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PPAR- γ 通过PI3K/Akt及NF- κ B通路调控猪颈动脉支架重构

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《第三军医大学学报》[ISSN:1000-5404/CN:51-1095/R] 卷: 36 期数: 2014年第09期 页码: 888-892 栏目: 论著 出版日期: 2014-05-15

Title: PPAR- γ regulates vascular remodeling after pig carotid artery stent implantation through PI3K/Akt and NF- κ B pathway

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关键词: 过氧化物酶体增殖物激活受体- γ ; Akt; p-Akt; NF- κ B; 血管重构; 猪

Keywords: peroxisome proliferator activated receptors- γ ; Akt; p-Akt; NF- κ B; vascular remodeling; swine

分类号: R322.121; R363.22; R394.3

文献标志码: A

摘要: 目的 观察过氧化物酶体增殖物激活受体- γ (peroxisome proliferator activated receptors- γ , PPAR- γ)在猪颈动脉支架植入术后血管重构中的作用。 方法 将12只小型猪分为正常对照组($n=4$)和介入手术组($n=8$)，介入手术组采用猪颈动脉支架植入建立动物模型，分为支架组($n=4$)和罗格列酮(ROSI)组($n=4$)。术后3个月，应用数字减影血管造影(digital subtraction angiography, DSA)技术及HE染色评估血管形态学变化，应用免疫组织化学方法检测p-Akt及NF- κ B的表达变化，应用Western blot方法检测支架段血管PPAR- γ 、Akt、p-Akt及NF- κ B的表达变化。 结果 ①猪颈动脉DSA和HE染色结果发现支架组血管狭窄较ROSI组明显($P<0.01$)。②免疫组织化学染色结果发现支架组颈动脉血管p-Akt和NF- κ B表达较正常对照组明显增加($P<0.01$)，ROSI组颈动脉血管p-Akt和NF- κ B表达较支架组明显降低($P<0.01$)。③Western blot检测结果显示：支架组PPAR- γ 蛋白表达水平较正常对照组明显降低($P<0.01$)，NF- κ B和p-Akt蛋白表达水平较正常对照组明显增加($P<0.05$)；ROSI组PPAR- γ 蛋白表达水平较支架组明显增加($P<0.01$)，NF- κ B和p-Akt蛋白表达水平较支架组明显降低($P<0.01$)；Akt在正常对照组、支架组和ROSI组中的蛋白表达量差异无

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统计学意义 ($P>0.05$)。 结论 PPAR- γ 可能通过调节PI3K/Akt及NF- κ B的表达, 从而改善支架植入术后的血管重构。

Abstract: Objective To determine the role of peroxisome proliferator activated receptors- γ (PPAR- γ) in swine vascular remodeling after carotid artery stent implantation. Methods Totally 12 miniature pigs were randomly divided into normal control group ($n=4$) and intervention group ($n=8$). The animals of intervention group were adopted to establish the animal model of carotid artery stenting, and then randomly divided into stent subgroup ($n=4$) and rosiglitazone treatment (ROSI) subgroup ($n=4$). In 3 months after surgery, digital subtraction angiography (DSA) and HE staining was used to assess vascular morphology, immunohistochemical staining was applied to detect the expression of P-akt and NF- κ B, and Western blotting was employed to detect the expression of PPAR- γ , Akt, p-Akt and NF- κ B. Results DSA and HE staining demonstrated that vascular stenosis was more severe in the stent subgroup than the ROSI group ($P<0.01$). Immunohistochemical staining showed that the expression of p-Akt and NF- κ B were significantly increased in the carotid artery stent groups than normal control group ($P<0.01$), and that in ROSI subgroup was significantly lower than in the stent subgroup ($P<0.01$). Western blot analysis showed that the protein expression level of PPAR- γ was significantly lower ($P<0.01$), and those of NF- κ B and p-Akt were obviously higher in the stent subgroup than the normal control group ($P<0.05$), while that of PPAR- γ was significantly enhanced ($P<0.01$), and those of NF- κ B and p-Akt were markedly reduced in the ROSI subgroup than in the stent subgroup ($P<0.01$). However, there was no significant difference in the protein expression of Akt in normal control group, stent subgroup and ROSI subgroup ($P>0.05$). Conclusion PPAR- γ might mediate the expression of PI3K/Akt and NF- κ B, and then improve the vascular remodeling after stent implantation.

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