Why (irculogene?

CIRCULOGENE's comprehensive tumor DNA and RNA sequencing is the only plasma testing available that combines the most advanced next-generation sequencing (NGS) and polymerase chain reaction (PCR) technology to detect and monitor cfDNA and cfRNA within well-characterized, well-documented, actionable cancer-associated genes.

SOMATIC RNA FUSION NGS						
ABL1	CCNB3	FGFR1	MYB	POU5F1	STAT6	
АСТВ	CCND1	FGFR2	MYC	PPARGC1A	STRN	
AFAP1	CD74	FGFR3	NAB2	PPP1CB	SUZ12	
AGK	CIC	FLI1	NCOA1	PRKACA	TACC1	
AKAP12	CLTC	FN1	NCOA2	PRKAR1A	TACC3	
AKAP4	CNTRL	FOXO1	NCOA4	PTPRZ1	TAF15	
AKAP9	COL1A1	FOXO4	NFIB	QKI	TCF12	
AKT2	CREB1	FUS	NOTCH2	RAF1	TERT	
AKT3	CREB3L1	GLI1	NPM1	RANBP2	TFE3	
ALK	CREB3L2	GOPC	NR4A3	RARA	TFG	
ASPSCR1	CRTC1	GPR128	NRG1	RELA	THADA	
ATF1	DDIT3	HMGA2	NRG2	RELCH	TMPRSS2	
ATP1B1	DNAJBI	JAZF1	NSD3	RET	ТРМЗ	
ATRX	EGFR	KIAA1549	NTRK1	ROS1	TPR	
BAG4	EML4	KIF5B	NTRK2	RREB1	TRIM24	
BCL2	EPC1	LMNA	NTRK3	RSP02	TRIM33	
BCOR	ERBB2	LPP	NUTM1	RSP03	TRIO	
BCORL1	ERBB4	MAGI3	PAX3	SDC1	VGLL2	
BCR	ERG	MAML1	PAX7	SDC4	WT1	
BICC1	ESR1	MAML2	PAX8	SHTN1	WWTR1	
BRAF	ETV1	MAML3	PDGFB	SLC34A2	YAP1	
BRD3	ETV4	MET	PDGFRA	SND1	YWHAE	
BRD4	ETV5	MGA	PDGFRB	SQSTM1	ZMYM2	
CAMTA1	ETV6	MGMT	PHF1	SS18	ZNF703	
CCAR2	EWSR1	MIR143	PIK3CA	SSX1	ZFTA	
CCDC6	EZR	MITF	PLAG1	SSX2		
CCDC88A	FEV	MKL2	PML	SSX4		

#### SOMATIC DNA SEQUENCING

	FULL GENE					
AKT1	CDH1	EZH2	JAK3	NRAS	SETD2	
ALK	CDK4	FBXW7	KDR	NTRK1	SMAD4	
AR	CDK12	FGFR1	KEAP1	NTRK2	SMARCA4	
ARAF	CDK6	FGFR2	КІТ	NTRK3	SMARCB1	
ARID1A	CDKN2A	FOXL2	KRAS	PALB2	SMO	
ATM	CHEK1	FGFR3	MAP2K1	PDGFRA	SRC	
ATR	CHEK2	GNA11	MAP2K2	PIK3CA	STAT3	
AXL	CRKL	GNAQ	МАРКЗ	POLD1	STK11	
BAP1	CSF1R	GNAS	MET (Incl.	POLE	TERT	
BARD1	CTNNB1	HNF1A	Exon 14 Skipping)	PTEN	TOP1	
BRAF	DDR2	HRAS	MLH1	PTPN11	TP53	
BRCA1	EGFR	IDH1	MTOR	RAF1	TSC1	
BRCA2	ERBB2	IDH2	MYC	RB1	TSC2	
CCND1	ERBB4	IGF1R	NF1	RET	VHL	
CCNE1	ESR1	JAK2	NOTCH1	ROS1		

CNV					
AR	CDK4	ERBB2	КІТ		
CCND1	CDK6	FGFR1	MET		
CCNE1	EGFR	FGFR2	МҮС		

FUSION BY RT-PCR				
ALK	NTRK2	RET		
NTRK1	NTRK3	ROS1		

IMMUNOTHERAPY				
MSI	PD-L1 RNA Expression			

HEREDITARY GENES							
	APC	CDH1	FLCN	NBN	RAD51C	TP53	
	ATM	CDK4	HOXB13	NF1	RAD51D	TSC1	
	AXIN2	CDKN2A	MET	NTHL1	RECQL	TSC2	
	BAP1	CHEK2	MITF	PALB2	SCG5	VHL	
	BARD1	CTNNA1	MLH1	PMS2	SDHB		
	BMPR1A	EPCAM	MSH2	POLD1	SDHC		
	BRCA1	FANCC	MSH3	POLE	SDHD		
	BRCA2	FANCM	MSH6	POT1	SMAD4		
	BRIP1	FH	MUTYH	PTEN	STK11		

# When to Test with Liquid Biopsy

CIRCULOGENE offers the most advanced NGS and PCR methods to both detect and continually monitor cfDNA and cfRNA.





Talk to your **CIRCULOGENE** representative to request our collection kit and requisition form.



EMPOWERING relision

WWW.CIRCULOGENE.COM INFO@CIRCULOGENE.COM 855-614-7083



LIQUID PD-L1 NCCN GUIDELINES PCR + NGS ALL STATES

# Comprehensive Molecular Testing for Lung Cancer

GETTING LUNG CANCER PATIENTS ON THE RIGHT TREATMENT FASTER

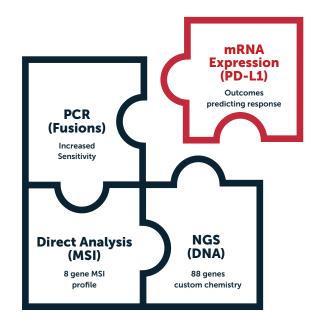
### The Circulogene Best-in-class

CIRCULOGENE combines the best-in-class preanalytical, instrumentation, and technology to provide oncologists {and their patients} with the most comprehensive results. Having results in as little as 7 days could lead to personalized treatment, faster.

- Combines NGS and PCR
- Increases RNA yields with proprietary LISA technology
- Provides the ONLY liquid PD-L1
- Offers Enhanced MSI
- Clinical utility to detect at all cancer stages
- Early indication of metastasis

#### INTERNATIONAL ASSOCIATION FOR THE STUDY OF LUNG CANCER (IASLC) RECOMMENDATION:

Liquid biopsy is emerging as not only complementary to tissue-based analysis but also acceptable as the initial approach ("plasma first") for biomarker evaluation at the time of diagnosis and for monitoring the efficacy of targeted therapies.



#### 80% OF CANCER PATIENTS START TREATMENT WITHOUT NUCLEAR PROFILE

ASCO (2019) reported that only 20% of NSCLC patients have a molecular profile or biopsy to maximize information gathered for optimal treatment decisions.

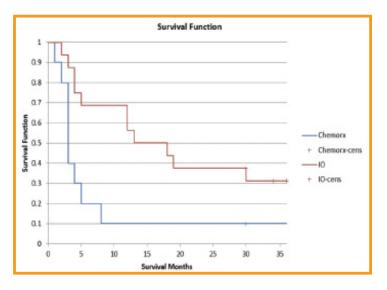


#### CIRCULOGENE OFFERS THE ONLY LIQUID PD-L1

PD-L1 expression has been correlated with clinical effectiveness and improved survival rates when treated with immunotherapy. Utilizing KEYNOTE-042 as a baseline comparison, plasma PD-L1 parallels tissue PD-L1 clinical trial outcomes with a 30% survival over 3 years.

Study demonstrates plasma cfRNA PD-L1 is predictive of immunotherapy benefit in advanced NSCLC of (compared to chemotherapy).

The ECU study demonstrated that Plasma PD-L1 expression was predictive of significant survival benefit of immunotherapy treatment over chemotherapy in advanced NSCLC patients. Using pembrolizumab monotherapy study as a baseline comparison, plasma PD-L1 parallels tissue PD-L1 clinical trial outcomes with a 30% survival over 3 years.



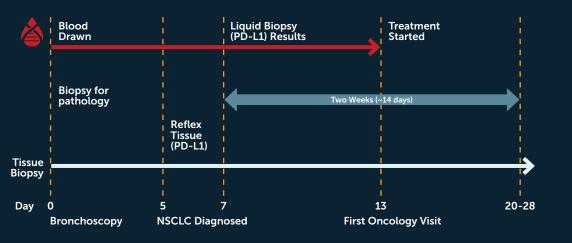
## Patients deserve the Best!

Each patient's cancer is unique, understanding the origins and severity of the cancers aids oncologists to personalize the patient's treatment plan at the time of diagnosis and throughout their cancer journey.

Patients deserve the right treatment for their cancer and within a time frame that improves their outcomes and chance for survival. Understanding the patient's tumor(s) molecular profile aids oncologists to prescribe the right first-line treatment.

#### LIQUID v. TISSUE TIME TO TREATMENT (NSCLC)

AS QUICK AS 7 DAYS TO RESULTS - FASTER TIME TO TREATMENT



# Combination of Liquid and Tissue Biopsy

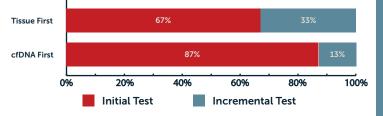
#### THE COMBINATION OF LIQUID AND TISSUE BIOPSY IDENTIFIES MORE ACTIONABLE MUTATIONS

Tissue-only molecular testing only detects 67% of National Comprehensive Cancer Network (NCCN) guideline mutations, missing 33%. Liquid biopsy detects 87% of guideline mutations.

Used together, liquid and tissue biopsy provides a more complete picture of the tumor's molecular makeup.

Liquid biopsy has the advantage of detecting a multitude of biomarkers which may be an indication of metastasis away from the primary tumor.





#### COMBINING PCR AND NGS LEADS TO MOST COMPREHENSIVE ASSAY

Highly sensitive RNA fusions are best detected by PCR versus NGS. CIRCULOGENE's best-in-class approach provides a multimodality approach combining the breadth of NGS for DNA and accuracy of PCR for RNA.

The fusion detection sensitivities of the cfRNA with PCR and cfDNA with NGS were compared. cfRNA with PCR detected 78% (7/9) of the fusions while cfDNA with NGS only detected 33% (3/9).

