Investigation of prolactin-related vasoinhibin in sera from patients with diabetic retinopathy

Jakob Triebel, Dirk Raddatz, Michael Huefner¹ and Giuliano Ramadori European Journal of Endocrinology 161 345–353

Department of Gastroenterology and Endocrinology, Center of Internal Medicine, Georg-August-University, Robert-Koch-Straße 40, 37075 Göttingen, Germany and ¹Endokrinologikum Göttingen, Von-Siebold-Straße 3, Göttingen, Germany (Corresponding Author: J Triebel; Email: jakob.triebel@gmx.de)

Abstract

Objective: In vitro experiments and in vivo studies on rodents demonstrate that Nterminal 14, 15, 16, 17, and 18 kDa fragments (prolactin-related vasoinhibin, PRL-V) of human PRL are natural inhibitors of neovascularization in the retina and elsewhere. These N-terminal PRL fragments belong to a family of peptides named vasoinhibins, which act as endogenous regulators of angiogenesis and vascular function. These observations led to the hypothesis that PRL-V could play a role in the pathophysiology of diabetic retinopathy in humans. The purpose of this study was to investigate whether patients with diabetes mellitus and diabetic retinopathy have aberrant concentrations of PRL-V in the circulating blood.

Research design: We performed a case-control study and developed a new technique to semiquantitatively determine PRL-V in serum samples from 48 male subjects. The case group consisted of 21 patients with diabetes mellitus and proliferative or non-proliferative diabetic retinopathy. The control group consisted of 27 healthy subjects with no history of diabetes mellitus.

Methods: For the detection of PRL-V, we developed a new analytical method, consisting of immunologic and laser-induced fluorescence techniques.

Results: The case group had significantly lower PRL-V serum concentrations than the control group (P=0.041). There was no significant difference between patients with proliferative and those with non-proliferative diabetic retinopathy.

Conclusion: We conclude that given the antiangiogenic and antivasopermeability actions of PRL-V, the decreased serum levels of PRL-V in patients with diabetes mellitus could contribute to the development and progression of diabetic retinopathy.

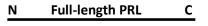
1 Background

Patients with Diabetes Mellitus are at risk for angiogenesis-dependent complications such as proliferative diabetic retinopathy



2 Objective

Detection and semi-quantitative measurement of the systemic levels of angiogenesis-inhibiting prolactin-related vasoinhibin



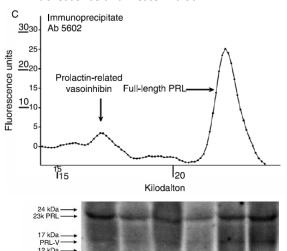
Proteases



N PRL-Vasoinhibin

4 Methods

Immunoprecipitation and immunocomplex constituent analysis with laser induced fluorescence and western blot

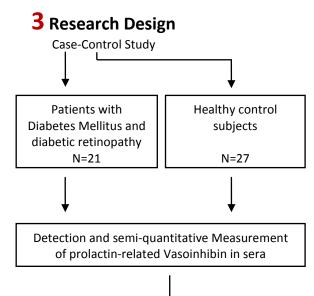


5 Results

The case group had significantly lower PRL-V serum concentrations than the control group (P=0.041). There was no significant difference between patients with proliferative and those with non-proliferative diabetic retinopathy.

6 Conclusion

The decreased levels of prolactinrelated Vasoinhibin in sera from patients with diabetic retinopathy could contribute to the development and progression of diabetic retinopathy.



Case group (n=21)

Subgroup (RDP) (n=14) 2.51 ± 0.85

Serum PRL (mU/l)

2.39±0.59 Subgroup (RDS) (n=7) 2.15±0.56 238.6±19.38 248.9±27.49 3.64+0.51

222.1 + 19.02

