NUCLEAR LOCALIZATION OF INTRONIC *RASSF1* TRANSCRIPTS IN HUMAN TISSUES USING *IN SITU* HYBRIDIZATION

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A new class of long intronic noncoding RNAs (ncRNAs) transcribed in the human genome with sense and/or antisense orientation relative to the corresponding protein-coding gene was recently identified. These intronic ncRNAs are postulated to be involved in processes related to posttranscriptional control of gene expression in normal and diseased tissues. Among them, intronic antisense transcripts (GenBank accession AY545527 and AY545528) originated in the RASSF1 locus, a tumor suppressor gene, were shown to have levels correlated to tumor malignancy in prostate cancer. In this work we investigated the expression pattern and sub-cellular localization of RASSF1 intronic transcripts using RNA in situ hybridization. We detected the presence of both sense and antisense intronic RASSF1 transcripts in histological sections of kidney tumor and non-tumor tissue adjacent to surgically removed prostate tumor. The sense transcript RASSF1 was more abundant than its corresponding antisense transcript, as previously observed in quantitative PCR experiments performed by our group. Interestingly, intronic RASSF1 transcripts seem to predominantly localize at the cell nucleus. These results support a mechanism where intronic RASSF1 transcripts exert their function by acting in *cis* through base-pairing with unprocessed sense RASSF1 pre-mRNA in the nucleus, thus affecting the levels of Rassf1 protein.

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