

An Implantable Wireless Multi-Channel Neural Prosthesis for Epidural Stimulation

Dai Jiang¹, Clemens Eder¹, Timothy A. Perkins², Anne Vanhoostenberghe³, Matthew Schormans¹, Fangqi Liu¹, Virgilio Valente¹, Nick Donaldson², and Andreas Demosthenous¹

¹Department of Electronic and Electrical Engineering, ²Department of Medical Physics and Bioengineering, ³Institute of Orthopaedics and Musculoskeletal Sciences, University College London, London WC1E 6BT, United Kingdom
d.jiang@ucl.ac.uk, a.demosthenous@ucl.ac.uk

Abstract—This paper presents a fully implantable multi-channel neural prosthesis for epidural stimulation. The prosthesis features three telemetry-operated independent stimulators providing in total eighteen stimulation channels. The stimulator circuits were implemented in a 0.6- μm CMOS technology. The prosthesis is protected in a hermetically sealed ceramic enclosure and encapsulated in medical grade silicone rubber. *In-vitro* measured results with electrodes in saline are presented.

I. INTRODUCTION

Research has demonstrated the potential of using epidural stimulation to restore coordinated locomotion, such as standing and walking, to patients suffering from paralysis due to severe spinal cord injury (SCI) [1]-[4]. As illustrated in Fig. 1, in the absence of input from the brain as a result of lesion in the spinal cord, the natural sensory information required for locomotion is projected onto the spinal cord by applying modulated electrical stimuli via an epidural electrode array implanted along the spinal cord. The intrinsic sensorimotor neural circuits embedded in the spinal cord can generate organised motor patterns for standing, walking and climbing in response to the stimulation. Research in animal models has shown a linear relationship between the frequency or amplitude of the electrical stimuli in pulsatile stimulation and the step height. A neural modulation scheme developed based on this result has enabled rats to walk on a treadmill for 1000 steps without failure [4]. Clinical trials in several human subjects also achieved weight-bearing standing and assisted stepping behaviour under epidural stimulation [1]-[3].

Advances in the epidural stimulation research demand a purpose built, multi-mode stimulator that is able through the implantable multi-electrode array to selectively target multiple sites of the spinal circuitries with precisely controlled pulse timing and intensity [4]-[5]. In addition, the stimulator should support simultaneous stimulation in multiple channels with advanced neural modulation schemes to coordinate variable motor outputs into unique functional movement such as standing and stepping [6]. The stimulator should also be suitable for chronic implantation for long-term studies. The stimulators used in existing studies, are either external devices with percutaneous

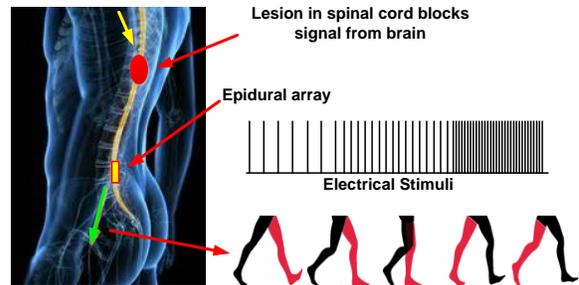


Fig. 1. Epidural stimulation facilitates gait cycles by modulating the electrical stimuli.

connections to implanted electrodes [4], [5], [7] that are only suitable for acute study, or modified commercial medical implants for off label use that only support simple on-off neural modulation [1]-[3].

A wireless, fully implantable, multi-site, multi-mode epidural prosthesis is presented which has three independent stimulators providing uncorrelated simultaneous stimulation in similar or different modulation schemes, targeting multiple sites along the spinal cord. Each stimulator supports multiple sites for selecting the most effective “hot spots”. The prosthesis is hermetically sealed in a ceramic package and encapsulated for chronic implantation. The implant is wirelessly powered by an inductive link and is controlled by a remote (programming) computer over a 900 MHz radio link, primarily for research studies with freely moving animals. The rest of the paper is organised as follows. Section II describes the overall architecture of the neural prosthesis, the operation of the telemetry, the stimulation procedure and the implant packaging. Section III presents measured results from *in-vitro* tests, followed by conclusions in Section IV.

II. SYSTEM AND CIRCUIT ARCHITECTURE

A. Overall System

The architecture of the implantable prosthesis system is shown in Fig. 2(a). The system comprises a wearable transmitter, an implantable prosthesis and a remote host computer. The

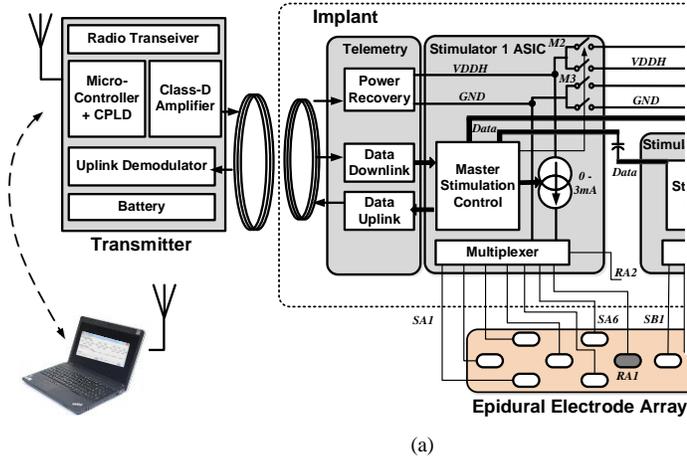


Fig. 2. (a) Architecture of the implantable multi-channel stimulator system; (b) Photo of the implantable stimulator comprising 5M (digital and analog), 18V (stimulator output supply voltage), 3M (digital and analog), 1.8V (stimulator output supply voltage), 3M (digital and analog), 1.8V (stimulator output supply voltage). 1-Telemetry, 2-Stimulator 1, 3-Stimulator 2, 4-Stimulator 3.

wearable transmitter receives instructions from the remote host via a 900 MHz radio link and forwards these instructions to the implant via a 9.6 MHz inductive link. A class-D amplifier in the transmitter drives the primary coil of the inductive link to power the implant. The inductive link also serves as a bi-directional half-duplex communication link. The processor in the transmitter translates received instructions into command frames that are sent to the implant for setting the stimulation characteristics. The frames are sent via downlink transmission whose carrier is modulated by on-off keying (OOK) at 400 kbit/s. The uplink acknowledgement from the implant to the transmitter uses passive-phase-shift keying (PPSK) modulation with a data rate at 600 kbit/s.

The implantable prosthesis comprises a receiver coil unit and the hybrid, as shown in Fig. 2(b), connected with Cooner Wire AS632. The receiver coil unit consists of only passive components, including the secondary coil of the inductive link and half-wave rectifiers to convert received power into dc voltages. There is a rare earth magnet at the center of the receiver coil unit for alignment with another magnet in the primary coil, also located its center. The hybrid comprises a telemetry for power regulation and data communication, and three parallel integrated stimulators (ASICs), as shown in Fig. 2(a) and 2(b). Each stimulator drives six stimulating electrodes (SA1-SA6, SB1-SB6 and SC1-SC6) and two return electrodes (RA1-RA2, RB1-RB2 and RC1-RC2) on the epidural electrode array. A stimulation channel is formed with one stimulating electrode and one return electrode. In total eighteen channels can be formed between the three stimulators.

B. Implant Electronics

The implant electronics are separated into the receiver coil unit and the hybrid. As shown in Fig. 3, the receiver coil unit has six connections to the hybrid, including a high voltage supply, a low voltage supply, the ground (0 V) rail, the carrier frequency as a 9.6 MHz clock, and two communication ports for the uplink and downlink data. The secondary coil, L_2 , is tuned with C_2 at 9.6 MHz. The coupled oscillating signal on L_2C_2 is rectified by D_5C_4

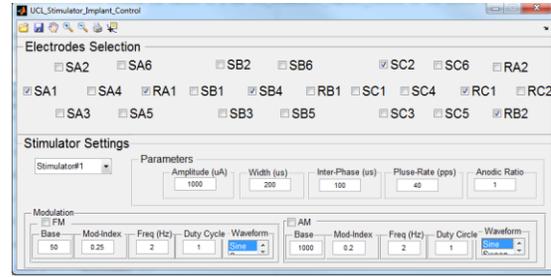


Fig. 4. Graphical user interface (GUI) for setting stimulation parameters.

TABLE I. FEATURES AND PERFORMANCE OF THE IMPLANT SYSTEM

Hybrid dimensions	46 mm × 42.8 mm × 8.8 mm
Stimulator ASICs	0.6- μ m HV CMOS (b)
Stimulation type	Biphasic constant current pulsatile stimulation
Stimulation current	≤ 3 mA, step size 12 μ A
Pulse rate	500 pps, resolution ≤ 0.5 pps
Pulse duration	Cathodic phase: 10–500 μ s
Anodic phase	1–8 times cathodic width
Stimulating electrodes	3, each driving 6 stimulating electrodes and 2 return electrodes
Primary coil	2 mm diameter, 0.5 mm gauge, 5 turns, 1.8 μ H
Secondary coil	28 mm diameter, 0.1 mm gauge, 7 turns, 2.52 μ H
working distance	3–11 mm

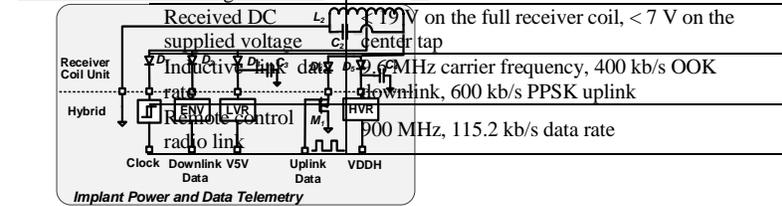


Fig. 3. Schematic of the power and data telemetry, with photos of the external transmitter (top and bottom sides).

into a dc voltage, which provides a stable 18 V supply using a high voltage regulator (HVR, MIC5233) in the hybrid. Another half-wave rectifier D_3C_3 is connected to the centre tap on L_2 to extract a lower dc voltage, which is regulated by a low voltage regulator (LVR) in the hybrid to a 5 V supply for the stimulation control logic. The LVR is implemented in the Stimulator 1 ASIC. An envelope detector (ENV) implemented in the Stimulator 1 ASIC in the hybrid is connected via D_2 to the centre tap on L_2 to demodulate the downlink data in the OOK scheme. The signal on the centre tap is also connected via D_1 to a Schmitt trigger (74AHCT1G14) in the hybrid to serve as a RF clock for the uplink data modulation (using PPSK [8]). In this scheme, the control logic in Stimulator 1 ASIC shorts L_2 by turning on a transistor switch, M_1 , also implemented in the same ASIC, for exactly 1.5 RF clock cycles from the time instant when the voltage on L_2 crosses zero. This action causes a transient voltage surge on the primary coil, L_1 . Uplink data transfer is achieved by applying the uplink bit-stream to the gate of M_1 at a speed of 600

kbit/s. Similar to load-shift keying (LSK), PPSK is also a passive modulation scheme but provides a faster data rate because of its transient nature.

The hybrid derives both high and low dc voltages as well as the downlink data from the receiver coil unit for the operation of the three parallel stimulators. The circuits of the stimulators are integrated into three ASICs. Each ASIC comprises a control logic, a 5 V regulator to generate a stable dc voltage for the logic, a 8-bit linear current digital-to-analog converter (DAC) with a dynamic range between 0 mA and 3 mA, an ‘‘H’’-bridge output stage to form biphasic current pulses, and a multiplexer to select the stimulating and return electrodes. Stimulator 1 ASIC also contains demodulator circuits for the downlink and a modulator for the uplink as previously described. The control logic in the Stimulator 1 ASIC also handles the bi-directional communication with the transmitter and manages the stimulation settings on all the three stimulators. A power-isolation method [9] was implemented to provide efficient isolation between stimulators, so that the crosstalk between the stimulators is minimised and thus ensuring the independence of the modulation scheme and pulse timing on each stimulator. In this arrangement, the power supplies to Stimulator ASIC 2 and 3 are gated in Stimulator ASIC 1, as shown in Fig. 2(a). During a biphasic pulse on ASIC 2 or ASIC 3, the switches to the ASIC ($M2$ or $M3$) are switched off and the ASIC is supplied by the storage capacitor, $C2$ or $C3$, respectively. After the current pulse, the ASIC is reconnected to the power supply for charging the storage capacitor. Stimulation settings to ASIC 2 and ASIC 3 are sent via a capacitive data link from Stimulator ASIC 1. This power isolation ensures no potential crosstalk paths so that current pulses can occur simultaneously on multiple stimulators without interfering with each other.

C. Implant Packaging and Encapsulation

The components in the hybrid were mounted on a ceramic (96% alumina) substrate in a size of 36 mm \times 38 mm with gold and platinum-gold tracks thick-film printed. The circuits were hermetically sealed under a metallised ceramic lid, as shown in Fig. 2(b). Six contacts to the receiver coil unit and twenty-four contacts to the epidural electrode array outside the lid were connected to the circuits via hermetic feedthroughs. The components in the receiver coil unit were mounted on a FR4 printed circuit board. The two units were joined by Cooner wires. The whole implant was encapsulated in medical grade silicone rubber EPM2420.

D. External Transmitter and Remote Control

The hardware architecture of the transmitter is shown in Fig. 3. The operation of the transmitter is managed by a microcontroller (MSP430). The microcontroller communicates with a remote computer via a 900 MHz radio link implemented with easyRadio advanced ISM radio module eRA900TRS [10]. Stimulation parameters, including the pulse settings and the modulation schemes, are defined through a graphical user interface, shown in Fig. 4, and sent to the microcontroller via the radio link. The settings are then be passed to a CPLD device XC2V256, where command frames are formed with additional

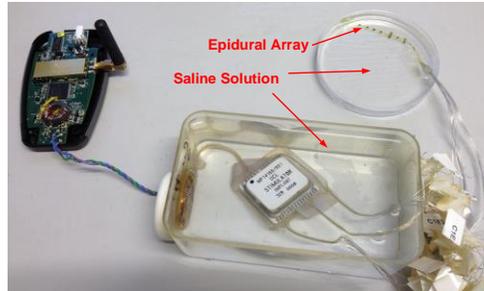


Fig.5. Setup for *in-vitro* testing.

CRC bits. The CPLD also applies OOK modulation to a 9.6 MHz carrier. A class-D amplifier drives the primary coil, L_1 , with the modulated carrier signal to power the implant. The class-D amplifier is powered from a dc-dc converter LT1615, whose output voltage is programmed by the microcontroller via a potentiometer AD5220. According to the feedback from the implant, the transferred power is adjusted to improve power efficiency. The uplink data are demodulated from the voltage on C_b , which together with C_a tunes L_1 to 9.6 MHz. The voltage surge on the carrier is extracted with a low-pass filter (LPF, built with Op-Amp $A2$, AD8656) preceded by an envelope detector (ENV) and an amplifier (built with Op-Amp $A1$, AD8656). The demodulated bit-stream is then converted to digital by a Schmitt trigger (74LVC1G17). The CPLD extracts information from the bit-stream and passes it to the microcontroller. The transmitter is supplied by a rechargeable lithium polymer battery LP-443440 (3.7 V, 600 mAh).

III. MEASUREMENTS

The stimulator ASICs were fabricated in a 0.6- μ m high-voltage CMOS technology. The stimulators used separate chips to prevent a common substrate that would prevent power isolation. The implant features and the measured specifications are summarised in Table I.

The neural prosthesis system was tested *in-vitro* as shown in Fig. 5. The implant was submerged in saline solution (16.7 mS/cm²). The secondary coil in the implant is aligned with the primary coil through the wall of the container. The implant is connected to an epidural electrode array (Courtesy of Fraunhofer ICT-IMM) that is also submerged in saline solution. The connection was made at an adaptor for measuring the current pulses. Fig. 6 shows the performance of alternative stimulations on multiple channels from a single stimulator. Three electrodes on an array were connected to three stimulating outputs on Stimulator 1. Symmetrical biphasic pulses with amplitude of 800 μ A and a pulse rate of 100 pps were delivered to the three electrodes in a sequence where, in a cycle with a length of one second, stimuli on Electrode $SA1$ started from 0 s to 0.4 s, on Electrode $SA2$ from 0.3 s to 0.7 s and on Electrode $SA3$ from 0.5 s to 0.9 s. Fig. 7 demonstrates independent modulation schemes on different stimulators. On Stimulator 1, 2 Hz sinusoidal amplitude modulation was applied with a base amplitude of 300 μ A and a swing of 400 μ A. On Stimulator 2 and Stimulator 3, 2 Hz sinusoidal frequency modulation was applied with a base

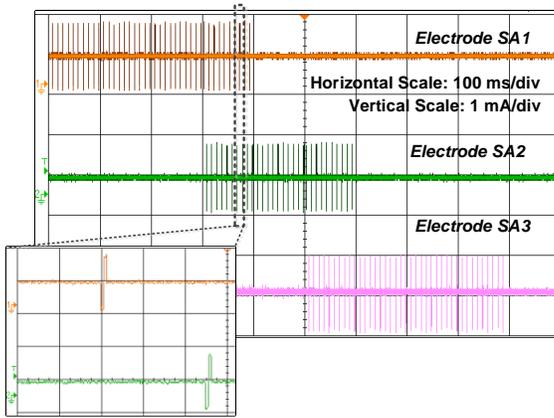


Fig. 6. Measured waveforms of an exemplar interleaved pulses from one stimulator for sequentially stimulating multiple “hot spots”.

pulse rate of 250 pps and a swing of 400 pps.

IV. CONCLUSIONS

A wirelessly powered and controlled, fully implantable, multi-channel, multi-mode neural prosthesis for epidural spinal cord stimulation has been presented. The prosthesis has been developed for research on restoring locomotion to patients after spinal cord injury. The details of the system implementation have been presented, and the functionality has been demonstrated with *in-vitro* tests.

ACKNOWLEDGEMENT

The authors would like to thank Prof David Borton for the discussions on the device specifications and Dr Peter Detemple for providing the epidural electrode array for the *in-vitro* evaluation.

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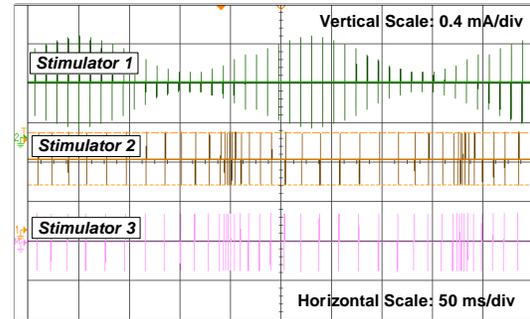


Fig. 7. Measured waveforms of independent stimulation from the three stimulators with different modulation schemes.

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