

# **Master's Thesis**

Medical Informatics Master

Universität Heidelberg / Hochschule Heilbronn

---

## **Predicting Blood Pressure from Photoplethysmogram Waveform Data: A Signal Processing and Machine Learning Approach**

# Contents

<b>1</b>	<b>Abstract</b>	<b>3</b>
<b>2</b>	<b>Introduction</b>	<b>3</b>
2.1	Subject and Motivation . . . . .	3
2.2	Tasks and Objectives . . . . .	4
<b>3</b>	<b>Theoretical Background</b>	<b>4</b>
3.1	Medical Background . . . . .	4
3.1.1	Blood Pressure . . . . .	4
3.1.2	Photoplethysmography . . . . .	6
3.1.3	MIMIC-III . . . . .	6
3.1.4	MIMIC-IV . . . . .	7
3.2	Computing Background . . . . .	7
3.2.1	Signal Processing . . . . .	7
3.2.2	Machine Learning . . . . .	8
<b>4</b>	<b>Methods</b>	<b>9</b>
<b>5</b>	<b>Results</b>	<b>10</b>
<b>6</b>	<b>Discussion</b>	<b>10</b>
<b>7</b>	<b>Conclusion</b>	<b>10</b>
	<b>References</b>	<b>10</b>

# 1 Abstract

## 2 Introduction

### 2.1 Subject and Motivation

Cardiovascular diseases (CVDs) are the leading cause of death worldwide, according to WHO publishing statistics [17]. One of the main factors contributing to CVDs is Hypertension. It is the leading risk factor for mortality, and is ranked third as a cause of disability-adjusted life-years [6]. Currently, there is a significant need for continuous blood pressure (BP) monitoring due to various factors. Primarily, while hypertension is a manageable condition, the availability of accurate high BP detection remains scarce, especially in low-resource environments [3]. Additionally, blood pressure (BP) is subject to rapid fluctuations influenced by various factors, including stress, emotions, dietary intake, physical activity, and medication usage [19]. Continuous monitoring of blood pressure, rather than relying on isolated measurements, plays a vital role in the early detection and treatment of hypertension [5].

The current accurate methods for measuring BP continuously are either invasive or involving a cuff-mechanism. Catheterization is internationally recognized as the "gold standard" for obtaining the most accurate measurement of continuous blood pressure [22]. However, due to its invasive nature and limited applicability to hospital settings, this method requires medical intervention, which renders it inconvenient for everyday use.

While cuff-based devices are commonly utilized for this objective, it is worth noting that over 30% of home blood pressure monitors are found to be inaccurate, rendering continuous measurement unfeasible [14]. Moreover, this approach relies on the individual consciously and intentionally engaging in manual blood pressure monitoring, which poses limitations and might be often overlooked.

An ideal technology for measuring blood pressure should have the following attributes: non-invasiveness, cuffless operation, optical functionality, wearable design, and cost-effectiveness [5]. One approach satisfying these requirements is the estimation of BP from a single measurement PPG sensor. This approach, using two modes, reflectance and transmission, has gained an increasing amount of attention in the literature due its simplicity, and ability to provide continuous and cuffless measurement [5]. Typically, the photoplethysmography (PPG) technique has been traditionally employed in healthcare settings to measure heart rate [20] and blood oxygen saturation using a pulse oximeter

[28].

Nevertheless, establishing a straightforward, distinct, and continuous relationship between these characteristics and blood pressure (BP) has proven to be challenging. To address this, the approach heavily depends on signal pre-processing techniques, extracting PPG features, and utilizing machine learning algorithms to estimate BP based on these features [5]. A recent scoping review by Knight et al. concluded that PPG can be successfully used to continuously measure BP, by evaluating latest publications and finding over 80% accuracy in detecting hypertension [11].

This study examines the current methods and aims to develop efficient approaches for the continuous and accurate measurement of blood pressure using PPG and addresses the following research questions:

1. **What is the relationship between photoplethysmogram (PPG) data and blood pressure among ICU patients?**
2. **Can PPG-based data be used to estimate blood pressure accurately?**

This thesis is organized as follows:

## **2.2 Tasks and Objectives**

The tasks of the thesis are as follows:

1. **Signal Processing:** to find an optimal data fetching and filtering approach from available MIMIC Databases. Likewise to create a consistent algorithm for key feature extraction.
2. **Machine Learning:** to develop a model based on the resulting features from Signal Processing, to reliably predict BP from PPG.

# **3 Theoretical Background**

## **3.1 Medical Background**

### **3.1.1 Blood Pressure**

1. Basic information about BP. How is it measured. What are the main methods. What is the significance of measuring BP. [1]

Arterial blood pressure (ABP) is a dynamic parameter that can change with each heartbeat, and it is typically represented as a waveform rather than a single numeric value. The arterial blood pressure waveform consists of two main components: systolic pressure (peak pressure during ventricular contraction) and diastolic pressure (lowest pressure during ventricular relaxation).

Continuous monitoring of arterial blood pressure is often done in clinical settings, especially in critical care environments, using an arterial line connected to a pressure transducer. The resulting waveform is displayed on a monitor in real-time. However, for ease of interpretation and documentation, numeric values are commonly extracted from the waveform at specific time intervals.

In situations where high temporal resolution is crucial, arterial blood pressure can be recorded beat-to-beat, providing a value for each heartbeat. This is particularly important in situations where rapid changes in blood pressure need to be closely monitored, such as during certain medical procedures or in critically ill patients.

For routine monitoring and documentation, numeric values are often averaged over a specific time interval, such as every 1, 5, or 15 minutes. This averaged value may be reported as the mean arterial pressure (MAP), which is a weighted average of the systolic and diastolic pressures over a cardiac cycle. Some monitoring systems may also provide systolic and diastolic blood pressure readings at regular intervals.

Blood pressure measured using a cuff, also known as non-invasive blood pressure (NIBP), is typically recorded as numeric values at specific time intervals. The process involves inflating the cuff to temporarily stop blood flow and then gradually releasing the pressure to detect the sounds associated with the flow of blood through the brachial artery. The two primary values obtained are systolic pressure (maximum pressure during heartbeats) and diastolic pressure (minimum pressure between heartbeats).

During the process of cuff inflation and deflation, each heartbeat generates characteristic sounds (Korotkoff sounds) that are detected by a stethoscope placed over the brachial artery. Systolic pressure is recorded when the first tapping sounds are heard, and diastolic pressure is recorded when the sounds disappear or change character. This beat-to-beat approach provides information about individual fluctuations in blood pressure.

Many automated blood pressure monitors, especially those used in clinical settings, provide averaged values over a specific time interval. The monitor may calculate mean arterial pressure (MAP) or provide systolic and diastolic blood pressure readings averaged over a set period, such as every 1, 3, or 5 minutes. Averaging over time helps

to smooth out transient changes and provides a more stable representation for clinical assessment.

Alternative approaches for measuring BP have emerged over the past years. Volume clamping [10] and tonometry [8] are some of the other methods. These non-invasive techniques offer continuous monitoring of blood pressure values. Volume clamping, which involves the use of a small finger cuff and a photoplethysmography (PPG) sensor, is one method for continuous blood pressure measurement. Tonometry, on the other hand, is a cuffless approach that utilizes a manometer-tipped probe pressed directly on an artery. The volume clamping approach allows for instantaneous and prolonged blood pressure measurement. However, it is associated with high costs and still necessitates the use of a cuff, which can be inconvenient and uncomfortable. Conversely, the tonometry method is sensitive to movement of the arm and probe, making it challenging to maintain accuracy in practical applications. Additionally, constant calibration with a cuff blood pressure device is required [18].

### **3.1.2 Photoplethysmography**

1. How does PPG work?

Basic information about PPG. How does it work. Main use cases. Potential use cases.

The PPG is an optical sensor, consisting of a light-emitting diode (LED) paired with a photodetector (PD), hence it is simple, inexpensive and can be easily build into a wearable device. The PPG waveform can be obtained using two modes, reflectance and transmission. This waveform corresponds to the blood volume in blood vessels. The PPG is primarily traditionally is utilised in healthcare for measuring heart rate and bloodoxygen saturation using a pulse oximeter [2].

Peripheral volumetric changes and BP are correlated [13]. Some characteristic PPG features can be used to estimate Systolic BP (SBP) and Diastolic BP (DBP) using machine learning functions. However, there is no simple, clear and continuous relationship between these features and BP. This approach relies heavily on signal pre-processing, extracting PPG features and employing machine learning algorithms for estimating BP as a function of these features.

### **3.1.3 MIMIC-III**

origin of MIMIC. When was it created? How is it structured?

### 3.1.4 MIMIC-IV

Multiparameter Intelligent Monitoring in Intensive Care IV (MIMIC-IV) [?].

1. How is the database structured?
2. Which patients and how many of them have both PPG and BP data?
3. Arterial Blood Pressure (ABP)

How does it differ from MIMIC3?

## 3.2 Computing Background

### 3.2.1 Signal Processing

1. What are the methods of signal processing for reading PPG?

Approaches for processing the given PPG data in correlation to BP:

**Time (PTT) based on PPG & ECG - Inverse correlation between BP and PTT.** PTT is the time delay for the pressure wave to travel between sites on the body. It can be calculated as the time difference between proximal and distal waveforms indicative of the arterial pulse.

**Pulse Arrival Time (PAT) = PTT + pre-ejection period.** It is defined as the time that takes the pulse wave to travel from the heart to a peripheral site e.g. finger, toe, etc. It can simply be estimated as the time delay between the R peak of the ECG waveform and a point on the rising edge of a distal PPG waveform.

These methods require simultaneous measurement at two different sites on the body, hence two measurement sensors (ECG and PPG) are needed for recording the signals in order to estimate these parameters.

2. What are the different types of filters for processing the PPG signal?

Signal filtering types include: Chebyshev filter, Butterworth filter [15].

Savitzky-Golay (SG) filter [7]

Second derivative & Age analysis [25]

Hemodynamics and vascular age [4]

### 3.2.2 Machine Learning

1. What are the methods of machine learning for estimating BP from PPG?

Approaches for estimating BP from PPG:

BP estimation using ML techniques is data driven, unlike the traditional PTT/PAT only models. Several studies attempted to fit regression models, such as **multilinear regression, support vector machine and random forest**, for estimating BP using PTT/PAT based approach with some degree of success, but the results did not always satisfy the international standards.

Teng and Zhang [26] tried to fit a **linear regression** model to study the relationship between four PPG features and BP. It was reported that the diastolic time has higher correlation with SBP and DBP than the other features.

Suzuki and Oguri [24] used **AdaBoost** classifier for the estimation of BP. In this technique, SBP values were classified according to a threshold and afterwards the nonlinear machine learning model was employed for estimating SBP.

Ruiz-Rodriguez et al. [21] employed a probabilistic generative model, **Deep Belief Network Restricted Boltzmann Machine**, for predicting SBP, DBP and mean arterial pressure simultaneously. The results of this study were highly variable, and therefore was not reliable.

Kurylyak et al. [12] extracted 21 characteristic features from the PPG waveform. These features were used for estimating SBP and DBP using a **feed forward neural network**. The results were promising towards an accurate cuffless BP monitoring.

Xing and Sun [27] applied **Fast Fourier Transformation** for selecting frequency domain features from the PPG waveform followed by a **feed forward neural network** for BP estimation. However, the authors suggested that these features are not sufficient for effective BP estimation.

Liu et al. [16], added 14 features extracted from the PPG's second derivate, in addition to the 21 features used in Kurylyak et al. A **support vector machine (SVM)** was then applied for estimating SBP and DBP. The authors reported that these 14 features further improved the estimation.

The relationship between BP and PPG features is not always linear. Therefore, linear models are inappropriate and often fail to model the relationship between BP and PPG



when tested on a large dataset collected from a diverse population. Other classical machine learning models, such as **SVM**, and **random forest**, provide better precision. Estimation using these models requires establishing one model per objective, hence, SBP and DBP are estimated separately. However, DBP strongly correlates with SBP and improve its estimation [49], thus should be modelled simultaneously using one model architecture. This can be achieved using neural networks. **Neural network** models can leverage large amount of data faster and more accurately compared to classical machine learning models.

El-Hajj and Kyriacou proposed using **Bidirectional Long Short-term memory (Bi-LSTM)** and **Bidirectional Gated Recurrent Units (Bi-GRU)** with **attention mechanisms** [5].

In Su et al. [23], a four layers **LSTM** with bidirectional structure and residual connections has been employed for BP estimation using the PTT approach. The reported results outperform other PTT based BP regression models.

A study by Joung et al [9] was conducted to evaluate a learning-based cuffless BP estimation system with calibration in challenging circumstances. A one dimensional CNN-based network was designed, that could efficiently extract BP from PPG signals using a comparative paired 1D-CNN structure with calibration.

To precisely design a learning-based BP estimation model such that its estimation accuracy obtained during the test is sustained after being built upon a practical cuffless BP monitoring system, the following delicate yet realistic experimental principles are applicable: i) the number of subjects should be sufficiently large, ii) subject independent training and test datasets are required, and iii) the intrasubject BP variation should be carefully scrutinized in the model design [9].

## 4 Methods

1. Data fetching and reading using WFDB and NumPy libraries.
2. Digital signal filtering using Savitzky-Golay and Butterworth Lowpass filters.
3. Beat detection algorithms from MIMIC WFDB Tutorial used for primary estimation. Beat detection improved with manual implementation of SciPy library.
4. Fiducial Point calculation based on the algorithm provided in the MIMIC WFDB Tutorial.

5. Feature extraction with personally created code to extract time domain features, and NumPy library to extract frequency domain features (FFT). Median value calculation using NumPy library.
6. Machine Learning Model creation using PyTorch and Scikit-Learn libraries.

## 5 Results

1st iteration:

306 records of 10 minutes each were extracted from the MIMIC-III Database for creation (training and testing) of a Machine Learning model. Signaling processing approaches were included (see 4.2). Feature extraction: - For target reference ABP Systolic and Diastolic values were extracted from the waveform data. - For Machine Learning, 34 features were extracted from the PPG waveform A total of 161269 values were extracted, resulting in a 34x161269 matrix.

## 6 Discussion

## 7 Conclusion

## References

- [1] What is blood pressure and how is it measured? In *InformedHealth.Org [Internet]*. Institute for Quality and Efficiency in Health Care (IQWiG).
- [2] J. Allen. Photoplethysmography and its application in clinical. 28(3).
- [3] V.L. Burt, P. Whelton, E.J. Roccella, C. Brown, J.A. Cutler, M. Higgins, M.J. Horan, and D. Labarthe. Prevalence of hypertension in the US adult population: Results from the third National Health and Nutrition Examination Survey, 1988-1991. 25(3):305–313.
- [4] Peter H. Charlton, Birutė Paliakaitė, Kristjan Pilt, Martin Bachler, Serena Zanelli, Dániel Kulin, John Allen, Magid Hallab, Elisabetta Bianchini, Christopher C. Mayer, Dimitrios Terentes-Printzios, Verena Dittrich, Bernhard Hametner, Dave Veerasingam, Dejan Žikić, and Vaidotas Marozas. Assessing hemodynamics from

- the photoplethysmogram to gain insights into vascular age: A review from VascAgeNet. 322(4):H493–H522.
- [5] C El-Hajj and given-i=PA family=Kyriacou, given=PA. Deep learning models for cuffless blood pressure monitoring from PPG signals using attention mechanism. 65:102301.
  - [6] Majid Ezzati, Alan D Lopez, Anthony Rodgers, Stephen Vander Hoorn, and Christopher JL Murray. Selected major risk factors and global and regional burden of disease. 360(9343):1347–1360.
  - [7] given-i=Abraham family=Savitzky, given=Abraham. and M. J. E. Golay. Smoothing and Differentiation of Data by Simplified Least Squares Procedures. 36(8):1627–1639.
  - [8] B.P.M. Imholz, W. Wieling, G.A. Van Montfrans, and K.H. Wesseling. Fifteen years experience with finger arterial pressure monitoring: Assessment of the technology. 38(3):605–616.
  - [9] Jingon Joung, Chul-Woo Jung, Hyung-Chul Lee, Moon-Jung Chae, Hae-Sung Kim, Jonghun Park, Won-Yong Shin, Changhyun Kim, Minhyung Lee, and Changwoo Choi. Continuous cuffless blood pressure monitoring using photoplethysmography-based PPG2BP-net for high intrasubject blood pressure variations. 13:8605.
  - [10] Chang-Sei Kim, Andrew M. Carek, Omer T. Inan, Ramakrishna Mukkamala, and Jin-Oh Hahn. Ballistocardiogram-Based Approach to Cuffless Blood Pressure Monitoring: Proof of Concept and Potential Challenges. 65(11):2384–2391.
  - [11] Sheida Knight, Jessica Lipoth, Mina Namvari, Carol Gu, Mojtaba Hedayati Ch, Shabbir Syed-Abdul, and Raymond J. Spiteri. The Accuracy of Wearable Photoplethysmography Sensors for Telehealth Monitoring: A Scoping Review.
  - [12] Yuriy Kurylyak, Francesco Lamonaca, and Domenico Grimaldi. A Neural Network-based method for continuous blood pressure estimation from a PPG signal. In *2013 IEEE International Instrumentation and Measurement Technology Conference (I2MTC)*, pages 280–283.
  - [13] G.J. Langewouters, A. Zwart, R. Busse, and K.H. Wesseling. Pressure-diameter relationships of segments of human finger arteries. 7(1):43–55.

- [14] Alexander A. Leung, Kara Nerenberg, Stella S. Daskalopoulou, Kerry McBrien, Kelly B. Zarnke, Kaberi Dasgupta, Lyne Cloutier, Mark Gelfer, Maxime Lamarre-Cliche, Alain Milot, Peter Bolli, Guy Tremblay, Donna McLean, Sheldon W. Tobe, Marcel Ruzicka, Kevin D. Burns, Michel Vallée, G. V. Ramesh Prasad, Marcel Lebel, Ross D. Feldman, Peter Selby, Andrew Pipe, Ernesto L. Schiffrin, Philip A. McFarlane, Paul Oh, Robert A. Hegele, Milan Khara, Thomas W. Wilson, S. Brian Penner, Ellen Burgess, Robert J. Herman, Simon L. Bacon, Simon W. Rabkin, Richard E. Gilbert, Tavis S. Campbell, Steven Grover, George Honos, Patrice Lindsay, Michael D. Hill, Shelagh B. Coutts, Gord Gubitz, Norman R. C. Campbell, Gordon W. Moe, Jonathan G. Howlett, Jean-Martin Boulanger, Ally Prebtani, Pierre Larochelle, Lawrence A. Leiter, Charlotte Jones, Richard I. Ogilvie, Vincent Woo, Janusz Kaczorowski, Luc Trudeau, Robert J. Petrella, Swapnil Hiremath, Denis Drouin, Kim L. Lavoie, Pavel Hamet, George Fodor, Jean C. Grégoire, Richard Lewanczuk, George K. Dresser, Mukul Sharma, Debra Reid, Scott A. Lear, Gregory Moullec, Milan Gupta, Laura A. Magee, Alexander G. Logan, Kevin C. Harris, Janis Dionne, Anne Fournier, Geneviève Benoit, Janusz Feber, Luc Poirier, Raj S. Padwal, Doreen M. Rabi, and CHEP Guidelines Task Force. Hypertension Canada’s 2016 Canadian Hypertension Education Program Guidelines for Blood Pressure Measurement, Diagnosis, Assessment of Risk, Prevention, and Treatment of Hypertension. 32(5):569–588.
- [15] Yongbo Liang, Mohamed Elgendi, Zhencheng Chen, and Rabab Ward. An optimal filter for short photoplethysmogram signals. 5(1):180076.
- [16] M. Liu, X. Zhan, J. Tu, B. Liu, and Z.H. Zhu. Integrated navigation for tethered nano-satellite system by modified input-delay neural networks and PROSAC. volume 2017-May, pages 202–210.
- [17] World Health Organization. *World Health Statistics 2023: Monitoring Health for the SDGs, Sustainable Development Goals*. World Health Organization.
- [18] L. Peter, N. Noury, and M. Cerny. A review of methods for non-invasive and continuous blood pressure monitoring: Pulse transit time method is promising? 35(5):271–282.
- [19] C.C.Y. Poon and Y.T. Zhang. Cuff-less and Noninvasive Measurements of Arterial Blood Pressure by Pulse Transit Time. In *2005 IEEE Engineering in Medicine and Biology 27th Annual Conference*, pages 5877–5880.

- [20] Ivan Reyes, Homer Nazeran, Mario Franco, and Emily Haltiwanger. Wireless photoplethysmographic device for heart rate variability signal acquisition and analysis. In *2012 Annual International Conference of the IEEE Engineering in Medicine and Biology Society*, pages 2092–2095.
- [21] Juan C. Ruiz-Rodríguez, Adolf Ruiz-Sanmartín, Vicent Ribas, and Jesús Caballero. Innovative continuous non-invasive cuffless blood pressure monitoring based on photoplethysmography technology. 39(9):1618–1625.
- [22] Manuja Sharma, Karinne Barbosa, Victor Ho, Devon Griggs, Tadesse Ghirmai, Sandeep K. Krishnan, Tzung K. Hsiai, Jung-Chih Chiao, and Hung Cao. Cuff-Less and Continuous Blood Pressure Monitoring: A Methodological Review. 5(2):21.
- [23] Peng Su, Xiao-Rong Ding, Yuan-Ting Zhang, Jing Liu, Fen Miao, and Ni Zhao. Long-term blood pressure prediction with deep recurrent neural networks. In *2018 IEEE EMBS International Conference on Biomedical & Health Informatics (BHI)*, pages 323–328.
- [24] Satomi Suzuki and Koji Oguri. Cuffless blood pressure estimation by error-correcting output coding method based on an aggregation of AdaBoost with a photoplethysmograph sensor. In *2009 Annual International Conference of the IEEE Engineering in Medicine and Biology Society*, pages 6765–6768.
- [25] K. Takazawa, N. Tanaka, M. Fujita, O. Matsuoka, T. Saiki, M. Aikawa, S. Tamura, and C. Ibukiyama. Assessment of vasoactive agents and vascular aging by the second derivative of photoplethysmogram waveform. 32(2):365–370.
- [26] X.F. Teng and Y.T. Zhang. Continuous and noninvasive estimation of arterial blood pressure using a photoplethysmographic approach. In *Proceedings of the 25th Annual International Conference of the IEEE Engineering in Medicine and Biology Society (IEEE Cat. No.03CH37439)*, volume 4, pages 3153–3156 Vol.4.
- [27] X. Xing and M. Sun. Optical blood pressure estimation with photoplethysmography and fft-based neural networks. 7(8):3007–3020.
- [28] Gilwon Yoon, Jong Youn Lee, Kye Jin Jeon, Kun-Kook Park, Hyung S. Yeo, Hyun Tae Hwang, Hong Sig Kim, and In-Duk Hwang. Multiple diagnosis based on photoplethysmography: Hematocrit, SpO<sub>2</sub>, pulse, and respiration. In *Optics*

*in Health Care and Biomedical Optics: Diagnostics and Treatment*, volume 4916, pages 185–188. SPIE.