# A High TCMRR, Inherently Charge Balanced Bidirectional Front-End for Multichannel Closed-Loop Neuromodulation

José Luis Valtierra, Rafaella Fiorelli, Norberto Pérez-Prieto, Manuel Delgado-Restituto and Ángel Rodríguez-Vázquez

Institute of Microelectronics of Seville and University of Seville Avda. Americo Vespucio s/n, 41092-Seville,SPAIN Email: {valtierra,fiorelli,norberto,mandel,angel}@imse-cnm.csic.es

Abstract—This paper describes a multichannel bidirectional front-end for implantable closed-loop neuromodulation. Stimulation artefacts are reduced by way of a 4-channel H-bridge current source sharing stimulator front-end that minimizes residual charge drops in the electrodes via topology-inherent charge balancing. A 4-channel chopper front-end is capable of multichannel recording in the presence of artefacts as a result of its high total common-mode rejection ratio (TCMRR) that accounts for CMRR degradation due to electrode mismatch. Experimental verification of a prototype fabricated in a standard 180 nm process shows a stimulator front-end with 0.059% charge balance and 0.275 nA DC current error. The recording front-end consumes 3.24  $\mu$ W, tolerates common-mode interference up to 1 Vpp and shows a TCMRR > 66 dB for 500 mVpp inputs.

Index Terms—closed-loop neuromodulation, TCMRR, charge balance, stimulation artefacts.

# I. INTRODUCTION

Implantable closed-loop neuromodulation has been proposed as a form of treatment for Parkinson's and epilepsy, among other neural disorders [1]. The idea is that the dynamics of the brain inform the state of the stimulators, thus establishing closed-loop control of region-specific brain activity. In a practical implementation, this scheme presents an intrinsic self-interference caused by stimulation artefacts that saturate the recording front-end of the neuromodulator. Stimulation artefacts are a product of residual charge accumulation in the electrodes which is in turn caused by mismatch in the stimulation cathodic and anodic phases. Artefact magnitude is proportional to the stimulation current amplitude and proximity between stimulation and recording electrodes. A commonmode artefact can reach up to hundreds of milivolts [2] and also exhibits a differential component as well as a ratcheting effect [3]. Fig. 1 shows a typical power spectrum for artefacts that reveals the presence of harmonics and an increase in magnitude due to time-dependent residual charge accumulation. An accurate charge-balancing scheme is therefore fundamental for the chronic operation of closed-loop neuromodulators. Traditional multichannel recording front-ends are specially susceptible to artefacts. A significant degradation in commonmode rejection presents itself under the conventional electrode

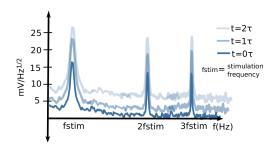


Fig. 1. Frequency domain (Log scale) view of artefacts and their inputreferred magnitude increase due to residual charge accumulation over time.

reference sharing configuration due to impedance mismatch between reference and signal inputs. This situation gives rise to the concept of total common-mode rejection ratio (TCMRR) that accounts for impedance mismatch in a multichannel fontend as follows [4]:

$$TCMRR = \left(\frac{1}{ICMRR} + \left(\frac{1 + 2(|Z_{IN}/Z_e| + N\epsilon)}{2(N\epsilon - 1)}\right)^{-1}\right)^{-1},$$
(1)

where N is the number of channels sharing a single reference electrode,  $\epsilon$  is the mismatch factor between reference and signal electrodes ( $\epsilon$ =1 when both electrodes are matched),  $|Z_{IN}/Z_e|$  is the impedance ratio between input and electrode impedances and ICMRR is the intrinsic common-mode rejection ratio (i.e. the CMRR of a single channel). In order to maintain a sufficiently high TCMRR,  $|Z_{IN}/Z_e|$  should be high, ICMRR high and N should be modest. This phenomenon was originally described by [4] with common-mode interference from capacitive coupling of electrical supply voltages in mind (as high as 100 mVpp). Evidently, it acquires greater importance in the presence of large common-mode stimulation artefacts (as high as 1 Vpp). A model for a single recording channel is shown in Fig. 2. Previously reported closed-loop neuromodulators deal with artefacts by input blanking [5], pole-shifting [6] or by placing stimulation and recording electrodes far from each other [7] [8]. These strategies somewhat

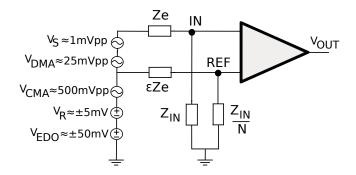


Fig. 2. Model for a single channel in a multichannel recording interface in the presence of self-interference (extended from [4]).  $V_S$  is the neural signal of interest,  $V_{DMA}$  is the differential-mode artefact,  $V_{CMA}$  is the common-mode artefact,  $V_R$  is the ratcheting effect and  $V_{EDO}$  is the electrode DC offset. Amplitudes for  $V_{DMA}$ ,  $V_{CMA}$ ,  $V_R$  will increase over time if the stimulation pulses are not balanced.

defeat the purpose of closed-loop neuromodulation since they imply either loss of information (during and right after stimulation) or a non-local control loop. The works reported by [9]-[11] do manage same electrode array closed-loop operation, however, they make no account for TCMRR meaning that simultaneous multichannel recording during stimulation might be degraded. Moreover, only [9] considers stimulator charge balance even though the link between residual charge in the electrodes (i.e. stimulator charge imbalance) and artefact generation has been well established. In this work we present a bidirectional front-end for multichannel closed-loop neuromodulation. The role in artefact generation by the charge balance scheme of the stimulators is acknowledged from the very beginning: a 4-channel stimulator front-end based on an H-bridge topology that shares the same current source for both phases of biphasic stimulation is proposed. This topology inherently balances the stimulation current in both phases. Since artefacts are unavoidable, the aim is to produce a stimulation system that generates non-saturating artefacts. A 4-channel low-noise low-power front-end based on chopper amplifiers comprises the recording side of the front-end. In order to tolerate artefacts, the chopper amplifiers are designed for large ac common-mode interferer range, high differential input range and large DC offset rejection. To maintain high TCMRR, the front-end includes input impedance boosting and high ICMRR for large common-mode inputs.

# II. BIDIRECTIONAL FRONT-END FOR MULTICHANNEL CLOSED-LOOP NEUROMODULATION

Figure 3 shows the block diagram of the proposed 8-channel bidirectional front-end. The circuit is comprised by a 4-channel stimulator front-end and a 4-channel recording front-end. The system is powered by  $V_{DDH}$ =3.3 V (stimulator) and  $V_{DDL}$ =1.8 V (recording) supply.

### A. H-Bridge Inherently Charge-Balanced Stimulator

In a biphasic stimulation scheme, charge imbalance comes from unavoidable mismatch between the stimulator current sources. Additional circuitry to compensate for residual charge

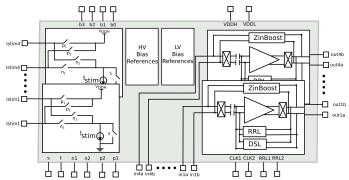


Fig. 3. Proposed bidirectional front-end.

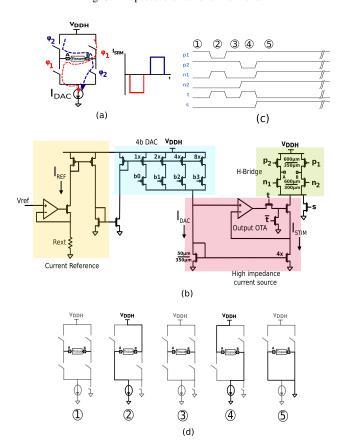


Fig. 4. (a) H-bridge (b) Proposed Stimulator (c) Time diagram (d) Stimulator Operation.

in the electrodes is necessary since good layout matching can only do so much. In order to avoid mismatch between current sources, we implement an H-bridge (Fig. 4(a)) operation to produce a biphasic pulse from a single current source, thus obviating the need for a charge compensation scheme. The proposed neural stimulator is shown in Fig. 4(b). An additional transistor is added to the H-bridge to short the output electrodes to ground after each biphasic pulse (passive discharge). The output H-bridge sinks the stimulation current to a high-output impedance current driver that also accurately copies  $I_{DAC}$ , which is set by a 4-b DAC. The timing diagram of the stimulator is shown in Fig. 4(c). The stimulator is turned

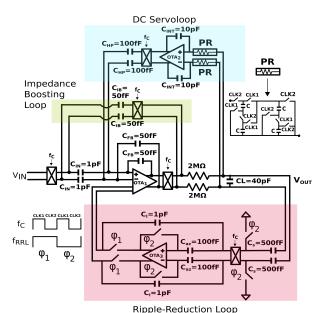
off in State 1. State 2 sees the current flow from A to B, thus creating the cathodic pulse. State 3 defines an interphase delay, usually much shorter than the width of each pulse. The anodic pulse is created in State 4 where the current flows from B to A. State 5 shorts the outputs to discharge the electrodes after each biphasic pulse, thus providing a DC current error that is safe for both the neural tissue and microelectrodes. Fig. 4(d) depicts the operation of the stimulator.

# B. Chopper Amplifier for High TCMRR Recording

The proposed chopper amplifier is shown in Fig. 4. Chopper amplifiers typically exhibit high ICMRR [12]. However, the input impedance defined as  $Z_{IN} = \frac{1}{2f_C C_{IN}}$ , jeopardizes its direct implementation in a high TCMRR scheme. This can be compensated by setting the chopping clock  $(f_C)$  at 2.5 kHz to minimize its impact on input impedance. An input capacitor  $C_{IN}$ = 1 pF is chosen, which translates as a theoretical  $Z_{IN}$ = 200 M $\Omega$ , nevertheless, a positive feedback impedance boosting loop has been included to obtain higher input impedance. The gain is set to 20 V/V by way of  $C_{IN}/C_{FB}$  in order to amplify both  $V_S$  and  $V_{DMA}$  without saturating the frontend. A current mirror amplifier with partial positive feedback has been chosen for  $OTA_1$  [13]. This OTA features a rail-torail output and high enough open-loop gain so that a second OTA in the forward path can be avoided. The bandwidth is set by a passive low-pass filter. A DC servoloop introduces a high-pass frequency corner in the front-end by feedbacking an integrator with a very large time constant. Its specific role is then, to reject  $V_R$  and  $V_{EDO}$ . Chopper modulation introduces an output ripple whose magnitude is dependent on the offset of the first stage. This was attenuated by sensing it in the output with a switched-capacitor integrator that produces a proportional voltage which in turn generates a compensating current by means of an auxiliary differential pair in  $OTA_1$ [12].

# III. MEASUREMENT RESULTS

Fig. 6 shows a microphotograph of the fabricated prototype. Fig. 7(a) shows the stimulation waveform for a R=5 k $\Omega$  C=100 nF load. The maximum current of a single stimulator (Fig. 7(b)) is 300  $\mu$ A, for a 5 k $\Omega$  load. More than one stimulator can be combined in a single load for higher current capability. Charge balance was characterized by stimulating 10 biphasic pulses of 600  $\mu$ s. The residual voltage was measured right after the last pulse. The final shorting phase was removed for this test. Fig. 7(c) shows the residual voltage for each stimulator. In all cases, a residual voltage under the water window is achieved. Fig. 7(d) shows the residual charge for each stimulator. The worst case residual charge is about 106 pC. The worst case charge mismatch is (residual/injected) x 100, which is 0.059%. The recording front-end shows a 600 Hz bandwidth and draws 3.24 µW/channel from a 1.8 V supply. Fig. 8(a) shows the measured input-referred noise, which is about 250 nV/rtHz. Differential input range, measured in terms of total harmonic distortion (Fig. 8(b)) is 2.85% for a 120 mVpp input. Fig. 8(c) shows a 5% gain drop for 1 Vpp



Ripple-Reduction Loop

Fig. 5. Proposed chopper amplifier.

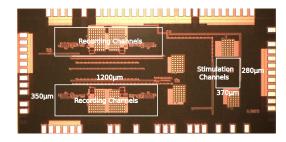


Fig. 6. Die microphotograph.

common-mode inputs while ICMRR measured for 500 mVpp inputs is kept above 80 dB in the amplifier bandwidth. Input impedance and ICMRR measured for each channel for 500 mVpp inputs at 200 Hz is shown in Table I. The calculated TCMRR at 200 Hz for each channel and electrode impedance =  $100 \text{ k}\Omega$  is > 66 dB. The specifications are summarized and compared with previous art in Table II.

# IV. CONCLUSIONS

This work presented an 8-channel bidirectional front-end for closed-loop neuromodulators. Previously reported front-ends generally ignore large ac common-mode interference, CMRR degradation due to electrode mismatch and charge balance in

 $\begin{tabular}{ll} TABLE\ I \\ TOTAL\ COMMON-MODE\ REJECTION\ RATIO\ AT\ 200\ Hz. \end{tabular}$ 

Recording channel	$Z_{IN}$ (M $\Omega$ )	ICMRR (dB)	TCMRR (dB)
A	826	81	66.9
В	816	83	67.1
С	795	80	66.4
D	884	88	68.4

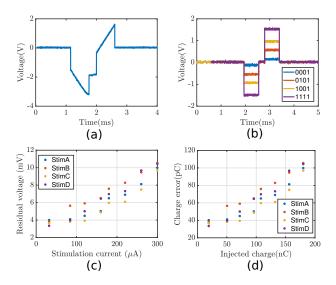


Fig. 7. Stimulator front-end. (a) Waveform for an R+C load (b) Waveform for an R load (c) Residual voltage after 10 biphasic pulses (d) Charge error.

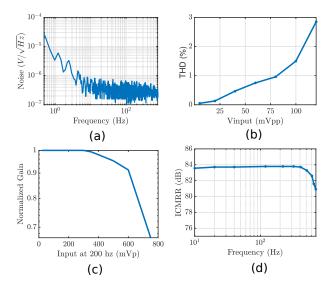


Fig. 8. Recording front-end. (a) Noise (b) Total harmonic distortion (c) Input ac common-mode interferer range (d) Intrinsic common-mode rejection ratio.

TABLE II
COMPARISON WITH CLOSED-LOOP NEUROMODULATORS

	[10]	[11]	[9]	This work
Process / Year	40 nm, 180 HV / 2018	65 nm / 2018	180 nm HV / 2017	180 nm / 2019
Channels Recording / Stimulation	64 / 8	8 / 2	64 / 4	4 / 4
Power (Recording) (µW)	8.2	2.7	8	3.24
Bandwidth (Hz)	250	8.3 k	500	600
Noise	$2.2~\mu Vrms$	8.2 μVrms	71 nV/rtHz	250 nV/rtHz
THD	-81 dB at 100 mVpp	-77 dB at 200 mVpp	0.7% at 100mVpp	2.85% at 120 mVpp
AC CMI range	-	-	-	1 Vpp
ICMRR (dB)	-	-	-	> 80
TCMRR (dB)	-	-	-	> 66
Charge Balance	-	-	<1%	0.059%
DC current error	-	-	-	0.275 nA

relation to artefact generation. This work is therefore, the first one to account for these specifications concurrently. Future work will deploy a complete characterization of the front-end.

#### **ACKNOWLEDGEMENTS**

This work has been supported by the Spanish Ministry of Economy and Competitiveness under grant TEC2016-80923-P, the ONR under grant N000141110312 and the FEDER Program. The first author was supported by CONACyT-México.

#### REFERENCES

- [1] M. Parastarfeizabadi and A. Z. Kouzani, "Advances in closed-loop deep brain stimulation devices," *Journal of neuroengineering and rehabilitation*, vol. 14, no. 1, p. 79, 2017.
- [2] P. Single and J. Scott, "Cause of Pulse Artefacts Inherent to the Electrodes of Neuromodulation Implants," *IEEE Transactions on Neural* Systems and Rehabilitation Engineering, vol. 26, no. 10, pp. 2078–2083, 2018
- [3] J. Lim, P. T. Wang, A. K. Bidhendi, O. M. Arasteh, S. J. Shaw, M. Armacost, H. Gong, C. Y. Liu, P. Heydari, A. H. Do et al., "Characterization of Stimulation Artifact Behavior in Simultaneous Electrocorticography Grid Stimulation and Recording," in 2018 40th Annual International Conference of the IEEE Engineering in Medicine and Biology Society (EMBC). IEEE, 2018, pp. 4748–4751.
- [4] K. A. Ng and Y. P. Xu, "A low-power, high CMRR neural amplifier system employing CMOS inverter-based OTAs with CMFB through supply rails," *IEEE Journal of Solid-State Circuits*, vol. 51, no. 3, pp. 724–737, 2016.
- [5] C.-H. Cheng, P.-Y. Tsai, T.-Y. Yang, W.-H. Cheng, T.-Y. Yen, Z. Luo, X.-H. Qian, Z.-X. Chen, T.-H. Lin, W.-H. Chen et al., "A Fully Integrated 16-Channel Closed-Loop Neural-Prosthetic CMOS SoC With Wireless Power and Bidirectional Data Telemetry for Real-Time Efficient Human Epileptic Seizure Control," *IEEE Journal of Solid-State Circuits*, no. 99, pp. 1–13, 2018.
- [6] X. Liu, M. Zhang, A. G. Richardson, T. H. Lucas, and J. Van der Spiegel, "Design of a closed-loop, bidirectional brain machine interface system with energy efficient neural feature extraction and PID control," *IEEE Transactions on Biomedical Circuits and Systems*, vol. 11, no. 4, pp. 729–742, 2017.
- [7] A. E. Mendrela, J. Cho, J. A. Fredenburg, V. Nagaraj, T. I. Netoff, M. P. Flynn, and E. Yoon, "A Bidirectional Neural Interface Circuit With Active Stimulation Artifact Cancellation and Cross-Channel Common-Mode Noise Suppression," *IEEE Journal of Solid-State Circuits*, vol. 51, no. 4, pp. 955–965, 2016.
- [8] E. Greenwald, E. So, Q. Wang, M. Mollazadeh, C. Maier, R. Etienne-Cummings, G. Cauwenberghs, and N. Thakor, "A Bidirectional Neural Interface IC with Chopper Stabilized BioADC Array and Charge Balanced Stimulator," *IEEE Transactions on Biomedical Circuits and Systems*, vol. 10, no. 5, pp. 990–1002, 2016.
- [9] B. C. Johnson, S. Gambini, I. Izyumin, A. Moin, A. Zhou, G. Alexandrov, S. R. Santacruz, J. M. Rabaey, J. M. Carmena, and R. Muller, "An implantable 700μW 64-channel neuromodulation IC for simultaneous recording and stimulation with rapid artifact recovery," in 2017 Symposium on VLSI Circuits. IEEE, 2017, pp. C48–C49.
- [10] D. Rozgić, V. Hokhikyan, W. Jiang, I. Akita, S. Basir-Kazeruni, H. Chandrakumar, and D. Marković, "A 0.338 cm<sup>3</sup>, Artifact-Free, 64-Contact Neuromodulation Platform for Simultaneous Stimulation and Sensing," *IEEE Transactions on Biomedical Circuits and Systems*, vol. 13, no. 1, pp. 38–55, 2018.
- [11] S. Jung, P. Kwon, D. Piech, M. Maharbiz, J. Rabaey, and E. Alon, "A 2.7-µW Neuromodulation AFE With 200 mVpp Differential-Mode Stimulus Artifact Canceler Including On-Chip LMS Adaptation," *IEEE Solid-State Circuits Letters*, vol. 1, no. 10, pp. 194–197, 2018.
- [12] Q. Fan, F. Sebastiano, J. H. Huijsing, and K. A. Makinwa, "A  $1.8\mu$  W 60 nV  $/\sqrt{Hz}$  Capacitively-Coupled Chopper Instrumentation Amplifier in 65 nm CMOS for Wireless Sensor Nodes," *IEEE Journal of Solid-State Circuits*, vol. 46, no. 7, pp. 1534–1543, 2011.
- [13] R. Wang and R. Harjani, "Partial positive feedback for gain enhancement of low-power CMOS OTAs," in Low-Voltage Low-Power Analog Integrated Circuits. Springer, 1995, pp. 21–35.