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## 综述

### lncRNAs参与基因表达调控机制的研究进展

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摘要:

长链非编码RNAs (long non-coding RNAs, lncRNAs)是一组内源性、长度超过200 nt、缺少特异完整的开放阅读框(open reading frame, ORF)、无或很少有蛋白编码功能的RNAs分子。近年来的研究发现, lncRNAs与许多重要的生物学过程相关, 如基因组印记、细胞分化、免疫反应等。lncRNAs在表观遗传水平、转录水平和转录后水平等多个

层面调节基因的表达, 通过介导染色质重塑和组蛋白修饰、干扰转录、调节选择性剪接模式、生成小RNAs、调节蛋白质活性、改变蛋白质定位等方式, 参与机体生长、发育、衰老及死亡等重要生命活动的调控。关于lncRNAs参与基因表达调控的机制有待进一步研究, 这将有助于深入理解疾病的发病机制, 为寻找疾病分子标记物、药物靶点提供新的方向。

关键词: lncRNAs 表观遗传调控 转录调控 转录后调控

### Advances in regulation of gene expression mediated by lncRNAs

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Abstract:

Long non-coding RNAs (lncRNAs) are a group of endogenous RNA molecules which exceed 200 nt in length, lack complete specific open reading frame, and completely lack or possess very limited protein-coding capacity. Recent studies have revealed that lncRNAs participate in critical processes such as genomic imprinting, cell differentiation, and immune reaction, etc. lncRNAs regulate gene expression at the epigenetic, transcriptional and post-transcriptional levels by modulating chromatin remodeling and histone modifications, interfering the transcription, regulating patterns of alternative splicing, generating small RNAs, and modulating protein activation and localization. Through their numerous functions, lncRNAs play critical roles in the growth, development, senescence, death, and other important physiological and pathological processes. Further investigation into the regulation of gene expression mediated by lncRNAs will be of great value in the thorough understanding of pathogenies and provide new molecular markers and drug targets of diseases.

Keywords: lncRNAs epigenetic regulation transcriptional regulation post-transcriptional regulation

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## 参考文献：

1. Ponting CP, Oliver PL, Reik W. Evolution and functions of long noncoding RNAs[J]. *Cell*, 2009, 136(4): 629-641.
2. Carninci P, Kasukawa T, Katayama S, et al. The transcriptional landscape of the mammalian genome[J]. *Science*, 2005, 309(5740): 1559-1563.
3. Kapranov P, Cheng J, Dike S, et al. RNA maps reveal new RNA classes and a possible function for pervasive transcription[J]. *Science*, 2007, 316(5830): 1484-1488.
4. Guttman M, Rinn JL. Modular regulatory principles of large noncoding RNAs[J]. *Nature*, 2012, 482(7385): 339-346.
5. Maass PG, Rump A, Schulz H, et al. A misplaced lncRNA causes brachydactyly in humans[J]. *J Clin Invest*, 2012, 122(11): 3990-4002.
6. Askarian-Amiri ME, Crawford J, French JD, et al. SNORD-host RNA Zfas1 is a regulator of mammary development and a potential marker for breast cancer[J]. *RNA*, 2011, 17(5): 878-891.
7. Cabianca DS, Casa V, Gabellini D. A novel molecular mechanism in human genetic disease: A DNA repeat-derived lncRNA point-ofview[J]. *RNA Biol*, 2012, 9(10): 1211-1217.
8. Torarinsson E, Yao Z, Wiklund ED, et al. Comparative genomics beyond sequence-based alignments: RNA structures in the ENCODE regions[J]. *Genome Res*, 2008, 18(2): 242-251.
9. Arun G, Akhade VS, Donakonda S, et al. mrhl RNA, a long noncoding RNA, negatively regulates Wnt signaling through its protein partner Ddx5/p68 in mouse spermatogonial cells[J]. *Mol Cell Biol*, 2012, 32(15): 3140-3152.
10. Beaulieu YB, Kleinman CL, Landry-Voyer AM, et al. Polyadenylation-dependent control of long noncoding RNA expression by the poly(A)-binding protein nuclear 1[J]. *PLoS Genet*, 2012, 8(11): e1003078.
11. Aguiló F, Zhou MM, Walsh MJ. Long noncoding RNA, polycomb, and the ghosts haunting INK4b-ARF-INK4a expression[J]. *Cancer Res*, 2011, 71(16): 5365-5369.
12. Gutschner T, Diederichs S. The hallmarks of cancer: a long non-coding RNA point of view[J]. *RNA Biol*, 2012, 9(6): 703-719.
13. Wilusz JE, Sunwoo H, Spector DL. Long noncoding RNAs: functional surprises from the RNA world[J]. *Genes Dev*, 2009, 23(13): 1494-1504.
14. Umlauf D, Fraser P, Nagano T. The role of long non-coding RNAs in chromatin structure and gene regulation: variations on a theme[J]. *Biol Chem*, 2008, 389(4): 323-331.
15. 胡春燕, 周建华. DNA甲基化修饰异常与肺癌[J]. 国际病理科学与临床杂志, 2009(5): 398-402.  
HU Chunyan, ZHOU Jianhua. Abnormality of DNA methylation modification and lung cancer[J]. International Journal of Pathology and Clinical Medicine, 2009, (5): 398-402.
16. Qureshi IA, Mattick JS, Mehler MF. Long non-coding RNAs in nervous system function and disease[J]. *Brain Res*, 2010, 1338(18): 20-35.
17. Redrup L, Branco MR, Perdeaux ER, et al. The long noncoding RNA Kcnq1ot1 organises a lineage-specific nuclear domain for epigenetic gene silencing[J]. *Development*, 2009, 136(4): 525-530.
18. Imamura T, Yamamoto S, Ohgane J, et al. Non-coding RNA directed DNA demethylation of Sphk1 CpG island[J]. *Biochem Biophys Res Commun*, 2004, 322(2): 593-600.
19. Wu SC, Kallin EM, Zhang Y. Role of H3K27 methylation in the regulation of lncRNA expression[J]. *Cell Res*, 2010, 20(10): 1109-1116.
20. Braconi C, Kogure T, Valeri N, et al. microRNA-29 can regulate expression of the long non-coding RNA gene MEG3 in hepatocellular cancer[J]. *Oncogene*, 2011, 30(47): 4750-4756.
21. Li M, Wen S, Guo X, et al. The novel long non-coding RNA CRG regulates Drosophila locomotor behavior[J]. *Nucleic Acids Res*, 2012, 40(22): 11714-11727.
22. Guttman M, Amit I, Garber M, et al. Chromatin signature reveals over a thousand highly conserved large non-coding RNAs in mammals[J].

- Nature, 2009, 458(7235): 223-227.
23. Qi W, Song X, Li L. Long non-coding RNA-guided regulation in organisms[ J]. Sci China Life Sci, 2013, 56(10): 891-896.
24. Feng J, Bi C, Clark B S, et al. The Evf-2 noncoding RNA is transcribed from the Dlx-5/6 ultraconserved region and functions as a Dlx-2 transcriptional coactivator[ J]. Genes Dev, 2006, 20(11): 1470-1484.
25. Rapicavoli NA, Poth EM, Zhu H, et al. The long noncoding RNA Six3OS acts in trans to regulate retinal development by modulating Six3 activity[ J]. Neural Dev, 2011, 6(1): 32.
26. Martianov I, Ramadass A, Serra B A, et al. Repression of the human dihydrofolate reductase gene by a non-coding interfering transcript[ J]. Nature, 2007, 445(7128): 666-670.
27. Yik JH, Chen R, Nishimura R, et al. Inhibition of P-TEFb (CDK9/Cyclin T) kinase and RNA polymerase II transcription by the coordinated actions of HEXIM1 and 7SK snRNA[J]. Mol Cell, 2003, 12(4): 971-982.
28. Wang X, Arai S, Song X, et al. Induced ncRNAs allosterically modify RNA-binding proteins in cis to inhibit transcription[ J]. Nature, 2008, 454(7200): 126-130.
29. Mariner PD. Human Alu RNA is a modular transacting repressor of mRNA transcription during heat shock[ J]. Mol Cell, 2008, 29(4): 499-509.
30. Beltran M, Puig I, Pena C, et al. A natural antisense transcript regulates Zeb2/Sip1 gene expression during Snail1-induced epithelialmesenchymal transition[ J]. Genes Dev, 2008, 22(6): 756-769.
31. Jalali S, Jayaraj GG, Scaria V. Integrative transcriptome analysis suggest processing of a subset of long non-coding RNAs to small RNAs[ J]. Biol Direct, 2012, 7(1): 25.
32. Okamura K, Chung WJ, Ruby JG, et al. The Drosophila hairpin RNA pathway generates endogenous short interfering RNAs[ J]. Nature, 2008, 453(7196): 803-806.
33. Aravin AA, Hannon GJ, Brennecke J. The Piwi-piRNA pathway provides an adaptive defense in the transposon arms race[ J]. Science, 2007, 318(5851): 761-764.
34. Ebert MS, Neilson JR, Sharp PA. MicroRNA sponges: competitive inhibitors of small RNAs in mammalian cells[ J]. Nat Methods, 2007, 4(9): 721-726.
35. Clemson CM, Hutchinson JN, Sara SA, et al. An architectural role for a nuclear noncoding RNA: NEAT1 RNA is essential for the structure of paraspeckles[ J]. Mol Cell, 2009, 33(6): 717-726.
36. Blower MD, Nachury M, Heald R , et al. A Rae1-containing ribonucleoprotein complex is required for mitotic spindle assembly[ J]. Cell, 2005, 121(2): 223-234.
37. Kloc M, Bilinski S, Dougherty MT. Organization of cytokeratin cytoskeleton and germ plasm in the vegetal cortex of *Xenopus laevis* oocytes depends on coding and non-coding RNAs: three-dimensional and ultrastructural analysis[ J]. Exp Cell Res, 2007, 313(8): 1639-1651.
38. Willingham AT, Orth AP, Batalov S, et al. A strategy for probing the function of noncoding RNAs finds a repressor of NFAT[J]. Science, 2005, 309(5740): 1570-1573.
39. 何丹, 段朝军. 长链非编码RNA在肿瘤研究中的进展[ J]. 国际病理科学杂志, 2012, 32(4): 297-301.
- HE Dan. Progress with long non-coding RNA in cancer research [ J]. International Journal of Pathology and Clinical Medicine, 2012, 32(4): 297-301.

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