Aim: To characterize behavioral and cardiorespiratory changes in a new, intermittent low-level lead exposure animal model.

Introduction: Lead (Pb) is a cumulative toxic metal affecting all body systems that are particularly vulnerable during developmental phase. Permanent lead exposure has been defined as a cause of behavioral changes, cognitive impairment, sympathoexcitation, tachycardia, hypertension and autonomic dysfunction. However, no studies have been performed to describe a new, intermittent low-level lead exposure profile, that has been increased in the past years.

Methods: Foetuses were intermittently (PbI) exposed to water containing lead acetate (0.2%, w/v) throughout life until adulthood (28 weeks of age). A control group (without exposure, CTL) matching in age and sex was used. At 26 weeks, behavioural tests were performed for anxiety (Elevated Plus Maze Test) and locomotor activity (Open Field Test) assessment. Blood pressure (BP), electrocardiogram (ECG), heart rate (HR) and respiratory frequency (RF) were recorded at 28 weeks of age. Baroreflex gain (BRG) and chemoreflex sensitivity (ChS) were calculated. Student’s T-test was used (significance p < 0.05) for statistical analysis.

Results: An intermittent lead exposure causes hypertension (increased diastolic and mean BP), increased RF, decreased baroreflex function and increased ChS, without significant changes in HR, when compared to CTL group. Regarding behavioral changes, the intermittent lead exposure model showed an anxiety-like behaviour without changes in locomotor activity.

Conclusion: Intermittent low-level lead exposure induces changes on the cardiorespiratory profile characterized by hypertension, carotid chemosensitivity and baroreflex impairment. According to behavioural tests results, this study also shows that the exposure to lead during developmental phases causes anxiety in adult animals without locomotor activity impairment.

In summary, this study brings new insights on the environmental factors that influence nervous and cardiovascular systems during development, which can help creating public policy strategies to prevent and control the adverse effects of Pb toxicity.

http://dx.doi.org/10.1016/j.pbj.2017.07.125

PS120

Antihypertensive effects of two novel dihydropyridine derivatives

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Aim: Treatment of hypertension.

Introduction: Mebudipine and dibudipine are two novel derivations of dihydropyridine (DHP) Ca2+ channel blockers. Previous studies have shown that these two compounds have relaxant effects on vascular smooth muscles. In addition, DPHs are able to reduce contraction force of cardiac muscle in rat. In this study we decided to evaluate the antihypertensive effects of these two novel DHPs in hypertensive rat.

Methods: Male Sprague-Dawley rats were used in the study (8–10 weeks old). The rats were randomly divided to 4 groups of 10 rats (one control and 3 test groups). Blood pressure was measured by Tail cuff method. Left kidneys of the rats were removed by nephrectomy and sodium chloride 1% was added to the drinking water of animals and desoxycorticosterone acetate 20 mg/kg (SC) were injected twice a week. During and after 4 weeks, blood pressure of animals was evaluated to confirm the hypertension. Blood pressure of the animals was measured before i.p. injection of mebudipine and dibudipine (1–8 μmole/kg) and 2 min after the drug administration.

Results: Mebudipine and dibudipine significantly reduced the systolic blood pressure. Mebudipine was more effective than dibudipine and nifedipine in hypertensive animals and has significant results.

Conclusion: Previous studies showed that i.p. injection and oral usage of mebudipine and dibudipine decrease systolic hypertension in normotensive animals, on the other hand vasodilation effects of DHPs have been proved on aorta. Both novel drugs showed significant reduction in systolic blood pressure in hypertensive animals and mebudipine was more potent than dibudipine and nifedipine (as a standard drug uses). It is remarkable that, two new DHPs have similar efficacy and safety profile, but have higher efficacy compared to nifedipine in present study. The brilliant point is that DHPs as calcium channel blockers are more effective in hypertensive animals compared to normotensive animals.

http://dx.doi.org/10.1016/j.pbj.2017.07.126

PS062

Biological processes of polyphenols in the cardiovascular system: A bioinformatics approach

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Aim: In this study, we aimed to evaluate the cardiovascular system-related biological processes (BP) modulated by polyphenols in rodents and humans, and to verify which of them are specie-
specific, in order to understand which outcomes for cardiovascular diseases (CVD) could be translated from animal to human studies.

**Introduction:** CVD stand as a great cause of morbidity-mortality worldwide and polyphenol-rich diets have been associated with improved cardiovascular risk profiles. Although rodent models have been a resourceful means of understanding the CVD mechanisms and possible outcomes of the use of polyphenols in that context, most experimental models do not fully reproduce human CVD.

**Methods:** Database searching was carried out on PubMed and Google Scholar using specific keywords concerning CVD, retrieving close to 300 publications. After excluding irrelevant results, proteome data was organized in Excel® spreadsheets and the Cytoscape platform, ClueGo + CluePedia and Venny 2.1.0 were used to explore the biological processes influenced by flavonoids in the approached CVD.

**Results:** This study was mainly focused in the species Rattus norvegicus and Homo sapiens and in flavonoids, a polyphenol subgroup. Only about 5% of the BP influenced by flavonoids were common to both species and they were mostly related to the main group. Only about 5% of the BP influenced by flavonoids were considered when planning clinical trials.

**Conclusion:** Our research highlights the need for a careful translation of the flavonoids’ effects observed in rat models to clinical trials, since different proteins and subgroups of flavonoids mediated the observed actions. Though this type of studies can provide insights to help choosing the most adequate polyphenols as preventive approaches or therapies for human CVD, further investigation should be performed to clarify the described effects. Besides, pharmacokinetic aspects of the flavonoids’ action should also be considered when planning clinical trials.

**Acknowledgements:** This work was supported by Portuguese Foundation for Science and Technology grants PEst-OE/SAU/UI0038/2014; UID/BIM/04293/2013, UID/IC/00051/2013 (financed by Fundo Europeu do Desenvolvimento Regional through COMPETE 2020 – Programa Operacional Competitividade e Internacionalização) and The European Foundation for Alcohol Research (ERAB) (EA 14 23).

**Aim:** The aim of the study was thus to identify Lm proteins capable to bind cAMP.

**Introduction:** Infectious diseases are still a major cause of death worldwide. To infect a host and survive the environment, bacteria have to sense their surrounding and adjust their behaviour. In this adaptation process, cAMP (cyclic adenosine monophosphate) is known to be an important player in pathogens such as *Pseudomonas* spp., *Vibrio* spp. or *Mycobacterium* spp. The small molecule cAMP is a cyclic nucleotide that relays information from receptors to one or more effector proteins within a bacterial cell, functioning as a second messenger. To mediate a response, cAMP allosterically interacts with cAMP-binding proteins. Understanding how this happens is fundamental to predict how bacteria will adapt/act in a given context.

**Methods:** We recently showed that the human foodborne pathogen *Listeria* monocytogenes (Lm) produces cAMP. The aim of this study was thus to identify Lm proteins capable to bind cAMP. To do this, four candidate proteins selected by bioinformatics analyses were expressed, purified and studied biochemically. Three approaches were used: cAMP affinity chromatography; competitive cAMP affinity chromatography; and isothermal titration calometry (ITC).

**Results:** Among the four tested proteins, CbpA displayed cAMP-binding ability on the three approaches used.

**Conclusion:** Hence, our preliminary results showed that CbpA binds to cAMP. It is now mandatory to understand the relation between cAMP and Cbpa, to determine the function of the protein itself and in complex with cAMP, and to understand the importance of this signalling system for virulence.

http://dx.doi.org/10.1016/j.pbj.2017.07.128