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(Methodological Review)

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The results are the same as a number of the data. The results are the same as a number of the data. The results are the same as a number of the data. The results are the same$ *Abstract***—The human ear has emerged as a bidirectional gateway to the brain's and body's signals. Recent advances in around-the-ear and in-ear sensors have enabled the assessment of biomarkers and physiomarkers derived from brain and cardiac activity using ear- electroencephalography (ear-EEG), photoplethysmography (ear-PPG), and chemical sensing of analytes from the ear, with ear-EEG having been taken beyond-the-lab to outer space. Parallel advances in non-invasive and minimally invasive brain stimulation techniques have leveraged the ear's access to two cranial nerves to modulate brain and body activity. The vestibulocochlear nerve stimulates the auditory cortex and limbic system with sound, while the au- ricular branch of the vagus nerve indirectly but significantly couples to the autonomic nervous system and cardiac output. Acoustic and current mode stimuli delivered using discreet and unobtrusive earables are an active area of research, aiming to make biofeedback and bioelectronic medicine deliverable outside of the clinic, with remote and continuous monitoring of therapeutic responsivity**

Manuscript received 4 September 2024; revised 10 November 2024; accepted 17 November 2024. Date of publication; date of current version. This work was supported by the UCSD Center for Wearable Sensors. *(Yuchen Xu, Abhinav Uppal and Min Suk Lee contributed equally to this work.) (Corresponding author: Gert Cauwenberghs.)*

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Digital Object Identifier 10.1109/RBME.2024.3508713

and long-term adaptation. Leveraging recent advances in 28 **ear-EEG, transcutaneous auricular vagus nerve stimulation** 29 **(taVNS), and unobtrusive acoustic stimulation, we review** 30 **accumulating evidence that combines their potential into** 31 **an integrated earable platform for closed-loop multimodal** 32 **sensing and neuromodulation, towards personalized and** 33 **holistic therapies that are near, in- and around-the-ear.** 34

*Index Terms***—Earables, ear-EEG, ear-PPG, biofeedback,** 35 **auditory neurofeedback, transcutaneous auricular vagus** 36 **nerve stimulation, closed-loop neuromodulation.** 37

I. INTRODUCTION 38

E ARABLES or hearables [1], [2] are devices that can be 39
worn inside or around the ears, and that provide additional 40
functionality beyond audio input and output [3]. Earables have functionality beyond audio input and output [3]. Earables have ⁴¹ emerged as a transformative innovation in the domain of wear- ⁴² able health monitoring $[4]$, $[5]$, and as a neuromodulation plat- 43 form for applying non-invasive stimulation to remedy a target ⁴⁴ pathology, such as using bimodal therapy combining auditory ⁴⁵ and electrical stimulation to the ear, with the goal of inducing 46 plasticity in the auditory cortex of tinnitus patients $[6]$. 47

Because of its anatomy and physiology, the ear is uniquely 48 positioned for multimodal sensing. It provides access to sounds ⁴⁹ and vibrations in the ear, to a rich network of vasculature and ⁵⁰ innervation [5], [7], [8], [9], to the eyes [10], [11], [\[12\],](#page-15-0) [\[13\],](#page-15-0) ⁵¹ [14], to the muscles of the jaw $[14]$, and to the brain $[15]$, 52 [16], [17], [18], especially the temporal cortex $[19]$, $[20]$. This 53 opens the door to acoustic, optical, electrophysiological (ExG), ⁵⁴ and electrochemical sensing. The semi-flexible cartilage of the ⁵⁵ auricle (outer ear) provides a convenient structure for comfort- ⁵⁶ ably supporting in-ear and around-the-ear devices. Ubiquitious 57 examples such as wireless earbuds and hearing aids demonstrate 58 that earables are stable and suitable for extended wearability. ⁵⁹ In particular, ear electroencephalography (ear-EEG) technology 60 has even been taken into orbit for sleep monitoring on the Huginn 61 space mission $[21]$. 62

With over a decade since the first report of the ear-EEG 63 sensing concept $[22]$, the ear electrography (ear-ExG) field has 64

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is contributed by the method interaction of the method in the method interaction of the method interaction of the method interaction of the method interaction of the method interaction and the method interaction and the accumulated over 250 publications as of August 2024 (as seen by searching the Web of Science database for "ear-eeg OR ear-ppg OR ear-ecg OR ear-eog OR around-the-ear EEG OR behind-the-ear EEG"). As a sampling of the rich and growing ear-ExG literature, studies have reported innovations in sensor designs [13], [17], [23], [24], [25], [26], [27], [28], [29], [30], [31], [32], [33], [34], and custom integrated circuits and data acquisition systems optimized for ear-EEG [35], [36], [37], [38], [39]. Characterization and validation studies have per- formed simultaneous ear-EEG and scalp-EEG recordings [16], [19], [40], or used phantom models [41] and computational forward models to characterize the signal propagation from cardiac or cortical sources to the ear [19], [20], [42], [43], [44]. Recent reports of ear-EEG applications have included sleep stag- ing [45], [46], [47], [48], [49], [50], epilepsy monitoring [51], [52], [53], brain-computer interfaces using speech imagery [54] or steady-state visual evoked potentials (SSVEP) [55], [56], [57], [58], audiometric assessment [59], and auditory at- tention decoding [60], [61] towards neuro-steered hearing aids [62].

 Recent reviews are available that summarize the sensing capabilities of earables: Röddiger et al. [3] organized earables research by fundamental phenomenon that can be sensed from the ears, spanning physiological and health-related sensing, acivity-monitoring, human-computer interaction, and biometric applications. Masè et al. [5] reviewed in-ear hearables mea- suring body temperature, pulse rate, and blood oxygen satu- ration. Ne et al. [63] extended this criterion to hearables ac- quiring electrophsyiological signals. For the subset of earables research focusing on ear-EEG, Kaongoen et al. [64] reviewed ear-EEG studies including applications and analysis methods. Juez et al. [65] further narrowed their focus to in-ear EEG studies (that is, excluding around-the-ear devices), tabulating biomarkers validated against scalp-EEG, along with in-ear EEG applications and computational modeling approaches.

 In addition to robust sensing, the ear provides opportunities to deliver stimuli for modulating brain and body activity. Two stimulation modalities well-suited to the ear are acoustic (sound) or electric (current) given its access to multiple cranial nerves, leading to downstream modulation of the brainstem and higher areas, often resulting in measurable biomarkers to gauge the effectiveness of the therapy or responsiveness of subjects. Given this duality of stimulating sensory nerves and sensing down- stream effects from the brain and body, Ruhnau and Zaehle wrote a perspective in 2021 [66] suggesting that ear-EEG could be combined with transcutaneous auricular vagus nerve stimulation (taVNS) in a wearable, closed-loop neuromodulation device targeting alpha activity as a biomarker of attention. Although the design or validation of such a device has not been reported in the literature thus far, and the modulation of alpha activity has had mixed reports given the evolving mechanistic understanding of the field [\[67\],](#page-16-0) [\[68\].](#page-16-0) Our goal, as highlighted in Fig. 1, is to position earables as a more comprehensive approach, with multimodal sensing of brain and body activity, integrated with bimodal stimulation capability leveraging the ear's access to both the aurciular branch of the vagus nerve (ABVN) and the vestibocochlear (auditory) nerve.

Fig. 1. Overview of a biofeedback earable system. Left: the brain and body form a closed-loop control system using electrical and chemical messaging through neuronal and vascular networks. Right: components of the earable, consisting of sensing and stimulation (Stim) systems, also form a closed-loop control system for adapting the stimulation given sensed changes in the user's state, thus providing biofeedback to the user.

The rest of this paper is organized as follows: Section II 122 enumerates the anatomical and physiological properties of the ¹²³ ear to highlight its unique access to a plethora of physiological 124 and neural signals. Sections III and IV provide the necessary 125 background for sensing and stimulation principles, respectively, ¹²⁶ as applicable for various modalities in the ear. Section \overline{V} provides 127 an overview of considerations for system-level integration in ¹²⁸ earables. Section VI develops key steps towards closing the ¹²⁹ loop between earable sensing and stimulation, followed by our 130 conclusions in Section VII. 131

II. EAR AS A BIDIRECTIONAL GATEWAY 132

The human ear's unique anatomical and physiological proper- ¹³³ ties make it an ideal site for various sensing modalities. Beyond ¹³⁴ its primary role in hearing, the ear's structure, location, and ¹³⁵ vascularization offer significant advantages for physiological, ¹³⁶ chemical, and brain activity monitoring. This section briefly ¹³⁷ highlights how these characteristics enable multiple sensing 138 opportunities. ¹³⁹

Fig. 2 shows an input-output (I/O) map of the ear, with a rich ¹⁴⁰ diversity of inputs that can be provided to the ear as stimuli, ¹⁴¹ and outputs that can be sensed from the ear. The following ¹⁴² sub-sections highlight some key enabling features that uniquely 143 position the ear for both sensing from and stimulating the brain ¹⁴⁴ and body. The same state of the state of

A. Biosignals ¹⁴⁶

1) Proximity to the Brain: The ear's closeness to the brain ¹⁴⁷ makes it an optimal site for monitoring neural activity through 148 biopotential electrodes. This proximity minimizes signal degra- ¹⁴⁹ dation and allows for the detection of brain waves with higher 150 fidelity compared to peripherally worn sensors (wrist-watches, ¹⁵¹ finger tips, chest straps, etc). 152

2) Stable Blood Flow: The ear's robust vascularization and ¹⁵³ lower susceptibility to peripheral vasoconstriction, compared to 154 the limbs, is advantageous for photoplethysmography (PPG) ¹⁵⁵

Fig. 2. The human ear as a bidirectional gateway to brain and body signals. Left: examples of sensory stimulation that can be delivered through the ear include acoustic, electric, and vibrotactile stimuli. Center: the ear provides access to innervation for delivering stimuli, and its proximity to vasculature and multiple sources of biosignals enable sensing. Right: examples of brain and physiological signals, with colors corresponding to sources of origin. EEG: electroencephalogram, EOG: electrooculogram, EDA: electrodermal activity, Lac: lactate, Na+ : sodium, PPG: photoplethysmogram, EMG: electromyogram, and ECG: electrocardiogram.

¹⁵⁶ sensors measuring heart rate and oxygen saturation. This sta-¹⁵⁷ bility ensures consistent readings even under varying environ-¹⁵⁸ mental conditions.

159 *B. Stability*

 1) Minimal Motion Artifacts: Compared to other body parts, the ear remains relatively stable due to less muscle move- ment and natural damping of vibrations due to the structure of the skull. Additionally, the placement of the sensors in the ear provides better protection from external environmental factors. However, the ears are still subject to certain movements such as chewing or talking. In order to mitigate these artifacts for mobile brain imaging (MoBI), careful sensor design [69], [70], [71], [72], [73] and sensor placement [14], [74], [75] are needed to increase measurement accuracy and reduce post-processing requirements for artifact removal.

 2) Mechanical Anchoring Point: The ear provides a natural and secure anchoring point for wearable devices. The pinna's grooves allow for stable attachment of sensors and devices with- out the need for additional securing mechanisms. This mechan- ical anchoring ensures that the devices remain in place during various activities, enhancing the reliability of the collected data.

¹⁷⁷ *C. User Adoption*

 1) Pervasiveness and Social Acceptability of Ear- phones: The widespread use of earphones and their social acceptance in daily life make the ear a familiar and non-intrusive location for future earables that can maintain the earphone form-factor. Users are accustomed to wearing devices in their ears, which improves compliance with experimental protocols and retention over longer studies.

2) Comfortable and Discreet Placement: Earables are ¹⁸⁵ generally more comfortable for long-term wear, compared to ¹⁸⁶ other body-worn sensors like headset, chest straps or wristbands, ¹⁸⁷ which requires minimal adjustment and are less intrusive to daily 188 activities. Their discreet placement within/near the ear canal or ¹⁸⁹ around-the-ear makes them less noticeable to others, alleviating ¹⁹⁰ any social awkwardness often associated with conventional EEG 191 systems. 192

D. Innervation 193

The ear's access to multiple sensory nerves including the ¹⁹⁴ vestibulocochlear nerve carrying acoustic, motion, and posi- ¹⁹⁵ tioning information, and the auricular branch of the vagus ¹⁹⁶ nerve (ABVN) carrying somatosensory information allow for ¹⁹⁷ acoustic, electric, and bimodal stimulation of the brainstem and ¹⁹⁸ higher brain areas, that can induce electrical, chemical (through 199 neurotransmitters), and physiological changes (by modulating ²⁰⁰ the cardiac system). ²⁰¹

III. MULTIMODAL EARABLE SENSING ²⁰²

As mentioned in Section [II,](#page-1-0) the ears offer unique advantages 203 over other body parts due to the ear's distinct anatomical and ²⁰⁴ physiological characteristics. This section focuses on sensing ²⁰⁵ technologies for earables, comparing ear-based sensing with ²⁰⁶ sensing from other common body parts such as the scalp, arm, ²⁰⁷ chest, wrist, fingers, and legs. ²⁰⁸

A. Electrophysiological Sensing 209

Neurophysiological sensing systems are essential for moni- ²¹⁰ toring and understanding the electrical activities of the nervous ²¹¹

 system. These systems utilize various modalities to capture brain activity, muscle activity, and eye movements, provid- ing valuable insights into cognitive functions, motor control, and sensory processing. The primary sensors used in wearable neurophysiological sensing systems are biopotential elec- trodes [77], which detect electrical potentials generated by neural and muscular activity. These electrodes are commonly made from materials like silver/silver chloride (Ag/AgCl), and are designed to ensure a stable and reliable interface between the skin and the sensor.

an anaromay for each interaction and the same state in the Electrography (ExG) encompasses a range of techniques, including electroencephalography (EEG), electromyography (EMG), electrocardiography (ECG), and electrooculography (EOG), which measure the electrical activity of the brain, mus- cles, heart, and eyes, respectively. In addition to these ExG signals, electrodermal activity (EDA) can also be measured using the same electrophysiological measurement setup. EDA characterizes the skin's conductance response [78], which varies with sweat gland activity and is commonly associated with physiological arousal and stress levels. These techniques rely on biopotential electrodes that are placed on the skin's surface to detect small biopotentials generated by neural or muscular activity. The electrodes capture these signals, which are then am- plified, filtered, and digitally recorded for analysis. A differential architecture is often used to minimize noise and interference by comparing signals from paired electrodes, forming a bipolar channel for more accurate measurement. The feasibility of using ear biopotential sensors to measure ExG has been validated by previous research with simultaneous recording of comparison data from reference locations such as the scalp (EEG), chest (ECG), and finger (PPG) [5], [63].

 Each type of ExG signal has distinct characteristics. EEG 244 signals, typically ranging from 20 to 150 μ V with a bandwidth of 0.5–60 Hz, reflect the brain's electrical activity and vary both temporally and spatially across the scalp [79]. EMG signals, which are generally larger than EEG signals, capture the elec- trical activity of muscles during contraction, with amplitudes ranging from a few microvolts to millivolts, and bandwidths typically between 10 Hz and 500 Hz [38]. EOG signals, used to measure eye movements, fall within the range of 0.1 to 252×5 mV with a bandwidth of 0–35 Hz, reflecting the potential differences generated by eye movements. EOG signals can be further categorized based on their origin: eye blinks and eye movements. Eye blinks produce transient, high-amplitude sig- nals that are typically short in duration, while eye movements generate more sustained signals with lower amplitudes [\[80\].](#page-17-0) Together, these ExG modalities provide a comprehensive ap- proach to monitoring and analyzing various physiological processes.

 1) Biopotential Sensors: Electrodes for electrophysiology are conductive materials that enable electrical conduction be- tween the subject and the recording electronics. However, the choice of electrode affects the design, durability, maintenance, biocompatibility, signal quality, comfort, longevity, usability, and other features. Such considerations are especially impor- tant when devising miniaturized wearables such as ear-EEG devices [\[53\],](#page-16-0) [\[78\].](#page-16-0)

Biopotential electrodes including widely used Ag/AgCl elec- ²⁶⁹ trodes capture EEG signals through the electrochemical inter- ²⁷⁰ face between the electrode surface and the skin. When neurons 271 in the brain fire, they produce electrical signals that propagate ²⁷² through the brain and skull, reaching the surface of the scalp ²⁷³ and the ear. Biopotential electrodes convert these ionic currents ²⁷⁴ in the body to electronic currents that can be measured. The ²⁷⁵ Ag/AgCl material is particularly effective due to its low and ²⁷⁶ stable impedance, and high signal fidelity, making it suitable ²⁷⁷ for picking up the relatively weak EEG signals. The electrodes ²⁷⁸ act as transducers, capturing the voltage fluctuations caused by ²⁷⁹ brain activity, which can then be amplified and recorded by the ²⁸⁰ EEG system. ²⁸¹

Currently, there are three major interface methods: gel, dry, ²⁸² and non-contact electrodes. Gel-contact utilizes conductive gel ²⁸³ which ensures stable physical contact and lower impedance. 284 Previous work has demonstrated stable and low electrode-skin ²⁸⁵ impedance values maintained for several hours using cEEGrids, ²⁸⁶ where adhesive tape seals the electrode-skin interface, minimiz- 287 ing air exposure and preventing the gel from drying out, thus ²⁸⁸ allowing prolonged recording sessions [81]. Such configurations 289 are particularly suitable in clinical or research settings where ²⁹⁰ stable, high-quality signals are prioritized. However, these sys- ²⁹¹ tems often require extended maintenance, cleaning, and careful ²⁹² application to achieve optimal results. ²⁹³

For user comfort and ease of long-term use, especially in wear- ²⁹⁴ able applications, a system that operates without adhesives and ²⁹⁵ gel would be more ideal, minimizing discomfort and simplifying ²⁹⁶ usability. Dry-contact electrodes do not require conductive gel, ²⁹⁷ but still have conductive material directly in contact with the ²⁹⁸ skin. Such electrodes are less obtrusive and more convenient ²⁹⁹ for long-term recording. However, they typically have higher ³⁰⁰ impedance than gel-based electrodes, and requiremechanical ³⁰¹ force or adhesives to fixate the electrodes on the skin to ensure 302 good contact. This can often lead to discomfort. ³⁰³

Non-contact electrodes utilize capacitive coupling between ³⁰⁴ conductive electrode material and the skin to detect electrical ³⁰⁵ signals without physically touching the body [82], [\[83\].](#page-17-0) Simi- ³⁰⁶ larly to dry-contact, non-contact enables ease-of-use, long-term 307 monitoring, and reusability. Additionally, it can be considered ³⁰⁸ more hygienic, which reduces skin irritation or infections due to 309 the conductive material. However, non-contact has several cons: ³¹⁰ it typically has higher impedance than dry-contact, which leads 311 to lower signal quality; it requires more complex electronics to ³¹² amplify the signal; it is more prone to motion artifacts as the gap 313 between the skin and the electrode may change due to movement, ³¹⁴ affecting the capacitance that the non-contact electrodes rely on. ³¹⁵

Although there are these three contact options, in-ear EEG 316 literature predominantly uses dry-contact. One of the major ³¹⁷ benefits of in-ear EEG is the eventual wearable applications for ³¹⁸ consumer use. Wet-contact electrode characteristics are not ben- ³¹⁹ eficial for ease-of-use and long-term recording. For non-contact, ³²⁰ the ear devices typically have limited space, making it difficult to 321 incorporate amplifiers needed to boost the signal. Additionally, ³²² wearables require mobility. Therefore, non-contact electrodes ³²³ which are more susceptible to motion artifacts will make it ³²⁴ less favorable for wearable applications [\[84\].](#page-17-0) Although dry ³²⁵

Fig. 3. An overview of earables for different sensing modalities. For all subfigures from bottom to top: physiological sources, illustrative devices, and devices as worn by users. Bottom row from left to right: (a) half-cell model of the skin-electrode interface for electrophysiological signals, (b) optical interface for pulse plethysmography consisting of a light-emitting diode (LED) and a photo detector (PD), (c) sweat glands generating chemical analytes, and (d) mechano-acoustic sources visualized as a pulsing artery (other possible sources of ear canal motion and vibrations including sound not shown). Sources of device images from left to right: (a) Kappel et al. [26], ©2017 IEEE, (b) Budidha and Kyriacou [\[76\],](#page-16-0) (c) Xu et al. [13], (d) Goverdovsky et al. [1]. Device images (b)–(d) were modified to remove annotations, and are under the CC-BY 4.0 International License: http://creativecommons.org/licenses/by/4.0/.

 contact requires mechanical pressure or adhesives to ensure good contact, the geometric enclosures of the ear enable mechanical fitting for stable fixture. Additionally, many studies and methods have also been performed to mechanically stably fit objects to ³³⁰ the ear.

 A dry-contact electrode model is illustrated at the bottom of the column Fig. 3(a). This model represents the skin-electrode interface as a combination of resistive and capacitive com- ponents that together form the overall impedance of the sys- tem [84]. The skin's resistive properties are represented by a 336 resistance R_e (conductance $G_e = 1/R_e$). This resistance is a function of the electrode's contact area with the skin and the inherent resistivity of the skin's outer layer (stratum corneum). The capacitive component C*^e* arises from the dielectric prop- erties of the skin and the insulating layer of the electrode. This capacitance is influenced by factors such as the dielectric constant of the skin, the thickness of the stratum corneum, and the distance between the electrode and the underlying conductive tissues. The capacitive coupling allows the electrode to detect biopotential signals even in the presence of a non-conductive layer, but it also introduces a frequency-dependent impedance. The total impedance at the skin-electrode interface is modeled as a parallel RC circuit:

$$
Z_e = 1/(G_e + j\omega C_e). \tag{1}
$$

 The impedance at the skin-electrode interface directly influences the noise levels in the recorded signals. The importance of a low impedance of the skin-electrode interface is twofold; firstly, the impedance generates thermal noise as described by the Johnson- Nyquist equation. Secondly, the current noise of the amplifier is converted to voltage noise through the impedance [\[85\].](#page-17-0) Previous research show that the impedance of ear biopotential electrodes ranges from 1.2 MΩ at low frequencies to lower than 100 kΩ 356 at high frequencies for dry electrodes, and from $34 \text{ k}\Omega$ at low 357 frequencies to 5.1 k Ω at high frequencies for wet electrodes, 358 characterized across a frequency range of 0.1 Hz to 2 kHz [\[85\].](#page-17-0) ³⁵⁹ Another challenge in the ear is the variation of impedance at ³⁶⁰ the ear electrode-skin interface due to environmental factors ³⁶¹ like cerumen presence and electrodermal activity [\[78\],](#page-16-0) which 362 requires careful consideration of biopotential sensor designs. ³⁶³

A key factor to affect the impedance of biopotential elec- ³⁶⁴ trodes is the material. Key features for the electrode material ³⁶⁵ should be low impedance and biocompatible. Low impedance 366 will enable better signal quality and biocompatible will pre- 367 vent toxic exposure to the user after prolonged skin contact ³⁶⁸ and have hypoallergenic properties to minimize the risk of ³⁶⁹ skin reactions. The types of materials used in literature for ³⁷⁰ in-ear EEG are but not limited to conductive polymers [\[53\],](#page-16-0) ³⁷¹ [86], gold [87], CNT/PDMS [88], IrO2 [27], [89], composite ³⁷² silicone, and predominantly, silver [13], [25], [90], [\[91\],](#page-17-0) [\[92\],](#page-17-0) ³⁷³ [93], [94], [95], [96] and Ag/AgCl [22], [29], [38], [\[97\],](#page-17-0) [\[98\].](#page-17-0) ³⁷⁴ Additional features such as material flexibility, design/shape, ³⁷⁵ durability and maintenance are important considerations that ³⁷⁶ vary among sensors. The fabrication techniques of these sensors 377 also widely vary yet are integral to optimize and balance these ³⁷⁸ features. Examples found in literature for in-ear EEG sensors ³⁷⁹ are electroplating [\[25\],](#page-15-0) coating [\[13\],](#page-15-0) [\[38\],](#page-16-0) [\[53\],](#page-16-0) [\[91\],](#page-17-0) [\[92\],](#page-17-0) [\[99\],](#page-17-0) ³⁸⁰ solid metal working [\[24\],](#page-15-0) [\[27\],](#page-15-0) [\[89\],](#page-17-0) [\[95\],](#page-17-0) [\[97\],](#page-17-0) [\[98\],](#page-17-0) conductive ³⁸¹ threading [\[25\],](#page-15-0) [\[94\],](#page-17-0) [\[100\],](#page-17-0) [\[101\],](#page-17-0) and molding [\[88\].](#page-17-0) Neverthe- ³⁸² less, the choice of fabrication techniques should accommodate ³⁸³ the unique anatomical features of the ear canal while ensuring ³⁸⁴ high signal quality. Apart for the impedance, it is also crucial that 385 the fabrication of in-ear EEG devices ensures no structures in- ³⁸⁶ cluding edges that could potentially damage the ear canal. These 387

 geometrical constraints make designing low-contact impedance electrodes more challenging, as the need to ensure a safe fit can limit the surface area and optimal positioning required for maintaining stable, low impedance contact. Researchers have proposed different electrode designs that can adapt to the anatomy of different subjects by adding degrees of adaptability through mechanical designs [13], [39].

 The contact impedance between the electrode and skin is typically measured using the electrical impedance spectroscopy (EIS) method [79], [88], [102]. This characterization involves using three-electrode or four-electrode measurement configu- rations, where electrodes with similar contact areas are placed at specific distances (e.g., 1cm apart) on a skin surface, such as the forearm, to simulate conditions similar to their intended application site or the phosphate-buffered saline (PBS) solution as a simulated environment. The impedance is measured across a range of frequencies, typically from 1 Hz to 1000 Hz, and the contact impedance is derived by measuring the current resulted from the applied voltage.

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maximum symmetric matrix and the state of the For ear-EEG measurements, three main types of electrodes are typically used: measuring electrodes, reference (REF) elec- trodes, and ground (GND) electrodes. Measuring electrodes are placed in [27] or around [81], [103] the ear to detect brain activity. The reference electrode provides a baseline for the measurements, ensuring that the signals from the measuring electrodes are recorded relative to a consistent point. The refer- encing configuration can be categorized as contralateral when the reference electrode is located at the opposite side of the sagittal plane from the measuring electrode or ipsilateral when 417 it is placed within the same ear or surrounding area [104]. The ground electrode stabilizes the electrical environment by providing a common return path for the electrical current and reducing noise from external sources. For ear-EEG recordings REF and GND are usually located at the concha [31], [39], 422 [104] or mastoid [99], [105]. For ear-ECG, due to the relatively farther distance to the source of signal, REF and GND are usually located non-cephalic to capture ECG signals of good 425 quality $[106]$. Ear-ECG measured completely from the ear has also been explored. Single-ear ECG is feasible but challenging for cardiac rhythm monitoring. The limitations particularly are lower signal amplitude and higher susceptibility to noise com- pared to a cross-ear ECG setup, due to the smaller potential difference and closer proximity of electrodes, which reduce signal quality and reliability [42], [43].

 2) ExG Signal Characteristics: Ear-ExG sensing employs similar sensing mechanisms to those ExG sensing from other parts of the body. However, ear-ExG mainly differs by its limited size and placement options, leading to differences in signal characteristics. To quantify these characteristics for ear-ExG sensing, previous research has built forward models to simulate the mapping of brain sources and compares the difference of electrical potential distributions between the scalp and ear [\[19\],](#page-15-0) [\[107\].](#page-17-0) Forward models specifically refer to the transfer function from sources in the brain volume to biopotential electrodes. Here we describe using a simplified brain signal dipole model to illustrate such differences with the most widely reported ear-ExG sensing modality: ear-EEG. EEG signals are generated by the synchronous activity of large populations of neurons, ⁴⁴⁵ primarily in the cerebral cortex. When neurons fire, they create ⁴⁴⁶ current dipoles due to the movement of ions across cell mem- ⁴⁴⁷ branes, generating an electrical field through volume conduc- ⁴⁴⁸ tion. This field can be described by the primary current source J 449 and the secondary volume currents induced in the surrounding ⁴⁵⁰ conductive medium (brain tissue, skull, scalp). Scalp or ear-EEG 451 measures the potential difference between two points: the mea- ⁴⁵² suring electrode and the reference electrode. The potential $V(\mathbf{r})$ 453 at an electrode placed at position **r** due to a current dipole source ⁴⁵⁴ **p** at position **r^p** in an infinite, homogeneous medium with con- ⁴⁵⁵ ductivity σ can be described by simplified dipole analysis [\[20\]](#page-15-0) 456 as shown in Fig. 4: $\frac{457}{457}$

$$
V(\mathbf{r}) = \frac{1}{4\pi\sigma} \mathbf{p} \cdot \frac{\mathbf{r} - \mathbf{r_p}}{\|\mathbf{r} - \mathbf{r_p}\|^3}
$$
(2)

The potential difference measured by the ear or scalp-EEG setup 458 between a measuring electrode at **r** and a reference electrode at ⁴⁵⁹ **r**_{ref} is: 460

$$
\Delta V = \frac{1}{4\pi\sigma} \mathbf{p} \cdot \left(\frac{\mathbf{r} - \mathbf{r}_{\mathbf{p}}}{\|\mathbf{r} - \mathbf{r}_{\mathbf{p}}\|^3} - \frac{\mathbf{r}_{\text{ref}} - \mathbf{r}_{\mathbf{p}}}{\|\mathbf{r}_{\text{ref}} - \mathbf{r}_{\mathbf{p}}\|^3} \right)
$$

$$
\approx \frac{1}{4\pi\sigma} \mathbf{p} \cdot \frac{\mathbf{r} - \mathbf{r}_{\text{ref}}}{\|\mathbf{r} - \mathbf{r}_{\mathbf{p}}\|^3}; \ \|\mathbf{r} - \mathbf{r}_{\text{ref}}\| \ll \|\mathbf{r} - \mathbf{r}_{\mathbf{p}}\| \tag{3}
$$

from which we derive that distance and the angle between ⁴⁶¹ electrodes and dipole moment are the main factors for signal ⁴⁶² characteristics. A first and important consideration is that for ⁴⁶³ closely spaced electrodes located far away from the source, the ⁴⁶⁴ magnitude of the measured potential is directly proportional to ⁴⁶⁵ the distance $D = ||\mathbf{r} - \mathbf{r}_{ref}||$ between the electrodes. Specifi- 466 cally the signal amplitude decreases by a factor proportional ⁴⁶⁷ to the relative difference, $\|\mathbf{r} - \mathbf{r}_{ref}\| / \|\mathbf{r} - \mathbf{r}_{p}\|$, which further 468 depends on the orientation of the electrode geometry $\mathbf{r} - \mathbf{r}_{ref}$ 469 relative to the dipole **p**, producing a null in the measured signal 470 relative to the dipole p, producing a null in the measured signal where $\mathbf{r} - \mathbf{r}_{ref}$, rather than $\mathbf{r} - \mathbf{r}_{p}$, is perpendicular to **p**. The 471 implication for in-ear electrode geometries with mm-scale inter- ⁴⁷² electrode distances is that they can pick up signals originating 473 from the cortical surface that are typically observed by scalp- ⁴⁷⁴ EEG with cm-scale distances, but with attenuated signal levels 475 further aggravated by higher noise levels due to smaller-size ⁴⁷⁶ electrodes resulting in 10–20 dB loss in signal-to-noise dynamic 477 range. 478

However, an equally important consideration is that ear-EEG 479 is able to resolve signals different from scalp-EEG with greater 480 specificity. Specifically, if **r** and **r**ref are relatively closer to the ⁴⁸¹ source **rp**, the measured potential difference will be larger and ⁴⁸² more specific to the source. This preliminary analysis can be ⁴⁸³ applied to contrast the relative merits of ear- and scalp-EEG. ⁴⁸⁴ Scalp-EEG places electrodes over the entire scalp, providing a ⁴⁸⁵ comprehensive view of brain activity, while ear-EEG electrodes ⁴⁸⁶ are placed inside or around-the-ear, providing a "keyhole" view ⁴⁸⁷ of activity from the temporal lobe [\[108\].](#page-17-0) The distance from ⁴⁸⁸ cortical sources to ear electrodes is generally greater than the ⁴⁸⁹ distance to scalp electrodes, potentially reducing signal am- ⁴⁹⁰ plitude in the ear. The exception here is the temporal lobe, ⁴⁹¹ which will be closer to the ear electrodes when placed in the 492 ear canal. The angle of measurement is limited to the relative 493

Fig. 4. Characteristics of brain ExG signals between the scalp and ear. (a) EEG is measured by placing electrodes on the surface of the scalp or in the ear to measure volume currents that yield potential differences. These potential differences are generated by a large number of simultaneously active neurons, which produce current dipoles across a small cortical area, often summarized as an equivalent current dipole. (b) Ear-EEG electrodes are placed in the ear canal or around the ear to measure the brain's electrical activity. The distance between the source and the measuring electrode is smaller than in scalp-EEG setups. While the distance between the ear-EEG electrodes and the source dipoles is generally larger than that of the scalp electrodes, there is an exception for signal sources from the auditory cortex on the same side as the ear-EEG electrodes. (c) Illustration of three common types of EEG measurement referencing setups. Scalp-EEG commonly employs references along the midline of the scalp (Cz shown in the figure) or mastoid. Ear-EEG commonly uses ipsilateral referencing, where the reference electrode is placed in the same ear as the measuring electrodes and contralateral referencing, where the reference electrode is placed in the opposite ear to the measuring electrodes. (d) Sensitivity map for brain sources analyzed by Yarici et al. [20]: (A), (B) Sensitivity map for a left ear unilateral (ipsilateral) ear-EEG montage. (C), (D) Sensitivity map for a bilateral (contralateral) ear-EEG montage. (E, F) Relative sensitivity map for a left ear unilateral montage and a 64-channel scalp-EEG montage. (G), (H) Relative sensitivity map for a bilateral ear-EEG montage and a 64-channel scalp-EEG montage. (A)–(D) The sensitivities displayed for each individual brain sources (dipoles) are extracted from the optimal differential pair of electrodes within the montage (for that dipole). High and low sensitivities are represented by magenta and cyan shading, respectively. (E)–(H) Severe and moderate signal losses are displayed in gray and white, respectively. Signal gains are displayed in red and yellow. (A), (C), (E), (G) Left brain surface. (B), (D), (F), (H) Inferior surface of the brain [20]. (e) Ear-EEG measurement data demonstrate that ear-EEG can pick up even stronger signals than scalp-EEG for sources close to the ear electrodes, such as the auditory cortex in the temporal lobe [13]. (f) Ear EOG measurement in ipsilateral and contralateral referencing. For eye blinks, both referencing setups record the blinking signal, while contralateral referencing records a much larger amplitude. For eyeball movement, ipsilateral referencing records very little signal component, while contralateral referencing records a much larger EOG amplitude [13]. Sensitivity maps A-H in subfigure (d) are ©2023 Yarici, Thornton and Mandic [20].

 position of the ear, primarily capturing activity from lateral and inferior regions of the brain. The much smaller distance between electrodes also decreases the signal amplitude. The analysis here is in line with results from finite element modeling of the brain, which also indicates that ear-EEG only produces an increase in signal amplitude in limited regions in the temporal lobe, while adjacent regions mostly exhibited a moderate decrease in signal amplitude [\[19\],](#page-15-0) [\[26\].](#page-15-0) It has also been shown that despite the limited spatial resolution and lower SNR of ear-EEG, there is a high degree of mutual information between signals captured by ear-EEG and those recorded by scalp-EEG [\[109\].](#page-17-0)

 The difference in amplitude between scalp and ear-EEG has significant implications for the design of sensors and analog front ends (AFE) used in earable EEG devices. The lower amplitude of ear-EEG signals necessitates the use of high-gain, low-noise amplifiers to ensure accurate and reliable signal cap-510 ture. The gain required can be calculated using $G = V_{out}/V_{in}$, where V*in* is the lower amplitude ear-EEG signal and V*out* is the desired output voltage for the analog front end. Additionally,

the signal-to-noise ratio (SNR) is a critical factor, given that ⁵¹³ ear-EEG signals are weaker, the SNR must be maximized by ⁵¹⁴ minimizing noise through the use of low-noise amplifiers, which 515 have a low noise figure calculated as $NF = SNR_{in}/SNR_{out}$. 516 In terms of the biopotential sensors, electrodes must have low 517 impedance to ensure minimal signal loss and high-quality signal ⁵¹⁸ acquisition. The impedance of biopotential electrodes plays a ⁵¹⁹ critical role in the noise performance of earable systems. The ⁵²⁰ thermal noise, which contributes significantly to the overall ⁵²¹ noise in such systems, can be modeled using the equation ⁵²² $V_{n\,rms}^2 = 4kT(G_e + G_{amp}) + V_n^2$. Here k is Boltzmann's con- 523 stant, T is the absolute temperature, G_e is the skin-electrode 524 coupling conductance from the skin-electrode impedance,G*amp* ⁵²⁵ is the amplifier input conductance, and V_n^2 is the input-referred 526 noise of the amplifier. The reduction in noise is crucial for ⁵²⁷ maintaining a high SNR in the AFE. High-resolution analog- ⁵²⁸ to-digital converters (ADCs) are also necessary to accurately ⁵²⁹ digitize the low-amplitude signals, with the resolution given ⁵³⁰ by Resolution = $V_{ref}/2^n$, where V_{ref} is the reference voltage 531

532 and n is the number of bits. The literature have shown that 533 scalp-EEG signals typically range from 10 μ V to 100 μ V, 534 while ear-EEG signals are usually lower, ranging from 1 μ V 535 to 10 μ V [\[110\].](#page-17-0) This lower amplitude necessitates the use of high-gain amplifiers, with ear-EEG requiring a gain of 10 times more than the scalp-EEG. The noise figure for the amplifiers must be exceptionally low to maintain a high SNR due to the smaller signal amplitude of ear-EEG [111]. From an energy standpoint, driven by these requirements of the analog front-end earable sensing systems typically need to maintain a low noise efficiency factor (NEF) to reduce energy consumption while preserving signal fidelity. The higher resolution required by the ADC also asks for optimization of the energy per con- version level figure of merit (FoM) to balance energy effi- ciency with the need for accurate digitization of low-amplitude ⁵⁴⁷ signals.

 3) Ear-EEG Devices: Kaongoen et al. [64] have previously noted the variability in nomenclature in the ear-EEG field. In this review, we use "in- and around-the-ear EEG" shortened to "ear-EEG" to jointly refer to the devices and methods for recording EEG from inside and close to the external ear. When there is a need to refer to only one of these two sub-sets, we use "in-ear EEG" to describe sensors that fit within the auricle of the ear (see Looney et al. [22] for an early example), and "around-the-ear EEG" for devices that contact the hairless scalp 557 behind the ear (for example, the device from Debener et al. [17]). Whereas around-the-ear EEG devices typically use electrode gel to make wet contact with the skin, in-ear devices have been reported as being wet- [15] or dry-contact [27], depending on whether electrode gel is applied before recording. We also note that electrode gel should not be assumed to refer to hydrogel, as alternatives are available that do not dry out, which is essential for long recordings as typical in sleep studies[112]. For electode density, in-ear EEG devices may use high-density montages to characterize spatial variations in voltage [26] or impedance [78] over the ear surface, or the ear canal [31], although most in-ear devices use 8 or fewer electrodes per ear [65]. Around-the-ear devices have mostly adopted a standard montage, with the cEEGrid [81] being the only ear-EEG device in our knowledge to provide an open-source plugin for visualizing topomaps of around-the-ear-EEG activity [113] for EEGLAB [114], an open- source EEG analysis and visualization toolbox that is frequently adopted by ear-EEG studies. Besides the cEEGrid montage, high-density around-the-ear montages have also been used to compare the signal quality of bipolar configurations for record-ing evoked EEG activity from around-the-ear [\[18\].](#page-15-0)

⁵⁷⁸ *B. Optical Sensing*

 Vital signals, including the arterial pulse, blood pressure, and blood oxygen, can be captured through optical approaches. Optical vital sign sensing techniques, such as PPG and pulse oximetry, utilize light to measure changes in blood volume and oxygen saturation, providing a non-invasive and continuous method for monitoring these critical parameters. Optical sensing methods, particularly PPG and pulse oximetry are widely used in in-ear sensors to monitor pulse and blood oxygen saturation $(SpO₂)$ levels. The working principle of PPG involves emitting 587 light from a source, typically a light-emitting diode, into the skin 588 and measuring the amount of light that is absorbed by arteries. ⁵⁸⁹ As blood pulses through these arteries, the varying blood volume 590 changes the amount of light absorbed, which is then detected by 591 a photodetector. This variation in light absorption corresponds ⁵⁹² to the pulse cycle, allowing the measurement of pulse rate [\[115\],](#page-17-0) ⁵⁹³ [116]. ⁵⁹⁴

associates the proposal matrix constrained the constrained tensor in the proposal matrix constrained the system constrained the s *1) Photoplethysmography for Blood Oxygenation, Car-* 595 *diovascular, and Respiratory Monitoring:* For pulse oxime- ⁵⁹⁶ try, the PPG technique is extended by using two light ⁵⁹⁷ sources with different wavelengths—usually red and infrared. ⁵⁹⁸ Hemoglobin in the blood has different absorption rates to these 599 wavelengths depending on whether it is oxygenated or deoxy- 600 genated. By comparing the absorption of the two wavelengths, ⁶⁰¹ the sensor can calculate the ratio of oxygenated to deoxygenated 602 hemoglobin, providing an estimate of $SpO₂$. Fingertip sensors 603 are widely used for $SpO₂$ but can be affected by peripheral 604 vasoconstriction, especially in cold environments. While the ⁶⁰⁵ ear provides a reliable site for $SpO₂$ measurement due to its 606 stable blood flow, offering consistent readings. It has also been 607 shown that in-ear $SpO₂$ response is faster than measurement 608 from the finger. In a previous study, the known phenomena of 609 time delay between central circulation and peripheral circulation 610 has been measured with a mean delay of 12.3 s between the ear 611 and finger when the subjects performed breath holds $[117]$. PPG 612 data are further used to extract additional physiological infor- ⁶¹³ mation, such as heart rate variability (HRV) [118], pulse rate 614 (PR) [116], blood pressure (combined with an air pump) [\[119\],](#page-17-0) ⁶¹⁵ blood glucose $[120]$, and $SpO₂[121]$. Among them, HRV is a key 616 indicator of autonomic nervous system activity, reflecting the 617 balance between the sympathetic and parasympathetic branches 618 of the autonomic nervous system. PR is a fundamental vital sign ⁶¹⁹ that provides insights into cardiovascular health, while blood ⁶²⁰ pressure is a critical indicator of cardiovascular function. $SpO₂$ 621 is a measure of the oxygen saturation level in the blood, reflecting 622 the efficiency of oxygen delivery to tissues. These parameters are 623 essential for monitoring cardiovascular health and stress levels, ⁶²⁴ making PPG a valuable tool for health and wellness applica- ⁶²⁵ tions [1], [115], [122]. Beyond, one work led by Hammour ⁶²⁶ and Mandic further expanded the principle of earable optical ⁶²⁷ sensing to continuous, non-invasive blood glucose monitoring 628 using a pulse oximeter, which is then combined with machine 629 learning models to estimate blood glucose levels [\[120\].](#page-17-0) These 630 parameters are essential for monitoring cardiovascular health ⁶³¹ and stress levels, making PPG a valuable tool for health and ⁶³² wellness applications [\[1\],](#page-15-0) [\[115\],](#page-17-0) [\[122\].](#page-17-0) 633

> Another main direction of research for ear PPG is respiration 634 monitoring, which is critical for understanding and managing 635 various health conditions, including respiratory diseases, car- ⁶³⁶ diac ailments, and stress. The fluctuations of absorption of the ⁶³⁷ two wavelengths are influenced by respiratory cycles, creating ⁶³⁸ modulations in the PPG signal that can be analyzed to extract 639 respiratory biomarkers such as respiratory rate (RR), breathing 640 phases, and tidal volume $[123]$, $[124]$. Apart from PPG, one 641 study from Taniguchi and Nishikawa also investigated using ⁶⁴² infrared light to detect shape changes in the ear canal caused 643

⁶⁴⁴ by breathing movements, offering a non-invasive and motion-⁶⁴⁵ resilient alternative for optical respiratory sensing [\[125\].](#page-17-0)

 However, PPG based optical sensing methods have limita-647 tions as well. The accuracy of PPG and $SpO₂$ measurements 648 can be affected by factors such as skin tone $[126]$, ambient light interference, and motion artifacts. Traditionally, vital signs including blood pressure requires cuff-based monitors for direct measurements, which are bulky and not suitable for continuous monitoring. Earable PPG alone is hard to make accurate estimate on BP. The common practice is to combine PPG with ECG at the ear to measure ECG-to-PPG pulse transit time (PTT) to provide better estimate of blood pressure non-invasively [127], [128], leveraging the ear's stable environment. Though ECG is not easily obtained in an integrated manner in the ear especially using a single ear ECG setup [43]. Additionally, in-ear place- ment poses unique challenges as the ear canal is a less stable measurement site compared to the fingertip or wrist, requiring sophisticated algorithms to mitigate motion artifacts and ensure reliable readings [119], [129].

In a time section in the section of the section of the section in the section of the *2) Body Temperature:* Infrared thermometry at the ear (tympanic membrane) is commonly used due to its proximity to the carotid artery and hypothalamus, making ear a viable location for estimating core body temperature. Previous studies demonstrate that tympanic thermometers can provide real-time, continuous temperature monitoring through infrared sensors integrated into earable devices [124], [130]. These devices, designed with customizable 3D printing techniques, aim to maintain a close fit in the ear canal, enhancing the core body temperature measurement accuracy. However, the accuracy of tympanic temperature measurements can be significantly af-674 fected by various factors. Cárdenas-García et al. $[131]$ found that environmental conditions such as ambient temperature and hu- midity can introduce errors, as the ear canal is exposed to external influences that may not accurately represent the core body tem- perature. Additionally, changes in local blood flow and sensor positioning within the ear canal can cause discrepancies. Chaglla et al. [132] further illustrate this by showing how non-thermal equilibrium conditions can lead to thermal shock errors, necessi- tating a waiting period for the sensor to stabilize before accurate readings can be obtained. Addressing these issues is critical for developing reliable, non-invasive temperature sensing systems for practical and clinical use. Despite these challenges, ear-based core temperature sensing continues to evolve, leveraging ad- vanced materials and designs to offer increasingly reliable and personalized monitoring solutions, particularly for clinical and at-home health applications. Advancements like graphene-inked infrared thermopile sensors have been developed to enhance ac- curacy by improving thermal conductivity and reducing IR light scattering. One study demonstrated that while these materials improve performance, continuous monitoring remains sensitive to positioning and user activity, which can affect the consistency of measurements [\[132\].](#page-18-0)

⁶⁹⁶ *C. Chemical Sensing*

⁶⁹⁷ The in-ear sweat is a rich source of health-related analytes. ⁶⁹⁸ Sweat, produced by eccrine glands, contains water, electrolytes, hormones, and metabolites, playing key roles in thermoregula- 699 tion, stress response, and waste excretion [\[133\].](#page-18-0) In-ear sweat, ⁷⁰⁰ though less studied, has significant potential for advancing our ⁷⁰¹ understanding of human physiology and health monitoring. The ⁷⁰² ear canal, with its unique environment and continuous exposure 703 to external elements, produces perspiration that can provide ⁷⁰⁴ critical insights into the body's biochemical state [\[134\].](#page-18-0) Given ⁷⁰⁵ its proximity to the brain, in-ear sweat may also offer more ⁷⁰⁶ precise indicators of neurological conditions and stress levels ⁷⁰⁷ compared to other sweat sources. In addition, the proximity ⁷⁰⁸ of the ear to the brain implies that in-ear sweat might provide ⁷⁰⁹ a more precise indication of neurological disorders and stress ⁷¹⁰ levels compared to sweat from other parts of the body. Previous 711 research has explored the use of optical sensing [\[120\],](#page-17-0) and 712 electrochemical sensing of biomarkers like glucose, lactate or 713 sodium ion concentrations in the ear [13], [135], [136]. 714

The metabolic profiles in the ear sweat can reflect the body's 715 physiological and pathological state, making it a valuable, non- ⁷¹⁶ invasive medium for health monitoring [137], [138]. Among ⁷¹⁷ the metabolism related biomarkers, one of the most prominent ⁷¹⁸ biomarkers is lactate which is indicative of tissue oxygenation 719 and metabolic stress. Elevated sweat-lactate levels can signal ⁷²⁰ anaerobic metabolism, often associated with strenuous phys- ⁷²¹ ical activity or certain medical conditions such as sepsis and ⁷²² ischemia [139]. Continuous monitoring of lactate can be partic- ⁷²³ ularly beneficial for athletes to optimize training and recovery, ⁷²⁴ as well as for patients in critical care settings [140]. Glucose ⁷²⁵ is another biomarker, which is crucial for monitoring metabolic 726 health and managing diabetes. Sweat glucose levels, although 727 lower than blood and ISF glucose levels, can be correlated ⁷²⁸ with them and offer continuous, non-invasive monitoring for 729 diabetic patients, aiding in better glycemic control and early ⁷³⁰ detection of hypo- or hyperglycemic events [141]. Electrolytes, ⁷³¹ including sodium, potassium, and chloride, are vital for main- ⁷³² taining fluid balance, nerve function, and muscle contractions. ⁷³³ Abnormal levels of these electrolytes in sweat can indicate ⁷³⁴ dehydration, electrolyte imbalances, and disorders such as cystic 735 fibrosis, which is characterized by elevated sweat chloride levels. ⁷³⁶ Monitoring these electrolytes in real time can help manage ⁷³⁷ conditions like dehydration and electrolyte imbalances, ensuring ⁷³⁸ proper hydration and electrolyte replenishment, especially in ⁷³⁹ athletes and individuals exposed to extreme environmental con- ⁷⁴⁰ ditions [133]. Cortisol, the primary stress hormone, is another 741 key biomarker found in sweat. Cortisol levels can provide in- ⁷⁴² sights into an individual's stress response, adrenal function, and 743 circadian rhythms. Abnormal cortisol levels are associated with ⁷⁴⁴ conditions such as Cushing's syndrome, Addison's disease, and ⁷⁴⁵ chronic stress. Continuous monitoring of cortisol through sweat ⁷⁴⁶ can aid in the management of these conditions by providing ⁷⁴⁷ a non-invasive means to track hormonal fluctuations [\[142\].](#page-18-0) In ⁷⁴⁸ addition, in-ear sweat contains a variety of biomarkers such as ⁷⁴⁹ pH levels, proteins, peptides, lipids like cholesterol and squa- ⁷⁵⁰ lene, and neuropeptides. These biomarkers can provide valuable ⁷⁵¹ diagnostic information for conditions such as skin disorders, ⁷⁵² infections, metabolic acidosis, immune responses, inflamma- ⁷⁵³ tion, hypercholesterolemia, oxidative stress, and neurological ⁷⁵⁴ and psychological health [\[143\],](#page-18-0) [\[144\],](#page-18-0) [\[145\],](#page-18-0) [\[146\],](#page-18-0) [\[147\].](#page-18-0) ⁷⁵⁵

79 Ontario and the state of the state Sweat-based lab analysis is commonly used for diagnosing pathophysiological states, but biosensing approaches are gain- ing attention for their real-time monitoring of metabolites and clinically relevant biomarkers [\[148\].](#page-18-0) The common mechanisms that are employed in this regard rely on optical, electrochemical, and mechanical-based biosensing to detect and quantify various biomarkers present in sweat, each offering unique advantages in terms of sensitivity, specificity, and integration [149], [150]. Electrochemical biosensors typically consist of electrodes made from advanced materials such as graphene, carbon nanotubes, and metal nanoparticles, which enhance the conductivity and surface area for analyte interaction [151]. Enzyme-based sensors for glucose and lactate, for example, use enzymes (glucose oxidase and lactate oxidase) that catalyze reactions with the target molecules to generate quantifiable electrical currents com- mensurate with their concentrations. In addition, measuring the potential difference across a selective membrane, ion-selective electrodes detect electrolytes such as sodium and potassium, therefore providing information on hydration state and elec- trolyte balance. So far, several such platforms have been reported representing an advancement of wearable health technology such as continuous monitoring of glucose, ketone, lactate and sodium. [13], [152], [153], [154] platforms, typically designed as flexible, skin-adherent patches, utilize advanced microflu- idic and electrochemical sensing technologies to facilitate the analysis of the various analytes in sweat. Other than that, opti- cal biosensing mechanisms, such as fluorescence, colorimetric, and chemiluminescence detection, complement electrochemical sensors by detecting changes in light properties due to biomarker interactions[155]. Mechanical biosensing, though less common, detects physical changes like pressure or volume associated with sweat production or specific biomarkers. Electrochemical biosensing is widely adopted for its high sensitivity and real-time measurement capabilities.

 Several attempts have been made to detect health parameters in in-ear or proximally located locations. However, sweat-based biochemical monitoring studies are limited in specific ear lo- cations, possibly due to the limitations of sweat harvesting technologies in the delicate sensory organ and the lower den- sity of sweat glands. So far, several chemical biomarkers have been reported using earable sensing platforms. Gil et al. [135] have reported on an ear-worn device that can monitor sweat parameters, including pH, lactate, and cardiovascular param- eters. The electrochemical techniques, amperometry, and po- tentiometry, were employed for monitoring the lactate and pH, respectively. Using this ear-worn device, the temporal profile has been successfully tested on the human subject for lactate and pH. Using a similar concept for lactate electrochemical moni- toring, Xu et al. [\[13\],](#page-15-0) have reported, an in-ear flexible sensing patch that can be installed on the earbuds. This multimodal sensor was coupled with the EEG for synchronous monitor- ing of brain activity and physiological lactate levels in human subjects.

 Despite the enormous attention and advantages associated with in-ear sweat-based sensing such as non-invasiveness and continuous monitoring, various challenges are yet to be ad-dressed to employ these strategies for comprehensive health monitoring. One of the intrinsic challenges is sweat produc- ⁸¹³ tion variability, which changes due to the change in physio- ⁸¹⁴ logical state, hydration status, and environmental conditions. ⁸¹⁵ Inter-individual sweat composition variability occurs evidently 816 due to the diverse genetic setup and the weather conditions 817 they live in, which can severely impact the consistency and ⁸¹⁸ reliability of the sensing data. As the ear locations are prone ⁸¹⁹ to contamination with dust, earwax, cosmetics, etc., these can ⁸²⁰ interfere with the sensor's analytical performance. Comfort, ⁸²¹ fit, and user acceptance are other challenges, that may limit ⁸²² its use for monitoring longer intervals to obtain significant ⁸²³ health information. Considering the potential of in-ear sweat in 824 healthcare monitoring, future works would be directed toward 825 its collection and the enrichment of the analytes for sensitive ⁸²⁶ detection/monitoring of clinically important analytes. 827

D. Mechano-Acoustic Sensing **Example 228**

Mechano-acoustic sensing in wearable ear devices offers an 829 innovative approach to detecting mechanical and acoustic vibra- ⁸³⁰ tions using integrated accelerometers, gyroscopes, and micro- ⁸³¹ phones. These sensors capture physiological activities, such as 832 occlusal force and tongue, jaw, and head movements, transform- ⁸³³ ing these vibrations into meaningful data for health monitoring, ⁸³⁴ human-computer interaction, and motion detection [\[156\],](#page-18-0) [\[157\],](#page-18-0) 835 [158], [159]. In-ear devices are particularly effective at tracking 836 head gestures and subtle movements, providing insights into 837 posture, balance, and even facial expressions [158], [160], [161], 838 [162]. ⁸³⁹

Applications of mechano-acoustic sensing in earables hold 840 significant promise across various domains. In health monitor- ⁸⁴¹ ing, earables have the potential to continuously track physio- ⁸⁴² logical signals such as respiratory patterns [163], [\[164\],](#page-18-0) heart 843 rate [165], [166] and gait analysis. Monitoring these signals 844 holds significant potential for health applications, such as using 845 in-ear mechano-acoustic sensors for gait tracking, which could 846 indicate diseases like Parkinson's $[167]$, and aid in rehabilitation 847 for seniors to improve mobility and prevent falls $[162]$. Another 848 area of application is tracking human activities such as tongue ⁸⁴⁹ movement, chewing, head and body motion, and facial expres- ⁸⁵⁰ sions, which can be utilized for human-computer interaction, ⁸⁵¹ including hands-free control via teeth gestures $[168]$, as well 852 as fitness assessments [169], [170]. For example, BreathPro ⁸⁵³ demonstrates the capability of in-ear microphones to monitor ⁸⁵⁴ breathing modes during running, employing a sophisticated ⁸⁵⁵ signal processing pipeline and machine learning-based classi- ⁸⁵⁶ fication model to enhance accuracy that can be used for fitness 857 assessment [\[171\].](#page-18-0) 858

However, there are limitations and trade-offs with this tech- ⁸⁵⁹ nology. Mechano-acoustic sensors are sensitive to noise from ⁸⁶⁰ external sources and non-relevant body movements, such as 861 head shakes or environmental sounds, which can affect their 862 accuracy in distinguishing between signal types. For example, ⁸⁶³ when trying to detect gait movement, other motion such as 864 chewing or talking will impact the accuracy to distinguish gait ⁸⁶⁵ movement from other movement [\[159\].](#page-18-0) In addition, just like 866 other motion-tracking wearables, earables fallshort in detecting 867

relief to user [\[180\].](#page-18-0) Moreover, ABR is used for assessment of 922 tinnitus [\[183\].](#page-19-0) ⁹²³

B. Transcutaneous Auricular Vagus Nerve Stimulation ⁹²⁴

2003 and determine and the based of the state and the state and the state and the state and determine and the state and the state and the state and the state of the state and the state and the state and the state and the *1) Anatomy and Brainstem Targets:* Vagus nerve (VN) ⁹²⁵ or the 10th cranial nerve is the longest cranial nerve in the ⁹²⁶ body, forming 75% of the parasympathetic nervous system ⁹²⁷ that mediates a state of "rest and digest" [184]. VN emerges ⁹²⁸ bilaterally from the brainstem, connecting the brain to multiple 929 body structures including the heart, lungs, and the gastroin- ⁹³⁰ testinal system (vagus is Latin for wandering). Just under the ⁹³¹ cranium (skull), VN sends off an auricular branch that receives ⁹³² somatosensory input from the auricle, innervating especially the 933 cymba conchae [7], [9], [185], although the literature on auricle 934 innervation in humans is sparse $[186]$ and nerve locations could 935 vary between subjects. As all nerve fibers in the auricle (includ- ⁹³⁶ ing ABVN and other, non-vagal cranial and cervical nerve fibers) 937 run only 1 mm to 1.5 mm deep between the skin and cartilage, ⁹³⁸ the auricle provides easy access for transcutaneous electrical ⁹³⁹ ABVN stimulation (taVNS) $[187]$. taVNS at the cymba conchae 940 recruits sensory ABVN fibers that project directly to the nucleus ⁹⁴¹ of the solitary tract (NTS) in the brainstem, and higher order ⁹⁴² brain structures as evidenced by fMRI studies [\[188\],](#page-19-0) [\[189\].](#page-19-0) ⁹⁴³ A key target of taVNS is the locus coeruleus (LC), the main ⁹⁴⁴ source of norepinephrine (NE) in the brain [190]. Although 945 the mechanisms underlying taVNS's modulatory effects are not ⁹⁴⁶ fully understood, the pathway for taVNS to affect a distant 947 organ or pathology can be considered indirect, as sensory input ⁹⁴⁸ through taVNS could either be directly modulating parasym- ⁹⁴⁹ pathetic vagal efferents with downstream targets [9], or pro- ⁹⁵⁰ ducing systemic (body-level) changes by influencing multiple ⁹⁵¹ neurotransmitters including gamma-aminobutyric acid (GABA) 952 and Norepinephrine (NE) [191], similar to invasive vagus nerve 953 stimulation (VNS) [66], [67]. 954

2) Stimulation Optimization and Dosage: Stimulation pa- ⁹⁵⁵ rameters for taVNS can be set with the goal of delivering a ⁹⁵⁶ target dose of electrical charge [192] at a stimulation intensity ⁹⁵⁷ that could excite ABVN fibers [193]. Optimizable parameters 958 include electrode size and stimulation site [194], and stimulation ⁹⁵⁹ level as determined by current intensity (amplitude, mA), pulse 960 width (μs) , frequency (Hz), duty cycle (%, from pulse width 961 and frequency), pulse-pause ratio (ON/OFF time, distinguish- ⁹⁶² ing tonic and phasic stimulation), pulse shape (monophasic or ⁹⁶³ biphasic), and the total duration of taVNS [195]. Optimization of 964 stimulation parameters has been attempted by recording fMRI, 965 far-field vagus somatosensory evoked potentials (VSEPs), heart 966 rate variability, and pupil diameter, but stimulation protocols are 967 not yet standardized and can lead to mixed results. fMRI findings 968 have provided support for stimulating the cymba conchae or the 969 inner tragus [\[188\],](#page-19-0) [\[189\],](#page-19-0) [\[196\],](#page-19-0) [\[197\].](#page-19-0) Earlier VSEP studies ⁹⁷⁰ have compared pulse amplitudes [\[198\]](#page-19-0) and frequencies near 1 971 Hz [\[199\],](#page-19-0) whereas other VSEP and fMRI studies have run faster 972 stimulation in the range 20 Hz with customized pulse amplitude 973 (current intensity) between each subject's sensitivity and pain ⁹⁷⁴ thresholds $[9]$, $[200]$. If optimizing stimulation parameters using 975 VSEP, it should be noted that muscular artifacts can confound ⁹⁷⁶

 broader human motion compared to leg and torso-worn wearable motion tracking devices [\[158\],](#page-18-0) [\[172\].](#page-18-0) Still, this is a shortcoming to all motion detecting wearables. Wrist-worn devices, although great in detecting arm movement, tendto be inaccurate for gait tracking during slow movement, or when the subject uses a walk- ing aid [173],[174]. Leg and torso-mounted wearables tend to do better for gait, but are not very useful when the subject is doing a stationary task such as driving and VR/AR [175]. Earables are no exception. Therefore, for future research, earables can be a crucial component of a multimodal system which combines data from earables with other body-worn sensors using advanced machine learning algorithms for improving activity monitor- ing [3], [159], [176]. This enables earables to contribute to both head-based and whole-body motion analysis, offering a more complete picture of a user's physical activity and physiological state [177], [178].

884 IV. EARABLE STIMULATION

 Earables can be used to deliver stimuli for neuromodulation using various modalities. In this section, we consider acoustic and electric (current) stimulation. Acoustic stimulation, deliv- ered through sound waves or tones, can be used for therapeu- tic purposes, such as sound therapy in hyperacusis and tinni- tus [180]. Transcutaneous current stimulation of the auricular branch of the vagus nerve at the ear has shown promise in managing conditions like epilepsy, depression, and chronic pain by modulating neural activity in the brainstem and higher brain regions. Other stimulation modalities have also been explored in literature, such as vibrotactile taVNS for improving working memory [181], or rigid ear canal inserts for providing biofeed-back as pressure in bruxism [182].

898 *A. Acoustic Stimulation*

 One of the key advantages of wearable in-ear technology is acoustic stimulation. The close proximity and occlusion of the ear canal enable both discreet hearing and noise cancellation. Beyond everyday use cases, audio stimulation can also target brain modulation. Research has demonstrated that acoustic stim- ulation can influence brain activity using methods like auditory steady-state response (ASSR) and auditory brainstem response (ABR). Furthermore, studies of various audio patterns have revealed potential therapeutic applications for users. In terms of applications, studies have demonstrated promising applications with acoustic stimulation such as tinnitus management, hearing health assessment, cognitive enhancement and relaxation, neu- romodulation for pain and mood disorders, sleep induction and maintenance, and monitoring otoacoustic emissions. Tinnitus is a common symptom where the user hears a sound in the absence of an external source, often associated to damage in the inner ear or an underlying neurological issue. The intensity of tinnitus can vary from mild to severe with brief ringing that is easily masked and doesn't interfere with daily life, to a constant noise that disrupts sleep and affects various activities, respectively. Re- search has shown sound therapy utilizing acoustic stimuli such as white noise, pink noise, and other types of soothing sounds can potentially help mask the ringing from tinnitus providing 977 VSEP results [\[9\],](#page-15-0) and appropriate artifact cancellation is needed to separate brainstem sources from artifactual responses [\[201\].](#page-19-0) Reports of taVNS affecting pupil diameter (as a biomarker of LC-NE activity) have been mixed [\[68\],](#page-16-0) [\[202\],](#page-19-0) [\[203\],](#page-19-0) and a recent study comparing tonic (30s ON, 30s OFF) and phasic (1 s ON, 29 s OFF) stimulation protocols with all other parameters kept the same found transient pupil dilation for both tonic and phasic stimulation [204]. Similarly, inconsistencies have also been observed in taVNS's modulation of resting-state EEG band powers [67], [68] and P300 evoked potentials [192], [203], which may also be attributable to differences in stimulation protocols.

as the contract of the contra Summarizing variations in other taVNS stimulation parame- ters, a systematic review of 41 taVNS randomized clinical trials until July 2020 found interquartile range of stimulation current amplitudes to range from 0.2 mA to 5 mA, pulse width from 0.2 ms to 0.5 ms, and stimulation frequency from 10 Hz to 26 Hz. The choice of pulse width and frequency was seen to be consistent with implantable vagus nerve stimulation (VNS) protocols, which could be further optimized for ABVN as its fiber composition is different from the cervical vagus nerve targeted in implantable VNS [193]. The range of chosen current amplitudes across studies is attributable to possible differences in the stimulation electrode size, impedance of the skin-electrode interface, and procedure for determining the subjects' perceived sensory and pain thresholds [192]. Current-controlled stim- ulation (current clamp) is preferable over voltage-controlled stimulation to accommodate for variations in the electrode-skin interface impedance across subjects and hardware [192], [195]. *3) Safety of taVNS:* Overall, taVNS is considered safe, with the most frequently reported adverse effects being mild and transient ear pain, headache, and tingling [205]. Additionally, there is evidence that taVNS can be self-administered with remote supervision as needed [206]. However, accurate dosage for taVNS is currently unknown and raises potential safety concerns, as invasive VNS studies have reported opposing neu- romodulatory effects with changing dosage, for instance, in protocols targeting inflammation [193]. Furthermore, as effects of stimulation seen in a clinical population may not carry over to a healthy population, multimodal sensing and analysis is crucial for monitoring downstream effects of taVNS, for instance, on multimodal physiological markers assessing stroke volume and contractility in a taVNS study targeting stress [207].

V. EARABLE SYSTEM INTEGRATION

A. Sensing Pipeline

 The signal flow in a multimodal earable system begins at the sensors, where physiological data is detected. ExG signals, being typically weak, require immediate amplification by an analog front-end, which could include amplifiers and analog-to-digital converters as visualized in Fig. 6(a). For chemical sensors, a potentiostat controls the sensor operation and measures the resultant signals. Integrated digital sensors, like those for PPG and temperature, provide direct digital output through standard

Fig. 5. Acoustic and electric stimulation modalities targeting the vestibulocochlear nerve and the auricular branch of vagus nerve (ABVN) respectively, with downstream targets. Left: stimulation modalities with time domain representation of illustrative stimulation patterns: a burst for sound, and a biphasic current pulse. Actual stimulation patterns used vary across studies and pathology. Right: pathway of the stimuli to subcortical and cortical targets. Sound stimulation travels through multiple nuclei including the inferior colliculus (IC) before reaching the auditory cortex in the temporal lobe [179]. ABVN projects to the neucleus tractus solitarius (NTS), with downstream targets including locus coeruleus (LC) and other brain regions [67]. Sources of anatomical drawings: auditory pathway © 2022 Jacxsens, De Pauw, Cardon, van der Wal, Jacquemin, Gilles, Michiels, Van Rompaey, Lammers and De Hertogh [\[179\],](#page-18-0) ABVN pathway © 2021 Sharon et al. [67].

Fig. 6. Schematic overview of the earable biofeedback system. (a) Sensing system showing electrodes, amplifier, and ADC for in-ear EEG acquisition, combined with an integrated PPG sensor. (b) Auditory stimulation system consisting of a digital to analog converter (DAC) and speaker driver (taVNS not shown), (c) Biofeedback earable combining the sensing and stimulation pipelines from (a) and (b) through a modulator which could be realized on an integrated circuit for controlling the biofeedback. taVNS stimulation electrodes are also shown, targeting cymba concha on the top with earlobe reference on bottom (anatomical features not visualized).

serial interfaces such as serial peripheral interface (SPI), inter- ¹⁰³⁰ integrated circuit (I2C), or universal asynchronous receiver- ¹⁰³¹ transmitter (UART). All these signals, whether analog or digital, ¹⁰³² are then processed by a computational unit. Preprocessing or ¹⁰³³ conditioning of signals, such as filtering and noise reduction, ¹⁰³⁴ can be carried out either via dedicated hardware or within ¹⁰³⁵ the computational unit for real-time applications. Alternatively, ¹⁰³⁶

¹⁰³⁷ more extensive processing can be performed on a host ma-¹⁰³⁸ chine via a wireless interface, leveraging greater computational ¹⁰³⁹ power.

¹⁰⁴⁰ *B. Stimulation Pipeline*

 The stimulation pipeline can be thought of as the sens- ing pipeline, but in reverse. To consider the case of acoustic stimulation, sound is delivered by a speaker, driven by a digital- to-analog converter and amplifier for driving the speaker as shown in Fig. 6(b). Current stimulation (not visualized) is conceptually analogous, with electrodes to deliver the current impulses, driven by DACs and amplifiers. Design considerations for DACs and amplifiers will be different for audio or current stimulation.

¹⁰⁵⁰ *C. Multimodal Synchronization*

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The same th[ro](#page-15-0)ugh the same through the same through the same three same that the same three same three same that the same that the same that the same that the same th Ear-EEG devices typically record from one or more channels from one or both ears, and the ear-EEG streams may further be combined with other data streams from scalp-EEG for valida- tion [105] Data synchronization is crucial for integrative analy- sis, and a popular open-source software library that address the synchronization problem is Lab Streaming Layer (LSL) [208]. LSL ensures that all incoming data, regardless of the source, are time-stamped with high precision and synchronized across different data streams. As an illustration, a study evaluating the synchronization of audio streaming from a hearing aid development platform, and a separate ear-EEG stream from a cEEGrid Smarting acquisition could be synchronized within a jitter (standard deviation of latencies across trials) of 3 ms using LSL, making it suitable for developing closed-loop au- dio and ear-EEG processing systems [209], [210]. However, it is important to acknowledge that the complexities of LSL still require careful attention to timing tests of experimental setups, especially for time-sensitive analysis. For more accurate synchronization between streams, system-specific latency and jitter tests such as clock skew and network delay checks are recommended to verify proper synchronization.

¹⁰⁷² *D. Mechanical Shell Design*

 The mechanical design of ear sensor shells plays a crucial role in the effectiveness and comfort of earable devices, particularly when it comes to embedding sensors for the modalities such as ExG, PPG, temperature monitoring, and motion detection. These shells can be broadly categorized into subject-customized and generic designs, each with distinct considerations for accom- modating the necessary electronics, including circuit boards, radio frequency (RF) components, and batteries.

 1) Subject-Customized Earpiece: These are tailored specifically to an individual's ear anatomy. The process begins with a scan of the user's ear canal, capturing the unique contours and dimensions. Using this data, a 3D-printed mechanical shell is fabricated, designed to fit the ear for one particular subject [\[22\],](#page-15-0) [\[51\],](#page-16-0) [\[93\]](#page-17-0) This customized approach is particularly beneficial for ear-ExG, where precise electrode placement and stable contact with the skin are essential for high-quality signal acquisition. The snug fit minimizes ¹⁰⁸⁹ movement artifacts, enhances comfort, and allows for the ¹⁰⁹⁰ reliable long-term monitoring of neural activity. Additionally, ¹⁰⁹¹ customized shells can accommodate other sensors such as PPG ¹⁰⁹² sensors for heart rate and oxygen saturation, which benefit ¹⁰⁹³ from stable contact with the richly vascularized areas of the ear. ¹⁰⁹⁴ Temperature sensors can be integrated to measure core body ¹⁰⁹⁵ temperature from within the ear canal, taking advantage of the ¹⁰⁹⁶ shell's close fit and thermal insulation properties. 1097

2) Generic Earpiece: These, on the other hand, are not ¹⁰⁹⁸ tailored to any specific individual but are instead created to fit ¹⁰⁹⁹ a broad range of users. These designs typically utilize flexible ¹¹⁰⁰ or adjustable components to accommodate various ear shapes ¹¹⁰¹ and sizes, making them more versatile and cost-effective for ¹¹⁰² mass production. Generic earpieces may adopt an earbud-like ¹¹⁰³ structure [13], [25], [89], [91], [211] that includes multiple ¹¹⁰⁴ embedded sensors, such as accelerometers for motion detection, ¹¹⁰⁵ PPG sensors, and temperature sensors. While generic designs ¹¹⁰⁶ may not offer the precise fit of customized shells, advancements ¹¹⁰⁷ in material science and ergonomic design have allowed these ¹¹⁰⁸ shells to achieve a balance between usability, comfort, and ¹¹⁰⁹ performance. 1110

3) Printed Circuit Board (PCB): When considering the in- ¹¹¹¹ tegration of electronics, polyimide flexible printed circuit boards 1112 (PCB) are usually employed to conform to the unique shape of ¹¹¹³ the ear, ensuring that the electronic components are seamlessly ¹¹¹⁴ embedded without compromising the fit or comfort. Flexible ¹¹¹⁵ PCBs are advantageous in these designs as they can be molded to 1116 the intricate curves of the ear, providing reliable connections be- ¹¹¹⁷ tween sensors, amplifiers, and other electronic components. The ¹¹¹⁸ placement of RF components, such as Bluetooth transmitters, ¹¹¹⁹ is carefully considered to minimize interference and preserve ¹¹²⁰ signal quality, often being positioned in areas of the shell that ¹¹²¹ are less likely to experience attenuation due to proximity to skin ¹¹²² or bone. ¹¹²³

Battery placement is another critical factor, the battery is ¹¹²⁴ typically positioned in a location that balances weight distri- ¹¹²⁵ bution and thermal management, often in the outer portion ¹¹²⁶ of the ear where heat dissipation is more effective, thus pre- ¹¹²⁷ venting discomfort during prolonged use. In both customized ¹¹²⁸ and generic designs, the careful integration of electronics is ¹¹²⁹ essential to maintain the overall functionality, comfort, and ¹¹³⁰ performance of the earable device. The design must account for ¹¹³¹ the unique thermal, mechanical, and electronic challenges posed ¹¹³² by the small, complex environment of the ear, ensuring that all ¹¹³³ components work together harmoniously to deliver accurate, ¹¹³⁴ reliable health monitoring. Lastly, sound hole structures are ¹¹³⁵ usually included in either customized or generic earpiece to ¹¹³⁶ preserve the fundamental acoustic transmission functionality of ¹¹³⁷ the earable system. 1138

VI. CLOSING THE LOOP 1139

This section motivates building a closed-loop biofeedback ¹¹⁴⁰ system by combining the sensing and stimulation systems into 1141 one earable, as visualized in Fig. [6\(c\).](#page-11-0) Here we focus on some key ¹¹⁴² ingredients that could help optimize the biofeedback's control ¹¹⁴³

Fig. 7. Various configurations for combining sensing, stimulation, and biofeedback in an earable. Brain and body coupling visualized here through cervical vagus nerve (VN) originating from the brainstem. (a) Earable as a monitoring system in the user's natural acoustic environment (Env.), with no additional stimulation provided by the earable. Monitoring targets include brain activity as projected to in-ear and around-the-ear electrodes (as a pink beam), and cervical vagus nerve activity projecting to lower around-the-ear electrodes (as a yellow beam), motivated by results from [212]. (b) Earable with open-loop stimulation and continuous monitoring. Stimuli include electrical (taVNS) and acoustic stimulation to the ear. (c) Closing the loop through a biofeedback controller. Visualized ear-EEG devices are croc V2 [31] (©2023 IEEE), and cEEGrid [17].

¹¹⁴⁴ policy to be used for adapting the stimulation given incoming ¹¹⁴⁵ sensory information.

¹¹⁴⁶ *A. Continuous Monitoring With Environmental Stimuli*

 Given the potential of earables to collect long durations of unlabelled data, and analysis methods to assess changes in a subject's state, for example, in ear-EEG sleep studies[\[46\],](#page-16-0)[\[213\],](#page-19-0) we first consider earables for continuous assessment of biomark- ers and physiomarkers in the presence of only environmental stimuli, that is, where no stimulation is applied by the device, as 1153 shown in Fig. $7(a)$.

 In a series of three studies, Hölle et al. have demonstrated that ear-EEG can be recorded beyond-the-lab using a wearable setup [\[214\],](#page-19-0) [\[215\],](#page-19-0) [\[216\].](#page-19-0) These studies measured ERPs either in response to auditory oddball stimuli [\[214\]](#page-19-0) or naturalistic sounds [\[215\],](#page-19-0) [\[216\],](#page-19-0) delivered either in a lab, office, cafeteria, or home-office environment. Larger P300 ERP responses to target stimuli were seen, compared to standard stimuli, but

naturalistic sounds did not evoke strong ERPs [\[216\].](#page-19-0) For oddball 1161 stimuli, the test tones were not deemed disruptive by sub- ¹¹⁶² jects performing office work, but ear-EEG responses to natu- ¹¹⁶³ rally occurring sounds that are ecologically meaningful to the ¹¹⁶⁴ participants, such as their names or ringtones, could also be ¹¹⁶⁵ considered [214]. 1166

To enable real-life recording of synchronized EEG with con- ¹¹⁶⁷ current soundscapes: AFEx (Audio Feature Extraction Frame- ¹¹⁶⁸ work) and Record-a. AFEx enables real-time audio capture, ¹¹⁶⁹ privacy-preserving feature extraction from the audio, and LSL ¹¹⁷⁰ streaming of three features: power spectral density, root-mean ¹¹⁷¹ square power, and sound onsets. Audio can be captured using 1172 wired microphones, which could be worn binaurally behind ¹¹⁷³ the ears (to not occlude environmental sounds from entering ¹¹⁷⁴ the ear canals), connected to a smartphone running AFEx. ¹¹⁷⁵ Simultanseously, ear-EEG is recorded and streamed using a ¹¹⁷⁶ wearable data acquisition device. The smartphone then runs ¹¹⁷⁷ Record-a for synchronized recording (using LSL) of the incom- ¹¹⁷⁸ ing streams carrying audio features and ear-EEG. Although new ¹¹⁷⁹ audio features could be implemented, for instance, as extracted ¹¹⁸⁰ by models of the auditory periphery [217], [218], for tuning ¹¹⁸¹ feature extraction for pathologies such as hearing impairment. ¹¹⁸² Any new features, however, will have to be evaluated for ¹¹⁸³ privacy [215]. 1184

B. Open-Loop Stimulation 1185

It may seem trivial to first deploy a biofeedback earable with- ¹¹⁸⁶ out the feedback, that is, in open-loop, but open-loop stimulation ¹¹⁸⁷ can have beneficial outcomes. This configuration is visualized ¹¹⁸⁸ \ln Fig. 7(b). 1189

As evidence of potential benefits of open-loop stimulation, ¹¹⁹⁰ we summarize accumulating findings from the Gamma EN- ¹¹⁹¹ trainment Using Sensory Stimuli (GENUS) program. Sensory ¹¹⁹² stimulation aimed at entraining gamma oscillations in a mouse ¹¹⁹³ model of Alzheimer's disease (5XFAD) has shown a reduction ¹¹⁹⁴ in amyloid plaques in the visual cortex when using 40 Hz light ¹¹⁹⁵ stimulation $[219]$, in the auditory cortex and hippocampus when 1196 using 40 Hz audio stimulation (1 ms long, 10 kHz tones), ¹¹⁹⁷ and more widespread reduction of plaques in the neocortex ¹¹⁹⁸ when audio and visual stimulation is combined with aligned ¹¹⁹⁹ onsets [220]. Intriguingly, the same rate of 40 Hz was found ¹²⁰⁰ to be the most effective for both modalities, individually and ¹²⁰¹ when delivered together. The choice of 40 Hz was motivated by ¹²⁰² noting that reduced gamma power is reported for Alzheimer's ¹²⁰³ mouse models with a clearance mechanism identified as 40 Hz ¹²⁰⁴ stimulation recruiting the glymphatic system, critical for re- ¹²⁰⁵ moving metabolic waste (and plaque) from the brain [\[221\].](#page-19-0) ¹²⁰⁶ 40 Hz sensory stimulation has also been applied to mouse ¹²⁰⁷ models of neurodegeneration, with 40 Hz visual [\[222\],](#page-19-0) or 40 Hz ¹²⁰⁸ vibrotactile [\[223\]](#page-19-0) stimulation entraining gamma activity and ¹²⁰⁹ reducing pathology. Finally, a feasibility pilot in humans with ¹²¹⁰ mild Alzheimer's has shown positive outcomes for 40 Hz au- ¹²¹¹ diovisual stimulation [\[224\].](#page-19-0) Therefore, open-loop stimulation ¹²¹² across multiple sensory modalities can activate pathways that ¹²¹³ may still be beneficial in certain pathologies and target brain ¹²¹⁴ areas. ¹²¹⁵

Fig. 8. Closed-loop acoustic/auditory stimulation (CLAS) concept and its effects on slow-wave sleep from a validation study in an animal model (rat) by Moreira et al. [225]. Top: Acoustic stimulation's phase with respect to ongoing slow oscillations in slow-wave sleep. Stimulation targeted either the (a) up-phase, (c) down-phase, or (b) no stimulation was provided (mock). Bottom: Grand-averaged time course of delta activity (0.5 Hz to 4 Hz) over 12 days of training with CLAS, where (d) targeting up-phase enhanced delta activity, (e) no stimulation (mock) had no significant effect, and, (f) targeting down-phase decreased delta activity. Gray shadows below plots show significant time-points (multiple t-tests, Holm–Sidak corrected). Time: 0–12 correspond to light, and 12–24 are dark periods. PT: Pre-Training (8 days), MT: Motor-Training (4 days). Rat subjects were trained to perform a single-pellet reaching task (SPRT), timing shown. ©2021, Moreira et al. [225].

1216 *C. Earables for Closed-Loop Biofeedback*

 As depicted in Fig. 7(c), earables can be used to deliver closed-loop acoustic and current stimulation, where sensory monitoring is used to control stimulation parameters and dosage, thus closing the loop. In this section, we provide examples of stimulation strategies that have shown promising results using acoustic stimulation or taVNS in the biofeedback literature, positioning them for follow-up translational work using earables to provide the biofeedback.

 1) Closed-Loop Acoustic Stimulation: Sleep and stress management can have an important impact on improving peo- ple's quality of life. With increasing urbanization, there is more exposure to external noise, and such ambient sounds typically have an adverse effect on sleep. Although barriers such as ear- muffs can help reduce disturbance from external noise, studies have demonstrated that various forms of acoustic stimulation can also help mask the undesirable sounds [\[226\].](#page-19-0) Acoustic stimulation also has the potential to alleviate stress, promoting relaxation, utilizing calming soundscapes to lower anxiety levels and encourage relaxation [\[227\].](#page-19-0) Next, we review supporting evidence for one particular strategy for closed-loop acoustic stimulation and its impact on enhancing slow wave sleep.

 Closed-loop Auditory (Acoustic) Stimulation (CLAS) is to play acoustic stimuli, such as brief 1/f pink noise bursts, in-phase with ongoing slow oscillations (0.1 Hz–1 Hz) observed during non-rapid eye movement (NREM) sleep.This stimulation strat- egy is visualized in Fig. 8, reproduced from a recent validation study in an animal model [\[225\].](#page-19-0) For human subjects, a recent review of CLAS by Esfahani et al. [\[228\]](#page-20-0) summarizes stimulation 1244 parameters and results from 14 CLAS studies from 2013 to 2022, ¹²⁴⁵ all providing encouraging evidence for increasing the amplitude ¹²⁴⁶ of slow oscillations, and a potential for improving memory ¹²⁴⁷ consolidation, although reports have been mixed for memory ¹²⁴⁸ effects, possibly confounded by stimulation parameters, target ¹²⁴⁹ group, and off-target stimulation applied to delta wave activity ¹²⁵⁰ (smaller amplitude local events) instead of slow oscillations ¹²⁵¹ (larger amplitude global slow waves) [229]. 1252

2) Earables for Closed-Loop taVNS: Given the accessibil- ¹²⁵³ ity of ABVN for electrical stimulation from the auricle, and the ¹²⁵⁴ feasibility of ear-EEG for measuring attention biomarkers such ¹²⁵⁵ as alpha power, merging taVNS and ear-EEG for closed-loop ¹²⁵⁶ (CL) attention modulation of taVNS stimulation parameters us- ¹²⁵⁷ ing simultaneous ear-EEG has been suggested previously [\[123\].](#page-17-0) ¹²⁵⁸ The form factor of in-ear EEG devices could provide access ¹²⁵⁹ to taVNS stimulation sites including the conchae (cymba and ¹²⁶⁰ cavum), tragus, and the ear canal, while around-the-ear devices ¹²⁶¹ such as the cEEGrid could be adapted for tragus stimulation. 1262

CL-taVNS systems could support phasic taVNS protocols ¹²⁶³ by time-multiplexing ear-EEG recording and taVNS to avoid ¹²⁶⁴ stimulation artifacts from corrupting ear-EEG. For tonic pro- ¹²⁶⁵ tocols, stimulation artifact reduction could be achieved using ¹²⁶⁶ real-time compatible Generalized Eigenmode Decomposition ¹²⁶⁷ (GED) [230], or device constraints permitting, a separate stim- ¹²⁶⁸ ulation reference channel could be added behind the earlobe for ¹²⁶⁹ artifact removal [231]. 1270

The matrix [of](#page-20-0) the matrix of the matrix The optimal EEG biomarker of LC activity as mediated by ¹²⁷¹ taVNS can be expected to evolve as the mechanistic under- ¹²⁷² standing of taVNS advances. For instance, using alpha power ¹²⁷³ as a biomarker of LC activity as suggested previously [\[66\]](#page-16-0) was ¹²⁷⁴ not reproducible in a replication study [68], but alpha activity ¹²⁷⁵ could still be modulated by taVNS during cognitive tasks [\[191\],](#page-19-0) ¹²⁷⁶ [232]. In addition to alpha power, ear-EEG devices have been ¹²⁷⁷ validated for recording other brain responses including the P300, ¹²⁷⁸ and extending ear-EEG to multimodal earables could addition- ¹²⁷⁹ ally provide ear-ECG, ear-PPG, electrochemical sensing, and ¹²⁸⁰ derivative biomarkers such as heart rate [233], HRV [\[118\],](#page-17-0) ¹²⁸¹ and breathing phase [123]. Breathing phase could especially ¹²⁸² be relevant for protocols aligning taVNS stimulation with the ¹²⁸³ expiration phase [234], or their invasive VNS counterparts [\[235\].](#page-20-0) 1284

D. Earables With On- and Off-Target Nerve Activity ¹²⁸⁵ *Monitoring* 1286

Earables also harbor the possibility of measuring neural ¹²⁸⁷ biomarkers of vagus nerve activity. Cervical electroneurography ¹²⁸⁸ is a recent non-invasive method for recording cervical VN ¹²⁸⁹ activity from the neck using an adhesive array of Ag/AgCl ¹²⁹⁰ electrodes[\[212\].](#page-19-0) Two of the rostrally placed electrodes of the re- ¹²⁹¹ ported electrode array appear visibly close to the L5, L6, R5, and ¹²⁹² R6 electrodes of around-the-ear cEEGrid devices, suggesting ¹²⁹³ that cEEGrids could be evaluated for non-invasive monitoring of ¹²⁹⁴ cervical VN activity as a downstream target for acoustic/taVNS ¹²⁹⁵ biofeedback. Non-invasive monitoring of ABVN activity has ¹²⁹⁶ also been attempted using in-ear electrodes to assess the auto- ¹²⁹⁷ nomic nervous system's response under physiological stressors ¹²⁹⁸

 (cold face test and cold pressor test) [\[236\],](#page-20-0) but given the pre- liminary stage of in-ear ABVN monitoring, follow-up source localization studies can eliminate possible confounds through in-silico modeling [\[237\],\[238\],](#page-20-0) or minimally-invasive recording such as microneurography to measure simultaneously from the cervical VN [206] and other, off-target nerves in the auricle, such as greater auricular nerve in the ear lobe [239].

 Regardless of the in-ear electrodes picking ABVN activity or sympathetic efferents, the possibility of accessing a neural biomarker from the ear could help monitor the efficiacy of biofeedback, analogous to using evoked compound action poten- tials (eCAPs) measured from cervical VN for dosing VNS [240]. To summarize, earables combining stimulation with monitoring through ear-ExG, chemical sensing, and potentially VN activity, could become a candidate platform for optimizing biofeedback dosage with simultaneous stimulation and monitoring of down-stream effects through the same earable.

1316 VII. CONCLUSION

As the contribution of the state of the The ear offers a rich source of brain and body biosignals that can be unobtrusively tapped as a highly versatile and powerful means for continuous cognitive and metabolic health monitor- ing, and further combined with equally unobtrusive stimulation applied to the ear for biofeedback and neuromodulation therapy. The ability of earables to integrate stimulation mechanisms with sensing capability provides a non-invasive, comfortable, and socially acceptable way to deliver therapeutic interventions. This integration not only enhances the functionality of earable technology but also opens new avenues for personalized health management and neurotherapy, leveraging the ear's unique anatomical and neural connections for effective and unobtrusive stimulation. Follow-up studies are needed to establish the longer- term outcomes and optimization of stimulation control based on fused biosignals, but the potential of earables for person- alizing bioelectronic therapeutics with continuous monitoring may lead to engineered naturalistic remediation of drug-resistant pathologies.

1335 **ACKNOWLEDGMENT**

 The authors would like to thank Virginia R. de Sa, Fiza Singh, Lara M. Rangel, Andrea Chiba, Benjamin L. Smarr, Shlomo Dubnov, Vikrant Jaltare, M. Florencia Assaneo, Shihab Shamma, John R. Iversen, Dick Lyon, Simon Carlile, Jason Mikiel-Hunter, Tzzy-Ping Jung, and attendees of the 2021 Tel- luride Neuromorphic Cognition Engineering Workshop, Sim- ula Summer School in Computational Physiology, Australian Hearing Hub's Signal Processing for Hearing – Lecture Series, and the van Vreeswijk Theoretical Neuroscience Seminars for helpful discussions and insights that directly and indirectly shaped our outlook on neuromodulation.

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