P0082
Pain and bladder dysfunction in an animal model of multiple sclerosis
H. Cavaleiro 1,*, R. Silva 1, R. Oliveira 1,2,3, A. Coelho 1,2,3, F. Cruz 2,3,4, C.D. Cruz 1,2,3
1 Department of Biomedicine - Experimental Biology Unit, Faculty of Medicine, Porto, Portugal
2 Translational NeuroUrology Group, Instituto de Investigação e Inovação em Saúde, Porto, Portugal
3 Instituto de Biologia Molecular e Celular, Porto, Portugal
4 Dept. of Urology Hospital S. João, Porto, Portugal
E-mail address: helenacavaleiro@hotmail.com (H. Cavaleiro).

Aim: Here, we investigated if MS-induced pain and bladder dysfunction can be attenuated by TRPV1 desensitization with RTX.

Introduction: Multiple sclerosis (MS) is the most prevalent neurological disorder in young people, causing irreversible disability and producing substantial economic and social impact. Among the most incapacitating symptoms, neuropathic pain and bladder dysfunction are reported by the majority of patients. The transient receptor potential vanilloid 1 (TRPV1) is a receptor described to have an important role in neuropathic pain, bladder dysfunction and inflammation. TRPV1 desensitization with agonists, such as resiniferatoxin (RTX), has been shown to improve bladder function and reduce behavioural signs of pain in various animal models of disease. In the context of MS, a recent study showed that TRPV1 knockout mice were protected from disease progressions, presenting delayed disease onset, myelin preservation and reduced clinical scores.

Methods: Experimental Auto-immune Encephalitis (EAE) was induced by a single injection in the flank of a solution of myelin basic protein (MBP) in Complete Freund’s adjuvant (CFA). Behavioural tests were performed to evaluate symptoms. One month after MS-induction, animals were anesthetized and cystometries performed. Two other groups of MS animals received intrathecal RTX or vehicle and also submitted to behavioural tests and cystometries. At the end of experiments, tissue was collected and processed.

Results: EAE rats developed neuropathic pain, as shown by the presence of mechanical allodynia and hypersensitivity to thermal stimuli. Cystometries performed at this time point showed signs of neurogenic detrusor overactivity. These clinical signs were accompanied by decreased spinal expression of MBP and increased activity of astrocytes and microglia. Preliminary observations suggest that intrathecal RTX improved cutaneous hypersensitivity and bladder function. These results suggest that TRPV1 might be involved in pain bladder dysfunction accompanying MS and that its modulation could have therapeutic relevance.

Conclusion: These results suggest that TRPV1 might be involved in pain bladder dysfunction accompanying MS and that its modulation could have therapeutic relevance.

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PS227
Neurogenesis in a rat model of sporadic Alzheimer’s disease
Sara L. Paulo 1,2,*, Rui S. Rodrigues 1,2, Liana Shvachiy 3, Filipa F. Ribeiro 1,2, Susana Solá 4, Ana M. Sebastião 1,2, Sara Xapelli 1,2
1 Instituto de Farmacologia e Neurociências, Faculdade de Medicina, Universidade de Lisboa, Lisboa, Portugal
2 Instituto de Medicina Molecular (iMM), Faculdade de Medicina, Universidade de Lisboa, Lisboa, Portugal
3 Cardiovascular Autonomic Function (CAF) lab, Cardiovascular Center of the University of Lisbon (CCUL), Faculdade de Medicina, Universidade de Lisboa, Lisboa, Portugal
4 Research Institute for Medicines (Med.ULisboa), Faculdade de Farmácia, Universidade de Lisboa, Lisboa, Portugal
E-mail address: sara.lnp@gmail.com (S.L. Paulo).

Aim: Characterize adult hippocampal neurogenesis in a rat model of the initial stages of sporadic Alzheimer’s disease (AD).

Introduction: Sporadic late-onset AD is the most common cause of dementia, that can be characterized by a progressive cognitive decline, with a noteworthy episodic long-term memory impairment at early stages, accompanied by an excess accumulation of amyloid beta (Aβ) peptide in the brain. Present treatment options are very limited, so understanding AD pathophysiology is essential for exploring efficient therapies. Adult hippocampal neurogenesis is thought to play a crucial role in hippocampus-dependent cognitive abilities, namely learning/memory, although how this process is modulated in AD remains unclear.

Methods: An Aβ1–42 peptide solution was intracerebroventricularly injected into the rats’ lateral ventricle (the same volume of vehicle was injected to controls). Moreover, rats were injected with 5-bromo-2′-deoxyuridine (BrdU) intraperitoneally to study cell proliferation and differentiation. Two weeks after Aβ1–42 injection, the open field (OF) test and the novel object recognition (NOR) test were performed. Further behaviour tests are currently being performed, including the elevated plus maze (EPM), the Y-maze forced alternation test, and the Morris water maze (MWM) test. Focusing on the dentate gyrus, immunohistochemical analysis is presently being performed to investigate cell proliferation, neuronal differentiation and neuroblast/neuron morphology. Additionally, the presence of Aβ1–42 monomers and oligomers will be assessed by western-blot and the eventual occurrence of Aβ1–42 aggregates by histology.

Results: Our results show that the Aβ1–42 injection did not affect locomotor activity, as assessed by the OF test. Furthermore, this injection did not affect exploratory drive or episodic long-term memory performance, as indicated by the NOR test.

Conclusion: Since the NOR test is dependent from several brain regions besides the hippocampus that might not be affected in our model, additional behaviour tests as well as cellular and molecular analysis are needed to further characterize this model.

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PS086
Hydroalcoholic extract of Dorema aucheri leaves prevents weakening of the brain antioxidant defense system and inhibits oxidative damage in rat model of ischemic stroke
Mohammad Ehsan Bayatpoor 1,*, Javad Rasouli Vani 1, Mohammad Taghi Mohammadi 3
1 Baqiyatallah Research Center for Gastroenterology and Liver Diseases (BRCGL), Baqiyatallah University of Medical Sciences, Tehran, IR Iran

Abstract: To investigate the effect of Dorema aucheri leaves extracts on brain antioxidant defense system and oxidative damage in a rat model of ischemic stroke.

Introduction: Dorema aucheri (L.) F. Schmidt, commonly known as the yellow Hospitality tree, belongs to the family Anacardiaceae. Previous studies have shown anti-inflammatory, anti-oxidant and anti-microbial effects of this plant. However, no reports have addressed the effect of this plant on cognitive function and ischemic stroke.

Methods: Thirty-two adult male Sprague-Dawley rats (250 ± 20 g) were randomly divided into four groups as following: sham, positive, and two experimental groups. Ischemic stroke was induced by occluding the left middle cerebral artery; the sham group received only an incision. Three weeks after treatment, passive avoidance test (PAT) and Morris water maze (MWM) were used to evaluate cognitive function. For oxidative damage markers, the levels of superoxide dismutase (SOD), glutathione (GSH), total antioxidant capacity (TAC) and malondialdehyde (MDA) were measured in the cortex, hippocampus, and striatum of the brain tissue. Moreover, immunohistochemistry was performed for the neuronal marker NeuN and the microglia marker Iba-1.

Results: Administration of the hydroalcoholic extract of Dorema aucheri leaves prevented weakening of the brain antioxidant defense system and inhibited oxidative damage. Additionally, it improved cognitive function as measured by the PAT and MWM tests.

Conclusion: The findings of the present study provide evidence for the potential of Dorema aucheri leaves to protect against oxidative damage and improve cognitive function in a rat model of ischemic stroke.

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Aim: The aim of this article is Hydroalcoholic extract of Dorema aucheri leaves prevents weakening of the brain antioxidant defense system and inhibits oxidative damage in rat model of ischemic stroke

Introduction: The production of free radicals is the principal mechanism of brain injury in ischemic stroke. The present study tried to identify whether pretreatment with hydroalcoholic extract of Dorema aucheri (DA) leaves potentiates the brain antioxidant system and decreases brain infarction and 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 DA (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 rats. Treatment with DA also decreased 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.

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PS150

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

Torrado Marília 1,∗, 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 whole mount staining for Synapsin-L, 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.

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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-1β. 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,1 activated by a two-signal process: 1) a priming signal (as lipopolysaccharides – LPS), which enhances the expression of NLRP3 and pro-IL-1β; and 2)