

CTC、ctDNA在结直肠癌诊断中的研究进展

《现代肿瘤医学》[ISSN:1672-4992/CN:61-1415/R] 期数: 2019年10期 页码: 1840-1844 栏目: 综述 出版日期: 2019-04-08

Title: Research progress of circulating tumor cells and circulating tumor DNA in the diagnosis of colorectal cancer

作者: 肖泽文; 陶冀

哈尔滨医科大学附属肿瘤医院消化内科，黑龙江 哈尔滨 150081

Author(s): Xiao Zewen; Tao Ji

Department of Gastroenterology, Harbin Medical University Cancer Hospital, Heilongjiang Harbin 150081, China.

关键词: CTC; ctDNA; 结直肠癌; 诊断; 肿瘤标记物

Keywords: CTC; ctDNA; CRC; diagnosis; tumor markers

分类号: R735.3

DOI: 10.3969/j.issn.1672-4992.2019.10.044

文献标识码: A

摘要: 结直肠癌 (colorectal cancer,CRC) 是世界上第三大最常见的癌症类型,也是导致癌症死亡的第四大原因。CRC的常规治疗策略包括手术、新辅助治疗和辅助治疗。不幸的是,大约50%的CRC患者仅在晚期被诊断,因此显著降低了不同治疗方案的可用性。学者们不断寻找更准确的诊断结直肠癌的方法,除了通过蛋白质组学技术研究新的血清肿瘤生物标记物外,还引入循环肿瘤细胞 (CTC) 和循环肿瘤DNA (ctDNA) 等新概念。本文就血液中CTC、ctDNA在结直肠癌诊断中的研究进展作一综述。

Abstract: Colorectal cancer (CRC) is the third most common type of cancer in the world and the fourth leading cause of death from cancer. Conventional treatment strategies for CRC include surgery, neoadjuvant therapy, and adjuvant therapy. Unfortunately, approximately 50% of CRC patients are diagnosed only late and therefore significantly reduce the availability of different treatment options. Scholars continue to search for more accurate methods of diagnosing colorectal cancer. In addition to new serum tumor biomarkers through proteomics, new concepts such as circulating tumor cells (CTC) and circulating tumor DNA (ctDNA) have been introduced. This article reviews the research progress of CTC and ctDNA in the diagnosis of colorectal cancer.

参考文献/REFERENCES

- [1] Jia S,Zhang R,Li Z,et al.Clinical and biological significance of circulating tumor cells,circulating tumor DNA, and exosomes as biomarkers in colorectal cancer [J]. Oncotarget,2017,8(33):55632-55645.
- [2] Chen W,Zheng R,Baade PD,et al.Cancer statistics in China,2015 [J]. CA:A Cancer Journal for Clinicians,2016,66(2):115-132.
- [3] Heitzer E,Auer M,Ulz P,et al.Circulating tumor cells and DNA as liquid biopsies [J]. Genome Medicine,2013,5(8):73.
- [4] Haber DA,Velculescu VE.Blood-based analyses of cancer:circulating tumor cells and circulating tumor DNA [J]. Cancer Discovery,2014,4(6):650-661.
- [5] Scher HI,Morris MJ,Larson S,et al.Validation and clinical utility of prostate cancer biomarkers [J]. Nature Reviews Clinical Oncology,2013,10(4):225-234.
- [6] Huang X,Gao P,Song Y,et al.Meta-analysis of the prognostic value of circulating tumor cells detected with the Cell Search System in colorectal cancer [J]. BMC Cancer,2015,15:202.
- [7] Galletti G,Sung MS,Vahdat LT,et al.Isolation of breast cancer and gastric cancer circulating tumor cells by use of an anti HER2-based microfluidic device [J]. Lab On A Chip,2014,14(1):147-156.
- [8] Chen P,Li GT,Liu HB.Research progress of hematological tumor cell markers CTC and ctDNA in early diagnosis of gastric cancer [J]. Northwestern Journal of Defense Medicine,2017,38(10):695-698. [陈鹏,李盖天,刘宏斌.血液肿瘤细胞标记物CTC、ctDNA在胃癌早期诊断中的研究进展 [J].西北国防医学杂志,2017,38(10):695-698.]
- [9] Chen X,Zhou F,Li X,et al.Folate receptor-positive circulating tumor cell detected by LT-PCR-based method as a diagnostic biomarker for non-small-cell lung cancer [J]. Journal of Thoracic Oncology:Official Publication of The International Association for the Study of Lung Cancer,2015,10(8):1163-1171.
- [10] Cristofanilli M,Fortina P.Circulating tumor DNA to monitor metastatic breast cancer [J]. The New England Journal of Medicine,2013,369(1):93.
- [11] Sastre J,Maestro ML,Puente J,et al.Circulating tumor cells in colorectal cancer:Correlation with clinical and pathological variables [J]. Annals of Oncology:Official Journal of the European Society for Medical Oncology,2008,19(5):935-938.
- [12] Gazzaniga P, Gianni W,Raimondi C,et al.Circulating tumor cells in high-risk nonmetastatic colorectal cancer [J]. Tumour Biology:The Journal of the

International Society for Oncodevelopmental Biology and Medicine,2013,34(5):2507-2509. [13] Grady WM,Pritchard CC.Molecular alterations and biomarkers in colorectal cancer [J]. Toxicologic Pathology,2014,42(1):124-139. [14] Skubitz APN,Taras EP,Boylan KLM,et al.Targeting CD133 in an in vivo ovarian cancer model reduces ovarian cancer progression [J]. Gynecol Oncol,2013,130(3):579-587. [15] Mao J,Song B,Shi Y,et al.ShRNA targeting Notch1 sensitizes breast cancer stem cell to paclitaxel [J]. Int J Biochem Cell Biol,2013,45(6):1064-1073. [16] Salnikov AV,Bretz NP,Perne C,et al.Antibody targeting of CD24 efficiently retards growth and influences cytokine milieu in experimental carcinomas [J]. Br J Cancer,2013,108(7):1449-1459. [17] Lianna Li,Charles F Bellows.Doublecortin-like kinase 1 exhibits cancer stem cell-like characteristics in a human colon cancer cell line [J]. Chinese Journal of Cancer Research,2013,25(02):134-142. [18] Sarkar S,O'Connell M,Kantara C,et al.Abstract 3903:A sub-set of DCLK1+ve colon cancer stem cells (CSCs) survive curcumin induced autophagy,while co-treatment with curcumin +DCLK1-siRNA eliminates CSCs:Role of long and short isoforms of DCLK1 [J]. Gastroenterology,2014,146(5):S-340. [19] Mirzaei A,Tavoosidana G,Modarressi MH,et al.Upregulation of circulating cancer stem cell marker,DCLK1 but not Lgr5,in chemoradiotherapy-treated colorectal cancer patients [J]. Tumor Biology,2015,36(6):4801-4810. [20] Valladaresayerbes M,Blancocalvo M,Reboreda M,et al.Evaluation of the adenocarcinoma-associated gene AGR2 and the intestinal stem cell marker LGR5 as biomarkers in colorectal cancer [J]. International Journal of Molecular Sciences,2012,13(4):4367-4387. [21] Mirzaei A,Tavoosidana G,Rad AA,et al.A new insight into cancer stem cell markers:Could local and circulating cancer stem cell markers correlate in colorectal cancer [J]. Tumor Biology,2016,37(2):2405-2414. [22] Fina E,Necchi A,Bottelli S,et al.Detection of circulating tumour cells in urothelial cancers and clinical correlations:Comparison of two methods [J]. Disease Markers,2017,2017(11):3414910. [23] Wu LL,Wen CY,Hu J,et al.Nanosphere-based one-step strategy for efficient and nondestructive detection of circulating tumor cells [J]. Biosensors & Bioelectronics,2017,94:219. [24] Wen-SY T,Ashish N,Oscar S,et al.Prospective clinical study of circulating tumor cells for colorectal cancer screening [C]. Chicaco:ASCO,2018. [25] Edge S,Byrd DR,Compton CC,et al.AJCC cancer staging manual 7th Edition [M]. New York:Springer,2010:589-628. [26] Xu RH,Wei W,Krawczyk M,et al.Circulating tumour DNA methylation markers for diagnosis and prognosis of hepatocellular carcinoma [J]. Nature Materials,2017,16(11):1155-1161. [27] Stroun M,Maurice P,Vasioukhin V,et al.The origin and mechanism of circulating DNA [J]. Annals of the New York Academy of Sciences,2000,906:161-168. [28] Frattini M,Gallino G,Signoroni S,et al.Quantitative and qualitative characterization of plasma DNA identifies primary and recurrent colorectal cancer [J]. Cancer Letters,2008,263(2):170-181. [29] Strickler JH,Loree JM,Ahronian LG,et al.Genomic landscape of cell-free DNA in patients with colorectal cancer [J]. Cancer Discovery,2018,8(2):164-173. [30] Mouliere F,El Messaoudi S,Gongora C,et al.Circulating cell-free DNA from colorectal cancer patients may reveal high KRAS or BRAF mutation load [J]. Translational Oncology,2013,6(3):319-328. [31] Flamini E,Mercatali L,Nanni O,et al.Free DNA and carcinoembryonic antigen serum levels:an important combination for diagnosis of colorectal cancer [J]. Clinical Cancer Research:An Official Journal of the American Association for Cancer Research,2006,12(23):6985-6988. [32] Perrone F,Lampis A,Bertan C,et al.Circulating free DNA in a screening program for early colorectal cancer detection [J]. Tumori,2014,100(2):115-121. [33] Taly V,Pekin D,Benhamim L,et al.Multiplex picodroplet digital PCR to detect KRAS mutations in circulating DNA from the plasma of colorectal cancer patients [J]. Clinical Chemistry,2013,59(12):1722-1731. [34] Oxnard GR,Pawletz CP,Kuang Y,et al.Noninvasive detection of response and resistance in EGFR-mutant lung cancer using quantitative next-generation genotyping of cell-free plasma DNA [J]. Clinical Cancer Research:An Official Journal of the American Association for Cancer Research,2014,20(6):1698-1705. [35] Xu T,Kang X,You X,et al.Cross-platform comparison of four leading technologies for detecting EGFR mutations in circulating tumor DNA from non-small cell lung carcinoma patient plasma [J]. Theranostics,2017,7(6):1437-1446. [36] Kwapisz D.The first liquid biopsy test approved.Is it a new era of mutation testing for non-small cell lung cancer [J]. Annals of Translational Medicine,2017,5(3):46. [37] Goodwin S,McPherson JD,McCombie WR.Coming of age:ten years of next-generation sequencing technologies [J]. Nature Reviews Genetics,2016,17(6):333-351. [38] Malapelle U,Mayord C,Rocco D,et al.Development of a gene panel for next-generation sequencing of clinically relevant mutations in cell-free DNA from cancer patients [J]. British Journal of Cancer,2017,116(6):802-810. [39] Tie J,Kinde I,Wang Y,et al.Circulating tumor DNA as an early marker of therapeutic response in patients with metastatic colorectal cancer [J]. Annals of Oncology:Official Journal of the European Society for Medical Oncology,2015,26(8):1715-1722. [40] Spindler KL,Pallisgaard N,Andersen RF,et al.Circulating free DNA as biomarker and source for mutation detection in metastatic colorectal cancer [J]. PloS One,2015,10(4):e0108247. [41] Bo BL,Lee EJ,Jung EH,et al.Aberrant methylation of APC,MGMT,RASSF2A, and wif-1 genes in plasma as a biomarker for early detection of colorectal cancer [J]. Clinical Cancer Research,2009,15(19):6185-6191. [42] Church TR,Wandell M,Loftonday C,et al.Original article:Prospective evaluation of methylated SEPT9 in plasma for detection of asymptomatic colorectal cancer [J]. Gut,2014,63(2):317-325. [43] Garrigou S,Perkins G,Garlan F,et al.A study of hypermethylated circulating tumor DNA as a universal colorectal cancer biomarker [J]. Clinical Chemistry,2016,62(8):1129-1139. [44] Rasmussen SL,Krarup HB,Sunesen KG,et al.Hypermethylated DNA,a circulating biomarker for colorectal cancer detection [J]. Plos One,2017,12(7):e0180809. [45] Nadal C,Winder T,Gerger A,et al.Future perspectives of circulating tumor DNA in colorectal cancer [J]. Tumor Biology,2017,39(5):101042831770574. [46] Guo S,Diep D,Plongthongkum N,et al.Identification of methylation haplotype blocks aids in deconvolution of heterogeneous tissue samples and tumor tissue-of-origin mapping from plasma DNA [J]. Nature Genetics,2017,49(4):635-642.

备注/Memo: 吴阶平医学基金会临床科研专项资助基金 (编号: 370675017202)

更新日期/Last Update: 1900-01-01