Aim: The goal of this article is to investigate whether pretreatment with hydroalcoholic extract of Dorema aucheri leaves potentiates the brain antioxidant defense system and inhibits oxidative damage during cerebral ischemia–reperfusion.

Methods: Three groups of rats were randomly selected (each group; n=12): sham, control ischemic and ischemic pretreatment groups. Treated rats received freshly hydroalcoholic extract of D. aucheri (200 mg/kg/day) for 14 days. Then, cerebral ischemia–reperfusion was achieved by 90 minutes middle cerebral artery (MCA) occlusion followed by 24 h reperfusion. Infarct volume and contents of malondialdehyde (MDA), glutathione and nitrate (NOx) as well as superoxide dismutase (SOD) and catalase activities were assessed after 24 h reperfusion.

Results: The contents of MDA and nitrate significantly increased in the ischemic hemispheres by 34% and 14%, respectively. Brain ischemia decreased the glutathione content (20%) and activities of catalase (38%) and SOD (14%) in ischemic hemispheres compared to sham rats. Treatment with DA before MCA occlusion significantly decreased the infarction in cortex and striatum by 63% and 75%, respectively, compared to control. DA considerably reduced the contents of MDA and nitrate in ischemic hemispheres by 28% and 11%, respectively, compared to control rats. Treatment with DA also increased the glutathione content (7%) and activities of catalase (46%) and SOD (16%) of ischemic hemispheres.

Conclusion: The present study revealed that pretreatment with hydroalcoholic extract of DA leaves prevents weakening of the brain antioxidant defense system and decreases the brain damage during cerebral ischemia–reperfusion.

http://dx.doi.org/10.1016/j.pbj.2017.07.073

PS150

The Levator Auris Longus (LAL) muscle as an accessible system to study the effects of Botulinum Toxins in vivo

Torrado Marília 1,2,*, Cruz Célia Duarte 1,2,3, Avelino António 1,2,3

1 Departamento de Biomedicina – Unidade de Biologia Experimental, Faculdade de Medicina da Universidade do Porto, Portugal
2 Translational NeuroUrology, IBMC – Instituto de Biologia Molecular e Celular, Universidade do Porto, Portugal
3 Instituto de Investigação e Inovação em Saúde - i3S, Universidade do Porto, Portugal
E-mail address: marilia.torrado@gmail.com
(T. Marília).

Aim: In the present work, we aimed to find a reproducible model to study the effects of Botulinum neurotoxins (BoNTs), that allowed a widespread visualization of the intoxicated nerve terminals.

Introduction: Despite the well-established successful use of BoNTs to treat a variety of human conditions, their mechanism of action is still not fully understood. Thus, there is an emergent need of new and accurate models to study the effects of BoNTs. However, considering their potential lethality, it is challenging to find reproducible models to study the local application of BoNTs in living animals that allow a widespread visualization of the intoxicated nerve terminals.

In these work, we studied the innervation pattern and the effect of BoNTs in a group of small subcutaneous cranial muscles that are responsible for moving the pinna in rodents. Although all are easily accessible and manipulated, we focused on the levator auris longus (LAL).

Methods: Animals were injected subcutaneously with the indicated doses of BoNT/A, in the cranial muscles area. Muscles were then dissected and prepared for wholemount staining for Synapsin-I, cleaved SNAP-25 (synaptosome-associated protein of 25 kDa) and β3-tubulin.

Results: Detection of cleaved SNAP-25, the end-product of the catalytic action of BoNT/A, was possible even with injections as low as 0.1 ng. Mapping of the injected muscle showed the effect of BoNT/A in the majority of the endplate population. Also, seven days after BoNT/A injection, a sprouting process was evident, a landmark of regeneration.

Conclusion: BoNTs delivery to the LAL is a sensitive, simple and reproducible model to study the mechanisms of action of these toxins as it allowed the evaluation of BoNT/A effects throughout the entire muscle, without sampling bias. Thus, we forward that the LAL manipulation may constitute an excellent model to clarify the mechanisms of action of BoNTs in the neuromuscular system.

Acknowledgements: This study has been funded by FEDER - Fundo Europeu de Desenvolvimento Regional funds through the COMPETE 2020 - Operational Program for Competitiveness and Internationalization (POCI), Portugal 2020, and by Portuguese funds through FCT – Fundação para a Ciência e a Tecnologia/Ministério da Ciência, Tecnologia e Ensino Superior in the framework of the project “Institute for Research and Innovation in Health Sciences” (POCI-01-0145-FEDER-007274).

http://dx.doi.org/10.1016/j.pbj.2017.07.074

PS188

NLRP3 inflammasome as a potential target to reduce epileptic-like activity


Faculdade de Medicina, Universidade de Lisboa e Instituto de Medicina Molecular, Faculdade de Medicina, Universidade de Lisboa, Portugal
E-mail address: leonor_rr@hotmail.com
(L. Ribeiro-Rodrigues).

Aim: Decipher how inflammation drives epilepsy and how NLRP3 targeting impacts epileptic-like activity.

Introduction: Epilepsy is one of the most common neurological diseases in worldwide. Inflammation was linked to the presence of inflammasomes, cytosolic multiprotein complexes, which promote the release of proinflammatory cytokines, namely IL-1ß. Although a feedback loop has been described between inflammation and epilepsy, the role of inflammasomes in epilepsy is still unknown. NLRP3 is the most studied inflammasome, activated by a two-signal process: 1) a priming signal (as lipopolysaccharides – LPS), which enhances the expression of NLRP3 and pro-IL-1ß; and 2)