

The need for data harmonization, multimodal integration and multi-objective loss functions to identify longitudinal changes in aging

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Abstract

Age-related cardiovascular changes, such as arterial stiffening, increased blood pressure, and decreased heart rate variability, contribute to a higher risk of cardiovascular diseases (CVDs), including hypertension, heart failure, atrial fibrillation and stroke. Detecting these changes early is essential for timely intervention and improved outcomes, but current monitoring methods often fall short in detecting subtle age-related changes before overt disease symptoms appear. Non-invasive tools like photoplethysmography (PPG), integrated into wearable devices, offer a promising approach for continuous cardiovascular monitoring. However, significant challenges remain in developing generalizable models that can handle diverse and noisy physiological data from various devices and populations. Addressing these limitations requires advanced, multimodal frameworks that harmonizes data from diverse sources and leverages state-of-the-art deep learning techniques to detect age-related cardiovascular changes.

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Original Manuscript

The need for data harmonization, multimodal integration and multi-objective loss functions to identify longitudinal changes in aging

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1 Introduction

Age-related cardiovascular changes, such as arterial stiffening, increased blood pressure, and decreased heart rate variability, contribute to a higher risk of cardiovascular diseases (CVDs), including hypertension, heart failure, atrial fibrillation and stroke. Detecting these changes early is essential for timely intervention and improved outcomes, but current monitoring methods often fall short in detecting subtle age-related changes before overt disease symptoms appear. Non-invasive tools like photoplethysmography (PPG), integrated into wearable devices, offer a promising approach for continuous cardiovascular monitoring. However, significant challenges remain in developing generalizable models that can handle diverse and noisy physiological data from various devices and populations. Addressing these limitations requires advanced, multimodal frameworks that harmonizes data from diverse sources and leverages state-of-the-art deep learning techniques to detect age-related cardiovascular changes.

2 The cardiovascular system changes with age

As people age, the cardiovascular system experiences progressive structural and functional changes that contribute to an increased risk of cardiovascular diseases (CVDs). Key cardiovascular changes include arterial stiffening, increased systolic blood pressure, decreasing heart rate variability (HRV), and elevated pulse wave velocity [1, 2]. These physiological changes increase the likelihood of adverse cardiovascular outcomes, including stroke, heart failure (HF), hypertension (HTN), and atrial fibrillation (AF) [3, 4]. Given the importance of early detection and intervention, there is a critical need for advanced tools that can detect and monitor these cardiovascular changes before they manifest as overt disease [5]. The challenge is that subtle, longitudinal cardiovascular changes associated with aging can be difficult to detect with infrequent measurements taken using standard monitoring tools like blood pressure cuffs or electrocardiograms (ECGs) in the clinic [6]. Continuous monitoring tools are therefore critically needed to identify at-risk individuals and provide a window for early therapeutic intervention.

Data Harmonization

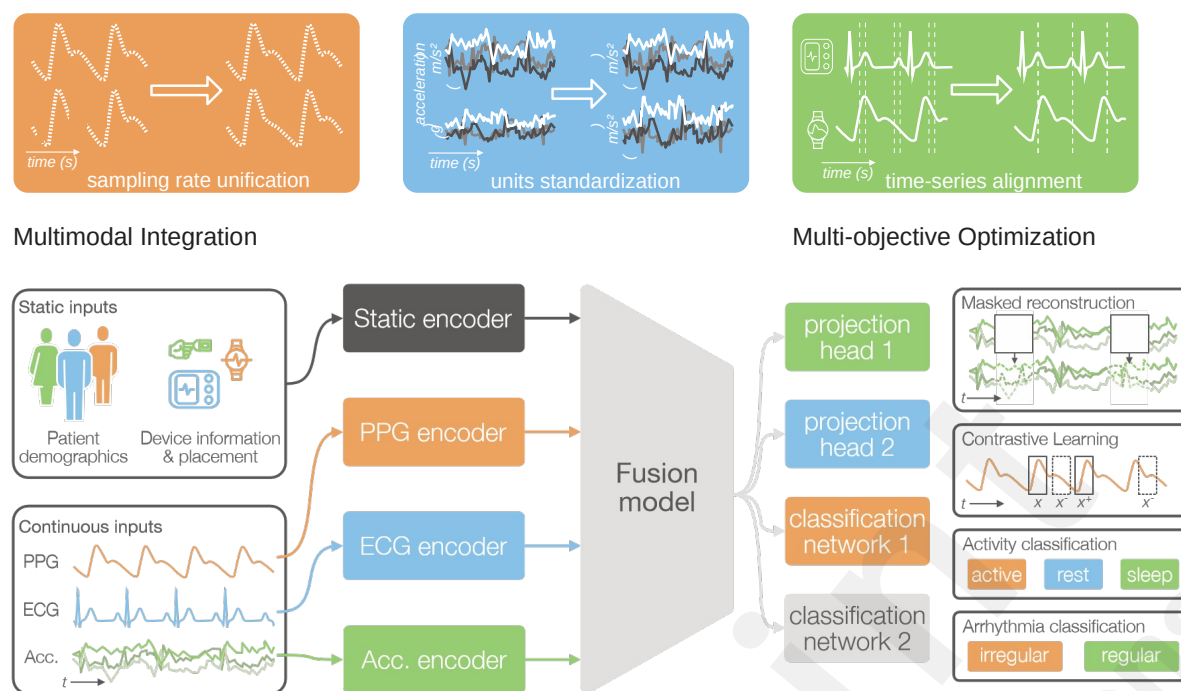


Figure 1: The roles of data harmonization, multimodal integration and multi-objective optimization in creating generalizable models.

3 PPG signals reflect changes in the cardiovascular system

Photoplethysmography (PPG) is a non-invasive optical technique that measures changes in blood volume in the microvascular bed, typically through light absorption at the skin surface [7, 8]. PPG has become widely adopted in wearable devices such as smartwatches and fitness trackers due to its simplicity and ease of integration into everyday settings [9]. Importantly, PPG signals can provide insights into key cardiovascular parameters, including heart rate, heart rate variability, and vascular health markers like pulse wave velocity (a surrogate marker for arterial stiffness) [10]. The ability of PPG to reflect real-time cardiovascular dynamics makes it a promising tool for monitoring age-related cardiovascular changes [11, 12]. For example, changes in PPG waveforms can indicate alterations in arterial compliance or peripheral vascular resistance, which are commonly observed in older adults as a result of arterial stiffening [13]. By capturing these subtle changes in vascular tone and heart rate dynamics, PPG offers the potential to detect early cardiovascular changes that might precede clinical symptoms.

4 Current approaches to interpreting PPG signals

To date, most PPG interpretation methods rely on traditional signal processing techniques. These include feature extraction methods that analyze various aspects of the PPG waveform (e.g., peak-to-peak intervals, amplitude) and time-domain or frequency-domain analysis to derive parameters like HRV [11]. While these approaches are effective for basic cardiovascular monitoring, they are less suited for detecting the complex, multi-faceted changes in cardiovascular function associated with aging. For instance, traditional methods often focus on short-term HRV or pulse characteristics, missing broader trends that could signal early cardiovascular deterioration. There is

increasing interest in using machine learning methods to enhance the robustness of PPG signal interpretation. For example, deep learning models have shown promise in detecting specific cardiovascular conditions, such as AF, from PPG signals with high accuracy [14]. However, these models require large amounts of clean, annotated data and are typically trained to detect specific disease states, limiting their ability to generalize to broader, age-related cardiovascular changes. Furthermore, the reliance on single, well-curated datasets poses a challenge in real-world applications, where PPG signals are often noisy and collected under suboptimal conditions [9]. Additionally, many existing models are application-specific and fail to generalize across different types of wearable devices [15]. These limitations highlight the need for new approaches that can generalize across different populations, devices, and environments while being robust to the noise and variability inherent in real-world data collection.

5 The need for large, diverse and harmonized datasets

Most existing PPG datasets are collected from young, healthy individuals in controlled conditions, limiting the generalizability [16]. To develop robust models capable of detecting cardiovascular changes in older adults, it is essential to utilize diverse datasets that reflect the variability found in the real-world populations. The integration of multi-modal data, such as PPG, ECG, and motion sensors, is vital to capturing the complex interactions between physiological systems. Harmonizing diverse datasets is necessary to ensure standardization and compatibility. Recent studies provide frameworks for addressing these challenges [17]. Harmonization of sleep and circadian data through standardization of sampling rates and alignment of time-series data has been shown to improve cross-study comparisons and reduce noise and variability [18]. The development of pipelines such as MASH have addressed issues of inconsistent data notations and varying sampling rates across multimodal datasets, facilitating the alignment of diverse datasets collected from different devices [19]. Furthermore, the importance of modular and flexible data preprocessing pipelines has been emphasized, demonstrating that scalable harmonization can be achieved through frameworks that enable seamless integration of new datasets [20]. These advancements in data harmonization techniques provide a foundation for the integration of diverse datasets in cardiovascular monitoring research.

6 Modern neural network architectures for multimodal data

Recent advances in deep learning offer significant potential for improving cardiovascular monitoring, particularly through attention-based and deep state space models [21, 22]. These models have shown considerable promise in handling multi-modal, sequential data such as health timelines, but have not yet incorporated PPG, ECG, or motion data. Modern architectures are well-suited to processing long sequences of data, allowing for the detection of temporal patterns and trends in cardiovascular signals that may indicate early deterioration [23]. Their ability to integrate multi-modal inputs enables them to learn from the complex interactions between different types of physiological signals, providing a comprehensive view of cardiovascular health. Furthermore, these architectures can process multiple high-frequency signals, such as PPG and ECG, simultaneously. These signals, though rich in cardiovascular information, are susceptible to noise, and traditional models often treat them independently, missing critical interactions [16]. Recent advances in self-supervised

learning offer the potential to design architectures that can integrate multiple high-frequency signals, enhancing the model's ability to distinguish between artifacts and true physiological changes [24]. By incorporating motion sensor data alongside PPG and ECG signals, the model can better interpret physiological states and detect subtle cardiovascular changes in older adults, even in noisy, real-world conditions [15].

7 The need for a multi-objective loss function

In machine learning, the loss function is a critical component that directs how models learn from data. For complex applications such as detecting cardiovascular changes using multi-modal, noisy data, conventional loss functions that focus purely on accuracy or classification often fall short [25, 26]. Multi-objective loss functions offer a powerful solution by enabling models to optimize multiple objectives simultaneously, such as imputing missing data, handling noise, and learning from diverse signals like PPG, ECG, and motion data [27]. By guiding the model to predict missing signals or adjust for signal quality variations, multi-objective loss functions will it possible to detect cardiovascular deterioration more accurately, even in challenging real-world scenarios. Advanced loss functions that balance tasks like imputing missing data and detecting subtle cardiovascular changes will significantly improve model performance across heterogeneous datasets, ultimately enhancing personalized care for older adults.

Recent advancements in multi-task learning have provided valuable frameworks for addressing complex, multi-objective optimization problems. The concept of framing multi-task learning as a multi-objective optimization problem, utilizing Pareto optimality, has been introduced as a method to balance competing objectives without compromising performance on individual tasks [27]. This approach is particularly relevant in cardiovascular monitoring, where multiple tasks such as autoregressive prediction of PPG/ECG signals, cross-modality predictions, and classification must be optimized simultaneously. Building on this foundation, frameworks for dynamically balancing tasks during training have been developed, enabling models to adapt and prioritize tasks based on their relative importance [27, 28]. This adaptive capacity is crucial when dealing with varied data types and the complex nature of cardiovascular changes. Furthermore, the significance of loss functions that capture interactions between tasks has been emphasized, offering strategies for refining multi-objective optimization approaches [29]. These developments in multi-task learning and multi-objective optimization provide a robust theoretical foundation for designing complex loss functions capable of handling the multifaceted challenges inherent in cardiovascular monitoring using multi-modal data.

8 Conclusion

The growing burden of age-related cardiovascular diseases highlights the urgent need for advanced tools that can detect and monitor subtle, longitudinal changes in cardiovascular function. While photoplethysmography (PPG) holds great promise for non-invasive, continuous monitoring, existing approaches fall short in their ability to generalize across diverse populations, devices, and realworld conditions. The future of cardiovascular health monitoring relies on a concerted effort to harmonize data across multiple sources and modalities, integrating diverse datasets that reflect the true variability of the human population.

To make meaningful progress in this field, researchers and industry stakeholders

must prioritize three key areas:

1. **Data Harmonization:** Without standardization and harmonization, the wealth of data collected from wearables, healthcare systems, and diverse populations remains underutilized. Collaborative efforts across disciplines are needed to establish common frameworks for integrating multimodal physiological data, such as PPG, ECG, and motion sensors. These frameworks must ensure compatibility across devices and environments, enabling more robust models that generalize to real-world scenarios.
2. **Multimodal Integration:** Cardiovascular changes do not occur in isolation. Integrating multiple physiological signals—PPG, ECG, and others—provides a more comprehensive picture of cardiovascular health. Advanced deep learning models, particularly those using modern architectures like attention-based and state-space models, are well-positioned to capture the complex interactions between these signals. However, future research must prioritize models that can learn across multiple modalities, handle noisy data, and address the unique challenges of older populations.
3. **Multi-Objective Loss Functions:** In real-world applications, the challenges of noisy data, missing signals, and high variability must be met with sophisticated loss function designs that balance multiple objectives. Multi-task learning and multi-objective optimization frameworks offer a promising path forward by enabling models to learn from diverse data types while prioritizing key health metrics. Implementing these techniques will pave the way for the early detection of cardiovascular changes, personalized care, and more effective interventions for aging populations.

This is a critical juncture in cardiovascular research and health monitoring. By advancing in these three areas, we can create the infrastructure needed for scalable, real-time monitoring solutions that empower healthcare providers to intervene earlier and more effectively. The integration of data harmonization, multimodal learning, and advanced loss functions has the potential to transform cardiovascular care, improving outcomes for millions of older adults worldwide.

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References

- [1] Alberto U. Ferrari, Alberto Radaelli, and Marco Centola. "Invited Review: Aging and the cardiovascular system". In: *Journal of Applied Physiology* 95.6 (Dec.

- 2003). Publisher: American Physiological Society, pp. 2591–2597. issn: 8750-7587. doi: 10.1152/japplphysiol.00601.2003.
- [2] Apostolos Karavidas, George Lazaros, Dimitris Tsiachris, and Vlassios Pyrgakis. “Aging and the cardiovascular system”. In: *Hellenic J Cardiol* 51.5 (2010), pp. 421–7.
- [3] Samuel L’evy. “Factors Predisposing to the Development of Atrial Fibrillation”. en. In: *Pacing and Clinical Electrophysiology* 20.10 (1997). eprint: <https://onlinelibrary.wiley.com/doi/pdf/10.1111/j.15408159.1997.tb06115.x>, pp. 2670–2674. issn: 1540-8159. doi: 10.1111/j.1540-8159.1997.tb06115.x.
- [4] Mikhail S. Dzeshka, Alena Shantsila, Eduard Shantsila, and Gregory Y.H. Lip. “Atrial Fibrillation and Hypertension”. In: *Hypertension* 70.5 (Nov. 2017). Publisher: American Heart Association, pp. 854–861. doi: 10.1161/HYPERTENSIONAHA.117.08934.
- [5] A. Moore, A. A. Mangoni, D. Lyons, and S. H. D. Jackson. “The cardiovascular system in the ageing patient”. en. In: *British Journal of Clinical Pharmacology* 56.3 (2003). eprint: <https://onlinelibrary.wiley.com/doi/pdf/10.1046/j.0306-5251.2003.01876.x>, pp. 254–260. issn: 1365-2125. doi: 10.1046/j.0306-5251.2003.01876.x.
- [6] Majd AlGhatrif, James B. Strait, Chris H. Morrell, Marco Canepa, Jeanette Wright, Palchamy Elango, Angelo Scuteri, Samer S. Najjar, Luigi Ferrucci, and Edward G. Lakatta. “Longitudinal Trajectories of Arterial Stiffness and the Role of Blood Pressure: The Baltimore Longitudinal Study of Aging”. en. In: *Hypertension* 62.5 (Nov. 2013), pp. 934–941. issn: 0194-911X, 1524-4563. doi: 10.1161/HYPERTENSIONAHA.113.01445.
- [7] Harikumar Rajaguru and Sunil Kumar Prabhakar. “A Comprehensive Review on Photoplethysmography and Its Application for Heart Rate Turbulence Clinical Diagnosis”. In: *Advanced Science Letters* 21.12 (Dec. 2015), pp. 3602–3604. doi: 10.1166/asl.2015.6541.
- [8] Hui Wen Loh, Shuting Xu, Oliver Faust, Chui Ping Ooi, Prabal Datta Barua, Subrata Chakraborty, Ru-San Tan, Filippo Molinari, and U Rajendra Acharya. “Application of photoplethysmography signals for healthcare systems: An in-depth review”. In: *Computer Methods and Programs in Biomedicine* 216 (Apr. 2022), p. 106677. issn: 0169-2607. doi: 10.1016/j.cmpb.2022.106677.
- [9] Denisse Castaneda, Aibhlin Esparza, Mohammad Ghamari, Cinna Soltanpur, and Homer Nazeran. “A review on wearable photoplethysmography sensors and their potential future applications in health care”. In: *International journal of biosensors & bioelectronics* 4.4 (2018), pp. 195–202. issn: 2573-2838. doi: 10.15406/ijbsbe.2018.04.00125.
- [10] Aymen A. Alian and Kirk H. Shelley. “Photoplethysmography”. In: *Best Practice & Research Clinical Anaesthesiology*. Hemodynamic Monitoring Devices 28.4 (Dec. 2014), pp. 395–406. issn: 1521-6896. doi: 10.1016/j.bpa.2014.08.006.
- [11] Mohamed Elgendi, Richard Fletcher, Yongbo Liang, Newton Howard, Nigel H. Lovell, Derek Abbott, Kenneth Lim, and Rabab Ward. “The use of photoplethysmography for assessing hypertension”. en. In: *npj Digital Medicine* 2.1 (June 2019). Publisher: Nature Publishing Group, pp. 1–11. issn: 2398-6352. doi: 10.1038/s41746-019-0136-7.
- [12] Gilene de Jesus Pereira, Rodrigo Miguel-dos-Santos, Valter Joviniano de Santana-Filho, Jos’e Augusto Soares Barreto-Filho, Cristiane Kelly Aquino dos Santos, and Mylena

Maria Salgueiro Santana. *Influence of antihypertensive pharmacological treatment on the acute cardiovascular responses to the resistance exercise in hypertensive middle-aged women*. 2020. doi: 10.6084/ m9.figshare.14290322.v1.

- [13] Dwaipayan Biswas, Neide Simões-Capela, Chris Van Hoof, and Nick Van Helleputte. “Heart Rate Estimation From Wrist-Worn Photoplethysmography: A Review”. In: *IEEE Sensors Journal* 19.16 (Aug. 2019). Conference Name: IEEE Sensors Journal, pp. 6560–6570. issn: 1558-1748. doi: 10.1109/JSEN.2019.2914166.
- [14] Tania Pereira, Nate Tran, Kais Gadhouri, Michele M. Pelter, Duc H. Do, Randall J. Lee, Rene Colorado, Karl Meisel, and Xiao Hu. “Photoplethysmography based atrial fibrillation detection: a review”. en. In: *npj Digital Medicine* 3.1 (Jan. 2020). Publisher: Nature Publishing Group, pp. 1–12. issn: 2398-6352. doi: 10.1038/s41746-019-0207-9.
- [15] Gideon Vos, Kelly Trinh, Zoltan Sarnyai, and Mostafa Rahimi Azghadi. “Generalizable machine learning for stress monitoring from wearable devices: A systematic literature review”. In: *International Journal of Medical Informatics* 173 (May 2023), p. 105026. issn: 1386-5056. doi: 10.1016/j.ijmedinf.2023.105026.
- [16] Karim Bayoumy, Mohammed Gaber, Abdallah Elshafeey, Omar Mhaimed, Elizabeth H. Dineen, Francoise A. Marvel, Seth S. Martin, Evan D. Muse, Mintu P. Turakhia, Khaldoun G. Tarakji, and Mohamed B. Elshazly. “Smart wearable devices in cardiovascular care: where we are and how to move forward”. en. In: *Nature Reviews Cardiology* 18.8 (Aug. 2021). Number: 8 Publisher: Nature Publishing Group, pp. 581–599. issn: 1759-5010. doi: 10.1038/s41569021-00522-7.
- [17] Supriyo Choudhury, Genko Oyama, and Hrishikesh Kumar. “Chapter 20 - Harmonization of data sets: basic principles and ethical aspects”. In: *Handbook of Digital Technologies in Movement Disorders*. Ed. by Roongroj Bhidayasiri and Walter Maetzler. Academic Press, Jan. 2024, pp. 315–328. isbn: 978-0-323-99494-1. doi: 10.1016/B978-0-323-99494-1.00007-1.
- [18] Diego R Mazzotti, Melissa A Haendel, Julie A McMurry, Connor J Smith, Daniel J Buysse, Till Roenneberg, Thomas Penzel, Shaun Purcell, Susan Redline, Ying Zhang, Kathleen R Merikangas, Joseph P Menetski, Janet Mullington, Eilis Boudreau, and on behalf of the Sleep Research Network Task Force. “Sleep and circadian informatics data harmonization: a workshop report from the Sleep Research Society and Sleep Research Network”. In: *Sleep* 45.6 (June 2022), zsac002. issn: 0161-8105. doi: 10.1093/sleep/zsac002.
- [19] Erin E. Dooley, J. F. Winkles, Alicia Colvin, Christopher E. Kline, Sylvia E. Badon, Keith M. Diaz, Carrie A. Karvonen-Gutierrez, Howard M. Kravitz, Barbara Sternfeld, S. Justin Thomas, Martica H. Hall, and Kelley Pettee Gabriel. “Method for Activity Sleep Harmonization (MASH): a novel method for harmonizing data from two wearable devices to estimate 24-h sleep–wake cycles”. en. In: *Journal of Activity, Sedentary and Sleep Behaviors* 2.1 (Apr. 2023), p. 8. issn: 2731-4391. doi: 10.1186/s44167-023-00017-5.
- [20] Sylvia Cho, Chunhua Weng, Michael G. Kahn, and Karthik Natarajan. “Identifying Data Quality Dimensions for Person-Generated Wearable Device Data: Multi-Method Study”. EN. In: *JMIR mHealth and uHealth* 9.12 (Dec. 2021). Company: JMIR mHealth and uHealth Distributor: JMIR mHealth and uHealth Institution: JMIR mHealth and uHealth Label:

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- [21] Adibvafa Fallahpour, Mahshid Alinoori, Arash Afkanpour, and Amrit Krishnan. *EHRMamba: Towards Generalizable and Scalable Foundation Models for Electronic Health Records*. arXiv:2405.14567 [cs] version: 1. May 2024. doi: 10.48550/arXiv.2405.14567.
- [22] Pawel Renc, Yugang Jia, Anthony E. Samir, Jaroslaw Was, Quanzheng Li, David W. Bates, and Arkadiusz Sitek. "Zero shot health trajectory prediction using transformer". en. In: *npj Digital Medicine* 7.1 (Sept. 2024). Publisher: Nature Publishing Group, pp. 1–10. issn: 23986352. doi: 10.1038/s41746-024-01235-0.
- [23] Paola Daniore, Vasileios Nittas, Christina Haag, Juergen Bernard, Roman Gonzenbach, and Viktor von Wyl. "From wearable sensor data to digital biomarker development: ten lessons learned and a framework proposal". en. In: *npj Digital Medicine* 7.1 (June 2024). Publisher: Nature Publishing Group, pp. 1–8. issn: 2398-6352. doi: 10.1038/s41746-024-01151-3.
- [24] Kexin Zhang, Qingsong Wen, Chaoli Zhang, Rongyao Cai, Ming Jin, Yong Liu, James Zhang, Yuxuan Liang, Guansong Pang, Dongjin Song, and Shirui Pan. *Self-Supervised Learning for Time Series Analysis: Taxonomy, Progress, and Prospects*. en. arXiv:2306.10125 [cs, eess, stat]. July 2023.
- [25] C. El-Hajj and P. A. Kyriacou. "A review of machine learning techniques in photoplethysmography for the non-invasive cuff-less measurement of blood pressure". In: *Biomedical Signal Processing and Control* 58 (Apr. 2020), p. 101870. issn: 1746-8094. doi: 10.1016/j.bspc.2020.101870.
- [26] Guangkun Nie, Jiabao Zhu, Gongzheng Tang, Deyun Zhang, Shijia Geng, Qinghao Zhao, and Shenda Hong. *A Review of Deep Learning Methods for Photoplethysmography Data*. arXiv:2401.12783 [cs, eess]. Jan. 2024. doi: 10.48550/arXiv.2401.12783.
- [27] Ozan Sener and Vladlen Koltun. "Multi-Task Learning as Multi-Objective Optimization". In: *Advances in Neural Information Processing Systems*. Vol. 31. Curran Associates, Inc., 2018.
- [28] Jian-Yu Li, Zhi-Hui Zhan, Yun Li, and Jun Zhang. "Multiple Tasks for Multiple Objectives: A New Multiobjective Optimization Method via Multitask Optimization". In: *IEEE Transactions on Evolutionary Computation* (2023). Conference Name: IEEE Transactions on Evolutionary Computation, pp. 1–1. issn: 1941-0026. doi: 10.1109/TEVC.2023.3294307.
- [29] Shubhkirti Sharma and Vijay Kumar. "A Comprehensive Review on Multi-objective Optimization Techniques: Past, Present and Future". en. In: *Archives of Computational Methods in Engineering* 29.7 (Nov. 2022), pp. 5605–5633. issn: 1134-3060, 1886-1784. doi: 10.1007/s11831-022-09778-9.

Supplementary Files

Figures

The roles of data harmonization, multimodal integration and multi-objective optimization in creating generalizable models.

