



Sample Information

Patient Name: 劉國台  
Gender: Male  
ID No.: K120115468  
History No.: 38145787  
Age: 67  
  
Ordering Doctor: DOC1242E 劉峻宇  
Ordering REQ.: OCRVRHA  
Signing in Date: 2023/10/13

Path No.: M112-00265  
MP No.: F23074  
Assay: Oncomine Focus Assay  
Sample Type: FFPE  
Block No.: S112-84501A  
Percentage of tumor cells: 20%

Reporting Doctor: DOC5466K 葉奕成 (Phone: 8#5466)

Note:

Sample Cancer Type: Non-Small Cell Lung Cancer

<table><tr><td>Table of Contents</td><td>Page</td></tr><tr><td>Variant Details</td><td>2</td></tr><tr><td>Biomarker Descriptions</td><td>2</td></tr></table>	Table of Contents	Page	Variant Details	2	Biomarker Descriptions	2	<table><tr><td>Report Highlights</td></tr><tr><td>0 Relevant Biomarkers</td></tr><tr><td>0 Therapies Available</td></tr><tr><td>0 Clinical Trials</td></tr></table>	Report Highlights	0 Relevant Biomarkers	0 Therapies Available	0 Clinical Trials
Table of Contents	Page										
Variant Details	2										
Biomarker Descriptions	2										
Report Highlights											
0 Relevant Biomarkers											
0 Therapies Available											
0 Clinical Trials											

Relevant Non-Small Cell Lung Cancer Variants

Gene	Finding	Gene	Finding
ALK	None detected	NTRK1	None detected
BRAF	None detected	NTRK2	None detected
EGFR	None detected	NTRK3	None detected
ERBB2	None detected	RET	None detected
KRAS	None detected	ROS1	None detected
MET	None detected		

Relevant Biomarkers

No relevant biomarkers found in this sample.

Prevalent cancer biomarkers without relevant evidence based on included data sources

MYC amplification

## Variant Details

DNA Sequence Variants

Gene	Amino Acid Change	Coding	Variant ID	Locus	Allele Frequency	Transcript	Variant Effect	Coverage
JAK1	p.(P733=)	c.2199A>G	.	chr1:65310489	51.65%	NM_002227.4	synonymous	2000
ALK	p.(D1529E)	c.4587C>G	.	chr2:29416366	50.75%	NM_004304.5	missense	1998
ALK	p.(G1125=)	c.3375C>A	.	chr2:29445458	50.10%	NM_004304.5	synonymous	1998
PDGFRA	p.(G313=)	c.939T>G	.	chr4:55133726	52.91%	NM_006206.6	synonymous	1996
PDGFRA	p.(V824=)	c.2472C>T	.	chr4:55152040	54.48%	NM_006206.6	synonymous	1997
FGFR4	p.(P136L)	c.407C>T	.	chr5:176517797	99.65%	NM_213647.3	missense	2000

Copy Number Variations

Gene	Locus	Copy Number
MYC	chr8:128748885	30.6

## Biomarker Descriptions

### MYC (MYC proto-oncogene, bHLH transcription factor)

**Background:** The MYC gene encodes the MYC proto-oncogene (c-MYC), a basic helix-loop-helix transcription factor that regulates the expression of numerous genes that control cell cycle progression, apoptosis, metabolic pathways, and cellular transformation<sup>1,2,3,4</sup>. MYC is part of the MYC oncogene family that includes related transcription factors MYCN and MYCL that regulate transcription in 10-15% of promoter regions<sup>5</sup>. MYC functions as a heterodimer in complex with the transcription factor MAX<sup>2,6</sup>.

**Alterations and prevalence:** Recurrent somatic alterations are observed in both solid and hematological cancers. Recurrent somatic mutations in MYC, including codon T58, are infrequent and hypothesized to increase the stability of the MYC protein<sup>7,8</sup>. MYC gene amplification is particularly common in diverse solid tumors. MYC amplification is observed in 30% of serous ovarian cancer, 20% of uterine serous carcinoma, 15% of esophageal and breast cancers, and is common (1-10%) in numerous other cancer types<sup>9,10,11</sup>. MYC is the target of the t(8;14)(q24;32) chromosomal translocation in Burkitt's lymphoma that places MYC coding sequences adjacent to immunoglobulin region regulatory sequences, which results in increased MYC expression<sup>12,13</sup>.

**Potential relevance:** B-cell lymphoma with MYC translocations that co-occur with BCL2 or BCL6 are referred to as double hit lymphoma, while co-occurrence with BCL2 and BCL6 rearrangements is referred to as triple-hit lymphoma<sup>14,15</sup>. MYC translocations are also indicative of high risk for multiple myeloma and is associated with poor risk in acute lymphoblastic leukemia<sup>16,17</sup>. Currently, no therapies are approved for MYC aberrations. Due to the high frequency of somatic MYC alterations in cancer, many approaches are being investigated in clinical trials including strategies to disrupt complex formation with MAX, including inhibition of MYC expression and synthetic lethality associated with MYC overexpression<sup>1,18,19,20</sup>.

## Clinical Trials in Taiwan region:

## References

1. Chen et al. Targeting oncogenic Myc as a strategy for cancer treatment. *Signal Transduct Target Ther*. 2018 Feb 23;3:5. doi: 10.1038/s41392-018-0008-7. eCollection 2018. PMID: 29527331
2. Dang. MYC on the path to cancer. *Cell*. 2012 Mar 30;149(1):22-35. PMID: 22464321
3. Dominguez-Sola et al. Non-transcriptional control of DNA replication by c-Myc. *Nature*. 2007 Jul 26;448(7152):445-51. PMID: 17597761
4. Wahlström et al. Impact of MYC in regulation of tumor cell metabolism. *Biochim. Biophys. Acta*. 2015 May;1849(5):563-9. PMID: 25038584
5. Dang et al. The c-Myc target gene network. *Semin. Cancer Biol*. 2006 Aug;16(4):253-64. PMID: 16904903
6. Blackwood et al. Myc and Max function as a nucleoprotein complex. *Curr. Opin. Genet. Dev*. 1992 Apr;2(2):227-35. PMID: 1638116
7. Chakraborty et al. A common functional consequence of tumor-derived mutations within c-MYC. *Oncogene*. 2015 Apr 30;34(18):2406-9. PMID: 24998853
8. Xu-Monette et al. Clinical and Biologic Significance of MYC Genetic Mutations in De Novo Diffuse Large B-cell Lymphoma. *Clin. Cancer Res*. 2016 Jul 15;22(14):3593-605. PMID: 26927665
9. Kalkat et al. MYC Deregulation in Primary Human Cancers. *Genes (Basel)*. 2017 May 25;8(6). PMID: 28587062
10. Beroukhi et al. The landscape of somatic copy-number alteration across human cancers. *Nature*. 2010 Feb 18;463(7283):899-905. doi: 10.1038/nature08822. PMID: 20164920
11. Cerami et al. The cBio cancer genomics portal: an open platform for exploring multidimensional cancer genomics data. *Cancer Discov*. 2012 May;2(5):401-4. PMID: 22588877
12. Taub et al. Translocation of the c-myc gene into the immunoglobulin heavy chain locus in human Burkitt lymphoma and murine plasmacytoma cells. *Proc Natl Acad Sci U S A*. 1982 Dec;79(24):7837-41. PMID: 6818551
13. Ott et al. Understanding MYC-driven aggressive B-cell lymphomas: pathogenesis and classification. *Hematology Am Soc Hematol Educ Program*. 2013;2013:575-83. PMID: 24319234
14. NCCN Guidelines® - NCCN-B-Cell Lymphomas [Version 5.2023]
15. Beham-Schmid. Aggressive lymphoma 2016: revision of the WHO classification. *Memo*. 2017;10(4):248-254. PMID: 29250206
16. NCCN Guidelines® - NCCN-Multiple Myeloma [Version 3.2023]
17. NCCN Guidelines® - NCCN-Acute Lymphoblastic Leukemia [Version 2.2023]
18. Posternak et al. Strategically targeting MYC in cancer. *F1000Res*. 2016;5. PMID: 27081479
19. Carabet et al. Therapeutic Inhibition of Myc in Cancer. *Structural Bases and Computer-Aided Drug Discovery Approaches*. *Int J Mol Sci*. 2018 Dec 29;20(1). PMID: 30597997
20. Shahbazi et al. The Bromodomain Inhibitor JQ1 and the Histone Deacetylase Inhibitor Panobinostat Synergistically Reduce N-Myc Expression and Induce Anticancer Effects. *Clin. Cancer Res*. 2016 May 15;22(10):2534-44. PMID: 26733615