# A Fully Differential CMOS Potentiostat

Meisam Honarvar Nazari Department of Electrical Engineering California Institute of Technology Pasadena, CA 91125, USA Email: meisam@caltech.edu

Abstract—A CMOS potentiostat for chemical sensing in a noisy environment is presented. The potentiostat measures bidirectional electrochemical redox currents proportional to the concentration of a chemical down to pico-ampere range. The fully differential architecture with differential recording electrodes suppresses the common mode interference. A  $200\mu m \times 200\mu m$  prototype was fabricated in a standard  $0.35\mu m$  standard CMOS technology and yields a 70dB dynamic range. The in-channel analog-to-digital converter (ADC) performs 16-bit current-to-frequency quantization. The integrated potentiostat functionality is validated in electrical and electrochemical experiments.

## I. INTRODUCTION

Electrochemical amperometric methods are widely employed in chemical and biochemical sensing. In these methods a current corresponding to the concentration of a chemical is generated as a result of a voltage applied to the chemical solution. A potentiostat is a circuit that records this redox current. Constant-potential amperometry and fast-scan cyclic voltammetry (FSCV) are two commonly employed amperometric methods. In constant-potential amperometry a fixed potential, known as the redox potential, is applied between two conductive electrodes, the reference electrode (RE) and the working electrode (WE). This method offers high temporal resolution at the cost of poor selectivity [1]. In FSCV a cyclic potential is applied between recording electrodes. It offers high selectivity at the cost of lower temporal resolution. The high selectivity in this method comes as a result of the cyclic redox potential [2] that generates a unique cyclic voltammogram corresponding to each chemical.

The cyclic potential employed in FSCV generates a large periodic background current due to charging and discharging of the double layer capacitance at the electrode-electrolyte interface [1], [2]. Such a current is typically much larger than the redox current. Any other electrical signals present in the solution can also cause large unwanted interference signals. The proposed differential working electrode configuration followed by a fully differential current acquisition circuit, as shown in Fig. 1, removes all of the offset and interference currents. It also rejects any common-mode potentiostat circuit noise. In addition to suppressing common-mode noise, a differential architecture provides a voltage range twice that of a singleended approach. In particular, the factor of two improvement in the voltage range is extremely beneficial when supply voltage shrinks due to CMOS technology scaling.

Roman Genov Department of Electrical and Computer Engineering University of Toronto Toronto, ON M5S 3G4, Canada Email: roman@eecg.toronto.edu



Fig. 1. Differential electrode amperometric sensing.



Fig. 2. Single-ended version of the potentiostat channel architecture.

A number of single-ended integrated potentiostats have been reported [3], [4], [5], [6]. As discussed above, the performance of a single-ended potentiostat is very likely to degrade in presence of common-mode interference. Several differential integrated potentiostats for multi-channel recording in noisy environments have also been reported. An array of 50 potentiostats which employ a pseudo-differential architecture is presented in [7]. Each cell requires an external load to perform the current-to-voltage conversion. The pseudodifferential architecture requires a larger area compared to that of a fully differential architecture. A  $24 \times 16400\mu$ m-pitch array of integrated potentiostats for DNA detection is presented in [8] in which a front end circuit converts the charge generated at differential electrodes to an analog voltage.

In this paper we present a potentiostat that detects chemicals directly on on-chip recording electrodes and provides a digital output. The channel sized  $200\mu$ m $\times 200\mu$ m can be easily tiled for a large sensory array area. The differential electrode configuration and signal acquisition circuit provide high common-



Fig. 3. Circuit diagram of the fully differential potentiostat.

mode interference rejection. In order to cover a wide dynamic range of input currents a current-to-frequency ADC is utilized. The proposed potentiostat provides over 70dB dynamic range of a differential current-mode input.

## II. CHANNEL VLSI ARCHITECTURE

The top level architecture of the potentiostat is shown in Fig. 2. The input redox current is first converted to a voltage which is subsequently quantized by the ADC.

Amperometry and cyclic voltammetry detection techniques rely on the accuracy of the potential difference between the working and reference electrodes. The negative capacitive feedback around an operational transconductance amplifier (OTA) and the input common-mode feedback circuit set a fixed voltage at the two working electrodes, as detailed in the next section. An external voltage source drives the reference electrode to set the redox potential (shown connected to ground in Fig. 1 for simplicity).

The ADC utilizes the current-to-frequency quantization architecture. During the conversion period the redox current is integrated across a capacitor,  $C_{INT}$ . As soon as the capacitor voltage reaches a known value,  $V_{REF}$ , a reset pulse is generated by the comparator in order to discharge the integrating capacitor, as shown in Fig. 2. The input current is related to the number of pulses generated as

$$I_{IN} = \frac{NC_{INT}V_{REF}}{T_{conv}},\tag{1}$$

where  $I_{IN}$  is the input current,  $T_{conv}$  is the conversion time, N is the number of comparator pulses,  $C_{INT}$  is the integrating capacitance, and  $V_{REF}$  is the comparator reference voltage.

## **III. VLSI CIRCUIT IMPLEMENTATION**

An array of 96 integrated potentiostats was designed and implemented in a  $0.35\mu$ m double-poly four-metal standard CMOS process. The redox current generated on on-chip working electrodes is accumulated on a capacitor and converted to the digital domain by a current-to-frequency ADC, as detailed in Fig. 3.



Fig. 4. Input common-mode feedback amplifier.



Fig. 5. Fully differential folded-cascode OTA along with the switched capacitor output CMFB circuit.

#### A. Current-to-Voltage Converter

A capacitive transimpedance amplifier (TIA) is employed to convert the input redox current to a voltage. The capacitive TIA has an averaging behavior and acts as a low-pass filter to remove the high-frequency noise. A common-mode regulation loop is necessary to prevent the high-impedance input node voltages from drifting to the positive or negative voltage levels. Fig. 3 shows an amplifier configured in a negative feedback to ensure a fixed input common-mode voltage. The amplifier circuit diagram is depicted in Fig. 4. The input differential pair transistors,  $M_{2,3}$ , sense the common-mode voltage. The common-mode voltage is compared with a desired value,  $V_{CM}$ , by transistors  $M_{1-3}$  in order to set the input commonmode voltage of the TIA.

Fig. 5 depicts the schematic of the fully differential OTA with the output common-mode feedback circuit. A simple switched capacitor circuit controls the output common-mode voltage. The RST signal is high once in a conversion period to refresh the voltage of the capacitors  $C_{CM}$ . The gain-enhanced folded-cascode topology improves the gain of the OTA and minimizes errors due to its finite gain. The closed-loop gain



Fig. 6. Dependency of the OTA gain on the differential output swing.

TABLE I OTA Simulated Electrical Characteristics

DC gain	108dB
Unity Gain Frequency (0.5pF load)	20MHz
Slew Rate	13V/µs
Differential output Voltage Swing	$2V_{pp}$
Total Bias Current	$16\mu A$
Supply Voltage	3.3V

error due to the finite gain of the OTA can be expressed as [?]

$$\Delta G = \left[1 - \frac{1}{1 + \frac{C_{IN}}{C_{INT}A}}\right] \times 100,\tag{2}$$

which for  $\frac{C_{IN}}{C_{INT}A} \ll 1$  reduces to

$$\Delta G \approx \frac{C_{IN}}{C_{INT}A} \times 100, \tag{3}$$

where A is the finite gain of the OTA,  $C_{INT}$  is the integrating capacitance and  $C_{IN}$  is the sum of the capacitances at the input of the OTA. The simulated characteristics of the fully differential OTA are presented in Table I.

#### B. Analog-to-Digital Converter

The current-to-frequency conversion technique offers a wide dynamic range at the cost of low conversion rate. The ADC is comprised of a comparator and a counter. In order to achieve a fast response, a multi-stage comparator is employed in the ADC. A differential difference pair at the input of the first stage compares a differential input with a differential reference voltage. The second stage increases the overall gain of the comparator and the third stage acts as a level shifter to make the output voltage compatible with CMOS logic.

To save silicon area a compact linear feedback shift register (LFSR) is employed as a counter. During the conversion time it counts the number of pulses generated by the comparator and streams out the one-bit digital output at the end of each conversion time.



Fig. 7. Potentiostat front end with noise sources and an electrode equivalent model.

## IV. NOISE ANALYSIS

In this section the noise performance of the proposed potentiostat is investigated. Fig. 7(a) shows a simplified circuit model of the potentiostat front end. The three noise sources in this circuit are:  $V_{n1}$  which represents the noise of the electrode,  $V_{n2}$  which accounts for the input-referred noise of the OTA, and  $I_{n1}$  that models the noise contribution of the input common-mode feedback amplifier. At low frequencies the series resistance of the electrode,  $R_S$ , is negligibly small, resulting the input-referred current noise equal to

$$\overline{I_{n,IN}^2} = \left| j\omega C_P + \frac{1}{R_P} \right|^2 \left( \overline{V_{n1}^2} + \overline{V_{n2}^2} \right) + \overline{I_{n1}^2}.$$
 (4)

In order to minimize the input-referred noise and accordingly improve the sensitivity of the potentiostat, all the three terms in equation (4) should be minimized.

The first stage of the common-mode amplifier including input transistors  $M_{1-3}$  and the current mirror transistor  $M_4$ , as well as the output stage including transistors  $M_{6-9}$  are the two noise sources in the common-mode feedback circuit. The noise of the first part appears as a common-mode signal at the input of the OTA which is removed due to its high commonmode rejection. As a result, the only sources of the noise are the output stage transistors  $M_{6-9}$ . This noise is composed of the thermal and flicker  $(\frac{1}{t})$  noises as follows

$$\overline{I_n^2} = 4kT\gamma \left(g_{m7} + g_{m9}\right) + \frac{K}{C_{OX}f} \left(\frac{g_{m7}^2}{W_7 L_7} + \frac{g_{m9}^2}{W_9 L_9}\right),$$
(5)

where k is the Boltzmann constant, T is the absolute temperature,  $\gamma$  is a technology-dependent constant,  $g_m$  is the transconductance of the transistor, K is the  $\frac{1}{f}$  noise constant,  $C_{OX}$  is the gate oxide capacitance, and W, L are the width and length of a transistor, respectively.

According to equation (5) for a fixed current, transistors operating in the strong inversion region have minimum noise contribution. The operation of transistors in the strong inversion region entails use of long transistors which also causes smaller  $\gamma$  and less  $\frac{1}{f}$  noise [?]. The transistors utilized in the OTA and the input common-mode feedback amplifier are chosen based on the above calculations.



Fig. 8. Potentiostat die micrograph (depicted twice) showing the electrodes location (left) and the circuits floorplan (right). The channel size is  $200\mu$ m ×  $200\mu$ m in a 0.35 $\mu$ m CMOS process.



Fig. 9. Experimentally measured current transfer characteristic of the potentiostat.

# V. EXPERIMENTAL RESULTS

Fig. 8 shows the die micrograph of the fully differential integrated potentiostat fabricated in a  $0.35\mu$ m standard CMOS technology. The channel has differential on-chip working electrodes. In order to validate the functionality and utility of the potentiostats, electrical and chemical characterization experiments were performed. A testing PCB interfaces with a computer through FIFOs implemented on an FPGA. A low-level current was generated by applying the output of a voltage-mode digital-to-analog converter (DAC) across a large resistor. By sweeping the digital input of the DAC a ramp current is applied to the potentiostat. Fig. 9 shows the transfer characteristic of the integrated potentiostat.

In order to test the utility of the integrated potentiostat, a cyclic voltammetry experiment was also performed. Fig. 10 shows the experimentally recorded cyclic voltammogram of a phosphate buffered silane solution with a scan rate of 100V/sec recorded by one channel of the proposed potentiostat.



Fig. 10. Cyclic voltammogram performed with one channel on a phosphate buffered silane solution.

#### VI. CONCLUSIONS

A fully differential integrated potentiostat with differential recording electrodes for biochemical sensing in noisy environments is presented. Experimental results validate the utility of the potentiostat in wide dynamic range recording. Inchannel analog-to-digital converter facilitates interfacing the microsystem with a digital signal processing unit and offers the parallelism in recording and data acquisition in a multichannel setting.

#### REFERENCES

- D.J. Michael and R.M. Wightman, "Electrochemical monitoring of biogenic amine neurotransmission in real time," *Journal of Pharmaceutical* and Biomedical Analysis, vol 19, pp 33-46, 1999.
- [2] D.L. Robinson, B.J. Venton, M.L.A. Heien, and R.M. Wightman, "Detecting subsecond dopamine release with fast-scan cyclic voltametry in vivo," *Clinical Chemistry*, vol 49, pp 1763-1773, 2003.
- [3] R.J. Reay, S.P. Kounaves, and G.T.A. Kovacs, "An integrated CMOS potentiostat for miniaturized electroanalytical instrumentation," *IEEE Int. Solid-State Circuits Conference (ISSCC'94)*, pp. 162-163, 1994.
- [4] R. Genov, M. Stanacevic, M. Naware, G. Cauwenberghs, and N. Thakor, "16-channel Integrated potentiostat for distributed neurochemical sensing," *IEEE T. Circuits and Systems I*, vol. 53, no. 11, pp. 2371-2376, November 2006.
- [5] M. Stanacevic, K. Murari, G. Cauwenberghs, and N. Thakor, "16channel wide-range VLSI potentiostat array," *IEEE Trans. Biomedical Circuits and Systems*, 2004.
- [6] A. Bandyopadhyay, G. Mulliken, G. Cauwenberghs, and N. Thakor, "VLSI potentiostat array for distributed electrochemical neural recording," *IEEE Int. Circuits and Systems (ISCAS'02)*, vol. 2, pp. 740-743, May 26-29, 2002.
- [7] A. Hassibi and T.H. Lee, "A Programmable electrochemical biosensor array in 0.18µm standard CMOS," *IEEE Int. Solid-State Circuits Conference*, vol 1, pp 564-565, 2005.
- [8] M. Augustyniak, C. Paulus, R. Brederlow, N. Persike, G. Hartwich, D. Schmitt-Landsiedel, and R. Thewes, "A 24×16 CMOS-based chronocoulometric DNA Microarray," *IEEE Int. Solid-State Circuits Conference*, vol 1, pp 59-68, 2006.
- [9] K. Murari, M. Stanacevic, G. Cauwenberghs, and N. Thakor, "Integrated Potentiostat for Neurotransmitter Sensing," *IEEE Engineering in medicine and biology magazine*, vol. 24, 2005.
- [10] R.F.B. Turner, D.J. Harrrison, and H.P. Baltes, "A CMOS potentiostat for amperometric chemical sensors," *IEEE Journal of Solid-State Circuits*, vol. 22, 1987.