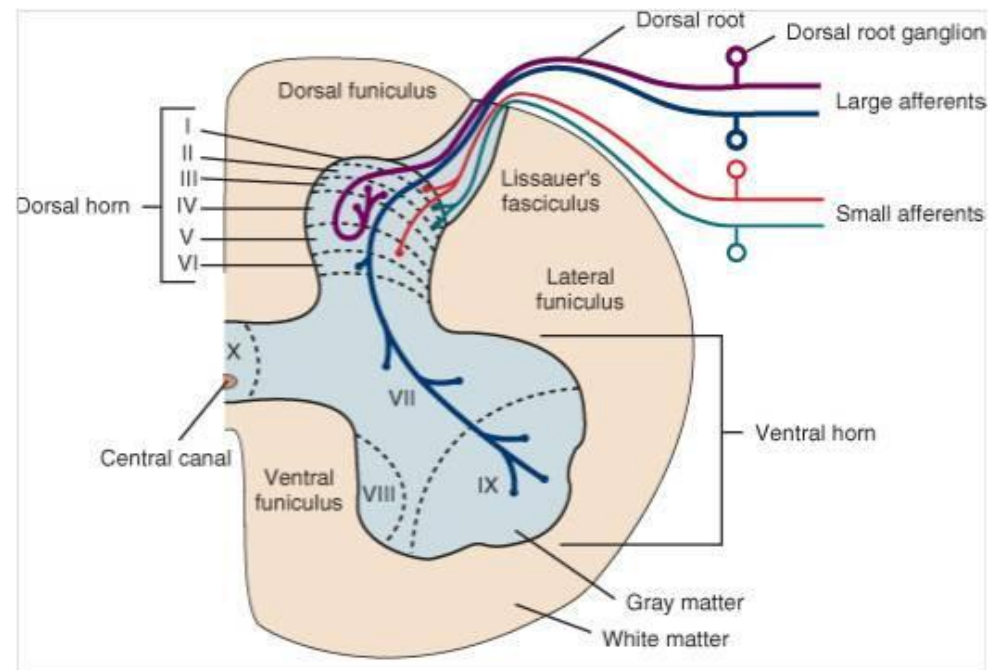




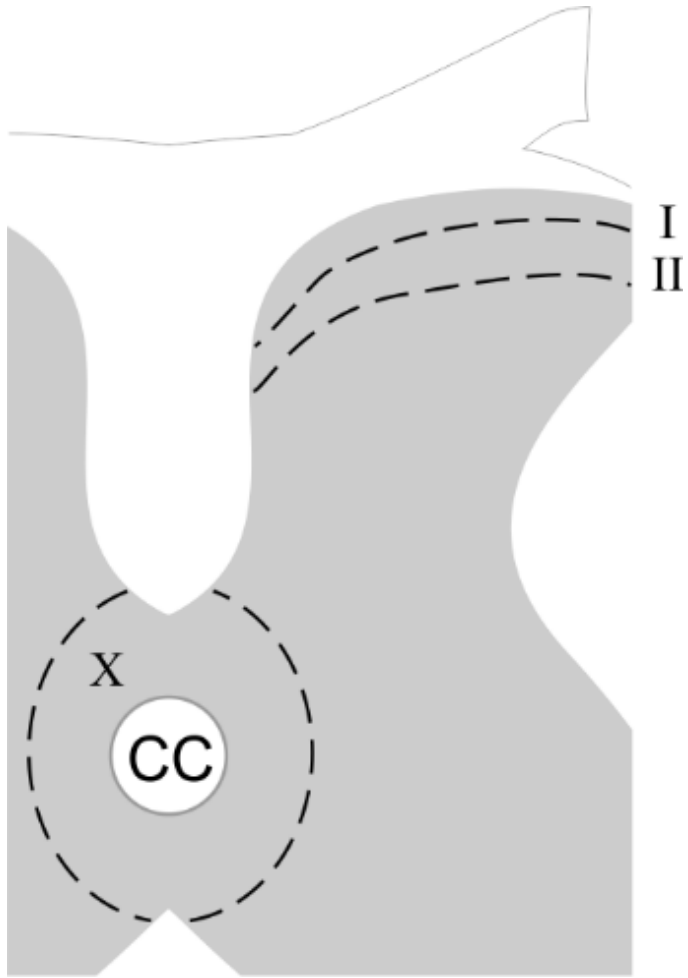
The role of lamina X neurons in nociception

Volodymyr Krotov



Source: www.studyblue.com

Multiple physiological functions of spinal lamina X neurons

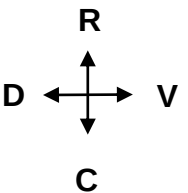
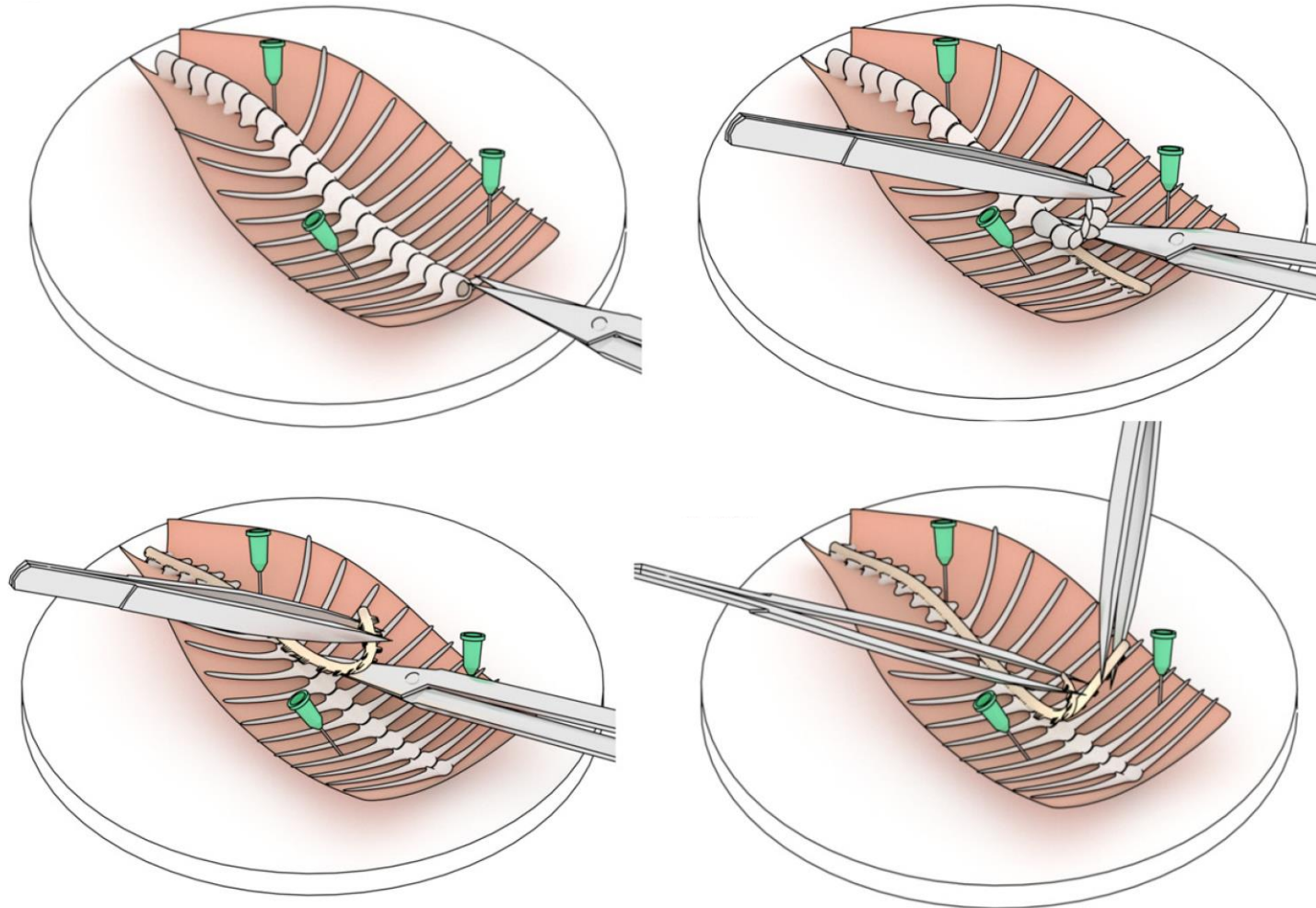


- **Visceral nociception**
- Somatosensory integration
- Autonomic regulation
- Modulation of motor neuron output

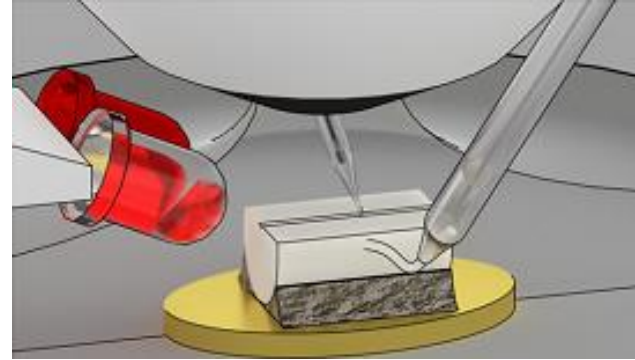
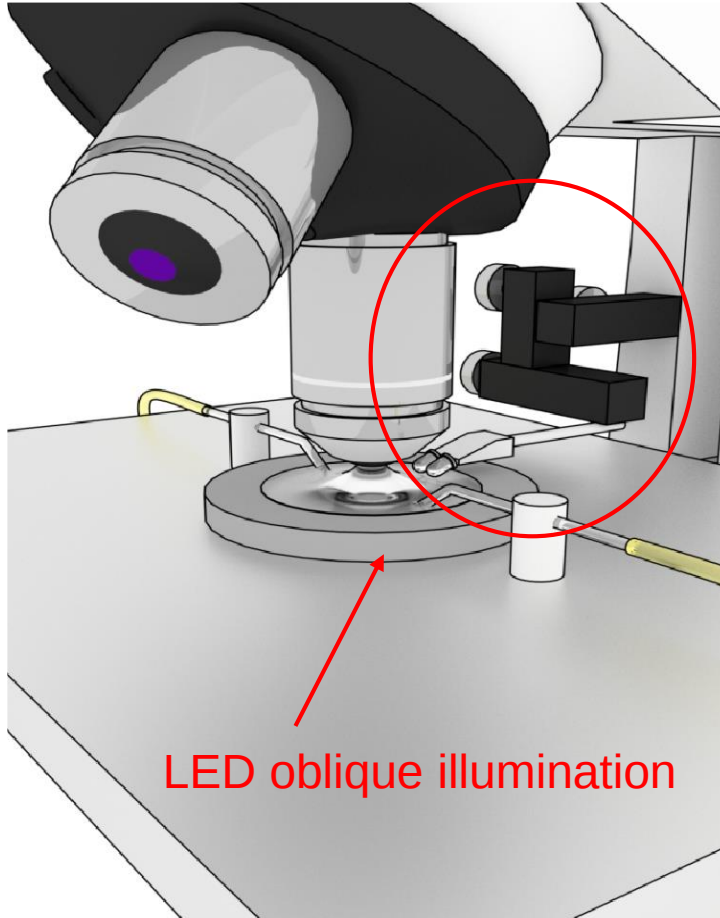
Experimental approach

Experimental approach: *ex-vivo* spinal cord preparation

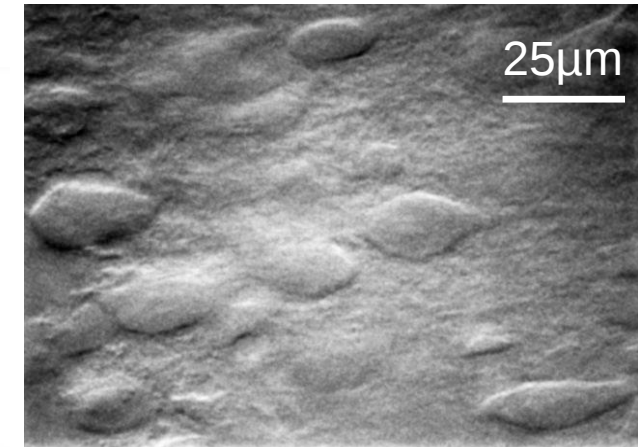
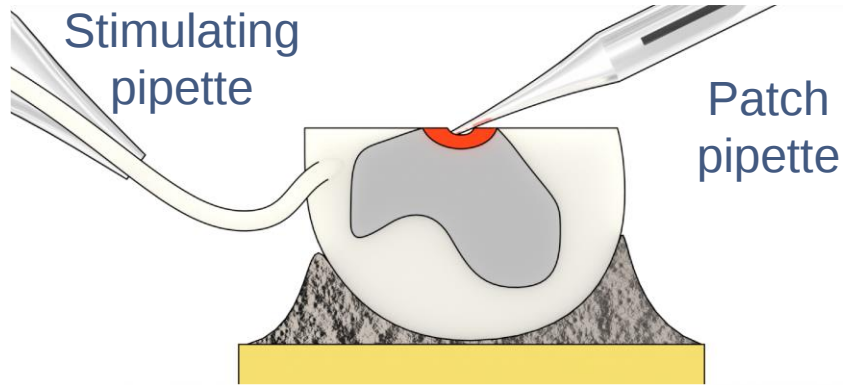
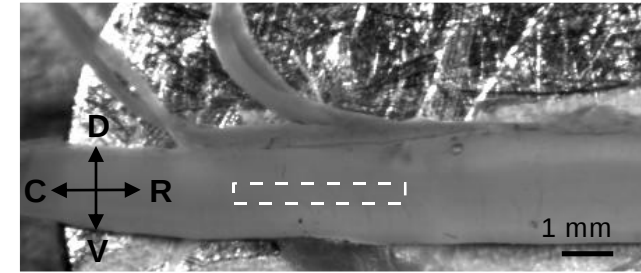
Hemisected spinal cord with spared dorsal & ventral roots



Experimental approach: LED oblique illumination technique

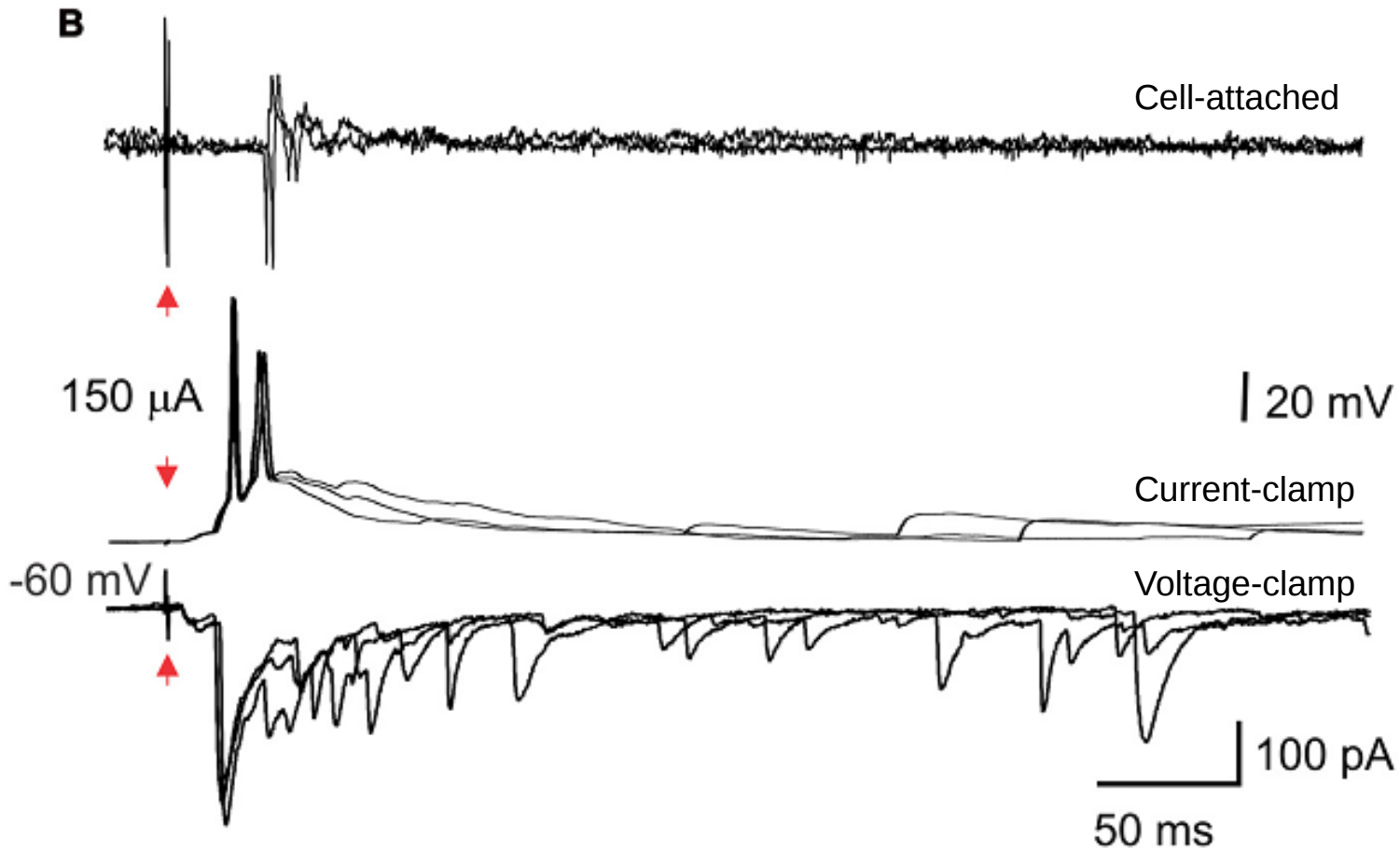


Hemisected spinal cord with spared dorsal roots

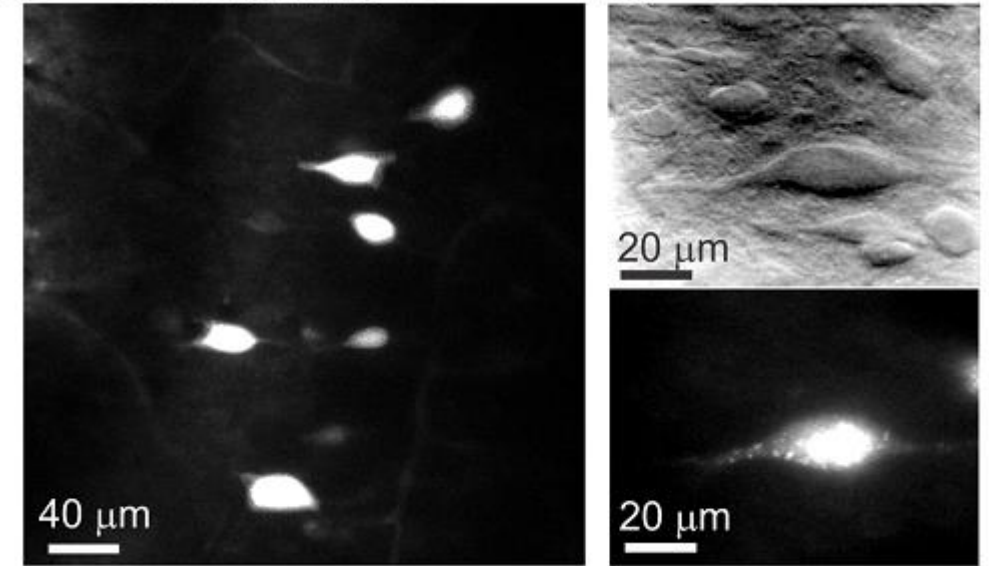


Experimental approach: electrophysiology & fluorescent labeling

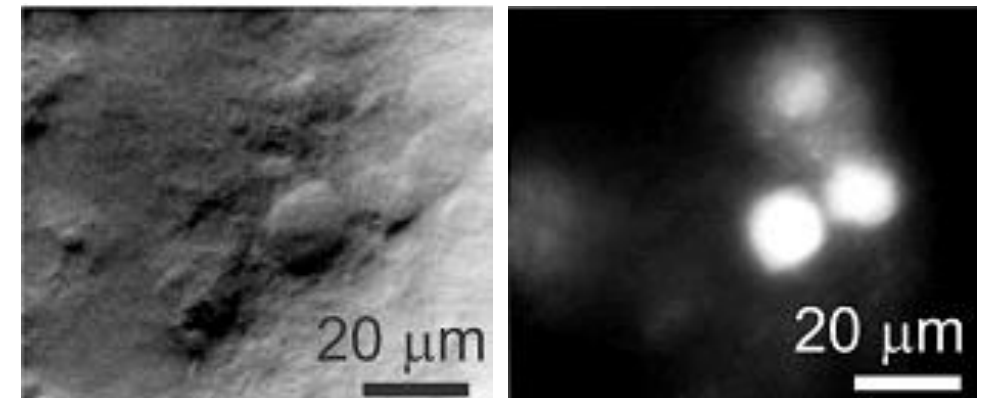
Electrophysiological experiments



Spino-thalamic projection neurons

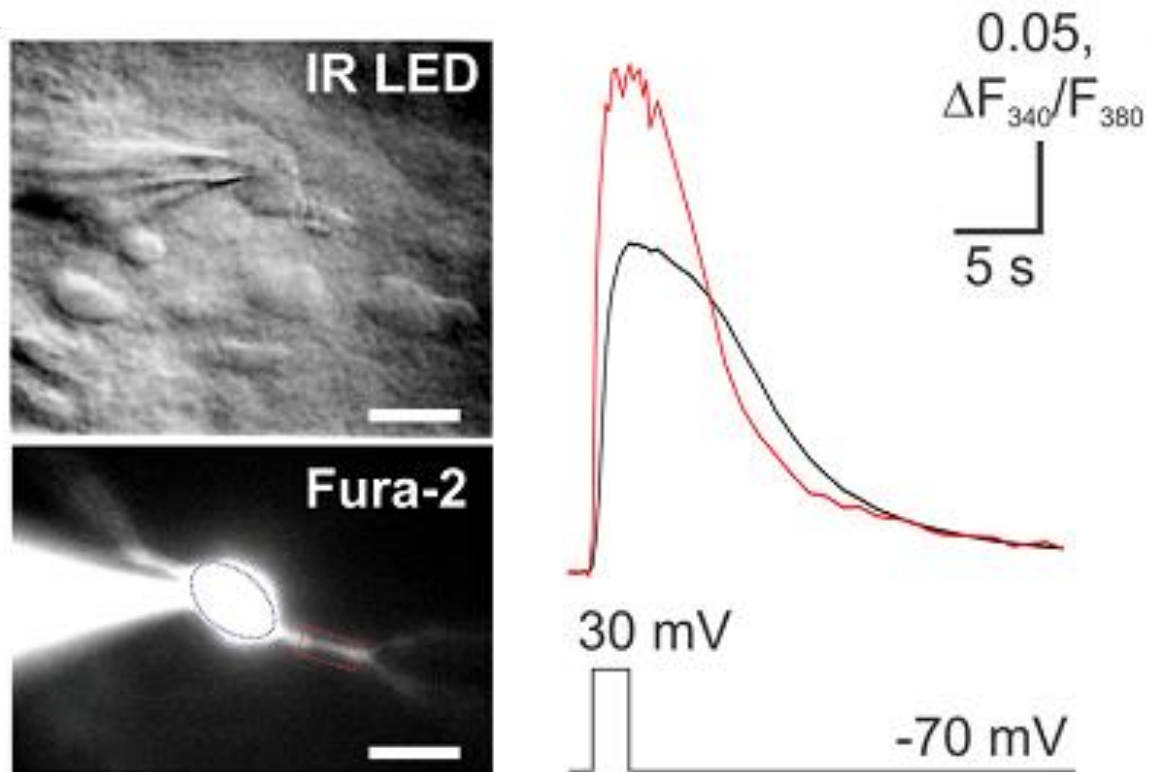


Sympathetic preganglionic neurons

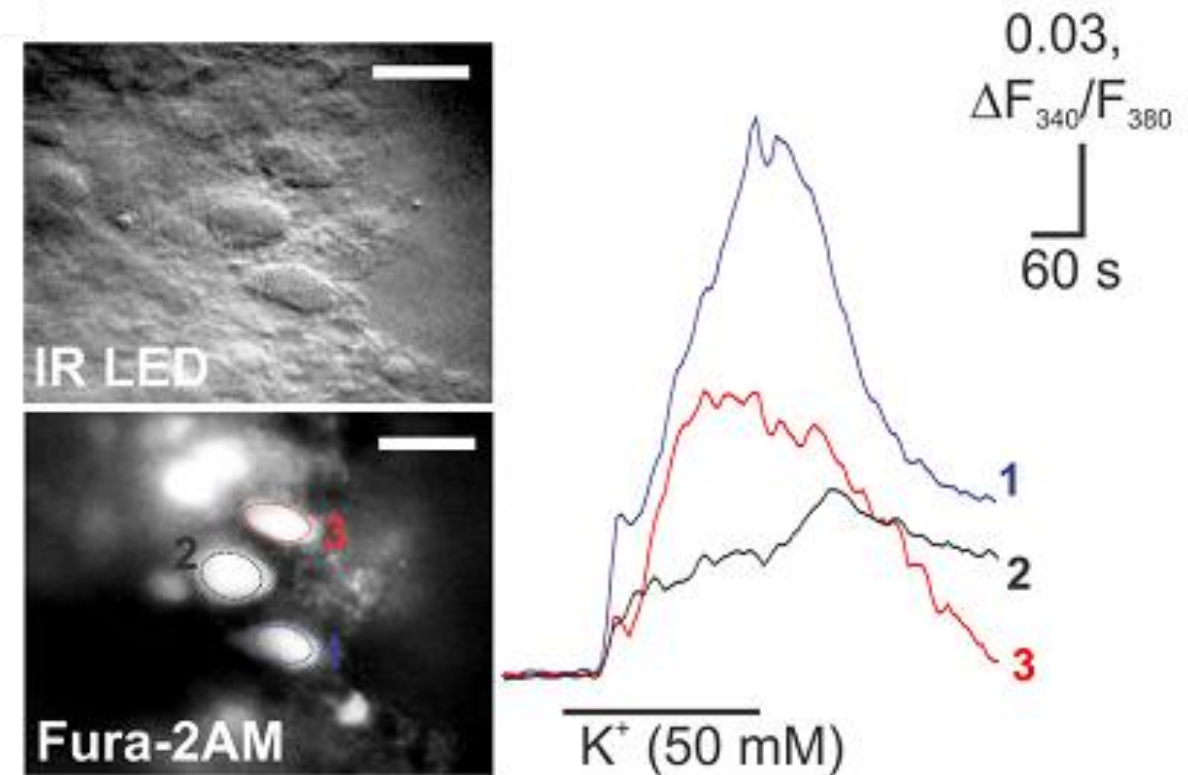


Experimental approach: calcium imaging

Cell-impermeable calcium dye

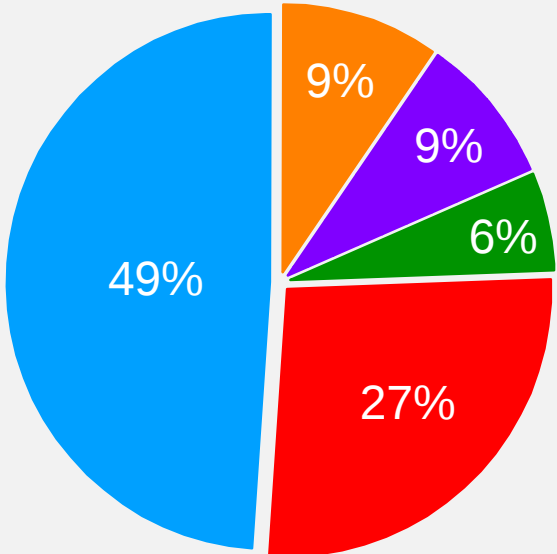
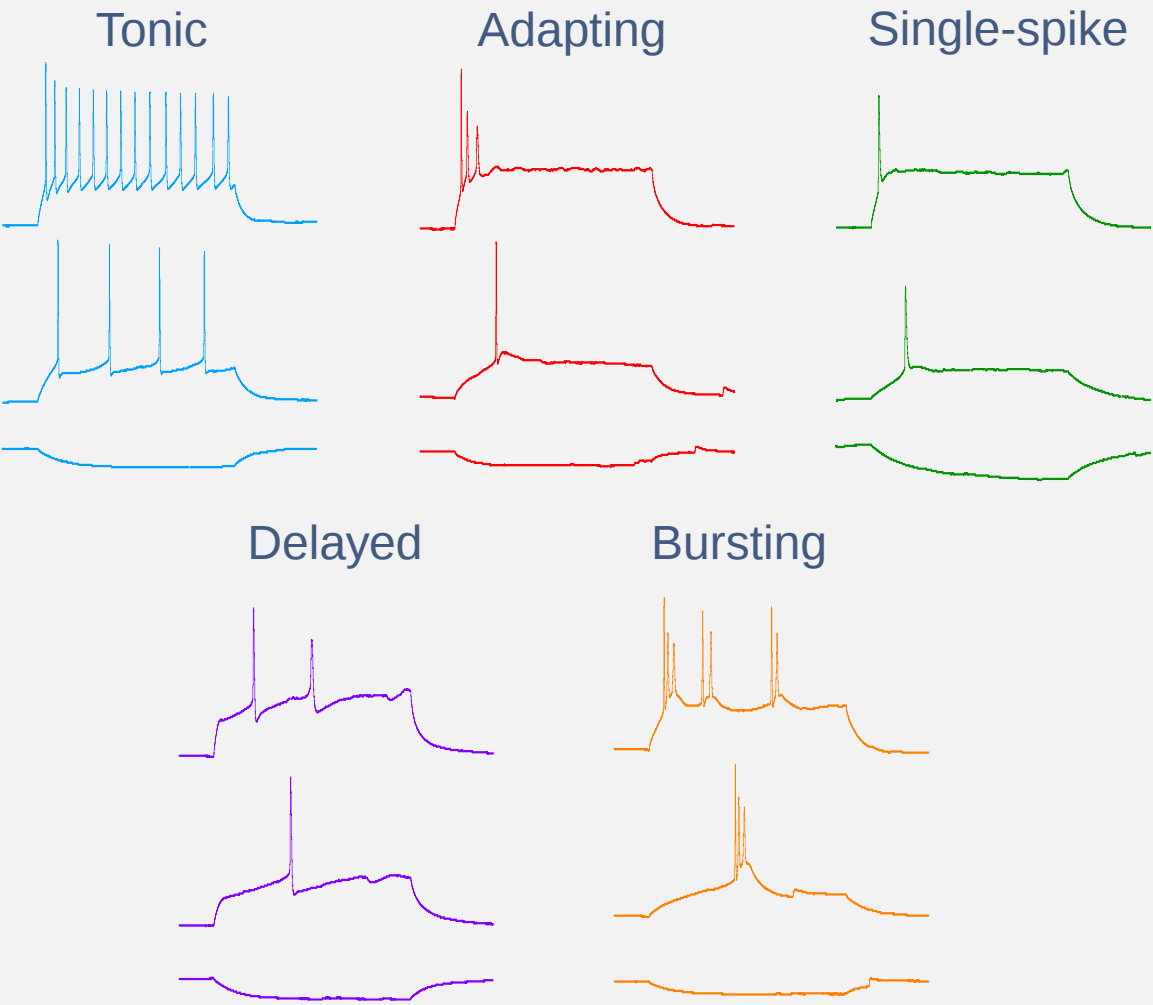


Cell-permeable (AM) calcium dye



Intrinsic properties and afferent supply of lamina X neurons

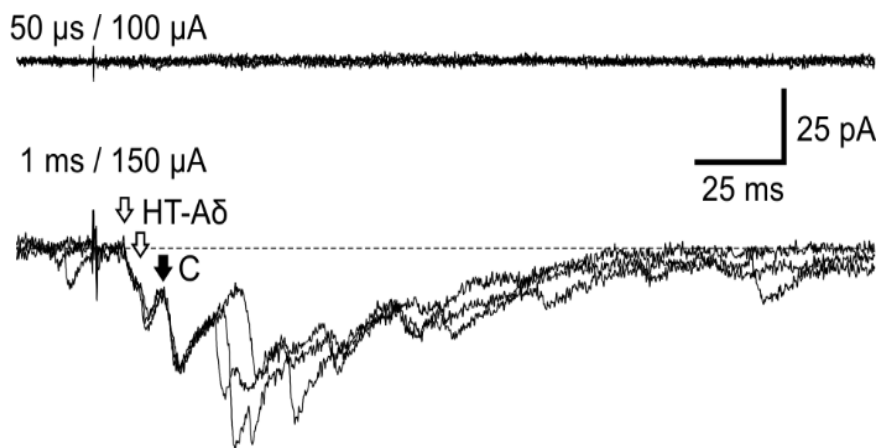
Diversity of lamina X neurons



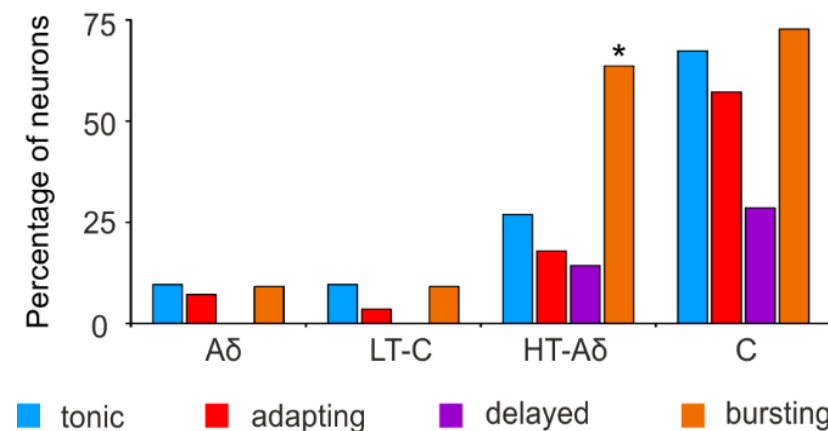
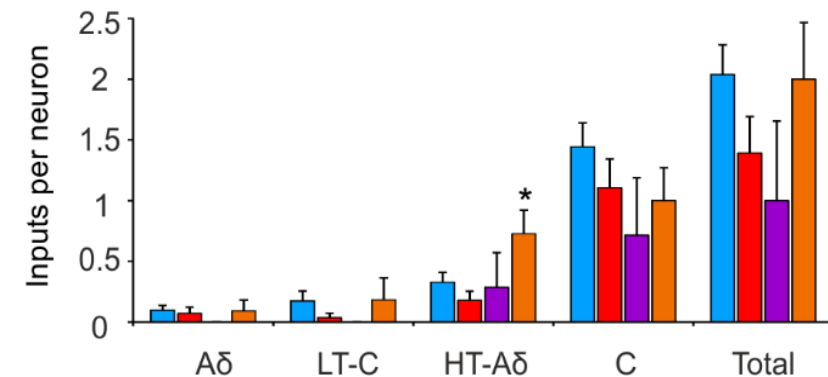
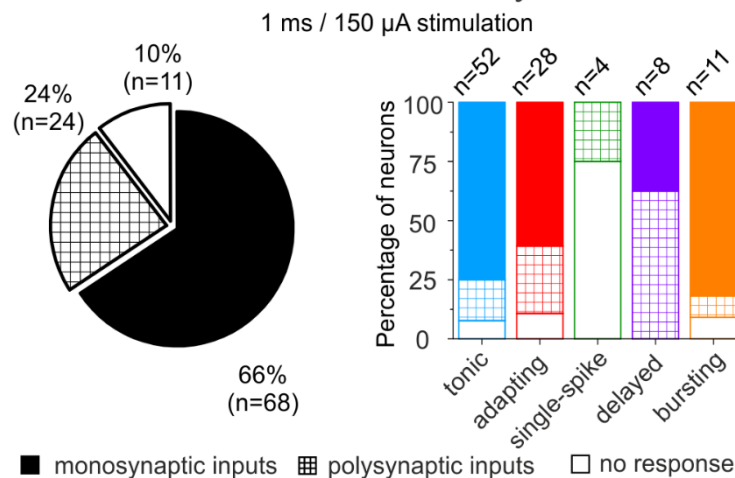
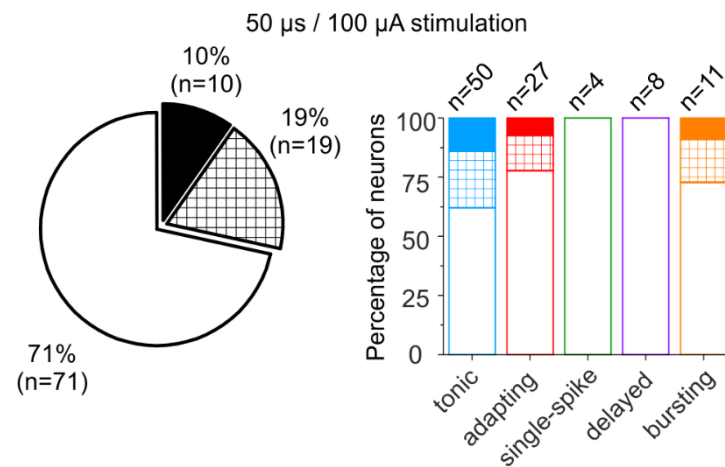
Lamina X neurons show many features common to the pain-processing neurons from the superficial dorsal horn.

Firing pattern	Capacitance	Resistance	RMP
Tonic (n=66)	43 ± 2 pF	0.69 ± 0.06 GΩ	-59 ± 1mV
Adapting (n=36)	27 ± 3 pF	1.03 ± 0.15 GΩ	-54 ± 2 mV
Single-spike (n=8)	32 ± 2 pF	0.76 ± 0.11 GΩ	-51 ± 2 mV
Delayed (n=12)	64 ± 12 pF	0.48 ± 0.15 GΩ	-56 ± 2 mV
Bursting (n=13)	71 ± 9 pF	0.45 ± 0.07 GΩ	-60 ± 1 mV

Primary afferent inputs to lamina X neurons



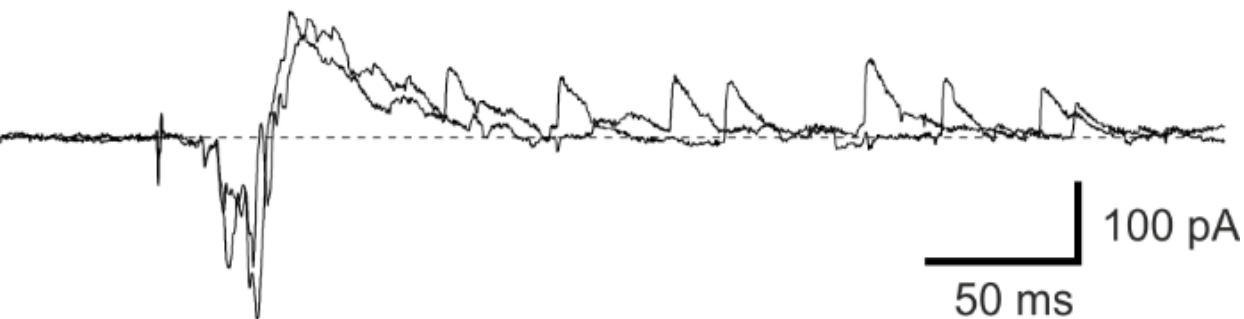
The majority of lamina X neurons receive functional monosynaptic inputs from the high-threshold A δ and C afferents.



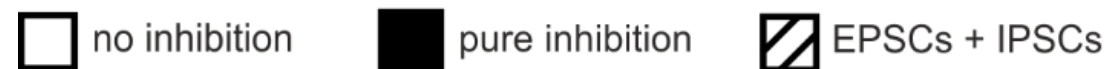
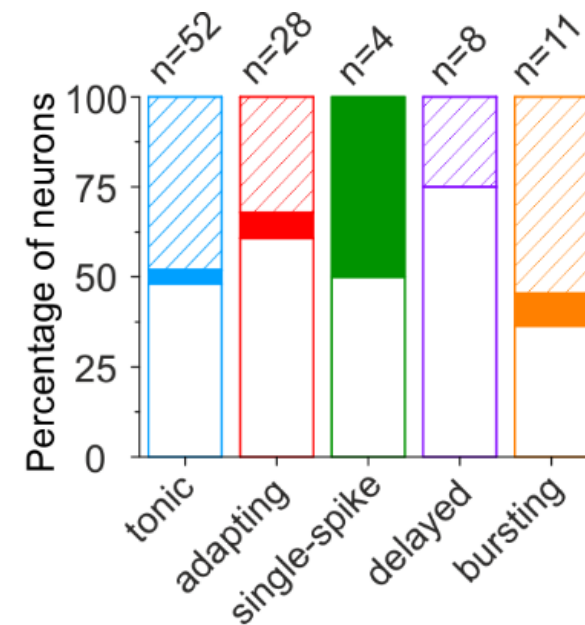
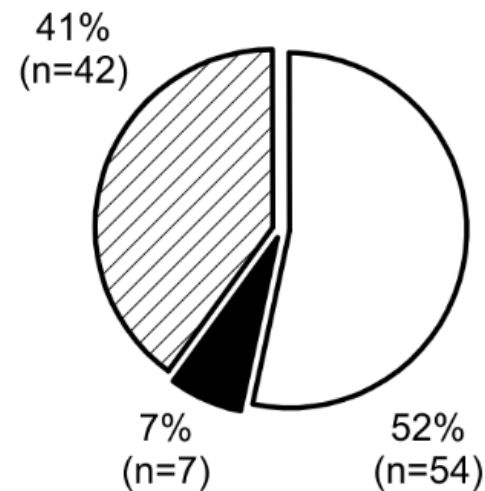
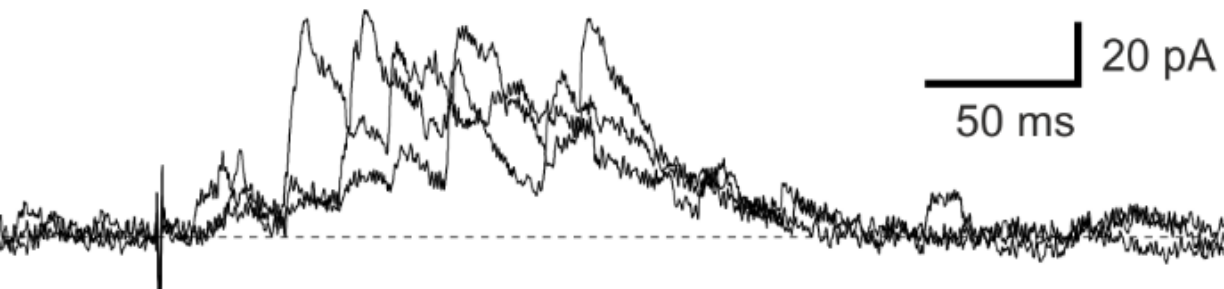
Inhibitory components of primary afferent input

About a half of lamina X neurons show primary-afferent-driven polysynaptic inhibition.

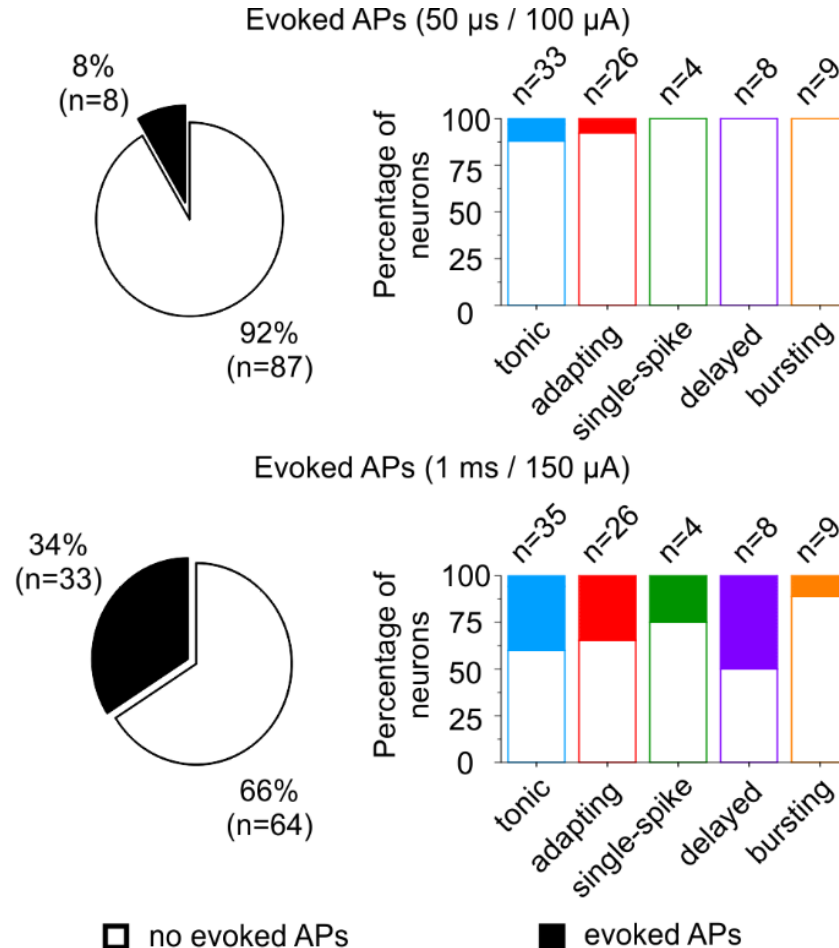
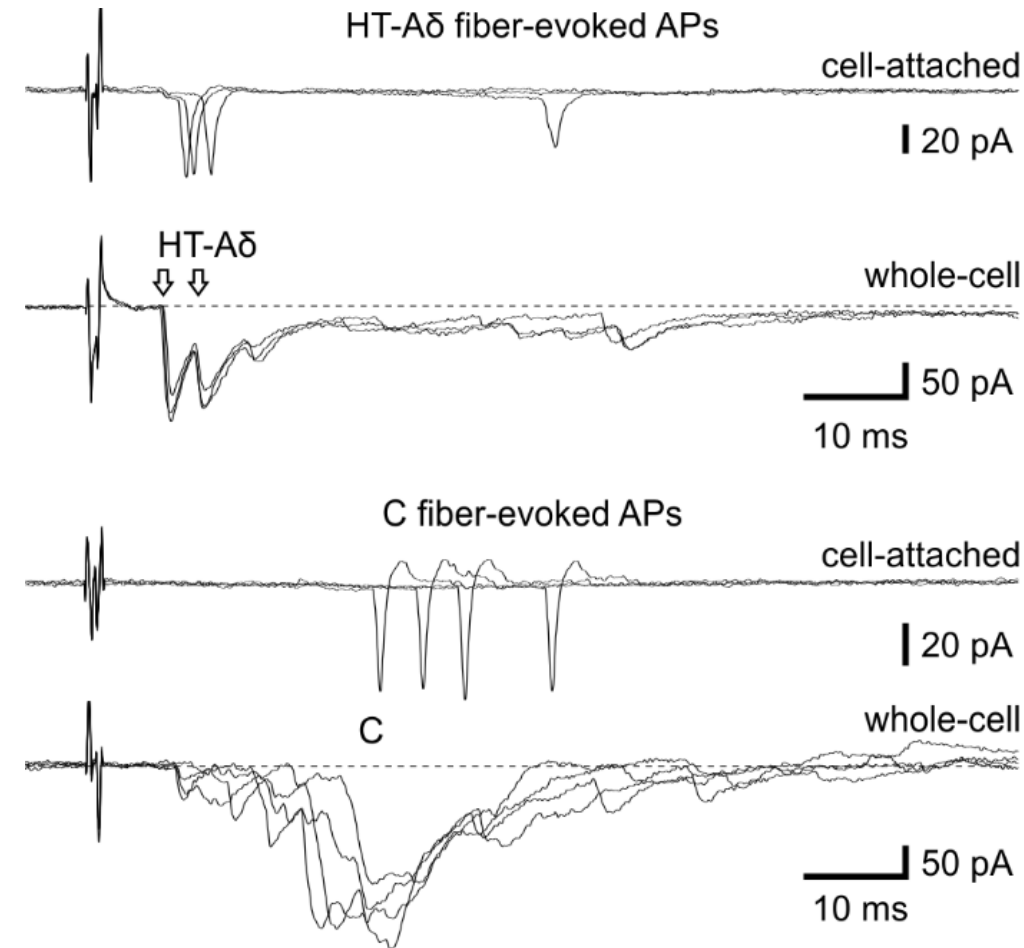
EPSCs + IPSCs



Pure inhibition

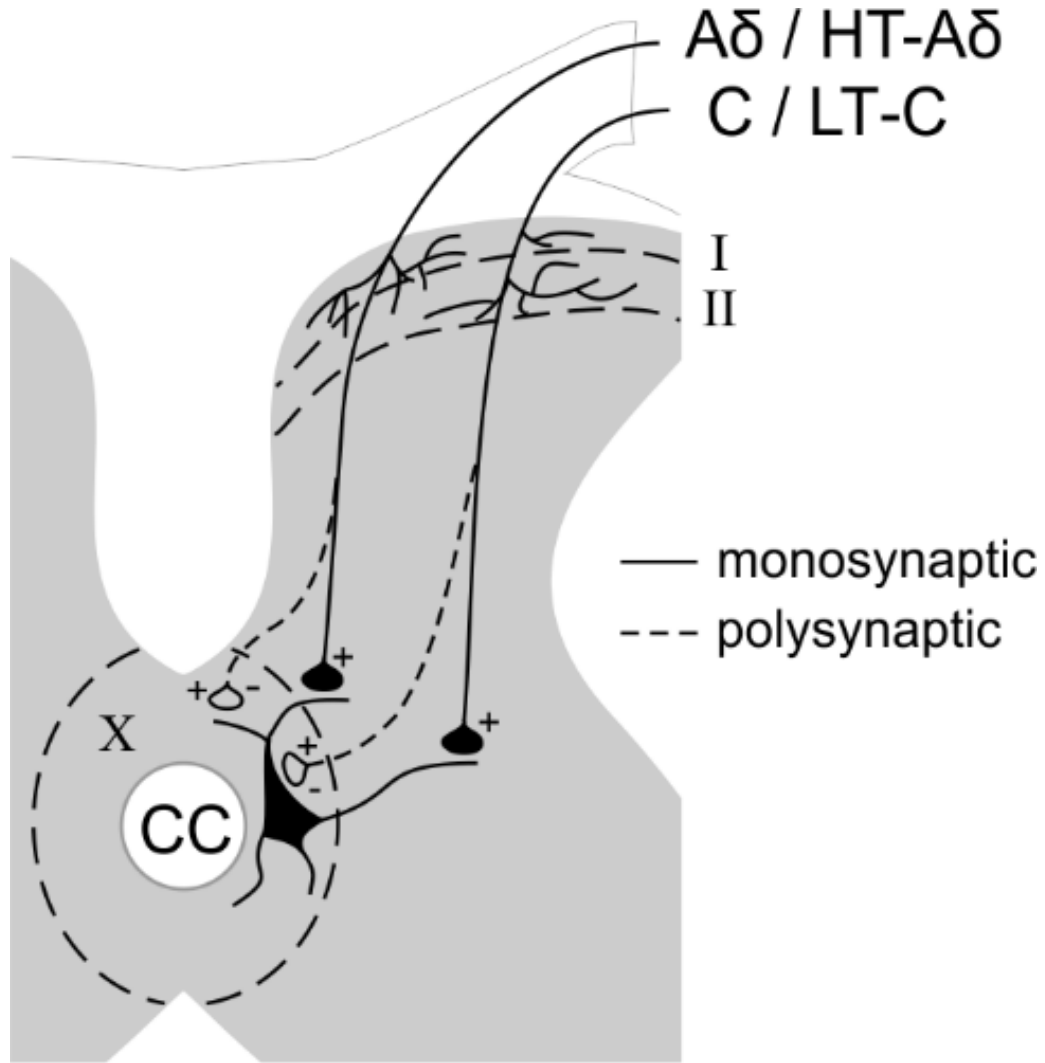


Afferent-driven firing of lamina X neurons



About 1/3 of lamina X neurons fire in response to the activation of high-threshold primary afferents.

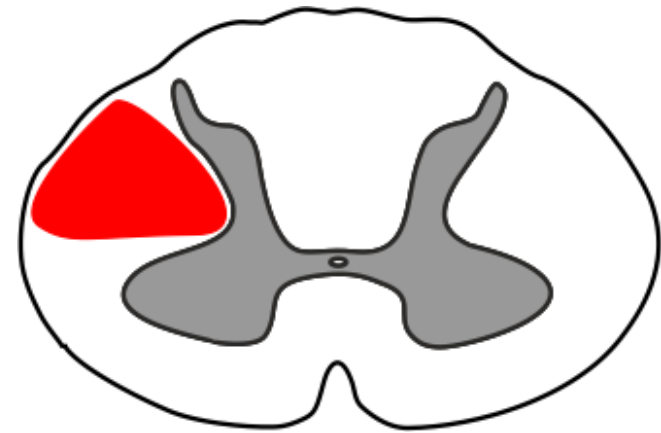
Organization of primary afferent inputs to lamina X neurons



Lamina X neurons are active elements of the intrinsic spinal network, that are directly targeted by nociceptive afferent fibers and process diverse modalities of peripheral input.

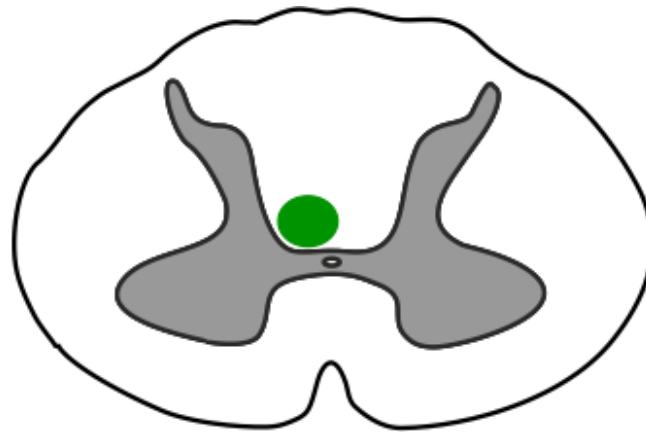
Descending modulation of lamina X neurons

Descending tracts of the spinal cord



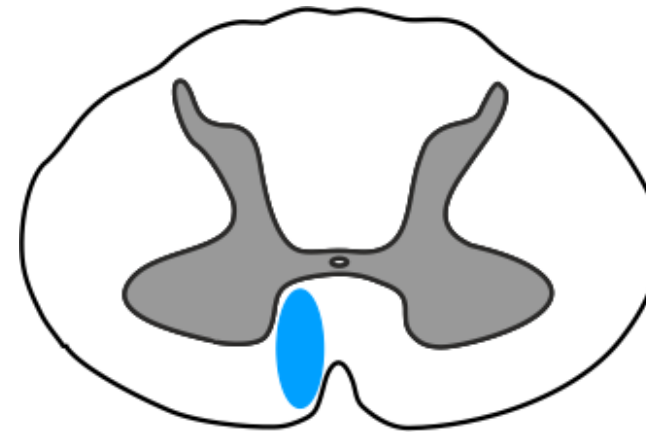
**Dorsolateral
funiculus (DLF)**

- Lateral vestibulospinal tract
- Rubrospinal tract
- Reticulospinal tract
- Axons from RVM and PBNs (GABA, 5-HT, NE, enkephalins)



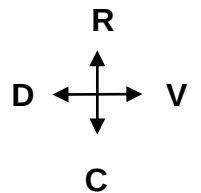
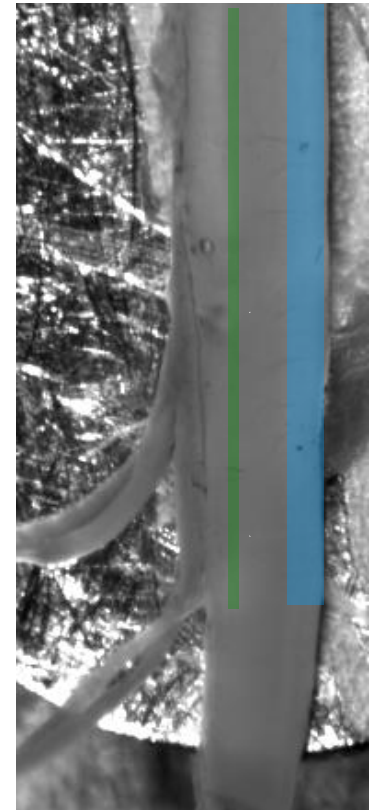
**Posterior
funiculus**

- Main corticospinal tract (CST)



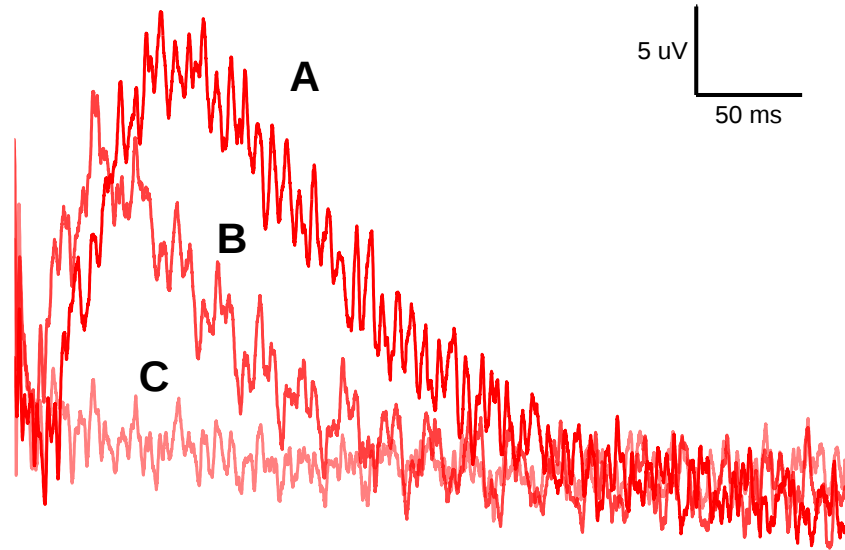
**Anterior
funiculus (AF)**

- Corticospinal tract (CST)
- Reticulospinal tract
- Medial vestibulospinal tract



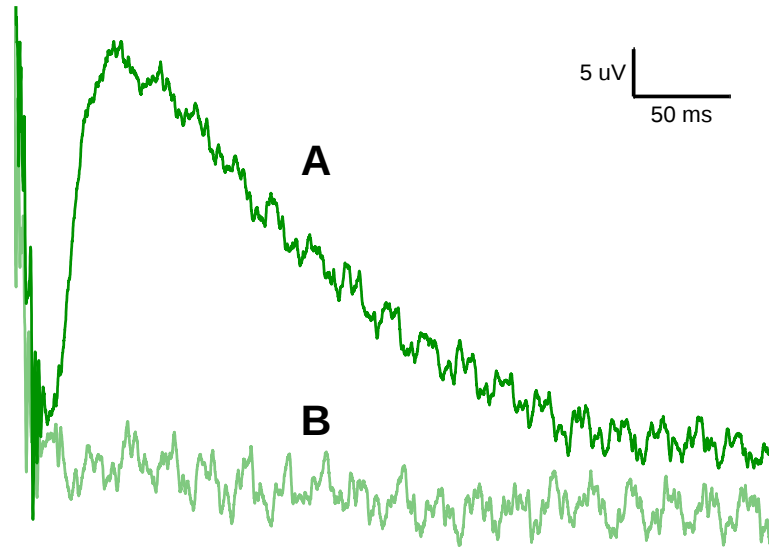
DRPs in response to stimulation of descending tracts

DLF



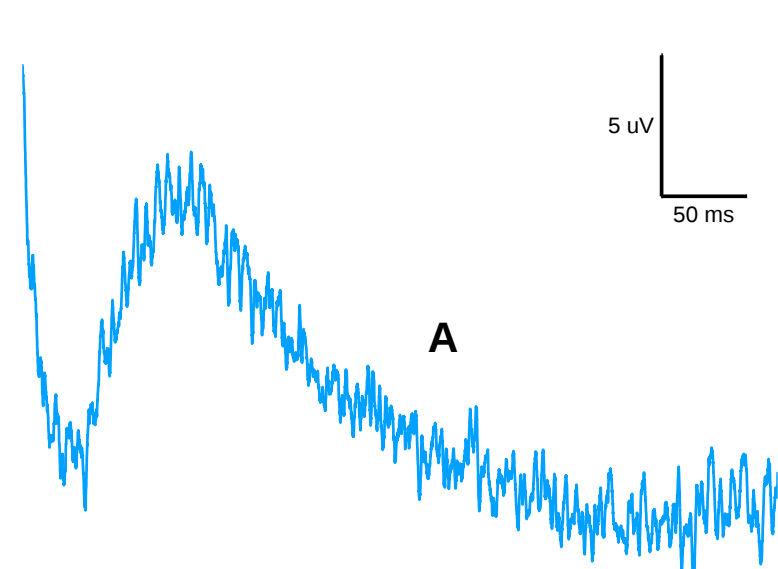
A - Krebs

Main CST



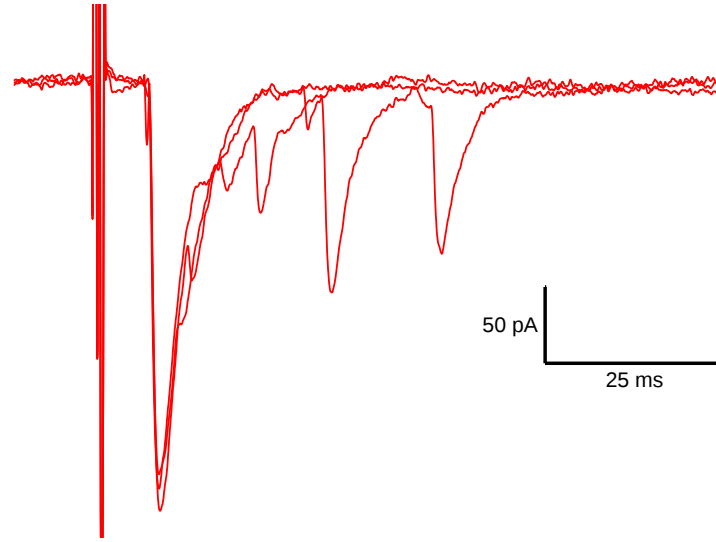
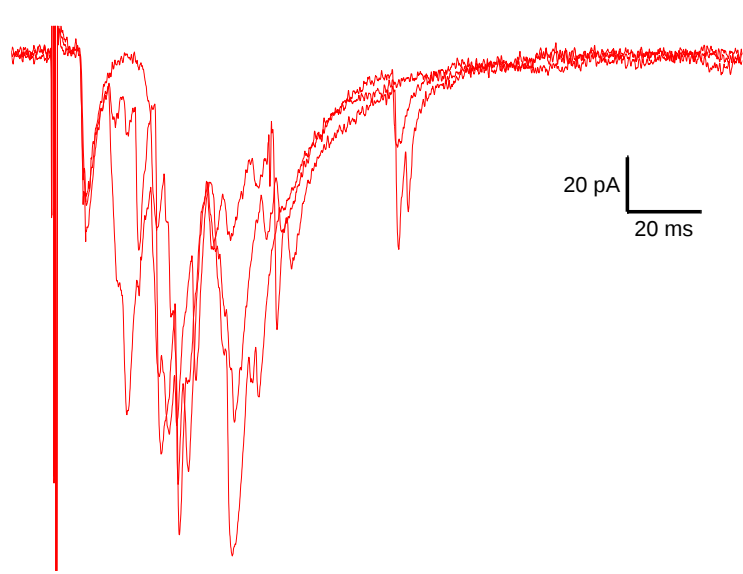
B – Krebs+CNQX+AP5

AF

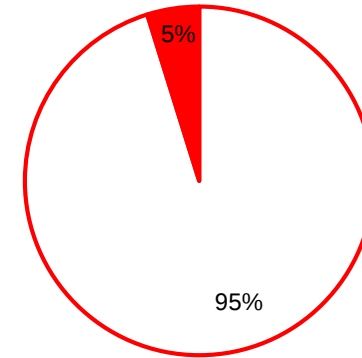


C – Krebs+CNQX+AP5+bicuculline

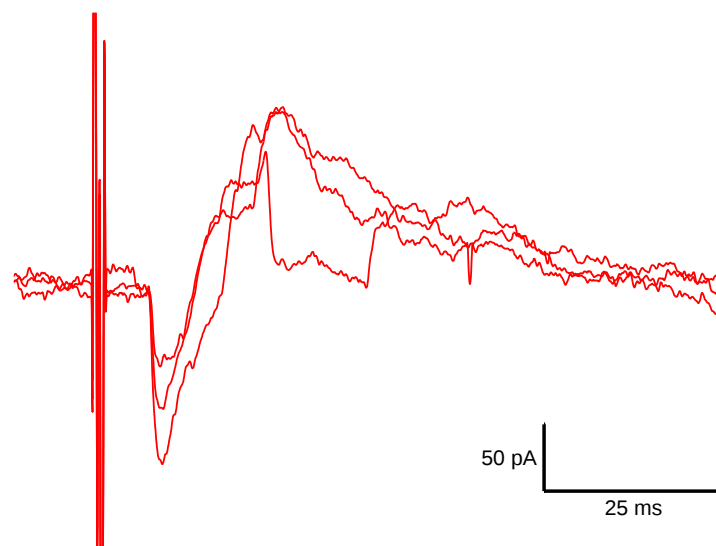
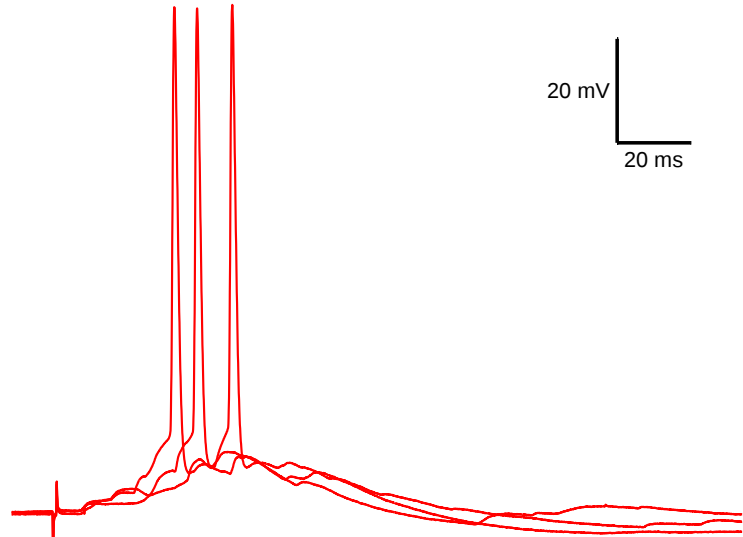
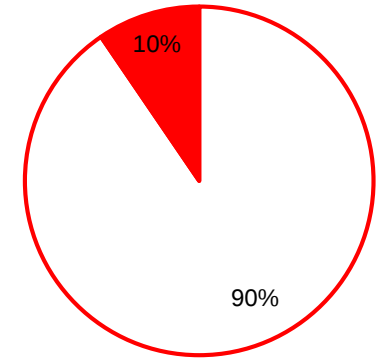
Responses of lamina X neurons to DLF stimulation



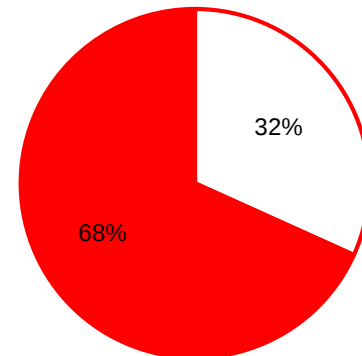
APs



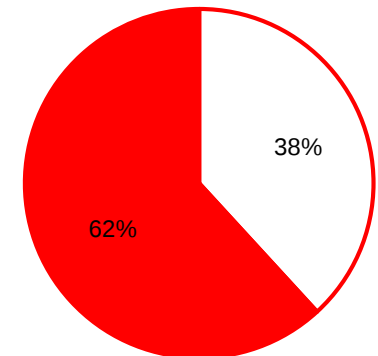
Monosynaptic EPSCs



Polysynaptic EPSCs

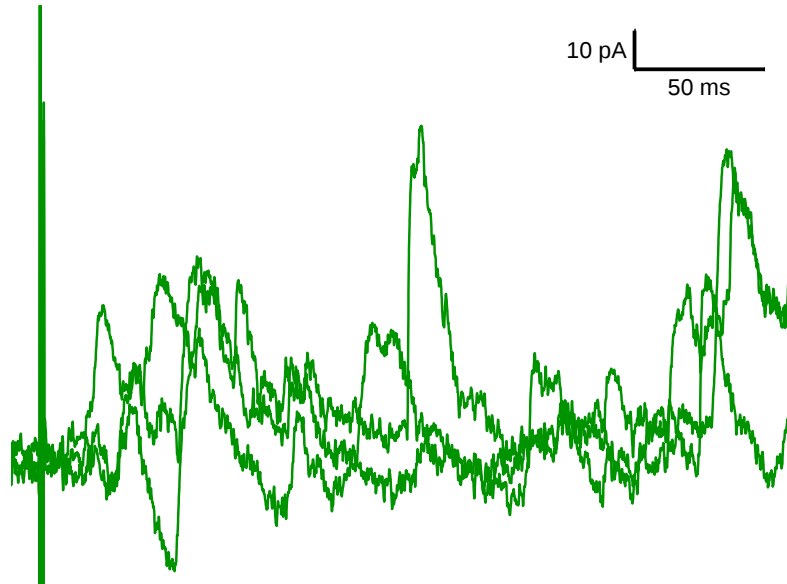
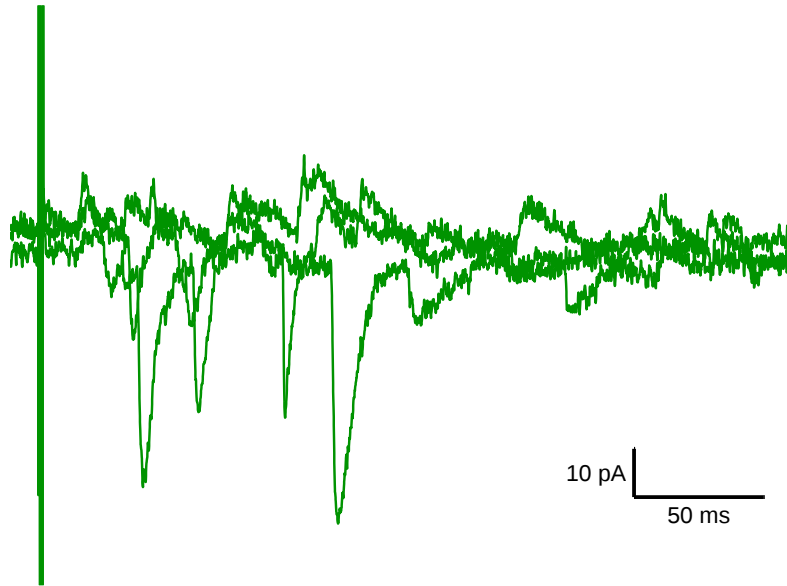


Polysynaptic IPSCs



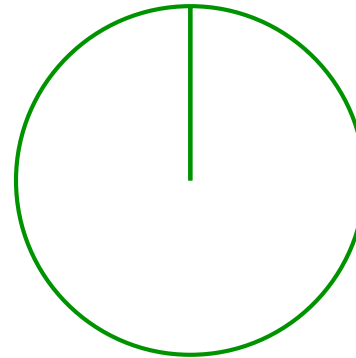
Stimulating pulse: 1ms / 200 uA / 0.1 Hz

Responses of lamina X neurons to main CST stimulation

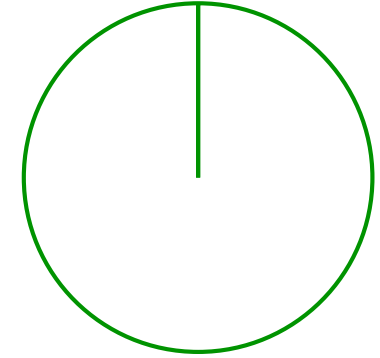


Stimulating pulse: 1ms / 200 uA / 0.1 Hz

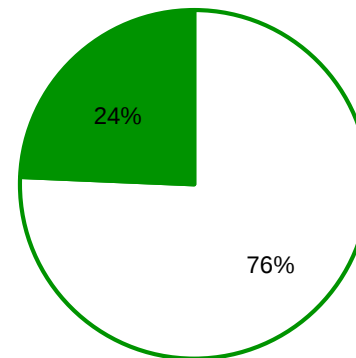
APs



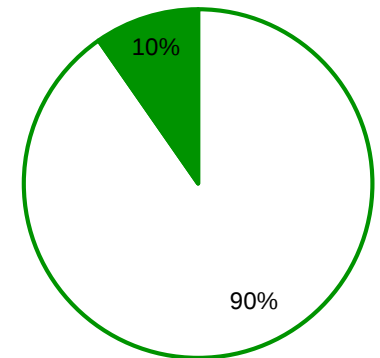
Monosynaptic EPSCs



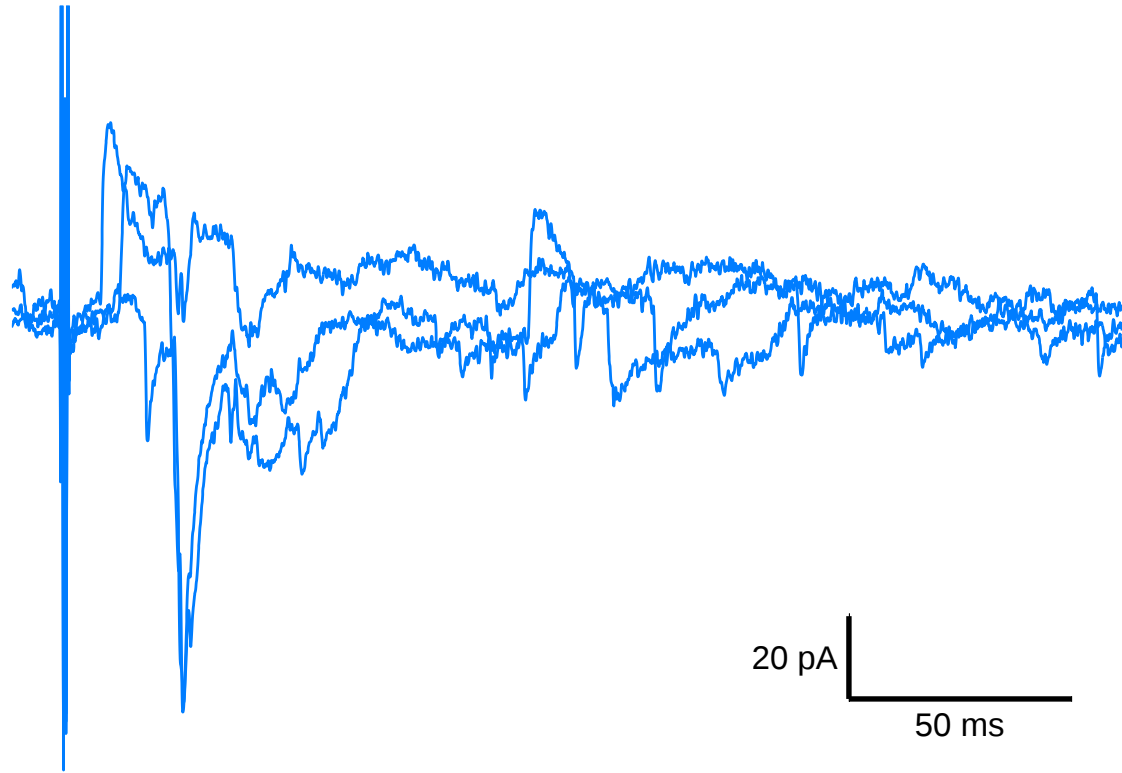
Polysynaptic EPSCs



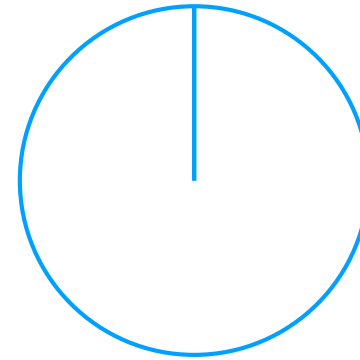
Polysynaptic IPSCs



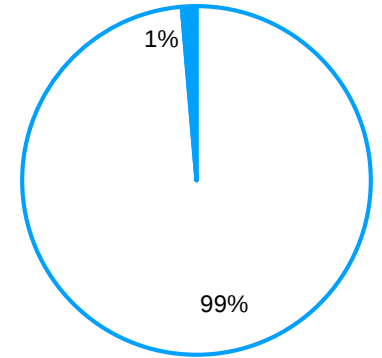
Responses of lamina X neurons to AF stimulation



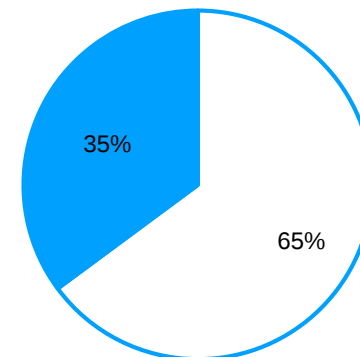
APs



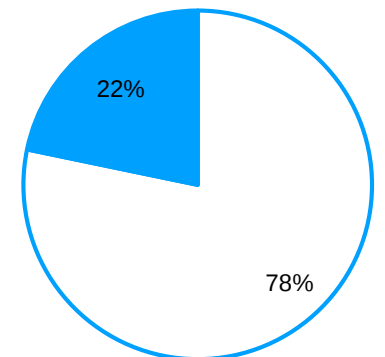
Monosynaptic EPSCs



Polysynaptic EPSCs

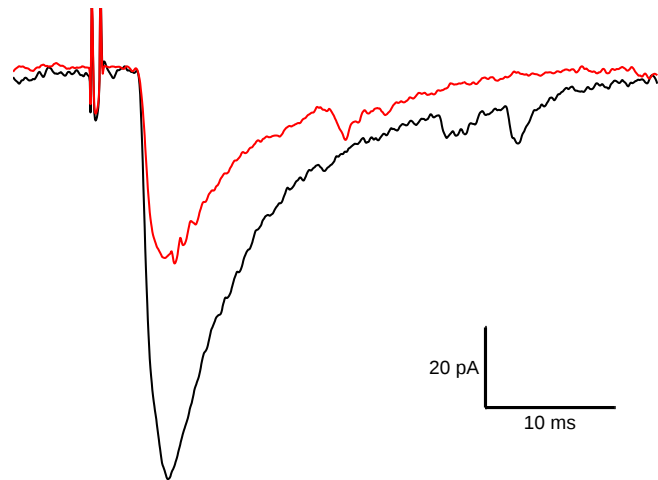
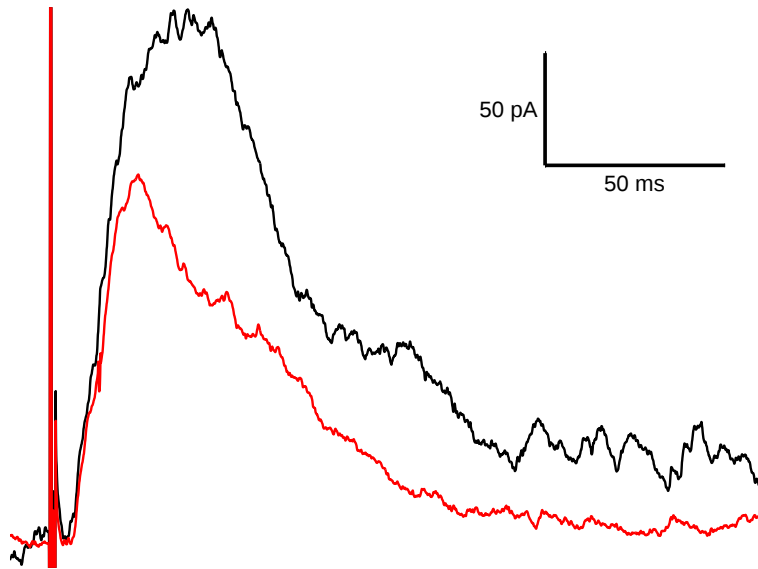
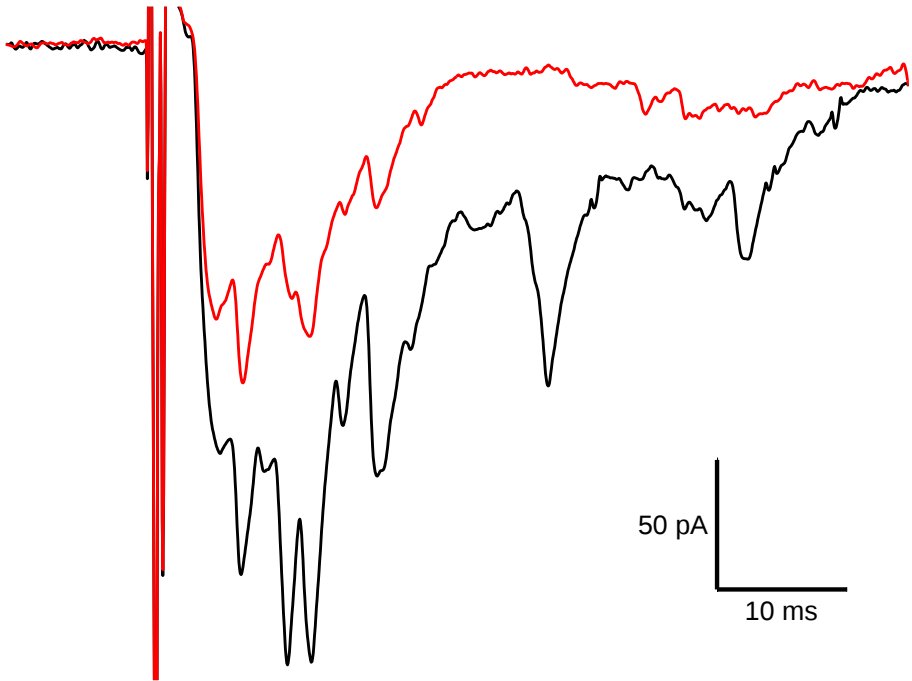


Polysynaptic IPSCs

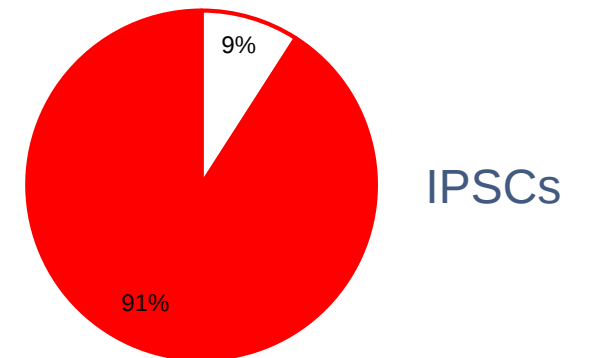
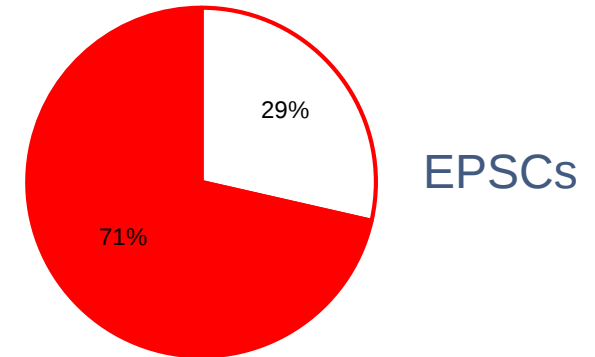
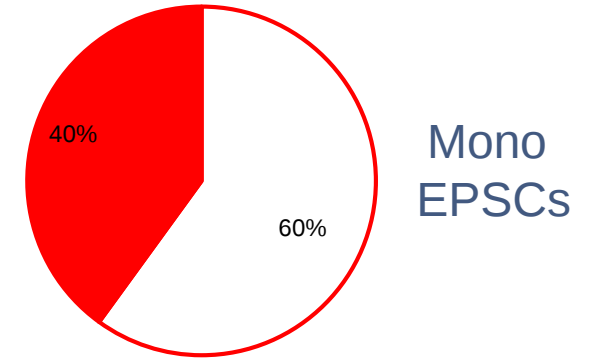
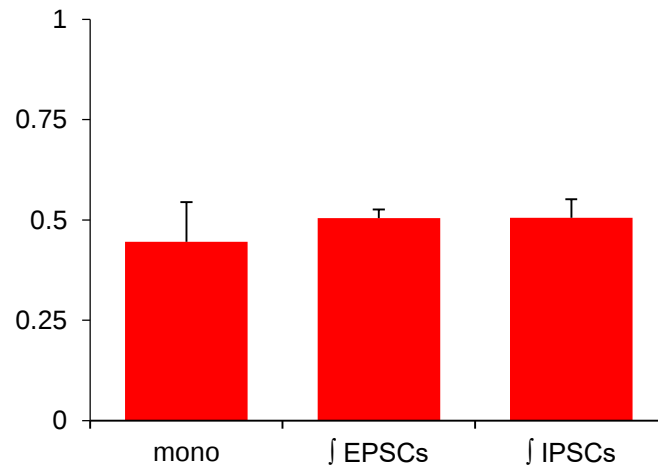


Stimulating pulse: 1ms / 200 uA / 0.1 Hz

Effects of DLF stimulation on PA-induced currents

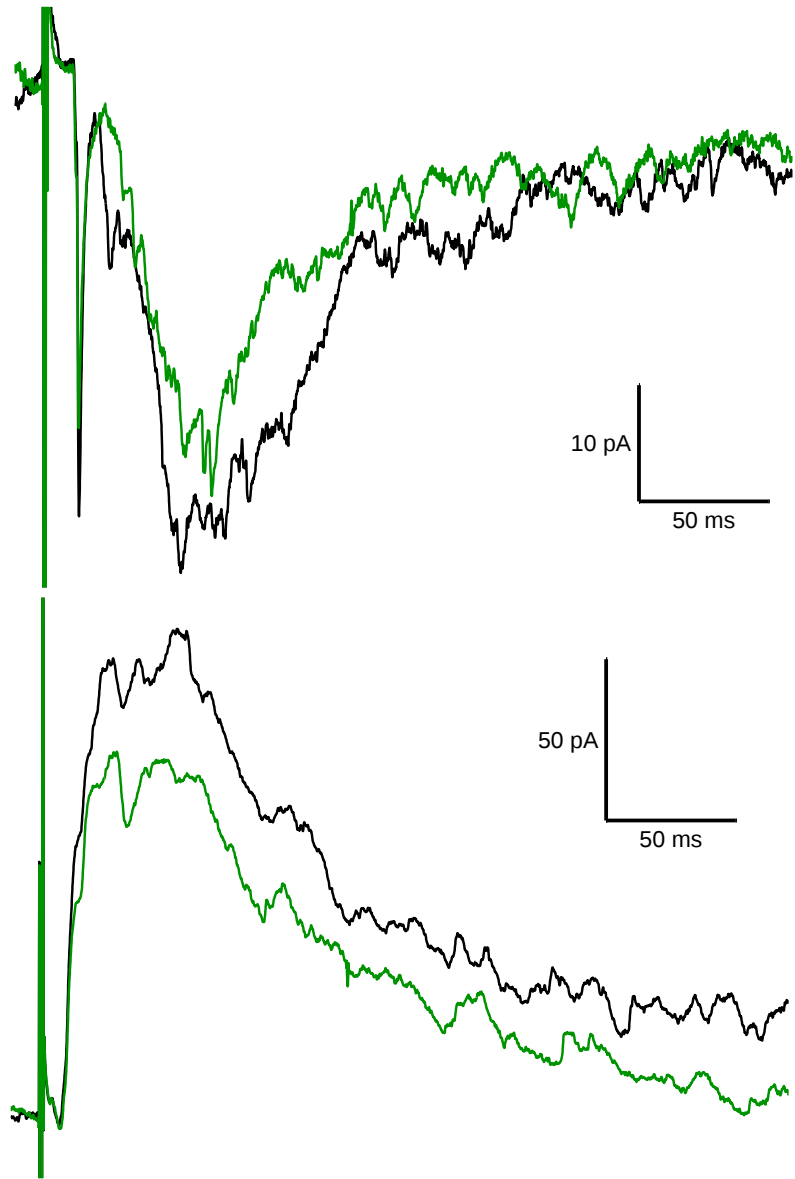


DLF-induced decrease of PA-EPSCs/IPSCs

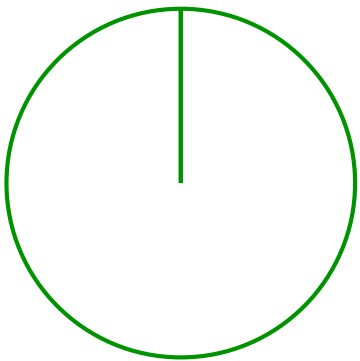
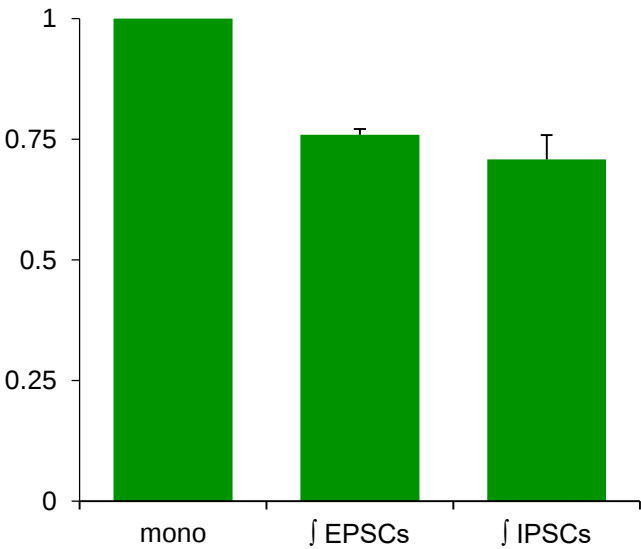


Stimulating pulses: PA -1ms / 150 uA / 0.1 Hz; DLF - 1ms / 200 uA / 5 Hz

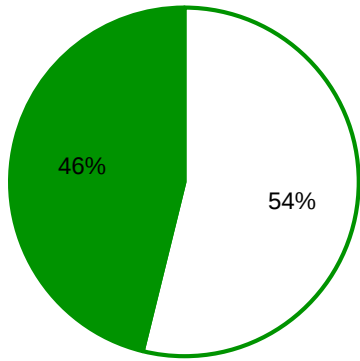
Effects of main CST stimulation on PA-induced currents



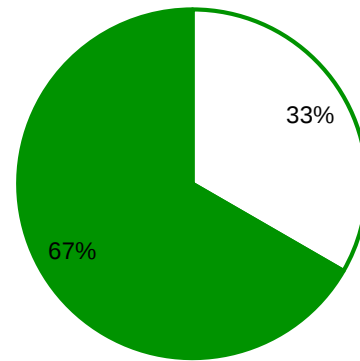
Main CST-induced decrease of PA-EPSCs/IPSCs



Mono EPSCs



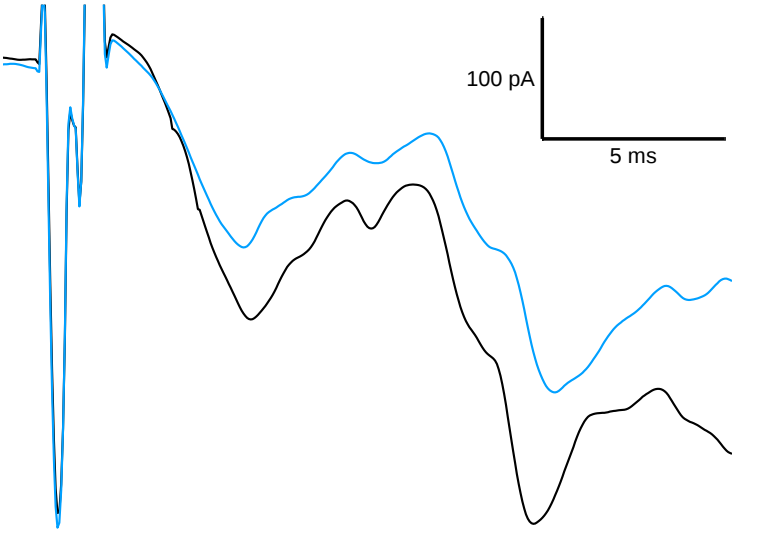
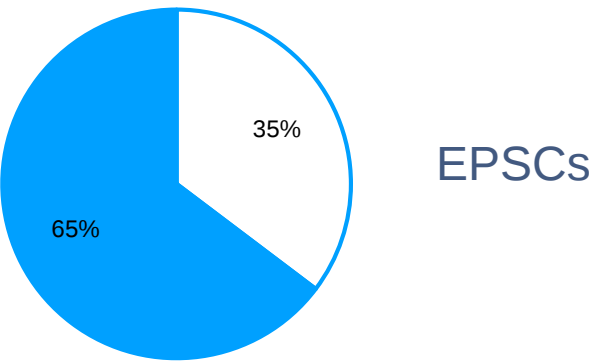
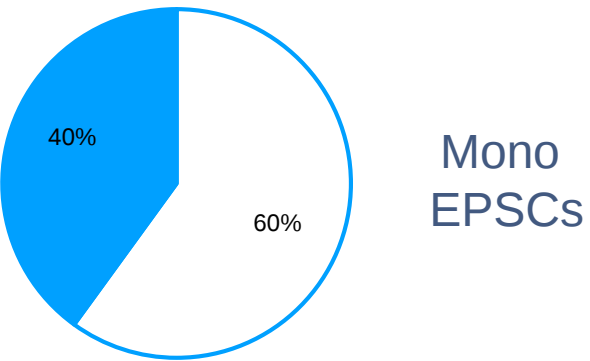
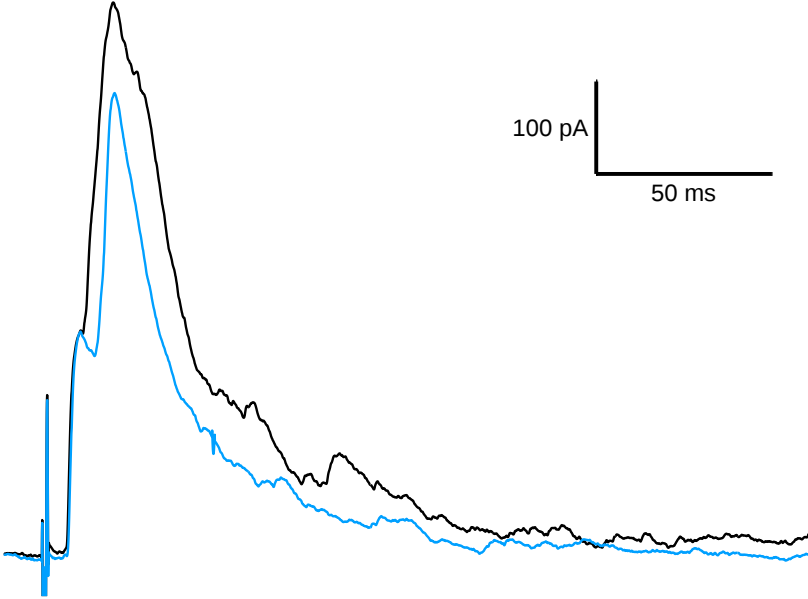
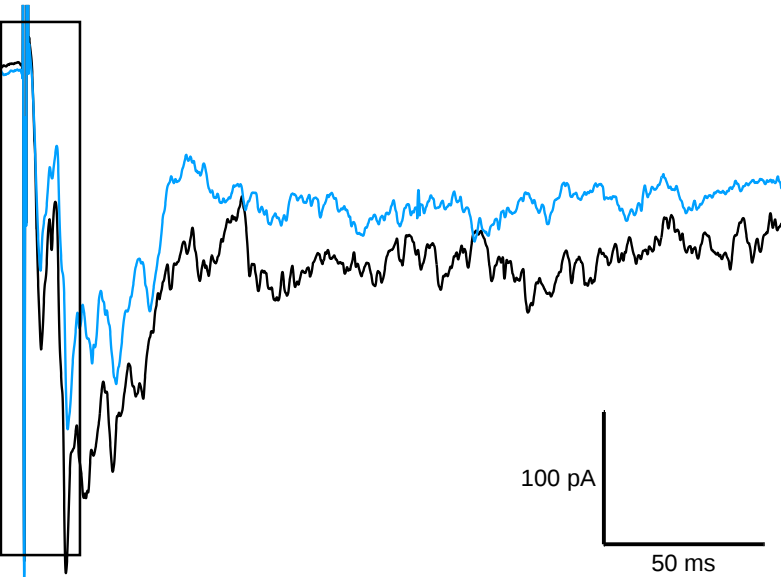
EPSCs



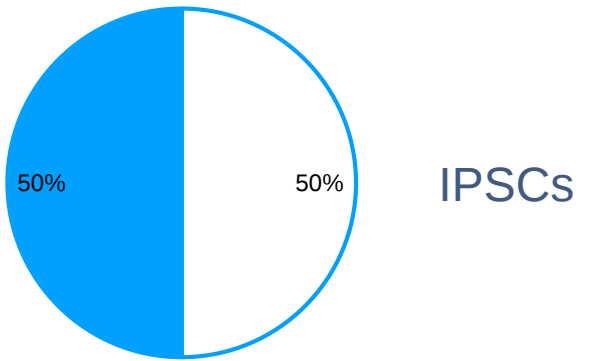
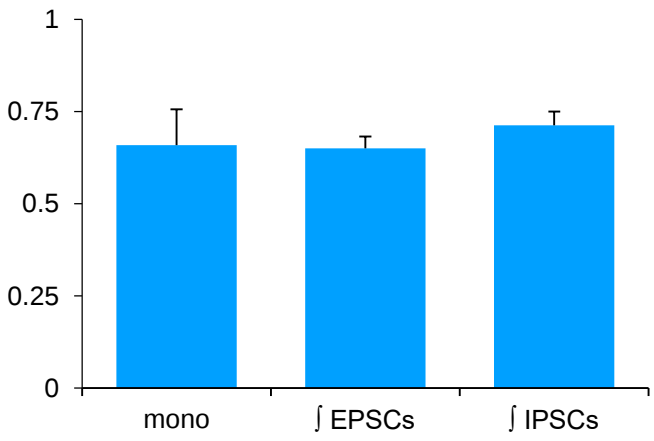
IPSCs

Stimulating pulses: PA - 1ms / 150 uA / 0.1 Hz; main CST - 1ms / 500 uA / 5 Hz

Effects of AF stimulation on PA-induced currents



AF-induced decrease of PA-EPSCs/IPSCs



Stimulating pulses: PA -1ms / 150 uA / 0.1 Hz; AF - 1ms / 200 uA / 5 Hz

Contributors



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Olga Kopach, PhD



Andrew Dromaretsky



Anastasia Tokhtamysh

**Thank you
for your attention**