Cuff-Less Continuous Blood Pressure Monitoring System Using Pulse Transit Time Techniques

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Abstract: This paper describes the development of a continuous cuff-less blood pressure system based on the pulse transit time (PTT) technique. In this study, PTT is defined by two different approaches denoted as PTT_1 and PTT_2. PTT_1 is the time difference between the R-wave peak of the Electrocardiogram (ECG) and the peak of the Photoplethysmogram (PPG). PTT_2 is the time difference between two peak PPG signals on same cardiac cycle at different positions on the body. The ECG is acquired on the chest using 3 lead electrodes and a reflection mode optical sensor is deployed on brachial artery and fingertip to monitor the PPGs. These data were synchronized using a National Instruments data acquisition card along with Matlab software for subsequent analysis. A wrist-type cuff-based blood pressure device was used to measure blood pressure on the right hand. Brachial blood pressure was measured on the upper left arm using oscillometric blood pressure monitor. Experiments were conducted by elevating the right hand at different position to investigate variability of PTT under the effects of hydrostatic pressure. Next the variability of PTT due to blood pressure changes during a Valsalva maneuver was investigated. The result shows that the PTT_1 is inversely proportional to blood pressure in both experiments. Meanwhile, there is weak correlation between PTT_2 and blood pressure measurement which suggests that by excluding the pre-ejection period (PEP) time in PTT calculation may reduce the accuracy of PTT for blood pressure measurement. In conclusion, PTT measurement between ECG and PPG signals has potential to be a reliable technique for cuff-less blood pressure measurement.

Keywords: photoplethysmogram, electrocardiogram, pulse transit time, blood pressure

1. Introduction

Blood pressure (BP) is the pressure measured in millimetres of mercury (mmHg) exerted on the wall of a blood vessel as the blood passes through it. Two pressures are measured during a blood pressure reading: systolic blood pressure is a measure of blood pressure while the heart is beating and diastolic pressure is a measure of blood pressure while the heart is relaxed. Human blood pressure varies over 24 hours due to the changes of the human biological system. The blood pressure is usually high in the daytime and will slightly drop on the night. A 24 hour blood pressure reading can be obtained by an ambulatory blood pressure monitor (ABPM). This device basically is a small digital blood pressure machine that is attached to belt around body and a cuff connected on the upper arm. The machine takes blood pressure every 15-20 minutes during the daytime and 30 minutes at night [1]. The measurement data will be used by doctor to monitor hypertension [2] and to see the reaction of patients to specific drugs [3]. Some patients may feel discomfort, be distracted and be inconvenienced to have their blood pressure measured by the cuff based ABPM [2,4]. Furthermore, the cuff based blood pressure measurement device gives BP reading over a 3 minute interval with no information on continuous blood pressure values. A continuous BP measurement is desirable for applications such as for assessing orthostatic hypotension [5] and during caesarean section surgery [6].

Pulse Transit Time (PTT) is one of promising method for cuff-less continuous blood pressure measurement. Most researchers define PTT as the delay time between the R-wave peak of the Electrocardiogram (ECG) and the peak of the Photoplethysmogram (PPG). The delay time between these two signals is shown to be correlated with blood pressure [7-10]. Increase in blood pressure causes vascular tone to increase and hence the arterial wall becomes stiffer causing the PTT to shorten. Conversely, a drop in blood pressure causes arterial wall tension and stiffness decrease thus increasing the PTT. This phenomenon can be explained by referring to the Moens-Korteweg Equation 1 [11]:

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$$PWV = \frac{D}{PTT} = \sqrt{\frac{Ea}{\rho d}} \qquad (1)$$

where *PWV* is pulse wave velocity (m/s), *D* is length of the vessel (m), *PTT* is pulse transit time (s), *E* is elastic modulus of wall tissues (pa), *a* is wall thickness (m), ρ is the density of the contained blood within the vessel (kg/m^3) and *d* is the interior diameter of the vessel (m). Huges et al. reported that *E* increases exponentially with increasing blood pressure according to the Equation 2 [12]:

$$E = E_O e^{\alpha P} \qquad (2)$$

where α is constant (mmHg^{-1}), E_0 is the zero-pressure modulus of the vessel wall and *P* is the BP (mmHg) within the vessel. This assumes that, in practice, the change in elasticity is much greater than the change in wall thickness and diameter of the vessel. Thus, PWV will increase with increase of blood pressure and the PTT inversely proportional to PWV.

In general, the ECG signal is acquired on the chest and the PPG signal at the finger for pulse transit time measurement [13-17]. This arrangement has a great attention due to availability of the measurement devices and to minimize error in the time domain due to distance between the heart and finger. Furthermore, these two signals are often simultaneously recorded by patient monitors in hospital and can be accessed through patient's database system for PTT analysis [18]. The major drawback from this technique is the motion artefact on the PPG signal due to finger movement [19] and movement of arm position relative to vertical body which causes error in PTT measurement [20].

Therefore, other researchers used different locations on the body to acquire the PPG signal. One of the most promising parts of the body to obtain a PPG signal using transmission mode optical sensor is at the earlobe. The PPG signal at earlobe gives smaller error in PTT measurements because there is less effect of movement and the hydrostatic pressure is much more constant when the person moves from supine to standing position [21]. Studies by Jeong et al. show that the PTT between ECG on the chest and PPG on the earlobe have a linear response with systolic BP, but requires individual calibration to estimate BP [20]. However it is reported that PPG on the ears comprised more artefact and noise when compared with PPG at the finger [22]. This problem could be overcome with specifically designed PPG circuit with signal processing techniques for monitoring PPG signal at the earlobe site.

The transmission mode optical sensor for PPG detection shows relatively good signal compared to reflection mode optical sensor, but it is limited only to the certain body parts. Using a reflection mode optical sensor, the PPG signal can be obtained almost anywhere on the body. This has led to the development of small wearable devices consisting of ECG and PPG sensors for calculating PTT on the chest [23]. The device has a compact size and is able to monitor changes in blood pressure with time during exercise. The only concern for the device is the measurement accuracy as it is still has a large measurement error when compared with available devices on the market [23].

Another way to determine PTT is by measuring the delay time between two PPGs signal at different locations on body. Through this method, PTT measurement devices will have simple interface system compared with the technique that uses ECG measurement which requires electrodes and additional circuitry. Jeong et al. had proposed PTT measurement using two PPG signals at earlobe and sternoclavicular and correlated this measurement with systolic blood pressure (SBP) during body postural change [9]. They estimated SBP in different body posture with a single equation. Furthermore, Amin et al. have shown that BP is highly correlates with PTT of two PPG signals during exercise [10]. They developed two identical optical sensor circuits to monitor PPG signals at finger with the distance between the sensor set to 55mm. PTT measurement between two PPGs pulses means that the pre-ejection period (PEP) measured in the ECG is excluded in PTT calculations. PEP is the time between the R-peak of the ECG and the opening of the aortic valve. Reports show that PEP has great contribution to the PTT change during daily activities such as posture change and dynamic exercise [3, 16]. Findings from Proenca et al. show that there is poor correlation between PTT and systolic BP in healthy young subjects during a sequence of short-term physical exercise if PEP is excluded from the PTT measurements [24]. This finding contradicts with that reported by Jeong et al. and Amin et al.. Therefore, further research is needed to clarify uncertainty relation between PTT of two PPGs pulses and blood pressure measurement.

Therefore, the aim of this study is to determine the best approach in determining PTT for blood pressure estimation. In this work, we defined PTT using two different approaches denoted as PTT_1 and PTT_2. PTT_1 is the time difference between the R-wave peak of the Electrocardiogram (ECG) and the peak of the Photoplethysmogram (PPG). PTT_2 is the time difference between two peak PPG signals on same cardiac cycle on the same arterial pathway. These positions were chosen to eliminate the different arrival time of PPG signals due to gravitation [25]. Besides, using the same arterial pathway will satisfy the assumption made in Moens-Korteweg model which used an infinitely long and straight vessel with elastic, isotropic, and homogenous walls for estimating pulse wave velocity [26]. Based on Equation 2, it is noted that vessel elasticity, *E* largely depends on blood pressure, thus make PTT decrease with increase of blood pressure.

The advantage and novelty of this study is that two different approaches of PTT measurement using cuff-less blood pressure can be compared simultaneously. All the experiments were conducted on one subject. The result from this study is an initial step in clarifying the contradicting findings in the literature on the PTT techniques for blood pressure estimation.

2. Experimental setup

The system block diagram for measuring PTT is shown in Figure 1. A reflectance mode optical sensor was used to capture the PPG signal on the brachial artery and fingertip. The PPG circuit consists of a green LED with wavelength of 530nm, ambient light photo sensor (APDS-9008) and operational amplifier. The LED was driven by 5 volt dc voltage. The green light that is transmitted to the skin will be absorbed by the pulsatile blood, and reflected light will be captured by the photo sensor. The sensor converts the light intensity into voltage and an operational amplifier is used to amplify the signal. Meanwhile, ECG board (STM32 ECG/EMG, Olimex) is used to monitor ECG signals on chest. The circuit consists of an instrumentation amplifier and active filter to convert electrical potential generated by electrical activity in cardiac tissue into readable voltage. A National Instruments data acquisition card is used to transfer the data to a computer and Labview software is used to visualize the waveform and recorded the signals. PTT_1 and PTT_2 was computed offline using Matlab software. Peak detection algorithm was developed to determine the peak of the ECG and PPG signals for PTT measurements.
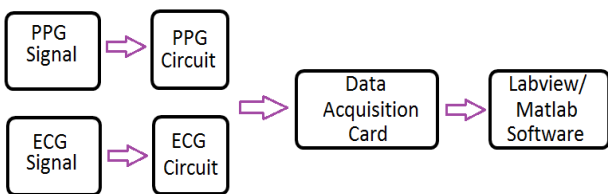

Fig.1 System block diagram

In these studies, two experiments were performed by a healthy male adult, age of 35 years to investigate the correlation of PTT with BP. First experiment was conducted by elevating hand above heart level. This method will change the blood pressure due to the effect of hydrostatic pressure [27]. This experiment was conducted in seated position with the arm kept constant at the heart level. The ECG electrodes were placed on the chest and measured the ECG signal according to lead II configuration. PPG sensors were attached on the brachial artery and on the middle finger of the right arm. A wrist-type cuff-based blood pressure (OSTAR M100) was used to measure blood pressure on the right hand. A cuff based blood pressure monitor (OSTAR P2) was worn on the left upper arm for monitoring brachial blood pressure as a reference measurement. After 5 minutes seated in relaxed condition, the subject's right hand was stretched out without bending and lifted to different positions above heart level without support. The height level was set to be 0, 15, 30, 45 and 60 cm, with 0 cm representing the base level of wrist position level with the heart. The right arm was maintained at these positions for 1 minute to record the ECG and PPGs signal. Simultaneously, radial blood pressure and brachial blood pressure were recorded.

In the second experiment, the subject performs a Valsalva maneuver to vary blood pressure. This technique used force exhalation against a closed airway to change

the blood pressure and has been applied by other researchers in their experiments [28, 29]. Shin et al. proposed that the R–J interval from ECG signal on the chest and feet can be used to estimate systolic BP continuously [28]. Authors correlated R-J interval with BP using Valsalva techniques. The same technique was used by Junior et al. to estimate the change of the blood pressure using smartphone [29]. In previous works the PTT was defined as the delay time between the heart sound and the ECG signal. In this research, we propose to measure PTT as the time delay between ECG to PPG signals and PPG to PPG signals simultaneously.

There are 4 phases involved in blood pressure changes throughout the execution of Valsalva maneuver [28]. In first phase, systolic BP will increase due to the initial inspiration. In second phase, the systolic BP will be maintained as the subject maintains the expiratory strain. In the third stage, the systolic BP drop to minimum level as breath released. In the last stage, systolic BP will return to pre-maneuver level again. The variability of the blood pressure during each phase of Valsalva maneuver was monitored using PTT techniques. In this experiment, the subject will rest in a seated position for 5 minutes before performing the Valsalva maneuver. The left and right hand are in rest positon on a table that is level with the heart. Blood pressure monitor (OSTAR P2) is used to measure the blood pressure on the upper left arm before and during the Valsalva maneuver process. The ECG signal was recorded on the chest using lead II configuration. PPGs signal on the brachial artery and on the middle finger of the right arm were recorded using reflection type optical sensor. The experiment was repeated 10 times and results are presented in the next section.

3. Results and discussions

Figure 2 shows the ECG waveform acquired on the chest (green trace) and the PPG waveforms acquired on the brachial artery (pink trace) and middle finger (blue trace). All signals were measured simultaneously and plotted in voltage and time scale.
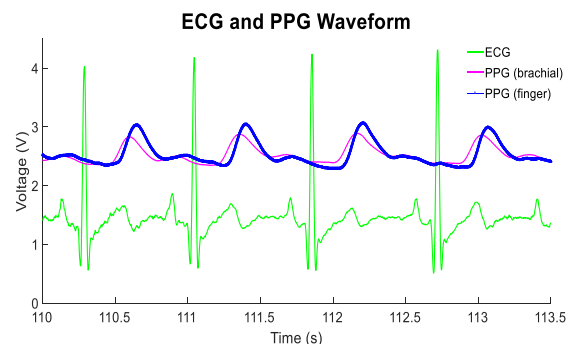

Fig. 2 ECG and PPGs waveforms

The ECG signal represents the electrical activity of the heart contraction. As the heart contracts, the oxygenated blood will pump out from the heart to the body through the aorta. For the upper body path, the aorta leads the blood to the brachial arteries and fingertips via subclavian arteries.

The blood flow at the body part can be measured using reflectance mode PPG sensor. As shown in Figure 2, the ECG signal is triggered first followed by the PPG brachial signal that occurs before the PPG at the finger. For 30 pulses, the average time of PTT_1 for ECG to PPG finger and ECG to PPG brachial artery was 367 ± 7.7ms and 312 ± 6.7ms respectively. The average time for PTT_2 which is the time difference between the two PPG pulses was 54 ± 5.6ms. In general, the PTT values are different between each subject and depend on physiological factors and the measurement circuit used [7]. For instance, it has been reported that arterial walls become thicker and the elastic properties of the large arteries will reduced as the age increases [7]. This phenomenon leads to a decrease in Young's modulus of arterial wall thus increasing the PTT. Besides, different filter design for ECG and PPG circuit may generate phase lag between those signals, thus lengthening the PTT [30]. From the literature, the PTT between ECG to PPG at the fingertips is in the range 300ms to 400ms [30] and PTT of two PPG signals between brachial and fingertip is reported around 50ms [31-32]. The position and the pressure applied on the sensor to the skin may also affect the PTT readings. Teng et al. have reported that the PTT will decrease as the contact force on the sensor probe increases [33]. Therefore, this force pressure on the sensor should be taken into account during PTT measurement to increase the accuracy. Moreover, individual calibration procedure is needed to obtain absolute blood pressure measurement using PTT techniques.

Figure 3 shows the experimental results for elevating hand at five different positions. The relationship between PTT_1 and radial systolic blood pressure (SBP) is shown in Figure 4. In this experiment, the right hand was elevated starting from 0 cm to 60 cm above heart level. The PTT_1 between ECG to PPG at finger and ECG to PPG at brachial artery were recorded simultaneously.
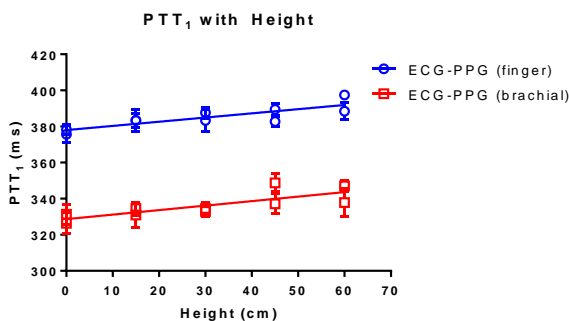

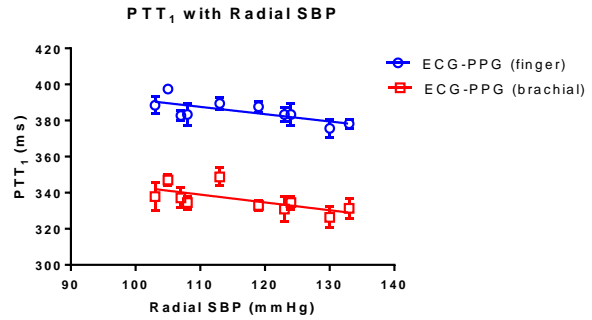Fig. 3 PTT with different wrist height level

Fig. 4 PTT with radial systolic BP

The results in Figure 3 show that PTT_1 steadily increases when the right hand is elevated above heart level. This condition resulted from the increase of hydrostatic pressure due to the increase of height level. A similar pattern was reported by Poon et al. in their experiments on 11 subjects [34]. Their findings suggest that this technique can be used for PTT calibration in order to perform continuous BP measurements. During the PTT measurement, the radial blood pressure on the right wrist and brachial blood pressure on the upper left arm were recorded simultaneously. The brachial BP measurement is used as reference BP and result shows that the BP reading was unchanged when the subject lifted up his right hand. In contrast, radial blood pressure slightly dropped as the hand is elevated above heart level due to hydrostatic pressure. The relationship between PTT_1 and radial SBP is shown in Figure 4. The result shows that the PTT_1 decrease with increase of systolic radial blood pressure during hand elevation experiments. The inverse linear relation between PTT_1 and BP is reported by many authors [35-37].

In this experiment the correlation analysis between PTT_1 and BP was performed using GraphPad Prism software. The correlation coefficient, R between radial BP and PTT_1 (ECG to PPG finger) was -0.7105 for the systolic BP and -0.7103 for diastolic BP. Meanwhile, for radial BP and PTT_1 (ECG to PPG brachial), the R values for the systolic BP was -0.6676 and -0.5308 for diastolic BP. This coefficient correlation value R is greater than -0.5 which indicates that PTT_1 have a good negative relation with BP. The different time between peak of PPG finger and peak of PPG brachial (PTT_2) was calculated and plotted with radial BP. Based on the plotting, correlation coefficient between radial BP and PTT_2 was determined. The result shows that the R value was 0.0065 for systolic BP and -0.1676 for diastolic BP. These R values indicated that there is no correlation between BP and PTT_2. A second experiment was conducted by using Valsalva maneuver techniques. Figure 5 shows the PTT_1 responses with time during Valsalva maneuver process.
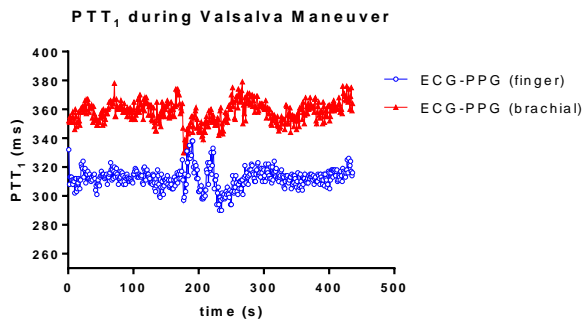
Fig. 5 PTT during Valsalva maneuver

From the Figure 5, both traces that represent PTT_1 changes with the time during the Valsalva maneuver period which occurred from 180s to 220s. The changes of PTT_1 indicated that there were changes in BP values. In this experiment, the variability of PTT with BP measurement before and during the Valsalva maneuver was investigated. The results from this study are shown in Figure 6 and Figure 7. Figure 6 shows the PTT_1 of ECG to PPG finger with SBP and Figure 7 shows the PTT_1 of ECG to PPG brachial with SBP. The blood pressure was recorded before and during Valsalva maneuver which is denoted on the graph as relaxes and release strain respectively.
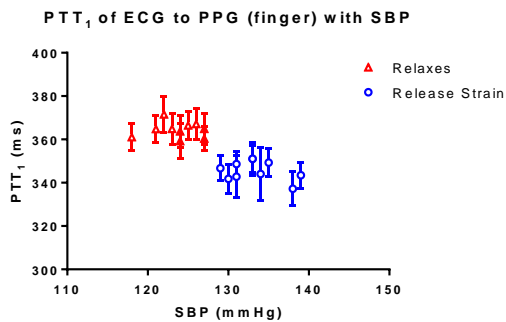

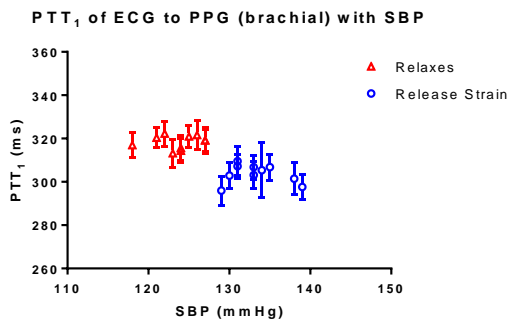Fig. 6 PTT_1 of ECG to PPG finger with SBP

Fig. 7 PTT_1 of ECG to PPG brachial with SBP

These results show that the PTT_1 has an inversely relation with SBP. According to the correlation analysis, the correlation coefficient R in Figure 6 and Figure 7 was -0.8255 and -0.7789 respectively. The same analysis was applied for the PTT_1 and diastolic BP and the result shows that the correlation coefficient R was -0.7270 for ECG to PPG finger and -0.6463 for ECG to PPG brachial. Next,

the time difference between PPG finger and PPG brachial (PTT_2) was determined and was plotted against systolic and diastolic BP. The results show that the R value for PTT_2 and systolic BP was -0.3336 and -0.4412 for diastolic BP. These R values indicate that there is weak relationship between BP and PTT_2. Both experiments revealed that there is correlation between BP and PTT_1 and these results agree with findings from the other authors [34-36]. Meanwhile, there is a weak correlation between BP and PTT_2 and this argument had been reported in the literature [3,16,24]. One possible reason for this non dependency was related to PEP. The exclusion of PEP in the PTT_2 calculation may reduce the accuracy of BP measurement with PTT [38].

In this work, there was a small data range of SBP of 25 mmHg for first experiment and 21 mmHg for second experiment. Nevertheless, the results obtained demonstrate that the PTT correlates well with the blood pressure changes. One of the possible applications of this system is development of a wearable device to monitor rapid modulations of blood pressure in falls detection system. The fall detection reliability can be increased by incorporating the blood pressure reading along with an accelerometer.

4. Summary

In conclusion, the PTT between ECG and PPG signal correlates with BP change in hand elevating and Valsalva maneuver experiments. Meanwhile the PTT between PPG to PPG signal shows a weak correlation with BP in both experiments. This finding suggests that PEP has a major contribution to PTT changes with BP. Studies in a large numbers of subject will be conducted to support this argument for the future work. Moreover, results obtained from this study will be the basis for future research in designing wearable continuous blood pressure monitors.

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