

# 下调TPX2抑制喉癌细胞生长的体外实验研究

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**Title:** Down regulation of TPX2 inhibits growth of laryngeal carcinoma cells in vitro

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**关键词:** 喉癌; 增殖; 凋亡; 细胞周期; TPX2

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**摘要:** 目的:研究TPX2对喉癌细胞增殖、克隆形成能力及细胞周期的影响。方法:Hep-2细胞中转染TPX2 siRNA、siRNA control记为TPX2 siRNA、siRNA -NC组,以不做转染的细胞为Con组。荧光定量PCR和Western blot分别测定细胞中TPX2 mRNA和蛋白水平,MTT测定各组细胞增殖,平板克隆实验测定各组细胞克隆形成能力,流式细胞术测定各组细胞周期情况,Western blot测定各组细胞中增殖细胞核抗原(PCNA)、细胞核增殖抗原(Ki-67)、细胞周期依赖性蛋白激酶4(CDK4)、细胞周期蛋白D1(Cyclin D1)蛋白水平。结果:siRNA control中TPX2 mRNA和蛋白水平、细胞存活率、克隆形成数目、细胞周期以及细胞中PCNA、Ki-67、CDK4、Cyclin D1蛋白水平与Con相比均没有明显变化( $P > 0.05$ )。TPX2 siRNA细胞中TPX2 mRNA和蛋白水平均明显低于Con( $P < 0.05$ )。TPX2 siRNA细胞存活率、克隆形成数目均明显低于Con,细胞G0/G1期比例明显高于Con,细胞中PCNA、Ki-67、CDK4、Cyclin D1蛋白水平明显低于Con( $P < 0.05$ )。结论:TPX2敲低可以降低喉癌细胞增殖、克隆形成能力,将细胞周期阻滞在G0/G1期,降低细胞中PCNA、Ki-67、CDK4、Cyclin D1蛋白表达。

**Abstract:** Objective: To study the effect of TPX2 on proliferation, clone formation and cell cycle of larynx cancer cells. Methods: The transfection of TPX2 siRNA and siRNA control in Hep-2 cells was recorded as TPX2 siRNA and siRNA-NC, and the cells that were not transfected were as Con. The levels of TPX2 mRNA and protein in cells were measured by fluorescence quantitative PCR and Western blot, respectively. MTT was used to determine the proliferation of cells in each group. The clone assay was used to determine the cell clone formation ability of each group. The cell cycle of each group was measured by flow cytometry. PCNA, Ki-67, CDK4 and Cyclin D1 in each group of cells were measured by Western blot. Results: There was no significant change in TPX2 mRNA and protein level, cell survival rate, colony formation number, cell cycle and PCNA, Ki-67, CDK4 and Cyclin D1 protein levels in siRNA control and Con ( $P > 0.05$ ). The levels of TPX2 mRNA and protein in TPX2 siRNA cells were significantly lower than that of Con ( $P < 0.05$ ). The survival rate of TPX2 siRNA cells and the number of clones were significantly lower than that of Con. And the proportion of G0/G1 phase in cell was significantly higher than that of Con, the level of PCNA, Ki-67, CDK4 and Cyclin D1 protein in the cells were significantly lower than that of Con ( $P < 0.05$ ). Conclusion: TPX2 knockdown can reduce the proliferation and clone forming ability of laryngeal carcinoma cells, block cell cycle at G0/G1 phase, and reduce the expression of PCNA, Ki-67, CDK4 and Cyclin D1.

## 参考文献/REFERENCES

- [1] Starska K, Forma E, Jóźwiak P, et al. Gene and protein expression of glucose transporter 1 and glucose transporter 3 in human laryngeal cancer-the relationship with regulatory hypoxia-inducible factor-1 $\alpha$  expression, tumor invasiveness, and patient prognosis [J]. *Tumour Biol*, 2015, 36(4): 1-13.
- [2] Sun X, Liu B, Zhao XD, et al. MicroRNA-221 accelerates the proliferation of laryngeal cancer cell line Hep-2 by suppressing Apaf-1 [J]. *Oncology Reports*, 2015, 33(3): 1221-1226.

- [3] Liu WW, Li EL, Wu LQ. Research progress on intercellular DNA damage response and its relationship with tumor occurrence by TPX2 [J]. *Guangdong Medical Journal*, 2016, 37(8):1245-1248. [刘万伟, 李恩亮, 鄂林泉. TPX2在细胞间期参与DNA损伤反应及其与肿瘤发生关系的研究进展 [J]. *广东医学*, 2016, 37(8):1245-1248.]
- [4] Liu HC, Liu YH, Zhao PR, et al. Small interfering RNA-mediated downregulation of targeting protein for Xklp2 suppresses the proliferation of human esophageal carcinoma EC9706 cells [J]. *World Chinese Journal of Digestology*, 2009, 17(28):2927-2930. [刘红春, 刘玉含, 赵培荣, 等. siRNA沉默TPX2基因对食管癌细胞EC9706的增殖和TPX2基因表达的影响 [J]. *世界华人消化杂志*, 2009, 17(28):2927-2930.]
- [5] Bo L, Zheng W, Lu F, et al. Overexpressed targeting protein for Xklp2 (TPX2) serves as a promising prognostic marker and therapeutic target for gastric cancer [J]. *Cancer Biology & Therapy*, 2016, 17(8):824-832.
- [6] Jiang P, Shen K, Wang X, et al. TPX2 regulates tumor growth in human cervical carcinoma cells [J]. *Molecular Medicine Reports*, 2014, 9(6):2347-2351.
- [7] Guo SS. TPX2 and Aurora-A expression and significance in laryngeal squamous cell carcinoma [D]. Hebei Medical University, 2015. [郭珊珊. TPX2和Aurora-A在喉鳞状细胞癌中的表达及意义 [D]. 河北医科大学, 2015.]
- [8] Petry S, Groen AC, Ishihara K, et al. Branching microtubule nucleation in *Xenopus* egg extracts mediated by augmin and TPX2 [J]. *Cell*, 2013, 152(4):768-777.
- [9] Fu J, Bian M, Xin G, et al. TPX2 phosphorylation maintains metaphase spindle length by regulating microtubule flux [J]. *J Cell Biol*, 2015, 210(3):373-383.
- [10] Roostalu J, Cade NI, Surrey T. Complementary activities of TPX2 and chTOG constitute an efficient importin-regulated microtubule nucleation module [J]. *Nature Cell Biology*, 2015, 17(11):1422-1434.
- [11] Roostalu J, Cade NI, Surrey T. Corrigendum: Complementary activities of TPX2 and chTOG constitute an efficient importin-regulated microtubule nucleation module [J]. *Nature Cell Biology*, 2015, 17(11):1512.
- [12] Song SF, Chang HP, Yang CR, et al. Effect and mechanism of lentivirus-mediated silencing of TPX2 gene expression on proliferation, migration and cell cycle of HeLa cells [J]. *Carcinogenesis, Teratogenesis & Mutagenesis*, 2017, 29(5):329-334. [宋淑芳, 常海平, 杨彩容, 等. 慢病毒介导的TPX2基因沉默对人宫颈癌HeLa细胞系增殖, 迁移和细胞周期的影响及机制 [J]. *癌变·畸变·突变*, 2017, 29(5):329-334.]
- [13] Zhu WB, Wu GY, Fang Y, et al. Expression and clinical significance of RHAMM and TPX2 in ovarian epithelial carcinoma [J]. *Jiangsu Medical Journal*, 2014, 40(7):803-806. [朱微波, 吴桂云, 方艳, 等. RHAMM及TPX2在卵巢上皮性癌中的表达及临床意义 [J]. *江苏医药*, 2014, 40(7):803-806.]
- [14] Smith LT, Mayerson J, Nowak NJ, et al. 20q11.1 amplification in giant-cell tumor of bone: array CGH, FISH, and association with outcome [J]. *Genes, Chromosomes and Cancer*, 2006, 45(10):957-966.
- [15] Liu Y. Effect of silencing TPX2 gene by shRNA on apoptosis and paclitaxel sensitivity of lung adenocarcinoma cell line A549 [D]. Chongqing Medical University, 2012. [刘莹. shRNA沉默TPX2基因对肺腺癌细胞A549凋亡及紫杉醇敏感性的影响 [D]. 重庆医科大学, 2012.]
- [16] Boehm EM, Powers KT, Kondratyck CM, et al. The proliferating cell nuclear antigen (PCNA)-interacting protein (PIP) motif of DNA polymerase mediates its interaction with the C-terminal domain of Rev1 [J]. *Journal of Biological Chemistry*, 2016, 291(16):8735-8744.
- [17] Billon P, Li J, Lambert JP, et al. Acetylation of PCNA sliding surface by Eco1 promotes genome stability through homologous recombination [J]. *Molecular Cell*, 2017, 65(1):78-90.
- [18] Hoster E, Rosenwald A, Berger F, et al. Prognostic value of Ki-67 index, cytology, and growth pattern in mantle-cell lymphoma: results from randomized trials of the European mantle cell lymphoma network [J]. *Journal of Clinical Oncology*, 2016, 34(12):1386-1394.
- [19] Qiu X, Mei J, Yin J, et al. Correlation analysis between expression of PCNA, Ki67 and COX-2 and X-ray features in mammography in breast cancer [J]. *Oncology Letters*, 2017, 14(3):2912-2918.
- [20] He XM, Huang R, Yu Y, et al. CDK2-AP1 inhibits breast cancer growth by regulating cell cycle [J]. *Zhejiang Clinical Medical Journal*, 2015, 17(7):1059-1061. [何向明, 黄润, 俞洋, 等. CDK2-AP1通过调控细胞周期抑制乳腺癌生长 [J]. *浙江临床医学*, 2015, 17(7):1059-1061.]
- [21] Hubbi ME, Semenza GL. An essential role for chaperone-mediated autophagy in cell cycle progression [J]. *Autophagy*, 2015, 11(5):850-851.
- [22] Xia B, Yang S, Liu T, et al. miR-211 suppresses epithelial ovarian cancer proliferation and cell-cycle progression by targeting Cyclin D1 and CDK6 [J]. *Molecular Cancer*, 2015, 14(1):57.
- [23] Blanco I, Kuchenbaecker K, Cuadras D, et al. Assessing associations between the AURKA-HMMR-TPX2-TUBG1 functional module and breast cancer risk in BRCA1/2 mutation carriers [J]. *PloS One*, 2015, 10(4):e0120020.

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备注/Memo: -

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