Curie-LC: copy number alteration - Affymetrix Genome-Wide Human SNP Array 6.0, WGS

Molecular Methods Description:

PDX were profiled using Affymetrix genomics array with SNP 6.0 or with Cytoscan HD array. Genome-wide copy number analysis was conducted by means of Affymetrix SNP arrays, as previously described (Laurent et al., 2013 and Crepin et al. 2017).

500 and 250ng of gDNA was respectively used for SNP 6.0 and Cytoscan HD as starting material as recommended by the supplier.

Analysis Description:

Raw data were normalized with Genotyping console for SNP 6.0 and Chromosome Analysis Suite for Cytoscan HD.

Focal amplification of oncogenes was defined by log ratio>1.58 (6 copies per diploid genome) and maximum size <10 megabases. Biallelic inactivation of tumor suppressor genes was defined by homozygous deletion or truncating mutation associated with heterozygous deletion.

All PDX copy numbers were represented by the Circular Binary Segmentation algorithm as implemented in the DNAcopy package for R using a minimum width of 3, alpha less than 0.01 and up to 10,000 permutations.

The BRCAness signature was defined with large-scale state transitions, defined as chromosomal break between adjacent regions of at least 10Mb initially described by T. Popova **

Table:

Gene List				
AKT1	AKT2	AKT3	ALK	
APC	ARHGAP35	ARID1A	ARID2	ATM
BRAF	BRCA1	BRCA2	CDK12	CDKN1B
CDKN2A	CTNNB1	DDR2	EGFR	ESR1
FAT1	FBXO16	FBXW7	FGFR1	FGFR2
FGFR3	FRFR4	GENE LIST	H3F3A	HER2/Erbb2
HER3/Erbb3	HER4/Erbb4	HRAS	IDH1	IDH2
KDR	KEAP1	KIT	KRAS	MAP2K1
MAP2K2	MAP2K4	MAP3K1	MBD1	MEK1
MET	MLL3	MTOR	NBPF1	NF1
NFE2L2	NRAS	PALB2	PBLD	PDGFRA
PIK3CA	PIK3R1	POLE	PPP2R1A	PTEN
PTPN11	RAD51B	RAD51C	RB1	RBM10
RET	RIT1	ROS1	SLC39A6	SLC4A5

SMAD4	SMARCA4	STK11	STX2	TP53
TSC1	TSC2	U2AF1	VHL	ZNF774