





# MEA2100-Lite-System

**USER MANUAL** 

# TABLE OF CONTENTS

Important Safety Advice
Guarantee and Liability
ABOUT THIS MANUAL
WELCOME TO THE MEA2100-Lite-System
The Headstage of the MEA2100-Lite-System
The Interface Board of the MEA2100-Lite-System7
SOFTWARE
System Requirements
Multi Channel Experimenter
MC_Rack
Setting up Cardio2D Software
Hardware
Hardware Configuration
MEA2100-Lite Interface Board (MCS-Lite-IFB)
MEA2100-Lite Headstage
Setting up the MEA2100-Lite-System25
Setting up the MEA
Temperature Controller TC01 / 02
Perfusion Cannula PH01
Troubleshooting
About Troubleshooting
MEA2100 Configuration Software
Main Window of the MCS Filter Configuration Program
Technical Support
MEA2100-Lite-System
Pin Layout
Changing Gold Spring Contacts
Contact Information

# IMPRINT

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#### **IMPORTANT SAFETY ADVICE**



Warning: Make sure to read the following advice prior to installation or use of the device and the software. If you do not fulfill all requirements stated below, this may lead to malfunctions or breakage of connected hardware, or even fatal injuries.



Warning: Always obey the rules of local regulations and laws. Only qualified personnel should be allowed to perform laboratory work. Work according to good laboratory practice to obtain best results and to minimize risks.

The product has been built to the state of the art and in accordance with recognized safety engineering rules. The device may only be used for

- its intended purpose;
- be used when in a perfect condition.

Improper use could lead to serious, even fatal injuries to the user or third parties and damage to the device itself or other material damage.



Warning: The device and the software are not intended for medical uses and must not be used on humans. MCS assumes no responsibility in any case of contravention.

Malfunctions which could impair safety should be rectified immediately.

#### Grounding

This product is grounded through the grounding conductor on the power cord. To avoid electric shock, the grounding conductor must be connected to earth.

#### **Orient the Equipment Properly**

Do not orient the equipment so that it is difficult to manage the disconnection device.

#### Explanation of the Symbol used



DC, direct current

#### **High Voltage**

Electrical cords must be properly laid and installed. The length and quality of the cords must be in accordance with local provisions. Only qualified technicians may work on the electrical system. It is essential that the accident prevention regulations and those of the employers' liability associations are observed.

Each time before starting up, make sure that the power supply agrees with the specifications of the product.

- Check the power cord for damage each time the site is changed. Damaged power cords should be replaced immediately and may never be reused.
- Check the leads for damage. Damaged leads should be replaced immediately and may never be reused.
- Do not try to insert anything sharp or metallic into the vents or the case.
- Liquids may cause short circuits or other damage. Always keep the device and the power cords dry. Do not handle it with wet hands.

#### **Requirements for the Installation**

Make sure that the device is not exposed to direct sunlight. Do not place anything on top of the device, and do not place it on top of another heat producing device, so that the air can circulate freely.

#### **Guarantee and Liability**

The general conditions of sale and delivery of Multi Channel Systems MCS GmbH always apply.

The operator will receive these no later than on conclusion of the contract. Multi Channel Systems MCS GmbH makes no guarantee as to the accuracy of any and all tests and data generated by the use of the device or the software. It is up to the user to use good laboratory practice to establish the validity of his / her findings.

Guarantee and liability claims in the event of injury or material damage are excluded when they are the result of one of the following.

- Improper use of the device.
- Improper installation, commissioning, operation, or maintenance of the device.
- Operating the device when the safety and protective devices are defective and / or inoperable.
- Non-observance of the instructions in the manual with regard to transport, storage, installation, commissioning, operation, or maintenance of the device.
- Unauthorized structural alterations to the device.
- Unauthorized modifications to the system settings.
- Inadequate monitoring of device components subject to wear.
- Improperly executed and unauthorized repairs.
- Unauthorized opening of the device or its components.
- Catastrophic events due to the effect of foreign bodies or acts of God.

# ABOUT THIS MANUAL

#### ABOUT THIS MANUAL

This short manual is an addendum to the elaborate MEA2100 Manual. It is strongly recommended to read the MEA2100 manual first, if you utilize the MEA Signal Generator for introducing yourself to the MEA2100-System. Being already familiar with MEA2100-System, the MEA-SG will be easy to mount and to handle.

It is assumed that you already have a basic understanding of technical and software terms.

If you are using the device for the first time, please read the important safety advice before installing the hardware and software, where you will find important information about the installation and first steps.

The device and the software are part of an ongoing developmental process. Please understand that the provided documentation is not always up to date. The latest information can be found in the help.

Check also the MCS web site (www.multichannelsystems.com) for downloading up-to-date manuals and help files.

#### WELCOME TO THE MEA2100-LITE-SYSTEM



The MEA2100-Lite-System is a compact and portable stand-alone system for extracellular recordings from microelectrode arrays (MEAs) with integrated amplification, data acquisition, online signal processing, integrated stimulus generator as well as temperature control. The MEA2100-Lite-System consists of two main devices: The interface board with integrated signal processor and the headstage equipped with amplifier, A / D converter and stimulus generator.

It is a flexible system, using a headstage that will fit on inverted and upright microscopes. The MEA2100-Lite-System acquires data from up to 60 channels per headstage and a sampling frequency of up to 32 kHz per channel with MC\_Rack, up to 25 kHz with Multi Channel Experimenter software. Four additional analog IN channels are available and 16 bit digital IN / OUT channels. The digitized electrode data is transmitted to the connected interface board via eSATAp cable. The connection between interface board and computer uses a super speed USB 3.0 connection. Thus, it is possible to use any computer as a data acquisition computer, also a laptop.

Data is recorded, stimulated, analyzed, graphed and reviewed with the powerful and easy-to-use Multi Channel Experimenter software or the MC\_Rack program from Multi Channel Systems MCS GmbH. The control functions of the internal stimulator are also integrated into the software. The data format can be converted in other standard formats with the software Multi Channel Data Manager or MC\_DataTool.

#### The Headstage of the MEA2100-Lite-System

The MEA2100-Lite-System is equipped with one headstage for microelectrode arrays with 60 electrodes.

The data acquisition integrated in the headstage supports 60 electrode channels. The sampling rate, signal range and bandwidth can be adjusted via software control and is suitable for a broad range of applications, from single unit spike recordings to field potentials from whole heart preparations.

A special advantage are the three integrated stimulus generator units, which deliver monophasic or biphasic rectangular pulses, as single pulses or pulse trains. You can choose between current or voltage stimulation and you can select each MEA electrode for stimulation within the data acquisition software.

As usual in MEA-Systems, the MEA2100-Lite-System features a heating element with a Pt100 temperature sensor. If you connect a temperature controller TC02 to the heating element via D-Sub 9 connector, the heating element guarantees constant temperature conditions for the biological sample on the chip. A perfusion system with heating is available for the MEA2100-Lite as optional accessory.

All MEA types with 60 channels can be used. A modified base plate is available for the headstage to be able to work with perforated MEAs also. Because of the compact designed headstage you can use most inverted or upright microscopes, for the optical control of the sample on the MEA chip.

#### The Interface Board of the MEA2100-Lite-System

The Digital IN / OUT connector (68-pin MCS standard connector) for 16 digital in- and output bits guarantees the communication with other devices. You can use the digital TTL inputs, for example, for synchronizing the recording in MEA2100-Lite-System with other systems. The digital TTL outputs can be used for triggering other devices, for example, an imaging setup.

Four digital inputs and four digital outputs (bit 0 to bit 3) are separately available via Lemo connectors on the front of the interface board. The up to four analog inputs (serial connector) are intended for recording additional information from external devices, for example, for recording patch clamp in parallel to the MEA recording, for monitoring the temperature, or for fluidic control.

Two of the analog inputs are separately available via Lemo connectors on the front of the interface board. You can connect an audio system to the Audio Out (3.5 mm phone jack) to make electrical activity audible. This audio output is in real-time. Headphones or a speaker can be connected directly to the Audio Out. Two channels at a time can be converted into sound.

#### SOFTWARE

#### System Requirements

For the Multi Channel Experimenter software Microsoft Windows <sup>®</sup> 10 or 8 operating systems is required. English and German versions supported. Other language versions may lead to software errors.

One of the following Microsoft Windows operating systems is required for MC\_Rack: Windows 10, 8 or 7 (32 or 64 bit), English and German versions supported. One of the following Microsoft Windows operating systems is required for Cardio2D: Windows 8 or 7 (32 or 64 bit), English and German versions supported.

Please note that there are sometimes hardware incompatibilities of the MEA2100-Lite-System and computer components; or that an inappropriate computer power supply may lead to artifact signals.

Please contact your local retailer for more information on recommended computer hardware.

Important: You need to have installed the latest MEA2100-Lite-System driver to operate the MEA2100-Lite-System, which is automatically installed with MC\_Rack. The installation may be invalid if the MEA2100-Lite-System does not respond. Please contact Multi Channel Systems or your local retailer in this case.

#### **Recommended BIOS settings**

#### Recommended operating system settings

The following automatic services of the Windows operating system interfere with the data storage on the hard disk and can lead to severe performance limits in the software. These routines were designed for use on office computers, but are not very useful for a data acquisition computer.

- $\rightarrow$  Turn off Screensaver.
- → Turn off Windows Indexing Service for all local disks.
- → Turn off Optimize hard disk when idle (automatic disk fragmentation).
- → Manual management of automatic Windows Update.
- → Power Options: Power scheme: Never turn off monitor, hard disk and system standby.

It is also not recommended to run any applications in the background when using the data acquisition software. Remove all applications from the Autostart folder. Be careful when using a Virus Scanner. These programs are known to disturb the data acquisition software, and even data loss may occur. When using a MEA2100-Lite-System it is recommended to connect a high performance computer with a separate hard disk for program files and data storage. The provided possibility to use up to 60 channels with up to 32 kHz sampling rate needs high memory capacity. Please remove data and defragment the hard disk regularly to ensure optimal performance.

#### **Multi Channel Experimenter**

Please read the manual "Multi Channel Experimenter" for detailed information!

#### **Building a Rack**

The "Multi Channel Experimenter" software operates with virtual instruments, which can be combined and saved as so called "Experiments" for later use. When opening the software, all available instruments are shown as blue icons on the left hand side of the screen. Each icon can be dragged and dropped in the main window. Most instruments have color coded data ports. Ports on the upper side represent data input to the instrument, ports on the lower side represent data outputs of the instrument. Only ports with matching colors can be connected by drawing a connection line between a data output and one or more data inputs. Data flows along those lines from the data source through the instruments. An instrument without connected data inputs is not functional.



Clicking the button of any instrument with the right mouse provides three options: "Delete Instrument" for deleting the tool, "Edit Instrument Name" for changing the name and "Show Info" for information about the instrument. It is not possible to loop instruments.



Hovering the mouse pointer above an output port will give you a "Tooltip" with the data type the port provides. A right click on a data connection allows to delete the connection. A right click on an instrument brings up a menu which allows to delete or rename the instrument.

### Instrument Control Window

A double click on each instrument opens a tab with the control interface for that instrument. Each instrument has its' own control window tab. The tab of the selected data source always shows the raw data. Most instruments can be selected more than once and be connected in parallel or in series. Exceptions are the data source, the recorder, and the stimulator, which can be selected only once. Once a configuration of instruments has been designed, it can be saved for later use. Such a configuration is called "Experiment" and can be saved and loaded from the main menu bar. The file extension for an experiment file is \*.mse.



The connecting lines between the instruments represent the flow of data. In the example above, there is a stream of raw data coming from the data source MEA2100 into the recorder directly, and will consequently be recorded in the file. The same raw data stream is processed through a 200 Hz high pass filter, and the filtered data is further processed by the Cross-Channel Tool. The data processed by those two instruments is then connected to the recorder, and will generate a second data stream with filtered AND referenced data in the recorded file. A third data stream will be the raw data filtered by a 50 Hz notch filter.

Instruments operate as independent units. They process the data from the respective input port and generate one or more data streams in the output port(s). Settings have to be done separately for each instrument in the respective control window tab.

# Multiple Data Sources, Multiple Software Instances

If multiple data sources are connected to the computer, they are indicated as separate icons in the Multi Channel Experimenter software. In case of a MEA2100-HS2x60, each individual MEA slot is a separate data source. Only one data source can be used in one instance of the software. Multiple instances can be opened simultaneously to control different data sources independently, for example two instances to control the two MEAs of a MEA2100-HS2x60. The active data source is shown in a light blue color. Data sources already occupied by another instance of the software are shown in grey. Not selected but available data sources are darker blue.



# SOFTWARE

#### Data Types

The types of output data occurring in the MEA2100-Lite-System, as indicated by the color-coded ports are:

Blue: Electrode data, raw or filtered, Cross-Channel data. These data types have the same structure and the same compatibility with instruments, hence they use the same data port color.

Green: Auxiliary analog data from the additional analog inputs on the Interface Board.

Red: Digital data, the 16 bit digital data stream from the Interface Board.

Pink: Trigger events.

#### File Types

The types of files generated with the Multi Channel Experimenter and the MEA2100-Lite-System are:

\*.mse: Experiment file, a configuration of instruments. Created manually by the user.

\*.msrd: Data file, generated by the Recorder, contains the raw data of all data streams.

\*.msrs: Settings file, generated by the Recorder, must also be present to open a data file in the Multi Channel Analyzer.

\*.html: Instrument settings, one \*.html file is generated by the Recorder with each data file. Each \*.html is linked to the \*.xml files of the individual instruments.

\*. xml: One \*.xml per instrument is generated for each recording. It contains all settings of the respective instrument. Stimulation patterns in the Stimulator instrument can also be saved as \*.xml files by the user.

Please read the manual "Multi Channel Experimenter" for detailed information!

### MC\_Rack

For operating the MEA2100-Lite-System by MC\_Rack software, please refer to the MC\_Rack manual.

# **Data Source Setup**

Open the option "Data Source Setup" in "File" menu.

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Amp	A MEA
MEA2100H-60	MEA 8x8 200/30

### Hardware Settings for the MEA2100-Lite-System

Add the MEA2100-Lite-System to the rack. When using the MEA2100-Lite-System, the "Hardware" tab of the MC\_Rack software is modified. Please see also chapter "Defining a Data Source". In the "Hardware" tab of the Data Source, the gain setting must be adjusted to ensure a correct scaling of the data in MC\_Rack. The gain is a fixed hardware property, while sampling frequency and input voltage range of the data acquisition board can be selected according to the demands of the experiment.

III Recorder ■ MEA2100-Lite (S/NES0002-A) ■ Display 1	Rack         Hardware Info         Audio         Audio           Stratigs         Scalage         -625 mV         -           Siguta Volapa França         -625 mV to +625 mV         -         -           Angilare Gan         500         Errein         -           Ipent Volapa França         -         -         -         -           Dark Acquistion Bread         -         -         -         -           Samplag Françaver,         XXX000         -         +         -           Device MEA2150 Line         SN 60002 A PH 152 (117) HW CESQO-         -         Number of Consmits.         Anside 54. Digita 1
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The appropriate "Amplifier Gain" factor of 5 will automatically be entered. The "Input Voltage Range of Data Acquisition Board" divided by the gain is the "Signal Voltage Range", the maximum size of signals you can record before they get clipped. Please adjust the "Input Voltage Range of Data Acquisition Board" according to the expected size of your signals. It is advisable to select the input voltage range as small as possible, to ensure an optimal A/D resolution of the analog input signals. The "Sampling Frequency" should also be selected as low as possible, depending on the signals you like to measure, to minimize data file size.

# Hardware Filter Settings

While the MEA2100-Lite-System in principle can measure DC signals, this is for most purposes very impractical, as the baseline would constantly drift. Therefore, the system is equipped with software configured hardware filters. By default, a second order high pass filter with a cut off frequency of 1 Hz is used to prevent a baseline drift. The upper cut off frequency is 3.3 kHz. It is possible to change the settings of these filters without modification of firmware or hardware. Please read chapter "MEA2100 Configuration Software". Additionally, you can use the software filters of MC\_Rack, if necessary.

Note: It is possible to change the settings of these filters without modification of firmware or hardware. Please read chapter "MEA2100 Configuration Software".

# Setting up MC\_Rack

### Start the Data Acquisition

Click the "Start" button to start the data acquisition. No data is recorded. You see the raw data streams of all channels. You may have to adjust the scaling of the y-axes until you can clearly see the noise level. You should see the baseline with a maximum noise level of  $+/-20 \mu$ V. Please note that the system has a broad frequency range by default, so slow wave noise and drifting baselines may occur. This can be solved by using virtual filters.

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### Setting up the Internal Stimulus Generator

The headstage of the MEA2100-Lite-System is equipped with an integrated 3-channel stimulus generator. The three stimulation channels operate independently.

It is possible to choose between current or voltage controlled stimulation, and any electrode can be selected for stimulation. The control functions for electrode selection and definition of the stimulation parameters are integrated into MC\_Rack, you do not need any additional software. However, it is possible to import stimulation patterns generated in MC\_Stimulus II as ASCII data.

Add the internal stimulus generator to your rack: Click menu "Edit" and choose "Add MEA2100 Stimulator".

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File	Edit	Measurement	View	Windo	w Help		
2	Â	id MEA2100 (S/I	V xxxxx	-A) Ct	rl+⊂		
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	Ad	dd MEA2100 Stin	nulator				
	Ad	dd Trigger		Ct	rl+T		
	Ad	dd Spike Sorter		Ct	trl+K		
	Ad	Ct	trl+I				
	Ad	dd Longterm Disp	Ct	rl+M			
	Add Parameter Display			Ct	Ctrl+P		
	Add Digital Display			Ct	Ctrl+G		
	Add Filter			Ct	Ctrl+F		
	Add Sound				Ctrl+N		
	Add Analyzer				rl+A		
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The following dialog appears.

Recorder     MEA2100 (5(N:coccc-A)     Preday 5	Rack Stimulus 1 Stimulus 2 Stimulus 3 Channels				
MEA2100 Stimulator 1	multichannel*				
	Status: created				
	Lag				

Select the tab of any of the three stimulation channels (Stimulus 1, 2 or 3). In the "Stimulus" tab page, you can define the stimulation pulse and select how the stimulation is started and stopped.

Click "Stimulus 1" tab page, for example.

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#### Defining the stimulation pulse

In "Output Mode" you can choose whether you want to stimulate in voltage or current mode. To set up the stimulus pulse or pulse train, please use the eight provided up down boxes.

Pulse							
U/mV	Τ1/με	U/nW	T2/µs	IPI/µs	Repeat	ITI/ms	Repea
1600	100 -	-1600 -	100 -	1000 -	10	1000	- 1

The individual stimulation pulse is defined in the "Pulse" window. Set up a monophasic pulse by selecting the voltage or current in  $mV/\mu A$ and the duration time T1 in  $\mu s$ . For setting up a biphasic pulse you have to select the stimulation strength and time also for the second phase. The shape of the pulse is displayed immediately in the window below. Once started, a programmed paradigm will run to its' end, or till it is manually terminated.



Warning: Anodic (positive) pulses can lead to a formation of titanium oxide on the MEA electrodes. When using MEA electrodes of TiN material, use only negative voltages pulses or biphasic current pulses applying the negative phase first. Always regard the safe-charge injection limits. Otherwise, electrodes can be irreversibly damaged by electrolysis.

When using the "Train" window for setting up a pulse train, please select the "Inter Pulse Interval IPI" in µs and the number of pulses in the train in "Repeat". If no pulse train is needed, set the IPI to zero and the Repeat to one.

The "Inter Train Interval ITI" function can be used to repeat a single pulse or a pulse train. The interval time can be set in ms, and also the number of repeats. The complete programmed paradigm is shown in the graphical display below the stimulus window.

Once started, a programmed paradigm will run to its' end, or till it is manually terminated.

If the "Continuous Mode" is activated, the paradigm will be repeated indefinitely, till manually stopped. Please note that stimulation does not stop when MC\_Rack is stopped.



#### Importing a stimulation file created with MC\_Stimulus II

To generate more complex stimulation patterns, it is possible to import a stimulation file setup with the MC\_Stimulus II software as ASCII data. Please read the respective stimulus generator STG manual for detailed information about creating a stimulus file. You can import any pulse type available in MC\_Stimulus II.

Program the file in MC\_Stimulus II as usual: Channel 1 will become stimulus 1 in MC\_Rack, channel 2 will become stimulus 2, and channel 3 will become stimulus 3.

Export the file as ASCII file (MC\_Stimulus II: File - Export ASCII)

Import this file in MC\_Rack (Stimulus Tab: Import File)

It is not necessary to program Sync Out pulses. Information in MC\_Stimulus Channel 4 and up and all Sync Out channels will be ignored.

The "Stimulus" interface will be disabled and the imported pattern will be shown in the graphical stimulus display. To skip the imported pattern and reactivate the "Stimulus" interface, click the "Unload File" button.

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# Starting and stopping the Stimulation

In the "Trigger" drop down menu it is possible to choose different types of triggers to start the stimulation. The LED next to the Start / Stop button indicates the status of stimulation; the LED turns orange when the stimulation is running, the LED turns grey when the stimulation is stopped.

Important: It is **not** necessary for MC\_Rack to run in "Play" or "Record" mode to be able to start a stimulation paradigm, and a running paradigm will continue even after MC\_Rack is stopped. Once started, a programmed paradigm will run to its' end, or till it is manually terminated.

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If "Manual" is selected in the "Trigger" drop down menu, the "Start" and "Stop" buttons can be used to control the stimulation manually.

If "Start" is selected, the stimulation starts as soon as MC\_Rack is started (in "Play" or "Record" mode). A delay between start of MC\_Rack and start of stimulation in ms can be selected in the "Pre" time window. You can stop the stimulation paradigm by pressing the "Stop" button.

If a "DigIn" is selected, the stimulation starts if a TTL pulse is delivered to the respective connectors Digital In 1 to 4 on the front of the MEA2100-Lite-System interface board.

If the internal STG is connected to one or more "Trigger Detectors" in the MC\_Rack virtual tree, it is also possible to start the stimulation on any of these Triggers (Trigger 1, 2 ...).

If a new trigger arrives before the programmed stimulation paradigm is finished, the paradigm starts again from the beginning. All three stimulus channels can be started and stopped independently.

To repeat a stimulation pulse in a loop, activate the "Continuous Mode". The pulse will be applied continuously until it is stopped by manually with the "Stop" button.

#### **Selection of Stimulation Electrodes**

Click the "Channels" tab to assign the programmed stimulation patterns to individual electrodes, and to activate the blanking circuit BC, and to operate the "List Mode".



To apply the stimulation pattern of a specific stimulation channel (Stim. 1, 2 or 3) to one or more electrodes, select the desired stimulation channel (Stim. 1 to 3) and click the electrodes you want to apply this paradigm to. To apply the stimulation pattern to all electrodes, click the button "All" below the grid. Electrodes can be grounded or selected as regular recording electrodes (Not Stim.) in the same way.

Electrodes selected for stimulation pattern 1 "Stim.1" are colored in blue, for stimulation pattern 2 "Stim. 2" are colored in red, or stimulation pattern 3 "Stim. 3" are colored in yellow. To cancel a stimulation electrode, please use the radio button "Not Stim." and click on the selected electrode.

The blanking feature to avoid stimulation artifacts can be activated or deactivated via the check box "Blanking".

#### Blanking

A blanking signal transiently switches off the input stage of the amplifier during the stimulus, thus avoiding stimulus artifacts on non-stimulating electrodes.

Amplifier saturation is effectively prevented and the recovery time is greatly reduced. During the blanking period, a flat line is displayed. This blanking period extends a few hundred microseconds before and after the actual stimulus, currently 600  $\mu$ s, but might be subject to change. The pre- and post-time are experimentally optimized and cannot be changed by the user.

#### **Dedicated Stimulation Electrodes**

The function "Dedicated Stimulation Electrodes" can be used to further avoid stimulation artifacts on the electrodes surrounding the stimulation electrode, which are sometimes not completely removed by the blanking alone. Usually, the stimulation electrode will be connected to the stimulator output only during the stimulation pulse, and afterwards be switched back to the amplifier to be used as recording electrode. This sometimes causes switching artifacts independently from the actual stimulation.

However, if the function "Dedicated Stimulation Electrodes" is active, the selected stimulation electrodes will be constantly connected to the stimulator outputs. This causes permanent higher noise on the selected stimulation electrodes, and makes of course recording on the stimulation electrodes impossible, but almost completely removes residual artifacts. The images below show stimulation on electrode F3 with and without "Dedicated Stimulation Electrodes" activated.

Note: The artifact suppression additional to blanking via "Dedicated Stimulation Electrodes" is not available when using the "List Mode". "Dedicated Stimulation Electrodes" selected (X-Axis: 1000 ms, Y-Axis: +/- 500 μV)

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"Dedicated Stimulation Electrodes" deselected (X-Axis: 1000 ms, Y-Axis: +/- 500  $\mu\text{V})$ 

# SOFTWARE

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#### List Mode



Clicking the button List Mode opens an additional window: List of Stimulation Patterns. In List Mode it is possible to generate and download a list of up to 256 user defined stimulation patterns, which will then be applied automatically one after the other.

When the List Mode is activated, the three stimulator units are coupled to a single starting trigger. The starting trigger is set in the tab of the first stimulator unit, Stimulus 1. In Stimulus 2 and 3, the respective functions are disabled. To uncouple the stimulator units again, close the List Mode dialog. The step from one stimulation pattern to the next in the list happens when the start condition defined for Stimulus 1 occurs (for example, manual start, incoming trigger, real-time feedback trigger) and all three stimulator units are idle. Please see the red bars on the picture. However, the next pattern can only be started by the respective starting condition when the previous stimulation paradigm, including all breaks, is completely finished.



That means, if one stimulation pattern of any of the three stimulator units is still running when the condition occurs, nothing will happen. This also holds true when a stimulator unit is not actually used in the currently active stimulation pattern. Hence, the longest programmed stimulation pattern determines the minimum possible time between two starting triggers, even if the respective stimulator unit is not actually used. It is therefore advisable to set a stimulation pattern with a length of 0 on all unused stimulator units and keep all patterns as short as possible.

A stimulation pattern can contain single or multiple stimulation electrodes and may get an user defined name. To create a list of different stimulation patterns, please do the following.

# Use of the Digital Channel in the MEA2100-Lite-System

In contrast to the other MEA- and ME-Systems from Multi Channel Systems, the 16 input bits for the digital channel are not all accessible for external trigger signals, but are mostly used for internal communication between the different components of the MEA2100-Lite-System. Only the first four bits are connected to the physical Digin connectors on the front side of the MEA2100-Lite interface board. Most bits are permanently assigned to certain functions. However, these settings can be customized if necessary.

Please see the channels assignments in the following table.

Bit O	DigIn 1	Digital In channel 1
Bit 1	DigIn 2	Digital In channel 2
Bit 2	DigIn 3	Digital In channel 3
Bit 3	DigIn 4	Digital In channel 4
Bit 4	DigIn 5	Digital In channel 5
Bit 5	DigIn 6	Digital In channel 6
Bit 6	DigIn 7	Digital In channel 7
Bit 7	DigIn 8	Digital In channel 8
Bit 8	DigIn 9	Digital In channel 9
Bit 9	DigIn 10	Digital In channel 10
Bit 10	DigIn 11	Digital In channel 11
Bit 11	DigIn 12	Digital In channel 12
Bit 12	DigIn 13	Digital In channel 13
Bit 13	DigIn 14	Digital In channel 14
Bit 14	DigIn 15	Digital In channel 15
Bit 15	DigIn 16	Digital In channel 16

To monitor the digital data stream, you can use the "Digital Display" tool. If a MEA2100-Lite device is connected, the input bits will be labeled according to their function in the digital display.

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To record the digital data stream in the data file, please enable the respective check box in the "Channels" tab of the Recorder tool.



# **Triggering on Stimulation Pulses from Internal STGs**

It is possible to trigger on stimulation pulses from any of the three internal stimulus generators. The three internal STG units send a trigger signals with each stimulation pulse on bits 4, 5 and 6 of the digital channel, Stimulus 1 on bit 4, Stimulus 2 on bit 5 and Stimulus 3 on bit 6. Please see chapter "Use of the Digital Channel in the MEA2100-Lite" for more details. So, to trigger, for example on each pulse of Stimulus 1, a Trigger detector in MC\_Rack must be set to the Digital Channel, bit 4. Please see screenshot below.



It is also possible to set the trigger detector to react on a signal from any of the three STG units. This might for example be useful to start a triggered recording if a stimulus is applied from any of the STG units. All three bits must be unmasked, and the bit value must be defined as > 0. See chapter "Triggering on TTL Pulses" for more information.



Triggered recording and any instrument which is below the Trigger Detector in the virtual tree of MC\_Rack can now work in relation to the selected STG unit(s).

#### Setting up Cardio2D Software

When operating a MEA2100-Lite-System, you can connect the devices to any data acquisition computer. Make sure that the recommended system requirements are fulfilled.

The data acquisition computer from Multi Channel Systems MCS GmbH together with the MEA2100-Lite-System comes preinstalled and preconfigured for a flawless operation. You should contact your local retailer for assistance if you want to install additional hard- or software, or if you want to replace the computer, as incompatibilities of hardware components or software settings with the MEA2100-Lite-System or Cardio2D may occur.

Caution: You have acquired a high performance data acquisition and analysis computer. Do not modify the system, do not install new hard- or software, or another operating system without asking MCS or your local retailer for advice. Especially do not install virus scanners or firewalls because these programs are known to interfere with the data transfer to the hard disk. MCS cannot guarantee that a modified system is fully operational. Even data loss may occur.



Warning: The operating system settings of the data acquisition computer were preconfigured by MCS and should not be changed by the user. Changing these settings can lead to program instabilities and data loss.

#### Installing the Cardio2D Software

Make sure to have a valid license for the Cardio2D software! Please ask license@multichannelsystems.com.

You do not need a license for the Cardio2D software if you like to test the software for getting an idea of how it works and what is possible. You can test all functions, run the viability test to see your data, but you are not able to start an experiment.

Please contact license@multichannelsystems.com to receive a valid license key.

To enter the following dialog, select "License Key" in menu "Help". Please type the key into the window "Please enter license key" and confirm the key with "Set".



Doubleclick the Cardio2D.exe on the installation volume. The installation assistant will show up and guide you through the installation procedure. Follow the instructions of the installation assistant.

#### Note: The hardware driver is installed (or updated) automatically.

#### Installing the Cardio2D+ Analyzer Software

Doubleclick the Cardio2D+.exe on the installation volume. The installation assistant will show up and guide you through the installation procedure. Follow the instructions of the installation assistant.

You will again be asked for a path to put the log and data files. Cardio2D+ does not produce raw data files, result files only. Raw data files created by Cardio2D are never changed by Cardio2D+, but analyzed and saved as result files. It is advisable to use the same subfolders as used by the Cardio2D.

#### HARDWARE

#### Hardware Configuration

The MEA2100-Lite-System is a basic system, which is intended to be used with one headstage for microelectrode arrays with 60 electrodes (HS60). Please be aware that no upgrade is possible.



MEA2100-Lite-HS60-System

#### MEA2100-Lite Interface Board (MCS-Lite-IFB)

The MEA2100-Lite-System consists of two main devices: The interface board with integrated signal processor, and a headstage equipped with amplifiers, an integrated stimulator and analog-to-digital converter ADC.

#### **Front Panel**



#### From left to the right:

#### Status LED

The status LEDs indicate the link status of headstage 1. The left LED labelled with "Headstage 1" lights up when the MEA2100-Lite headstage is connected to the interface board via eSATAp cable. Please switch the device on.

#### **Auxiliary Channels**

Two reserve auxiliary channels are available for future use. They have no function at the moment.

#### Digital IN / OUT

A Digital IN / OUT for 16 digital in- and output bits is available (68-pin MCS standard connector) on the rear panel of the interface board. On the front panel four digital IN and four digital OUT connectors bits are also accessible via Lemo connector DIG IN bit 0 to bit 3 and DIG OUT bit 0 to bit 3. If access to more bits of the DIG /N / OUT channel is required, it is necessary to connect a Di/o board with a 68-pin standard cable. This Di/o board is available as optional accessory.

#### Power LED

The power LED near to the MCS logo on the front panel of the interface board should light up when the MEA2100-Lite-System is "ON", and the device is connected to the power line. If not, please check the power source and cabling.

#### **Rear Panel**



#### Toggle Switch On / Off

Toggle switch for turning the device on and off. The MEA2100-Lite-System is switched to status "ON" when the toggle switch is switched to the left. The device is switched "OFF" when the toggle switch is switched to the right. If the MEA2100-Lite-System is "ON", and the device is connected to the power line, the Power LED on the front panel of the interface board should light up. If not, please check the power source and cabling.

#### Power IN

Connect the power supply unit here. This power supply powers both, the headstage and the interface board of the MEA2100-Lite-System. The device needs 12 V and 2.5 A / 30 W.

# Ground

If an additional ground connection is needed, you can connect this plug with an external ground using a standard common jack (4 mm).

#### Digital IN / OUT

A Digital IN / OUT for 16 digital in- and output bits is available (68-pin MCS standard connector). Please read chapter "Pin Layout" (Digital IN / OUT Connector) in the Appendix for more information about the pin layout of the connector. The Digital IN / OUT connection accepts or generates standard TTL signals. TTL stands for Transistor-Transistor Logic. A TTL pulse is defined as a digital signal for communication between two devices. A voltage between 0 V and 0.8 V is considered as a logical state of 0 (LOW), and a voltage between 2 V and 5 V means 1 (HIGH).

The Digital OUT allows generating a digital signal with up to 16 bits and read it out, for example, by using a Digital IN / OUT extension Di/o from Multi Channel Systems MCS GmbH. You can utilize this digital signal to control and synchronize other devices with the MEA2100-System. Bit 0 to 3 of the Digital OUT are separated and available as Lemo connector DIG OUT 0 to 3 on the front panel of the interface board. So, if you need only one, two, three or four bits of the digital signal, you don't need the additional Di/o. Please read chapter "Pin Layout" (Digital IN / OUT Connector) in the Appendix for more information.

The Digital IN can be used to record additional information from external devices as a 16 bit encoded number. The Digital IN is most often used to trigger recordings with a TTL signal. The 16 bit digital input channels is a stream of 16 bit values. The state of each bit (0 to 15) can be controlled separately. Standard TTL signals are accepted as input signals on the digital inputs. Unused input bits, which have an undefined state, should be masked in the "Trigger Detector" of MC\_Rack.



Warning: A voltage that is higher than +5 Volts or lower than 0 Volts, that is, a negative voltage, applied to the digital input would destroy the electronics. Make sure that you apply only TTL pulses (0 to 5 V) to the digital inputs.

#### **Analog Channels**

Up to eight Analog IN channels are available via 10-pin connector. Please read chapter "Pin Layout" (Analog IN Connector) in the Appendix for more information.

The additional analog inputs are intended for recording additional information from external devices, for example, for recording patch clamp in parallel to the MEA recording, for monitoring the temperature, or for fluidic control. Two of these channels (No 1 and 2) are available via Lemo connectors on the rear panel of the interface board. You could also use the analog inputs for triggering, but please note that the digital inputs DIG IN 0 to 3 are intended for accepting TTL pulses. Signals on the analog channels are digitized and amplified with a gain factor of 2.

#### Analog Channels 1 and 2

Two of the analog channels described above (analog channel 1 and 2) are directly accessible via two Lemo connectors.

#### JTAG Connector

The 20-pin JTAG connector for use of the digital signal processor DSP is not in use in MEA2100-Lite Systems.

#### USB 3.0 Connector

The USB connector is used to transfer the amplified and digitized data from all data channels and the additional digital and analog channels to any connected data acquisition computer via USB 3.0 super speed (type A - Micro B) cable. Please use USB 3.0 connector labelled "A".

Important: It is recommended to connect the USB cables to the USB ports of the computer. Do not use an USB hub!

#### Audio OUT

To the Audio Out (3.5 mm phone jack) you can connect an audio system to make the electrical activity audible. This audio output is real-time. Headphones or a speaker can be connected directly to the AUDIO OUT. Only two channels at a time can be converted into sound (Stereo).

# Side Panel



Socket for connecting the headstage via eSATAp cable. The socket labelled "1" is available only.

### MEA2100-Lite Headstage

The rear part of the MEA2100-Lite-System headstage contains the amplifiers and the data acquisition for 60 channels and three independent stimulator units. The variable front part of the headstage contains the contact unit for different kinds of MEAs.

Headstage for microelectrode arrays (MEAs) with 60 electrodes.



The side part has connectors for the serial eSATAp connector to connect the headstage to the interface board, and for the connection between the integrated heating element) of the headstage and the Temperature Controller TCX in the base plate. The contact unit houses the MEA and features three grounding plugs for each MEA. These are internally interconnected and enable the user to ground an Ag/AgCl reference electrode or external components, like a perfusion system.

Headstage for microelectrode arrays (MEAs) with 60 electrodes.



Additionally, headstages for 60 channels electrodes are also available with a modified ground plate to accommodate perforated MEAs. The amplifier and data acquisition integrated in the headstage support 60 electrode channels.

The sampling rate, signal range and bandwidth can be adjusted via software control. Thereby the system is suitable for a broad range of applications, from single unit spike recordings to field potentials from whole heart preparations. Electrode raw data acquired from the MEA chip is amplified and digitized by the analog / digital converter that is integrated in the headstage at sampling rates of up to 32 kHz per channel. Data is transferred to the interface board via eSATAp cable.

A special advantage are the three integrated stimulus generator units, which deliver monophasic or biphasic rectangular pulses as single pulse and pulse trains. You can choose between current or voltage stimulation and you can select each MEA electrode for stimulation within the Multi Channel Experimenter and / or the MC\_Rack software. The integrated blanking circuit (BC) disconnects the electrodes from the amplifiers during the stimulation and avoids stimulus artifacts.

As usual in MEA-Systems, the MEA2100-System-Lite features a heating element with a Pt100 temperature sensor. If you connect a temperature controller TC02 to the heating element via D-Sub 9 connector, the heating element guarantees constant temperature conditions for the biological sample on the chip. A perfusion system with heating is available for the MEA2100-Lite as optional accessory.

Because of the compact designed headstage you can use most inverted or upright microscopes, for the optical control of the sample on the MEA chip.

#### **Bandwidth and Gain Specification**

The MEA2100-Lite amplifier with blanking circuit is a 60-channel headstage with a hardware defined bandwidth. Default settings for the hardware filter will be 1 Hz to 3 kHz. However, a different bandwidth can be achieved by an additional software program, called "MEA2100 Configuration", no hardware modification is necessary. The hardware defined bandwidth marks the upper and lower limit of the frequency range, fine tuning within this range can be done with software filters. Please see the manual for the "MEA2100 Configuration" software in the Appendix.

The MEA2100-Lite-System has a fixed hardware gain of 5. However, due to the large resolution of 24 bit (16 bit), it is possible to adjust the actual signal input range used for recording in a wide range from  $\pm$  812  $\mu$ V to a maximum of  $\pm$  227 mV. The 24 bits are distributed among this complete voltage range. In the MC\_Rack software, a window with 16 bit resolution can be adjusted on this range and display signals with high resolution in the following magnitudes:

 $\pm$  812 µV;  $\pm$  1727 µV,  $\pm$  3545 µV;  $\pm$  7090 µV;  $\pm$  14 mV;  $\pm$  28 mV,  $\pm$  57 mV;  $\pm$  114 mV;  $\pm$  227 mV

A smaller input range results in a better resolution in bits/ $\mu$ V, signals larger than the selected range will be clipped. In summary, the procedure has exactly the same functionality as a software selectable gain, even though, strictly speaking, it is not.

#### **DC Offset Correction**

The internal baseline restore routine is used to eliminate DC offsets of the electrode channels and set all baseline values to zero. This has to be a slow process. If the routine corrected too fast, slow signals would be eliminated as well.

All electrodes on an array have different DC offsets at the beginning of an experiment. The DC offsets are amplified by the headstage, and the amplified output signal of the preamplifier is likely to exceed the input range of the filter amplifier. In order to eliminate this DC offset, a so-called baseline restorer is integrated in the headstage. For signals in the usual measurement range, the baseline restore routine acts like a high pass filter with a very low cutoff frequency of roughly 0.01 Hz.

Due to the low cut-off frequency, the baseline restorer takes some time to stabilize after switching on the amplifier or after replacing the MEA. One to two minutes are quite normal. During this time, the filter amplifier input will be overloaded and you will see only a straight line with very low noise. This is considered a normal behavior and should not trouble you. Please wait until the baseline restore routine has been completed. After about one minute, the baseline restorer of the headstage has corrected the DC offset, and the output signals are now in the operating range of the filter amplifier. Now you should see a noise level of about 20 µV.

#### **Stimulus Generator**

The headstage of the MEA2100-Lite-System is equipped with an integrated three channel stimulus generator, which delivers rectangular pulses in three different stimulation patterns: monophasic, biphasic and pulse trains. You can choose between current or voltage stimulation and you can select each electrode for stimulation. The integrated blanking circuit (BC) disconnect the electrodes from the amplifiers during the stimulation and avoids stimulus artifacts. Control of stimulation patterns and selection of stimulation electrodes is done within the respective software.

Please read chapter "Setting up the Internal Stimulation Generator" for setting up and applying the stimulation parameters.



#### SETTING UP THE MEA2100-LITE-SYSTEM

#### **Connecting Headstages, Interface Board and Computer**

- Provide a power supply in the immediate vicinity of the installation site.
- Place all devices on a stable and dry surface, where the air can circulate freely and the devices are not exposed to direct sunlight.
- Set up the computer (with installed software program).
- Install the software program from the installation volume if it is not already installed. The MEA2100-Lite-System is a plug and play device. The driver is automatically installed together with the software program.
- Set up the MEA2100-Lite headstage(s) on a microscope or on a solid table or workbench.
- Connect the headstage/s via the provided eSATAp cable/s to the interface board.
- Connect the USB 3.0 port of the interface board to a free USB 3.0 port of the data proceeding acquisition computer. Do not use an
  USB hub and also avoid USB ports on the front panel of the computer, as they are often connected to the main board via an internal
  hub.
- Connect the Interface Board via power supply unit to a power outlet of the same electrical system (connected to the same ground / earth wire) as all other components of the setup, for example, the computer or shielding.
- If necessary, connect the system to an external ground.
- Connect the internal heating element of the MEA2100-Lite headstage to the temperature controller's output channel (D-Sub9 socket) with the provided cable. Do not connect the heating element cable to the computer!
- Turn the toggle switch at the rear panel of the interface board on.
- Check the power LED. It should light up as soon as the power line is connected, and the toggle switch is switched on. If not, check the
  power source and cabling.
- Start the software and select the MEA2100-Lite-System as the data source.
- Please see the software help or manual for more details on how to define the data source.

#### **Opening and Closing the Headstage**

Open the lid of the headstage by pushing the two rectangular panels on the left and right side of the contact unit sideward. The upper part of the headstage swings open. The base plate underneath the lid has either one or two slots for MEAs. Usually, each slot has a heating element with a hole in the middle to allow microscope access from the bottom. If equipped for perforated pMEAs, the heating element is replaced by a perfusion ground element with fluidic channels and an O-ring. Close the lid again by pushing the panels back in their original position.



# HARDWARE



#### **First functional Tests**

Each MEA2100-Lite-System has been thoroughly tested at the factory site before delivery. However, you may want to perform some tests yourself before you begin your experiment to exclude any damage that might have occurred during transportation, or to fulfill your own guidelines, for instance. Some of the tests will also help you to get to know the basic functions of the hard- and software, like a short tutorial. It will take only a few minutes time and can save time and trouble in the long run. Multi Channel Systems recommends running these tests after the setup of your system before you start your real experiments.

#### Setting up the MEA

Open the lid of the headstage. Place the MEA chip inside the MEA2100-Lite headstage. The MEA is not rotationally symmetrical, so take care for the orientation of the MEA.



Important: MEAs are not symmetrical! That is, why the writing (for example NMI, LEITER, MEA type) on the MEA chip should be on the right side viewed from the front, when the headstage is open. MEAs with one big internal reference electrode should be placed with reference electrode to the left side in the headstage. Otherwise, the MEA layout will not match with the pin layout of the channel map of the software.

You can use the engraved serial number as marker. The pin layout of a MEA is described in the respective "MEA Datasheet".



Warning: Spilled liquid can damage or even completely destroy the electronics of the MEA2100-Lite headstage. Please be careful when setting up your perfusion system and when starting the perfusion. Take care that the flow rates of the inlet and outlet flow match so that flooding of the amplifier is efficiently prevented.

# Grounding the Bath

Most MEA chips are equipped with an internal reference electrode, which will be automatically connected to the systems ground when the MEA2100-Lite headstage is closed. This has the advantage that you can keep the culture chamber closed and sterile (for example, with MEA-MEM semipermeable membranes).

If the MEA does not have an internal reference, please use the provided AgAgCl pellet or a silver wire to connect the bath to ground. The connectors beside the area for the MEA on the lid of the headstage are all internally connected to the system ground, so you can use any of them for grounding.

### Service and Maintenance

You should clean the contact pins of the MEA2100-Lite headstage carefully with a soft tissue and alcohol or isopropanol from time to time, especially if you have problems with the noise level. For handling and cleaning of the MEA, please refer to the MEA manual or to MEA Cleaning Quick Guide on the MCS web site <u>www.multichannelsystems.com</u>.



Warning: Please be very careful when handling the headstage, or cleaning the device! The pins can be damaged easily.

#### **General Performance / Noise Level**



Please use the provided test model probe to test the amplifier immediately after installation. It simulates the behavior of a MEA electrode in a bath with conductive solution by a resistor of 220 kOhm and a 1 nF capacitor between ground and amplifier. Use the data acquisition program to record from the test model probe and to check the headstage.

Please read also the datasheet "Test-60MEA"

A MEA Signal Generator is a convenient tool adapted for MEA2100-Lite-Systems with HS60. Use the 60MEA2100-SG instead of setting up an experiment with biological sample for training, controlling, and troubleshooting purposes. This reduces the number of animal experiments and saves laboratory equipment. Please see also datasheet "60MEA2100-SG".

Important: When using a 60MEA2100-SG which is not adapted to MEA2100-Lite-System, the contact to the internal reference must be disconnected. Otherwise, the delivered signals are not acceptable. Please insert the 60MEA2100-SG in correct orientation as shown on the following picture. Cover the designated contact pad connected to the reference electrode, for example with isolation tape.



# Temperature Controller TC01 / 02



The MEA2100-Lite-System features a heating element with a PT100 temperature sensor. If you connect a temperature controller TCX to the heating element via D-Sub 9 connector, the heating element guarantees constant temperature conditions for the biological sample, placed on the MEA.

#### **Perfusion Cannula PH01**



When using a perfusion system, the perfusion cannula with heating element and sensor PH01 can be connected to a temperature controller TC02. This is very useful for general perfusion purposes, whenever the fluid stream needs to be temperature controlled. Magnetic holders for the PH01 and a suction cannula and a steel plate fitting the headstage of the MEA2100 are available as optional accessory.

For using pMEAs it is necessary to have a MEA2100-Lite-System equipped with a perfusion element. The perfusion element replaces the regular ground plate with heating element. It is possible to change the elements, but the exchange has to be done by MCS or a qualified distribution partner.

#### TROUBLESHOOTING

#### About Troubleshooting

The following hints are provided to solve problems that have been reported by users. Most problems occur seldom and only under specific circumstances. Please check the mentioned possible causes carefully when you have any trouble with the product. In most cases, it is only a minor problem that can be easily avoided or solved.

If the problem persists, please contact your local retailer. The highly qualified staff will be glad to help you. Please inform your local retailer as well, if other problems that are not mentioned in this documentation occur, even if you have solved the problem on your own. This helps other users, and it helps MCS to optimize the instrument and the documentation. Please pay attention to the safety and service information in the separate user manuals of the related products and in the software help. Multi Channel Systems has put all effort into making the product fully stable and reliable, but like all high-performance products, it has to be handled with care.

#### No Computer Connection / No Recording Possible

You cannot establish a connection to the computer. The MEA2100-Lite-System channel layout is not available in MC\_Rack. When loading a previously saved virtual rack file, you will get an error message and the simulator will be started automatically. You get an error message when starting the recording in MC\_Rack after a successful computer connection.

#### Possible causes:

? The power LED is not lightning. The supply power is not connected or there is a technical problem with the instrument.

Check the power source and the cable connections. If this does not solve the problem, contact your local retailer for support.

? The power LED is lighting. The USB port might not support USB 3.0 or might not be working.

Check the USB port. Only high speed USB 3.0 ports can be used. Try another USB 3.0 port.

? You can establish a connection, but get an error message when starting the recording in MC\_Rack.

The bandwidth of the USB port is not sufficient for recording. This can be the case if the MEA2100-Lite-System is connected via USB hub, and a second device that sends or receives continuous data streams, for example, a web cam or USB speakers, is connected to the same USB port. Connect the MEA2100-Lite-System directly to a USB port, not via hub.

#### Triggering / Digital Input does not Work

You have connected a TTL source to one of the digital inputs of the MEA2100-Lite-System, and configured the virtual rack in MC\_Rack for triggering displays or data acquisition by the TTL source, but you do not see any sweeps.

#### Possible causes:

? The TTL source does not generate true TTL signals (> 1.5 to 5 V TTL level), or the TTL pulse duration is too short in combination with the sampling rate, so that the pulse is missed in-between two data points.

The MEA2100-Lite-System can only accept TTL signals as a digital input stream. The TTL pulse needs to be optimized according to the sampling rate. Otherwise, a detection of the trigger cannot be guaranteed.

? The software settings for the Trigger Detector do not match with the hardware configuration.

In MC\_Rack, add a Trigger Detector to your virtual rack, and select the Digital Data D1 input stream as the Trigger. Check the pin layout of the Digital IN / OUT connector and make sure that the same bit input that is connected is selected in the software. (The standard settings of the Trigger Detector are for using bit 0.) Mask all unused bits. Select the appropriate logical state (generally HIGH) for triggering.

Please see the MC\_Rack help or manual for more details.

#### **Noise on Single Electrodes**

The noise level on single electrodes is significantly higher than expected or you see artifact signals.

Possible causes:

? The electrode or the contact pin of the headstage may be defective. To test this, do the following:

Open the amplifier and turn the MEA by 90 degrees. Close the MEA2100-Lite headstage again and start the recording. If the same electrode in the MEA layout is affected, the amplifier's contact is not okay. If another electrode is now affected and the previously affected electrode is all right now, the MEA electrode is not okay, but the headstage is fine.

-OR-

Use the test model probe to test the MEA2100-Lite headstage. If the noise level is fine without the MEA, bad MEA electrodes cannot be the cause.

# MEA is defective

MEAs wear out after multiple uses or over a longer time of use, for example, for long-term cultures. This is considered a normal behavior. MEAs are also easily damaged by mishandling, for example, if wrong cleaning solutions or too severe cleaning methods are used or if the recording area is touched. If you observe a bad long-term performance of MEAs, consider a more careful handling.

Possible causes:

? The contact pads are contaminated.

Clean the contact pads carefully with a swab or a soft tissue and pure (100 %) alcohol.

? The contact pads or the electrodes are irreversibly damaged. You could have a look at the electrodes under a microscope: If they appear shiny golden, the TiN is gone and the electrode is irreversibly damaged. Electrodes may be damaged without changing their visual appearance, though.

Pick one of the bad channels after the other and ground it. See the MEA amplifier's manual for more information on grounding channels. In most cases, only one of the electrodes that appear bad is actually defective, and the other ones are only affected by the single defective electrode. Ground as many electrodes as you need for a good general performance. Grounded electrodes show a noise level that is lower than that of good electrodes. If too many electrodes are defective, use a new MEA.

#### **Overall Noise / Unsteady Baseline**

The baseline is unstable, signals are jumping or drifting.

Possible causes:

? Bath electrode is not connected to ground.

Check orientation of MEA with internal reference. The internal reference must be on the left side of the MEA when it is in the amplifier. If the MEA has an internal reference, use an Ag/AgCl bath electrode.

? AgCl bath electrode is not well-chlorided.

Rechloride the electrode or use a new one.

? 50 Hz hum: 50 Hz is the frequency of mains power in Europe.

If the shielding and grounding of the setup is not sufficient, electrical signals are picked up from the environment. Use a proper shielding. For example, you can place aluminum foil over and / or under the amplifier that is connected to the system ground (ground connectors of the headstage or ground connector in the back of the interface board). You can also use special shielding equipment like a Faraday cage.

# Missing Spikes or Strange Signal Behavior

MEAs wear out after multiple uses or over a longer time of use, for example for long-term cultures. The insulation layer gets thin over time. This is considered a normal behavior.

Possible causes:

? The insulation layer is too thin. As a result, the MEA gets the behavior of a low pass filter.

This means, that the signal frequency may be shifted to a lower frequency, and spikes are missing. Optically control the MEA with a microscope. If concentric colored rings (Newton rings) are visible (due to light interference), the insulation layer is too thin and you should use a fresh MEA.

? The insulation layer has been abraded and is missing in parts.

This will result in a short circuit between the electrodes or tracks and the bath. You will still see signals, but as an unspecific smear over the complete array. Use a fresh MEA.

#### MEA2100 CONFIGURATION SOFTWARE

The MEA2100 Configuration software is an add-on program for changing the hardware filter settings by the software. No firmware update is necessary. You can operate the software with MEA2100-Systems and ME-Systems.

Please install the MEA2100 hardware filter configuration software by clicking the "MEA2100\_Config.exe" file. Follow the installation guide as usual.

#### Main Window of the MCS Filter Configuration Program

MEA Devices:	MEA2100	(00087-A)			✓ Refresh	
Filter Settings	MEA2100	(00087-A)				
Headstage 1					Permanent Memory	
High F	ass Filter:	2. order 🗸	1 Hz	~	Store Coefficients	Clear Coefficents
	ass Filter	4. order ∨	3500 Hz	~	Store Coefficients	Clear Coefficents

The MCS Filter Configuration program for changing the hardware filter setting of the MEA2100-System is self-explanatory via tool tips. It is possible to control each connected headstage and each MEA slot separately.

Define the "MEA Devices" from the drop-down box if two headstages or headstages with two MEA slots are available. Press "Refresh" button to update the settings.

#### **Filter Settings**

The actual settings for each headstage separately are available. "High and Low Pass Filter" for the selected headstage are displayed. If the actual settings are not on the drop-down list, "Custom" will be displayed.

Select the order of the filter from the first drop-down box. The message "The filter order determines the steepness of the filter" is shown. The higher the order, the steeper the cut off.

🖸 MCS Filter Configuration - Version 1.1.3 🚽 🗖 🗙						
MEA Devices: MEA2100	) (00087-A)		✓ Refresh			
Filter Settings	Filter Settings					
Headstage 1	Headstage 1 The filter order determines the steepness of the filter.					
High Pass Filter:	2. order 🗸	1 Hz 🗸	Store Coefficients	Clear Coefficents		
Low Pass Filter:	4. order 🗸 🗸	3500 Hz 🗸 🗸	Store Coefficients	Clear Coefficents		
	1. order 2. order 3. order 4. order	Exit				

Select the cut off frequency of the filter from the second drop-down box. The message "The cut off frequency of the filter. The amplitude is reduced by 3 db at this frequency. A filter with Bessel characteristic is used." is shown.



If you need a frequency which is not available on the list, please contact <a href="support@multichannelsystems.com">support@multichannelsystems.com</a>. The selected settings are automatically downloaded to the ME2100 / MEA2100 device as soon as you select them. If the "Store Coefficients" button is not used, the device will be back to the last saved settings after a reset.

Click the button "Store Coefficients" to store the selected filter settings permanently in the Eeprom memory (electrically erasable programmable read-only memory).



Click the button "Clear Coefficients" to remove the coefficients from Eeprom and to refresh the factory default settings via power-cycle.



### **TECHNICAL SUPPORT**

Please read the chapter "Troubleshooting" of the manual first. Most problems are caused by minor handling errors. Contact your local retailer immediately if the cause of trouble remains unclear. Please understand that information on your hardware and software configuration is necessary to analyze and finally solve the problem you encounter. Please keep information on the following at hand:

- Description of the error (the error message text or any other useful information) and of the context in which the error occurred. Try to remember all steps you had performed immediately before the error occurred. The more information on the actual situation you can provide, the easier it is to track the problem.
- The serial number of the device. You will find it on the backside of the housing or click "About" in "Help" in the main menu of the software.
- The operating system and service pack number of the connected computer.
- The hardware configuration (microprocessor, frequency, main memory, hard disk) of the connected computer. This information is especially important if you have modified the computer or installed new hard- or software recently.
- The version of the recording software. On the "Help menu", click "About" to display the software version.

# MEA2100-Lite-System



Warning: The device may only be used together with MEA chips from Multi Channel Systems MCS GmbH, and only for the specified purpose. Damage of the device and even fatal injuries can result from improper use. Do not open the headstage or the interface board and do not change hardware configuration as it could lead to improper behavior of the system.

General Characteristics					
Operating temperature	10 °C to 50 °C				
Storage temperature	0 °C to 50 °C				
Relative humidity	10 % to 85 %, non-condensing				
Headstage	AFA AF4 AF				
Dimensions (W x D x H)	250 mm x 151 mm x 25 mm				
Weight	1000 g				
Integrated Amplifier					
Number of analog recording channels	60				
Data resolution	24 bit (16 bit, if operated with MC_Rack)				
Input voltage range	MC_Rack: from ± 4.9 mV to ± 500 mV Multi Channel Experimenter: ±1250 mV				
Bandwidth	0.1 Hz to 10 kHz				
Sampling frequency per channel	up to 32 kHz, MC_Rack software controlled up to 25 kHz, Multi Channel Experimenter controlled				
Input impedance	1 G [2]    10 pF				
Integrated Heating Element					
Temperature sensor type	Pt 100 with four wire connection, compatible with TCX				
Accuracy	± 0.1 °C				
Integrated Stimulus Generators					
Output current	± 1.5 mA				
Current output compliance voltage	± 16 V				
Output voltage	± 10 V				
Voltage output compliance current	± 20 mA				
Stimulation pattern	MC_Rack: rectangle (biphasic, monophasic, pulse trains) Multi Channel Experimenter: almost arbitrary patterns				
Number of stimulation channels	3 independent stimulation patterns per 60 channels				
Resolution	16 bit				
MCS-IFB 3.0 Interface Board					
Dimensions (W x D x H)	250 mm x 60 mm x 25 mm				
Weight	300 g				
Front panel					
4 Digital inputs	Lemo connector, EPL.00.250 NTN				
4 Digital outputs	Lemo connector, EPL.00.250 NTN				
2 Auxiliary channels (not in use)	Lemo connector, EPL.00.250 NTN				
Rear panel					
1 16 bit Digital In / Out	68-pin MCS standard connectors				
1 8-Channel Analog connector	10-pin connector (2,54 mm grid), dual row standard IDC				
2 Auxiliary channels (not in use)	Lemo connector, EPL.00.250 NTN				
2 Analog Inputs,	Lemo connector, EPL.00.250 NTN				
Channel 1 and Channel 2					
1 USB 3.0 micro port	USB 3.0 super speed cable (type A - Micro B)				
Power Supply	MPU 30, scope of delivery, PWR DC 0.65 x 2.75 mm				
Ground	Common jack 4 mm, banana plug				
1 Audio Output	Stereo jack 3.5 mm				

# Side panel

1 Interface board to headstage

# Power supply unit (MPU 30)

Input voltage Output voltage Max. Power Mark of conformity European standards

# Software

Operating system Microsoft Windows <sup>®</sup> Multi Channel Experimenter Data acquisition and analysis software Multi Channel Analyzer Data analysis software Multi Channel DataManager MC\_Rack program Data acquisition and analysis software MC\_DataTool program Data export software External powered over serial ATA cable (eSATAp)

90 – 264 VAC @ 47 – 63 Hz 11 – 13 V 30 W CE, TÜV, cUL EN60601

Windows 10, 8 or 7 (32 or 64 bit), English and German version supported

Version 2.6.0 and higher

Version 2.6.0 and higher

Version 1.5.6 and higher Version 4.6.0 and higher

Version 2.6.3 and higher Axon binary file (\*.abf), ASCII file (\*.txt), binary file (\*.raw)

# Pin Layout

# MEA2100-Lite Interface Board: 10-Pin Connector (Analog IN)

10-Pin Connector for Analog IN



The gain factor for amplification of all analog channels is 2.



#### 10-Pin Connector for Analog IN

Pin Number	1	2	3	4	5	6	7	8	9	10
Analog Channel Number	GND	GND	1	2	3	4	5	6	7	8

# MEA2100-Lite Interface Board: Lemo Connectors for Analog IN

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Analog IN 1 and 2, available via Lemo Connectors Analog IN 1 channel 1 Analog IN 2 channel 2

# MEA2100-Lite Interface Board: Digital IN / OUT Connector



#### 68-Pin MCS Standard Connector



Pin 1	GND
Pin 2	GND
Pin 3 - 10	Digital output channels 0 - 7
Pin 11 - 14	GND
Pin 15 - 22	Digital output channels bit 8 - 15
Pin 23 - 26	GND
Pin 27 - 34	Digital input channels 0 - 7
Pin 35 - 38	GND
Pin 39 - 45	Digital input channels bit 8 - 15
Pin 46 - 48	GND
Pin 49 - 63	Internal use (do not connect)
Pin 64 - 66	GND
Pin 67	Positive supply voltage output 3.3 V
Pin 68	GND

# Digital IN / OUT, separate available via Lemo Connectors on the front panel of the IFB

Separate Digital IN Lemo Connectors Digital In 1 Bit 0 of the 16 bit digital input channels (Pin 27) Digital IN 2 Bit 1 of the 16 bit digital input channels (Pin 28) Digital IN 3 Bit 2 of the 16 bit digital input channels (Pin 29) Digital IN 4 Bit 3 of the 16 bit digital input channels (Pin 30) Separate Digital OUT Lemo Connectors Digital OUT 1 Bit 0 of the 16 bit digital output channels (Pin 3) Digital OUT 2 Bit 1 of the 16 bit digital output channels (Pin 4) **Digital OUT 3** Bit 2 of the 16 bit digital output channels (Pin 5) Digital OUT 4 Bit 3 of the 16 bit digital output channels (Pin 6)

### **Changing Gold Spring Contacts**

Instruction for the replacement of gold spring contacts inside the lid of the MEA2100-Lite headstage.

Only experienced persons should replace the gold spring contacts, because they are small and lightweight buildings and can easily be destroyed!



Please follow the instructions with great care! Switch off the MEA2100-Lite headstage and unplug the device from power supply.

Define the pins which need to be changed with the software. Count the gold spring contacts in the lid of the amplifier to find the pin which need to be changed.

Please use the pin layout of a standard 60MEA200/30iR-Ti for the MEA2100-HS60. Please read also the respective data sheets of the MEA.

### Pin Layout: Headstage HS60

Please insert the 60MEA200/30iR-Ti as recommended. The internal reference electrode has to show to the left side, the writing "NMI MEA" to the right side when looking from the front of the headstage with the articulation in the back. Open the software, one instance for each MEA. Add the default map in the display and you find the designated pin layout. Count the pins to find the gold spring contact, you like to change.

#### Pin Layout of the 60MEA200/30iR-Ti



# TECHNICAL SPECIFICATIONS

# Changing Gold Spring Contacts in MEA2100 Headstages



- 1. Open the MEA2100 headstage, turn it upside down to have a better access to the gold spring contacts in the lid. Place the device on a soft surface.
- 2. Grab the defective gold spring contact with a needle nose pliers and remove it upwards without damaging the contacts on the left and on the right side.



- 3. Place a new gold spring contact into the hollow shaft of the old one.
- 4. Take the fixing tool for gold spring contacts and put the small end of the tool over the head of the new gold spring contact straight from above.
- 5. Push the gold spring contact with the fixing tool into the shaft until the gold spring contact is engaged.



# CONTACT INFORMATION

# Local retailer

Please see the list of official MCS distributors on the MCS web site.

# Mailing list

If you have subscribed to the Newsletter, you will be automatically informed about new software releases, upcoming events, and other news on the product line. You can subscribe to the list on the MCS web site.

www.multichannelsystems.com

