THESIS ABSTRACT

Evaluation of the use of long noncoding RNAs as biomarkers for diagnosis of breast cancer

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Background: FAM83H antisense RNA 1 (FAM83H-AS1) and long noncoding RNA activated by TGF β (IncRNA-ATB) are two IncRNAs that have tumor promoting functions in breast cancer (BC). Matrix metalloproteinase-9 (MMP-9) is another tumor marker that exerts an oncogenic role in BC and facilitate cancer invasion and metastasis. Aims: Our study aimed to 1) analyze serum levels of FAM83H-AS1, IncRNA-ATB, and MMP-9 in BC patients, 2) compare their diagnostic role with that of CA15-3 in BC patients with different stages (I-II, III, and IV), and 3) correlate the levels of the measured lncRNAs, MMP-9, and CA15-3 with the clinicopathological features of BC. Materials and Methods: Serum FAM83H-AS1 and IncRNA-ATB levels were analyzed in 90 BC patients and 30 healthy controls using RT-PCR. While serum levels of MMP-9 and CA15-3 were measured in the same studied groups by ELISA. Results: FAM83H-AS1, IncRNA-ATB, MMP-9, and CA15-3 levels were significantly elevated in sera of BC patients. ROC curve analysis showed that IncRNA-ATB and MMP-9 had higher AUC values than CA15-3 in diagnosis of stage I-II patients (AUC: 0.844 and 0.898, p= 0.000 for IncRNA-ATB and MMP-9, respectively versus 0.738, p= 0.002 for CA15-3). FAM83H-AS1 level was significantly correlated with higher stage, larger tumor size, positive lymph node metastasis, and distant metastasis. FAM83H-AS1 level was also significantly increased in PR-negative and ki67-rich tumors. Conclusion: Our study demonstrated that circulating IncRNA-ATB and MMP-9 could be used as diagnostic biomarkers for early BC. FAM83H-AS1 could also be a potential candidate for staging BC and monitoring disease progression.

Keywords: Breast cancer diagnosis; FAM83H-AS1; LncRNA-ATB; Long non-coding RNA; MMP-9

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