## Effect of H<sub>2</sub>S donor, NaHS on rat blood pressure and possible involvement of calcium RyR2 channel

Zuzana Tomaskova<sup>1</sup>, Frantisek Kristek<sup>2</sup>, Sona Cacanyiova<sup>2</sup>, Karol Ondrias<sup>1</sup>

<sup>1</sup>Institute of Molecular Physiology and Genetics, Slovak Academy of Sciences, Bratislava, <sup>2</sup>Institute of Normal and Pathological Physiology, Centre of Excellence for Cardiovascular Research, Bratislava. E-mail: <u>zuzana.tomaskova@savba.sk</u>

Hydrogen sulfide, H<sub>2</sub>S, is an endogenously formed gaseous transmitter. H<sub>2</sub>S is involved in a regulation of cardiovascular functions. It affects several cardiac ion channels (i.e. L-type  $Ca^{2+}$  channel,  $K_{ATP}$  channel, Cl<sup>-</sup> channel). Protective effects of H<sub>2</sub>S were shown for both ischemia/reperfusion preconditioning and postconditioning. It might be also beneficial in treatment of hypertension. Therefore, we decided to study the effects of H<sub>2</sub>S on blood pressure and heart rate of anaesthetized rats and on the activity of ryanodine receptor Ca<sup>2+</sup> release channels (RyR2) derived from cardiac sarcoplasmic reticulum.

NaHS (up to 64  $\mu$ mol/kg, i.v.), a donor of H<sub>2</sub>S, decreased transiently the blood pressure to (84.5–46.5)%, followed by its increase to (107.8–170.8)%. Single channel properties of RyR2 channels were studied in bilayer lipid membrane. NaHS modulated RyR2 channel activity. It increased 3.6-fold open probability of the channel at 50  $\mu$ mol/l. At higher concentrations (500 – 1000  $\mu$ mol/l), in 6/11 experiments NaHS decreased P-open, while in 5/11 experiments, it did not have any effect.

The results indicate that  $H_2S$  and/or  $HS^-$  effects on blood pressure are concentration dependent (biphasis), and that modulation of RyR2 by  $H_2S$  might be responsible, at least in part, for the increase of blood pressure.