

论著

MK801 对近视视网膜NO-cGMP 信号通路的调控

文丹¹, 刘双珍¹, 毛俊峰¹, 谭星平¹, 夏朝华¹, 尹楚南²

1. 中南大学湘雅医院 眼科,长沙 410008;
2. 中南大学湘雅医院 同位素室,长沙 410008

摘要: 目的: 观察MK801 对豚鼠近视的调节,探讨其在近视发病机制中的作用。方法: 3 周龄三色豚鼠分为6 组: A 组(正常空白对照组)、B 组(右眼遮盖3 周组)、C 组(右眼遮盖3 周+ 玻璃体腔生理盐水注射组)、D 组(右眼遮盖3 周+ 玻璃体腔注射1 ng MK801 组)、E 组(右眼遮盖3 周+ 玻璃体腔注射10 ng MK801 组)、F 组(右眼遮盖3周+ 玻璃体腔注射100 ng MK801 组)。实验前及实验3 周时对各组进行视网膜检影和A 超测眼轴,原位杂交法检测神经细胞性一氧化氮合酶(ncNOS) 的表达,放射免疫法检测cGMP 的含量,将D,E,F 组的屈光度、眼轴、ncNOS 及cGMP 含量与MK801 药物浓度进行直线相关分析。结果: 玻璃体腔药物注射C,D,E,F 组遮盖眼随注射浓度的升高近视屈光度数下降,眼轴延长减慢, ncNOS 及cGMP 含量下调,与MK801 注射浓度行相关分析呈直线相关,屈光度与注射浓度呈正相关($r=0.702, P<0.05$), 眼轴长度、ncNOS 表达、cGMP 表达与其呈负相关($r=-0.736, -0.637, -0.725, P<0.05$)。结论: 近视豚鼠MK801 玻璃体腔注射能通过下调NO-cGMP 表达减缓近视的进展, 呈剂量依赖性。

关键词: 形觉剥夺性近视 视网膜 一氧化氮 环磷酸鸟苷 MK801

MK801 controls form-deprivation myopia by nitric oxide-cyclic GMP signaling pathway in guinea pig

WEN Dan¹, LIU Shuangzhen¹, MAO Junfeng¹, TAN Xingping¹, XIA Chaohua¹, YIN Chunan²

1. Department of Ophthalmology, Central South University, Changsha 410008, China;
2. Department of Nuclide, Xiangya Hospital, Central South University, Changsha 410008, China

Abstract: Objective: To investigate the mechanism of myopia following intravitreal injection of MK801 (dizocipine maleate) intravitreal injected. Methods: Three-week-old guinea pigs were divided into six groups: group A (control), group B (3 weeks form-deprivation in right eye), group C (3 weeks form-deprivation in right eye + saline), group D (3 weeks form-deprivation in right eye + MK801 1ng), group E (3 weeks form-deprivation in right eye + MK801 10 ng), group F (3 weeks form-deprivation in right eye + MK801 100 ng). The refraction and axial length of the eyes were measured. ncNOS was measured by hybridization in situ, and cyclic GMP (cGMP) concentrations by radioimmunochemistry. The correlation between MK801 concentration and diopter degree, axial length of the eyes, and levels of ncNOS or cyclic GMP were analyzed with linear correlation in the groups C-F. Results: Diopter degree was decreased, axial eye length was shortened and levels of ncNOS and c-GMP were decreased in groups C, D, E and F dependent on the concentration of MK801. The diopter degree had positive correlation with MK801 concentration ($r=0.702, P<0.05$), while the axial eye length and the levels of ncNOS and cGMP were negatively correlated ($r=-0.736, -0.637, -0.725, P<0.05$) Conclusion: MK801 injected into the vitreous humor can restrain myopia by down-regulated the expression of the nitric oxide-cyclic GMP signaling pathway. The effect is concentration dependent.

Keywords: form deprivation myopia retina nitric oxide cyclic GMP MK801

收稿日期 2012-02-26 修回日期 网络版发布日期

DOI: 10.3969/j.issn.1672-7347.2012.07.016

基金项目:

国家自然科学基金(81100691); 湖南省自然科学基金(09JJ5033); 湖南省科技厅科技计划项目(2010SK3125, 2009FJ3175)。

通讯作者: 刘双珍,Email: liushzhxy@163.com

作者简介: 文丹,博士,主治医师,主要从事屈光及斜弱视研究。

作者Email: liushzhxy@163.com

参考文献:

1. Vladychenskaya E, Tyulina O, Urano S, et al. Rat lymphocytes express NMDA receptors that take part in regulation of cytokine production[J]. Cell Biochem Funct, 2011, 29(7): 527-533.
2. 文丹,刘双珍,毛俊峰,等.NO-cGMP信号通路对豚鼠形觉剥夺性近视的调控[J].眼科研究,2008,26(9):672-675.WEN Dan, LIU Shuangzhen, MAO Junfeng, et al. Regulation role of Nitric oxide-cyclic GMP signaling pathway in form-deprivation myopia of Guinea pig[J]. Chinese Ophthalmic Research, 2008, 26(9): 672-675.
3. Fischer AJ, Seltner RL, Stell WK. Opiate and N-methyl-D-aspartate receptors in form-deprivation myopia[J]. Vis Neurosci, 1998, 15(6): 1089-1096.
4. Lu F, Zhou X, Zhao H, et al. Axial myopia induced by a monocularly deprived facemask in guinea pigs: A non-invasive and effective model[J]. Exp Eye Res, 2006, 82(4): 628-636.

扩展功能

本文信息

- Supporting info
- PDF(1424KB)
- [HTML全文]
- 参考文献[PDF]
- 参考文献

服务与反馈

- 把本文推荐给朋友
- 加入我的书架
- 加入引用管理器
- 引用本文
- Email Alert
- 文章反馈
- 浏览反馈信息

本文关键词相关文章

- 形觉剥夺性近视
- 视网膜
- 一氧化氮
- 环磷酸鸟苷
- MK801

本文作者相关文章

- 文丹
- 刘双珍
- 毛俊峰
- 谭星平
- 夏朝华
- 尹楚南

PubMed

- Article by WEN Dan
- Article by LIU Shuangzhen
- Article by MAO Junfeng
- Article by TAN Xingping
- Article by XIA Chaohua
- Article by YIN Chunan

5. Boulton AD, Mishra N. Effects of the nitric oxide synthase inhibitor L-NAME on recognition and spatial memory deficits produced by different NMDA receptor antagonists in the rat[J]. *Neuropsychopharmacology*, 2010, 35(12): 2357-2366.
6. Fletcheer EL, Hack I, Brandstatter JH, et al. Synaptical localization of NMDA receptor subunits in the retina[J]. *Comp Neurol*, 2000, 420(1): 98-112.
7. Watanabe M, Mishina M, Inoue Y, et al. Differential distribution of the NMDA receptor channel subunit mRNAs in the mouse retina[J]. *Brain Res*, 1994, 634(2): 328-332.
8. 刘双珍, 文丹, 毛俊峰, 等. N-甲基-D-天冬氨酸受体1在形觉剥夺性近视豚鼠视网膜上的表达[J]. *眼视光学杂志*, 2008, 10(1): 1-5. LIU Shuangzhen, WEN Dan, MAO Junfeng, et al. The Expression of NMDAR1 in form-deprivation myopia of Guinea pig's retina[J]. *Journal of Optometry & Ophthalmology*, 2008, 10(1): 1-5.
9. Fischer AJ, Seltner RL, Stell WK. N-methyl-D-aspartate-induced excitotoxicity causes myopia in hatched chicks[J]. *Can J Ophthalmol*, 1997, 32(6): 373-377.
10. Li J, Henman MC, Tatlisumak T, et al. The pre-ischaemic neuroprotective effects of N1-dansyl-spermine in a transient focal cerebral ischaemia model in mice[J]. *Brain Res*, 2005, 1055(1/2): 180-185.
11. Wu J, Zou H, Strong JA, et al. Bimodal effects of MK-801 on locomotion and stereotypy in C57BL/6 mice[J]. *Psychopharmacology*, 2005, 177(3): 256-263.
12. Nickla DL, Danyanov P, Lytle G. Inhibiting the neuronal isoform of nitric oxide synthase has similar effects on the compensatory choroidal and axial responses to myopic defocus in chicks as does the nonspecific inhibitor L-NAME[J]. *Exp Eye Res*, 2009, 88(6): 1092-1099.
13. 朱玉广, 刘双珍, 吴小影. 视网膜内源性NO在形觉剥夺性近视眼中的作用研究[J]. *湖南医科大学学报*, 2003, 28(6): 631-634. ZHU Yuguang, LIU Shuangzhen, WU Xiaoying. Effects of nitric oxide on form-deprivation myopia in chicks[J]. *Bulletin of Hunan Medical University*, 2003, 28(6): 631-634.
14. 卢建, 余应年, 徐仁宝. 受体信号转导系统与疾病[M]. 济南: 山东科学技术出版社, 2001: 4731. LU Jian, YU Ying, XU Renbao. Receptor signal transduction system and disease[M]. Jinan: Shandong Science and Technology Publishing House, 2001: 4731.
15. Park CS, Pardhasaradhi K, Gianotti C, et al. Human retina expresses both constitutive and inducible isoforms of nitric oxide synthase mRNA[J]. *Biochem Biophys Res Commun*, 1994, 205(1): 85-91.
16. Koch KW, Duda T, Sharma RK. Photoreceptor specific cGMP guanylate cyclases in vertebrate phototransduction[J]. *Mol Cell Biochem*, 2002, 230(1/2): 97-106.
17. Zayas RM, Trimmer BA. Characterization of NO/cGMP mediated responses in identified motoneurons[J]. *Cell Mol Neurobiol*, 2007, 27(2): 191-209.

本刊中的类似文章

1. 姜志伟¹, 唐罗生¹, 朱晓华¹, 郭小健¹, 姜德咏², 龚玲². 孔源性视网膜脱离合并非继发性青光眼治疗的初步探讨[J]. *中南大学学报(医学版)*, 2008, 33(08): 737-740
2. 王燕¹, 董蕾¹, 邹百仑¹, 李慧鹏². Ghrelin 对大鼠十二指肠肌电活动的影响及作用机制[J]. *中南大学学报(医学版)*, 2008, 33(02): 93-98
3. 彭程^{1,2}, 肖涛¹, 罗远明¹, 刘夏君¹, 林绵辉¹, 胡金玺¹. IGF-1对IL-1诱导的兔关节软骨细胞NO和PGE2的影响[J]. *中南大学学报(医学版)*, 2008, 33(03): 197-203
4. 晋丽平; 王晨虹; 陈建林. 妊娠高血压综合征患者血浆 von Willebrand 因子与一氧化氮水平的变化[J]. *中南大学学报(医学版)*, 2001, 26(5): 477-
5. 吴静; 钟慧菊; 孙志湘; 易红; 雷闽湘. 灯盏花素治疗糖尿病周围神经病变的疗效观察[J]. *中南大学学报(医学版)*, 2002, 27(4): 337-
6. 秦波; 姜德咏; 唐朝珍; 刘湘平; 彭隆祥. 兔眼玻璃体视网膜界面正常结构与相关生理的实验研究[J]. *中南大学学报(医学版)*, 2002, 27(5): 413-
7. 谢晋良; 熊凤姣; 杨敬华; 齐范. 肾脏缺血预处理中一氧化氮作用的实验研究[J]. *中南大学学报(医学版)*, 2002, 27(5): 422-
8. 杨敬华; 周巧玲; 成小苗; 邓声莉; 吴彩玲. 川芎素对肾性高血压并慢性肾功能不全患者血浆ET-1和NO的影响[J]. *中南大学学报(医学版)*, 2002, 27(5): 445-
9. 陈胜喜; 王为; 黄凌瑾; 蒋海河; 张位星. 卡托普利对风湿性心脏病换瓣手术病人肺缺血-再灌损伤的延迟期保护作用[J]. *中南大学学报(医学版)*, 2002, 27(5): 451-
10. 倪江东; 丁仁奎; 吕国华; 王万春. 中药丹参注射液对兔脊髓损伤的影响[J]. *中南大学学报(医学版)*, 2002, 27(6): 507-
11. 王启常; 唐罗生; 姜德咏; 唐朝珍; 刘湘平. Coats病误诊原因分析[J]. *中南大学学报(医学版)*, 2003, 28(1): 56-
12. 唐勇军; 陈腊梅; 陈琼; 杨红忠; 尹本义. 肺心病患者血液NO和ET-1的改变及其意义[J]. *中南大学学报(医学版)*, 2003, 28(1): 59-
13. 刘豫¹, 许雪亮². 糖尿病性视网膜病变激光治疗的视力预后[J]. *中南大学学报(医学版)*, 2009, 34(03): 247-251
14. 籍雪颖, 张金嵩, 王艳婷, 孙宏亮, 贾沛生. Smad3信号通路及结缔组织生长因子在豚鼠西平抑制形觉剥夺性近视中的作用机制[J]. *中南大学学报(医学版)*, 2009, 34(04): 349-355
15. 伍校琼 蔡维君 罗学港.

猪后肢动脉生成过程中血管内皮细胞eNOS的表达

[J]. *中南大学学报(医学版)*, 2006, 31(01): 63-65