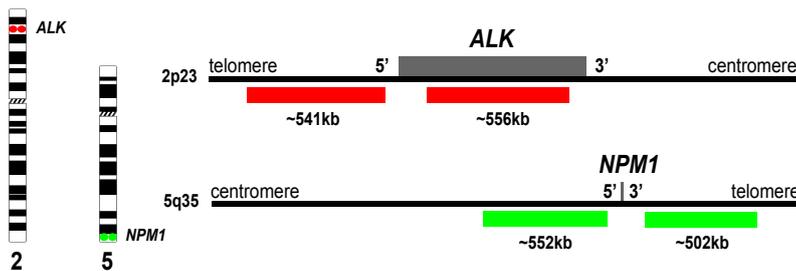


Intended Use

The *ALK/NPM1* DNA-FISH Probe is designed to detect the translocation between the *ALK* gene located at 2p23 and the *NPM1* gene located at 5q35, using fluorescence *in situ* hybridization (FISH); the translocation between the *ALK* and *NPM1* gene is designated as t(2;5) (p23;q35). By conventional cytogenetics, the translocation occurs in up to 50% of anaplastic large cell lymphoma (ALCL) cases.^[1] As assessed by immunohistochemistry, expression of the fusion protein *ALK/NPM1* that is generated by t(2;5), occurs more frequently in childhood ALCL with an occurrence rate of 83% versus adult ALCL which has an occurrence rate of 31%.^[1,2] The presence of t(2;5)(p23;q35) carries a better overall prognosis for ALCL patients.^[1]



Schematic representation of the *ALK/NPM1* DNA-FISH Probe:

Horizontal red and green bars indicate the regions covered by the probes (approximate to scale, GRCh37/Hg19/2009). The directly labeled *ALK* (red) & *NPM1* (green) probe flanks the respective genes.

Signal Interpretation

In normal diploid metaphase and interphase nucleus, two red and two green signals would be observed corresponding to the two normal homologous chromosomes 2 and 5, respectively (Figures 1 and 2). Upon translocation, the most commonly observed pattern is a single red and green signal, representing the normal chromosomes 2 and 5, and two fusion signals (red/green or yellow), which represent the translocated chromosomes (Figure 3). It is recommended to confirm variant pattern or atypical signal patterns by metaphase analysis whenever possible.

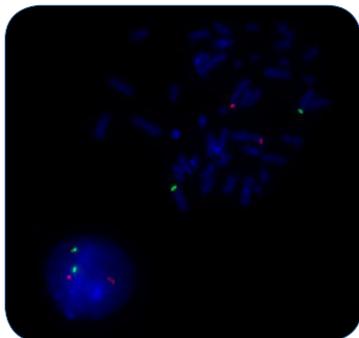


Figure 1: Normal diploid metaphase and interphase nucleus (from normal peripheral blood specimen) with 2 red (*ALK*) and 2 green (*NPM1*) signals.

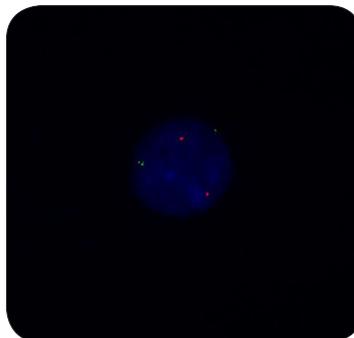


Figure 2: Normal diploid interphase nucleus (from bone marrow specimen) with 2 red (*ALK*) and 2 green (*NPM1*) signals.

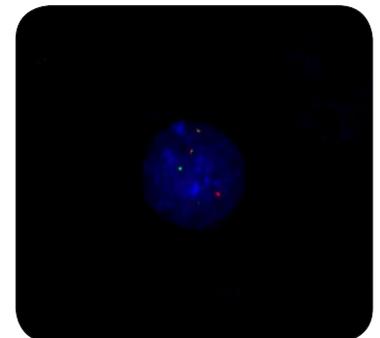


Figure 3: Interphase nucleus with 1 red (*ALK*), 1 green (*NPM1*), and 2 fusion (red/green or yellow) signals.

References

1. Drexler, H.G., et al. *Leukemia*, 2000. 14(9):p.1533-59.
2. Weitzman, S., et al. *Curr Oncol Rep*, 2002. 4(2):p.107-13.

Fluorescence Microscopy Filter Requirements

Fluorophore	Excitation _{max}	Emission _{max}
Green	496 nm	520 nm
Red	580 nm	603 nm
DAPI	360 nm	460 nm

Instructions for use are available at www.cancergeneticsitalia.com