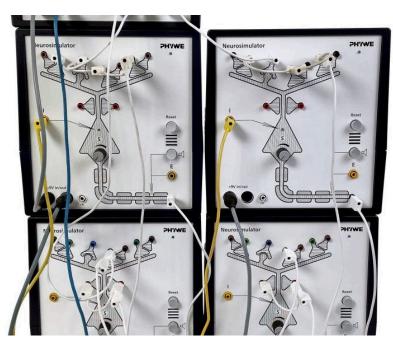
## **Neural networks with Cobra SMARTsense**



Experiments on the topic of neural networks like short-term memory, body clock and others. In preparation for these experiments, it is strongly recommended that the experiments on nerve cells with one nerve cell and nerve cell interactions with two nerve cells be carried out beforehand. The material required for this is already included in these experiments on neural networks.

| Biology          |                         | Nervous system, neurobiology |                |  |  |
|------------------|-------------------------|------------------------------|----------------|--|--|
| Difficulty level | <b>RR</b><br>Group size | D<br>Preparation time        | Execution time |  |  |
| hard             | -                       | 10 minutes                   | 45+ minutes    |  |  |

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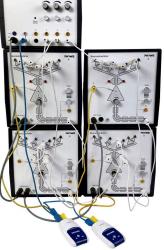


# **General information**

## **Application**

The human brain is made up of billions of neurons connected by synapses that transmit information in response to electrical and chemical signals. These interactions enable the brain to perform complex functions such as memory, perception, movement and thinking. Typical experiment setups with 3

and 4 nerve cells

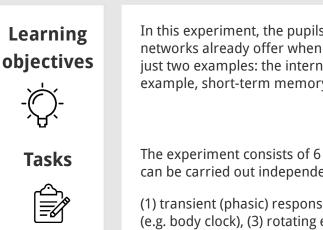


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# Other information (1/2) Second and a construction of the second and the se

## Other information (2/2)



In this experiment, the pupils and students learn what complex possibilities neuronal networks already offer when three or four nerve cells are interconnected. To name just two examples: the internal clock, which determines the sleep rhythm, for example, short-term memory and sensory learning.

The experiment consists of 6 examples from the wide world of neural networks that can be carried out independently of each other:

(1) transient (phasic) responses with emphasis on visual sense, (2) neuronal oscillator (e.g. body clock), (3) rotating excitation (short-term memory), (4) cerebral cortex and sensory learning, (5) directional selectivity through unilateral inhibition and self-calibration of paired sensory channels.



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## Safety instructions





The general instructions for safe experimentation in science lessons apply to this experiment.

#### Equipment

| Position | Equipment  | Item no. | Quantity |
|----------|--|----------|----------|
| 1        | Set Neurobiology with a nerve cell with Cobra SMARTsense | 65963-22 | 1        |
| 2        | Neurosimulator   | 65963-00 | 3        |

#### **Theory 1**



#### Transient (phasic) reactions in the sense of vision

One type of neuron responds only to changed stimulation, not to sustained stimulation. These are called ON and OFF neurons. They respond only to the onset and/or disappearance of a stimulus with an activation that is usually brief and more intense the stronger the stimulus change. These two types of neurons act on other neurons, e.g. ganglion neurons of the visual sensory system. In addition, such neurons also exist in the tactile and olfactory sensory systems. The circuit in this experiment - an example from the visual system - can be used to show the **interconnection of retinal ganglion cells with amacrine cells**:

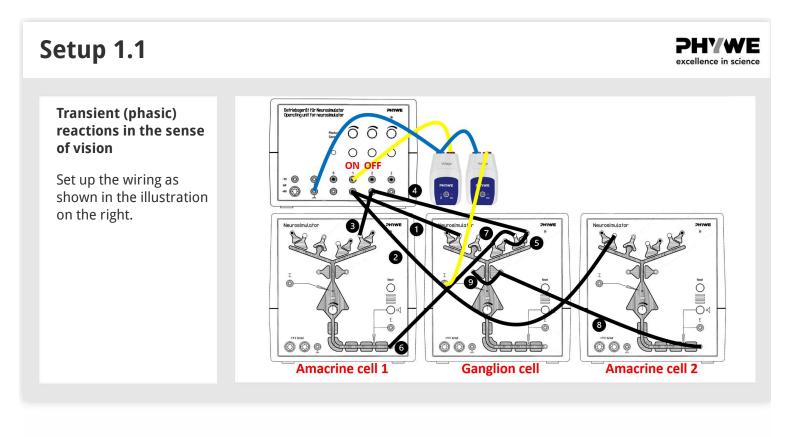
In the retina, the photoreceptors come into contact with bipolar cells, usually several of which are connected to one ganglion cell each. In addition, the photoreceptor cells are interconnected via horizontal cells and the ganglion cells are interconnected via amacrine cells. Each nerve cell thus receives signals from several sensory cells, and each sensory cell transmits signals to several nerve cells. Receptor potentials build up on the bipolar, horizontal and amacrine cells, whereas action potentials are formed first on the ganglion cells.





# Setup and procedure 1





## Setup 1.2

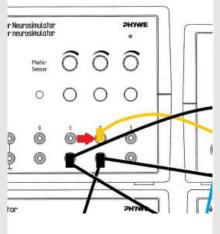




- Neurosimulators 1,2 and 3: Rotary knob firing threshold: 0%
- Power supply: Rotary knob stimulus intensity 1 (stimulus for ON): 100%
- $\circ~$  Power supply: Rotary knob stimulus intensity 2 (stimulus for OFF): 100%

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#### **Procedure 1**



Cable to be switched during the experiment

- Start measurement.
- Press stimulation button 1 for about 5 seconds. Wait until the voltage has reached the initial value.
- Now plug the yellow cable in the black socket in stimulus channel 1 into the black socket of channel 2 (red arrow in the illustration).
- Press stimulation button 2 for approx. 5 seconds. Wait until the voltage has reached the initial value.
- Press stimulation buttons 1 and 2 for approx. 5 seconds. Wait until the voltage has reached the initial value.
- Stop measurement as soon as the voltage has reached the initial value.
- Save and evaluate results.





# **Evaluation 1**



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#### Result 1.1

**ON**: Two cells are involved, the ganglion cell and the amacrine cell, which transmits a signal to the inhibitory synapses of the ganglion cell. The lower cells receive a stimulus at one of their excitatory synapses. The membrane potential of the ganglion cell is determined.

At the beginning of the signal, the membrane of the ganglion cell is depolarised for a short time, which leads to a decrease in the membrane potential. Due to the activity of the inhibitory synapses, hyperpolarisation can be observed.

#### cells receive a excitatory ne potential of the ned. signal, the on cell is time, which leads mbrane potential.

X

3

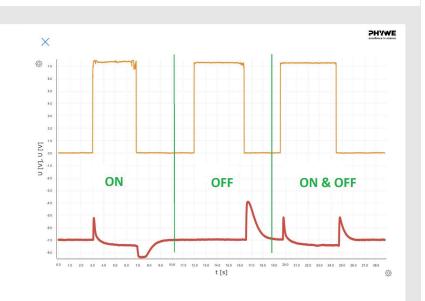
6.0



**OFF**: Two cells are involved: the ganglion cell and the amacrine cell, which transmits a signal to the veto synapses of the ganglion cell. Both cells receive a stimulus at their excitatory synapses. The membrane potential of the ganglion cell is determined.

At the beginning of the signal, nothing happens. However, the membrane of the ganglion cell is depolarised as soon as the stimulus is switched off.

**ON & OFF**: Combines the settings of ON and OFF neurons.



OFF

14.0 15.0 t[s]



**ON & OFF** 

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#### **Theory 2**



#### Neuronal oscillator (body clock)

Many animal and human behaviours have rhythmic characteristics. The periodicity of this rhythmicity, which originates in the central nervous system, can extend over months (e.g. seasonal rhythm), days (e.g. hormonal rhythm), hours (e.g. sleep-wake cycle) or seconds (e.g. the rhythmic movement of many animals). In any case, neuronal oscillators are necessary as timers for such behaviour. The circuit example shows how individual neurons can bring about **oscillating behaviour** when they are grouped together. The rhythmic behaviour of this neuronal network is based on the **time-delayed negative feedback via a veto synapse**. As a result, the input signal is switched off at regular time intervals. Due to the membrane time constant of the neuronal module (caused by its capacitive properties), there is no abrupt signal drop. With a time delay, which is essentially determined by the signal strength, the small excitation at the veto synapse also leads to such a small inhibition again that the stimulation signal can become effective again. I.e. the inhibition at the veto synapse becomes so weak after some time of increasing inhibition strength that it can no longer inhibit the excitatory synapse.





# Setup and procedure 2

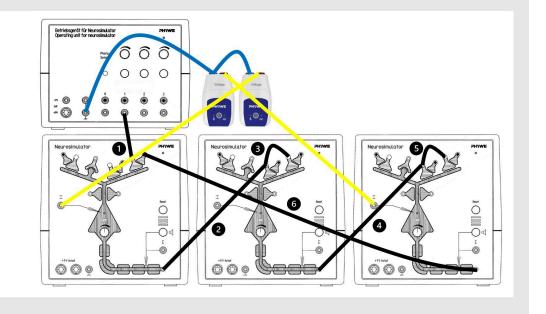


## Setup 2.1

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#### 1. Neural oscillator:

Set up the wiring as shown in the illustration on the right.



#### Setup 2.2

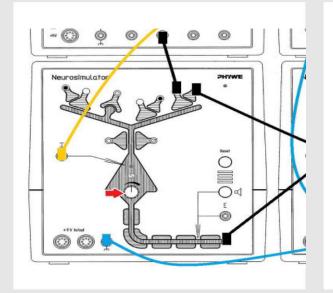
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- Neurosimulators 1, 2 and 3: Rotary knob firing threshold: 0%
- Power supply: Rotary knob stimulus intensity 1: 100%

#### **Procedure 2.1**





- Start measurement.
- Press stimulation button 1 for about 15 seconds.
- Stop measurement as soon as the voltage has reached the initial value.
- Save and evaluate results.

## 2. Transient response with different thresholds on Neurosimulator 1

• Change oscillation by different settings of the threshold on neurosimulator 1 (figure left, red arrow).

## **Procedure 2.2**

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- Start measurement.
- Change the fire threshold in ascending order. Press stimulus button 1 for approx. 5 seconds each time. Wait until the voltage has reached the initial value, then increase the fire threshold again, etc.
- Save and evaluate results.

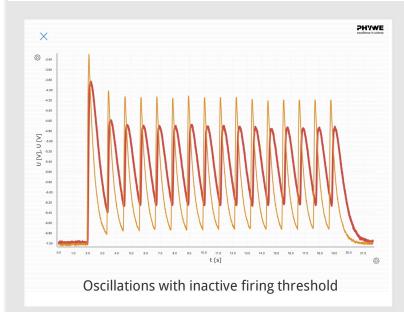
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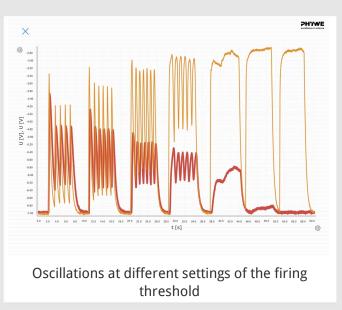
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# **Evaluation 2**

**Results 2** 







#### **Theory 3**



#### **Rotating excitation (short-term memory)**

This self-perpetuating excitation circling within a neuron group is exemplary for short-term memory, because a stimulus remains in this network for a while. Neurosimulators 1 (left) and 2 (in the middle) form a positive feedback loop, i.e. as soon as a brief stimulus has been triggered, the signal is transmitted further. The third Neurosimulator (right) acts as an inhibitory interneuron for Neurosimulator 2. With this setup, different variations can be investigated by changing the stimulus intensity and thresholds of neurons 1 or 2 and the inhibitory interneuron.





# Setup and procedure 3



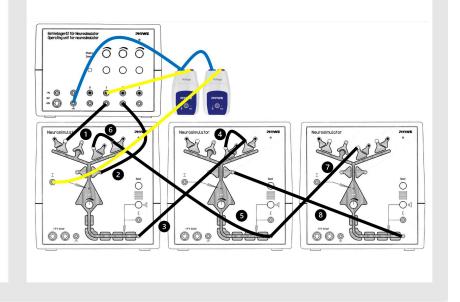
## Setup 3.1

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Notes: Since each person operates push buttons in a slightly different way, to obtain a similar signal duration, all parts of this experiment should be performed by the same person.

#### 1. Change in rotating excitation: Change in stimulus duration

Set up the experiment as shown in the figure on the right.



## Setup 3.2

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- Neurosimulators 1, 2 and 3: Rotary knob firing threshold: 0%
- Power supply: Rotary knob stimulus intensity 1: 50%
- Power supply: Rotary knob stimulus intensity 2: 100%
- Activate "Additional Y-axis" in the measureAPP



#### **Procedure 3.1**

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#### 1.1 Convulsive excitation

- $\circ~$  Start measurement and press stimulus button 1 for about one second.
- Stop measurement as soon as the voltage has reached the output value.
- Variation: Repeat the measurement and end the convulsive excitation by pressing stimulation button 2.
- Save and evaluate results.

#### 1.2 Damping

- Start measurement and stimulus button 1 less than one second (very short tap, like on a hot cooker top).
- Stop measurement as soon as the voltage has reached the output value.
- Save and evaluate results.

## **Procedure 3.2**



#### 2. Change in rotating excitation: Change in stimulus intensity

- Start measurement.
- Press stimulation key 1 for less than one second (very short tap, like on a hot cooker top).
- Stop measurement as soon as the voltage has reached the initial value. Save results.
- Repeat the measurement several times. Minimally change stimulus intensity for each measurement: Turn the knob for stimulus intensity 1 minimally, both directions are possible. The aim is to obtain measurement diagrams with convulsive excitation and with damping.
- Save and evaluate the results each time. Before each measurement, stop the convulsive excitation by pressing the stimulation button 2.

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#### **Procedure 3.3**

#### 3. Change in rotating excitation: Inhibition

- Start measurement.
- Press stimulation key 1 for approx. one second.
- $\circ~$  Press stimulation key 2.
- Stop measurement as soon as the voltage has reached the initial value. Save result.





# **Evaluation 3**



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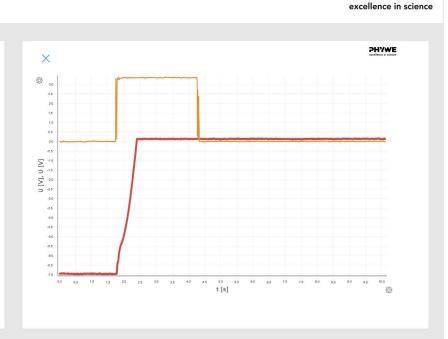
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#### **Result 3.1.1**

1. Variation of the rotating excitation: Variation of the stimulus duration

#### **1.1 Convulsive excitation**

The figure on the right shows the first example of convulsive excitation (upper curve: stimulus, lower curve: membrane potential of nerve cell 1).



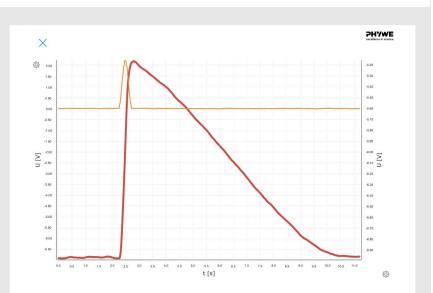
#### **Result 3.1.2**

#### 1.2 Damping

The illustration on the right shows an example of attenuation after a very short stimulus.

In this experiment, the stimulus channel is medium strong and only a short stimulus impulse is given. The nerve membrane is depolarised, but the stimulus is too weak and the membrane potential returns to baseline.

Since the threshold of the interneuron is low, its inhibition increases the attenuation of the neuron-neuron cycle.





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## Result 3.2

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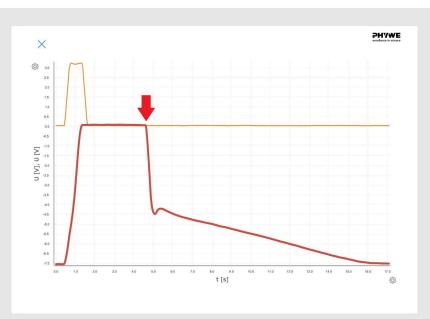
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| 2. Change in rotating<br>excitation: Change in<br>stimulus intensity | Number | signal intensity $U_{max}$ | signal duration $t_2-t_1$ | convulsive<br>excitation | dampening |
|--|--------|----------------------------|---------------------------|--------------------------|-----------|
|  | 1.     | 6.28                       | 0.12                      | x                        |           |
| The table shows examples of  | 2.     | 5.05                       | 0.11                      | x                        |           |
| different values for convulsive<br>excitation and attenuation        | 3.     | 5.24                       | 0.10                      | x                        |           |
| depending on stimulus  | 4.     | 5.09                       | 0.11                      | х                        |           |
| intensity and duration.  | 5.     | 4.96                       | 0.11                      |                          | x         |
|  | 6.     | 4.95                       | 0.11                      |                          | x         |
|  | 7.     | 4.89                       | 0.11                      |                          | x         |
|  | 8.     | 5.08                       | 0.09                      |                          | ×         |
|  | 9.     | 5.09                       | 0.08                      |                          | x         |
|  | 10.    | 3.65                       | 0.10                      |                          | x         |
|  |        |                            |                           |                          |           |

## Result 3.3

## 3. Change in rotating excitation: Inhibition

Depending on the length of the inhibitory stimulus (red arrow in the figure on the right), the convulsive excitation subsides more or less quickly.





#### **Theory 4**



#### Cerebral cortex and sensory learning

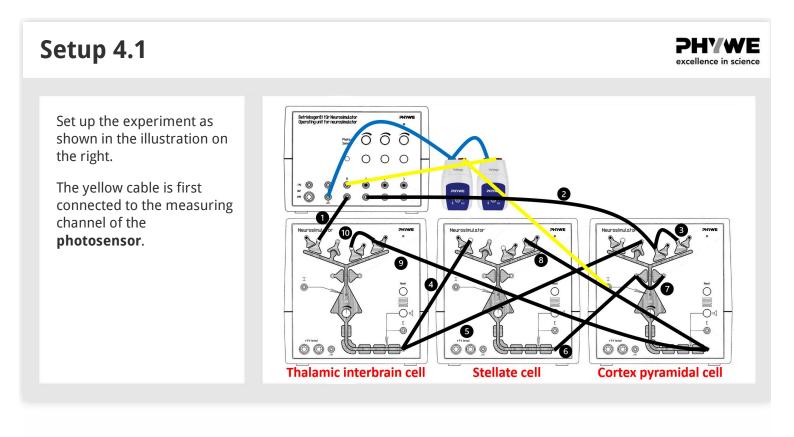
This experiment shows that a cortical pyramidal cell can only react in a stimulus-specific way if it has learned to do so through a previous correlation of the sensory signal with a non-specific alarm signal. In this way, only important signals are stored and processed, and an **overload of the cerebral cortex** is avoided. The photosensor provides the specific signal of a sensory organ, which is a **thalamic interbrain cell** which in turn excites the Hebbian synapse of a **cortex pyramidal cell**. The cortex pyramidal cell is also excited by a non-specific stimulus (channel 1). At the same time, a **stellate cell** inhibits the pyramidal cell. These **three cell types** form the **triad**. The mammalian cerebral cortex is where sensory signals are processed and linked to motor programmes. In addition, this is where **experience storage** takes place. In an early stage of development, the cerebral cortex is in a diffuse, unformed state in which signal processing does not function with the precision that it does in the adult organism. This ability is only slowly acquired through active engagement with the environment. Stimulus processing is also a result of **plastic adjustment**. In this process, coincidence between different stimuli is always necessary to reach a cortical pyramidal cell and stabilise Hebbian synapses there through sustained potentiation.





# Setup and procedure 4





#### Setup 4.2





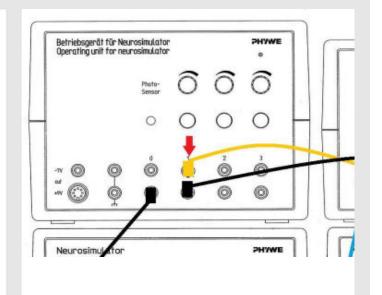
- Neurosimulators 1, 2, 3: Rotary knob firing threshold: 50%.
- Power supply: Rotary knob stimulus intensity 1: 100%

#### **Procedure 4.1**

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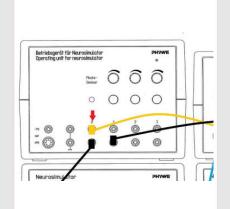
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- To set the Hebbian synapse to the default values, press the reset button on neurosimulator 3 (cortex pyramidal cell).
- The yellow cable is plugged into the measuring channel of the p**hotosensor**.
- Start measurement.
- Cover the photosensor for 3 seconds. Wait until the voltage has reached the initial value.
- Yellow cable on s**timulus channel 1** (red arrow in the illustration on the right).

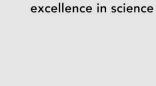


## **Procedure 4.2**

- Press stimulation button 1 for approx. 3 seconds. Wait until the voltage has reached the initial value. Reconnect the yellow cable to the measuring channel of the **Photosensor** (red arrow in the illustration on the right).
- Cover the photosensor and press stimulus button 1 (simultaneously) for
  60 seconds. Wait until the voltage has reached the initial value.
- Cover the photo sensor for 3 seconds. Wait until the voltage has reached the initial value. Finish the measurement, save and evaluate the results.
- To reset the Hebbian synapse to the default value, press the reset button on neurosimulator 3 (cortex pyramidal cell).





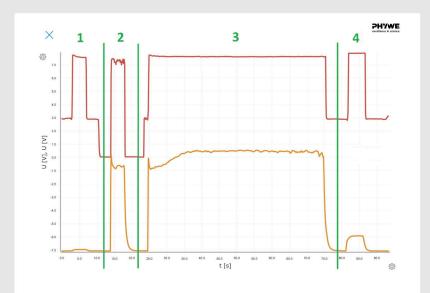


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# **Evaluation 4**

#### **Result 4**



The excitation by the specific stimulus (photosensor, measuring range 4) after simultaneous actuation of the photosensor and stimulus button 1 is greater than in measuring range 1 and decays more slowly than the excitation by stimulus button 1 in measuring range 2.

Conclusion: A cortex pyramidal cell (measuring range 1) can only react in a stimulus-specific manner (measuring range 4) if it has learned to do so through a previous correlation of the sensory signal with a non-specific alarm signal (measuring range 3).





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#### **Theory 5**



#### Directional selectivity through unilateral inhibition

Examples: Many nerve cells in sensory systems are direction-selective. For example, certain ganglion cells in the retina only react when a light stimulus moves in a certain direction, but not when it moves in the opposite direction. A similar behaviour is also known for the sense of touch. This neural circuit can be simulated by a circuit with one-sided inhibition between two stimulus channels that are activated one after the other. Local projection sensory organs, e.g. the retina of the eye or the surface of the body with its tactile receptors, are basically able to encode movements (changes in position over time). Accordingly, neurons are actually found in these sensory channels that respond selectively to stimulus movements (see also on-off response). Some of these cells do not simply respond to every movement, but respond only to certain directions of movement, while other directions remain unresponsive. Their response intensity is usually a function of the speed of movement, the direction of movement and - usually to a lesser extent - the intensity of the stimulus being moved. Such cells are already found in the retina of most vertebrates, but only in a few of them at a higher processing level (e.g. anthropoids).





# Setup and procedure 5



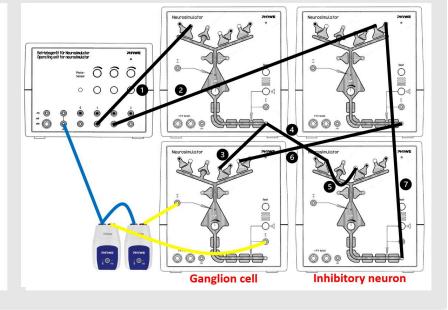
## Setup 5.1

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Set up the experiment as shown in the illustration.

The measuring channel for the action potentials in the ganglion cell can be connected to an oscilloscope. Alternatively, both voltage sensors can be connected to a computer via USB to increase the measuring frequency (5000 Hz per channel instead of 500 Hz with Bluetooth operation). However, the easiest way to carry out the experiment is to use the acoustic monitor to display the action potentials.



## Setup 5.2



- Neurosimulators 1, 2, 3, 4: Rotary knob firing threshold: 0%
- $\circ~$  Power supply: Rotary knob stimulus intensity 1 and 2: 50% each

## **Procedure 5**



- Activate only the acoustic monitor of the ganglion cell (Neurosimulator 4).
- Then increase the threshold of the ganglion cell by emitting a stimulus only from stimulus channel 1 so that no action potential is audible (acoustic monitor of the ganglion cell). Check whether an action potential is audible when stimulus channel 2 is activated. No acoustic signal should be heard.
- Part 1: Start measurement. Press **stimulus button 1** and immediately afterwards (almost simultaneously) press **stimulus button 2**. End measurement. Save and evaluate results.
- Part 2: Start measurement. Press **stimulus button 2** and immediately afterwards (almost simultaneously) press **stimulus button 1**. End measurement. Save and evaluate results.





# **Evaluation 5**



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#### Result 5.3

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Measurement with the measureLAB software, which can also be used.

Only in the second part of the measurement is an action potential (AP) generated, the consequence of direction selectivity.

## **Theory 6**



#### Self-calibration of paired sensory channels

Example: The embryonic formation of axisymmetric species is not perfect, resulting in slight irregularities in symmetry. Irregularities of the sensory epithelia, e.g. in the organ of equilibrium, can be compensated for by the nervous system: Hebb's principle offers the possibility of adjusting the output signals so that they are symmetrical when the sensory organs are asymmetrical.

Experimental setup: There are two sensory neurons and two interneurons (two sensory neuron - interneuron pairs). Asymmetrical signals are sent from stimulus channel 2 and stimulus channel 3 to the Hebbian synapses of the two sensory neurons (here: stimulus intensity 50% and 100% respectively). A signal originating from stimulus channel 1 of the operating unit is sent to the two interneurons, which transmit the signal via their efferent axon to their sensory neuron pair, where the signal is amplified by branching. Both sensory neurons inhibit their own interneurons.



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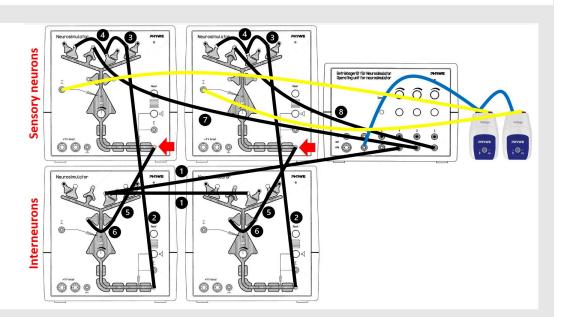


# Setup and procedure 6

## Setup 6.1

#### Self-calibration of paired sensory channels:

- Set up the wiring as shown in the illustration on the right.
- For the second measurement, leave one of the two cables marked with the red arrows unplugged.





#### Setup 6.2

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- Neurosimulator 1, 2, 3, 4: Rotary knob firing threshold: 0%
- Power supply: Rotary knob stimulus intensity 1: 33%
- Power supply: Rotary knob stimulus intensity 2: 50%
- Power supply: Rotary knob stimulus intensity 3: 100%

## **Procedure 6.1**

#### Self-calibration of paired sensory channels

- Start measurement.
- Press stimulation key 1, wait 1 to 2 seconds, keep key pressed and additionally press stimulation keys 2 and 3 simultaneously.
- Press and hold the three stimulus buttons simultaneously for a longer time (e.g. 30 seconds).
- End measurement when the voltage has reached the initial value.
- Save and evaluate results.

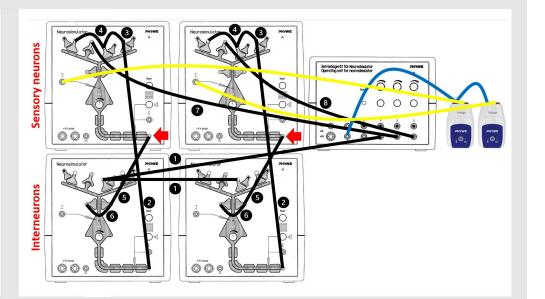


## **Procedure 6.2**

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#### No self-calibration

- Remove one of the cables marked with a red arrow from the black socket.
- Otherwise, perform the same procedure as for the experiment "Selfcalibration of paired sensory channels".



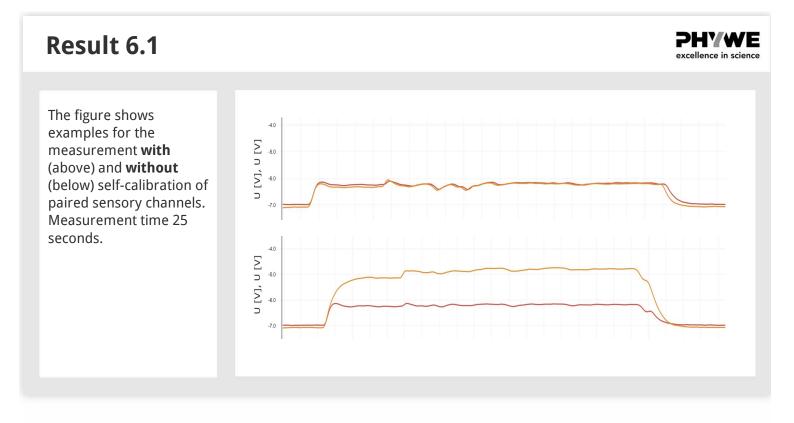




# **Evaluation 6**



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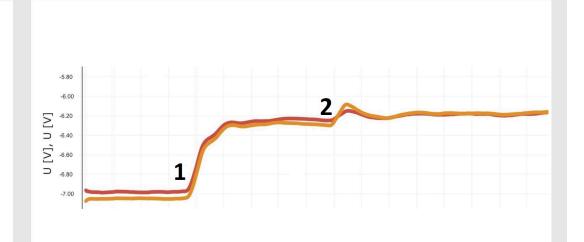


## Result 6.2

The illustration on the right shows the start of self-calibration:

Action 1: Pressing the stimulus button 1

Action 2: Stimulus button 1 remains pressed, additional simultaneous pressing of stimulus buttons 2 and 3





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# Outlook

## Outlook

#### Additional neural networks

More circuits can be created with data from various research projects. An internet search term such as "synaptic potentiation" yields a considerable amount of scientific papers. Their neuronal models can be converted into neurosimulator circuits.

Please note that if you want to use more than 4 Neurosimulators, you need another power supply to supply power to the additional Neurosimulators, i.e. 1 additional power supply per 4 Neurosimulators.



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