# **Endogenous Quinidine-like Immunoreactivity in the Serum of Rats after Acute Hypoxia**

## M. SCHREIBER, M. HOSTLOVSKÁ<sup>1</sup>, J. NEDVÍDKOVÁ<sup>2</sup>, V. SCHREIBER<sup>3</sup>

Institute of Physiology, First Faculty od Medicine, Charles University, <sup>1</sup>Department of Pharmaceutical Control and Bioanalysis, Postgraduate Medical and Pharmaceutical Institute, <sup>2</sup>Institute of Endocrinology and <sup>3</sup>Laboratory for Endocrinology and Metabolism, 3rd Medical Department, First Faculty of Medicine, Charles University, Prague

Received October 5, 1993 Accepted October 20, 1993

#### Summary

Quinidine-like immunoreactivity (Abbott TDx Quinidine fluorescence polarization immunoassay) was measured in blood serum of control rats and rats exposed to hypoxia in a hypobaric chamber (7 500 m for one hour). The mean "quinidine" levels (mean  $\pm$ S.D.) were 0.159 $\pm$ 0.058 and 0.260 $\pm$ 0.110 (µmol quinidine/l serum), respectively (p<0.01).

#### Key words

Quinidine-like immunoreactivity - Hypoxia - Blood - Rats

Recently Schreiber et al. (1993) observed presence of quinidine-like immunoreactivity in blood of rats made thyrotoxic by thyroxine feeding. This finding may fit to the knowledge of other endogenous ligands for drug receptors, e.g. endorphins for opiate receptors endogenous nitric (Snvder 1984), oxide for nitrosovasodilators (Khana and Furchgott 1992), endogenous digitalis-like factor (Schreiber et al. 1981, Gruber et al. 1979) and others (endogenous endogenous benzodiazepines, calcium channel blockers).

Since the endogenous quinidine-like factor could display antiarrhytmic activity, it is predictable that it could be present in blood of animals with a tendency to arrhytmia, such as thyroxine-induced thyrotoxicosis. Actually, its presence is linked with an increase in beta-adrenoceptor activity induced by thyroxine (Schreiber *et al.* 1993).

Another situation, in which elevated endogenous quinidine-like (antiarrhytmic?) activity could be expected is hypoxia. After several pilot experiments (examining the time interval after hypoxia was studied) we report here the results of an experiment, in which quinidine-like immunoreactivity was measured by the Abbott TDx quinidine fluorescence polarization immunoassay. Ten control and ten hypoxic Wistar male rats were examined. Mean body weight was  $108 \pm 7$  g in controls and  $107 \pm 9$  g in hypoxic rats, fed on a standard laboratory diet. The animals were exposed to high-altitude hypoxia in a hypobaric chamber at 7 500 m for one hour. After an additional hour the rats were killed by decapitation and quinidine-like immunoreactivity was measured in their serum. The animals of the control group were exposed to the same manipulations except of high-altitude hypoxia.

A highly significant 72 % increase of endogenous quinidine-like immunoreactivity was observed in the posthypoxic animals, as compared to controls ( $0.260 \pm 0.110$  vs  $0.159 \pm 0.058 \mu$ mol quinidine/l serum p < 0.01). The activity in sera of control rats was slightly higher than in controls of the previous experiment (Schreiber *et al.* 1993) but is the same as in slightly stressed rats. Thus the manipulation with the animals, especially their placing into the hypobaric chamber (without hypobaria), could play a role.

Thus, one hour after the end of acute hypoxia (7 500 m lasting for one hour) an increased quinidinelike immunoreactivity was observed in rat blood. The source and the nature of endogenous quinidine-like activity are unknown, as well as its possible antiarrhythmogenic action.

### References

- GRUBER K.A., WHITAKER J.M., PLUNKETT W.C., BUCKALEW W.M. Jr.: Detection of an endogenous digoxin-like substance (endoxin) in plasma of dogs, sheep and man. (Abstract) *Kidney Int.* 16: 817, 1979.
- KHAN M.T., FURCHGOTT R.F.: Similarities of behaviour of nitric oxide (NO) and endothelium-derived relaxing factor in a perfusion cascade bioassay system. Fed. Proc. 46: 385, 1992.
- SCHREIBER V., KÖLBEL F., ŠTĚPÁN J., GREGOROVÁ I., PŘIBYL T.: Digoxin-like immunoreactivity in the serum of rats with cardiac overload. J. Mol. Cell. Cardiol 13: 107-110, 1981.
- SCHREIBER V., KÖLBEL F., HOSTLOVSKÁ M., NEDVÍDKOVÁ J.: Endogenous quinidine-like immunoreactivity in the serum of rats with cardiac overload, stress or hyperthyroidism. Cor. Vasa 35: 144-146, 1993

SNYDER S.H.: Drug and neurotransmitter receptors in the brain. Science 224: 22-31, 1984.

#### **Reprint Requests**

Dr. M. Schreiber, Institute of Physiology, First Faculty of Medicine, Charles University, Albertov 5, 128 00 Prague 2, Czech Republic.