

A 16-Output 10V Compliant Stimulator ASIC with Sub-10nA Mismatch and Simultaneous ETI Sensing for Selective Neural Stimulation

Haoming Xin, Meiyi Zhou, Roland van Wegberg, Peter Vis, Konstantinos Petkos, Shrishail Patki, Nicolo Rossetti, Mark Fichman, Vojkan Mihajlovic, Carolina Mora Lopez, Geert Langereis, Mario Konijnenburg, Nick Van Helleputte

Abstract—This paper presents a 16-output high-voltage (HV) compliant stimulator ASIC for selective neural stimulation. The ASIC supports temporal interference stimulation (TIS) to achieve high-spatial selectivity without requiring nerve-penetrating electrodes. A novel on-the-fly active CB is proposed since existing charge balancing (CB) solutions cannot be directly applied to TIS. Simultaneous electrode-tissue impedance (ETI) sensing is realized by reusing the CB hardware. The stimulator ASIC, fabricated in 130nm Bipolar-CMOS-DMOS (BCD) technology, occupies 0.29mm² per output and achieves 10V compliance while supporting up to 10mA stimulation current and maintaining TIS steering flexibility. The proposed active CB approach compensates for electrode voltage drift during TIS based on a negative feedback loop, achieving a sub-10nA mismatch current over a wide range of ETIs. The ETI sensing reuses the stimulation current and CB hardware for simultaneous measurements during stimulation, achieving a sensing inaccuracy of $\pm 2\Omega$. Extensive saline experiments confirm the ability of the ASIC to achieve superior spatial selectivity for stimulation while maintaining proper active CB and simultaneous ETI sensing.

Index Terms—Neural stimulator, high-voltage stimulator, charge balancing, electrode-tissue interface, peripheral nerve stimulation, selective neural stimulation, temporal interference stimulation.

I. INTRODUCTION

NEUROMODULATION devices have been used in clinical practice for treating various neurological conditions and are expected to benefit from novel technological advances to address the rapid expansion of the patient population [1]. Modulating the activity of the vagus nerve (VN) as one prominent example of peripheral nerve (PN), can offer the possibility of regulating and repairing essential bodily functions, given that the VN is a key component of the parasympathetic nervous system [2-5]. Recognizing its importance, neural stimulation (e.g. VN stimulation) has been approved by the FDA for the treatment of drug-resistant

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Haoming Xin, Meiyi Zhou, Roland van Wegberg, Peter Vis, Konstantinos Petkos, Shrishail Patki, Nicolo Rossetti, Mark Fichman, Vojkan Mihajlovic, Geert Langereis, and Mario Konijnenburg are with the IMEC-Netherlands, 5656 AE Eindhoven, The Netherlands (email: haoming.xin@imec.nl, meiyi.zhou@imec.nl, roland.vanwegberg@imec.nl, peter.vis@imec.nl, konstantinos.petkos@imec.nl, shrishail.patki@imec.nl, nicolo.rossetti@imec.nl, mark.fichman@imec.nl, vojkan.mihajlovic@imec.nl, geert.langereis@imec.nl, mario.konijnenburg@imec.nl).

Carolina Mora Lopez and Nick Van Helleputte are with the IMEC, Leuven 3001, Belgium (email: carolina.moralopez@imec.be, nick.vanhelleputte@imec.be).

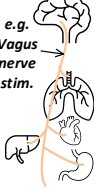
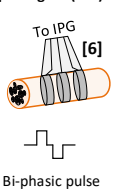
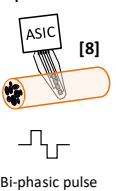
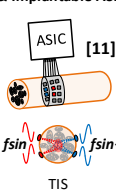
Application: PNS e.g. Vagus nerve stim.	Cuff electrode & implantable pulse gen. (IPG) [6]	Transverse intra-fascicular micro-electrode & Implantable ASIC [8]	High density cuff electrode & Implantable ASIC [11]
			
	Bi-phasic pulse	Bi-phasic pulse	TIS
Invasiveness	LOW	HIGH	LOW
Stim. selectivity	LOW	HIGH	HIGH

Fig. 1. Comparisons of different approaches for PNS.

epilepsy and depression [4] and is being explored for its potential benefits in treating chronic inflammation and arrhythmia, conditions that significantly impact patient quality of life [2-3]. However, current PN stimulation (PNS) techniques are not without limitations. Existing solutions using bipolar or tripolar cuff electrodes [6] stimulate the whole nerve with very poor spatial nerve fiber selectivity. Hence therapies based on traditional cuff electrode stimulation often have many undesired side effects due to the stimulation of non-target areas. To improve spatial selectivity, invasive methods based on transverse intra-fascicular micro-electrodes have been proposed [7]. While such methods can indeed achieve much more targeted stimulation, the necessity to insert electrodes into the nerve can pose significant risks, including nerve damage and subsequent complications, making them unsuitable for long-term chronic therapeutic use [8].

To address these challenges, emerging systems leveraging temporal interference stimulation (TIS) have been proposed. Initially applied in the brain [9-10], TIS has later been adapted for PNs [11] to enhance spatial selectivity using non-penetrating electrodes. While TIS is a comparatively new stimulation paradigm and hence its effectiveness is still actively being (pre)-clinically investigated, TIS shows promise in providing side-effect-free PNS [12]. TIS involves the use of 2 differential current stimulators that generate semi-continuous sinusoidal currents in the kHz range with a small frequency offset (e.g. $\Delta f=10\text{Hz}$) [9]. Each stimulator requires an independent current source and sink to reduce crosstalk between stimulators. This setup creates an interference pattern within the tissue. Since neural cells are more receptive to low frequency (LF), they would be entrained by the LF envelope [9]. Another attractive feature of TIS is its capability of steering

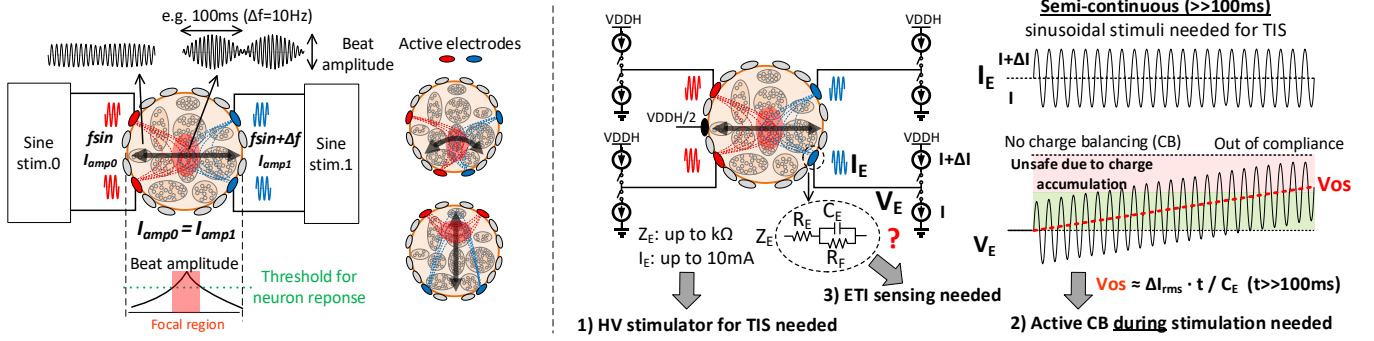


Fig. 2. (left) Illustration of the working principle of TIS (in 2D for simplicity); (right) ASIC requirements for nerve stimulation employing TIS.

the stimulation focal region (i.e. the LF envelope) within the tissue without moving the electrodes. This is achieved by changing the relative stimulation current amplitudes between the 2 stimulation pairs (Fig. 2 left), and the steering direction is determined by the position of the active electrodes. By utilizing a multi-output TIS ASIC with on-chip output selection switches and high-density cuff electrodes, and by carefully selecting the active electrodes at different locations, the stimulation focal region within the PN can be flexibly controlled.

Here we present a neuromodulation ASIC employing TIS that besides spatial selectivity meets the following requirements:

1) High-voltage (HV) compliance: For a broad range of therapies, stimulation currents up to 10mA could be needed [5, 13]. However, small cuff electrodes (mm-range diameter) used in PNS have high impedance. For instance, a 1mm diameter platinum black electrode has an impedance exceeding 1k Ω [14], even with the kHz carrier frequency used in TIS. Thus, the stimulator circuit should ideally provide an HV compliance of $\geq 10\text{V}$. Together with the multi-output requirement for flexible steering, an area-efficient HV TIS ASIC is essential.

2) Charge-balancing (CB) during TIS: CB is crucial for neural stimulators to prevent electrode and/or tissue damage. Unlike the traditional bi-phasic pulse stimulation, TIS is a semi-continuous stimulation method [9]. Taking $\Delta f = 10\text{Hz}$ as an example, a stimulation time $\gg 100\text{ms}$ would be needed to produce a train of LF envelopes. Such long durations mean that even a small mismatch current (ΔI) between the current sink and source would cause significant voltage drift (V_{os}) on the electrode (Fig. 2 right). Hence, CB during TIS is necessary for safety, and the mismatch current should be minimized (preferably $\ll 100\text{nA}$) to prevent V_{os} building up during stimulation [15].

3) Electrode-tissue impedance (ETI) sensing: ETI sensing provides vital information about the quality of the electrode as well as the electrode-tissue contact and is often used to assess optimal electrode selection and stimulation parameters. Furthermore, it offers a way to measure implant degradation over time due to for example scar tissue formation [16]. Therefore, ETI sensing with sufficient accuracy is a highly desirable feature.

While TIS ASICs have been demonstrated [10-11], they have low voltage compliance (3.3V), do not support CB during stimulation, nor allow ETI sensing during TIS. To tackle these

challenges, we present a 16-output HV TIS ASIC with a novel on-the-fly CB and simultaneous ETI sensing. This paper is an extension of [17]. A prototype is fabricated in 130nm Bipolar-CMOS-DMOS (BCD) technology. The proposed ASIC architecture and stimulator circuit are designed to achieve 10V compliance by leveraging HV transistors in BCD technology. It supports stimulation currents of up to 10mA, while occupies only 0.29mm² per output and maintaining flexibility for TIS steering. The proposed active CB approach monitors the electrode voltages and performs the compensation on the fly with a negative feedback loop at each active stimulator output, achieving a sub-10nA mismatch current. The proposed ETI sensing leverages stimulation current for measurement during stimulation by reusing the CB hardware, achieving $\pm 2\Omega$ accuracy.

The paper is organized as follows. Section II introduces the architecture of the proposed ASIC. Section III details the circuit implementation. Section IV presents the measurement results, and Section V concludes the paper.

II. ARCHITECTURE

As discussed previously, TIS needs 2 independently controlled differential sinusoidal current stimulators with 4 outputs to realize its basic function. Stimulators with adjustable stimulation current amplitude offer stimulation focal region steering capability. However, this simple system with 4 outputs allows steering of the stimulation focal region in only one direction, since the 4 stimulation electrodes connected to these outputs are fixed in place after implantation. To enable flexible steering of the focal region in multiple directions, the ASIC should support a significantly larger number of outputs (well beyond 4) that can be connected to high-density cuff electrodes and should have the ability to flexibly select the 4 active outputs for TIS. Since the stimulator needs to support HV compliance with a large stimulation current, the transistors in the output stage are large. Therefore, it is important to minimize the number of these components to reduce chip area while keeping the desired output count for flexible TIS steering.

Fig. 3 (top) gives 3 stimulator architecture options for an $N \times$ outputs TIS AISC. All the 3 options have 2 independently programmed current DACs (I-DACs), generating the desired sinusoidal currents for TIS. The I-DAC current is then copied

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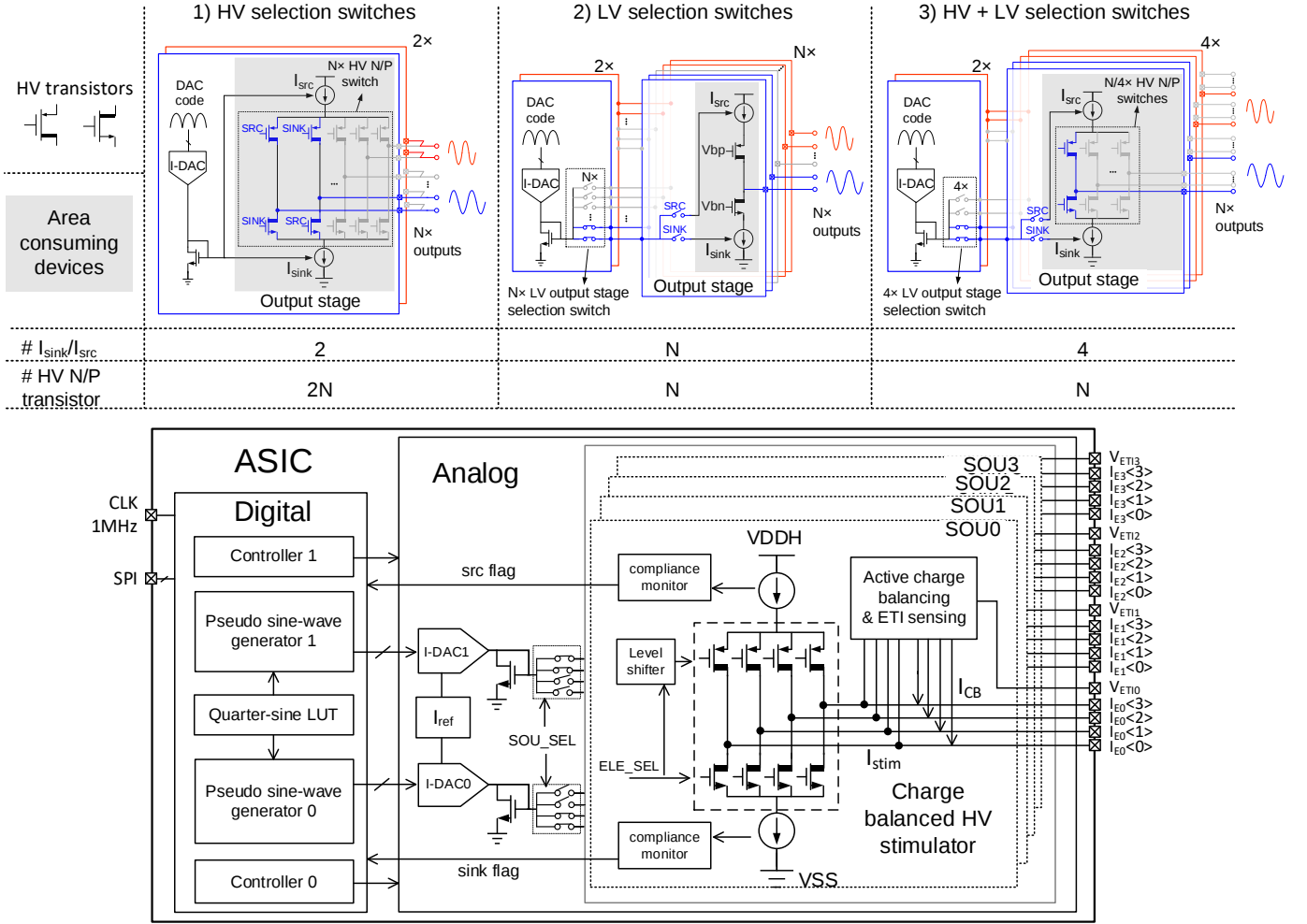


Fig. 3. Comparison of 3 TIS architecture options for $N \times$ stimulation outputs; (bottom) the proposed ASIC architecture.

and amplified to the stimulation output stage with current mirrors using low-voltage (LV) devices for better matching with smaller sizes as compared to HV devices. HV devices are then used as cascode transistors to protect the LV devices against HV during operation. The 130nm BCD technology used in this work offers LV (1.5V/3.3V/5V) complementary metal-oxide-semiconductor (CMOS) transistors and HV double-diffused metal-oxide-semiconductor (DMOS) transistors, which can tolerate high drain-source voltages for HV operation. Unless otherwise specified, the terms LV and HV devices refer to the 1.5V CMOS transistors and HV DMOS transistors, respectively. To offer a flexible electrode configuration such that any stimulation output can be selected to produce the sinusoidal current profile from either of the 2 I-DACs, analog switches are needed for active output selection. Note that the outputs that are not selected will remain inactive during stimulation. The first option includes $2 \times$ stimulation output stages, and each output stage has $N \times$ N- and P-type HV transistors to support $N \times$ outputs [18], from which 4 active outputs can be selected. Although this option uses the least number of current sources ($2 \times I_{\text{sink}}/I_{\text{src}}$), it requires $2N \times$ HV N/P transistors for flexible output selection, and each HV N/P transistor requires a level shifter for proper switching, which further increases design complexity and chip area. Instead of

using complex HV switches, LV switches can be implemented at the gate of the diode-connected transistor that copies the I-DAC current [19], as shown in Fig. 3, the second option. The LV switches then select four output stages, which are configured to operate in two pairs for TIS. As the LV switches only need to drive the gates of the current mirrors, their size is negligible compared to the transistors in the output stage. However, in this case, $N \times$ stimulation output stages are needed to support $N \times$ outputs. Each output stage contains a pair of I_{sink} and I_{src} and their corresponding HV devices for protection, resulting in a total number of $N \times I_{\text{sink}}/I_{\text{src}}$ and $N \times$ HV N/P transistors. The proposed option combines the usage of LV and HV switches to reduce the number of area-consuming devices while offering stimulation output selection flexibility. It includes $4 \times$ output stages that work in pairs for TIS, and each output stage contains a $1:N/4$ HV demultiplexer (DMUX), supporting total $N \times$ outputs. The LV switches select which 2 output stages work in a pair, while the HV DEMUX selects which output within each output stage is active. This approach requires a total of $4 \times I_{\text{sink}}/I_{\text{src}}$ and $N \times$ HV N/P transistors. Considering that the size of the HV devices is typically larger than (or at least comparable to) the LV devices used for $I_{\text{sink}}/I_{\text{src}}$ and that N needs to be much larger than 4 for flexible focal region steering, the proposed architecture is the most area-

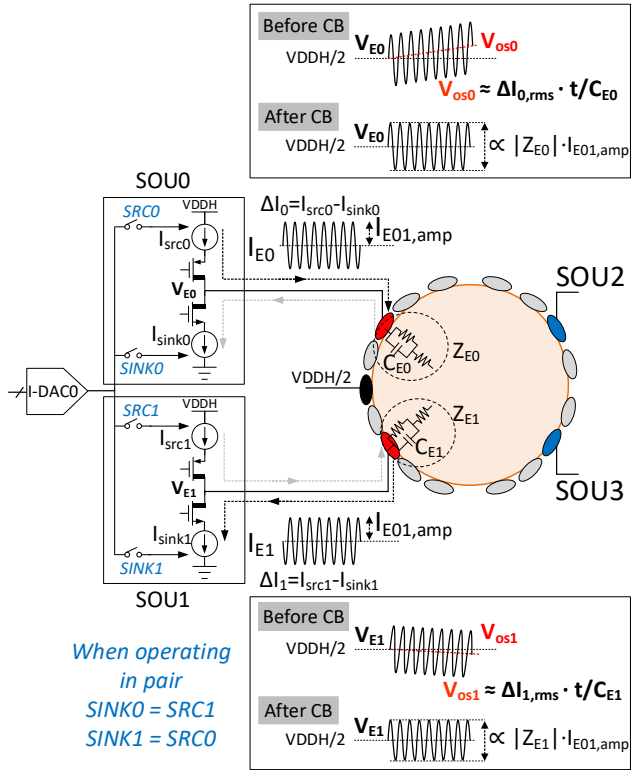


Fig. 4. TIS setup illustration using the proposed ASIC.

efficient choice among these 3 architecture options.

The complete ASIC architecture is shown in Fig. 3 (bottom). It contains 4 stimulation output units (SOU), SOU0-3, and each SOU has an output demultiplexer to 4 electrodes, supporting a total of $N=16$ electrodes. On-chip digital pseudo-sine synthesizers generate the stimulation current control signals based on a quarter-sine wave look-up table (LUT). Two 9-bit current-steering DACs (I-DAC0/1) similar as in [18], produce the frequency-offset currents which are mirrored into the SOUs. The I-DACs operate on a 1MHz clock to achieve sufficient frequency accuracy and to closely approximate the desired kilo-Hz sinusoidal stimulation current. To offer focal region steering capability, the amplitude of the sinusoidal stimulation current is programmable. This is done by adjusting the I-DAC reference current, which sets the I-DAC's full range from 255.5 μ A to 1.022mA in 255.5 μ A increments, thereby adjusting the stimulation current amplitude from 2.55mA to 10.22mA in 2.55mA steps. Optionally, finer adjustments to the stimulation amplitude can be made by modifying the amplitude of the pseudo-sine wave digital signal, though this reduces resolution as the current amplitude decreases. However, these smaller steps provide more precise control for focal region steering. High- and low-side compliance monitoring is implemented in each SOU, which raises a warning flag when the electrode voltage is excessively high or low.

Fig. 4 illustrates how the proposed ASIC is used for TIS. In this example, SOU0/1 and SOU2/3 are used in pairs respectively for TIS, and SOU2/3 and the HV DEMUX in SOU0/1 are omitted for simplicity. When working in a pair, the same stimulation current amplitude and frequency are used for SOU0/1 but with reversed stimulation polarity. This ensures a

well-defined current path within the tissue, which is crucial to reduce crosstalk between stimulation pairs for TIS [9]. 2 SOU pairs are synchronized using the same 1MHz clock and the timing of each pair can be independently programmed. A dedicated low-impedance body bias electrode sets the body to $V_{DDH}/2$. Due to the mismatch between I_{sink} and I_{src} , each SOU has a random mismatch current (ΔI) being injected into the tissue through the electrode, causing a voltage drift (V_{os}) on the electrode depending on the RMS value of ΔI and the ETI capacitance C_E . Therefore, each SOU has its dedicated CB circuit. The use of CB results in stable electrode voltages that can be further used for ETI sensing during stimulation, as described in Section III-C.

III. CIRCUIT IMPLEMENTATION

A. I-DAC and HV stimulator

The detailed circuits of the current steering DAC and the HV stimulator and its compliance monitoring circuits are shown in Fig. 5. The I-DAC is implemented with 9-bit resolution and employs a current-steering architecture to reduce spikes caused by code-switching. It is segmented with 3 unary and 6 binary bits to improve linearity, and the least significant bit (LSB) can be programmed from 0.5 μ A to 2 μ A by changing its reference current. The current from the LV I-DACs is mirrored (and amplified 10 $\times$) to either M_{sink} or M_{src} based on whether the stimulator is operating in sink or source mode. M_{sink} and M_{src} are implemented with LV core devices (1.5V) to save the area. M_{sink} is a standard NMOS device, while M_{src} is an LV PMOS device with extra HV protection rings. The 10 $\times$ amplification reduces the I-DAC full-scale current to 1.022mA for a maximum stimulation current of 10.22mA. As the I-DAC is operating under a 1.5V supply, its power consumption is thus negligible compared to the HV stimulator. The I-DAC current can be further reduced by increasing the current mirror gain. However, this approach could introduce settling issues with the gate voltage of the current mirror, which would degrade the linearity of the sinusoidal stimulation current.

To provide a sinusoidal current with sufficient linearity (setup with <0.4% THD was used in [9]), the stimulator should have a sufficiently high output impedance, such that the stimulation current is not distorted by the large voltage swing established on the electrode. To boost the output impedance, a regulated HV cascode $M1$ is used for M_{sink} . $M1$ serves two purposes as it at the same time also protects M_{sink} from the HV and forms the HV output DEMUX for electrode selection. To ensure adequate gate voltage to turn on $M1$, the operational amplifier used in the active cascode configuration operates under a 3.3V supply and utilizes thick-oxide LV transistors rated for 3.3V. In principle, the output impedance of M_{src} could similarly be boosted with a regulated cascode structure. However, this would require a floating ground rail (e.g., 3.3V below V_{DDH}) to implement the LV operational amplifier used for the regulated cascode device [14]. Instead, M_{src} is protected against HV with a normal cascode HV device $M3$, and $M2$ serves as the second cascode device to further boost the output impedance to a sufficient value for linearity and implements the HV DEMUX for electrode selection. It is worth noting that the

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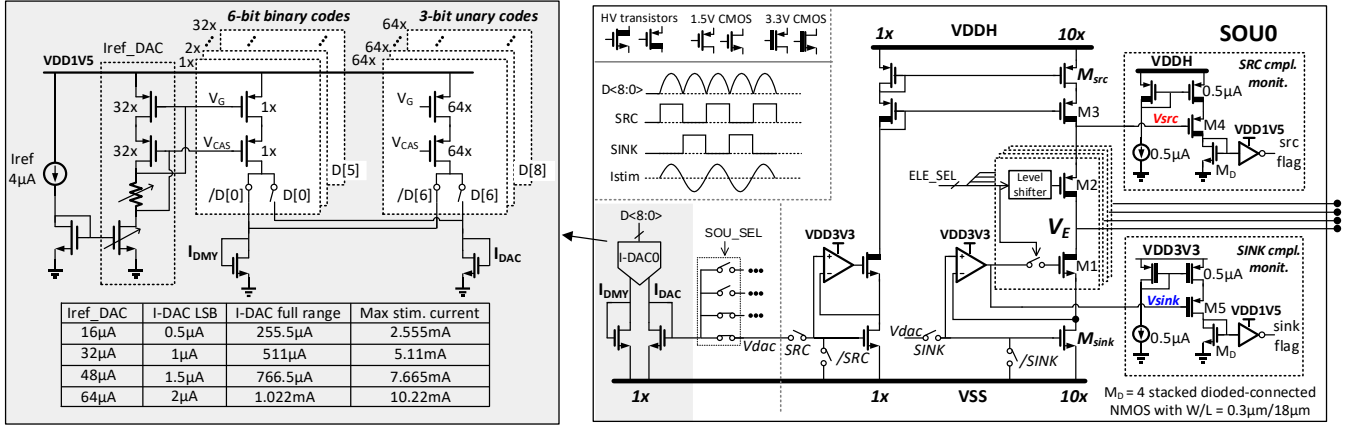


Fig. 5. Current steering I-DAC and HV stimulator.

output impedance mismatch between the current sink and source can cause current distortion. However, since both are designed with sufficiently high output impedance, the distortion caused by this mismatch is minimized. Using this topology, the stimulator circuit can operate flexibly from 5V to 12V by simply adjusting the stimulator supply $VDDH$, offering system-level flexibility to save power when possible. An additional HV device $M3$ is used in each stimulator as compared to the proposed option shown in Fig 3. However, the area overhead is limited, as the number of area-consuming HV devices is increased from $32\times$ to $36\times$ for the 16 outputs of this prototype. The overhead would be even smaller when a larger number of outputs within each SOU is used.

A reference-voltage-free compliance monitor with embedded HV-to-LV level shifting is proposed and implemented for each HV stimulator. When the electrode voltage V_E is abnormally high, V_{src} is pulled towards $VDDH$, turning off $M4$ and triggering the high-side compliance flag in the LV domain. The low-side compliance monitor works similarly but checks the output of the regulated cascode amplifier [14] without the need for HV devices. Using this simple structure results in the 2 compliance monitors occupying only 0.02mm^2 in total and each consumes only $1\mu\text{A}$ of current. Unlike the traditional inverter-based configuration [14], the proposed current source compliance monitoring circuit does not require a floating ground, allowing it to operate more flexibly with different $VDDH$ levels and ensuring compatibility with the HV stimulator circuit proposed in this work. While the inverter-based configuration may occupy less area due to its simplicity, the overall area-saving benefit is minimal since the chip area is primarily dominated by the stimulator output stage.

B. Charge balancing during TIS

Due to imperfections in the fabrication process, current mismatches between the I_{sink} and I_{src} must be considered for each SOU [14-15]. Conventional passive CB solutions (Fig. 6(a)), which rely on bulky external capacitors, are impractical for such a system with 16 electrodes due to space limitations. They are also unreliable for alternating current (AC) stimulators, as they can only reject direct current (DC) conduction into tissue [20]. An active CB approach is necessary

to effectively compensate for current mismatches.

Several active CB solutions have been proposed for biphasic stimulators [14], [18], [20-25]. Most of them monitor residual voltages at the stimulator outputs after each stimulus and then provide a means of current compensation (Fig. 6(a)). However, since stimulation bursts can be quite long ($>100\text{ms}$) in TIS, charge accumulation can reach unsafe levels during stimulation before it can be detected, as shown in Fig. 2. Detecting V_{os} is particularly challenging due to the continuous nature of sinusoidal waveforms. The unpredictable phase shift between I_E and V_E , induced by the ETI at different sinusoidal frequencies, further complicates the timing control for detection. Therefore, existing techniques developed for biphasic current stimulation cannot be directly applied to TIS. In this work, we propose a novel active CB method based on on-the-fly voltage monitoring. During stimulation phases, the voltage drift V_{os} at each active output is monitored and is maintained within safe limits by a feedback loop. Fig. 6(a) shows the implementation of the proposed CB loop. Each CB loop has 4 inputs/outputs, multiplexed to 4 outputs of each SOU. For optimal power and area efficiency, this loop begins with a capacitive divider that generates a scaled copy of the electrode voltage (V_E). Before the initiation of stimulation, the output of the capacitive divider is reset to V_{REF} , and the stimulator outputs are initially biased to $VDDH/2$, the same as the reference electrode. By choosing an appropriate attenuation factor (F), the voltage monitoring circuits can be fully implemented in the LV domain.

As shown in Fig. 2, V_{os} can drift at a rate of $\Delta I_{rms}/C_E$ (R_F is often large) during TIS, with a smaller C_E exacerbating this effect. Since an attenuation factor F is applied, the mean voltage of V_{SEN} will experience a voltage drift at a rate of $\Delta I_{rms}/(C_E \times F)$ accordingly. The voltage monitoring block needs to extract this near-DC change from the AC voltage signal. A low-pass filter (LPF) could be used for this purpose. However, with the lowest sinusoidal frequency of 1kHz in our application, extracting V_{os} change would require a high-order LPF with a cut-off frequency below 100Hz. The corresponding passive components at these frequencies are impractically large, making LPFs sub-optimal for this application. Instead, as shown in Fig. 6, two peak detectors with complementary architectures [26] are used to

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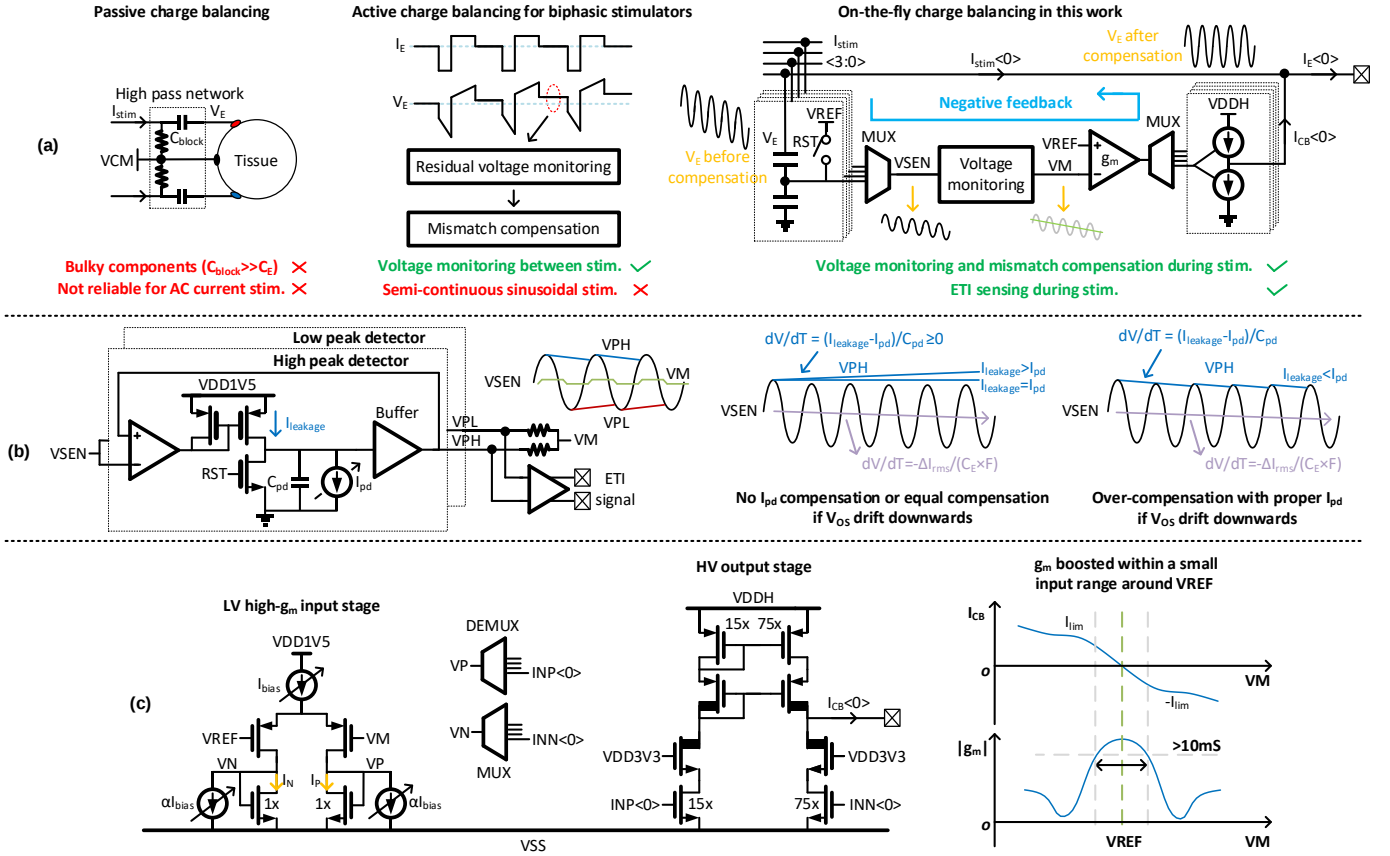


Fig. 6. (a) Comparison of the proposed CB approach with existing solutions in literature; (b) implementation of the voltage monitoring circuit; and (c) implementation of the current-bleeding g_m amplifier.

obtain the peak values (V_{PH}/V_{PL}) of V_{SEN} . The mean voltage of V_{SEN} can be simply generated at VM as $(V_{PH}+V_{PL})/2$. Moreover, the peak information can be further used for ETI sensing as will be explained later.

The operation of the peak detectors can be understood as follows (taking the high peak detector as an example): Initially, C_{pd} (and hence also V_{PH}) is reset to ground. As long as $V_{SEN} > V_{PH}$, the pre-amplifier, and current mirror will charge C_{pd} until $V_{SEN} = V_{PH}$, effectively implementing a peak detector. However, leakage from the current mirror ($I_{leakage}$) would continuously increase V_{PH} , resulting in an inability to properly track the peak of V_{SEN} . To counteract this, $I_{leakage}$ can be over-compensated by using a programmable I_{pd} ($I_{pd} > I_{leakage}$) [27]. In this way, V_{PH} always decreases slightly at a rate of $(I_{leakage} - I_{pd})/C_{pd}$, as shown in Fig. 6(b). The use of over-compensation ensures that

$$\frac{|I_{leakage} - I_{pd}|}{C_{pd}} > \frac{|\Delta I_{rms}|}{C_E \times F'} \quad (1)$$

guaranteeing that V_{PH} can accurately track the peak of V_{SEN} even if V_{OS} drifts downwards.

The stability of this feedback loop relies on timely updating V_{OS} . As shown in (1), a smaller C_E needs a larger I_{pd} due to the rapid change in V_{OS} ; thus, I_{pd} is programmable to support C_E down to 100nF. However, the use of over-compensation can introduce unwanted ripples at V_{PH} and V_{PL} , potentially

compromising the accuracy of VM generation. The ripple amplitude $V_{ripple,amp}$ can be estimated as

$$V_{ripple,amp} \approx \frac{|I_{leakage} - I_{pd}|}{C_{pd}} \times \frac{1}{f_{sin}}, \quad (2)$$

indicating increased ripple amplitude at a lower frequency. To address this issue, $I_{leakage}$ is minimized to be negligible compared to I_{pd} across PVT variations by using thick transistors, which ensures that the $V_{ripple,amp}$ is mainly influenced by I_{pd} and C_{pd} . I_{pd} and C_{pd} are then matched in the two peak detectors, allowing the introduced voltage ripples at VM to be suppressed by self-cancellation, as depicted in Fig. 6(b).

VM is then compared to a reference voltage (V_{REF}) in a transconductance (g_m) amplifier. The resulting output current I_{CB} can be expressed as $I_{CB} = g_m \times (V_{REF} - VM)$. Since the change of VM reflects $|V_{OS}/F|$, larger g_m minimizes V_{OS} as in $|I_{CB}/g_m| = |V_{OS}/F|$. It's worth mentioning that during the initial sinusoidal cycles of each stimulation, V_{PH}/V_{PL} will require 1-3 cycles (simulated over PVT variations) to track the peak values and stabilize VM . Therefore, the g_m amplifier is enabled with a programmable delay proportional to the sinusoidal periods. Once the CB loop stabilizes, I_{CB} compensates for the mismatch current ΔI of the core stimulator, thus $I_{CB} = \Delta I$. In this design, a g_m of more than 10mS over PVT variations is achieved to regulate V_{OS} within $\sim 100mV$, as per our simulations.

The g_m amplifier consists of two stages: an LV high- g_m input

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stage and an HV output stage. The input stage achieves a g_m of $\sim 0.2\text{mS}$, which is further amplified by a factor of $75\times$ by the output stage. Typically, achieving high g_m requires a large bias current (I_{bias}). However, this bias current will be amplified in the output stage through the NMOS current mirror, leading to significant static power consumption from the HV supply. To address this, two current-bleeding sinks (αI_{bias} , $\alpha=10/22$) are introduced at the input pair, which can boost the g_m within a small input range around V_{REF} while keeping the bias current in I_P/I_N low ($\sim 1\mu\text{A}$ only). As a result, most current consumed by the HV output stage can be used as output current to counteract ΔI . Since ΔI generally increases with the stimulation current I_{stim} , the input stage offers programmable flexibility, enabling power savings for smaller I_{stim} .

Fig. 6(c) illustrates the relationship between I_{CB}/g_m and VM . Outside of the boosted range, g_m decreases significantly, allowing only a limited output current around I_{lim} . I_{lim} is designed to be larger than the 3σ mismatch current of the SOU in the worst case. Therefore, VM will be limited around V_{REF} when the CB loop is stable and low V_{os} is ensured.

C. Electrode-tissue impedance sensing

As previously mentioned, ETI sensing offers extra insights into TIS therapies. Typically, ETI sensing involves applying a small current to the electrode and measuring the resulting voltage [28-29]. However, this approach may not be ideal for TIS. To avoid saturation due to stimulation artifacts, sensing is often performed before or after a stimulation pulse train, which could be too sporadic for TIS. Dedicated stimuli and readout circuits are often required, increasing power and area overhead. The additional current injection may also introduce unwanted interactions with the nerve.

In this work, we utilize an added benefit of the proposed CB approach for ETI sensing. Since the peak-to-peak value of the V_E signal inherently correlates with $|Z_E|$ (magnitude of the ETI) multiplied by the stimulation current I_E (Fig. 4), $|Z_E|$ can be derived by leveraging the outputs of the peak detectors as in

$$|Z_E| = \frac{(V_{PH} - V_{PL}) \times F}{2 \times I_{E,amp}}. \quad (3)$$

Since $V_{PH}-V_{PL}$ is nearly constant around DC with CB, an on-chip differential buffer (as shown in Fig. 6(b)) can easily output V_{PH} and V_{PL} as ETI signals without the need for a dedicated readout. Minimal power and area overhead are introduced, making this approach well-suited for TIS.

IV. MEASUREMENT RESULTS

The ASIC is implemented in a 130nm HV BCD technology. A chip photograph with size indications and area breakdown is shown in Fig. 7. The HV stimulator and the active CB & ETI sensing circuit for each SOU occupy 0.51mm^2 and 0.58mm^2 respectively for 4 outputs, translating to an area of 0.13mm^2 and 0.15mm^2 per output, respectively. The total active area, including all analog blocks, for 16 stimulation outputs is 4.64mm^2 , resulting in an area of 0.29mm^2 per output. The chip is first characterized with a set of electrical measurements. Saline experiments are then performed by connecting the chip to a nerve cuff electrode. Further, a TIS demonstration is

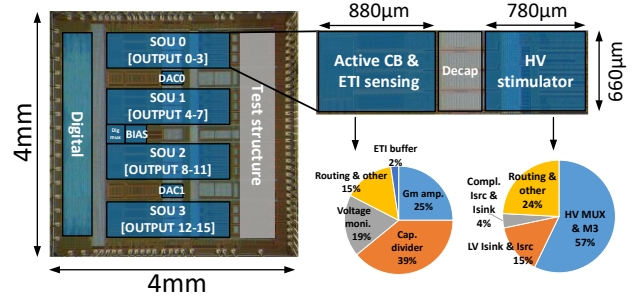


Fig. 7. Die photo with size indications and area breakdown.

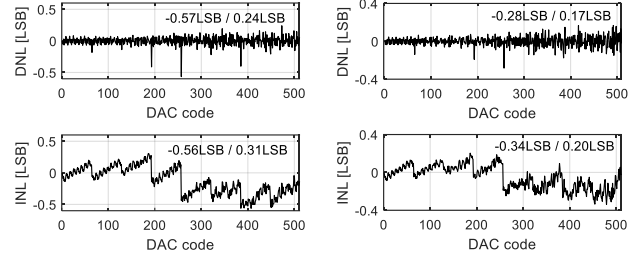


Fig. 8. Measured INL/DNL of I-DAC: (left) $I_{LSB} = 0.5\mu\text{A}$, and (right) $I_{LSB} = 2\mu\text{A}$.

performed in saline showcasing the flexible focal region steering capability of the proposed ASIC.

A. Electrical characterization

The static linearity performance of the I-DAC is characterized at its maximum and minimum LSB settings, as shown in Fig. 8. With an LSB current of $0.5\mu\text{A}$, the measured integral nonlinearity (INL) and differential nonlinearity (DNL) errors are within $-0.57/0.24$ LSB and $-0.56/0.31$ LSB, respectively. These numbers reduce to $-0.28/0.17$ LSB and $-0.34/0.20$ LSB when the LSB current is set to $2\mu\text{A}$. Fig. 9 (top) shows the DC measurement results of a single SOU by sweeping the stimulator output (V_E) from 0 to 12V with a source meter for different current amplitudes. As shown, stable output currents are achieved for both I_{sink} and I_{src} with a worst-case output impedance of $>1\text{M}\Omega$. This ensures a $<0.5\text{LSB}$ ($\text{LSB}=20\mu\text{A}$) error with a 10V swing on V_E , which is crucial for the linearity of the sinusoidal stimulation current and for ETI sensing accuracy. The compliance monitoring reliably triggers when the current drops by $\sim 0.5\%$ from its nominal value. The current source mismatch between I_{sink} and I_{src} is characterized (with CB disabled) at all the 9-bit settings, and the worst-case value is $29\mu\text{A}$ at 10.22mA (Fig. 9 (bottom left)). Furthermore, the current mismatch is quantified at 10.22mA across 24 SOUs from 6 different chips to obtain the statistical spread, as shown in Fig. 9 (bottom right). Without CB, the measured 3σ mismatch is $103\mu\text{A}$ at 10.22mA , which is well within the designed I_{lim} in Fig. 6(c) for the active CB.

To evaluate the proposed CB circuit, a sinusoidal stimulation current is generated with a 10.22mA amplitude. To test the CB under worst-case conditions, we deliberately set a 1% mismatch between I_{sink} and I_{src} (hence $\sim 3\sigma$ mismatch at 10.22mA). Under these conditions, the electrode voltage V_E is measured from the electrode model. Fig. 10 top demonstrates how the proposed CB regulates V_{os} with $C_E = 1\mu\text{F}$ as an example. With CB disabled,

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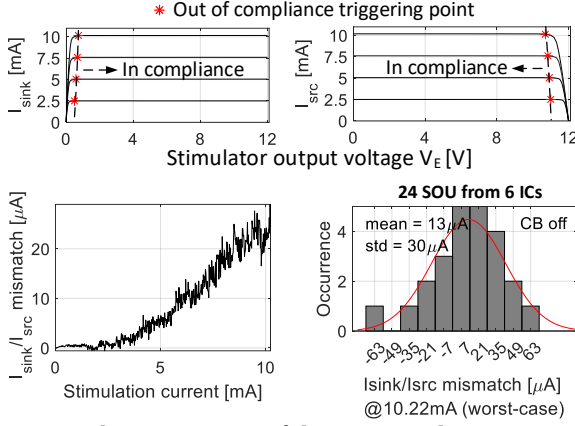


Fig. 9. DC characterization of the HV stimulator.

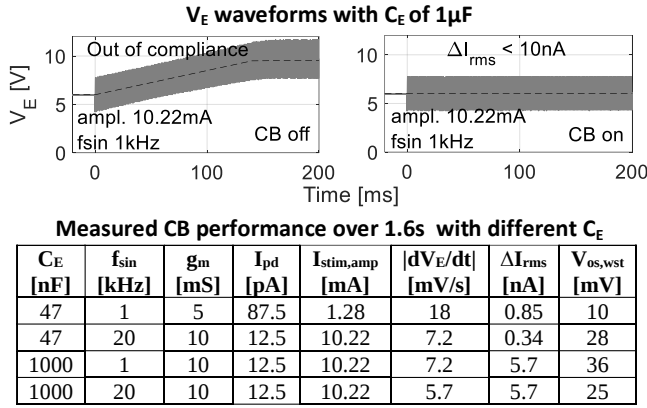


Fig. 10. Electrical characterization of the proposed CB with different ETI capacitances.

V_E drifts out of compliance within 200ms at a rate of $|dV_E/dt| = 25$ V/s, indicating an equivalent RMS mismatch current ΔI_{rms} of $25\mu A$, calculated as $\Delta I_{rms} = C_E \times dV_E/dt$. When CB is enabled, $|dV_E/dt|$ is reduced to 7.2 mV/s, corresponding to an ΔI_{rms} of 7.2 nA. As previously mentioned, the CB loop is optimized for a minimum C_E of 100 nF in simulations. Here we further evaluate CB performance for C_E values ranging from 47 nF to $1\mu F$, with each stimulation lasting up to 1.6 s (Fig. 10 bottom). For low frequency and small C_E , the amplitude of I_{stim} is reduced to ensure that V_E stays within compliance. I_{pd} is properly chosen based on a lookup table, ensuring that the lowest I_{pd} is provided to maintain good CB performance with minimal voltage ripples at VPH/VPL. Generally, $|dV_E/dt|$ below 20 mV/s are achieved, and $V_{os,wst}$ consistently remains below 50 mV within 1.6 s. For small C_E (47 nF) and low frequency (1 kHz), $|dV_E/dt|$ is slightly higher. One potential reason is that the voltage ripples introduced by the peak detectors are more significant according to (2), which degrades the feedback loop accuracy. It's important to note that V_{os} may continue to drift with $|dV_E/dt|$ if a longer stimulus is applied. Consequently, the maximum stimulus duration might be constrained by $|dV_E/dt|$ performance to ensure that $V_{os,wst}$ remains within safe limits.

To verify the ETI sensing, different resistors are added to the stimulator output to mimic various tissue impedances. Since $VPH-VPL$ is nearly a DC value when the CB loop is stable, its rms value can be measured using a multimeter. Note that as the

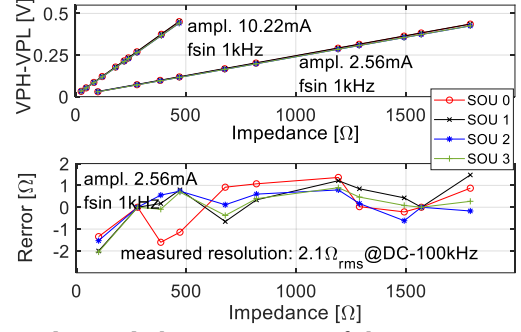


Fig. 11. Electrical characterization of the ETI sensing with stimulation current amplitude of 2.56 mA and 10.22 mA.

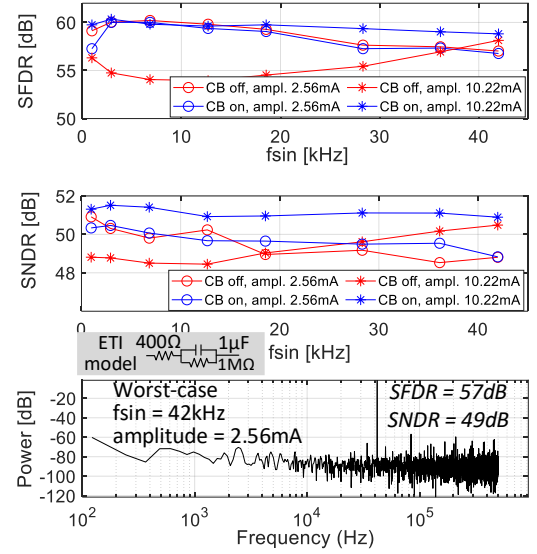


Fig. 12. Measured SFDR (top), SNDR (middle) and spectrum of the stimulation current at worst-case (bottom).

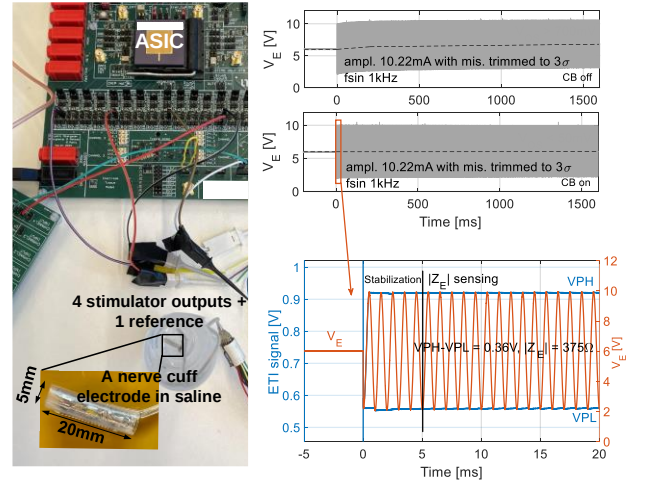


Fig. 13. Characterization of the proposed CB and ETI sensing with a nerve cuff electrode in saline.

proposed ETI sensing method measures only the magnitude of the entire ETI, adjusting the resistor value is equivalent to modifying the ETI magnitude in a more complex R&C model. As mentioned in Section III-B, low frequency leads to larger

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ripples at V_{PH}/V_{PL} , which can degrade the ETI sensing performance; the characterizations are therefore performed with the lowest sinusoidal frequency of 1kHz. Fig. 11 shows the measured results from 4 SOUs. Smaller current amplitude offers a wider sensing range with reduced accuracy. Impedance values up to $1.8k\Omega$ can be measured using 2.56mA amplitude. After applying a 2-point calibration, an error of $\leq 2\Omega$ is achieved.

Fig. 12 illustrates the dynamic performance of the generated sinusoidal stimulation current, which is measured using the indicated ETI model. SFDR and SNDR performance with and without enabling CB is compared. The current mismatch is worst at 10.22mA, which can degrade SFDR significantly. Thanks to the CB, SFDR can be further improved with a worst-case of 57dB for frequencies up to 42kHz.

During TIS operation, the chip power is largely dominated by the SOUs which operate at HV domain. For example, with a stimulation current of 10mA per pair, the total chip power is around 175mW, with the SOUs accounting for 96% of this total.

B. Saline experiments of CB with a nerve cuff electrode

A first saline test is performed with a nerve cuff electrode submerged in saline (0.9% NaCl). A sinusoidal stimulation current is generated at 1kHz with 10.22mA amplitude. Fig. 13 shows the CB and ETI sensing performance. The proposed CB approach ensures a small V_{os} ($< 50mV$) over a long stimulation time (1.6s). After the initial settling period, a consistent $V_{PH}-V_{PL}$ is established, which is recorded by an oscilloscope as shown in Fig. 13. Since the recorded voltage $V_{PH}-V_{PL}=0.36V$, the measured $|Z_E|$ is 375Ω , which is in line with the expected value for this platinum electrode (size: $1.2mm \times 1.3mm$).

It's worth mentioning here that the CB performance might be compromised due to the inherent characteristics of the ETI (e.g. self-discharging or electrode self-potential), which is a potential limitation for electrode voltage compensation. Therefore, the primary goal of this CB approach is to ensure that the charge on the electrode does not build up and that V_{os} does not increase over time, preventing both electrode and tissue damage.

C. TIS demonstration in saline

Fig. 14 (top) shows a more complicated saline measurement setup specifically designed to validate the stimulation focal steering capability of the proposed ASIC using TIS. A beaker is filled with saline water (0.9% NaCl). 16 stimulation electrodes, which are connected to the chip, and 1 reference electrode ($V_{DDH}/2$) are attached at the circumference of the beaker. The electrode and beaker sizes ($1.5cm/10cm$ diameter) are selected to match the proportions of the small cuff electrodes used around a VN (e.g. $\sim 0.75mm/5mm$ diameter). TIS is performed with carrier frequencies of 1kHz and 1.01kHz and varying amplitudes. The electric field inside the saline solution is measured in an X/Y fashion [9] with recording probes containing 4 stainless-steel wires spaced 9mm apart. For each grid point (0.5cm pitch) we measured the local differential voltage (E_x and E_y) between the recording electrodes X_1, X_2 and Y_1, Y_2 for X and Y direction, respectively. The envelope amplitude ($E_{x, AM}$ and $E_{y, AM}$) of E_x and E_y at the differential

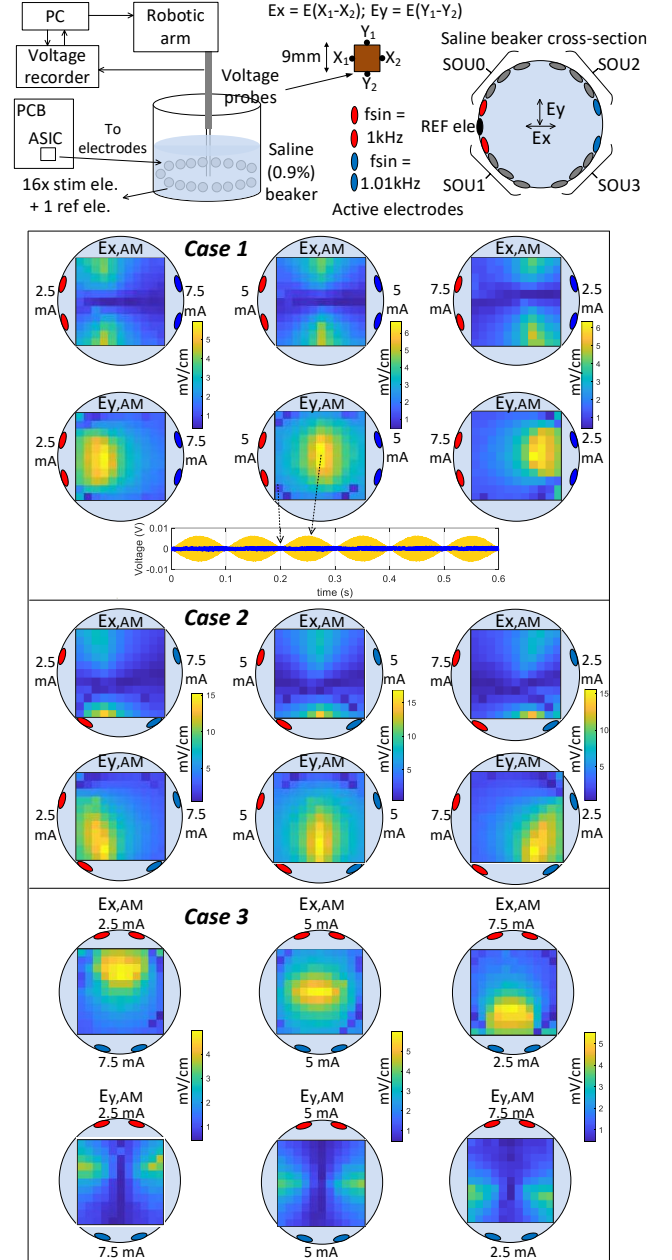


Fig. 14. TIS demonstration of flexible stimulation focal region steering with the proposed ASIC in saline.

frequency of 10Hz is extracted in postprocessing to evaluate the spatial focusing ability of TIS. As shown in Fig. 14, by using different stimulation current amplitudes and electrodes at different locations, the envelope amplitude stimulation focal region can be steered precisely within the area of the 16 electrodes.

It should be noted that, to create detailed TIS focal region steering heatmaps, the size of the phantom and electrodes used in this demonstration is significantly larger than what would be used in real-life applications. Since TIS fundamentally relies on electric fields to shape the desired stimulation focal region [9], the stimulation focal region remains consistent when both the phantom and electrode sizes are scaled together. Therefore, this

Table I. Performance comparison with the state-of-the-art.

		This work	[11] ISSCC23 ^d	[19] ESSCIRC23 ^d	[25] TBCAS24 ^d	[22] JSSC18	[18] JSSC22 ^d	[30] CICC21
System	Technology	130nm BCD	65nm	55nm HV CMOS	180nm HV CMOS	350nm HV	180nm HV	180nm HV
	Application	PNS	PNS	Brain stim	Brain stim	PNS	Brain stim	Brain stim
	Selective stim.	Yes	Yes	No	No	No	No	Yes
	Stim. method/ waveform	TIS/ Sinusoidal	TIS/ Sinusoidal	Multipolar/ Bi-phasic	Multipolar/ Bi-phasic	Unipolar/ Bi-phasic	Multipolar/ Bi-phasic	Vector/ Bi-phasic
	# Output / # SOU	16 / 4	64 / 64	128 / 128	4 / 4	1 / 1	16 / 4	46 / 46
Stimulator & I-DAC	Area ^e / output [mm ²]	0.29	<0.04 ^c	0.024	3.3 ^c	2.04 ^c	1.15 ^c	0.12
	Compliance voltage [V]	10	<3.3	<6	40	22	40	26
	Compliance monitor	Yes	No	No	No	No	No	No
	Max amplitude [mA]	10.22	2	0.127	14	5.12	12.75	2
	I-DAC bit	9b	8b	7b	8b	7b	8b	7b
	Area / output [mm ²]	0.13	<0.04 ^c	0.024	2.07 ^c	1.24 ^c	0.68 ^c	0.12
	Max fsin [Hz]	42k	>2k	--	--	--	--	--
Charge balancing	SFDR [dB]	>57	--	--	--	--	--	--
	TIS compatibility	Yes	--	No	No	No	No	No
	CB power / SOU [μW]	60 - 114 ^a	--	--	--	56	--	--
	Area / output [mm ²]	0.15	--	--	1.06 ^c	0.8 ^c	0.3 ^c	--
	CB method	Active	--	Passive	Active	Active	Active	Passive
ETI sensing	CB precision	$I_{mis,rms} < 10nA$, $V_{os} < \pm 50mV$	--	--	$V_E < \pm 2mV / \pm 7.5mV$	$V_E \leq \pm 20mV$	$V_E < \pm 2mV$	--
	Buffer power / SOU [W]	54μ	--	--	--	--	--	--
	Range [Ω]	0 - 1.8k ^b	--	--	--	--	--	--
	Inaccuracy [Ω]	± 2 ^b	--	--	--	--	--	--

^a For $\pm 3\sigma$ mismatch coverage @ 2.56-10.22mA stim. amplitude, excluding HV compensation current output, which is part of stim. current^b @ stimulation amplitude of 2.56mA^c Estimated from the die photo^d Only the stimulation circuitry for comparison^e Including all analog circuitry

scaled setup can still effectively verify the TIS focal region and its steering capability using our ASIC. However, the measured absolute values of the electrical fields in this scaled setup will be much smaller than in actual applications, as the strength of the electric field scales proportionally with $1/d^2$ on a 2D surface, where d is the diameter of the phantom, assuming the stimulation electrodes are scaled and kept at the same relative locations. For example, in case 1 shown in Fig. 14, the measured peak LF electrical field envelope amplitude (EFEA) is slightly above 6mV/cm. Extrapolating this to a high-density cuff electrode placed on a 5 mm diameter peripheral nerve, with the same stimulation parameters, the peak LF EFEA would be approximately 2.4V/cm. However, it is important to note that in real-life scenarios, the LF EFEA also depends on other factors, such as the nerve tissue conductivity (which could differ significantly from the 0.9% NaCl solution used here), stimulation current amplitude, and the relative location of the active electrodes. As a result, stimulation current amplitudes and electrode positions must be carefully adjusted for optimal stimulation efficacy, which remains an important area for further research in TIS [31-32].

D. Benchmark

The performance comparison of the proposed ASIC with prior arts is shown in Table I. This work is the only HV TIS ASIC with a wide programmability of stimulation current and carrier frequency. Thanks to the area-efficient architecture and circuit implementation, an active area of 0.29mm²/output is achieved despite the high stimulation current and HV compliance. An active CB method is proposed for TIS for the first time, achieving a sub-10nA mismatch. ETI sensing during TIS is realized with minimal design complexity.

V. CONCLUSION

A 16-output HV ASIC designed for spatially selective neuromodulation through TIS is presented in this work. Fabricated using 130nm BCD technology, the ASIC achieves an area-efficient design. The proposed active CB effectively

compensates for mismatch currents during stimulation across a wide range of ETIs. Furthermore, the ETI sensing method demonstrates sufficient accuracy with minimal power and area overhead. Extensive saline experiments validate the ASIC's capability to steer the stimulation focal region based on TIS. Integrating this ASIC in a neuromodulation system would enable novel and more effective electrical neural stimulation therapies, having fewer undesirable side effects, as stimulation can be steered towards the confined neural region of interest.

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Haoming Xin received the B.Sc. degree from Xi'an Jiaotong University, Xi'an, China, in 2013, and the M.Sc. and Ph.D. degrees from Eindhoven University of Technology (TU/e), Eindhoven, The Netherlands, in 2015 and 2020, respectively. His Ph.D. research focused on ultra-low-power, energy-efficient sensor interfaces for energy-constrained

IoT applications. From 2019 to 2020, he was a Postdoctoral Researcher with the Integrated Circuit Group at TU/e. Since 2021, he has been with imec, Eindhoven, where he is currently a Senior Researcher focusing on neural interface ASIC development. His research interests include low-power, energy-efficient analog and mixed-signal circuits and systems for biomedical and IoT applications. He is a recipient of the CICC 2024 Best Paper Award.



Meiyi Zhou received the B.Eng. degree in electronic information engineering from Zhejiang University, Hangzhou, China, in 2016. She completed the M.Sc. in electrical engineering with Cum Laude at Eindhoven University of Technology (TU/e), Eindhoven, the Netherlands, in 2017, sponsored by the ALSP full scholarship. From 2017 to 2021, she

pursued her Ph.D. at TU/e, specializing in integrated circuit (IC) design for medical ultrasound imaging systems, in close collaboration with Philips Research. In 2022, she received her Ph.D. and joined the IC-design group at IMEC. She is currently a senior Researcher at IMEC, focusing on implantable neural interface ASICs. Her research interests include the development of efficient and reliable biomedical circuits, systems, and devices for healthcare applications.



Roland van Wegberg received his M.Sc. degree in electrical engineering in 1997 from the Eindhoven University of Technology (TU/e), The Netherlands. He has worked for IMEC, Philips Semiconductors, NXP and Synopsys Inc. as an analog IC design engineer. From 2013, he re-joined IMEC in the analog IC design team as a Senior Analog R&D

Design Engineer. His research interests include high performance data converters, ultra-low power analog and mixed-signal circuits for biomedical applications and neural interfaces. He has authored or coauthored several patents and publications.



Peter Vis received the B. Eng. degree in electrical engineering from the AVANS University of Applied Sciences, 's-Hertogenbosch, The Netherlands, in 2005. He joined NXP semiconductors in 2008 and later Synopsys Inc. where he worked on the development of interface IP. In 2018, he joined IMEC- the Netherlands,

Eindhoven, The Netherlands, as a Digital R&D engineer. His research interests include the design of low-power analog mixed signal designs for biomedical and RF applications.



Konstantinos Petkos is currently a Researcher at IMEC working on portable/wearable and implantable medical electronics. Before joining IMEC, he was a Postdoctoral Research Associate at the Department of Bioengineering of Imperial College London. He received the 5-year diploma in Electrical and Computer

Engineering from Aristotle University of Thessaloniki, and the MRes in Bioengineering, MRes in Neurotechnology & Ph.D. in Bioinstrumentation design degrees from Imperial College London. He was the recipient of a competitive 4-year Ph.D. studentship on neuroelectroceuticals sponsored by Medtronic, under the aegis of the Centre for Doctoral Training (CDT) in Neurotechnology. His research focuses on: i) advanced analog signal processing techniques for recording high-quality neural signals in the presence of strong artefacts stemming from various types of electrical stimulation, and ii) the research and development of portable/wearable/miniatuised medical devices that can be used as diagnostic tools in the clinic.



Shrishail Patki received his bachelor degree in Instrumentation Engineering from University of Mumbai and MS degree from University of Miami in Biomedical Engineering. He spent one and half years working as an assistant electronics engineer at Bio-Signal Group Corp where he worked on EEG system development. In 2010, he joined imec as a researcher

working on design and development of different generations of miniaturized, low noise and low power wearable systems. He is currently R&D Engineer, focusing on system development for implantable systems.



Nicolò Rossetti received the B. Eng. degree Cum Laude in biomedical engineering from Politecnico di Milano (Milan, Italy) in 2016 and the M.Sc. double degree Cum Laude in biomedical engineering from Polytechnique de Montréal (Montréal, Canada) and Politecnico di Milano (Milan, Italy) in

2019. In 2019 he joined Thimus Inc. (Brescia, Italy) as R&D engineer where he worked on the development of wearable eeg systems and on the study of brain signals for applications in cognitive neuroscience. Since 2021 he has been a researcher at IMEC (Eindhoven, Netherlands), where he has been working on both non-invasive and implantable solutions for neuromodulation.



Mark Fichman is a technical lead with over 20 years of experience in active implantable medical devices (AIMDs) in industrial environments. As Principal Technical Lead at imec, he drives innovation in bioelectronic medicine and neuromodulation, with expertise in system architecture, strategic road mapping, and regulatory compliance. Mark has guided complex implant projects to clinical trials, mentored multidisciplinary teams, and provided technical leadership to bridge research and clinical applications. Recognized for his vision and technical expertise, he remains dedicated to advancing AIMD innovation through collaboration and strategic guidance.



Vojkan Mihajlović is a principal member of technical staff at Stichting imec Nederland. He received his Ph.D. in computer science from the University of Twente in 2002. He has more than 15 years of experience in the domain of neuromodulation technologies and applications. He worked within Philips Research (2008-2012), where he focused on algorithms and methods for improving signal integrity in EEG recordings. He has been leading the Wearable Brain Monitoring work package at Stichting imec Nederland (2016-2021), after which he fused on the creation of a roadmap on the selective peripheral nerve stimulation and establishing a research team to explore and develop required technology solutions. He is currently active as a technical lead for neuro applications and coordinates health innovation activities within Stichting imec Nederland. He has authored more than 80 peer reviewed publications and holds 15 patents.



Carolina Mora Lopez (Senior Member, IEEE) received her Ph.D. degree in Electrical Engineering in 2012 from the KU Leuven, Belgium, in collaboration with imec, Belgium. From 2012 to 2018, she worked at imec as a researcher and analog designer focused on interfaces for neural-sensing applications. She is currently the Scientific Director and team leader of the circuits for neural interfaces team in imec. Her research interests include analog and mixed-signal circuit design for sensor, bioelectronics and neural interfaces. Carolina is a senior IEEE member and serves on the technical program committee of the ISSCC and ESSCIRC conferences.



Geert Langereis received his PhD in lab-on-a-chip technologies from the University of Twente, the Netherlands. He worked at the industrial research laboratories of Philips and NXP on MEMS silicon technology and photonics in combination with data science. After that, he conducted a research and education line at the University of Applied Sciences in Eindhoven for smart sensors and associated data science for human behavior and physiology.

He is currently a Principal Member of Technical Staff in the Health Applications department of imec in Eindhoven on technologies for artificial organs and bioelectronic medicine. He holds a part-time position at the dept. of Electrical Engineering of the Technical University of Eindhoven.



Mario Konijnenburg (Member IEEE) received the M.S. degree in electrical engineering from Delft University of Technology in The Netherlands in 1993. A Ph.D. degree was received from Delft University of Technology in 1999 on Automatic Test Pattern Generation for Sequential Circuits. He joined Philips Research / NXP Semiconductors and worked on methodologies to improve testability of designs. Currently he is R&D manager of the IC-design group at imec in Eindhoven, The Netherlands, targeting chip research for bio-medical applications covering SoC design, sensors, stimulators, power and (RF) communication.



Nick Van Helleputte (Member, IEEE) received the MS degree in electrical engineering in 2004 from the Katholieke Universiteit Leuven, Belgium. He received his Ph.D. degree from the same institute in 2009 (MICAS research group). His PhD research focused on low-power ultra-wide-band analog front-end receivers for ranging applications. He joined imec in 2009 as an Analog R&D Design Engineer. He is currently Scientific director of Circuits & Systems for health. Nick served on the technical program committee of VLSI circuits symposium and ISSCC.