ZytoLight® SPEC WT1 Dual Color Break Apart Probe

Background

The ZytoLight ® SPEC WT1 Dual Color Break Apart Probe is designed to detect translocations involving the chromosomal region 11p13 harboring the WT1 (Wilms tumor 1) gene.

The WT1 gene is located on 11p13 and encodes a zinc finger DNA-binding protein that acts as a transcriptional activator or repressor depending on the cellular or chromosomal context. Inactivating mutations in the tumor suppressor gene WT1 have been identified in patients with Wilms' tumor and in a subset of sporadic cancers.

However, in desmoblastic small round cell tumors (DSRCT) recurrent translocations affecting the WT1 gene have been found. DSRCT is a highly aggressive mesenchymal tumor that primarily affects male adolescents and young adults. The translocation t(11;22)(p13;q12.2) is detectable in virtually all DSRCT tested and results in the fusion of the potent transcriptional activator domain of the EWSR1 gene and the DNA-binding zinc-finger domains of the WT1 gene. The EWSR1-WT1 chimeric protein acts as an oncogenic transcription factor as evidenced by its ability to transform cells in vitro.

While EWSR1 rearrangements are present in about 90% of DSRCT but are also frequently found in other small round blue cell neoplasms as e.g. Ewing sarcoma, WT1 translocations are exclusively found in DSRCT. Hence, detection of the t(11;22) by Fluorescence in situ Hybridization represents a valuable tool for the differential diagnosis of DSRCT.

References Gerald WL, et al. (1995) Pro Natl Acad Sci USA 92: 1028-32. Kim J, et al. (1998) Oncogene 16: 1973-9. Ladanyi M & Gerald W (1994) Cancer Res 54: 2837-40. Wang ZY, et al. (1993) J Biol Chem 268: 9172-5.

Probe Description

The SPEC WT1 Dual Color Break Apart Probe is a mixture of two direct labeled probes hybridizing to the 11p13 band. The orange fluorochrome direct labeled probe hybridizes distal and the green fluorochrome direct labeled probe hybridizes proximal to the WT1 gene.



SPEC WT1 Probe map (not to scale).

Results

In an interphase nucleus lacking a translocation involving the 11p13 band, two orange/ green fusion signals are expected representing two normal (non-rearranged) 11p13 loci. A signal pattern consisting of one orange/green fusion signal, one orange signal, and a separate green signal indicates one normal 11p13 locus and one 11p13 locus affected by a translocation.

Molecular diagnostics simplified



SPEC WT1 Dual Color Break Apart Probe hybridized to normal interphase cells as indicated by two orange/green fusion signals per nucleus.



Desmoblastic small round cell tumor tissue section with translocation affecting the 11p13 locus as indicated by one non-rearranged orange/green fusion signal, one orange signal, and one separate green signal indicating the translocation.

Prod. No.	Product	Label	Tests* (Volume)
Z-2142-50	Zyto <i>Light</i> SPEC WT1 Dual Color Break Apart Probe C E IVD	•/•	5 (50 µl)
Related Pro	lucts		
Z-2028-5	Zyto <i>Light</i> FISH-Tissue Implementation Kit C E IVD		5
	Incl. Heat Pretreatment Solution Citric, 150 ml; Pepsin Solution, 1ml; Wash Buffer SSC, 150 ml; 25x Wash Buffer A, 50 ml; DAPI/DuraTect-Solution, 0.2 ml		
lsing 10 µl probe solu	ion per test. CE [IVD] only available in certain countries. All other countries research use only! Please contact your local dealer for more information.		

ZytoLight[®] FISH probes are direct labeled using the unique ZytoLight[®] Direct Label System II providing improved signal intensity. Advanced specificity of the single copy SPEC probes is obtained by the unique ZytoVision[®] Repeat Subtraction Technique.

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