Mutation analysis of the FHIT gene in bronchoscopic specimens from patients with suspected lung cancer

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ABSTRACT

Aims and background. Lung cancer is a leading cause of cancer death worldwide. However, despite recent advances in molecular biology that have revealed various genetic changes in lung cancer, the prognostic outcome of lung cancer patients has improved only minimally. This situation has changed fundamentally with the identification of molecular abnormalities that are characteristic of premalignant changes, such as changes in tumor suppressor genes, loss of heterozygosity at crucial sites, and activation of oncogenes. Inactivation of the tumor suppressor gene Fragile Histidine Triad (FHIT) is a frequent genetic change in lung cancer. The aim of this study was to identify FHIT gene alterations in bronchoscopy specimens of patients with suspected lung cancer and to determine the molecular relevance, if any, of FHIT alterations in the development of cancer.

Patients and methods. Sixty-two patients with suspected lung tumors were screened for variations within exons 5-9 of the FHIT gene using intronic primer pairs and single-strand conformation polymorphism and sequencing analysis.

Results. FHIT gene alterations were detected in 27 out of 62 bronchoscopic specimens (43.54%). All of these alterations were identified as T to A alteration at position IVS8-17. This intronic variant also was identified in approximately half of control cases (45%).

Conclusions. Our findings showed that the FHIT IVS8-17 T to A alteration identified in bronchoscopy specimens from patients with clinically suspected lung cancer is a polymorphism found in the Turkish population. We think that this polymorphism does not affect gene function because it is located in the intron portion of the gene and is present in many cancer patients as well as healthy subjects. We suggest that the FHIT gene may be turned off in lung carcinogenesis via other genetic or epigenetic mechanisms rather than mutations.

Key words: lung cancer; bronchoscopy specimen; FHIT gene; sequence variant; single-strand conformation polymorphism; DNA sequencing.

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