the operational amplifier in the sensor assembly prevented interference problems from occurring within the 2.6 m length of nonshielded cable.

The LEDs are all rated between 10 to 20 mCd and provide more than adequate illumination at approximately 25 mA of current drain during normal operation. The on time is 100 μs for each LED and the overall cycle time is 1/300 s. Fig. 5 shows the circuit diagram for the sensor.

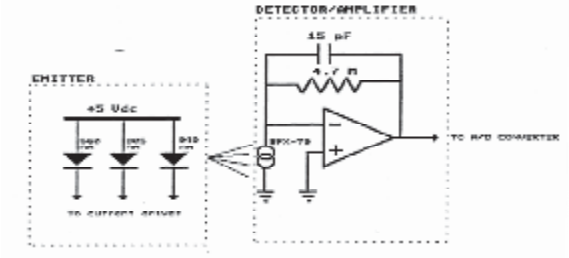

Fig. 5: The circuit diagram inside the sensor shows the three LED dies transmitting to a photovoltaic cell and amplifier which amplifies and filters the output.

9. Hardware

The entire system is designed around a Hitachi 64180 CMOS microprocessor which is supported by a 32-kbyte EPROM and 2 kbyte of RAM. Information is displayed on a customized LCD display. Data are retrieved via a 12 bit A/D converter which accepts the sensor output and digitizes it for input to the processor. The microprocessor controls the current driver. Each LED receives ramp inputs and self calibrates output of the detector amp to a preset dc level (1.25 V) which is controlled via the microprocessor. Each LED is connected to a multiplexed output of the current driver and strobed in sequence after which a delay time occurs for energy conservation. Fig. 6 shows the hardware layout.

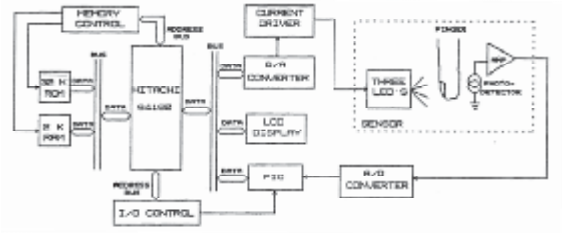

Fig. 6: Block diagram of the three-wavelength oximeter.

10. Software

Software controls the LCD display, the current drivers, the data storage and the subsequent data analysis. The received data are integrated to stabilize the resulting values and eliminate erroneous data points. The software also has self checking routines for determining whether the finger is inserted, whether power is low, detection of a low pulse, etc. It also has built in safety features detecting the presence of low or high pulse, high levels of HbCO and low levels of HbO_2. An alarm sounds when any of these conditions are present. The sensor receives full software control through the current driver. Each LED is stepped in increments as a self test to indicate whether the sensor is working properly. The software also is capable of determining whether finger, ear, or nose sensor is connected to the unit. Fig. 7 shows the flowchart of the software algorithm for calculating data.

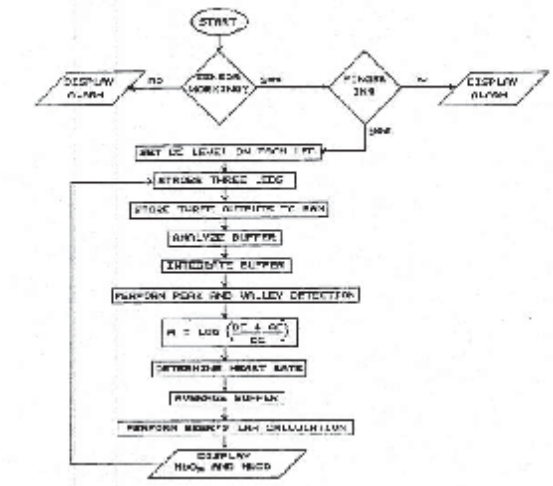

Fig. 7: Flowchart for general control and signal processing of the oximeter.

11. Absorption Coefficients

It would be easiest to use monochromatic sources of light to calibrate an oximeter. Any changes due to components and geometry could be made by simply scaling all the values by the same factor or making small changes in individual coefficients. For low cost and compact size, we use LEDs which are not monochromatic, but have a small to medium bandwidth.

Incorporating individual LEDs creates some problems, but these can be overcome. The use of nonfiltered LEDs has been reported previously [12] where it was stressed that the difference between absorption coefficients for oxygenated and deoxygenated hemoglobin was of great importance. However, we have found that the concentration of reduced hemoglobin and carboxyhemoglobin both make up the deoxyhemoglobin and that both have considerably different spectral characteristics. Thus, the absorption coefficients for both substances must be established along with the base value

for oxygenated hemoglobin. At 940 nm and 805 nm, carboxyhemoglobin has a much lower absorption coefficient than reduced hemoglobin and therefore using the equations for two components causes large error when HbCO contamination is present.

12. Clinical Procedure

Finding the absorption coefficients required clinical verification and actual drawn samples. The absorption coefficient values were preset to values found in the literature on oximetry [12–15] although values tend to vary for each article. A sample of blood was then drawn. Its blood gas concentration was analyzed by an ICL 282 oximeter. The values recorded from the ICL 282 were compared to displayed results on our own oximeter. By knowing the actual concentrations of Hb, HbO_2 and HbCO and also the different absorbance ratios, it was possible to solve for the absorption coefficients by plugging the known components into equations 9 and 10. The first blood sample was fully oxygenated. The patient breathed pure O_2 and blood was drawn by arterial puncture. This yielded ratios for e_{12}, e_{22}, and e_{32}. Since the absorption coefficient of Hb at 805 nm is near e_{22} then $e_{22} = e_{21}$. The absorption coefficient at 650 nm for HbO_2 and HbCO is similar so that $e_{12} = e_{13}$. Four coefficients remain to be determined: e_{11}, e_{23}, e_{31}, e_{33}. These are Hb at 660 nm and 940 nm and HbCO at 805 nm and 940 nm.

The patient was allowed to breathe a lung diffusion mixture containing 0.4% CO balance air. After 100 breaths the HbCO level of the subject had increased to approximately 15–17% HbCO. The strong binding of CO proved useful because very little Hb remained. Thus the ratios depended on only two components HbO_2 and HbCO. We used the absorption coefficient approximation of HbCO at 660 nm as a starting point. It was a simple matter of finding the necessary coefficient at 940 nm to make the ratio found equal to the theoretical ratio. Likewise at 805 nm, the ratio was used to find the absorption coefficients of HbCO at 805 nm. The absorption coefficient of HbCO at 805 nm and 940 nm is small and contributes little to the overall ratio. Thus it must be known accurately to avoid errors in other two absorption ratios.

13. Noise

During the testing, it was noted that the noise component of the transmitted signal was greatly amplified after the log of the absorbance was made. This effect is significant when considering the transparent qualities of HbCO at 805 nm and 940 nm. A noise free environment would be beneficial when making a measurement in the infrared region.

One possible solution to this problem would involve utilizing a lower wavelength LED at 600 nm. In this region, HbCO loses its transparent qualities and exhibits a significantly higher absorption coefficient, somewhere between Hb and HbO_2. Below the 600 nm region, the three components begin to exhibit very similar absorption coefficients and would therefore prove difficult to separate. Therefore, a 600 nm LED would take the place of the 940 nm LED. Whether this would remedy the noise problem remains a question.

14. Interfering Substances

Under certain circumstances the percentage of methemoglobin and sulfhemoglobin would be substantial and cause error in our oximeter. Typically, the levels are relatively low and do not cause error for normal healthy individuals. However, patients taking drugs may have their iron oxidized from a ferrous form to a ferric form [16]. This methemoglobin is unable to take up oxygen and therefore has a different absorption spectrum. It may also occur under normal conditions in healthy individuals. Sulfhemoglobin conversion also occurs for certain drugs and causes irreversible inactive O_2 transport. Detection of both would be useful, but would greatly increase the complexity. Such detection is already possible through invasive optical techniques [10],[17].

15. Conclusion

We have presented the method and design of an instrument to measure the concentration of hemoglobin, oxygenated hemoglobin, and carboxyhemoglobin. Beer's law defines the three absorption equations at three separate wavelengths for nonfocussed LED sources. The plots of the three-dimensional and two-dimensional solutions to Beer's law help show an efficient way to calibrate an oximeter based on a unique set of absorption coefficients inherent to a given instrument.

Observations made during this study showed it was necessary to add absorbing materials in the form of black plastic and black finger pads to eliminate scattering which caused fluctuating dc levels. Further, we noted that unusually high levels of sulfhemoglobin and methemoglobin would cause erroneous data to result.

Funding for this project ceased in the middle of the calibration procedure. For this reason, it has been impossible to validate the accuracy of the instrument to date. Nevertheless, we feel the information conveyed will help expand the field of pulse oximetry.

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