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(54) MINIATURE WIRELESS BIOMEDICAL TELEMETRY DEVICE

DRAHTLOSE BIOMEDIZINISCHE MINIATUR-TELEMETRIEVORRICHTUNG

DISPOSITIF MINIATURE DE TÉLÉMÉTRIE BIOMÉDICALE SANS FIL

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Description

BACKGROUND

1. Field of the Invention

[0001] The embodiments described herein relate to biomedical telemetry devices, and more particularly to miniature wireless biomedical telemetry devices and methods of their use.

2. Background

[0002] The untethered monitoring of disease and injury in animal models, primarily in rats and mice, and comparable monitoring of humans in cases of suspected or demonstrated disease or injury, is important for the development of new therapies, determining markers for seizures or other acute events, and the development and application of treatment protocols.

[0003] Currently, monitoring physiological signals such as electroencephalogram (EEG), electrocardiogram (ECG), blood pressure, etc. requires cumbersome monitoring systems that generally involve surgical implantation and/or have limited monitoring time because of power consumption and, in humans, require expensive hospital stays with numerous electrodes, and as a minimum, result in limited mobility and otherwise prevent a normal way-of-life.

[0004] Epilepsy is the second most prevalent neurological disorder with over 50 million people affected worldwide. Nearly three million individuals in the U.S. have epilepsy, and over 30% are refractory to treatment with antiepileptic drugs (AEDs). The medical, psychological, sociological and financial implications of refractory epilepsy can be devastating for both the patient and their families. Seizures can be debilitating and result in major irreversible morbidity, often with long-term consequences that may include brain damage from recurrent seizures, seizure-induced injuries and accidents associated with loss of consciousness, impairment of memory, and even death. The personal consequences of refractory epilepsy may include the adverse effects of AEDs, strained personal and family relations, and the inability to obtain and hold meaningful employment or even a driver's license. Thus, there is a need for better approaches and new technologies for discovering novel therapies for intractable epilepsy.

[0005] One of the most widely used techniques for recording epileptic seizures is the EEG. This technique remains the mainstay to diagnose epilepsy, and to help localize the seizure onset zone in people with intractable epilepsy. As in the clinical diagnosis of the epilepsies, the EEG has become indispensable for translational research in animal models of both genetic and acquired epilepsy. This technique is also widely used to provide fundamental information about the mechanisms of sleep and to diagnose sleep disorders. For some research,

"wired" or "tethered" recordings are quite acceptable and can be done in adult rats for weeks at a time. However, electrical noise, movement artifacts, and the risk that tethered animals will injure themselves by pulling on the cable have led to the need for radio-telemetry over the last several years. Current "wireless" systems, however, often involve extensive surgery to implant the transmitter and battery system, an expensive and time-intensive process. Other systems use a relatively large and awkward "backpack" on the animal, which causes animal stress similar to a tether.

[0006] Further, acquired epilepsy (i.e., after brain injury, of many possible types) affects up to about 3 percent of the human population, and acute seizures after various forms of brain injury impact a much higher proportion. For example, post-traumatic epilepsy affects up to about 53 percent of adults suffering moderate to severe traumatic brain injury (TBI). Importantly, seizures early after TBI, similar to stroke, are associated with increased injury and poorer outcomes. Thus, optimal acute therapy for TBI victims and other individuals with brain injuries includes prompt seizure detection and effective therapy leading to seizure cessation. Continuous EEG monitoring reveals that roughly 25 percent of adults suffering moderate or severe TBI exhibit electrographic seizures, despite therapeutic levels of anticonvulsive medications. Nearly 60 percent of post-traumatic seizures in adults suffering severe TBI are non-convulsive seizures and can be detected only by EEG monitoring.

[0007] Aggressive, monitor-guided, early anti-seizure medical therapy may mitigate secondary brain injury after TBI and reduce the probability and severity of subsequent cognitive deficits or post-traumatic epilepsy. Ideally, EEG monitoring would be initiated as soon as possible after TBI, in order to detect non-convulsive seizures and guide anti-seizure therapy in the transport or pre-hospital setting. Reliable EEG recording during medical transport or field care will require a small device that can be properly positioned on the head quickly and lacks cumbersome cables or receivers. Unfortunately, an easy-to-use, telemetric, portable recording device is not currently available.

[0008] What are needed, therefore, are devices and methods of use that overcome challenges found in the current state of the art, some of which are described above.

[0009] US 2006/020300 discloses a wireless communication system that includes an antenna structure adapted for coupling to a medical device antenna when the medical device is not implanted in a patient's body. The antenna structure effectively extends the medical device antenna length, thereby improving the efficiency and reliability of a communication link between the medical device and a programmer or monitor outside the implanted environment. The antenna structure is fabricated from any conductive material, which may be in the form of conductive wire, tape, ink, foil, film, adhesive or the like, and is attached to a portion of a medical device packaging

assembly or another accessory device or substrate such as a pouch or overlay.

[0010] US 2006/173259 discloses a system and method for an improved biological interface system that processes multicellular signals of a patient and controls one or more devices. The system includes a sensor that detects the multicellular signals and a processing unit for producing the control signal based on the multicellular signals.

SUMMARY

[0011] The invention is defined in claim 1. Further aspects and preferred embodiments are defined in the appended claims. Aspects, embodiments and examples of the present disclosure which do not fall under the scope of the appended claims do not form part of the invention and are merely provided for illustrative purposes. Furthermore, the methods presented in the present description are provided for illustrative purposes only and do not form part of the present invention. Provided herein are embodiments of a miniature wireless biomedical telemetry device along with systems and methods for its use. A miniature amplifier and transmitter allow recording of physiological signals from small animals, such as rats, mice and birds, as well as humans. The device is positioned externally and is easily replaceable, as is its battery, so surgical complications and other challenges are minimized.

[0012] In one aspect, a miniature, wireless, capacitive-coupled biomedical telemetry device and system having long-life and minimally-invasive surgical procedures configured to monitor physiological signals (e.g., EEG, ECG, blood pressure, etc.) as well as environmental sensors (e.g., temperature, pressure, etc.) is described.

[0013] In another aspect, a miniature, wireless, biomedical telemetry device and system comprised of a transmitter, electrodes and power source integrated on a single substrate having long-life and configured to monitor physiological signals (e.g., EEG, ECG, blood pressure, etc.) as well as environmental sensors (e.g., temperature, pressure, etc.) is described.

[0014] In yet another aspect, described herein are embodiments of a miniature telemetry system to permit continuous, uninterrupted EEG recordings; this will be accomplished with a capacitive-coupled telemetry system that uses commonly-available batteries (e.g., watch batteries) as a power source, which would allow straightforward replacement of either the battery or the entire unit without invasive surgery.

[0015] In one aspect the power source for the telemetry unit can be one or more photodiodes.

[0016] In another aspect, the wireless telemetry device can comprise a multi-channel transmitter such as, for example, a pulse-width modulated (PWM) two-channel transmitter.

[0017] Although the most pressing need is probably for research and clinical care during and after various

forms of brain injury, embodiments according to the present invention have potential use in many other experimental animal and human clinical studies. These include, but are not limited to antiepileptic drug testing and optimization in both animal and human studies, research and care of migraine headache, and analyses of sleep disorders, including development and optimization. Additional advantages will be set forth in part in the description which follows or may be learned by practice. The advantages will be realized and attained by means of the elements and combinations particularly pointed out in the appended claims. It is to be understood that both the foregoing general description and the following detailed description are exemplary and explanatory only and are not restrictive, as claimed.

BRIEF DESCRIPTION OF THE DRAWINGS

[0018] The accompanying drawings, which are incorporated in and constitute a part of this specification, illustrate embodiments and together with the description, serve to explain the principles of the methods and systems:

FIG. 1 is an exemplary computing device that can be used according to aspects of the present invention for processing and analysis of the received information or performing any other needed computing/processing functions.

FIGS 2A and 2B illustrate embodiments of transmitters that are small enough to mount externally on the head of a laboratory rat, mouse, or other small animal and that is able to transmit physiological signals according to an aspect of the present invention;

FIG. 2C illustrates an embodiment of a two channel PWM transmitter;

FIG. 2D illustrates an embodiment of a two channel receiver that can be used with a transmitter such as the one of FIG. 2C that provides one voltage proportional to the duration of the positive signal and one voltage proportional to the duration of the negative signal;

FIG. 2E illustrates an embodiment of a photodiode power source for supplying power to a transmitter;

FIG. 3A illustrates one design of a differential antenna high-impedance electric field receiver that can be used with single channel transmitters (such as those shown in FIG. 2A and 2B) that provide an output voltage proportional to the frequency of the transmitter;

FIG. 3B illustrates one design of a single antenna high-impedance electric field receiver according to an aspect of the present invention;

FIG. 4 illustrates an embodiment of a standard rat cage modified for capacitive-coupling of the transmitter and receipt of physiological data;

FIG. 5 is a schematic diagram of an embodiment of a miniature EEG telemetry system;

FIGS. 6-7A illustrate embodiments of a standard rat cage modified for capacitive-coupling of the transmitter and receipt of physiological data;
 FIGS. 7B illustrates an embodiment of a cage (e.g., a bird cage) having multiple antenna(s) such that a differential input in the FM receiver then provides a constant signal as the subject moves from one antenna toward the other;
 IGS. 8A-11B illustrate aspects of embodiments of the transmitter device according to the present invention that transmits physiological signals to an antenna that is connected to an amplifier near the cage/shielding;
 FIGS. 12 and 13 illustrate repetitive seizures induced in a rat treated with kainate;
 FIG. 14 illustrates an embodiment where multiple subjects can be simultaneously and continuously monitored for physiological activity as well as videoed;
 FIG. 15 is a schematic diagram of the top and side views of an embodiment of the EEG telemetry device;
 FIG. 16 is an illustration of a RF FM EEG transmitter according to the present invention; and
 FIG. 17 is a flowchart illustrating a method of wireless biomedical telemetry.

DETAILED DESCRIPTION

[0019] Before the present methods and systems are disclosed and described, it is to be understood that the methods and systems are not limited to specific synthetic methods, specific components, or to particular compositions, as such may, of course, vary. It is also to be understood that the terminology used herein is for the purpose of describing particular embodiments only and is not intended to be limiting.

[0020] As used in the specification and the appended claims, the singular forms "a," "an" and "the" include plural referents unless the context clearly dictates otherwise. Ranges may be expressed herein as from "about" one particular value, and/or to "about" another particular value. When such a range is expressed, another embodiment includes from the one particular value and/or to the other particular value. Similarly, when values are expressed as approximations, by use of the antecedent "about," it will be understood that the particular value forms another embodiment. It will be further understood that the endpoints of each of the ranges are significant both in relation to the other endpoint, and independently of the other endpoint.

[0021] "Optional" or "optionally" means that the subsequently described event or circumstance may or may not occur, and that the description includes instances where said event or circumstance occurs and instances where it does not.

[0022] "Exemplary" means "an example of" and is not intended to convey a preferred or ideal embodiment. Fur-

ther, the phrase "such as" as used herein is not intended to be restrictive in any sense, but is merely explanatory and is used to indicate that the recited items are just examples of what is covered by that provision.

[0023] Other definitions used herein include:

A/D: analog-to-digital;
 dbm: decibel power level in milliwatts;
 EEG: electroencephalogram;
 FCC: Federal Communication Commission;
 FM: frequency modulation;
 PC: personal computer;
 RF: radio-frequency;
 PWM: pulse-width modulation; and
 USB: universal serial bus.

[0024] The present methods and systems may be understood more readily by reference to the following detailed description of preferred embodiments and the Examples included therein and to the Figures and their previous and following description.

[0025] In one embodiment described herein is a micro-transmitter comprised of a physiological amplifier controlling a pulse-width (or frequency) modulation oscillator. The recording input is two leads connected to an AC-coupled amplifier (or DC for pressure and other sensor measurements). In one aspect, the amplitude of the physiological signal modulates the pulse width of a square-wave oscillator, which is transmitted via capacitive coupling to the antenna(s). Advantageously, this process is performed with one miniaturized device so that it can be fastened directly to the head (or elsewhere) of the animal. In one aspect, the miniaturized device is self-contained and includes a power source such as a small battery. In one aspect, the device presently operates from a single 1.4 V zinc air cell, and power consumption is <3 μ A, providing an estimated >270 days of operation with a #5 cell. In another aspect, photodiodes, as are known in the art, are used as the power source. A high-impedance receiver picks up, amplifies, and provides filtering of the signal. In one aspect, the signal can be leveled via an automatic gain control (AGC). An integrator (or a frequency-to-voltage converter) and a band-pass filter recover the original AC signal from the transmitter.

[0026] In another aspect, the device is coupled with a radio frequency (RF) transmitter for wirelessly transmitting monitored data to a remote receiver.

[0027] In some instances, the receiver can be operably connected with a computing device, such as that shown in FIGS. 1 and 5 for processing and analysis of the received information or performing any other needed computing/processing functions.

[0028] Functions as described herein can be performed by software, hardware, or a combination of software and hardware. A unit can be software, hardware, or a combination of software and hardware. The units can comprise the Analysis Software 106 as illustrated in

FIG. 1 and described below. In one exemplary aspect, the units can comprise a computer 101 as illustrated in FIG. 1 and described below.

[0029] FIG. 1 is a block diagram illustrating an exemplary operating environment for performing the disclosed method. This exemplary operating environment is only an example of an operating environment and is not intended to suggest any limitation as to the scope of use or functionality of operating environment architecture. Neither should the operating environment be interpreted as having any dependency or requirement relating to any one or combination of components illustrated in the exemplary operating environment.

[0030] The present methods and systems can be operational with numerous other general purpose or special purpose computing system environments or configurations. Examples of well known computing systems, environments, and/or configurations that can be suitable for use with the system and method comprise, but are not limited to, personal computers, server computers, laptop devices, and multiprocessor systems. Additional examples comprise set top boxes, programmable consumer electronics, network PCs, minicomputers, mainframe computers, distributed computing environments that comprise any of the above systems or devices, and the like.

[0031] The processing of the disclosed methods and systems can be performed by software components. The disclosed system and method can be described in the general context of computer-executable instructions, such as program modules, being executed by one or more computers or other devices. Generally, program modules comprise computer code, routines, programs, objects, components, data structures, etc. that perform particular tasks or implement particular abstract data types. The disclosed method can also be practiced in grid-based and distributed computing environments where tasks are performed by remote processing devices that are linked through a communications network. In a distributed computing environment, program modules can be located in both local and remote computer storage media including memory storage devices.

[0032] Further, one skilled in the art will appreciate that the system and method disclosed herein can be implemented via a general-purpose computing device in the form of a computer 101. The components of the computer 101 can comprise, but are not limited to, one or more processors or processing units 103, a system memory 112, and a system bus 113 that couples various system components including the processor 103 to the system memory 112. In the case of multiple processing units 103, the system can utilize parallel computing.

[0033] The system bus 113 represents one or more of several possible types of bus structures, including a memory bus or memory controller, a peripheral bus, an accelerated graphics port, and a processor or local bus using any of a variety of bus architectures. By way of example, such architectures can comprise an Industry

Standard Architecture (ISA) bus, a Micro Channel Architecture (MCA) bus, an Enhanced ISA (EISA) bus, a Video Electronics Standards Association (VESA) local bus, an Accelerated Graphics Port (AGP) bus, and a Peripheral Component Interconnects (PCI) bus also known as a Mezzanine bus. The bus 113, and all buses specified in this description can also be implemented over a wired or wireless network connection and each of the subsystems, including the processor 103, a mass storage device 104, an operating system 105, analysis software 106, received data 107, a network adapter 108, system memory 112, an Input/Output Interface 110, a display adapter 109, a display device 111, and a human machine interface 102, can be contained within one or more remote computing devices 114a,b,c at physically separate locations, connected through buses of this form, in effect implementing a fully distributed system.

[0034] The computer 101 typically comprises a variety of computer readable media. Exemplary readable media can be any available media that is accessible by the computer 101 and comprises, for example and not meant to be limiting, both volatile and non-volatile media, removable and non-removable media. The system memory 112 comprises computer readable media in the form of volatile memory, such as random access memory (RAM), and/or non-volatile memory, such as read only memory (ROM). The system memory 112 typically contains data such as received data 107 and/or program modules such as operating system 105 and analysis software 106 that are immediately accessible to and/or are presently operated on by the processing unit 103.

[0035] In another aspect, the computer 101 can also comprise other removable/non-removable, volatile/non-volatile computer storage media. By way of example, FIG. 1 illustrates a mass storage device 104 which can provide non-volatile storage of computer code, computer readable instructions, data structures, program modules, and other data for the computer 101. For example and not meant to be limiting, a mass storage device 104 can be a hard disk, a removable magnetic disk, a removable optical disk, magnetic cassettes or other magnetic storage devices, flash memory cards, CD-ROM, digital versatile disks (DVD) or other optical storage, random access memories (RAM), read only memories (ROM), electrically erasable programmable read-only memory (EEPROM), and the like.

[0036] Optionally, any number of program modules can be stored on the mass storage device 104, including by way of example, an operating system 105 and analysis software 106. Each of the operating system 105 and analysis software 106 (or some combination thereof) can comprise elements of the programming and the analysis software 106. Received data 107 can also be stored on the mass storage device 104. Received data 107 can be stored in any of one or more databases known in the art. Examples of such databases comprise, DB2®, Microsoft® Access, Microsoft® SQL Server, Oracle®, MySQL, PostgreSQL, and the like. The databases can be cen-

tralized or distributed across multiple systems.

[0037] In another aspect, the user can enter commands and information into the computer 101 via an input device (not shown). Examples of such input devices comprise, but are not limited to, a keyboard, pointing device (e.g., a "mouse"), a microphone, a joystick, a scanner, tactile input devices such as gloves, and other body coverings, and the like. These and other input devices can be connected to the processing unit 103 via a human machine interface 102 that is coupled to the system bus 113, but can be connected by other interface and bus structures, such as a parallel port, game port, an IEEE 1394 Port (also known as a Firewire port), a serial port, or a universal serial bus (USB).

[0038] In yet another aspect, a display device 111 can also be connected to the system bus 113 via an interface, such as a display adapter 109. It is contemplated that the computer 101 can have more than one display adapter 109 and the computer 101 can have more than one display device 111. For example, a display device can be a monitor, an LCD (Liquid Crystal Display), or a projector. In addition to the display device 111, other output peripheral devices can comprise components such as speakers (not shown) and a printer (not shown) which can be connected to the computer 101 via Input/Output Interface 110.

[0039] The computer 101 can operate in a networked environment using logical connections to one or more remote computing devices 114a,b,c. By way of example, a remote computing device can be a personal computer, portable computer, a server, a router, a network computer, a peer device or other common network node, and so on. Logical connections between the computer 101 and a remote computing device **114a,b,c** can be made via a local area network (LAN) and a general wide area network (WAN). Such network connections can be through a network adapter **108**. A network adapter **108** can be implemented in both wired and wireless environments. Such networking environments are conventional and commonplace in offices, enterprise-wide computer networks, intranets, and the Internet **115**.

[0040] For purposes of illustration, application programs and other executable program components such as the operating system **105** are illustrated herein as discrete blocks, although it is recognized that such programs and components reside at various times in different storage components of the computing device **101**, and are executed by the data processor(s) of the computer. An implementation of analysis software **106** can be stored on or transmitted across some form of computer readable media. Computer readable media can be any available media that can be accessed by a computer. By way of example and not meant to be limiting, computer readable media can comprise "computer storage media" and "communications media." "Computer storage media" comprise volatile and non-volatile, removable and non-removable media implemented in any method or technology for storage of information such as computer read-

able instructions, data structures, program modules, or other data. Exemplary computer storage media comprises, but is not limited to, RAM, ROM, EEPROM, flash memory or other memory technology, CD-ROM, digital versatile disks (DVD) or other optical storage, magnetic cassettes, magnetic tape, magnetic disk storage or other magnetic storage devices, or any other medium which can be used to store the desired information and which can be accessed by a computer.

[0041] The methods and systems can employ Artificial Intelligence techniques such as machine learning and iterative learning. Examples of such techniques include, but are not limited to, expert systems, case based reasoning, Bayesian networks, behavior based AI, neural networks, fuzzy systems, evolutionary computation (e.g. genetic algorithms), swarm intelligence (e.g. ant algorithms), and hybrid intelligent systems (e.g. Expert inference rules generated through a neural network or production rules from statistical learning).

[0042] Embodiments according to the present invention of the device are miniature, so it can be placed externally on rats, mice, birds or other small laboratory animals with quick and straight-forward surgical procedures. In one aspect, the bandwidth is 0.1-2.78 kHz (or DC for pressure and other measurements), which is suitable for most physiological signals. Other frequencies are contemplated within the scope of the invention as faster integrated circuits (IC's) would provide a higher frequency response (e.g., 20 kHz) to allow recording of action potentials, but can reduce battery life. Because the device is positioned externally and the system is shielded, it requires far less power than conventional radiotelemetry systems, so the power source is also much smaller than is used for other telemetry devices. Because the power source (e.g., battery) is smaller, it too is located externally rather than being implanted in the animal, so it can be replaced easily. If the power source is a battery, then the batteries are generally inexpensive (i.e., watch batteries, hearing-aid batteries, etc.). Alternative to the use of replaceable batteries, surface-mounted photodiodes using infrared illumination can be used with the purpose of providing a continuous power source that does not need to be changed or replaced. Embodiments of this device allow prolonged and continuous wireless recording of physiological signals.

[0043] In one embodiment, a transmitter (FIGS. 2A and 2B) that is small enough to mount externally on the head of a laboratory rat, mouse, or other small animal and that is able to transmit physiological signals continuously for a year or longer is described. FIG. 2A illustrates an embodiment of a FM transmitter 202 for receiving information from EEG or ECG electrodes, modulating the information, and transmitting the modulated information to a receiver. The antenna of the transmitter is capacitively coupled with the antenna of the receiver. The physiological signals can either be electrophysiological recordings (EEG, ECG, etc.) or signals from a transducer (e.g., air or blood pressure). In one aspect, implantable recording

electrodes (for example, stainless steel 0.005" wire) can be used for EEG measurement. However, it should be appreciated that any biological/medical transducer can be used with this transmitter system. For example, this system should be useful for monitoring slowly changing signals, such as temperature changes, in organisms ranging from small animals like reptiles in biological and ecological research to college and professional athletes engaged in competitive activities. Highfrequency signals, most notably action potentials, can also be recorded in a research or clinical setting from lab animals and human patients by simply altering the frequency response of the amplifier in the transmitter. For example, FIG. 2B illustrates a transmitter 206 that can be used to monitor, modulate and transmit a signal related to an environmental condition such as atmospheric pressure or temperature. In one aspect, the transmitter operates from, for example, a single "#5" 1.4 volt zinc air cell and is designed using a 600 nA per channel dual operational amplifier. The low-power design of the transmitter combined with "electric field" capacitive coupling to a high-impedance receiver (FIGs. 3A and 3B) is a basis for an embodiment of the device.

[0044] One exemplary embodiment of the miniaturized device comprising a transmitter and a power source such as a 1.4 V zinc air cell has a volume of less than 1 cc, for example about 0.64 cc, and a weight of less than 1 gram. For example, one embodiment weighs about 0.8 grams, with the power source (e.g., battery). The small weight and volume makes the miniaturized device suitable for mounting on the head or body of a rat, mouse, or other laboratory animal. The low-profile design reduces stress on the animal's headcap and provides little interference with movement of the animal.

ANIMAL CONTAINMENT FOR CAPACITIVE COUPLING

[0045] In one aspect, because of capacitive coupling between the antenna of the transmitter and the ground plane, the animal subject must remain oriented between the antenna and ground plane, which the containment cage provides. The maximum distance between the antenna and ground plane is also controlled, and the subject should not be allowed to cover the antenna. In one aspect, the antenna is isolated from external capacitive loads by a capacitive isolation shield (i.e., driven shield) because of the capacitance between the transmitter and receiver antenna is so small. Because capacitive coupling is used according to one aspect, a number of animals can be monitored in parallel in neighboring cages without the risk of crosstalk.

[0046] A standard rat cage provides such an environment when placed on the receiver antennae system, as indicated in FIGS. 4-7B. FIGS. 6 and 7A illustrate an embodiment of an integrated antenna, amplifier and receiver unit 602. FIG. 7A illustrates the integrated unit of FIG. 6 as used with a standard rat cage 700.

[0047] In another aspect, a differential antenna system is utilized. In larger animal containers with subjects that move in three dimensions such as birds, the receiving distance to a single antenna can be exceeded. One solution is the use of two antennas, one 702 on top and one 704 on the bottom of the container 706, as shown in FIG. 7B. A differential input 708 to the FM receiver 710 then provides a constant signal as the subject moves from one antenna toward the other. It is also contemplated that complex containers can utilize multiple antennas (greater than two).

TRANSMITTER

[0048] An embodiment of an FM transmitter (FIG. 2A) 202 for electroencephalogram (EEG) and electrocardiogram (ECG) operates from, for example, a 1.4 volt zinc air cell. Referring to FIG. 2A, U1A forms an AC-coupled amplifier with gain set by R4 and R3. C1 sets a low-frequency cut off pole. Resistors R1 and R2 set a voltage of $\frac{1}{2} V+$ and the signal electrode is coupled to this point through C2. Note that $\frac{1}{2} V+$ and the amplified signal are present on U1A pin 1.

[0049] U1B forms a square-wave oscillator with frequency set by R6 and C3. R7 provides positive feedback and R5 sets the trip point. Note that R5 sees $\frac{1}{2} V+$ plus a varying signal that modulates the trip point. Due to non-linear plus and minus slew rates of U1B, the trip point modulates the frequency of oscillation, which is coupled to the antennae on pin 7. U1 is selected for its extremely low current draw of 600 nA per section.

[0050] A second implementation is the FM transmitter for pressure (FIG. 2B) 206. R1 and R2 reduce the current through the MSI 1451 MEMS transducer and center the input to U1A-3 and R3 at $\frac{1}{2} V+$. The output from the MEMS bridge is amplified by U1A. U1B operates as described above. Other inputs contemplated within the scope of this invention include strain gauges, chemical transducers, microphones, light sensors, and many other types of transducers or sensors can be used as inputs.

[0051] FIG. 2C illustrates an embodiment of a PWM two channel transmitter 208. This configuration requires minimum number of parts and low power consumption which is ideal for very small devices. U3 is a modified square wave oscillator with frequency controlled by current through R12 and R11 charging and discharging C3. U1 is an amplifier biased to clamp positive voltage at R12 via D1 and U2 is an amplifier biased to clamp negative voltage at R12 via D2. Each amplifier is modulated by an input signal and controls the positive and negative duration of U3 independently. A receiver 302 such as the one shown in FIG. 3A provides a voltage proportional to the duration of the positive swing (channel one) and the negative swing (channel two).

[0052] FIG. 2D illustrates an embodiment of a two channel receiver 212 that can be used with a transmitter such as the one of FIG. 2C that provides one voltage proportional to the duration of the positive signal and one

voltage proportional to the duration of the negative signal. In FIG. 2D, a square waveform input (originating from a transmitter such as the one shown in FIG. 2C) is sent to U1, amplified, and sent through a band pass filter. Comparator U2 responds to the modulated positive duration of the waveform which is then integrated by U3 to recover the original signal. Comparator U4 responds to the modulated negative duration of the waveform which is then integrated by U5 to recover the original signal.

POWER SOURCE

[0053] As described herein, in one aspect the power source for the transmitter can be small, inexpensive batteries that can be replaced without surgical invasion of the subject which allows continuous or near continuous recording of information for extended periods. These batteries can be, for example, a #5, 1.4 V zinc air cell as are commonly available. In another instance, the transmitter can be powered by one or more photodiodes. An embodiment of a photodiode power source 210 is shown in FIG. 2E. Some experiments require that a signal be recorded for long periods of time. It is therefore desirable that a transmitter of these signals operate continuously and indefinitely. Because of the extremely low power requirement (< 1.5 micro amps at 1.25 volts) of transmitter embodiments described herein, such transmitters can be powered by a small array of surface mount photodiodes (D2, D3, D4 and D5). A typical photo diode operating in the 940 nanometer range can produce about 0.6 volts at about 10 micro amps. Four diodes connected in series to a storage capacitor will provide about 2.4 volts to a voltage regulator U1, which draws about 3.9 micro amps. The regulator U1 provides a stable 1.25 volts for the transmitter. The total power drain is about $1.5 + 3.9 = 5.4$ micro amps, well within the range of small surface mount photodiodes under typical 940 nanometer illumination.

FM RECEIVER/CONVERTER

[0054] FIG. 3B illustrates a single channel receiver and amplifier 304 that can be used to receive the modulated signal from the transmitter. An exemplary IC U3, R2 and R3 (FIG. 3B) form an embodiment of a high-impedance amplifier with a gain of two. R4 is selected to match the antennae. The gain of two allows R1 to drive a capacitive isolation shield used in some applications. R27 and C19 form a 6db/octave low-pass filter. U7, R28, R29, C20, R30, C21, and R31 form a 12db/octave high-pass filter. U5 is an inverting amplifier with gain set by R7 and R5. C8, R6, and C9 form high and low pass poles.

[0055] U6 along with R8, C10, R9, C12, C11, and R11 form a frequency-to-voltage converter that converts the frequency modulated signal from the transmitter back to the original signal.

[0056] U4A, R23, C13, and R13 form a DC offset adjustment. U4B, R16, R17, R14, R15, C15, and C16 form a 12 Db/octave low pass filter. U4D, R18, R20, R21, R19,

C18 and C17 form a 12db/octave low pass filter.

[0057] All filter frequency points and amplifier gains vary depending on the application.

[0058] R24 adjusts gain. U4C, R25, and R26 with associated jumpers configure the gain and polarity of the output signal. R22 and C14 form a 6db/octave low-pass filter.

USE OF THE TELEMETRY DEVICE

[0059] In one aspect, to record the EEG, an embodiment of the telemetry device 502 is implanted on the head of a laboratory animal 504 (e.g., a rat, FIGS. 5, 8A and 8B), that is housed in a standard animal cage 506 that is within a shielded enclosure 508. The telemetry device 502, which is only a few millimeters (FIGS. 8A-11B), transmits physiological signals 514 to an antenna 510 that is connected to an amplifier/receiver 512 near the cage/shielding.

[0060] In an experiment, an embodiment of the device has been used to record electrophysiological signals 516 for weeks from a rat treated with kainate, which induces repetitive seizures (FIG. 12 and 13). Since the kainate-induced status epilepticus involves repeated convulsive seizures, the likelihood that the telemetry device will remain operational during normal and even abnormal movement is high.

[0061] The recording device allows a myriad of physiological signals to be recorded from small experimental animals, particularly immature rats and mice. These recording procedures have been difficult if not impossible, but may become routine with variations of the proposed device. For example, embodiments according to the present invention allow the EEG to be recorded from small animals for chronic studies of epileptic seizures, which has important value in antiepileptic drug development. The EEG-recording device can also be useful for drug-development studies relating to promotion of sleep. A similar device can also be used for measurement of physiological signals from a small bird, as part of neuroscience research on the mechanisms of "bird song."

[0062] In another aspect, as shown in FIG. 14, multiple subjects can be simultaneously and continuously monitored for physiological activity as well as videoed for physical activity.

MINIATURE RF TELEMETRY DEVICE

[0063] Further described herein are embodiments of a miniature radio-frequency (RF) telemetry device to record, for example, electroencephalogram from persons with acute TBI, which allows remote monitoring and recording.

[0064] An embodiment of the device is comprised of a millimeter-sized EEG electrode/transmitter unit that wirelessly transmits EEG signals to a radio-frequency receiver connected to, for example, a laptop computer. The only component on the patient is a small electrode/trans-

mitter unit. In one aspect, the device can be used to reliably detect acute TBI-associated seizures during a medical emergency both in transit to and within a field hospital setting. EEG telemetry strategies and techniques described herein provide prompt detection and continued monitoring of seizures under conditions where traditional ambulatory EEG is not feasible. In addition, this approach will have other uses in both military and civilian settings (e.g. easy, prompt seizure detection after field-exposure to nerve gas agents).

[0065] Described herein is an embodiment of a single-channel telemetry-based EEG device that will allow detection of electrographic seizures in an emergency medical setting. It is to be appreciated that several independent channels can be used simultaneously. The device uses a pair of small EEG electrodes coupled with a miniature telemetry-based recording/transmitter system that will allow acquisition and storage of EEG data on a computing device such as the one described in relation to FIG. 1, above. A USB A/D acquisition hardware, laptop PC, and chart recorder/storage software can be used in coordination with the device, including analysis of received data by analysis software and transmission of received data over a network (e.g., the Internet) for review and/or further analysis by trained medical personnel or analysis software.

[0066] Described below is a small, lightweight transmitter that can be easily mounted on a human skull. In one embodiment, characteristics of an exemplary transmitter include a transmitting radius of about 50 feet, a battery life with, for example, a 3 volt lithium 1632 cell (125 MAH) of at least about 24 hours, and a bandwidth of approximately 0.1 to 300 Hz. FIG. 15 illustrates an embodiment of the device. FIG. 15 is a schematic diagram of the top and side views of an embodiment of the EEG telemetry device. As above, it is also to be appreciated that several such transmitters can be used simultaneously to monitor a single subject.

[0067] The embodiment of FIG. 15 can be constructed on a substrate 1502 such as a flexible circuit board with built-in EEG or other physiological recording electrodes and/or environmental sensors 1504 attached to both ends. In one aspect, the substrate is a single piece of material. In one aspect, the antenna can be incorporated into the flexible circuit board. The flexible circuit board can be, in one example, approximately 0.008" thickness. The rigid board 1506, as shown on FIG. 15, can, in one example, be 0.02" thickness. Each electrode can have a region for conductive gel 1508 to connect the two electrodes to the scalp or other body part. One electrode has a battery 1510 on the opposite side, and the other has the transmitter electronics on the opposite side. The device is designed to have a low profile so that it will not be easily dislodged or be uncomfortable to an awake individual. In one aspect, embodiments according to this device provide rapid detection of non-convulsive seizures after TBI and allow early, monitor-directed anti-seizure therapy.

Transmitter

[0068] FIG. 16 is an illustration of a RF FM EEG transmitter 1602 according to the present invention. In this embodiment, U1 is a MCP6142 dual operational amplifier in a MSOP surface mount package. It is biased at one-half of U2's output by divider R1 and R2. The EEG signal is AC coupled through C2 to U1A. Gain is set with R4/R3 and low frequency response set by C1. U1B is a square wave oscillator with frequency set by R6 and C3. R5 coupled to U1A-1 sets the pulse width at about 50% for both positive and negative saturation times of U1B. Resistor R5 also modulates the pulse width duration of U1B with the signal waveform from U1A. The non-linear properties between the rise and fall times of U1B provide significant frequency modulation allowing an FM type receiver to be used. U2 is a precision voltage reference providing approximately 1.247 volts from the 3 volt lithium cell "B". This powers U1 for stable operation and sets the center operating frequency of U3 through R9. R8 couples the FM oscillator U1B to modulate RF oscillator/amplifier U3. L1 sets center frequency of U3, L2 is a "pull up" for the open collector output of U3. U3 as shown operates at approximately 234.275 megahertz. However, the FCC allows power levels below 10 dbm on some parts of the FM band. Embodiments of the current transmitter produce about -21 dbm into 50 ohms, and is tunable from 40 to 450 MHz. Therefore frequency bands between 40 and 450 MHz that are not otherwise used, and are available for the telemetry transmission system, are within the scope of embodiments of this invention.

Receiver

[0069] In one aspect, an FM wideband receiver (e.g., a ICOM IC-R5 receiver) is used to pick up the RF signal from the transmitter. The audio output is then fed to bandwidth limits and a frequency to voltage converter (similar to FIG. 3B) to recover the original EEG signal.

EXAMPLES

[0070] The following examples are put forth so as to provide those of ordinary skill in the art with a complete disclosure and description of how the compounds, compositions, articles, devices and/or methods claimed herein are made and evaluated, and are intended to be purely exemplary and are not intended to limit the scope of the methods and systems. Efforts have been made to ensure accuracy with respect to numbers (e.g., amounts, temperature, etc.), but some errors and deviations should be accounted for.

[0071] In one example, aspects according to the present invention can be used for at-home, untethered, longer-term monitoring of a patient post-TBI and early prediction and treatment of seizure activity. Further examples of use include immediate detection of EEG activity soon after a TBI and proscribed treatment predicat-

ed on such activity. Another use is to monitor patients that have experienced a single seizure. In many instances, such a patient should not or cannot drive for weeks or months until it is clear that the patient has not experienced another seizure. Yet another use would be to help patients with intractable epilepsy obtain more reliable measures of their seizure frequency and severity, since these individuals are often amnesic after a seizure and thus self-reporting is generally inaccurate. Embodiments of the miniature telemetry device described herein can be used to determine whether the patient has or has not experienced another seizure. Embodiments according to the invention can also be used for remote monitoring of EEG activity in various situations.

[0072] FIG. 17 is a flowchart illustrating a method of wireless biomedical telemetry. The exemplary process comprises step 1702, at least partially implanting a miniaturized device weighing less than one gram and having a volume less than one cubic centimeter and comprising a wireless transmitter and a power source into a subject. The subject can be a laboratory animal such as a rat or a mouse, or can be a human. The miniaturized device is at least partially implanted into the subject with minimally invasive procedures and at least a portion of the miniaturized device is external to the subject. The wireless transmitter of the miniaturized device is operably connected to one or more electrodes or sensors that are attached to and/or implanted in the subject. At step 1704, the transmitter converts monitored physiological parameters or environmental conditions of the subject into a modulated signal. At step 1706, a receiver operably connected with one or more antennae receives the modulated signal from the wireless transmitter. The one or more antennae are each capacitively coupled with the wireless transmitter. The receiver demodulates the modulated signals from the wireless transmitter, thereby allowing the physiological parameters or environmental conditions of the subject to be monitored.

[0073] While the methods and systems have been described in connection with preferred embodiments and specific examples, it is not intended that the scope be limited to the particular embodiments set forth, as the embodiments herein are intended in all respects to be illustrative rather than restrictive.

[0074] Unless otherwise expressly stated, it is in no way intended that any method set forth herein be construed as requiring that its steps be performed in a specific order. Accordingly, where a method inventive concept does not actually recite an order to be followed by its steps or it is not otherwise specifically stated in the claims or descriptions that the steps are to be limited to a specific order, it is no way intended that an order be inferred, in any respect. This holds for any possible non-express basis for interpretation, including: matters of logic with respect to arrangement of steps or operational flow; plain meaning derived from grammatical organization or punctuation; the number or type of embodiments described in the specification.

[0075] Throughout this application, various publications are referenced. The disclosures of these publications may be consulted to more fully describe and understand the state of the art to which the methods and systems pertain.

[0076] It will be apparent to those skilled in the art that various modifications and variations can be made without departing from the scope. Other embodiments will be apparent to those skilled in the art from consideration of the specification and practice disclosed herein. It is intended that the specification and examples be considered as exemplary only, with a true scope being indicated by the following inventive concepts.

Claims

1. A wireless biomedical telemetry system comprising:

one or more electrodes or sensors configured for attachment to a living subject (504) that is moveable in three dimensions relative to or on a bottom of a cage (706), said one or more electrodes or sensors being configured to detect a signal generated by the living subject and to generate a physiological signal reflective of the signal generated by the living subject;

a telemetry device (502) for external attachment to the living subject, said telemetry device including a wireless transmitter (202) and a power source (210), said wireless transmitter being operably connected to the power source to receive power therefrom and to said one or more electrodes or sensors to receive said physiological signal, said wireless transmitter being configured to generate a modulated signal based on said physiological signal;

a transmitting antenna connected to said wireless transmitter for transmitting the modulated signal, said transmitting antenna being moveable with said living subject relative to or on said bottom of the cage;

a receiving antenna (510) sized and fixedly positioned away from said living subject to effect a capacitive coupling with said transmitting antenna to receive through said cage said modulated signal;

a receiver (212) connected to said receiving antenna and configured to demodulate said modulated signal to form a receiving signal; and

a means connected to said receiver to receive and selectively store, based on said receiving signal, data reflective of the signal generated by the living subject.

2. The system of claim 1, wherein the wireless transmitter is a multi-channel wireless transmitter.

3. The system of claim 1, wherein the wireless transmitter is a FM transmitter or a PWM transmitter.
4. The system of claim 1, wherein the power source is a battery. 5
5. The system of claim 1, wherein the battery can be replaced in the telemetry device without surgical invasion of the subject. 10
6. The system of claim 1, wherein the power source is one or more photodiodes.
7. The system of claim 1, wherein the physiological signal includes one or more of EEG, ECG, and blood pressure. 15
8. The system of claim 1, wherein the physiological signal includes one or more of temperature and pressure. 20
9. The system of claim 1, wherein the living subject is a human, a rat or a mouse.
10. The system of claim 1, wherein the telemetry device weighs less than one gram and has a volume of less than one cubic centimetre. 25
11. The system of any preceding claim, wherein the transmitting antenna is isolated from external capacitive loads by a capacitive isolation shield. 30
12. The system of claim 11, wherein the capacitive isolation shield is located between the receiving antenna and the ground plane. 35
13. The system of claim 11, wherein the receiving antenna is planar and a distance between the receiving antenna and the ground plane is controlled. 40
14. The system of any preceding claim, wherein the one or more electrodes or sensors, the telemetry device, and the transmitting antenna, are incorporated into a flexible circuit board. 45

Patentansprüche

1. Drahtloses biomedizinisches Telemetriesystem, Folgendes umfassend: 50
 - eine/n oder mehrere Elektroden oder Sensoren, welche zur Befestigung an einem lebenden Subjekt (504) konfiguriert sind, welches in drei Dimensionen in Bezug auf einen Käfig (706) oder auf seinem Boden beweglich ist, wobei die/der eine oder die mehreren Elektroden oder Sensoren konfiguriert sind, ein Signal zu detek-

tieren, welches durch das lebende Subjekt erzeugt wird, und ein physiologisches Signal zu erzeugen, welches das Signal wiedergibt, welches durch das lebende Subjekt erzeugt wird; eine Telemetrievorrichtung (502) zur externen Befestigung an dem lebenden Subjekt, wobei die Telemetrievorrichtung einen drahtlosen Sender (202) und eine Stromquelle (210) umfasst, wobei der drahtlose Sender mit der Stromquelle, um Strom daraus zu erhalten, und mit der/dem einen oder den mehreren Elektroden oder Sensoren betriebsfähig verbunden ist, um das physiologische Signal zu empfangen, wobei der drahtlose Sender konfiguriert ist, ein modulierte Signal auf der Grundlage des physiologischen Signals zu erzeugen; eine Sendeantenne, welche mit dem drahtlosen Sender zum Senden des modulierten Signals verbunden ist, wobei die Sendeantenne mit dem lebenden Subjekt in Bezug auf den Käfig oder auf seinen Boden beweglich ist; eine Empfangsantenne (510), welche von dem lebenden Subjekt entfernt bemessen und fest angeordnet ist, um eine kapazitive Koppelung mit der Sendeantenne zu bewirken, um das modulierte Signal durch den Käfig hindurch zu empfangen; einen Empfänger (212), welcher mit der Empfangsantenne verbunden ist und konfiguriert ist, das modulierte Signal zu demodulieren, um ein Empfangssignal auszubilden; und ein Mittel, welches mit dem Empfänger verbunden ist, um auf der Grundlage des Empfangssignals Daten zu empfangen und selektiv zu speichern, welche das Signal wiedergeben, welches durch das lebende Subjekt erzeugt wird.

2. System nach Anspruch 1, wobei der drahtlose Sender ein drahtloser Mehrkanalsender ist.
3. System nach Anspruch 1, wobei der drahtlose Sender ein FM-Sender oder ein PWM-Sender ist.
4. System nach Anspruch 1, wobei die Stromquelle eine Batterie ist.
5. System nach Anspruch 1, wobei die Batterie in der Telemetrievorrichtung ohne chirurgischen Eingriff an dem Subjekt ersetzt werden kann.
6. System nach Anspruch 1, wobei die Stromquelle eine oder mehrere Fotodioden ist.
7. System nach Anspruch 1, wobei das physiologische Signal ein oder mehrere Signale aus EEG, EKG und Blutdruck umfasst.
8. System nach Anspruch 1, wobei das physiologische

Signal ein oder mehrere Signale aus Temperatur und Druck umfasst.

9. System nach Anspruch 1, wobei das lebende Subjekt ein Mensch, eine Ratte oder eine Maus ist. 5
10. System nach Anspruch 1, wobei die Telemetrievorrichtung weniger als ein Gramm wiegt und ein Volumen von weniger als einem Kubikzentimeter aufweist. 10
11. System nach einem der vorhergehenden Ansprüche, wobei die Sendeantenne von externen kapazitiven Lasten durch eine kapazitive Abschirmung isoliert ist. 15
12. System nach Anspruch 11, wobei die kapazitive Abschirmung zwischen der Empfangsantenne und der Erdungsebene lokalisiert ist. 20
13. System nach Anspruch 11, wobei die Empfangsantenne planar ist und eine Entfernung zwischen der Empfangsantenne und der Erdungsebene gesteuert ist. 25
14. System nach einem der vorhergehenden Ansprüche, wobei die/der eine oder die mehreren Elektroden oder Sensoren, die Telemetrievorrichtung und die Sendeantenne auf einer flexiblen Schaltungsplatine einbezogen sind. 30

Revendications

1. Système de télémétrie biomédicale sans fil comprenant : 35
- une ou plusieurs électrodes ou un ou plusieurs capteurs conçus pour une fixation à un sujet vivant (504) qui est mobile en trois dimensions par rapport à un fond de cage (706) ou sur celui-ci, ladite ou lesdites électrodes ou ledit ou lesdits capteurs étant conçus pour détecter un signal généré par le sujet vivant et pour générer un signal physiologique réfléchissant le signal généré par le sujet vivant ; 40
- un dispositif de télémétrie (502) pour une fixation externe au sujet vivant, ledit dispositif de télémétrie comprenant un émetteur sans fil (202) et une source d'alimentation (210), ledit émetteur sans fil étant connecté fonctionnellement à la source d'alimentation pour recevoir une alimentation en provenance de celle-ci et à ladite ou auxdites électrodes ou audit ou auxdits capteurs pour recevoir ledit signal physiologique, ledit émetteur sans fil étant conçu pour générer un signal modulé sur la base dudit signal physiologique ; 50
- 55

une antenne émettrice connectée audit émetteur sans fil pour émettre le signal modulé, ladite antenne émettrice étant mobile avec ledit sujet vivant par rapport audit fond de la cage ou sur celui-ci ;

une antenne réceptrice (510) dimensionnée et positionnée à demeure à distance dudit sujet vivant afin d'effectuer un couplage capacitif avec ladite antenne émettrice pour recevoir à travers ladite cage ledit signal modulé ; un récepteur (212) connecté à ladite antenne réceptrice et conçu pour démoduler ledit signal modulé afin de former un signal de réception ; et un moyen connecté audit récepteur pour recevoir et enregistrer sélectivement, sur la base dudit signal de réception, des données réfléchissant le signal généré par le sujet vivant.

2. Système selon la revendication 1, dans lequel l'émetteur sans fil est un émetteur sans fil multicanal. 20
3. Système selon la revendication 1, dans lequel l'émetteur sans fil est un émetteur FM ou un émetteur MID. 25
4. Système selon la revendication 1, dans lequel la source d'alimentation est une batterie. 30
5. Système selon la revendication 1, dans lequel la batterie peut être remplacée dans le dispositif de télémétrie sans invasion chirurgicale du sujet. 35
6. Système selon la revendication 1, dans lequel la source d'alimentation est une ou plusieurs photodiodes. 40
7. Système selon la revendication 1, dans lequel le signal physiologique comprend un ou plusieurs parmi un EEG, un ECG et la pression artérielle. 45
8. Système selon la revendication 1, dans lequel le signal physiologique comprend un ou plusieurs parmi la température et la pression. 50
9. Système selon la revendication 1, dans lequel le sujet vivant est un humain, un rat ou une souris. 55
10. Système selon la revendication 1, dans lequel le dispositif de télémétrie pèse moins d'un gramme et a un volume inférieur à un centimètre cube. 60
11. Système selon l'une quelconque des revendications précédentes, dans lequel l'antenne émettrice est isolée de charges capacitives externes par un écran d'isolation capacitif. 65
12. Système selon la revendication 11, dans lequel l'écran d'isolation capacitif est situé entre l'antenne 70

réceptrice et le plan de sol.

- 13.** Système selon la revendication 11, dans lequel l'antenne réceptrice est plane et une distance comprise entre l'antenne réceptrice et le plan de sol est commandée. 5
- 14.** Système selon l'une quelconque des revendications précédentes, dans lequel la ou les électrodes ou le ou les capteurs, le dispositif de télémétrie et l'antenne émettrice sont intégrés dans une carte de circuits imprimés souple. 10

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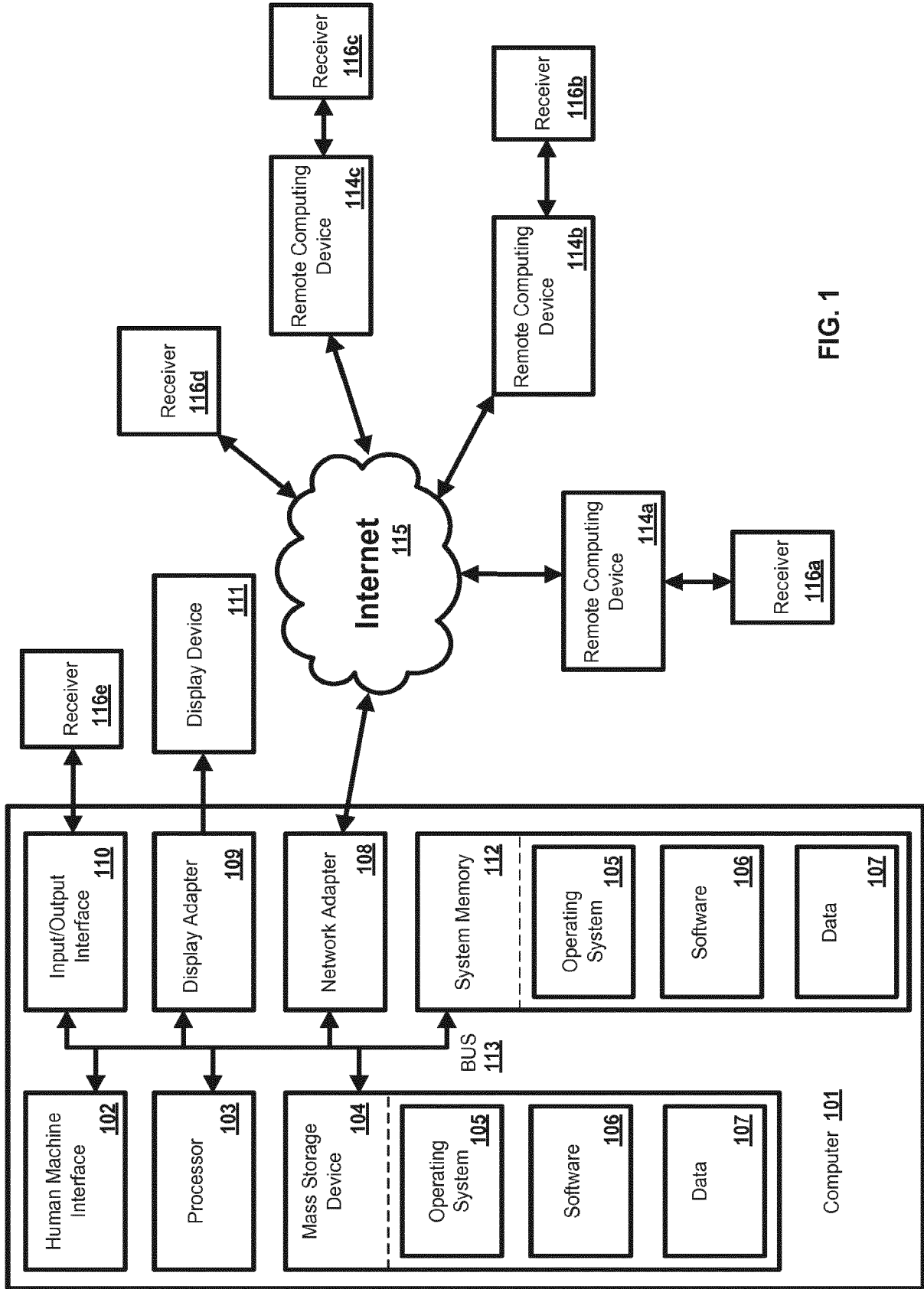


FIG. 1

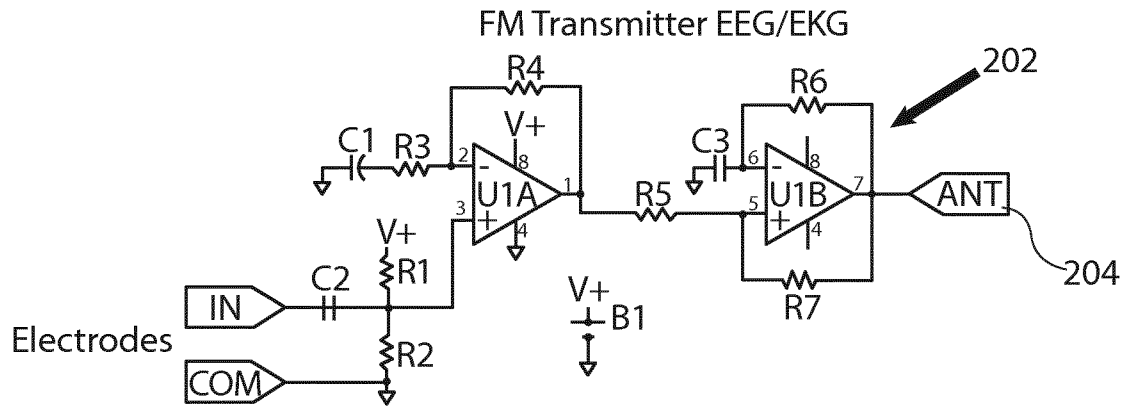


FIG. 2A

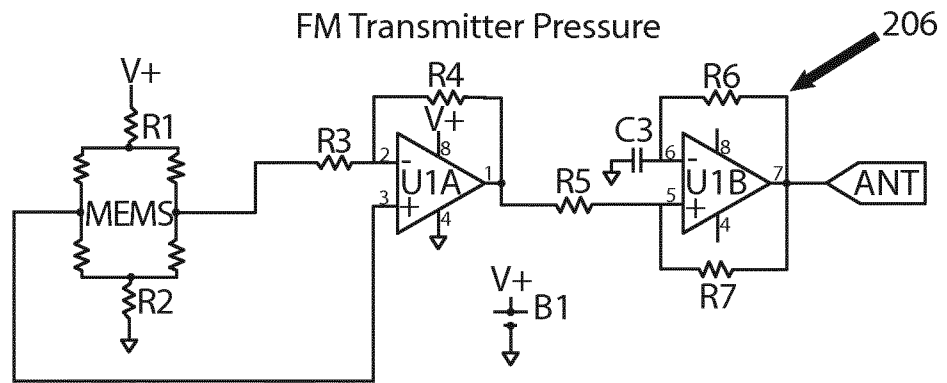


FIG. 2B

Two channel PWM Transmitter

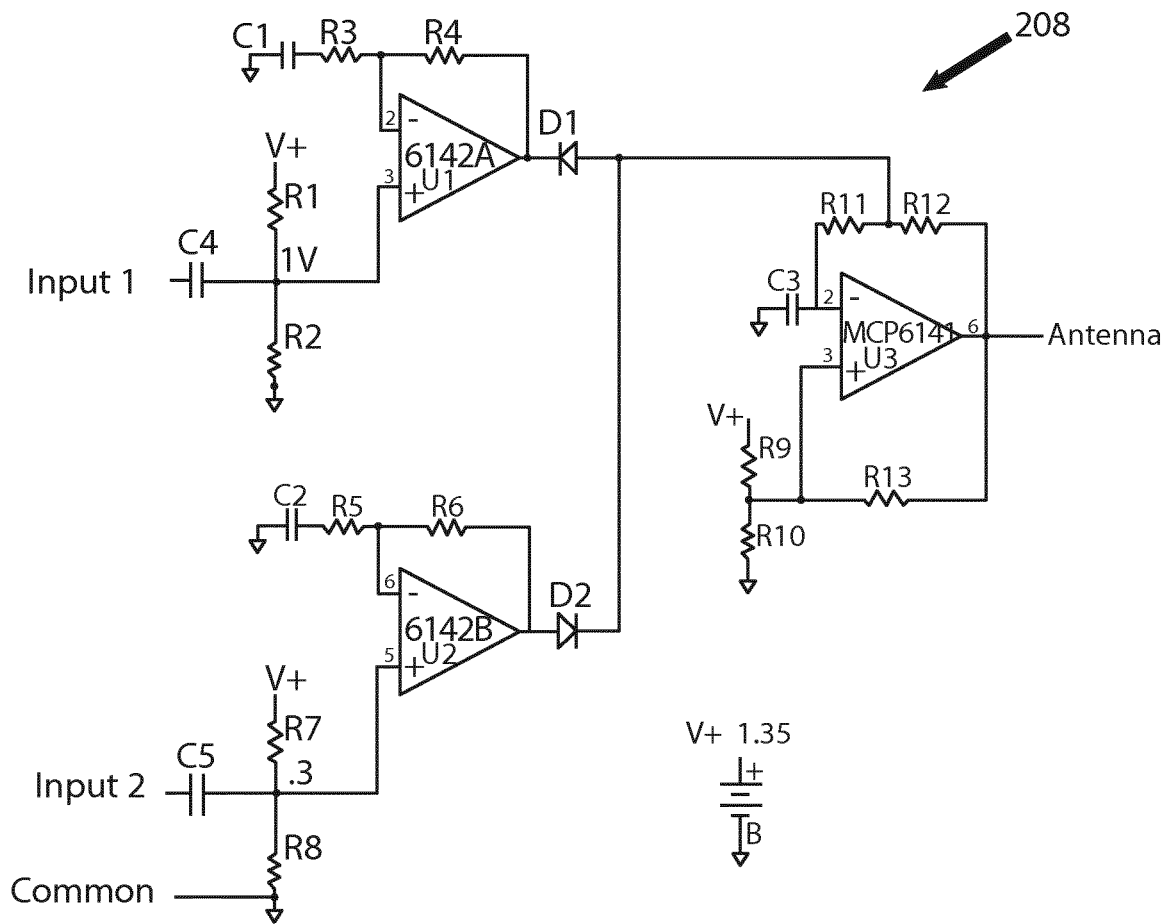


FIG. 2C

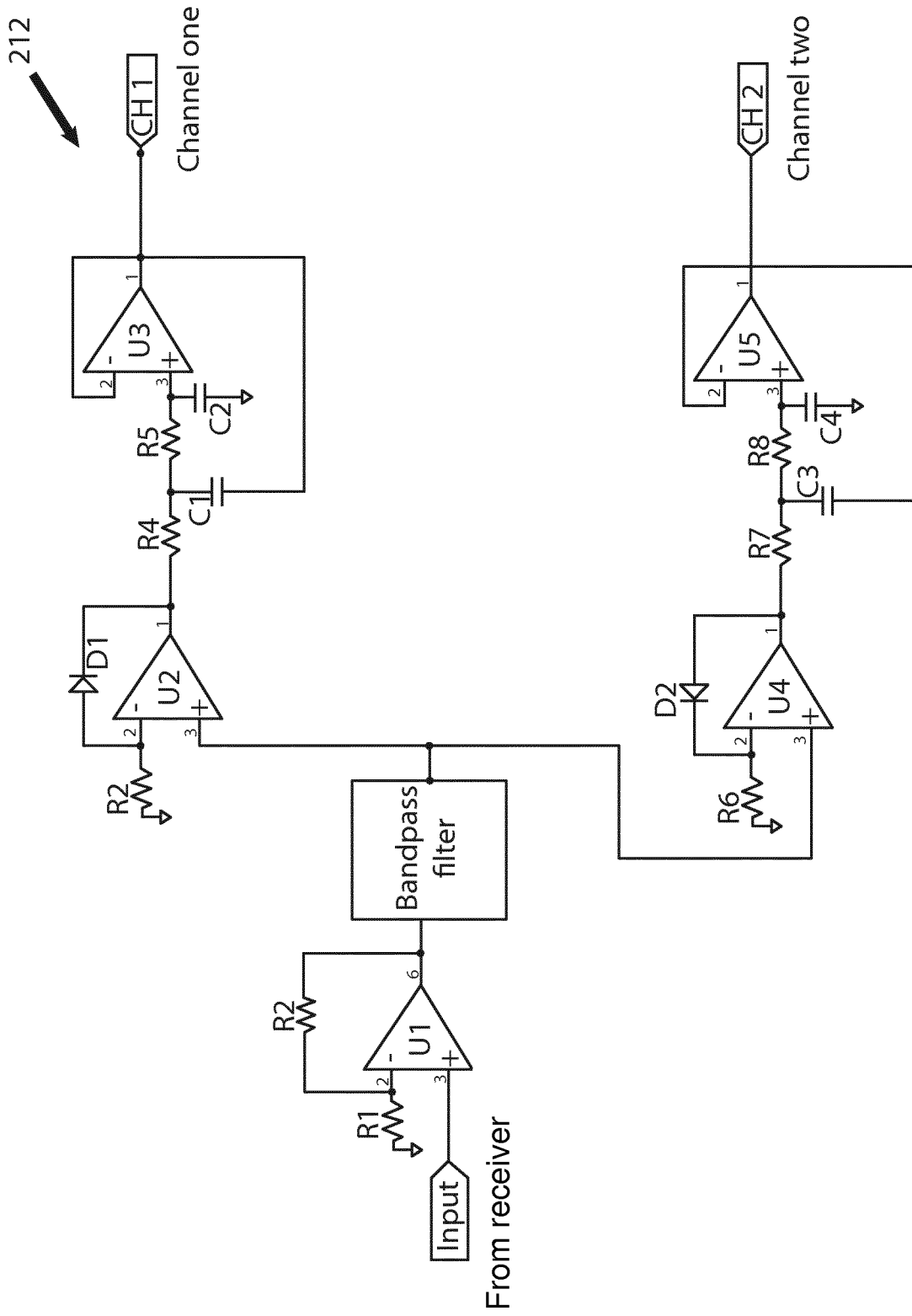


FIG. 2D

Photo diode power source

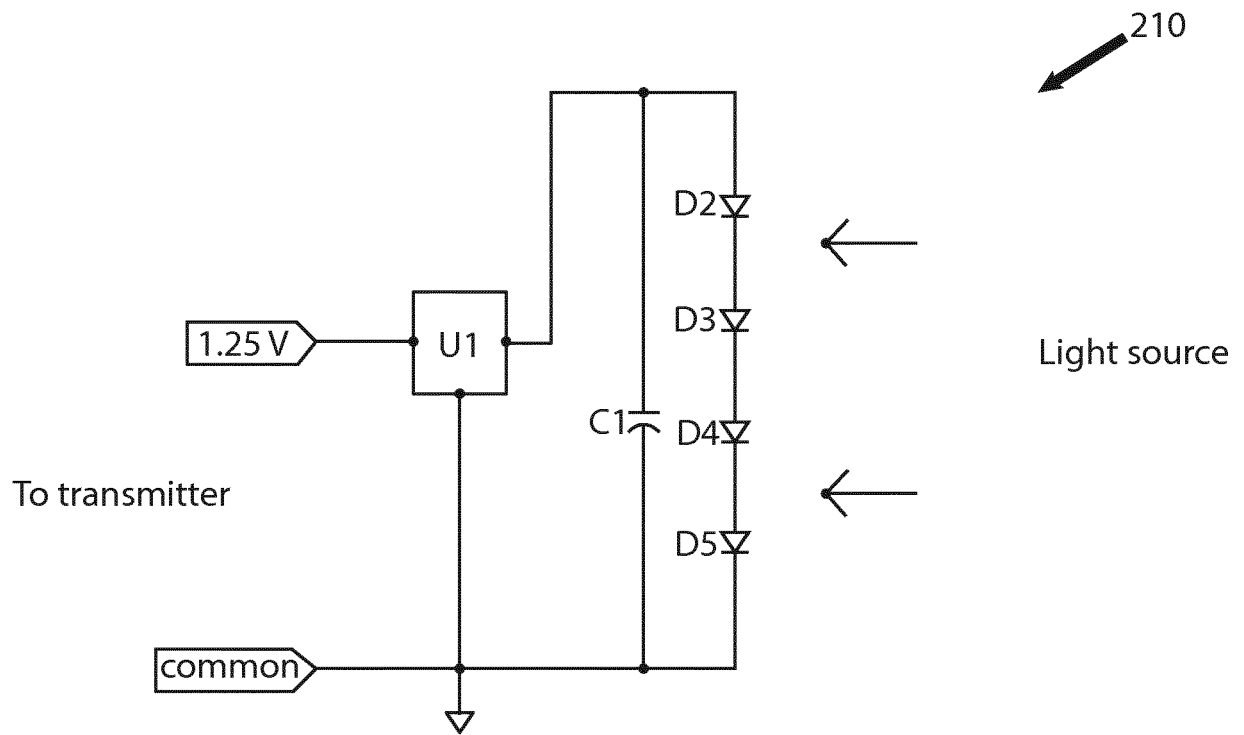
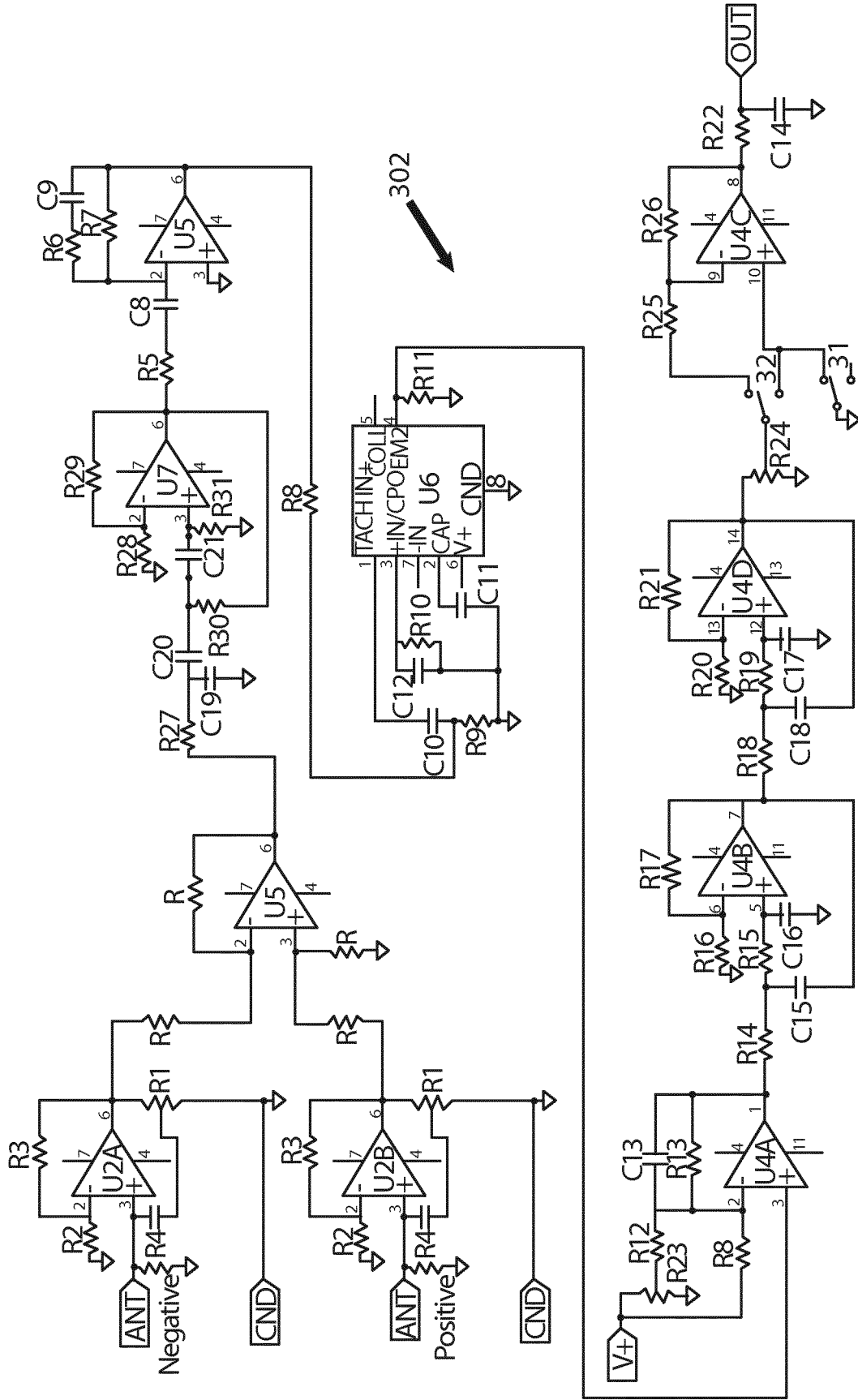


FIG. 2E



Configuration jumpers

FM receiver 2

FIG. 3A

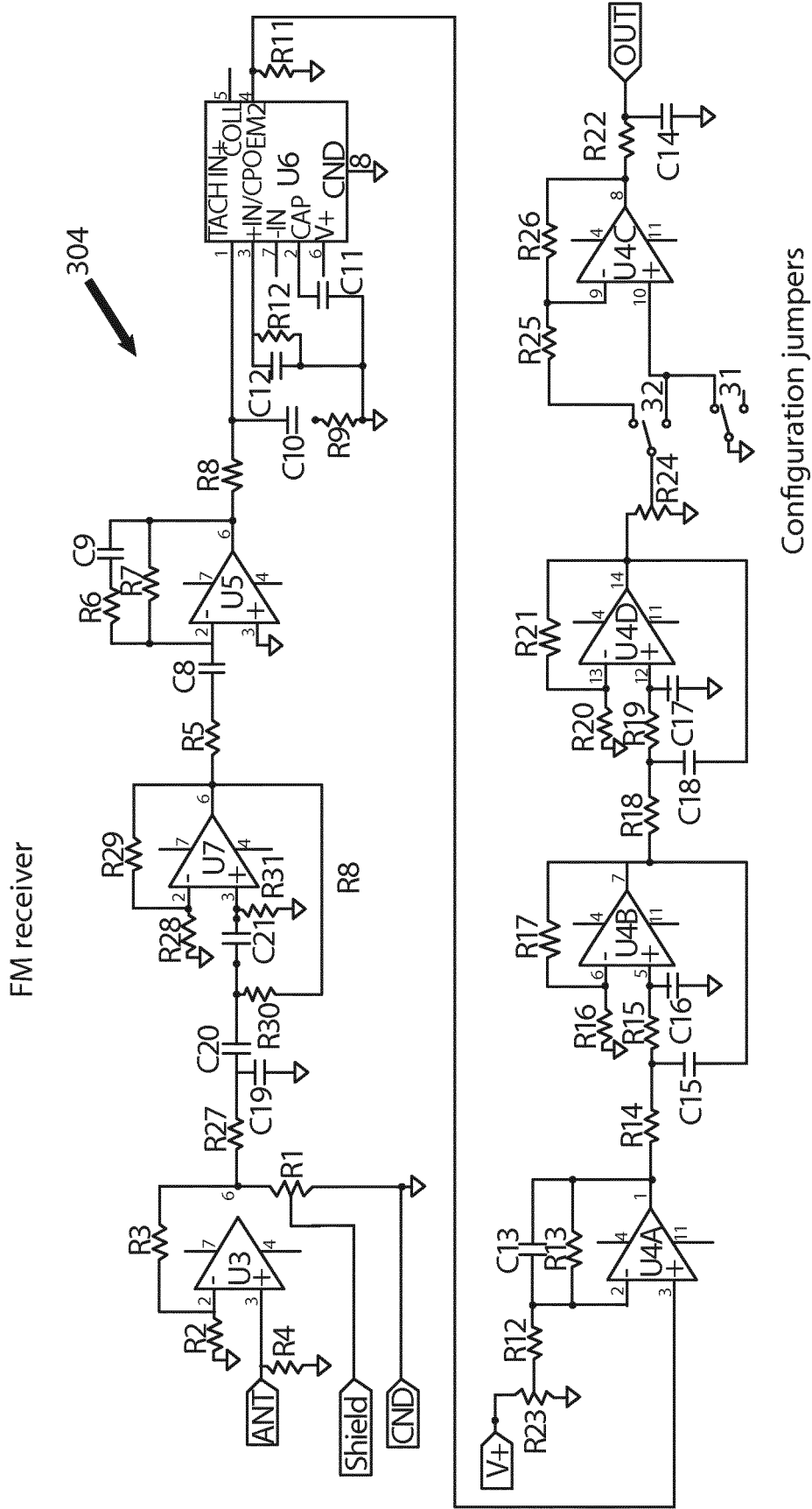


FIG. 3B

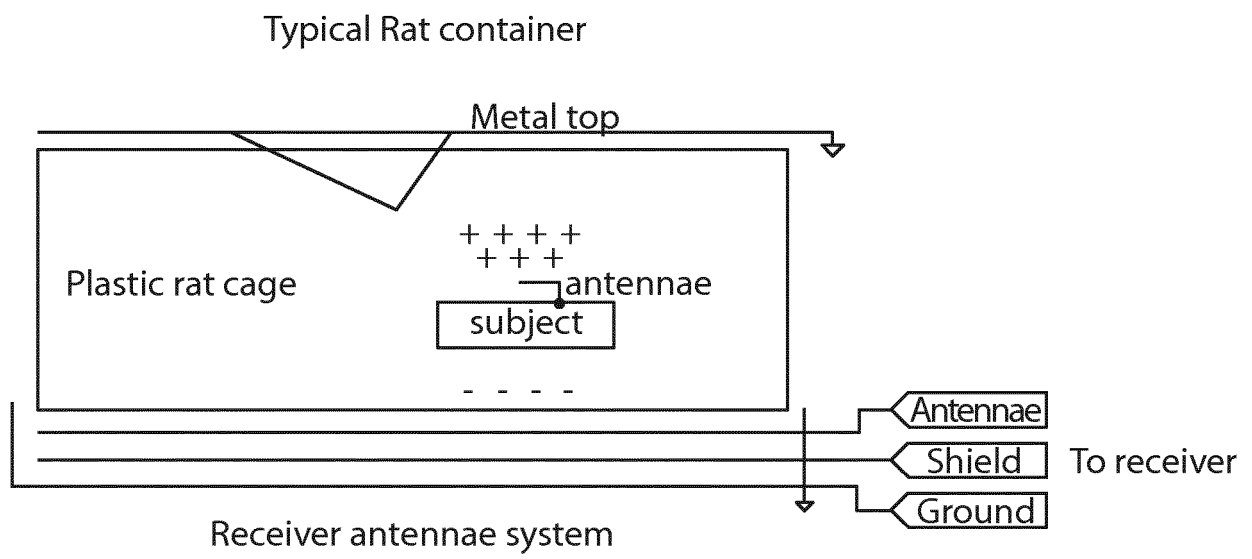


FIG. 4

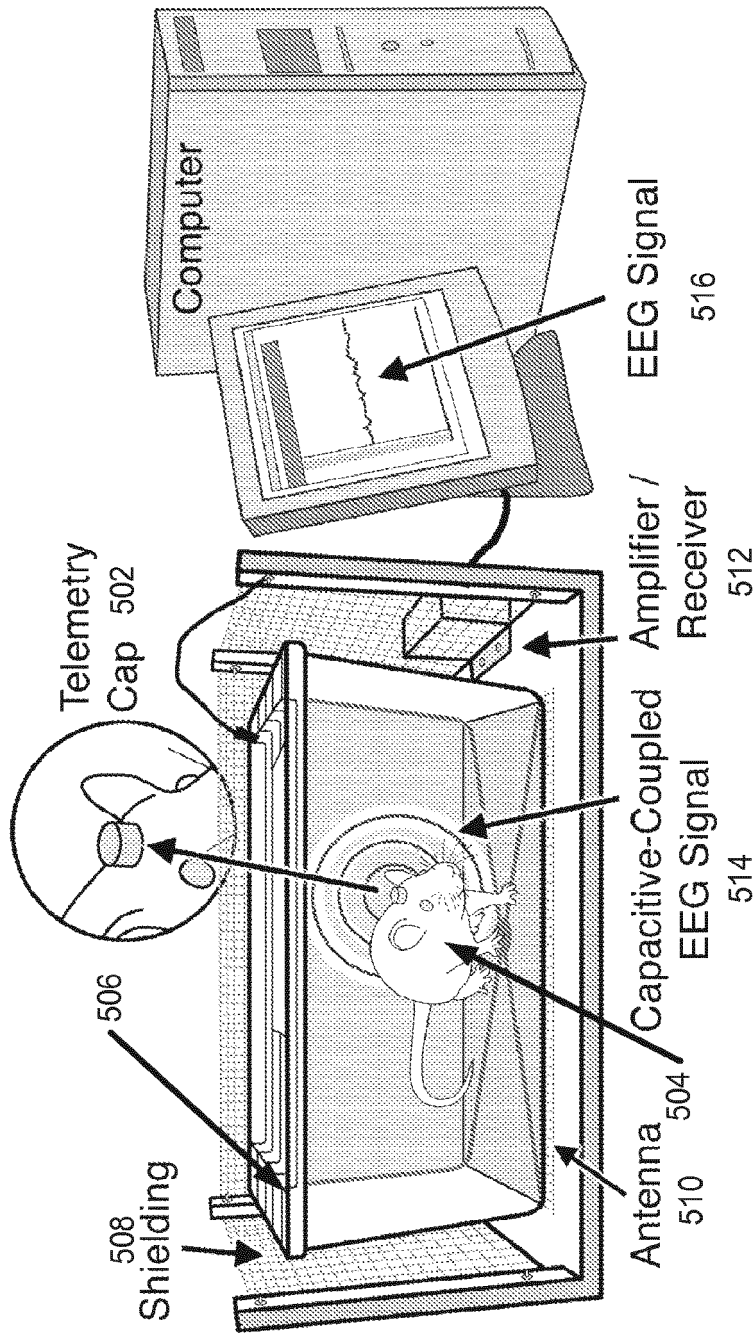


FIG. 5

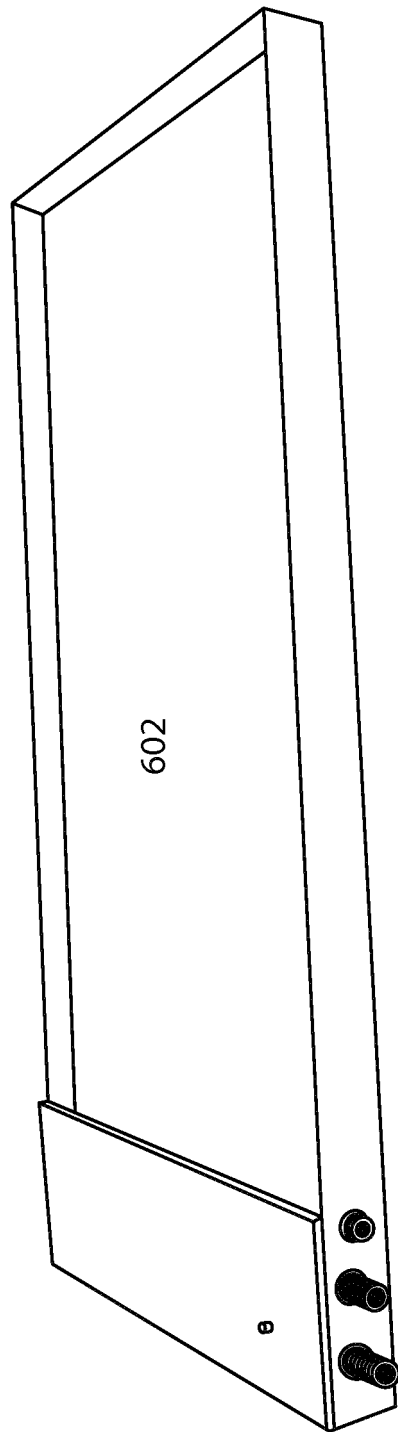


FIG. 6

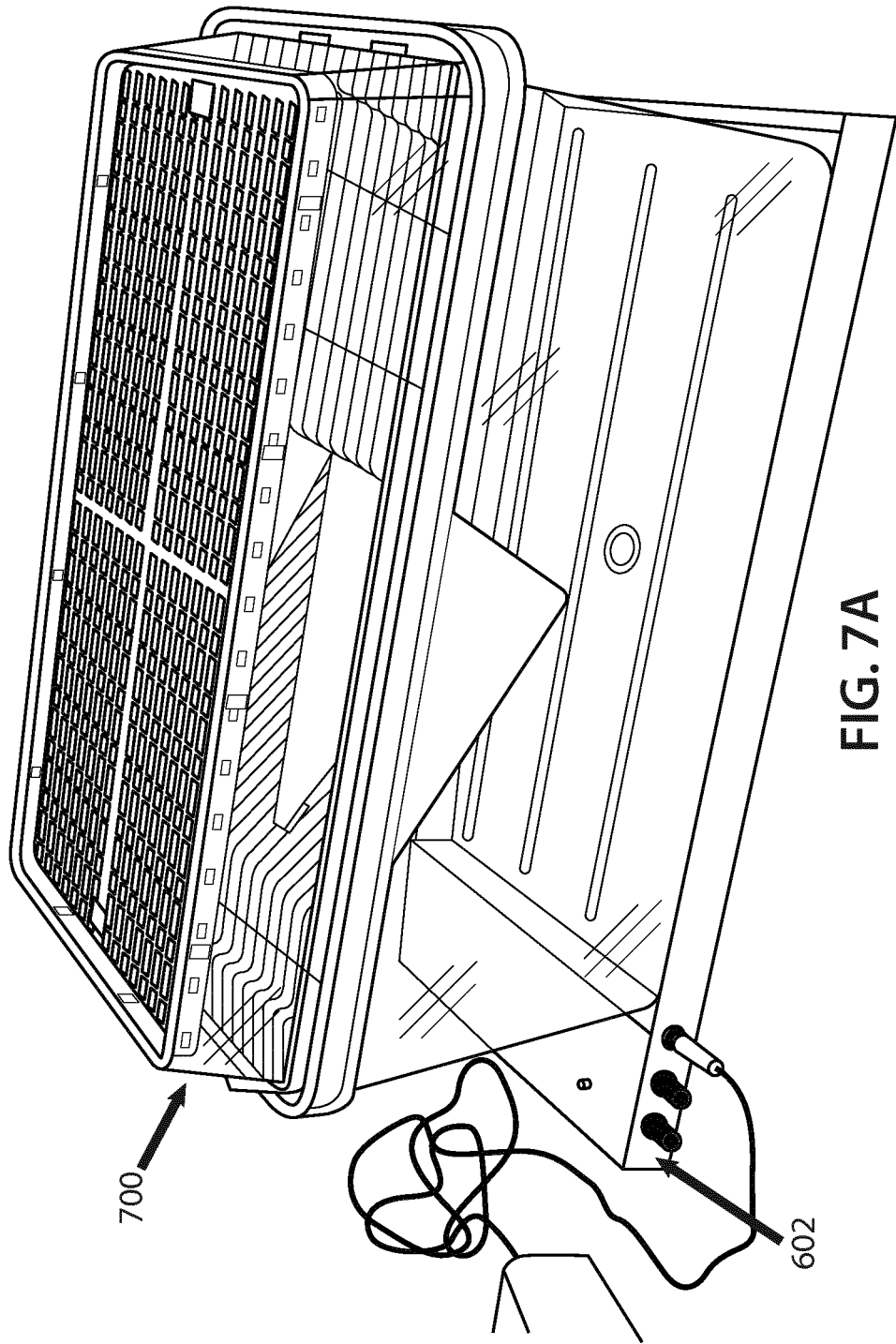


FIG. 7A

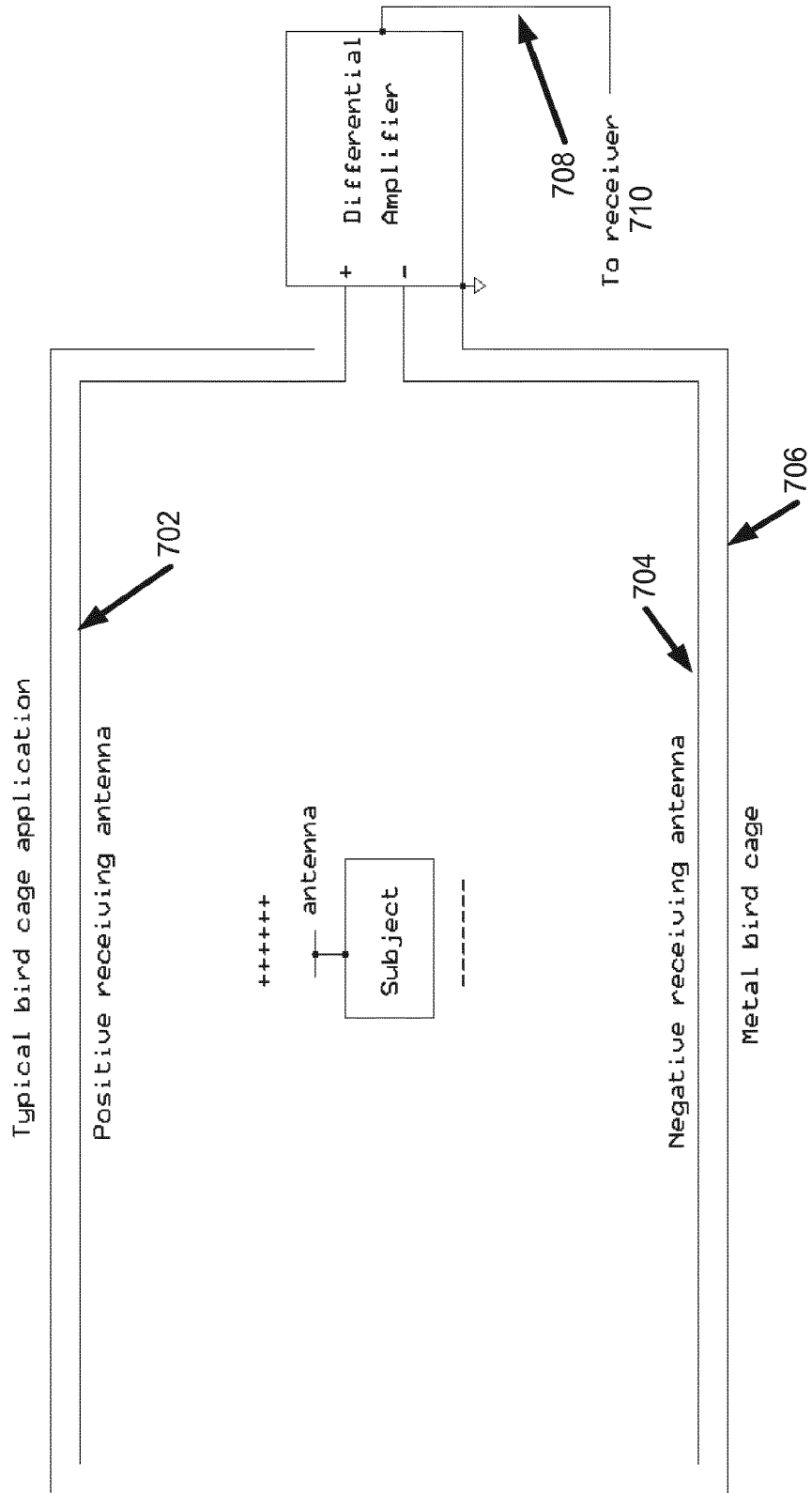


FIG. 7B

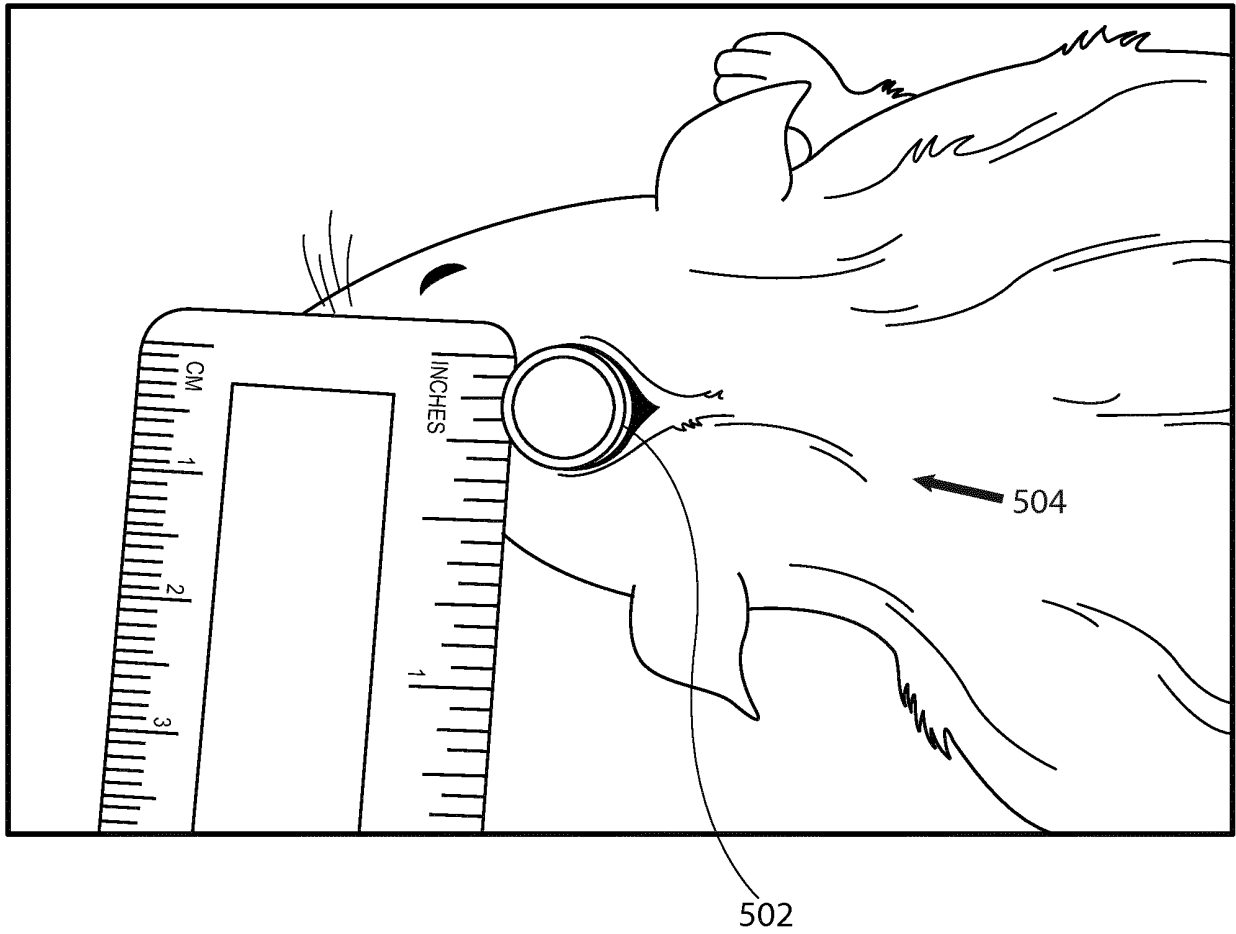


FIG. 8A

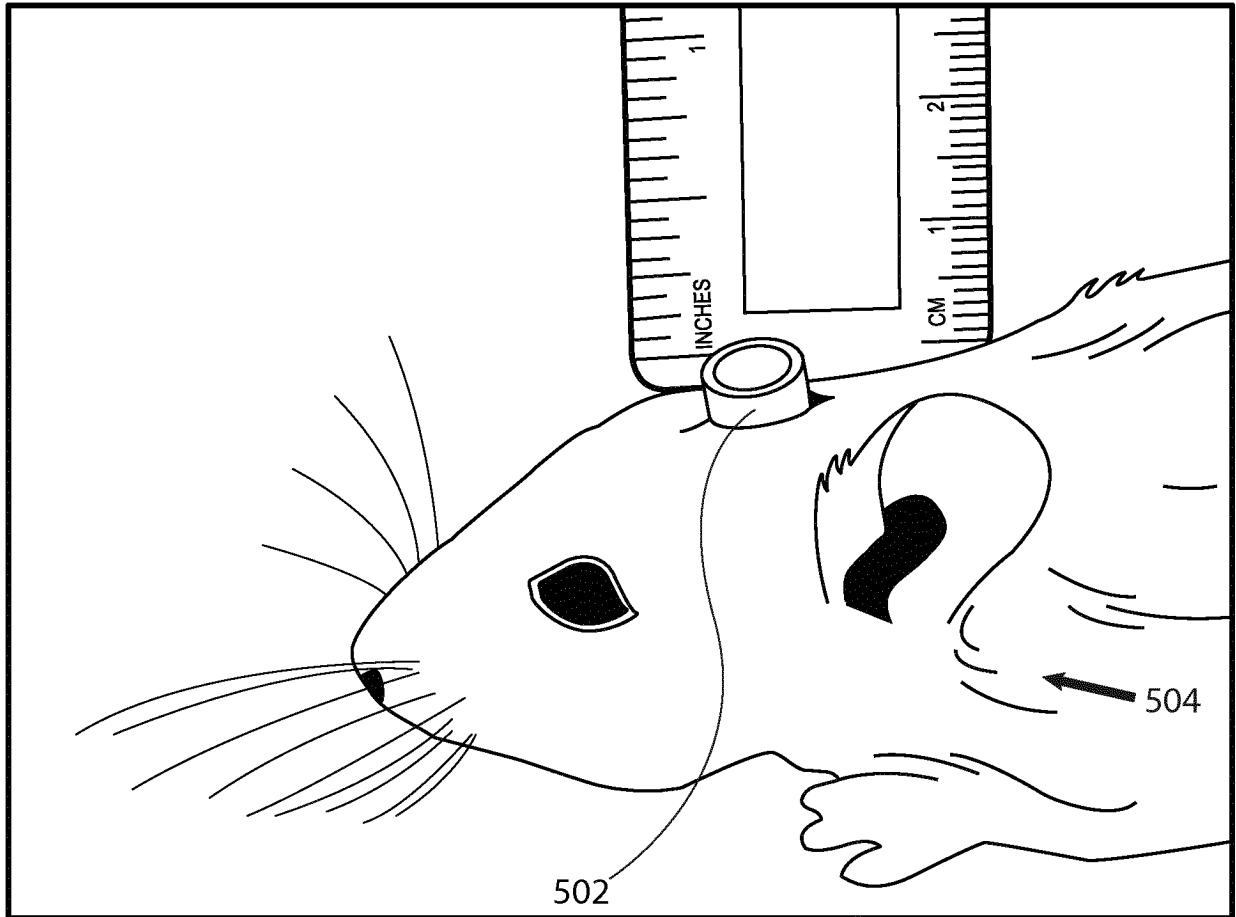


FIG. 8B

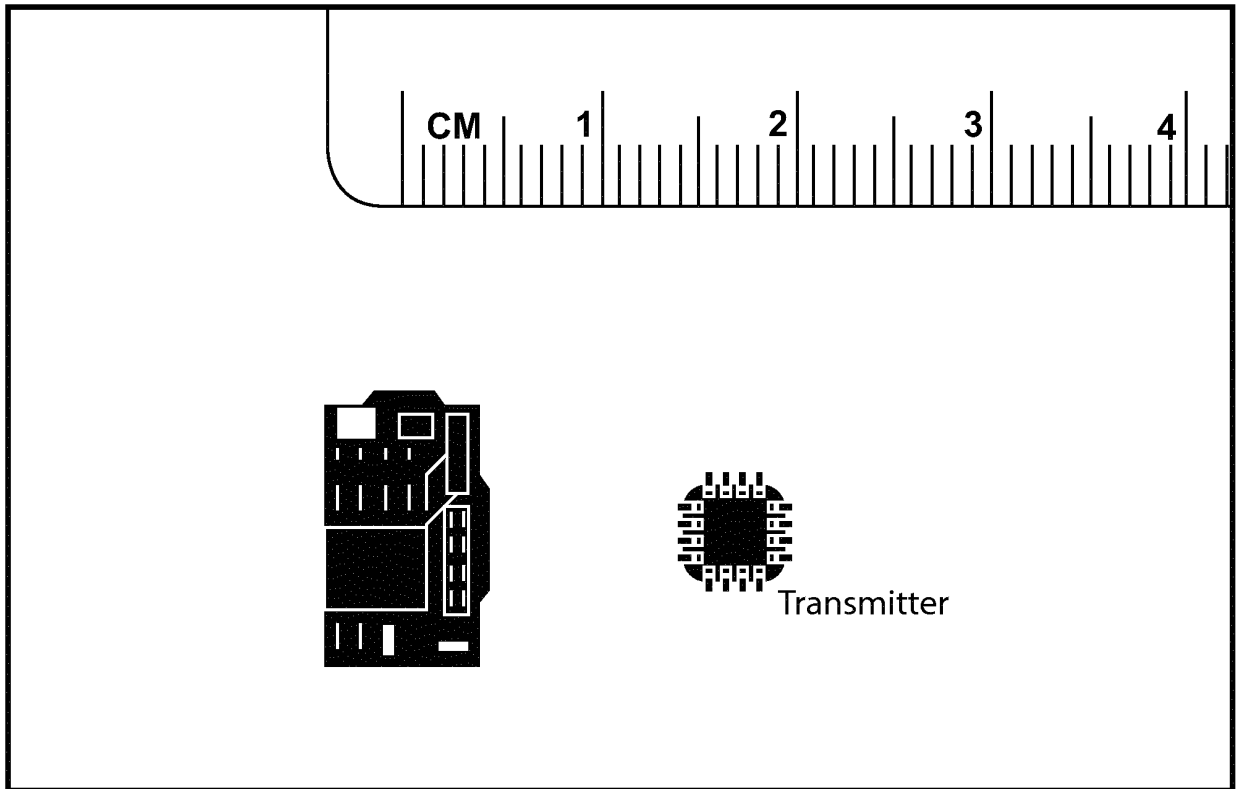


FIG. 9A

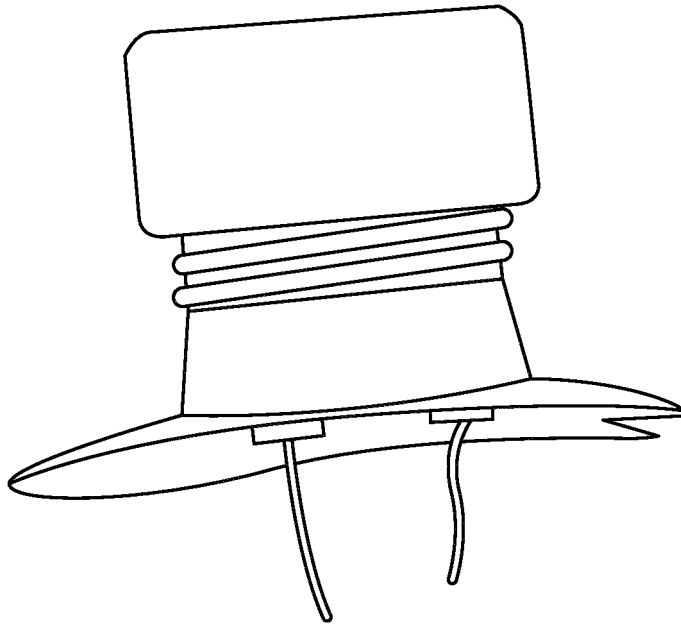


FIG. 9B

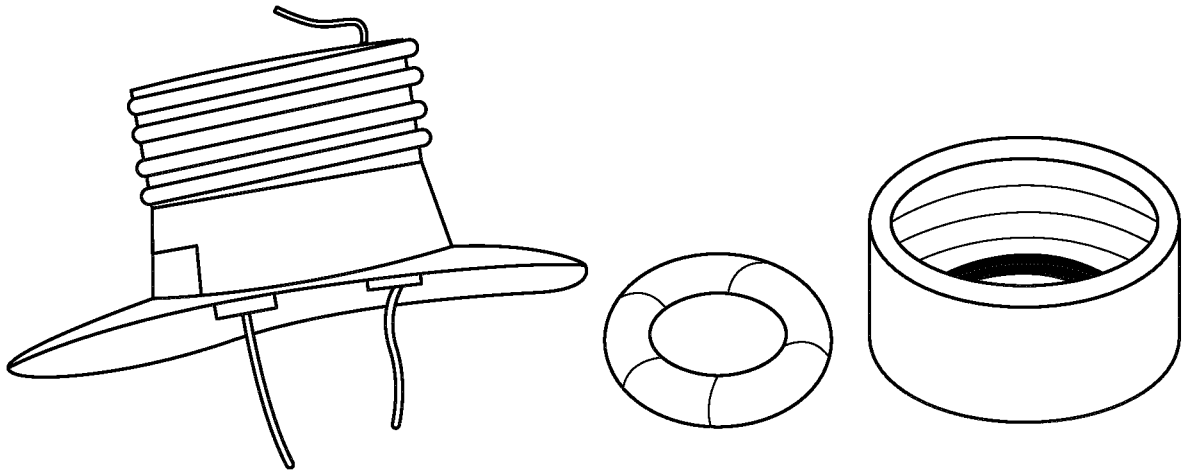


FIG. 9C

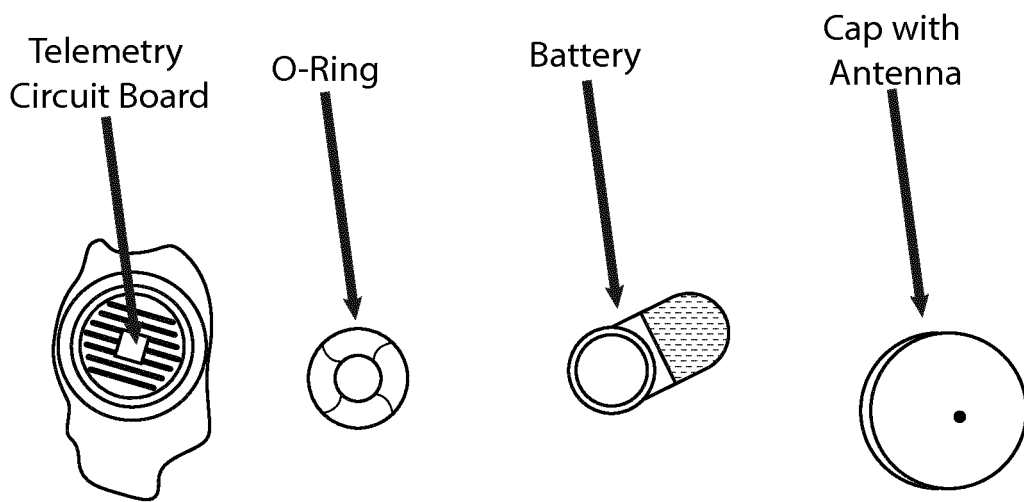


FIG. 9D

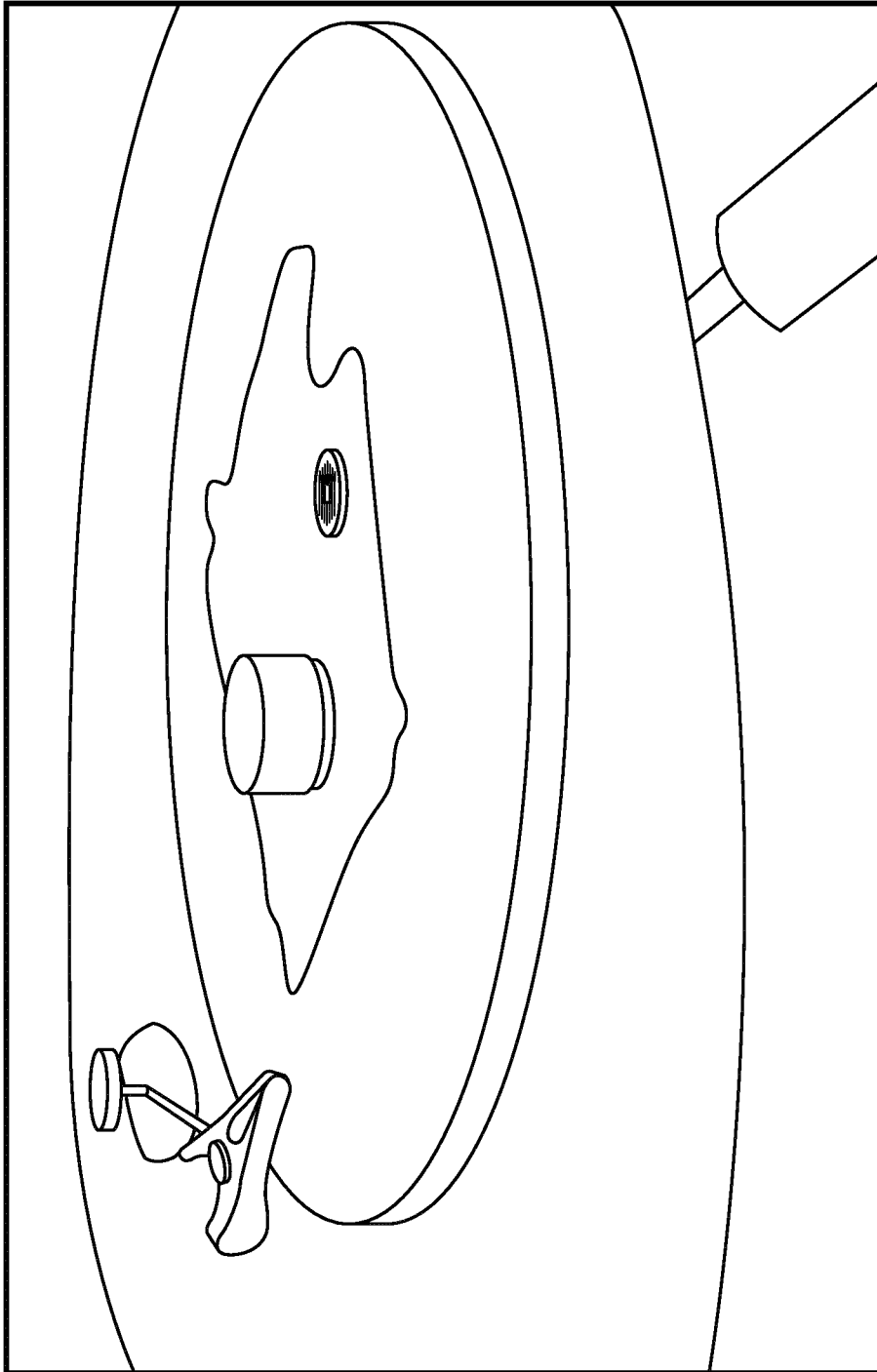


FIG. 10

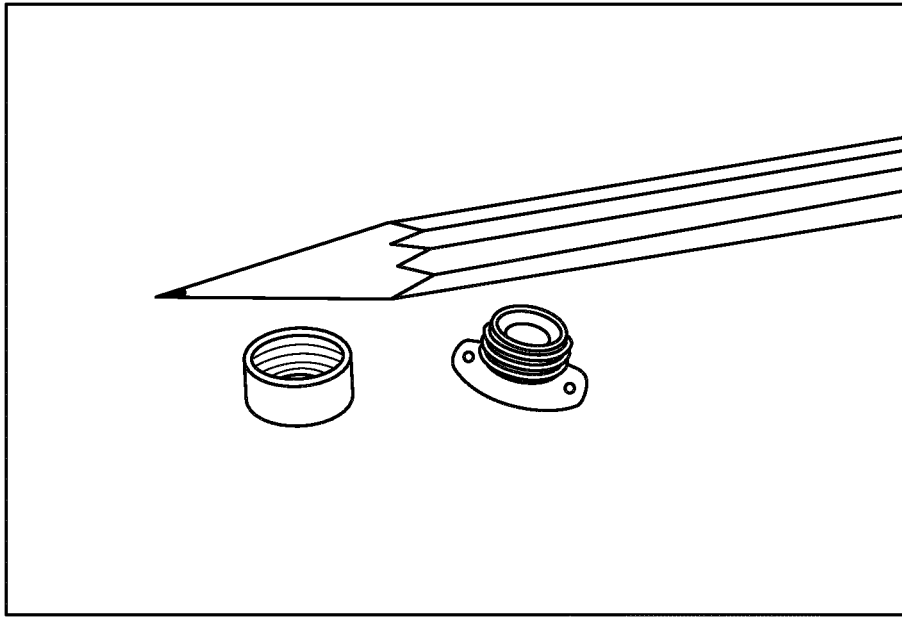


FIG. 11A

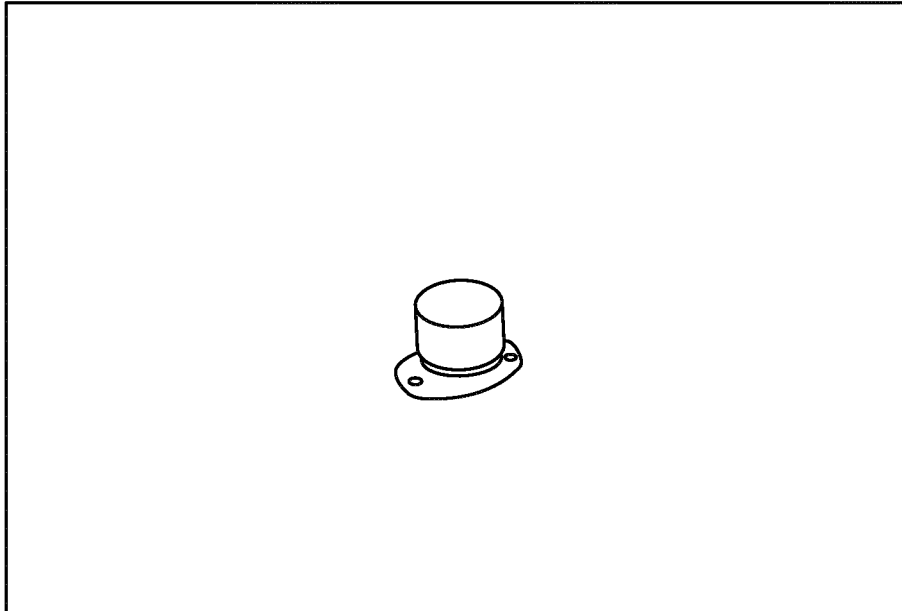


FIG. 11B

EEG Activity after Kainate Injections

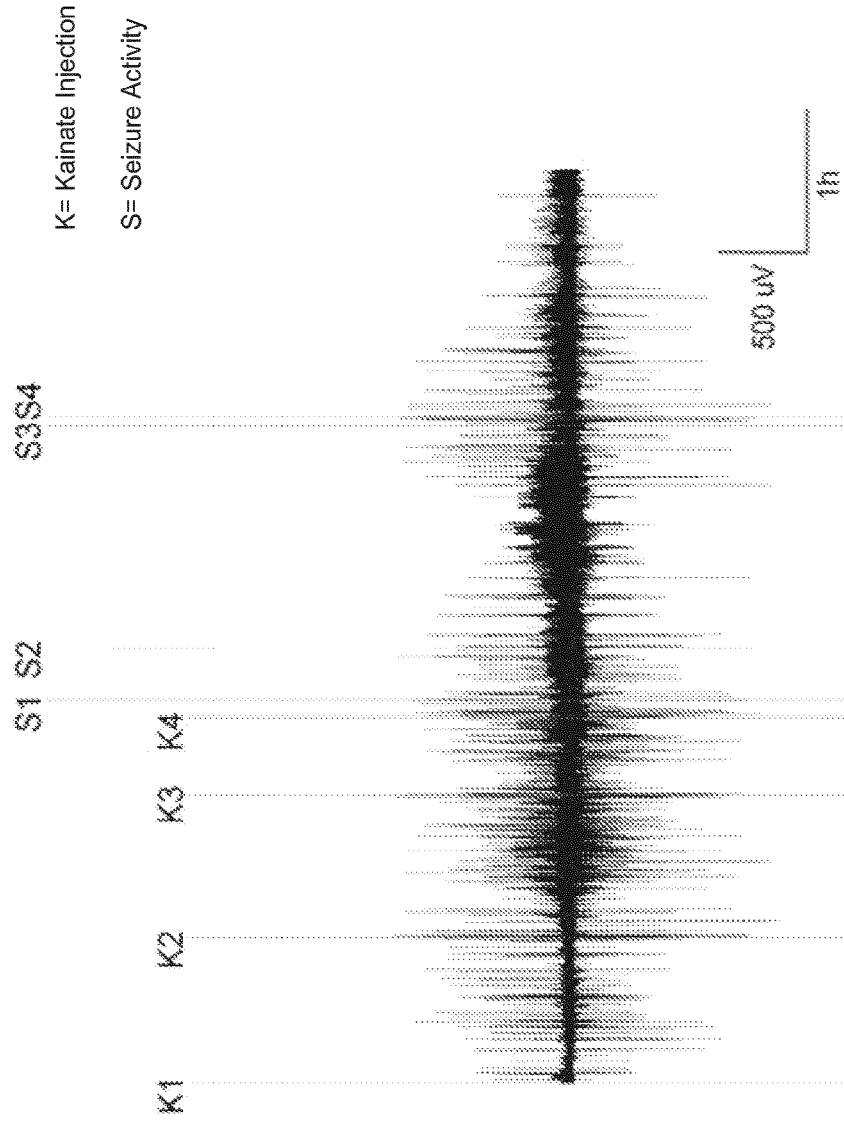


FIG. 12

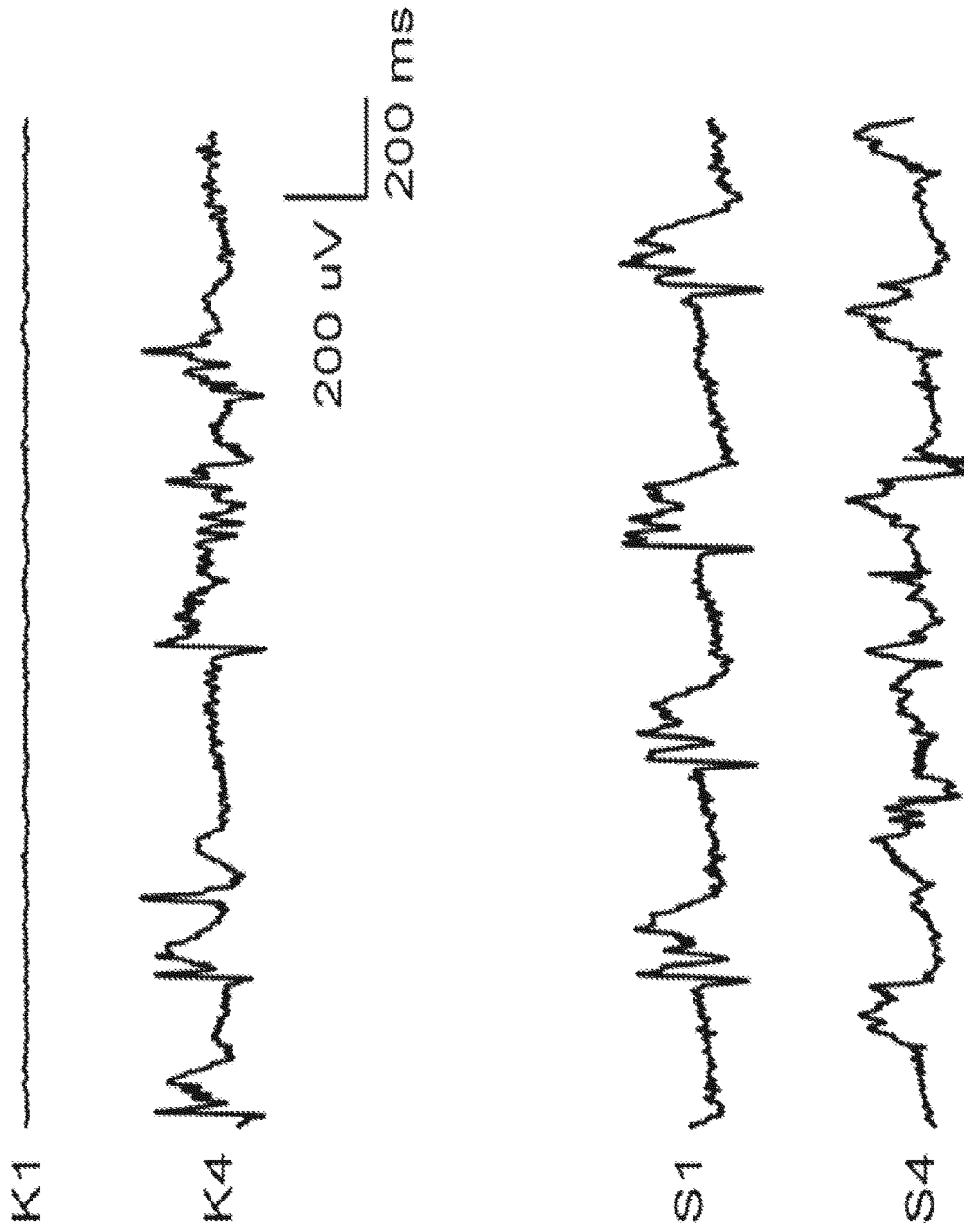


FIG. 13

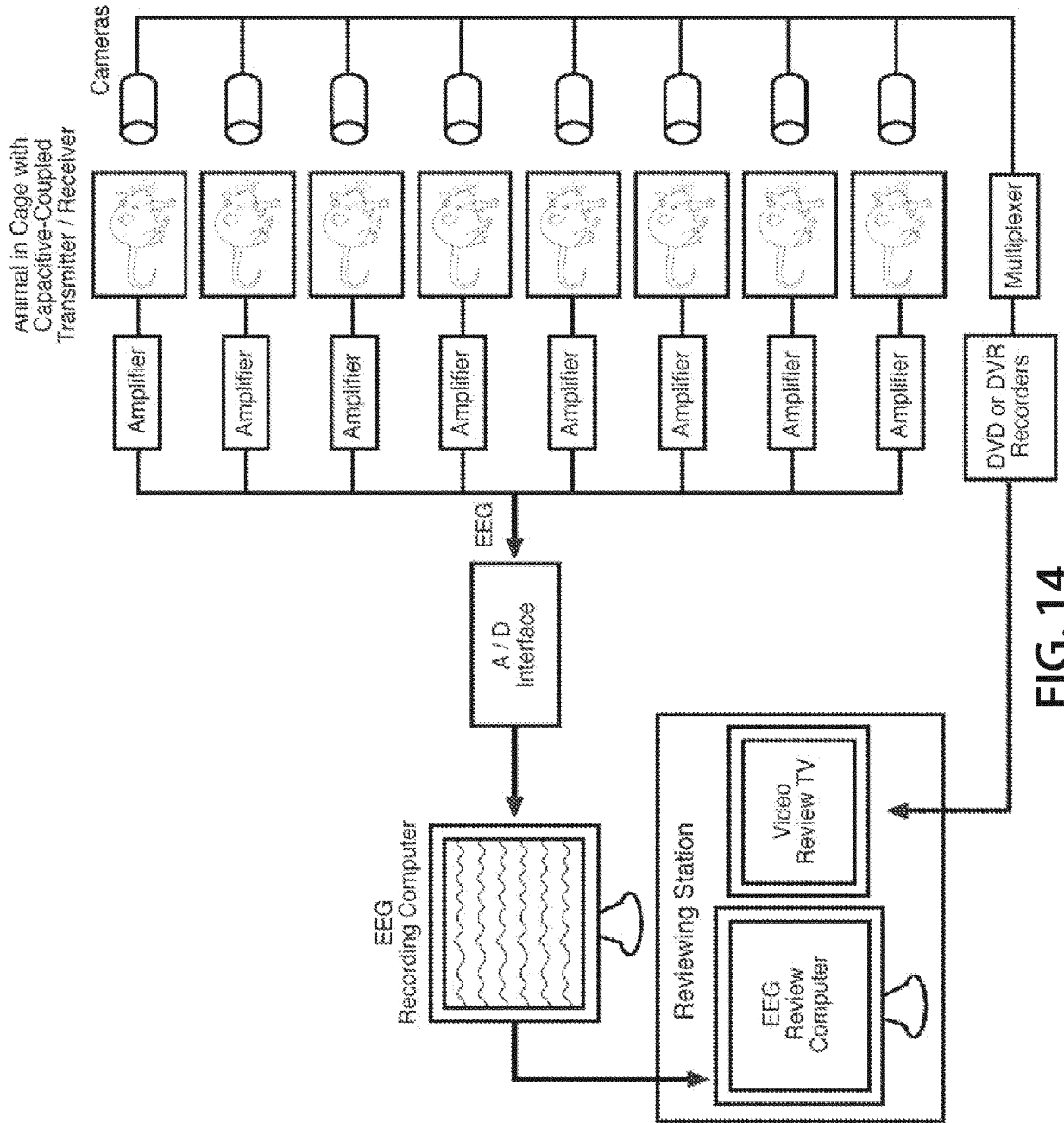


FIG. 14

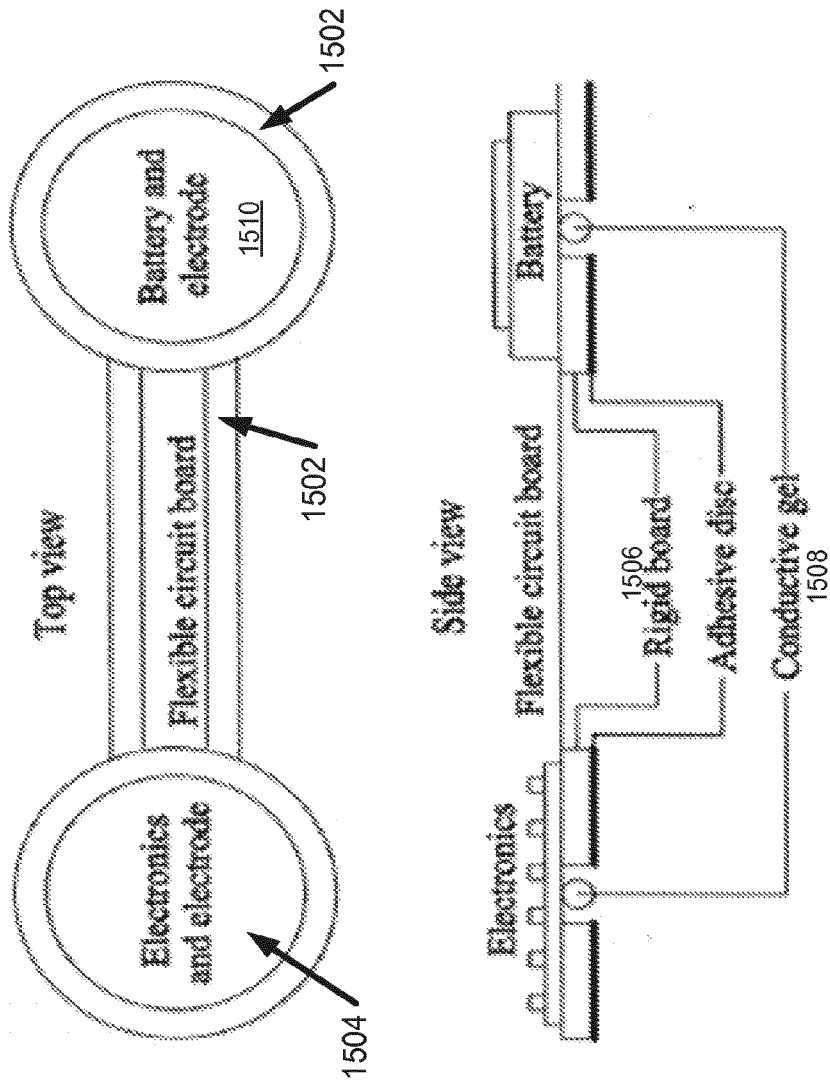


FIG. 15

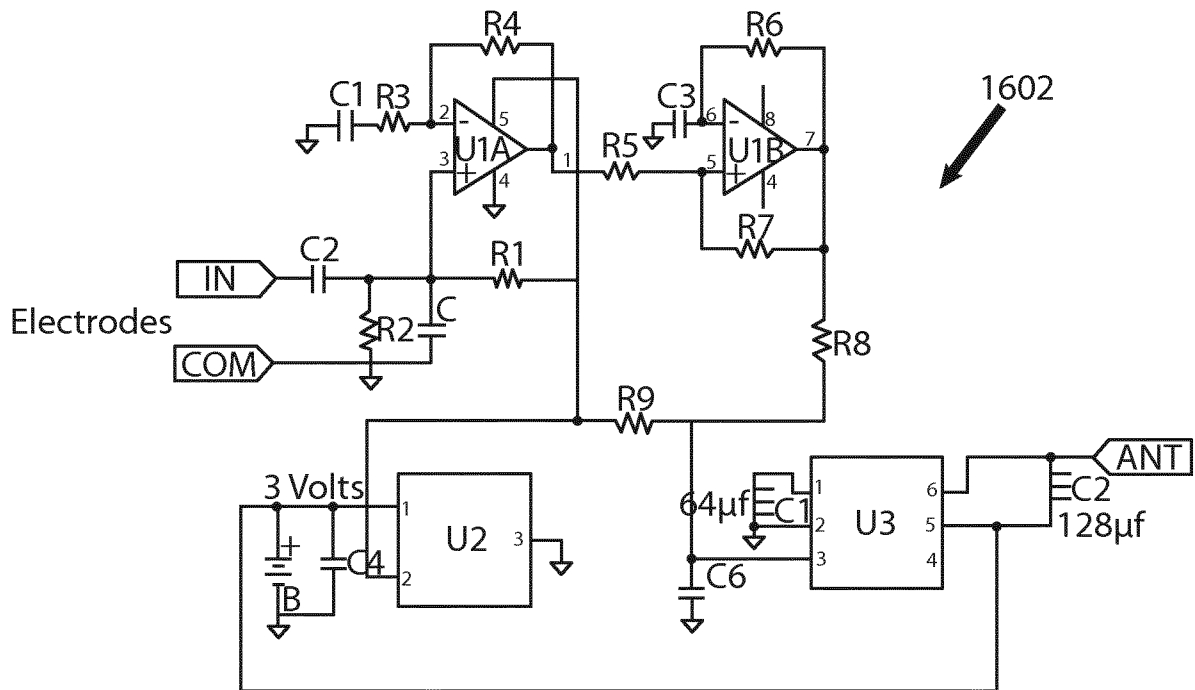


FIG. 16

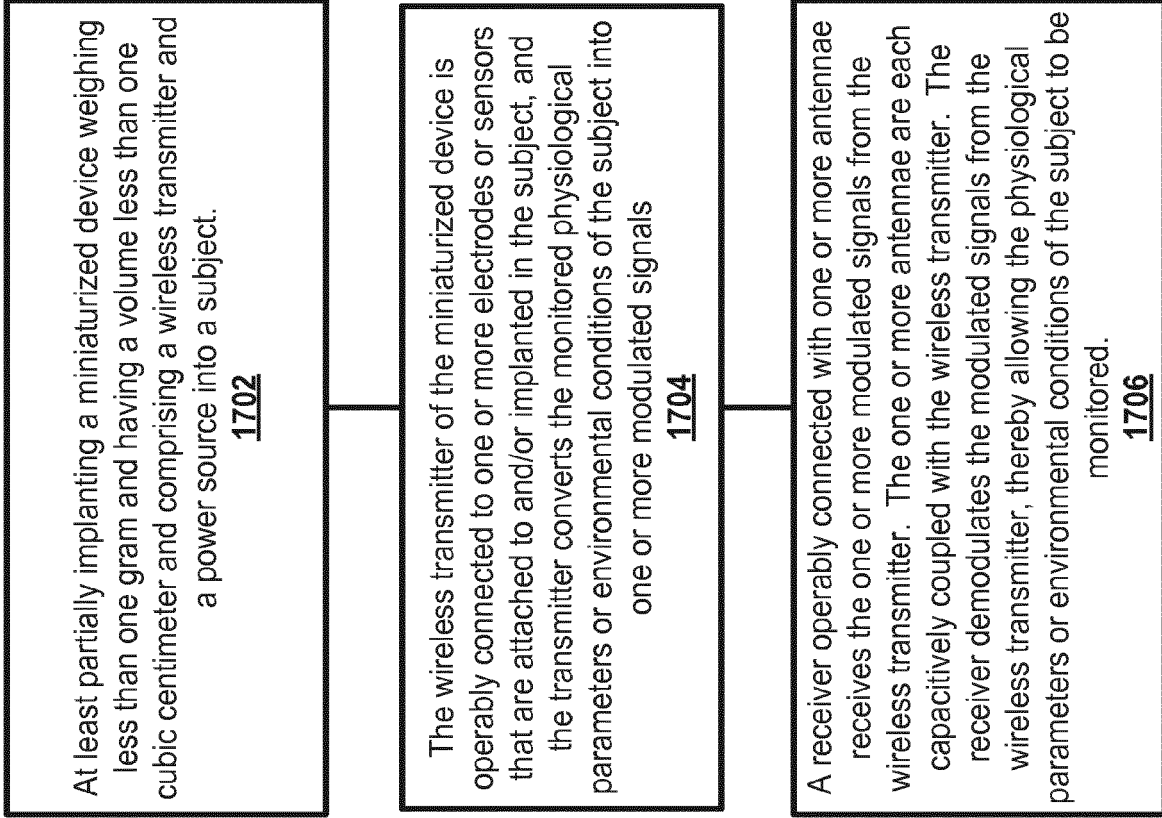


FIG. 17

REFERENCES CITED IN THE DESCRIPTION

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Patent documents cited in the description

- US 2006020300 A [0009]
- US 2006173259 A [0010]

专利名称(译)	微型无线生物医学遥测设备		
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摘要(译)

本文提供了微型无线生物医学遥测设备以及其使用的系统和方法的实施例。微型放大器和发射器允许记录来自小动物(如大鼠,小鼠和鸟类以及人类)的生理信号。该设备与电池一样位于外部,易于更换,因此手术并发症和其他问题得以最小化。