

## **Bibliografie**

1. DUTTON R. Current concepts of hemorrhagic shock. *Anesthesiology Clinic*, 2007, 25: 23-34.
2. CHAPMAN M. Hemorrhagic shock. *Critical Care*, 2004, 8: 373-84.
3. KLABUNDE RE. The pathophysiology of hemorrhagic shock. PPT, 2004.
4. LIU LL, Dubick MA. Hemorrhagic shock-induced vascular hyporeactivity in the rat: Relationship to gene expression of nitric oxide synthase, endothelin-1, and select cytokines in corresponding organs. *The Journal of surgical research*. 2005, 125(2): 128-136.
5. MAURITZ JL, Renedo J, Barrion JP et al. Experimental models of hemorrhagic shock. *Nutr Hosp*, 2007, 22: 190-8.
6. HIROAKI S, Kentaro K, Toshiko T et al. Role of tumor necrosis factor-alpha and interleukin-1beta on lung dysfunction following hemorrhagic shock in rats. *Inter Med J Exp Clin Res*, 2008, 14(5): 63-69.

## **REFLECTIONS ABOUT ESTIMATIVE METHOD OF THE ISOLATE PORTAL VEIN CONTRACTILITY**

**Aurel Saulea, Victor Rotaru, Victor Ojog**

Department of Physiology and Biophysics, State University of Medicine and Pharmacy,  
Chisinau, R. Moldova

### **Summary**

The current investigation intended to determine the conditions of portal vein contractions in rats in isometric regime. The proceeding of object preparation as well as experimental phases was described. Main parameters of contractility were highlighted, also the importance of portal vein muscle reactivity at different concentrations of Ca ions. Hypotheses about the origin and mechanism of portal vein muscle contraction were enounced.

### **Rezumat**

Prezenta investigație determină condițiile contracțiilor venei porte la șobolan în regim izometric. A fost descrisă desfășurat manopera de preparare de obiectului de studiu precum și etapele experimentale. S-au evidențiat indicii principali ai contractilității, cât și importanța reactivității mușchiului venos portal la diferite concentrații a ionilor de Ca<sup>2+</sup>. Au fost expuse ipoteze privind originea și mecanismul contracției venei porte izolate.

Keywords - Cajal cells, portal vein, smooth muscle

The present study was done on a group of 200 white male rats, Wistar line with about 180-200 g weight. All the painful manipulations were done under anesthesia, by using Nembutal in dose of 50 mg/kg of the animal's weight or thiopental of Na (0,1g/kg).

In fixed terms and after finishing the experiments the animals were slaughtered through beheading. In anaesthetized rat the thoracic cavity has been opened, was found the vena portae, after which nearby of the liver's entrance was applied the first tweezers which compressed plainly the vena (vein). In this way the vessel is stuffed with blood, becoming larger and comfortable for the manipulations. Then the second tweezers is applied in the lower part of the vessel a little bit higher of confluence of one of its ramification in vena portae. Then with ophthalmic scissors an excision is done below the first tweezers (just near by). The excision continues along the vein's route (longitudinal) making a concrete separation of the conjunctive tissue, without any harm to the vessel wall itself. The vein's excision is done just after tweezers.

All these manipulations had to be done without any additional tension of the vena, in order not to change its contractility, that is to keep the experimental purity.

So, a vena segment of about 4 mm was obtained. Once being extracted from the operating field the given isolated segment together with tweezers was removed into Petri's box which has been filled with Krebs-Henseleit solution, preventively being heated and oxygenated. This proceeding was done as quickly as possible with a maximal exclusion of unfavorable action of the air on the vena. It is necessary to underline this because vena portae in comparison with others represents a vessel which possesses a pace-maker and the harmful influence of the atmospheric air could modify its property. Being put into Petri's box with the given solution the segment of the vena portae was again separated, this time more thoroughly, but keeping its integrity.

After that the separated vena portae has been transported together with Petri's box to the definite station - the organ's bath thermostatically isolated. The whole "pilgrimage" of the rat's vena portae, extraction and separation inclusively, with an intermittent station in Petri's box must last as less as possible, for about 5-6 min. This term had to be kept strictly in each case in order to respect the experimental purity and for a minimum exposure to the atmospheric air.

So, being in the isolated thermostatic organ's bath an end of the vena portae together with the tweezers is fixed to an immobile hook near the bath's bottom. The second end with the respective tweezers is inserted by the mechanotron's arm 6MX1C, which serves as a translator of force. The position of the vena portae was exactly perpendicular to bath's bottom. The thermostatic bath with a volume of 10 ml was perfused with Krebs-Henseleit solution with the following content (mM/L): NaCl - 22,0; KCl - 4,7; NaHCO<sub>3</sub> - 15,5; KH<sub>2</sub>PO<sub>4</sub> - 1,2; CaCl - 2,5; MgCl<sub>2</sub>+6H<sub>2</sub>O - 1,2; glucoza-11,5; 37°C; the aeration being done with the gas composition O<sub>2</sub> - 95%; CO<sub>2</sub> - 5%; pH-7,3 [1].

The perfusion was prepared just before the experiment from a concentrated solution of the above mentioned salts. NaHCO<sub>3</sub> and glucose were added at the end. Until the Ca<sup>2+</sup> was added preventing the sedimentation of the obtained solution CO<sub>2</sub> was added.

The vascular isolated preparation was extended till the optimal power of contraction with 4 mN. The signal of the mechanotron was transmitted to the amplifier UBP2-03. The recording of the contractile function was done with autorecorder K 200 (RDG). The spontaneous contractions of the isolated vena portae were determined in isometric regime [2, 3], the calibration was done preventively. Before starting the experimental actions the vascular preparation was stabilized for about 45 min till the complete adaptation "in vitro" conditions. Subsequently the recording was done during 10 min with the film's speed of 1min/cm, then it was changed to 5sec/cm in order to proceed the respective with a 1 min length.

After the end of the recording during 2 min the solution replacement from the isolated bath organs was done and used another solution; Krebs one, which previously was heated and aerated in the trickling column, the concentration of some ions was modified in dependence and exigency of the studies at every stage of it. In some cases the ingredient or the substance which was studied immediately was introduced in the bath of the isolated organs with Krebs solution, as in the group with trypsin "in vitro". The introduced volume was corrected in such a way that the corresponding dose of the substance to be sufficient and the total volume from the bath isolated organs with the respective osmotic pressure to correspond exactly.

Just immediately after vena portae is put in the bath, the recording starts, the regime and exigencies being the same - all 15 min with the specific features which were described previously.

The total time for an experiment from the beginning till the end of the study was about 120 min; in such a way the possible metabolic tiredness being avoided in the vein's segment. The repeated apparatus calibration was inevitable at the end of all experiments. At last, the segment

of the vena portae was weighed in order that the contractility data to be reported to mg/mg of vein's weight.

The following indexes were studied (tab.1):

- the amplitude contraction (mg/mg) through the contraction force report to the contractile portion mass of the vena portae;
- the contraction's frequency (contraction/min);
- the contraction's surface ( $\text{mm}^2/\text{min}$ );
- the intensity of the structures functioning through the amplitude produce to the contractions frequency (mg/mg/min);
- the total contraction time (sec);
- the contraction phase (sec);
- without relaxations (sec);

Table 1. The contractile index value of the isolated vena portae.

Index	Value
Amplitude, N/mg	35,5±3,28
Frequency, contr/min	6,18±0,4
IFS, Nmg/min	219,4±23,7
Surface, $\text{mm}^2/\text{min}$	919,91±86,6
The total contraction time, sec	5,8±0,75
The contraction phase, sec	2,9±0,37
The relaxation phase, sec	2,9±0,37

### Discussions and conclusions

1. The mechanotron 6 MXIC, used for measuring the small movements, allows to determine the phasic contractions of the vena portae and less the tonic contractions, because the artifacts (errors) would confuse the correct estimation in this regime. Minimalization of the electric artifacts with the help of the electric transformer connected to the network which stops the tension's fluctuation as well as the presence of the isolated organ bath in the screening room can not be absolute in order for keeping the experimental purity in estimating of the tonic estimation.

2. Changing the solutions in the isolated organ bath may reflect initial recording tonic contractions. It is necessary to keep a strictly liquid volume in isolated organ bath.

3. The amplitude of contractions characterizes the force of contraction and is indissoluble bound by the  $\text{Ca}^{2+}$  ions reserve from outside, for the smooth vascular venous muscle in different  $\text{Ca}^{2+}$  ions concentration, is remarkable the change of the venous muscle reactivity of the vena portae in the groups of study in comparison with the control. Modification of the contractions frequency tells us about pacemaker's involvement, the source of the vena portae automatism – interstitial cells of Cajal [4, 5], which have been recently discovered.

4. The intensity of the structures functioning in the studies was changed in some cases on the base of amplitude's transformation in other only on the base of contractions frequency, but there were noticed groups in which the transformation was depended by both index.

5. The other contractile index – the contraction's surface, the total contraction's time, the phase of contraction and of relaxation contribute to supplementation of the contractility characteristic of the smooth vascular muscle of the vena portae.

## References

- [1] R. Macinelli, P. Tonali, R. Romani, A. Tringali, R. Vargiu, G.B. Azzena, *Mechanical properties of smooth muscle portal vein in normal and dystrophin-deficient (mdx) mice*, Experimental Physiology, vol. 84, 1999, pp. 929-940
- [2] Е.Б. Манухина, *Воротная вена и ее сократительная функция в норме и патологии*, Успехи физиологических наук, том 19, № 3, 1988, стр. 45-66
- [3] В.Г. Пинелис, М.О. Лоншамп, *Влияние эндотелина-I на  $Na^+/H^+$  обмен в гладкомышечных клетках сосудов*, Бюллетень экспериментальной биологии и медицины, № 12, 1992, стр. 594-596
- [4] T.B. Bolton, *Calcium events in smooth muscles and their interstitial cells; physiological roles of sparks*, Journal of Physiology, vol. 570, no. 1, 2006, pp. 5–11.
- [5] O.V. Povstyan, D.V. Gordienko, M.I. Harhun & T.B. Bolton, *Identification of interstitial cells of Cajal in the rabbit portal vein*, Cell Calcium, vol. 33, 2003, pp. 223–239.

## EFECTELE NESPECIFICE ALE ESTROGENILOR

Tatiana Cucu, Cristina Cucu

(Conducător științific: doctor în medicină, conferențiar universitar, Nicolae Demişcan)  
Catedra Fiziologia Omului și Biofizică, USMF “Nicolae Testemițanu”

### Summary

#### *The non-specific effects of estrogens*

The effects of estrogens upon the cardiovascular system are: prevention of atherosclerosis, improvement of the lipid profile and vasodilation. The effects of estrogens upon the digestive system are: reduction of the intestinal peristalsis and increase of absorption. Estrogens also have neurotrophic and neuroprotective effects, along with the prevention of brain ischemia, they have an antidiabetic action, increase the retention of water and sodium, protect the skeletal system and increase blood coagulation. Estrogens also maintain the normal structure of skin in women by promoting keratinocytes and fibroblasts proliferation, and by stimulating its vascularization.

### Rezumat

La nivelul sistemului cardiovascular, estrogenii au efect antiaterosclerotic, de ameliorare a profilului lipidic și de vasodilatare. La nivel digestiv, estrogenii micșorează peristaltismul intestinal, favorizând absorbția. Estrogenii mai posedă efect neurotrofic, neuroprotector și antiischemic cerebral, au acțiune antidiabetică, favorizează retenția de apă și sodiu, au efect protector asupra țesutului osos, majorează coagulabilitatea sîngelui. La nivelul pielii, estrogenii mențin structura ei normală la femei prin promovarea proliferării keratinocitelor, fibroblastelor, prin stimularea vascularizării pielii.

### Introducere

Estrogenii sunt hormoni sexuali feminini, de origine steroidă, fiind reprezentați de estradiol (cel mai activ), estronă și estriol (mai puțin activi). Prin penetrarea estrogenilor în celulele-țintă (din uter, vagin, glanda mamară, sistemul hipotalamo-hipofizar, oase, ficat etc.) și fiind transportați în nucleu, aceștia se cuplează cu receptorii estrogenici ( $ER\alpha$  și  $ER\beta$ ), interacționînd ulterior cu ADN și cu proteine celulare specifice, realizînd modularea expresiei și transcripției unor gene și inducînd sinteza proteinelor în celule.

Pe lîngă efectele specifice asupra organelor sexuale și asupra dezvoltării sexuale la femei (stimularea proliferării și dezvoltării uterului, vaginului și glandelor mamare, proliferarea