

¹Supplementary information

A Phase II exploratory study to identify biomarkers predictive of clinical response to regorafenib in patients with metastatic colorectal cancer who have failed first-line therapy.

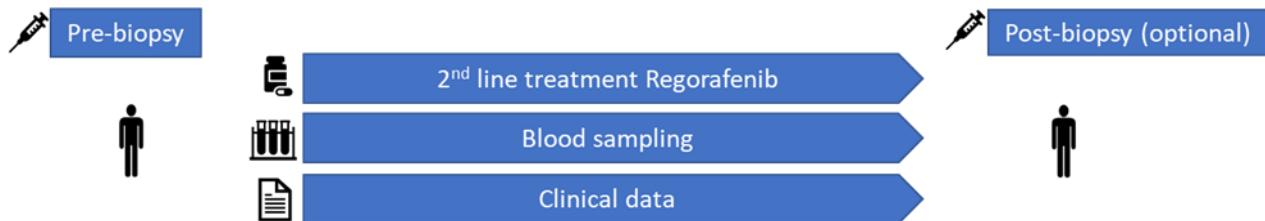


Figure S1. Overview of Study Design

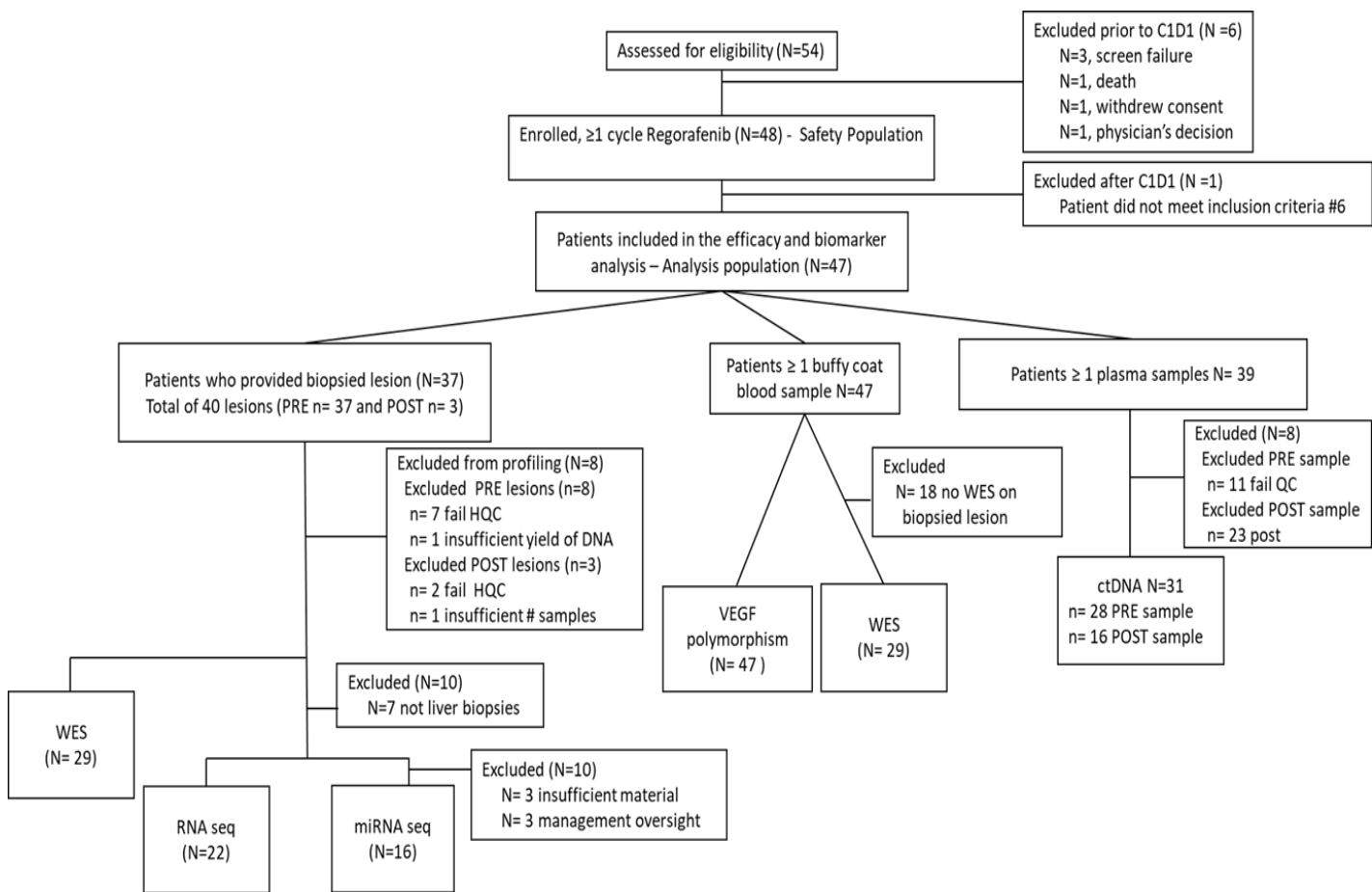


Figure S2. Consort Diagram of Subject Disposition. The number of patients is presented using N and the number of lesions is presented using n. C1D1, Cycle 1 Day 1; HQC, Histopathology Quality Control; WES, Whole Exome Sequencing, ctDNA, Circulating tumour DNA; Seq, Sequencing.

S1: Objective Response of Biopsied Lesions

At each radiological evaluation, the site staff was to capture the size of the lesion sampled at the time of the pre-treatment biopsy collection in order to follow its response to treatment. RECIST v. 1.1 guidelines were used to assess the response of the biopsied lesion as follows:

- PR: a lesion with $\geq 30\%$ decrease in the longest diameter compared to baseline
- SD:
 - a lesion with $< 30\%$ decrease in the longest diameter compared to baseline, or
 - a lesion with $< 20\%$ increase in comparison to the smallest diameter, or
 - a lesion with $\geq 20\%$ and $< 5\text{ mm}$ increase in the longest diameter in comparison to the smallest diameter.
- PD: a lesion with $\geq 20\%$ and $\geq 5\text{ mm}$ increase in the longest diameter in comparison to the smallest diameter.

Lesions were subsequently divided, when applicable, into two subcategories intrinsic resistant (IRES) lesions and acquired resistant (ARES) lesions which were defined as follows:

- IRES: a new lesion or a lesion with $\geq 20\%$ and minimally by $\geq 5\text{ mm}$ increase in the longest diameter at the first 8-week evaluation,
- ARES: a lesion with a PR or SD response followed by a PD response. When pre- and post-biopsies were from different liver segments, the response of the lesion sampled was used.

S2: Inclusion Criteria

Patients were to meet these inclusion criteria to be enrolled in this study:

- (1) Signed informed consent obtained before any study-specific procedures. Patients must have been able to understand and be willing to sign a written informed consent.
- (2) Male or female patients ≥ 18 years of age.
- (3) Histological documentation of adenocarcinoma of the colon or rectum, with at least one liver metastatic site available for biopsy (other metastatic sites may have been considered but must have been first approved by the Sponsor-Investigator).
- (4) Metastatic disease not suitable for upfront curative-intent surgery.
- (5) Patients must have received one (and no more than one) prior treatment regimen for metastatic CRC (i.e. FOLFOX, FOLFIRI or XELOX, with/without bevacizumab). Patients who had withdrawn from standard first-line treatment due to unacceptable toxicity before progression of disease were to be allowed into the study
- (6) Measurable disease according to RECIST v. 1.1.
- (7) ECOG status of ≤ 1 .
- (8) Life expectancy of at least 3 months.
- (9) Women of childbearing potential and men must have agreed to use adequate contraception since signing of the informed consent form until at least 3 months after the last study drug administration. The investigator or a designated associate was requested to advise the patient on how to achieve adequate birth control. Adequate contraception was defined in the study as any medically recommended method (or combination of methods) as per the standard of care.

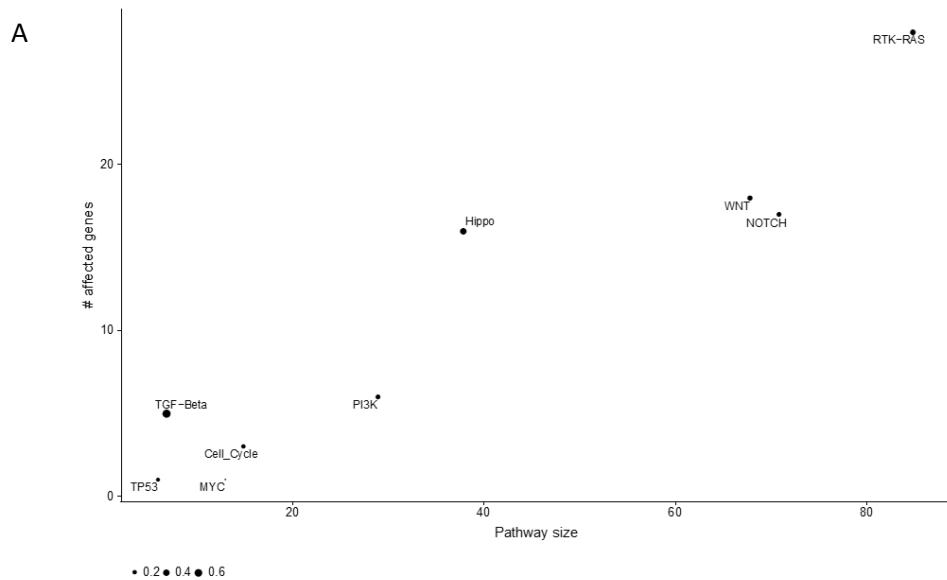
- (10) Adequate bone marrow, liver, and renal function as assessed by the following laboratory requirements conducted within 7 days of starting to study treatment:
- Total bilirubin $\leq 1.5 \times$ the upper limit of normal (ULN)
 - Alanine aminotransferase (ALT) and aspartate aminotransferase (AST) $\leq 2 \times$ ULN ($\leq 5 \times$ ULN for patients with liver involvement of their cancer)
 - Alkaline phosphatase limit $\leq 2.5 \times$ ULN ($\leq 5 \times$ ULN for patients with liver involvement of their cancer) • Amylase and lipase $\leq 1.5 \times$ ULN
 - Serum creatinine $\leq 1.5 \times$ ULN
 - International normalized ratio (INR) and partial thromboplastin time (PTT) $\leq 1.5 \times$ ULN.
 - Platelet count $\geq 100 \times 10^9 /L$, hemoglobin $\geq 90 \text{ g/L}$, absolute neutrophil count (ANC) $> 1.5 \times 10^9 /L$.
- 4.4.2.

Exclusion Criteria

Patients were to be excluded from this study if they met any of the following criteria:

- (1) Previous treatment with regorafenib.
- (2) Previous or concurrent cancer that was distinct in primary site or histology from colorectal cancer within 5 years before randomization, EXCEPT for curatively treated cervical cancer in situ, non-melanoma skin cancer and superficial bladder tumours.
- (3) Extended field radiotherapy within 4 weeks or limited field radiotherapy within 2 weeks prior to registration. Patients had to have recovered from all therapy-related toxicities. The site of previous therapy was to have evidence of progressive disease if this was the only site of disease.
- (4) Major surgical procedure or significant traumatic injury within 28 days before start of study treatment (day of first dose).
- (5) Pregnancy or breast-feeding.
- (6) Congestive heart failure \geq New York Heart Association (NYHA) class 2.
- (7) Unstable angina (angina symptoms at rest), new-onset angina (began within the last 3 months). Myocardial infarction less than 6 months before start of study treatment (day of first dose).
- (8) Cardiac arrhythmias requiring anti-arrhythmic therapy (beta blockers or digoxin were permitted).
- (9) Uncontrolled hypertension (systolic blood pressure $> 150 \text{ mmHg}$ or diastolic pressure $> 90 \text{ mmHg}$ despite optimal medical management).
- (10) Phaeochromocytoma.
- (11) Pleural effusion or ascites that caused respiratory compromise (\geq Grade 2 dyspnea per Common Terminology Criteria for Adverse Events [CTCAE] v. 4.0 or higher).
- (12) Arterial or venous thrombotic or embolic events such as cerebrovascular accident (including transient ischemic attacks), deep vein thrombosis or pulmonary embolism within the 3 months before start of study treatment (day of first dose).
- (13) Ongoing uncontrolled infection $>$ Grade 2 per CTCAE.
- (14) Known history of human immunodeficiency virus (HIV) infection.
- (15) Active hepatitis B or C, or chronic hepatitis B or C requiring antiviral therapy.
- (16) Seizure disorder requiring medication.
- (17) Any history of or currently known brain metastases (patients with stable brain metastases ≥ 3 months may be eligible for the study).
- (18) History of organ allograft.
- (19) Evidence or history of severe bleeding diathesis.
- (20) Non-healing wound, ulcer, or bone fracture.

- (21) Renal failure requiring hemodialysis, peritoneal dialysis or baseline GFR < 30 mL/min/1.73m² .
- (22) Dehydration ≥ Grade 2, as per CTCAE.
- (23) Substance abuse or medical, psychological, or social conditions that could have interfered with the patient's participation in the study or evaluation of the study results.
- (24) Known hypersensitivity to regorafenib, regorafenib class of drugs, or excipients in the formulation.
- (25) Any illness or medical conditions that were unstable or could have jeopardized the safety of the patient and his or her compliance in the study in the opinion of the investigator.
- (26) Interstitial lung disease with ongoing signs and symptoms at the time of informed consent.
- (27) Persistent proteinuria ≥ Grade 3 per CTCAE (i.e. >3.5 g/24 hours).
- (28) Inability to swallow oral medications.
- (29) Any malabsorption condition.
- (30) Unresolved toxicity ≥ Grade 1, attributed to any prior therapy/procedure, excluding alopecia and oxaliplatin neurotoxicity ≤ Grade 2 (CTCAE).



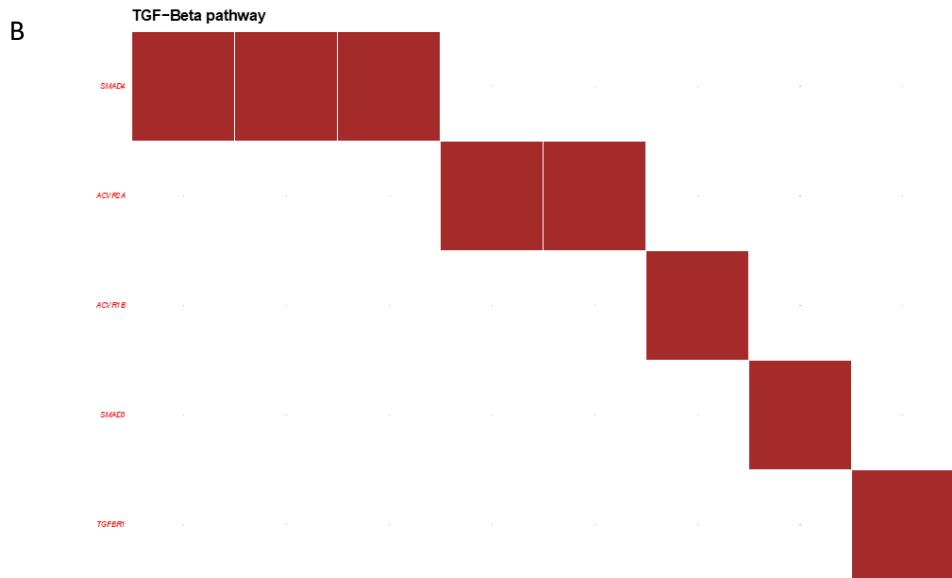
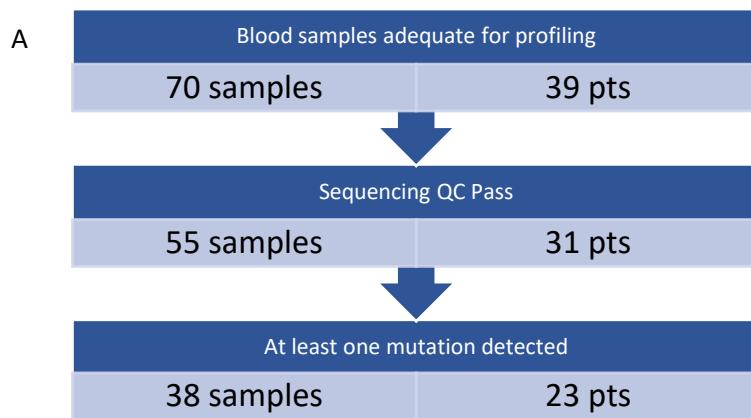


Figure S3. Visualization of mutations present in metastatic samples of the cohort by oncogenic pathways. A: The x-axis shows the number of genes in the pathways. The y-axis shows the number of genes altered in the present cohort. The size of the dot represents the proportion of genes mutated in Q-CROC-06 cohort per the total number of genes in the pathway (N=29). B. Visualization of the TGF- β pathways. Each column represents a patient and each row a gene in the TGF- β pathway. A red square is present when the gene is mutated in the patient sample.



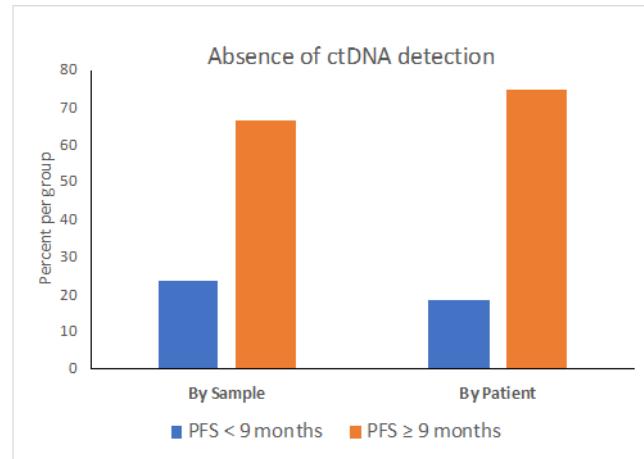


Figure S4. Cell-free DNA analysis.

- A. Attrition flowchart for cfDNA profiling assay; B. Proportion of samples and patients with no ctDNA detected within each group defined using a PFS cutoff of 9 months.

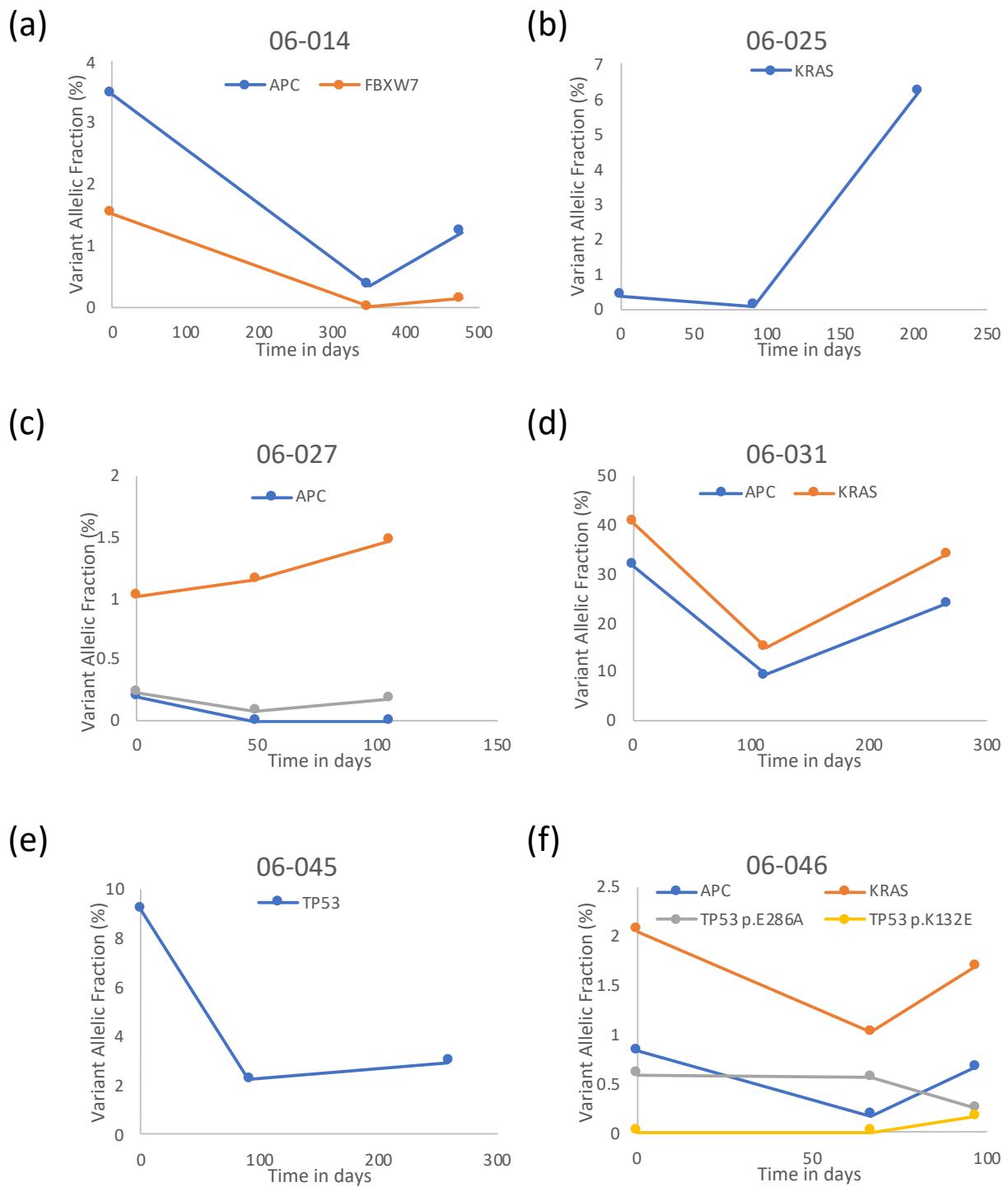


Figure S5. Variant allelic fraction dynamics during the course of treatment

a) patient 06-14, b) patient 06-025, c) patient 06-027, d) patient 06-031, e) patient 06-045, and f) patient 06-046. On the x-axis is the time in days and on the y-axis is the VAF. Each line represents a specific mutation detected as

shown in each patient legend. The first point represents time at baseline, the second time at response and the third, time of progression.

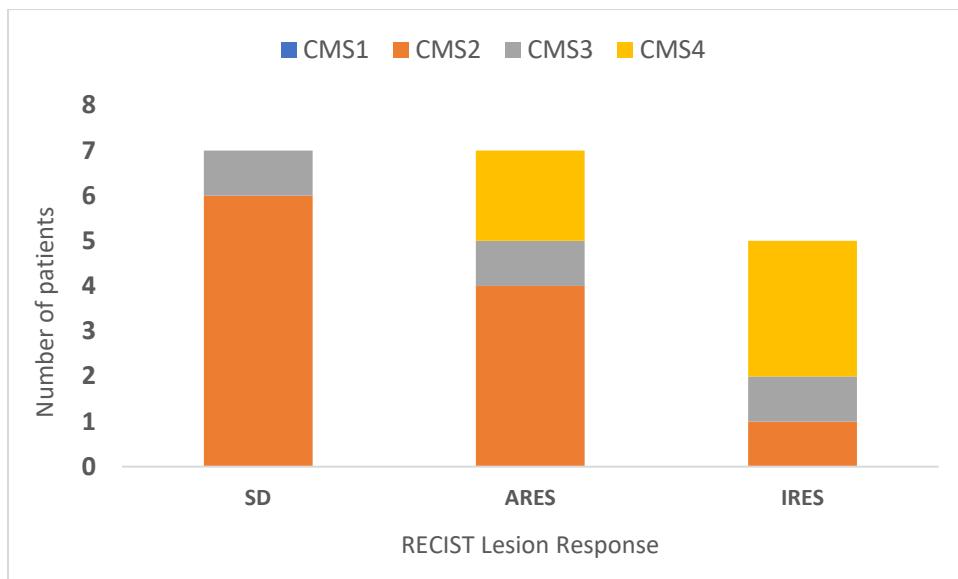


Figure S6: Distribution of Consensus Molecular Subtypes (CMS) within RECIST lesion Response Groups.

Table S1: Reasons for treatment discontinuation

Category	Overall
Dosed patients, N	48
Patients withdrawn due to progression of disease, n (%)	34 (70.8%)
Occurrence of an unacceptable AE/toxicity, n (%)	10 (20.8%)
Patients withdrawn due to clinical progression, n (%)	1 (2.1%)
Patients request (withdrew consent), n (%)	1 (2.1%)
Investigator decision, n (%)	1 (2.1%)
Other, n (%)	1 (2.1%)

Table S2: Summary of biopsied lesion and lesion response assessment

LESION ASSESSMENT	ANALYSIS POPULATION N=47	
	N or n ¹	%
# Patients with > 1 biopsy sample	N=47	
Yes	37	78.7%
No	10	21.3%

# PRE and POST Biopsied Lesions ²	n=40	
prior to regorafenib initiation	37	92.5%
after regorafenib discontinuation ³	3	7.5%
PRE Biopsied Lesion Location²	n=37	
Liver	27	73.0%
Other	9	24.3%
Unknown	1	2.7%
PRE Biopsied Lesion Response ^{3,4}	n=31	
SD	12	38.7%
ARES	11	35.5%
IRES	8	25.8%
N/A	6	N/A

¹ N refers to the number of patients and n refers to the number of biopsy samples

² Calculated from the total number of biopsied lesions (n=40)

³ Pre- and post-regorafenib biopsies of the same patient (n=3) are from different lesions. Post regorafenib biopsies responses were IRES, ARES, and SD and were not used for translational analysis

⁴ Calculated from the total number of pre-biopsied lesions (n=37)

⁵ Calculated from the total number of pre-biopsied lesions with an evaluable response (n=31)

Table S3. Summary of Treatment-Emergent Adverse Events by SOC Experienced by ≥10% of patients.

System Organ Class	Safety Population (N=48)
CTCAE Term	n (%)
Patients with at least one TEAE	48 (100 %)
Gastrointestinal Disorders	38 (79.2 %)
Diarrhea	21 (43.8 %)
Abdominal pain	16 (33.3 %)
Nausea	10 (20.8 %)
Vomiting	10 (20.8 %)
Mucositis oral	8 (16.7 %)
Constipation	7 (14.6 %)
Rectal hemorrhage	6 (12.5 %)
General disorders and administration site conditions	36 (75.0 %)
Fatigue	26 (54.2 %)
Fever	15 (31.3 %)
Chills	10 (20.8 %)

System Organ Class	Safety Population (N=48)
CTCAE Term	n (%)
Skin and subcutaneous tissue disorders	32 (66.7 %)
Palmar-plantar erythrodysesthesia syndrome	19 (39.6 %)
Rash maculo-papular	7 (14.6 %)
Alopecia	6 (12.5 %)
Dry skin	5 (10.4 %)
Pruritus	5 (10.4 %)
Other	5 (10.4 %)
Musculoskeletal and connective tissue disorders	28 (58.3 %)
Arthralgia	11 (22.9 %)
Back Pain	9 (18.8 %)
Pain in extremity	9 (18.8 %)
Myalgia	8 (16.7 %)
Metabolism and nutrition disorders	25 (52.1 %)
Anorexia	12 (25.0 %)
Hypophosphatemia	10 (20.8 %)
Dehydration	5 (10.4 %)
Respiratory, thoracic and mediastinal disorders	25 (52.1 %)
Hoarseness	12 (25.0 %)
Dyspnea	8 (16.7 %)
Epistaxis	6 (12.5 %)
Cough	5 (10.4 %)
Investigations	24 (50.0 %)
Blood bilirubin increased	8 (16.7 %)

System Organ Class	Safety Population (N=48)
CTCAE Term	n (%)
ALT increased	6 (12.5 %)
AST increased	6 (12.5 %)
Lipase increased	5 (10.4 %)
Platelet count decreased	5 (10.4 %)
Infections	21 (43.8%)
Urinary tract infection	8 (16.7 %)
Nervous system disorders	21 (43.8 %)
Headache	10 (20.8 %)
Paresthesia	7 (14.6 %)
Peripheral motor neuropathy	5 (10.4 %)
Vascular disorders	19 (39.6 %)
Hypertension	17 (35.4 %)
Renal and urinary disorders	11 (22.9 %)
Proteinuria	8 (16.7 %)

Table S4: Incidence of Treatment-Emergent Adverse Events by Maximum Severity by Patient

Parameter	Safety Population
	N = 48
	n (%)
Patients with at least one TEAE¹	47 (97.9 %)
Grade 1	2 (4.2%)
Grade 2	10 (20.8%)
Grade 3	29 (60.4 %)
Grade 4	7 (14.6%)
Grade 5	0

¹Each patient is counted once for the maximum severity experienced (any TEAE), unknown TEAEs were not included.

Table S5: Summary of Treatment-Emergent Serious Adverse Event.

Parameter	Safety Population N = 48
Serious Adverse Events reported (n)	19
Treatment-emergent serious adverse events (n)	19
Patients with at least one biopsy-related serious adverse event (n[%])	0 (0 %)
Treatment-emergent serious adverse events reported (n)	19
Patients with at least one TESAE (n[%])	16 (33.3 %)
Patients with at least one drug-related TESAE (n[%]) ¹	6 (12.5 %)
Patients with at least one TESAE leading to drug withdrawal, dose reduction or interruption (n[%])	16 (33.3 %)
Patients with at least one TESAE leading to dose interruption ¹ (n[%])	9 (18.8%)
Patients with at least one TESAE leading to drug withdrawal ¹ (n[%])	8 (16.7%)
Patients with at least one TESAE leading to dose reduction ¹ (n[%])	1 (2.1%)

¹Patients could have been included in more than one category.

Table S6: Distribution of Patients' Time of Death After Treatment Discontinuation (Safety Population N=48)

Months elapsed since regorafenib's last dose	Patients ¹ n (%)
< 1 month	3 (6.3%)
≥ 1 and < 3 months	6 (12.5%)
≥ 3 and < 6 months	5 (10.4%)
≥ 6 and < 12 months	1 (2.1%)
≥ 12 months	7 (14.6%)

¹For some patients, death was reported independently of study discontinuation.

Table S7: Univariate and multivariable analysis using PFS as outcome.

	N	UNIVARIATE		MULTIVARIABLE	
		HR (95% CI)	P	HR (95% CI)	P
Age					
< 65y	21	1			
≥ 65y	26	1.1 (0.58 – 2.2)	0.72		
Sex					
Female	15	1			
Male	32	1.2 (0.57 – 2.6)	0.6		
BMI					
< 30	30	1			
≥ 30	17	0.54 (0.27-1.1)	0.073		
ECOG					
0	12	1			
1	31	1.1 (0.54 – 2.4)	0.72		
Primary Resected					
No	9	1			
Yes	38	0.28 (0.12 – 0.67)	0.0042*	0.5 (0.19 – 1.3)	0.158
Sidedness					
Left	13	1			
Right	34	1.3 (0.48 – 3.5)	0.61		
Number of metastatic Sites					
< 3	31	1			
3 +	16	1.9 (0.97 – 3.8)	0.062		
Liver Metastases					
No	12	1			
Yes	35	3.1 (1.2 – 8.1)	0.023*	4.0 (1.40 – 11.7)	0.01*
Lung Metastases					
No	23	1			
Yes	24	1.2 (0.63 – 2.4)	0.55		
Peritoneum Metastases					
No	37	1			
Yes	10	1 (0.46 – 2.2)	0.98		
Other Metastatic Site					
No	24	1			
Yes	23	1.2 (0.63 – 2.3)	0.58		
Bevacizumab					
No	23	1			
Yes	22	2.2 (1.1 – 4.3)	0.023*	2.8 (1.26 – 6.1)	0.011*

* P ≤ 0.05

HR: Hazard Ratio

CI: Confidence interval

Table S8: Summary of the genotype and univariate analysis performed on 47 patients.

Gene	SNP	Genotype	# Patient	%	Univariate	HR (95% CI)	P
VEGFA	rs25648	TT	2	4.1	CC vs TT+TC	0.93 (0.47-1.9)	0.85
		CT	15	30.6			
		CC	30	61.2			
VEGFA	rs833061	CC	10	20.4	TT+TC vs CC	1.2 (0.54-2.6)	0.67
		CT	25	51.0			
		TT	12	24.5			
VEGFA	rs699947	AA	10	20.4	CC vs AA+AC	0.93 (0.44-2)	0.86
		AC	25	51.0			
		CC	12	24.5			
VEGFA	rs2010963	CC	4	8.2	CC vs GG+GC	1.4 (0.41-4.7)	0.59
		CG	20	40.8			
		GG	23	46.9			
VEGFC	rs4604006	CC	32	65.3	TT+TC vs CC	0.99 (0.48-2.1)	0.99
		CT	15	30.6			
		TT	0	0.0			
FLT1	rs664393	CC	38	77.6	CC+CT vs TT	0.51 (0.12-2.2)	0.36
		CT	7	14.3			
		TT	2	4.1			
VEGFR2	rs2071559	AA	18	36.7	GG+AG vs AA	0.9 (0.44-1.8)	0.76
		AG	17	34.7			
		GG	12	24.5			
KDR	rs2305948	CC	41	83.7	CC vs CT+TT	1.6 (0.61-4.2)	0.34
		CT	6	12.2			
		TT	0	0.0			

Table S9. GISTIC analysis showing region focally gained or lost in 29 CRC metastatic lesions.

Only significant regions with Q-bound (FDR adjusted p-value) value ≤ 0.05 are shown. G-score: measure of the frequency of occurrence of the aberration and the magnitude of the copy number change (log ratio intensity) at each location in the aggregate of all samples in the data set.

Region	Extended Region	Type	Q-Bound	G-Score
chr12:3,603,686-4,155,279	chr12:121,792-5,153,110	CN Gain	0.00824022	8.05155565
chr13:32,402,096-32,750,036	chr13:25,264,913-98,600,139	CN Gain	1.69E-07	14.4018502
chr20:25,365,450-25,657,629	chr20:24,163,644-25,893,207	CN Gain	7.26E-05	11.0571811
chr20:34,205,363-34,740,875	chr20:29,963,172-45,811,961	CN Gain	9.44E-11	17.9885341
chr7:130,526,248-131,233,775	chr7:127,484,260-142,041,334	CN Gain	0.01169375	7.82692424
chr7:77,471,344-78,075,687	chr7:63,518,434-82,578,860	CN Gain	0.00527846	8.35501357
chr7:27,285,501-27,449,776	chr7:9,289,724-38,261,159	CN Gain	1.09E-04	10.8448921
chr8:43,033,260-43,057,692	chr8:41,753,513-43,665,344	CN Gain	3.64E-04	10.0897836
chr8:85,861,666-86,351,201	chr8:79,174,768-103,918,616	CN Gain	1.99E-06	13.1085476
chr1:57,893,600-58,948,410	chr1:42,628,447-94,578,484	CN Loss	0.00413668	8.53380572
chr10:89,728,892-90,122,400	chr10:89,631,252-93,579,279	CN Loss	8.71E-04	9.44427241
chr14:20,201,425-20,219,310	chr14:19,123,663-20,586,298	CN Loss	0.00166615	9.09233989
chr17:11,701,132-11,900,314	chr17:2,966,482-18,292,633	CN Loss	1.68E-05	11.5199467
chr18:13,491,557-13,694,281	chr18:2,537,503-15,270,970	CN Loss	1.00E-04	10.5997541
chr18:48,474,120-48,584,469	chr18:44,336,731-50,686,159	CN Loss	4.96E-13	20.2771852
chr22:16,346,117-16,449,729	chr22:16,171,502-16,688,967	CN Loss	0.00142717	9.17590884
chr4:134,111,761-138,440,204	chr4:100,851,518-158,257,339	CN Loss	0.01710582	7.52708145
chr5:69,817,965-70,070,758	chr5:69,521,493-70,671,993	CN Loss	0.01335331	7.71814656
chr8:11,987,652-11,990,735	chr8:0-21,705,632	CN Loss	5.45E-08	15.1886766
chr9:21,623,432-22,169,612	chr9:13,945,142-22,370,932	CN Loss	0.02920405	7.05118589

Table S10: Copy number variation significantly associated with progression-free survival in 29 CRC metastatic lesions.

Displaying only significant regions with a permuted p-value ≤ 0.05 .

Region	Event	Length	Number of samples with the event	Median PFS (days) with the event	Number of samples without the event	Median PFS (days) without the event	Perm P-Value	Gene Symbols	miRNAs
chr18:34304267-34802552	CN Loss	498286	23	56	6	44	0.0019	FHOD3, TPGS2, LOC105372069, KIAA1328	
chr11:48367375-48370611	CN Loss	3237	3	28	26	55	0.006568144	OR4C45	

chr1:16890840-16901409	CN Gain	10570	3	25	26	55	0.010946907	NBPF1	
chr2:240098365-240497905	CN Loss	399541	3	28	26	55	0.013683634	MGC16025, HDAC4, MIR4269, MIR2467, LOC101928111	hsa-mir-4269
chr8:120918570-121473447	CN Gain	554879	18	53	11.5	137	0.0172	DEPTOR, COL14A1, MRPL13, MTBP	
chr10:30024628-32802256	CN Gain	2777629	5	28	24	55	0.0207	SVIL, JCAD, LOC101929279, MTPAP, GOLGA2P6, MIR7162, MAP3K8, LYZL2, SVIL2P, LOC105376480, ZNF438, LOC101929352, ZEB1-AS1, ZEB1, ARHGA12, KIF5B, EPC1, LOC102031319, LOC101929431, CCDC7	
chr1:169572294-169890531	CN Gain	318238	8	51	21	64	0.0208	SELP, SELL, SELE, METTL18, Clorf112, SCYL3, KIFAP3	
chr4:49098865-49244797	CN Gain	145933	9	49	20	64	0.0221		
chr1:155911616-156103255	CN Gain	191640	10	51	19	56	0.0229	RXFP4, MIR6738, ARHGEF2, SSR2, UBQLN4, LAMTOR2, RAB25, MEX3A, LMNA	
chr6:139036828-158520157	CN Loss	19483330	6	49	23	56	0.0233	GVQW2, CCDC28A, ECT2L, REPS1, ABRACL, HECA, TXLNB, CITED2, LINC01625, FILNC1, LOC103352541, LOC100507477, MIR3668, MIR4465, NMBR, GIE1, VTA1, ADGRG6, LOC153910, HIVEP2, LINC01277, AIG1, ADAT2, PEK3, FUC4A2, PHACTR2-AS1, PHACTR2, LTIV1, ZC2HC1B, PLAGL1, HYMAI, SF3B5, STX11, SNORA98, UTRN, EPM2A, FBXO30, LOC100507557, SHPRH, GRM1, RAB32, LOC101928661, ADGB, KATNB1L1P6, STXBPS5, SAMD5, SASH1, UST, UST-AS1, LOC105378047, TAB2-AS1, TAB2, SUMO4, ZCH12D, PP1L4, GINM1, RPS18P9, KATNA1, LAT51, LOC645967, NUP43, PCMT1, LRP11, RAET1E, RAET1E-AS1, RAET1G, LOC105378052, ULBP2, ULBP1, RAET1K, RAET1L, ULBP3, PPP1R14C, IYD, PLEKHG1, MTHFD1L, LOC102723831, AKAP12, ZBTB2, RMND1, ARMT1, CCDC170, ESR1, MIR3163, SYNE1, SYNE1-AS1, MYCT1, VIP, FBXO5, MTRFL1, RGS17, OPRM1, IPCEF1, CNKS3, SCAF8, MIR1273C, TIA1, CLDN20, TFB1M, NOX3, LOC105378068, MIR1202, SNORD28B, MIR4466, ARID1B, TMEM242, ZDHHC14, MIR3692, SNX9, SYN12-IT1, SYN12	hsa-mir-1273c, hsa-mir-1202
chr6:158733977-159548166	CN Loss	814190	6	49	23	56	0.0233	SNORA16, TULP4, TMEM181, MIR7161, DYNLT1, SYT13, MIR3918, EZR, EZR-AS1, OSTCP1, C6orf99, RSPH3, TAGAP	
chr4:49244857-49319646	CN Gain	74791	3.5	36	25.5	55	0.02583618		
chr6:51742394-53787700	CN Gain	2045309	12	51	17	56	0.0259	PKHD1, MIR206, LINCMD1, MIR133B, IL17A, IL17F, MCM3, PAQR8, EFHC1, TRAM2, TRAM2-AS1, LOC730101, TMEM14A, GSTA7P, GSTA2, GSTA1, GSTA5, GSTA3, GSTA4, ICK, FBXO9, GCM1, MIR5685, FLOVLS, RPS16P5, GLCL, LOC101927136, LINC01564, KHL31, LRC1	hsa-mir-206, hsa-mir-133b
chr1:175329942-178517389	CN Gain	3187449	7.5	50	21.5	60	0.0281	TNR, LINC01657, SCARNA3, COP1, PAPPA2, ASTN1, MIR482, BRINP2, LINC01645, LINC01741, SEC16B, CRYZL2P-SEC16B, CRYZL2P, RASAL2-AS1, RASAL2, TEX35, Clorf220	hsa-mir-488
chr1:181051640-182571579	CN Gain	1519941	7.5	50	21.5	60	0.0281	IERS5, LINC01732, LINC01699, CACNA1E, ZNF648, LINC01344, GLUL, TEDDM1, LINC00272, RGS11, RNASEL, RGS16	
chr8:85861666-86839229	CN Gain	977566	19	53	10	210	0.0289	LRRCC1, LOC102723322, E2F5, C8orf59, CA13, CA1, CA3, CA3-AS1, CA2, REKO1L2P	
chr7:100635850-100645140	CN Gain	9292	17	54	12	109	0.0294	MUC12	
chr8:126136286-135642910	CN Gain	9506634	20	53	9.5	175.2	0.02954	NSMCE2, TRIB1, LINC00861, LOC101927657, FAM84B, PCAT1, PCAT2, PRNCR1, CASC19, CCAT1, CASC21, CASC8, CCAT2, POU5F1B, CASC11, MYC*, MIR1204, TMEM75, PVT1, MIR1205, MIR1206, MIR1207, MIR1208, LINC00824, LINC00976, LINC00977, MIR3686, CCDC26, GSDMC, FAM49B, MIR5194, ASAP1-IT2, ASAP1, ASAP1-IT1, ADCY8, FFR3A, OC90, HHLA1, KCNQ3, HPYR1, LRC6, TMEM71, PHF20L1, TG, MIR7848, PTSC1, SLA, WISP1, NDRG1, ST3GAL1, LOC105375773, LOC101927798, LOC101927822, ZFAT, ZFAT-AS1	hsa-mir-1204, hsa-mir-1205, hsa-mir-1206, hsa-mir-1207, hsa-mir-1208
chr8:141568561-141889251	CN Gain	320693	17.3	53.7	11.7	210	0.029833333	AGO2, PTK2	hsa-mir-151
chr6:57213464-58609380	CN Gain	1395920	6.5	49	22.5	56	0.030925	PRIM2, MIR548U, GUSBP4, LINC00680-GUSBP4, LINC00680	hsa-mir-548u
chr18:350676761-38973978	CN Gain	3906308	3	28	26	55	0.031472359	CELF4, MIR4318, MIR924HG, MIR924, MIR5583-2, MIR5583-1, LINC01902, LINC01477	hsa-mir-4318, hsa-mir-924

chr7:101865172-101892455	CN Gain	27284	15	53	14	109	0.033	CUX1	
chr1:69736-232213	CN Gain	162478	4	49	25	55	0.0331	OR4F5, LOC729737	
chr19:11979616-12573085	CN Gain	593470	4	49	25	55	0.0331	ZNF439, ZNF69, ZNF700, ZNF763, ZNF433-AS1, ZNF433, ZNF878, ZNF844, ZNF788P, ZNF20, ZNF625-ZNF20, ZNF625, ZNF136, LOC100289333, ZNF44, ZNF563, ZNF442, ZNF799, ZNF443, ZNF709	
chr8:86841900-96166177	CN Gain	9324284	20	53	9.43	126.6	0.034614286	PSKH2, ATP8V0D2, SLC7A13, WWP1, RMDN1, CPNE3, CNGB3, CNBD1, DCAF4L2, MMP16, LOC101929709, RIPK2, OSGIN2, NBN*, DECR1, CALB1, LINC00534, LINC01038, TMEM64, NECAB1, C8orf88, PIP4P2, OTUD6B-AS1, OTUD6B, LRRK69, MIR4661, SLC26A7, RUNX1T1, LOC102724710, FLJ46284, TRIQK, MIR8084, C8orf87, LINC00535, FAM92A, RBM12B, RBM12B-AS1, TMEM67, MIR378D2, PDP1, CDH17, GEM, RAD54B, F5BP, VIRMA, LOC100288748, ESRP1, DPY19L4, INTS8, CCNE2, TP53INP1, NDUFAF6, MIR3150HG, MIR3150B, MIR3150A, PLEKH2	hsa-mir-3150
chr8:11990735-12009128	CN Loss	18394	19	53	10	115	0.0349	USP17L7, FAM66D, USP17L2	
chr19:8974148-10112806	CN Gain	1138661	3.7	49	25.3	55	0.035111731	MUC16, OR1M1, OR7G2, OR7G1, OR7G3, ZNF317, OR7D2, OR7D4, OR7E24, ZNF699, ZNF559, ZNF559-ZNF177, ZNF177, ZNF266, ZNF560, ZNF426, ZNF426-D1-ZNF121, ZNF561, ZNF561-AS1, ZNF562, ZNF812P, ZNF846, LOC100505555, FBXL12, UBL5, PIN1, OLFM2, COL5A3	
chr1:173056437-173154765	CN Gain	98329	7	49	22	56	0.0354	TNFSF4	
chr1:173953664-175126323	CN Gain	1172660	7	49	22	56	0.0354	RC3H1, LOC102724601, GPR52*, RABGAP1L, LOC101928696, CACYBP, MRPS14, TNN, KIAA0040	
chr1:186054498-186957787	CN Gain	903290	7	49	22	56	0.0354	HMCN1, PRG4, TPX, ODR4, OCLM, PDC, LOC102724919, PTGS2*, PACER, PLA2G4A	
chr19:22319640-24346108	CN Gain	2026470	3.5	49	25.5	55	0.036117597	ZNF676, ZNF729, ZNF98, LOC101929124, LINC01233, GOLGA2P9, LOC100996349, LINC01785, ZNF492, ZNF99, ZNF723, ZNF728, LINC01859, LINC01858, ZNF730, ZNF724, IPOS1, ZNF91, LINC01224, ZNF675, ZNF681, RP5AP58, ZNF726, LOC100505851, ZNF254, HAVCR1P1	
chr11:89406870-89777672	CN Gain	370803	5	49	24	55	0.0366	FOLH1B, TRIM77, TRIM49, TRIM53AP, TRIM51EP, TRIM64B, TRIM49D1, TRIM49D2, TRIM49D2, TRIM49D1, TRIM64, TRIM51EP, TRIM53AP, TRIM49C	
chr4:49244857-49320199	CN Loss	75343	3	49	26	55	0.037219485		
chr8:96289202-103404337	CN Gain	7115137	20	53	9.5	64	0.03865	C8orf37-AS1, LOC100500773, GDF6, UQCRB, MTERF3, PTDS1, LOC101927066, TSPY1L, SNORD3H, MTDH, LAPTM4B, MATN2, SNORAR72, RPL30, ERICH5, RIDA, POP1, NIPAL2, KCNS2, STK3, OSR2, VPS13B, MIR599, MIR875, COX6C, SNORD77B, RGS522, FBXO43, POLR2E, SPAG1, RNF19A, MIR4471, ANKRD46, SNX31, MIR7705, PABPC1, YWHAZ, FLJ42969, ZNF706, NACA4P, GRHL2, NCALD, LOC104054148, MIR5680, RRM2B, UBR5-AS1, UBR5	hsa-mir-599, hsa-mir-875, hsa-mir-1273
chr7:74580831-74954762	CN Gain	373933	15	52	14.5	82.5	0.0421	NCF1C, GTF2IP4, GTF2IP1, SPDYE10P, SPDYE17, SPDYE11, SPDYE13P, SPDYE14P, SPDYE15P, CASTOR2, PMS2P5, SPDYE8P	
chr8:114388948-116424864	CN Gain	2035917	18	54	11	210	0.0438	CSMD3, TRPS1	
chr18:121171-778209	CN Loss	657041	12.7	51.7	16.3	90	0.047933333	ROCK1P1, USP14, THOC1, COLEC12, LINC01925, CETN1, CLUL1, TYMSOS, TYMS, ENOSF1, YES1	
chr4:49095653-49098814	CN Loss	3162	5	49	24	55	0.048		

Permutated p-value calculated by permuting the survival time for each sample and comparing the log-rank statistic for the permuted data to the original data.

*Drug resistance genes, ** Clinically actionable gene (based on the DGIdb²¹)

Table S11. Distribution of patients per analysis and their associated PFS, lesion and overall objective response.

Pt ID	Best Overall Response	PRE_lesion	POST_lesion	PRE_lesion response	POST_lesion response	PRE_Lesion location	PFS (months)	PFS-Event Censored	WES	RNAseq	miRNA	VEGF	CTDNA_PRE	CTDNA_POST
01-001	SD	X	N/A	ARES	N/A	Liver, Seg. 8	3.78	No	Y	Y	N	Y	Y	Y
01-002	PD	X	N/A	SD	N/A	Liver, Seg. 4b	1.68	No	Y	Y	N	Y	Y	N
01-003	PD	X	N/A	SD	N/A	Liver, Seg. 2	0.92	No	Y	Y	N	Y	N	N
01-004	N/A	X	N/A	N/A	N/A	Liver, Seg. 1	0.46	Yes	Y	Y	Y	Y	N/A	N/A
01-005	SD	X	N/A	SD	N/A	Right Rectus Lesion	2.53	Yes	Y	Y	N	Y	Y	N
01-006	PD	X	N/A	IRES	N/A	Liver, Seg. 2 junction Seg. 3	1.84	No	Y	Y	N	Y	Y	N
01-007	PD	X	N/A	IRES	N/A	Liver, Seg. 6	1.61	No	Y	Y	Y	Y	Y	N
01-008	PD	X	N/A	IRES	N/A	Liver, Seg. 6	1.45	No	Y	Y	Y	Y	N	N
01-009	SD	N/A	N/A	N/A	N/A	N/A	7.95	No	N/A	N/A	N/A	Y	Y	Y
01-010	PD	X	N/A	N/A	N/A	Lymph node	1.61	No	Y	Y	Y	Y	Y	N
01-011	PD	X	N/A	IRES	N/A	Liver, Seg. 2/3	1.61	No	N/A	N/A	N/A	Y	Y	N
01-012	SD	N/A	N/A	N/A	N/A	N/A	3.58	No	N/A	N/A	N/A	Y	N	Y
01-013	PR	N/A	N/A	N/A	N/A	N/A	18.33	No	N/A	N/A	N/A	Y	Y	Y
01-014	SD	X	N/A	ARES	N/A	Liver, Seg. 5	15.64	No	Y	Y	N	Y	Y	Y
01-015	PD	X	N/A	IRES	N/A	Liver, Seg. 4b	1.74	No	N/A	N/A	N/A	Y	N/A	N/A
01-016	SD	X	N/A	ARES	N/A	Liver, Seg. 7	6.9	No	Y	Y	N	Y	Y	Y
16-017	PD	X	N/A	SD	N/A	Liver, Seg. 3	1.74	No	N/A	N/A	N/A	Y	N	N
15-018	N/A	X	N/A	N/A	N/A	Left upper quadrant peritoneal mass	0.33	Yes	Y	Y	Y	Y	N/A	N/A

Pt ID	Best Overall Response	PRE_lesion	POST_lesion	PRE_lesion response	POST_lesion response	PRE_Lesion location	PFS (months)	PFS-Event Censored	WES	RNAseq	miRNA	VEGF	CTDNA_PRE	CTDNA_POST
15-020	PD	X	N/A	IRES	N/A	Liver, right lobe	0.52	Yes	Y	Y	Y	Y	N/A	N/A
01-022	PD	X	N/A	SD	N/A	Liver, Seg. 6/7	1.81	No	Y	Y	Y	Y	N	N
01-023	N/A	X	N/A	N/A	N/A	Liver, Seg. 3	0.53	Yes	Y	Y	Y	Y	N/A	N/A
06-024	PD	X	N/A	SD	N/A	Abdominal wall	1.81	No	Y	Y	Y	Y	N	N
06-025	SD	X	N/A	ARES	N/A	Liver, Seg. 7/8	6.9	No	N/A	N/A	N/A	Y	Y	Y
01-026	SD	N/A	N/A	N/A	N/A	Liver, Seg. 5	9	Yes	N/A	N/A	N/A	Y	Y	N
01-027	SD	X	N/A	ARES	N/A	Liver, Seg. 6/7	3.45	No	N/A	N/A	N/A	Y	Y	Y
06-028	SD	N/A	N/A	N/A	N/A	N/A	0.49	Yes	N/A	N/A	N/A	Y	N/A	N/A
01-030	SD	X	N/A	SD	N/A	Left lung	7.79	No	Y	Y	N	Y	N	Y
01-031	PR	X	N/A	ARES	N/A	Liver, Seg. 2	7.29	No	Y	Y	Y	Y	Y	Y
06-032	PD	N/A	N/A	N/A	N/A	N/A	1.61	No	N/A	N/A	N/A	Y	Y	N
01-033	PD	X	N/A	IRES	N/A	Abdominal wall	0.92	No	Y	Y	N	Y	Y	N
01-034	PD	N/A	N/A	N/A	N/A	N/A	1.81	No	N/A	N/A	N/A	Y	Y	N
01-035	N/A	N/A	N/A	N/A	N/A	N/A	0.36	Yes	N/A	N/A	N/A	Y	N/A	N/A
01-037	SD	N/A	N/A	N/A	N/A	N/A	1.87	No	N/A	N/A	N/A	Y	Y	N
01-038	PD	X	N/A	N/A	N/A	Unknown	1.81	No	Y	Y	N	Y	Y	N
01-039	PD	N/A	N/A	N/A	N/A	N/A	1.71	No	N/A	N/A	N/A	Y	Y	Y
01-040	PD	X	N/A	IRES	N/A	Liver, Seg. 7/8	1.61	No	Y	Y	Y	Y	N	N
06-041	PD	X	X	ARES	IRES	Liver, Seg. 5	2.1	No	Y	Y	Y	Y	Y	N
01-042	N/A	X	N/A	N/A	N/A	Liver, Seg. 8	0.13	No	Y	Y	Y	Y	N/A	N/A
01-045	SD	X	X	ARES	ARES	Transgluteal	8.08	No	N/A	N/A	N/A	Y	Y	Y
06-046	SD	X	N/A	ARES	N/A	Liver, Seg. 8	3.52	No	Y	Y	Y	Y	Y	Y

Pt ID	Best Overall Response	PRE_lesion	POST_lesion	PRE_lesion response	POST_lesion response	PRE_Lesion location	PFS (months)	PFS-Event Censored	WES	RNAseq	miRNA	VEGF	CTDNA_PRE	CTDNA_POST
06-047	PD	X	X	ARES	SD	Liver, Seg. 4a/2	1.77	No	Y	Y	Y	Y	N	N
06-048	PD	X	N/A	SD	N/A	Liver, Seg. 8	0.82	No	Y	Y	Y	Y	Y	N
18-049	PR	X	N/A	ARES	N/A	Omentum	20.73	No	N/A	N/A	N/A	Y	Y	Y
18-050	PD	X	N/A	SD	N/A	Liver, Seg. 5	1.74	No	Y	Y	Y	Y	N	N
18-051	SD	X	N/A	SD	N/A	Liver, Seg. 5	3.58	No	Y	Y	Y	Y	Y	Y
06-052	SD	X	N/A	SD	N/A	Peritoneal carcinomatosis. Nodules along superficial linea alba	7.33	No	N/A	N/A	N/A	Y	N	Y
06-053	SD	X	N/A	SD	N/A	Liver, Seg. 6/7	1.64	Yes	Y	Y	Y	Y	Y	N

X: Biopsy collected

PR: Partial response

SD: Progression of disease

ARES: Acquired resistance lesion

IRES: Intrinsic resistance lesion

Y: Yes, profiled and passed sequencing QC

N: No, profiled and did not pass sequencing QC

NA: Not available, sample not profiled or collected

 Liver samples used for the RNAseq or miRNA analysis

Table S12: Significant differential expression of genes between each RECIST response group comparison

GROUP COMPARISON	# OF SIGNIFICANT GENES
SD (N=7) VS ARES (N=7)	42
SD (N=7) VS IRES (N=5)	76
ARES (N=7) VS IRES (N=5)	197
SD+ARES (N=14) VS IRES (N=5)	195

Table S13: List of genes differentially expressed

SD vs IRES			IRES vs ARES			SD+ARES vs IRES		
Genes	log2FC	FDR	Genes	log2FC	FDR	Genes	log2FC	FDR
DKK1	-8.06	4.15E-15	MAGEC1	25.24	6.12E-28	PEG10	-6.85	7.44E-11
KRT6A	-9.82	1.53E-10	MAGEC2	27.58	6.12E-28	DKK1	-6.21	3.65E-08
PEG10	-6.81	3.03E-08	MAGEA6	14.77	1.51E-14	TUBB2B	-5.61	3.49E-07
CPLX2	-6.76	5.21E-06	MAGEA2	10.67	1.08E-08	DYNC1I1	-3.70	4.52E-07
TUBB2B	-5.32	7.22E-05	PEG10	6.90	1.94E-08	LYPD3	-4.44	4.52E-07
DYNC1I1	-3.43	1.88E-04	CSAG1	12.97	7.72E-08	KRT6A	-7.73	4.97E-07
DIRAS2	-5.66	1.88E-04	MAGEA3	11.94	8.59E-08	S100A3	-5.45	1.45E-06
SLITRK6	-6.72	1.88E-04	S100A3	6.26	5.16E-07	SCG2	-4.47	2.22E-06
KRT16	-6.62	1.88E-04	DYNC1I1	4.09	8.46E-07	CPLX2	-6.09	2.24E-06
LYPD3	-4.16	1.89E-04	DKK1	5.41	1.38E-06	DIRAS2	-5.45	1.09E-05
S100A3	-5.03	2.63E-04	TUBB2B	5.96	1.49E-06	CALML5	-11.25	1.32E-05
SIX4	-3.96	2.80E-04	SCG2	5.00	4.44E-06	ASXL3	-4.87	1.96E-05
PLA2G4A	-3.21	3.01E-04	LYPD3	4.71	4.46E-06	CNTN1	-6.55	3.44E-05
CDR2L	-2.04	4.02E-04	FGF3	10.34	1.21E-05	NPTX1	-5.19	3.44E-05
ADRA2C	-4.93	5.16E-04	SLC9A4	7.32	1.89E-05	TDRD1	-6.51	4.00E-05
SCG2	-4.09	7.09E-04	ASXL3	5.46	1.89E-05	SLC1A5	-1.86	4.00E-05
TUBB2A	-3.36	9.57E-04	KRT6A	6.97	4.47E-05	TUBB2A	-3.33	5.14E-05
CALML5	-10.63	9.57E-04	IGFBPL1	5.55	4.80E-05	RIMKLA	-2.71	7.03E-05
SLC1A5	-1.85	9.57E-04	RTKN2	2.29	4.92E-05	TBX3	-1.89	9.30E-05
TBX3	-1.94	1.01E-03	TDRD1	7.30	4.92E-05	ADRB1	-4.84	2.08E-04
DEFA5	-8.08	1.35E-03	CALML5	12.32	8.09E-05	KRT5	-6.35	2.26E-04
PRSS56	-8.77	1.66E-03	IGSF1	4.73	1.17E-04	BMP5	-4.49	2.80E-04
FGF19	-6.09	1.74E-03	ANKRD34B	8.35	2.08E-04	SIX4	-3.49	2.80E-04
ASXL3	-4.45	1.74E-03	CHGA	8.29	2.08E-04	IFITM1	2.61	2.98E-04
RIMKLA	-2.63	1.99E-03	FAM46B	4.02	3.76E-04	CDR2L	-1.82	3.56E-04
B3GALT2	-4.48	2.28E-03	CPLX2	5.59	3.76E-04	SLC9A4	-5.77	4.09E-04

TDRD1	-6.04	2.28E-03	MLLT11	3.10	5.34E-04	ECEL1	-5.90	5.52E-04
INHBB	-3.97	2.56E-03	DIRAS2	5.24	5.50E-04	PCDH18	-3.66	5.52E-04
KRT5	-6.39	2.63E-03	RIMKLA	2.79	6.26E-04	FGF3	-7.65	5.52E-04
AKR1B10	-4.75	2.81E-03	COL11A2	4.95	6.26E-04	MT4	-7.14	5.52E-04
TGFB2	-4.12	3.01E-03	ADAMTS20	8.61	6.26E-04	SRRM3	-3.95	6.42E-04
ADRB1	-4.77	3.29E-03	SOX3	10.63	7.67E-04	CDX2	2.44	6.42E-04
IL17RD	-4.02	3.68E-03	IFITM1	-2.88	7.67E-04	VGLL3	-3.36	7.51E-04
MSX1	-3.98	3.89E-03	SLC1A5	1.88	7.67E-04	NOTUM	-4.98	7.66E-04
PIAS3	-2.21	7.46E-03	ECEL1	6.64	7.89E-04	VSTM2L	-4.14	7.66E-04
VGLL3	-3.37	7.46E-03	PCDH20	-3.68	7.95E-04	KRT16	-5.39	1.14E-03
NOTUM	-4.97	8.68E-03	NEB	3.99	8.23E-04	VSIG8	-4.55	1.21E-03
IGFBP3	-2.00	9.07E-03	TUBB2A	3.31	1.01E-03	CLDN5	-3.47	1.30E-03
PCDH18	-3.60	9.56E-03	PRDX5	-2.21	1.04E-03	TGFB2	-3.78	1.38E-03
DBNDD1	-1.56	1.09E-02	REXO1L1	10.79	1.26E-03	LYPD8	6.65	1.38E-03
TRIM62	-2.41	1.11E-02	LGSN	5.31	1.56E-03	FAM46B	-3.33	1.55E-03
SLC9A4	-5.14	1.15E-02	NEUROD1	6.39	1.63E-03	MLLT11	-2.62	1.63E-03
FGF3	-6.93	1.15E-02	ADRB1	4.92	1.72E-03	SGCE	-2.51	1.77E-03
HIST2H3D	-1.94	1.20E-02	ZNF8	1.17	1.72E-03	CT45A5	-7.94	1.80E-03
PROM2	-2.75	1.20E-02	TBX3	1.83	2.29E-03	RAC3	-2.21	1.90E-03
TMEM158	-4.20	1.31E-02	SOX21	7.62	2.30E-03	GPR155	-3.17	1.97E-03
ECEL1	-5.48	1.32E-02	RASL11B	4.01	3.04E-03	RELL1	-2.08	1.97E-03
GADD45A	-2.03	1.64E-02	KRT5	6.32	3.04E-03	BAMBI	-3.60	1.97E-03
GPR155	-3.12	1.64E-02	VSIG8	4.92	3.19E-03	PIGR	5.01	2.03E-03
FGFRL1	-2.13	1.64E-02	KCNK2	7.05	3.42E-03	TMEM158	-4.09	2.37E-03
RELL1	-2.06	1.64E-02	COMM3-BMI1	1.09	3.42E-03	PLA2G4A	-2.59	2.44E-03
FSCN1	-2.89	1.64E-02	GPR108	-0.96	3.42E-03	ALDH3B2	-3.90	2.44E-03
DUSP4	-2.59	1.64E-02	SCN5A	5.46	3.62E-03	ARTN	-2.78	2.56E-03
BAMBI	-3.56	1.64E-02	ZNF716	8.65	3.66E-03	PIAS3	-2.03	2.66E-03

VSIG8	-4.31	1.68E-02	CDX2	-2.51	4.70E-03	SLC39A2	5.00	2.66E-03
OSBPL6	-3.03	1.68E-02	ARTN	3.05	4.83E-03	ASPSCR1	-2.24	2.66E-03
SGCE	-2.47	1.68E-02	HHIPL2	3.43	5.12E-03	INHBB	-3.41	2.78E-03
ASPSCR1	-2.26	1.68E-02	SLC4A3	3.94	5.12E-03	C6orf222	2.91	2.81E-03
TTLL7	-3.30	1.82E-02	PCDH18	3.73	5.12E-03	FSCN1	-2.81	3.19E-03
ID3	-2.44	2.06E-02	C18orf54	1.77	5.12E-03	ZIC1	-5.22	3.31E-03
RAC3	-2.12	2.31E-02	SLC38A11	4.89	5.26E-03	GJB6	-4.81	3.41E-03
SMAD6	-1.56	2.43E-02	PTPRZ1	6.57	5.35E-03	APCDD1	-3.73	3.51E-03
SERPINI1	-3.28	2.44E-02	PCDHB2	4.02	5.35E-03	TRIM62	-2.24	3.55E-03
GNAI1	-3.11	2.44E-02	CASP5	-3.12	5.35E-03	AKR1B10	-4.07	3.56E-03
AGPAT9	-3.27	2.51E-02	CRB3	-1.03	5.35E-03	PRDX5	1.86	3.75E-03
RASL11B	-3.36	2.68E-02	COL4A6	7.66	6.23E-03	CYP24A1	-5.03	4.22E-03
APCDD1	-3.54	3.47E-02	ARHGEF18	-1.05	6.58E-03	CASP5	2.81	4.32E-03
FAM46B	-2.91	3.63E-02	VGLL3	3.35	6.88E-03	C18orf54	-1.58	4.39E-03
PRAME	-3.91	3.63E-02	CTPS1	1.31	7.18E-03	B3GNT3	1.76	4.46E-03
FAHD2B	-1.81	3.78E-02	DLX5	7.90	7.18E-03	OSBPL6	-2.88	4.60E-03
MLLT11	-2.27	3.83E-02	RAC3	2.29	7.32E-03	FAHD2B	-1.84	4.72E-03
ARTN	-2.55	4.25E-02	ZNF300	3.24	7.47E-03	PRAME	-3.98	4.72E-03
TMEM163	-2.82	4.25E-02	MRAP2	-3.75	7.80E-03	FIGN	-2.69	4.80E-03
CEP72	-2.48	4.61E-02	SIX4	3.15	7.80E-03	GPD1L	1.29	4.91E-03
GSTA1	-3.75	4.61E-02	NOTUM	4.99	8.02E-03	SLC4A3	-3.47	5.23E-03
TNFRSF19	-3.17	4.74E-02	MOBP	5.03	8.66E-03	PCDH20	3.13	5.30E-03
			SGCE	2.57	8.66E-03	FABP3	3.11	5.35E-03
			NXPE4	-5.51	8.66E-03	B3GALT2	-3.69	5.35E-03
			CDR2L	1.63	8.66E-03	ADAMTS14	-3.35	5.40E-03
			SULT1E1	5.54	9.04E-03	CTPS1	-1.16	5.45E-03
			OR13H1	4.90	9.64E-03	ADRA2C	-3.81	5.65E-03
			APCDD1	4.00	9.64E-03	HIST2H3D	-1.75	6.37E-03

			GPR155	3.24	1.05E-02	DBNDD1	-1.39	6.37E-03
			S100A7	7.00	1.06E-02	RASL11B	-3.65	6.75E-03
			SERPINB13	8.68	1.06E-02	LYZL2	-4.14	7.02E-03
			BEST3	4.78	1.09E-02	IGFBPL1	-3.91	7.09E-03
			C6orf222	-3.05	1.15E-02	POU3F2	-5.83	7.51E-03
			BAMBI	3.64	1.22E-02	LNP1	-1.29	8.81E-03
			RELL1	2.10	1.22E-02	DAZ4	-7.74	8.93E-03
			SLC39A2	-5.12	1.24E-02	TMEM163	-2.77	9.53E-03
			TTL	1.36	1.24E-02	AGPAT9	-3.05	9.53E-03
			ADAMTS14	3.61	1.25E-02	NXPE4	4.80	9.61E-03
			SPECC1L	1.21	1.26E-02	TSPYL5	-2.95	9.73E-03
			TMPRSS11E	5.32	1.38E-02	ID3	-2.25	9.93E-03
			LRRC31	-3.04	1.39E-02	IL3RA	1.53	1.04E-02
			SEMA3D	3.49	1.39E-02	FUT1	-1.03	1.04E-02
			SELENBP1	-1.98	1.39E-02	KRTAP19-1	-5.87	1.04E-02
			LRRN1	4.41	1.44E-02	SLC38A5	3.38	1.06E-02
			KITLG	2.11	1.44E-02	GADD45A	-1.81	1.07E-02
			OTOP1	8.27	1.50E-02	SPRR3	-6.76	1.07E-02
			HSD17B14	1.54	1.50E-02	FCGBP	4.71	1.07E-02
			S100A4	2.23	1.51E-02	KITLG	-1.88	1.11E-02
			PIR	1.95	1.56E-02	GRAMD2	2.55	1.11E-02
			B4GALNT3	-2.21	1.64E-02	TRIM40	2.72	1.11E-02
			FRMD1	-3.75	1.70E-02	C10orf99	4.44	1.11E-02
			SLC6A15	9.05	1.71E-02	MISP	1.44	1.13E-02
			ATP4A	5.49	1.71E-02	MSX1	-3.22	1.17E-02
			FUT1	1.10	1.71E-02	GPR98	-3.34	1.17E-02
			PIGR	-4.89	1.76E-02	FRMD1	3.36	1.20E-02
			CXorf48	4.96	1.76E-02	SERPINI1	-2.99	1.20E-02

		B3GNT3	-1.82	1.78E-02	FGF19	-4.64	1.32E-02
		TMEM158	4.00	1.78E-02	HOXC8	-3.69	1.32E-02
		ASPSCR1	2.22	1.78E-02	MCOLN2	2.53	1.40E-02
		PRAME	4.07	1.93E-02	SDR16C5	-3.37	1.40E-02
		FAHD2B	1.87	1.94E-02	G6PD	-1.75	1.40E-02
		SBSN	5.53	1.94E-02	SCN5A	-4.36	1.46E-02
		EPHB3	-2.53	1.96E-02	S100A7	-5.62	1.50E-02
		SRPK1	-0.94	1.96E-02	GNAI1	-2.80	1.50E-02
		AKR1B15	4.50	1.96E-02	NWD1	2.61	1.50E-02
		KRT16	4.71	1.96E-02	CDK5R1	-2.06	1.73E-02
		RAB3D	-1.49	2.10E-02	UGT2B17	4.02	1.74E-02
		S100PBP	1.13	2.11E-02	DLL4	1.21	1.76E-02
		CHRM5	3.20	2.11E-02	RPS28	1.42	1.83E-02
		CCDC88B	-2.41	2.19E-02	SOX21	-5.63	1.87E-02
		FCHO1	-1.93	2.24E-02	HIST1H3E	-2.30	1.93E-02
		CEP192	1.39	2.39E-02	DEFA5	-5.89	2.07E-02
		IL3RA	-1.60	2.53E-02	PTGDR	3.44	2.10E-02
		PTGDR	-3.85	2.53E-02	NAA40	-1.23	2.12E-02
		PDE4A	-1.68	2.53E-02	LMO4	-2.16	2.12E-02
		CERCAM	1.89	2.57E-02	LY6G6F	3.21	2.14E-02
		TGFB2	3.38	2.65E-02	TTLL7	-2.81	2.21E-02
		RFX8	2.12	2.65E-02	DLX5	-6.09	2.21E-02
		CXCR4	1.95	2.65E-02	SMAD6	-1.36	2.21E-02
		NRARP	-1.64	2.65E-02	CEACAM7	4.47	2.21E-02
		UNC13C	5.84	2.65E-02	CYP26B1	-2.18	2.31E-02
		FSCN1	2.69	2.71E-02	TRIP13	-1.24	2.31E-02
		NWD1	-2.79	2.73E-02	DNALI1	-2.39	2.33E-02
		INSM1	5.42	2.73E-02	PRSS56	-6.28	2.42E-02

			FBXL14	-1.78	2.74E-02	CDHR1	3.37	2.42E-02
			CHORDC1	1.16	2.82E-02	C17orf78	-4.88	2.49E-02
			C10orf71	7.52	2.82E-02	EREG	2.66	2.52E-02
			CT45A5	8.91	2.83E-02	AP3M2	-1.12	2.55E-02
			FAIM2	-2.99	2.89E-02	RPL36	1.35	2.55E-02
			TPM4	-0.94	2.91E-02	FCHO1	1.71	2.55E-02
			LNP1	1.33	2.91E-02	CXCR4	-1.71	2.60E-02
			PIAS3	1.87	2.97E-02	SOD3	2.24	2.63E-02
			CT45A4	8.27	2.97E-02	MICU3	-2.00	2.63E-02
			LYZL2	4.33	3.09E-02	FOXD4L2	-3.01	2.72E-02
			KREMEN2	3.09	3.09E-02	POF1B	1.05	2.77E-02
			TMPRSS5	3.86	3.11E-02	PLAGL1	1.88	2.79E-02
			CEACAM7	-4.92	3.14E-02	PTBP2	-1.17	2.85E-02
			DCAF4L2	9.17	3.22E-02	NAPG	-1.22	2.87E-02
			CDH19	3.49	3.25E-02	COL4A6	-5.81	2.88E-02
			SLC38A5	-3.49	3.30E-02	MRAP2	3.01	2.88E-02
			OPRD1	4.29	3.31E-02	CEP72	-2.21	2.90E-02
			TRIM62	2.08	3.31E-02	PPM1L	-1.84	2.94E-02
			NR2C1	1.10	3.56E-02	IL17RD	-3.02	2.95E-02
			CTLA4	3.41	3.70E-02	NOS2	2.80	2.97E-02
			TMEM173	-1.55	3.75E-02	LY6G6D	3.36	3.15E-02
			ANKRD35	4.07	3.80E-02	SLITRK6	-4.27	3.16E-02
			OSBPL6	2.71	3.80E-02	GPAM	-1.55	3.22E-02
			SMIM10	2.12	3.80E-02	CDC45	-1.08	3.33E-02
			NAPG	1.36	3.80E-02	ZDHHC11	-2.42	3.35E-02
			TPRN	-1.42	3.81E-02	TPRN	1.24	3.35E-02
			EMB	2.81	3.93E-02	DIO3	2.65	3.35E-02
			GRAMD2	-2.57	3.96E-02	LPAR2	-2.23	3.35E-02

		ZNF280B	1.56	3.96E-02	DYNC2H1	-1.17	3.44E-02
		GPD1L	-1.23	3.96E-02	GGT6	2.61	3.55E-02
		AKR1B10	3.68	3.96E-02	DUSP4	-2.08	3.62E-02
		TSPYL5	2.94	3.97E-02	NRARP	1.40	3.69E-02
		LCN2	-2.63	4.01E-02	SLC12A2	1.36	3.84E-02
		KLHL14	4.12	4.02E-02	SLC26A2	2.45	3.84E-02
		ARHGEF40	-1.36	4.11E-02	CSRNP3	-2.45	4.13E-02
		NDST4	5.64	4.13E-02	PIR	-1.59	4.13E-02
		C10orf99	-4.48	4.13E-02	EID3	-1.74	4.13E-02
		KHNYN	-0.95	4.17E-02	MTSS1L	-1.57	4.13E-02
		UNC80	4.22	4.26E-02	CDH19	-2.99	4.13E-02
		AP1M2	-1.25	4.26E-02	KIF1A	-4.73	4.16E-02
		CCNE1	1.93	4.26E-02	GPR64	3.80	4.16E-02
		SPRR3	6.97	4.30E-02	CLN3	1.36	4.40E-02
		TMEM163	2.70	4.30E-02	ARL4C	-1.35	4.51E-02
		SOWAHB	-1.25	4.30E-02	SERPINB4	-4.07	4.56E-02
		ATP1B4	7.84	4.30E-02	REEP1	2.90	4.56E-02
		CYP26B1	2.33	4.39E-02	TMPRSS3	-1.75	4.72E-02
		MCOLN2	-2.57	4.45E-02	UBA52	1.02	4.78E-02
		NRSN1	5.52	4.45E-02	HOMER1	-1.53	4.80E-02
		PLA2G4A	2.12	4.47E-02	BATF2	1.97	4.81E-02
		BMI1	1.10	4.50E-02	PTPRZ1	-4.71	4.84E-02
		INHBB	2.98	4.72E-02	CCDC88B	2.00	4.84E-02
		LMO4	2.27	4.79E-02	PRDM7	-3.91	4.84E-02
		B3GALT2	3.31	4.79E-02	LRRN1	-3.42	4.85E-02
		ZDHHC11	2.65	4.79E-02	NSG1	-3.29	4.85E-02
		CDC45	1.16	4.79E-02	ZNF439	1.59	4.85E-02
		DLL4	-1.25	4.89E-02	TRNP1	-2.10	4.88E-02

			NT5DC2	1.58	4.94E-02	SLC38A11	-3.52	4.95E-02
			KIF21B	-1.70	4.98E-02			
			GOLPH3L	-0.92	4.98E-02			

Table S14: List of genes in the gene set

Adherens_junction	Tight_junction	Cancer_mesenchymal_transition	EMT	WNT_bCatenin_signaling	TGFB_response_signature_of_fibroblasts	TGFB_response_signature_of_T_cells	TGFB_response_signature_of_macrophage_cells	TGFB_response_signature_of_endothelial_cells	Activation_of_MAPK_activity	PI3K_activity	VEGF_signaling
NECTIN1	CRB3	ACTA2	ABI3_BP	ADAM17	FLT1	TIMP1	C5orf23	ASAP1	ADAM8	ATM	FIGF
NECTIN2	CLDN4	ADAM12	ACTA2	AXIN1	FLT1	RAB31	RAI14	SAV1	ADORA2B	BTC	FLT1
NECTIN3	CLDN3	AEBP1	ADM12	AXIN2	COL10A1	CLIC4	NA	WWTR1	ADRA2A	CD19	FLT4
NECTIN4	CLDN7	ASPN	ANPEP	CCND2	IGFBP3	MXRA7	AHNAK2	ZNF532	ADRA2B	CD28	KDR
PARD3	CLDN19	BGN	APLP1	CSNK1E	NOX4	SERPINE1	ZNF532	ARL4C	ADRA2C	CD80	NRP1
SRC	CLDN16	C1QTNF3	AREG	CTNNB1	MEX3B	INPP5F	CEP170	PALM2-AKAP2	ALK	CD86	NRP2
FARP2	CLDN14	C7orf10	BASP1	CUL1	GAS1	GEM	POSTN	NRP1	ARRB1	EGF	PDGFC
CDC42	CLDN15	CDH11	BDNF	DKK1	INHBA	RAB31	PALM2-AKAP2	CEP170	AVPI1	EGFR	VEGFA
RAC1	CLDN17	COL10A1	BGN	DKK4	VEGFA	SERPINE1	CEP170	BNIP3L	AXIN1	ERBB2	VEGFB
RAC2	CLDN20	COL11A1	BMP1	DLL1	CDKN2B	ANXA5	SPP1	RBMS1	BIRC7	ERBB3	VEGFC
RAC3	CLDN11	COL1A1	CADM1	DVL2	NA	RAB31	NA	MYOF	BMP2	ERBB4	
WAS	CLDN18	COL1A2	CALD1	FRAT1	FBXO32	LMCD1	VEGFA	TGFB1	C1QTNF2	EREG	
WASL	CLDN22	COL3A1	CALU	FZD1	NA	CST6	ERRFI1	HIP1	C5	FGF1	
IQGAP1	CLDN5	COL5A1	CAP2	FZD8	CALB2	RBPJ	APOE	RBMS1	C5AR1	FGF10	
BAIAP2	CLDN10	COL5A2	CAPG	GNAI1	CTGF	RASGRP3	FBXO32	ANGPT2	CCL19	FGF16	
WASF1	CLDN8	COL6A2	CD44	HDAC11	VEGFA	CLIC4	PCOLCE2	CDH2	CD40LG	FGF17	
WASF2	CLDN6	COL6A3	CD59	HDAC2	VEGFA	PLAU	EHD2	VEGFA	CD74	FGF18	
WASF3	CLDN2	COMP	CDH11	HDAC5	KANK4	LOH3CR2A	NA	UGCG	CD81	FGF19	
AFDN	CLDN1	COPZ2	CDH2	HEY1	NET1	ITGAV	ENO2	PLOD2	CDK1	FGF2	
LMO7	CLDN9	CRISPLD2	CDH6	HEY2	HEY1	MXRA7	KCNJ8	IDS	CHRNA7	FGF20	
SSX2IP	CLDN23	CTSK	COL11A1	JAG1	SERPINE1	MAP4	PLAUR	RNF13	CSPG4	FGF22	

SORBS1	CLDN2	DCN	COL1 2A1	JAG2	ESM1	MAP4	TANC2	SPOCK1	CXCR4	FGF2 3	
ACTN1	CLDN2	EDNRA	COL1 6A1	KAT2A	NET1	KLF7	VCAN	WWC2	DAB2IP	FGF3	
ACTN4	OCLN	E PYC	COL1 A1	LEF1	NOX4	ABHD2	NA	FAP	DAXX	FGF4	
VCL	F11R	FAP	COL1 A2	MAML1	TIMP3	BMP1	NA	RBMS1	DBNL	FGF5	
TJP1	JAM2	FBN1	COL3 A1	MYC	SYNE1	KCNK1	PLAUR	EMP1	DRD4	FGF6	
CDH1	JAM3	FN1	COL4 A1	NCOR2	BHLHE40	BMP1	CD109	COL5A2	DUSP5	FGF7	
CTNND1	BVES	GLT8D2	COL4 A2	NCSTN	PLAUR	RGS16	NA	ALCAM	DUSP6	FGF8	
CTNNB1	CDC42	GREM1	COL5 A1	NKD1	APBB2	ATXN1	V CAN	V CAN	DUSP7	FGF9	
CTNNA3	PARD6 A	HNT	COL5 A2	NOTCH1	FGF1	CST6	CDKN1C	V CAN	DUSP9	FGFR 1	
CTNNA1	PARD6 G	INHBA	COL5 A3	NOTCH4	TIMP3	NMB	VEGFA	CALD1	EFNA1	FGFR 2	
CTNNA2	PARD6 B	ITGBL1	COL6 A2	NUMB	SERPINE1	KCNK1	PLA2G16	AKAP13	EGF	FGFR 3	
ACTB	MPP5	LGALS1	COL6 A3	PPARD	ANGPTL4	PLAU	AXL	CHMP2B	EPGN	FGFR 4	
ACTG1	MPP4	LOX	COL7 A1	PSEN2	IMCD1	RGS16	NA	STX7	ERCC6	FRS2	
RHOA	TJP3	LOXL2	COL8 A2	PTCH1	NA	MAP4	KANK4	MAP1B	ERN1	FYN	
PTPRM	PATJ	LRRC15	COM P	RBPJ	FGF1	BMP1	PLOD2	DCBLD2	ERP29	GAB1	
PTPRB	MPDZ	LUM	COP A	SKP2	SYNE1	EGR2	ULBP2	PTRF	FCER1A	GRB2	
PTPRF	PRKCZ	MFAP5	CRLF 1	TCF7	PGM2L1	IL1RN	RASSF8	FBN1	FGF10	HBEG F	
PTPN1	PRKCI	MMP11	CTGF	TP53	KAL1	MAP4	TBC1D8	LRRKIP1	FGF2	IRS1	
PTPN6	AMOT	MMP2	CTH RC1	WNT1	TNFAIP6	ABHD2	SERPINE1	SLC25A36	FPR1	IRS2	
PTPRJ	AMOT	MXRA5	CXCL 1	WNT5B	OSGIN2	ATXN1	SOX11	SULF1	FRS2	KIT	
CSNK2A 1	AMOT	NID2	CXCL 12	WNT6	PODXL	IL1RN	V CAN	10-Sep	GAB1	KITLG	
CSNK2A 2	ARHG AP17	NOX4	CXCL 6		PCDH9	CLIC4	DOCK4	CALD1	GHR	KL	
CSNK2A 3	RAC1	NUAK1	CYR6 1		C13orf33	KLF10	SULF1	ACVR1	GHRL	KLB	

CSNK2B	NF2	OLFML2B	DAB2		RASGRP3	APOD	IER5L	PAPPA	GNG3	LCK	
TCF7	LLGL2	PCOLCE	DCN		LOH3CR2A	IL1RN	C15orf52	GNB5	GRM1	NRG1	
TCF7L1	LLGL1	PDGFRB	DKK1		SPSB1	KLF7	FGD6	CAMSAP1L1	GRM4	NRG2	
TCF7L2	DLG1	PLAU	DPYS L3		FN1	LMCD1	PDLIM7	QKI	HGF	NRG4	
LEF1	SCRIB	POSTN	DST		GADD45B	PIK3IP1	EBF1	PALLD	IKBKG	PDGFA	
IGF1R	PPP2CA	PRRX1	ECM1		TRIB1	CRIP1	AMIGO2	TNFRSF21	IL1B	PDGFB	
INSR	PPP2CB	RAB31	ECM2		HEY1	ATXN1	NA	NRP2	INSR	PDGFR	RA
MET	PPP2R1B	RCN3	EDIL3		STK17B	SLC5A3	NA	UBE2H	IQGAP3	PDGFRB	
EGFR	PPP2R1A	SERPINF1	EFE MP2		KLF7	NR4A3	NA	FERMT2	IRAK1	PIK3C2A	
ERBB2	PPP2R2A	SFRP4	ELN		LRRC8C	EV12A	PTPRM	CDC42BPA	IRAK2	PIK3C2B	
FGFR1	PPP2R2B	SNAI2	EMP3		FNIP2	FAM102A	VCAN	ELK3	ITGA1	PIK3C2G	
FYN	PPP2R2C	SPARC	ENO2		TGFB2	TIAM1	BHLHE40	CALU	KIAA1804	PIK3C3	
YES1	PPP2R2D	SPOCK1	FAP		FRMD4A	ALOX5AP	BACH2	OSBPL1A	KIT	PIK3CA	
MAPK1	PARD3	SULF1	FAS		NA	ATXN1	OLFML2B	NRP1	LPAR1	PIK3CB	
MAPK3	TIAM1	THBS2	FBLN1		CNTN1	ZNF365	PLAUR	CLIC4	LPAR3	PIK3CD	
SNAI2	TJP1	THY1	FBLN2		TIMP3	MAP4	NA	SMAD3	LRRK2	PIK3CG	
SNAI1	TJAP1	TIMP3	FBLN5		APBB2	PIK3IP1	APBB2	ANGPT2	MADD	PIK3R1	
TGFBR1	DLG2	TMEM158	FBN1		NGF	SERPINE1	MFGE8	PICALM	MAP2K1	PIK3R2	
TGFBR2	DLG3	TNFAIP6	FBN2		NUAK1	MXRA7	MYADM	PLAU	MAP2K2	PIK3R3	
SMAD2	NEDD4	VCAN	FERMT2		NA	SLC5A3	SLC16A6	FN1	MAP2K3	PIK3R4	
SMAD3	NEDD4L		FGF2		TNFAIP6	RBPJ	NA	RHOQ	MAP2K5	PIK3R5	
SMAD4	CGN		FLNA		NA	TNFSF4	SERPINE1	LRP12	MAP2K6	PIK3R6	
CREBBP	CGNL1		FMOD		EFNB2	PPAP2A	ACTN1	FGFR1	MAP2K7	PTPN11	

EP300	ARHG EF2		FN1		TSPAN2	PKIA	OLR1	BAG2	MAP3K10	TLR9	
MAP3K7	RHOA		FOXC 2		NA	PKIA	DPP4	RHOQ	MAP3K11	TRAT 1	
NLK	GATA4		FSTL 1		FN1	FBXO11	CD109	CCDC88A	MAP3K2	VAV1	
FER	MARV ELD3		FSTL 3		CHST11	DCLRE1C	TRNP1	KLF7	MAP3K5		
ACP1	MAP3 K1		FUC A1		NA	ABHD2	IGFBP4	ANXA3	MAP3K6		
	MAPK 8		FZD8		STK17B	SOX4	B3GALNT1	UBE2H	MAP3K7		
	MAPK 10		GAD D45 A		VEGFA	C18orf1	PCDHB2	PTPRR	MAP3K9		
	MAPK 9		GAD D45B		BHLHE40	RBPJ	NA	RDX	MAP4K1		
	JUN		GAS 1		EGR2	SLC5A3	ABLIM3	MPDZ	MAP4K2		
	CD1A		GEM		DAAM1	BMP1	NA	TRAM1	MAP4K5		
	CD1B		GJA1		PALLD	NPTX1	LMTK3	COL4A2	MAPK1		
	CD1C		GLIP R1		FNIP2	ABHD2	NA	CALD1	MAPK10		
	CD1D		GLT2 5D1		GPR161	BMP1	HTRA1	STAG2	MAPK11		
	CD1E		GPC1		CALD1	SLC5A3	NA	TPM4	MAPK14		
	CFTR		GPX7		LOC728449	ABHD2	CLDN11	BGN	MAPK3		
	CDK4		GRE M1		TIMP3	CSGALNACT1	TMCC3	NOTCH2	MAPK8IP3		
	YBX3		HTR A1		FNIP2	IL1RN	DPYSL3	BMPR2	MAPKAPK2		
	SYMP K		ID2		NA	CSGALNACT1	ADAMTS2	TM6SF1	MAPKAPK3		
	PCNA		IGFB P2		MGC16121	TMOD1	LOC641518	RRAS2	MAPKAPK5		
	CCND 1		IGFB P3		HIVEP2	ABHD2	MYH10	MEF2A	MDFI		
	ERBB2		IGFB P4		RHOU	SOX4	CST6	ZNF281	MDFIC		
	RUNX 1		IL15		KDM6B	KCNK1	SKI	CALCRL	MOS		
	HSPA4		IL32		FOXP1	SOX4	ELK3	UBE2W	MUC20		

	SLC9A3R1		IL6		ELMOD1	ATP1B1	FGD6	PTPN14	MUL1		
	EZR		IL8		SEMA7A	CCR4	PHLDA1	DPYSL3	NOD1		
	RDX		INHBA		PTHLH	CD83	NA	IGFBP5	NOD2		
	MSN		ITGA2		PMEPA1	LRIG1	SLC16A6	GPR176	NTF3		
	PRKCE		ITGA5		APBB2	FBXO11	KAL1	FADS3	NTRK3		
	ACTB		ITGAV		CDKN2B	ADO	CCDC88A	TRIB2	P2RX7		
	ACTG1		ITGB1		DAAM1	LRBA	CYTH3	FN1	PAK3		
	CACNA1D		ITGB3		HAS2	LMCD1	ATP10A	COL8A1	PDE6G		
	MAP3K5		ITGB5		TGFB2	SOX4	ITGA5	ODZ3	PDE6H		
	MAP2K7		JUN		ANGPTL4	DIXDC1	PLOD2	MACF1	PEA15		
	SRC		LAM A1		KLF7	PPAP2A	SLC43A3	TMEFF1	PIK3CB		
	CTTN		LAM A2		KDM6B	SOX4	EBF1	AKT3	PKN1		
	HCLS1		LAM A3		LMCD1	IL1RN	ZNF281	MACF1	PLA2G1B		
	ACTR2		LAM C1		PGM2L1	SOX4	ADAM12	MBNL2	PLCE1		
	ACTR3B		LAM C2		SNORD114-3	SYNJ2	HSPG2	MACF1	PRKAA1		
	ACTR3C		LEPR E1		NA	DIXDC1	PTPRM	LAMA4	PROK1		
	ACTR3		LGALS1		GRB14	NR4A3	MSR1	TRAK1	PROK2		
	WHA MM		LOX		LIF	RIMS3	LAMC1	ABCA1	PTPLAD1		
	WAS		LOXL1		PTHLH	JUN	APOC1	EMCN	PTPN1		
	WASL		LOXL2		CHST11	CD96	PDLIM4	AIDA	PTPN11		
	VASP		LRP1		TSHZ3	DCLRE1C	NA	FAM129A	RIPK1		
	PRKCA		LRRC15		PDLIM4	CD96	VSIG10L	THBS1	RIPK2		
	PRKCB		LUM		TSPAN2	KLF7	PAC SIN3	RDX	RPS27A		

	PRKACG		MAGEE1		FOXP1	INPP5F	B3GALNT1	PIK3CA	S1PR2		
	RAB13		MATN2		LOC728264	PRKCD	LARP6	SERPINB2	SAA1		
	ARHGEF18		MATN3		TSHZ3	S100PBP	THBS1	PERP	SHC1		
	ROCK1		MCM7		ZNF365	LRBA	CXCR7	MCTP1	SHC2		
	ROCK2		MEST		HAS2	S100PBP	PDLIM7	PDP1	SOD1		
	MYL2		MFA P5		APBB2	HLF	GABARAPL1	OSMR	SPAG9		
	EPB41L4B		MGP		SERPINE1	MAP4	SLC11A1	POLR2K	SYK		
	STK11		MM P1		PDGFC	HLF	NA	MAP1LC3B	TAB1		
	PRKA A1		MM P14		JUNB	ABCC4	APOC1	EZR	TAB2		
	PRKA A2		MM P2		PMEPA1	HLF	GADD45B	FERMT2	TAB3		
	PRKA B1		MM P3		OSGIN2	INPP5F	BTG1	ADAM10	TDGF1		
	PRKA B2		MSX 1		CILP	SYNJ2	MIMP8	BMPR2	TGFA		
	PRKA G1		MXRA5		PGM2L1	KCNK1	ELK3	ARPP19	TGFB3		
	PRKA G3		MYL 9		NA	CXCL13	ITGAV	CLIC4	THBS1		
	PRKA G2		MYLK		KDM6B	PPARG	TANC2	LTBP1	TLR4		
	MYH15		NID2		NA	ABCC4	SPHK1	CLDND1	TNF		
	MYH1		NNMT		NA	ADO	GDF5	DUSP3	TNFSF11		
	MYH2		NOTCH2		BPGM	AHCYL2	ABHD2	TXNIP	TP73		
	MYH3		NTSE		ARHGEF3	FBXO11	HPGD	SUSD5	TRAF6		
	MYH4		NTM		PGBD5	SYNJ2	KIAA1217	ZEB1	UBA52		
	MYH8		OXT R		TAGLN3	AQP3	PECAM1	BACH1	UBB		
	MYH9		PCOLCE		PTHLH	PIK3IP1	APOE	EZR	UBC		
	MYH10		PCOLCE2		TUFT1	BMP1	DPP4	AZIN1	WNT5A		

	MYH1 1		PDG FRB		GPR183	AQP3	PECAM1	MAP4K5	ZAK		
	MYH7 B		PDLI M4		S1PR5	DIXDC1	PDE4DIP	DGKA			
	MYH1 4		PFN2		STK17B	C18orf1	GADD45B	LEPR			
	MYH1 3		PLAU R		CLDN4	JUN	NA	PRKCH			
	MYL6 B		PLO D1		MBOAT2	TSPAN13	FLCN	PLEC			
	MYL6		PLO D2		CNNM4	MAP4	THBS3	TCF7L2			
	MYL9		PLO D3		PMEPA1	AHCYL2	CXCR4	TPM4			
	MYL1 2B		PME PA1		CALD1	SLC1A4	GPX3	BCAT1			
	MYL1 2A		PMP 22		DNAJB5	ADO	TM6SF1	LBH			
	IGSF5		POST N		NA	TFEB	DLC1	PEA15			
	MAGI 1		PPIB		C3orf52	C18orf1	FKBP1B	LEPR			
	SYNP O		PRR X1		DHRS2	ABCC4	S100A2	SLC39A6			
	ACTN1		PRSS 2		SOX4	LRIG1	BEAN	DNAJB4			
	ACTN4		PTHL H		DNAJB5	TMOD1	NMB	APLP2			
	MICAL L2		PTX3		NA	SYNJ2	NAV1	PDLIM5			
	RAB8 A		PVR		LOC728264	SYNJ2	NA	SAMD9			
	RAB8B		QSO X1		EPHA4	ITGAE	TREM1	DEGS1			
	RAPG EF6		RGS4		COL27A1	TFEB	PDK4	STC2			
	RAP1 A		RHO B		SMAD7	NR4A3	MEF2A	MLLT10			
	ITGB1		SAT1		F2RL1	APOBEC3G	ST3GAL6	NMD3			
	AFDN		SCG2		NA	SYNJ2	NA	SLC29A1			
	TJP2		SDC1		NA	CDYL	CDK14	ATP6V1C1			
	RAPG EF2		SDC4		NA	ABCC4	DPYSL3	TLK1			

	RAP2C		SERPINE1		KDM6B	JUN	SPTBN1	PPAP2B			
	MARVELD2		SERPINE2		LOC100128178	SLC1A4	HPGD	FXR1			
	TUBA1B		SERPINH1		NA	FBXO11	CAMK2N1	TGFB2			
	TUBA4A		SFRP1		NA	APOBEC3G	PHLDA1	RANBP9			
	TUBA3C		SFRP4		RASL12	AQP3	RUNX2	PDLIM5			
	TUBA1A		SGCB		SLC35F2	LRIG1	SLIT3	CHMP1B			
	TUBA1C		SGCD		SETBP1	IL7R	HPGD	ATP6V1G1			
	TUBA8		SGCG		NA	ZNF365	USP46	MCL1			
	TUBA3E		SLC6A8		DOCK10	DCLRE1C	LRRC32	SPTBN1			
	TUBA3D		SLT2		NA	IL9	ANGPT2	BCAT1			
	TUBAL3		SLT3		C5orf13	RASGRP3	FADS3	NCOA3			
		SNAI2		DNAJC18	BMP1	ADAM9	CALU				
		SNTB1		DACT1	SLC1A4	CD151	SMARCA2				
		SPARC		WNT9A	RGS16	RGS16	MCFD2				
		SPOCK1		ETV6	ATP1B1	GPX3	TRIM23				
		SPP1		FGF18	IL7R	SLC44A2	NAB1				
		TAGLN		HBEGF	SLC1A4	GULP1	ZEB1				
		TFPI2		TNC	TIAM1	MID2	ADAM10				
		TGFB1		SDC1	CDYL	EBF1	MAP3K7				
		TGFB1		KIAA1755	ZNF238	ADAMTS2	TPP1				
		TGFB3		COL27A1	MYH6	LYNX1	FZD6				
		TGM2		EDN1	LRIG1	COL22A1	THBS1				
		THBS1		HBEGF	RIMS3	NA	MMP14				

			THBS2		ITGB6	SLC1A4	BNC2	CALU				
			THY1		MURC	C18orf1	NA	KIDINS220				
			TIMP1		PTGS2	GPR35	SDCCAG8	HMOX1				
			TIMP3		EPHA4	DCLRE1C	ORA12	SPSB1				
			TNC		PLEK2	CDYL	MMP19	BICD2				
			TNFAIP3		COL27A1	SYNJ2	THBS1	ZFHX3				
			TNFRSF11B		NA	MB	PDE4DIP	OSBPL8				
			TNFRSF12A		RHOU	ZNF238	DENND3	WSB1				
			TPM1		NA		BHLHE40	APPL2				
			TPM2		LOC201651		LAMC1	USO1				
			TPM4		NA		ZEB1	DDA1				
			VCAM1		STEAP2		HPGD	PRKAR1A				
			VCAN		EDN1		DNASE2	IL6ST				
			VEGFA		SLC46A3		NA	PPAP2B				
			VEGFC		IMCD1		NA	IFNGR1				
			VIM		SNX30		NA	SNX3				
			WIPF1		SOX4		NA	TAB2				
			WNT5A		NA		ADAM8	YWHAZ				
					RYBP		FST	SLC6A8				
					NA		IL1RAP	APBB2				
					SOX4		FST	MAF				
					TMEM49		GABARAPL1	LEPR				
					NA		DLC1	MMP14				
					MGC16121		AZI2	PICALM				

				SORBS2		ABHD2	CCNG2				
				KDM6B		THBS1	SAR1A				
				PTGS2		SOCS6	RC3H2				
				LOC100128178		MRC2	CCL2				
				HIC1		NA	PDLIM7				
				NEDD9		UBXN2B	HPS5				
				ARHGEF40		NA	SLC7A11				
				FN1		IL1RAP	ICAM1				
				IFIH1		HTRA4	FHL1				
				GZMK		BNC2	IFI16				
				VEPH1		SPOCD1	FAM129A				
				EPHA4		ADARB1	FHL1				
				FOXP1		ANO4	VAMP3				
				PIK3CD		DFNB31	FAM198B				
				IL6		RGS1	LAMA4				
				YIPF5		SDS	F2R				
				SKIL		DCLK1	RGS4				
				RASD1		OBSL1	DNASE2				
				JARID2		DPP4	JAG1				
				NEDD9		DFNB31	IL32				
				ETV6		TMEM180	SRGN				
				IL11		C21orf7	MAP3K7				
				SETBP1		GNG11	NCK1				
				SNAI1		NA	FHL1				
				NA		CCL7	DKK1				
				NA		UPP1	ICAM1				
				NA		FZD4	BEX4				
				SOX6		FCGR1B	TES				

				SYNE1		DPH3	MYO6				
				STK38L		UBXN2B	ANTXR1				
				NKX3-1		NA	NUPR1				
				CDH6		NA	TUG1				
				NA		N4BP2L2	TCF4				
				PELI1		GRAMD1B	RASA3				
				PRDM1		GLI3	ZMYM6				
				PDPN		FGD6	SS18				
				WNT2		FBLN5	RAB23				
				LMO4		KLF10	CELF2				
				C4orf26		CLEC5A	PRKCI				
				OSGIN2		MAMLD1	B3GNT2				
				CACHD1		NA	KIAA1033				
				PRR5L		TCEAL3	ARF6				
				SOX6		SEMA4B	GOLGA2				
				TMEM2		ST3GAL6	ANKRD12				
				DDX10		IRS1	PARVA				
				MTSS1		BNC2	NID1				
				ARHGEF40		GPC1	CLIP1				
				CLDN14		WBP5	11-Sep				
				JHDM1D		MYOZ1	EID1				
				SLC19A2		NA	MET				
				PLCE1		FPR3	PTPN12				
				PRR9		PDE4DIP	SCARB2				
				MEGF9		NA	C3orf52				
				GOPC		PROC	PROSC				
				NA		FNDC3B	HSP90AA1				
				NA		FZD10	ITGA2				

				NA		MRO	HIPK3				
				MSC		ADM	AOX1				
				PPP1R14C		NA	TJP1				
				NA		NA	PRNP				
				PKNOX2		MSR1	MET				
				MSX2		NA	LAPTM5				
				SNCAIP		PLXDC2	ARL6IP5				
				SLC35F3		NA	MET				
				ETV6		MBD6	SS18				
				PRR5L		PDLIM7	NA				
				LOC727930		LDLRAD3	PICALM				
				HS3ST3B1		NA	SQLE				
				IL11		SLC22A4	SEC23A				
				MEOX1		SH2D3C	FKBP1A				
				STK38L		KCNK10	SOAT1				
				SNCAIP		FRY	TNFSF18				
				NA		PMEPA1	NPTX1				
				E2F7		MSR1	PPAP2B				
				AUTS2		RIN3	RAB3B				
				FUT4		MOAP1	BCL2L1				
				DLX2		NA	CREG1				
				ETV6		TMEM223	ITGB3				
				TBX3		PECAM1	RIOK3				
				TBX3		PDK4	PKN2				
						DIO3	ST6GALNAC4				
						NA	PDLIM5				
						NA	IQGAP1				
						TTC6	IL6ST				

						FST	TRIO				
						IL1RL1	TPR				
						DIO3OS	DAB2				
						CPE	ITGB4				
						FAM89A	GNS				
						CD300LF	PPP1R2				
						SLC29A1	LIG4				
						FMNL2	LYPLA1				
						MTUS1	PAK2				
						PDLIM7	CLDN1				
						BNC2	NA				
						GRB10	RNF146				
						NA	EIF5A				
						NA	FHL1				
						FAM110B	PTK2				
						LOC646903	NBN				
						CRTAC1	CPNE3				
						RGS1	RAB14				
						BNC2	PANX1				
						C5orf62	ARNTL				
						DOCK9	ST6GALNAC4				
						TLR7	TWSG1				
						AXL	NCOA3				
						NMBR	TSPAN3				
						SDCCAG8	IGF2BP3				
						ALOX5	PVR				
						DNM3	YY1				
						LHFPL2	PHTF2				

							ITGA2	APLP2				
							FGF18	CDH2				
							IFIT2	CDK17				
							TGFBR1	SLC6A15				
							NA	HIST1H2BE				
							BTC	TBL1XR1				
							TANC1	BCAP29				
							CDK5RAP2	CSNK1A1				
							FLRT2	HDGFRP3				
							NA	CBS				
							FAM27E3	NRP2				
							NA	CD58				
							NA	GLUD1				
							MGC24103	SLC16A1				
							TSPAN2	SPAG9				
							SLC31A2	PPFIBP1				
							SPRY1	CREBL2				
							NA	PSEN1				
							NA	RYK				
							NA	F2RL1				
							TNFRSF12A	MAX				
							SH2D3C	ARF4				
							NA	KIAA1109				
							RGL3	B4GALT1				
							FHL3	KIAA1033				
							LOC283033	ROCK1				
							LDB2	HSPG2				
							NA	TANK				

							ATP11B	RAC2				
							DAPK1	PROSC				
							NA	VAMP3				
							GAL3ST4	BCL2L1				
							TMEM163	PLEC				
							ABCD1	PXN				
							LTB4R	TM9SF1				
							SLC6A8	ECE1				
							NAV1	EID1				
							ALOX5AP	SCARB2				
							MASP1	TRIO				
							TMEM51	IL8				
							XRCC6BP1	B4GALT1				
							RNF17	NBN				
							LAIR1	SPTBN1				
							NA	DSTYK				
							NA	SCAMP1				
							NA	PLD1				
							ZNF618	LUC7L3				
							ADAM17	TPM1				
							GRB10	PLAA				
							DUSP1	ARFGEF1				
							DHX58	KDELR1				
							PHLDA1	CCPG1				
							NCEH1	LAPTM4B				
							SERPINE1	GLUD2				
							ADAMTS2	SERPINB9				
							SLC22A15	PTP4A1				

						NA	ATP2B1				
						NA	IL6ST				
						NCOR2	HSPA13				
						PMEPA1	PTP4A1				
						ZNF555	WDFY3				
						HOXC10	NA				
						PDLIM7	RIOK3				
						ARL9	MDFIC				
						TSKU	TMCO3				
						NA	IFI27				
						NA	POLG				
						SPATA12	RALB				
						CD109	MTUS1				
						NA	NA				
						C1orf204	NA				
						PHLDA1	SOS2				
						CHD3	OGT				
						IQCG	NA				
						LOC100128501	ITGA6				
						SLFN5	HIPK3				
						NA	CD58				
						CCDC40	GNB1				
						NA	IGF2BP3				
						S100A11	ATP6V1C1				
						MAPK8IP3	MCL1				
						TM7SF4	CBFB				
						AICDA	SPIN1				
						PLEKHA7	SOS2				

						C1orf106	LMAN1				
						UNC5C	PPPDE1				
						MYOZ3	UBA6				
						ADCY2	ATP2B4				
						DNASE2	CDC73				
						NA	WNK1				
						KCNH3	ATP2B1				
						IL28RA	ZNF22				
						JAG1	IQGAP1				
						NA	LPP				
						MAP3K2	SCAMP1				
						COL8A2	FLI1				
						TPST1	NEDD4				
						B3GALNT1	PCDHGA3				
						FGF7	PPP1R12A				
						NA	IL6ST				
						DKK1	PON2				
						PRRG4	UBE2D3				
						NA	TTC3				
						PPARD	MEIS3P1				
						BTG1	LAPTM4B				
						GPR157	CSDA				
						AREG	TMOD3				
						FER1L4	PTPN11				
						UNC5B	SH3BGRL				
						LOC100240734	EIF3A				
						RP1L1	PHACTR2				
						DNM3	NA				

						NUPR1	CRK				
						PHLDA1	UBE2J1				
						PKIA	RYBP				
						IQCH	DOCK9				
						WRNIP1	SDPR				
						MCOLN3	NCKAP1				
						NA	CLDN11				
						TTC7A	CD58				
						RAB23	NFAT5				
						NA	NA				
						PHF17	SHC1				
						EVX1	UBE2G1				
						FPR1	MAP4K5				
						AQP9	RPRD1A				
						JAG1	TGFB2				
						SIM1	PPFIA1				
						CACNA1G	SCAMP1				
						NA	EPAS1				
						NFKBIL2	NFATC2IP				
						NA	EPB41L5				
						FAM84B	FGF2				
						FAM7A3	KLHL7				
						PEX14	SGPP1				
						MBOAT7	HERC4				
						GUCA1A	MICAL2				
						NA	SP110				
						RUNX1	PHACTR2				
						PMEPA1	GSN				

						BEST1	CCNA1				
						DUSP1	SEC14L1				
						SPAM1	RAB5C				
						ADAMTS18	RAD21				
						HCFC1R1	PTGS2				
						PPOX	MAP3K2				
						STOX2	UTP3				
						SPATS2L	GLIPR1				
						TOP1P2	NAA35				
						ARL4A	NBR1				
						VSIG1	TAF9B				
						NA	NEDD9				
						NA	TOX4				
						P4HA2	FLRT2				
						NA	MBNL1				
						KCP	LOXL2				
						TRIM55	EPB41L3				
						LOC283050	NFIB				
						RHOB	TPP1				
						ABHD2	LIMS1				
						NA	PVR				
						DISP1	CALM1				
						MMP14	TMX1				
						NA	SIRPA				
						MMP7	DCTD				
						ABCG2	CHP				
						ABHD8	NF1				
						ARL5A	PAFAH1B1				

						GOS2	PRKAR1A				
						CMTM1	ACBD3				
						FFAR1	TFDP1				
						MYH8	FEM1B				
						TES	HHLA3				
						NA	SYNM				
						SEMA6B	YIPF5				
						CES1	NFYA				
						NA	MGEA5				
						SLC10A2	PHACTR2				
						KLHL6	LYN				
						LAPTM5	KITLG				
						NA	CEACAM1				
						NA	SNAP23				
						CLDN12	TPP2				
						GPR26	NRAS				
						NA	ASF1A				
						PRRG4	XPO7				
						LOC151438	CDC5L				
						NA	RASA1				
						C1orf150	NA				
						PEX2	07-Mar				
						MAP3K2	LMAN1				
						NA	IL13RA1				
						NA	ITPR3				
						NA	GLG1				
						FAM113B	KPNA1				
						FABP6	NA				

						NA	ATP13A3				
						NA	EXOC5				
						SNTB1	LIPG				
						LIMS1	NA				
						SNTB1	ABCF2				
						KLF15	IL1RL1				
						SOAT1	ASNS				
						MTUS1	HSP90B1				
						SH2D1B	ATG5				
						RNASEH2B	ELL2				
						MET	WAC				
						MAP3K2	ATP6AP2				
						SLC39A12	FAIM				
						NA	ABI1				
						KRT13	ADH5				
						RAB3B	PDE4B				
						MAN2A2	PWP1				
						CNBD1	ASPH				
						VASH1	DNAJC10				
						SEC22C	SELT				
						UBQLNL	02-Sep				
						SUCLG1	GLUL				
						JAG1	ENO1				
						ZFYVE19	PTPRF				
						RIF1	CAPN7				
						RHOBTB2	RBM9				
						UNC80	NA				
						ARMC8	C5orf28				

						CADM2	EIF4G1				
						VASH1	HSF1				
						C3	HSPH1				
						SLC8A1	NONO				
						ABCC3	IKBKB				
						DPY19L1	NEDD9				
						TSPAN5	TNFRSF10C				
						MCOLN3	TMEM41B				
						TMEM163	PTPRO				
						NA	GNB1				
						AAA1	KPNA4				
						ABHD2	SPTLC1				
						NA	KIAA1109				
						SLC26A10	NRCAM				
						NA	RAB5A				
						JAG1	TCF25				
						SMAD7	NUCB1				
						NA	RCN2				
						ABCC3	ZNF148				
						NA	LAMP1				
						NA	NA				
						HAMP	SMC3				
						NA	SOS1				
						NA	GTF2I				
						MERTK	RNFT1				
						TRIM62	IL1A				
						TP53I11	ITGA4				
						NA	SLC16A3				

							CLDND2	NA				
							SLC11A1	RBM25				
							SLC16A10	USP34				
							NA	ANXA4				
							AMPH	UBXN4				
							SLC6A8	ETS1				
							ZNF687	DIAPH1				
							NA	SYPL1				
							CNTNAP3B	UBXN4				
							TPM1	EXTL3				
							GPX7	CALR				
							FNDC3B	MGAT2				
							SLC44A2	CREB1				
							NCRNA00113	G3BP2				
							ABHD2	SLC4A7				
							SLIT3	KLHL9				
							MTHFD1L	ELF1				
							NOS3	HDGFRP3				
							LOC286437	YWHAB				
							SLC24A2	PPFIA1				
							RNF6	SMARCA2				
							SERINC2	CCDC6				
							PSME4	SELE				
							C8orf55	MFSD6				
							NA	SORBS3				
							NA	CYP51A1				
							NA	KITLG				
							MRO	CD44				

						NA	RAPGEF2				
						NA	PIGC				
						NA	TOX4				
						NA	EDEM3				
						TSPAN3	ATP6V1A				
						CADM1	ZFR				
						SLC16A3	USP10				
						NA	CPD				
						LOC154822	WTAP				
						TFAP2C	TPR				
						NA	GDE1				
						FPR3	DHX9				
						NA	UTRN				
						FAM40B	CDC27				
						IL3RA	GFRA1				
						SDCCAG8	TWF1				
						NA	STIP1				
						NA	PSAT1				
						STOX2	GMFB				
						IFIT3	TNFRSF10D				
						SH3GL2	RERE				
						NA	COMT				
						GLCCI1	NUP50				
						NA	CD164				
						KRTAP11-1	TGOLN2				
						NA	WIZ				
						C1orf91	ENC1				
						EPAS1	M6PR				

							OR10D3	S100P				
							ASIP	PGRMC1				
							TDRD9	RAD23A				
							NA	NUDT4				
							YPEL2	PAFAH1B1				
							MIA3	LIN7C				
							NA	GPD2				
							BTBD19	IL13RA1				
							GPR84	RABEP1				
							NA	VDAC1				
							PHC2	ANXA4				
							BNC2	MED20				
							MGST1	DMD				
							NA	EML4				
							NA	SPRY2				
							NA	SLC11A2				
							NA	TRIO				
							MFAP3L	UTRN				
							NA	SLC33A1				
							NR4A2	SMAD5				
							NA	NA				
							GLUL	BCLAF1				
							TLE3	NONO				
							TLE3	BUB3				
							NA	NUS1P3				
							MAPKAPK2	SENP3				
							DDA1	DUSP3				
							NA	ZWILCH				

						TPD52L1	PSG6				
						NA	MAPK1				
						FOXD4	MANEA				
						GNB4	PPPDE1				
						LSR	SPAST				
						SMURF1	SNRNP27				
						C2orf40	RNF115				
						FKRP	PSPH				
						NA	TOR1AIP1				
						CLCN4	PTPN11				
						OSM	SMC3				
						SLC8A1	MED6				
						HECW1	MAX				
						NA	EPRS				
						DIRAS2	VPS41				
						CLVS1	USP46				
						MME	SMEK1				
						NA	YWHAE				
						COL22A1	FKBP1A				
						NA	SCYL2				
						ZNF549	SNAP23				
						01-Dec	CTH				
						FRY	HIPK2				
						SKIL	TOR1AIP1				
						SS18	ACLY				
						DGCR6L	TNFRSF10B				
						RWDD2A	PRKACB				
						MBD4	CD164				

						MBD4	CREB1				
						NA	GBX2				
						RUNX1T1	MT1E				
						PXN	FBXW11				
						LOC283104	MT1X				
						ECE1	PRKCI				
						FZD7	CEACAM1				
						RNASE4	DDX18				
						PRIM2	G3BP2				
						BTBD11	TRRAP				
						TLX3	CLPTM1				
						NA	GRB2				
						NA	PAPOLA				
						ABHD2	PREPL				
						HP	FNDC3A				
						FAM40B	MT1M				
						MTHFD1L	NFYB				
						TLR7	NA				
						ZNF589	POT1				
						TRIO	PRPF4				
						PRPF40B	GGA3				
						SOX18	SNAPC3				
						DDHD1	SRSF2IP				
						CLPS	ABCB1				
						ERMP1	CFLAR				
						NA	KIAA0494				
						SLC16A8	C1orf144				
						CDKN1A	PTMS				

						AK7	CBFB				
						STX6	KPNA6				
						PHYHD1	C6orf106				
						PCBP3	RPE				
						OLFML3	MAGT1				
						NA	SLC26A2				
						ZBTB16	ABCC1				
						C9orf117	TPD52				
						TPD52L1	LRP8				
						SOX4	SYNCRIP				
						NA	PCBD1				
						PCDHGB8P	PVR				
						EYA2	MAPKAPK2				
						SNTB1	SLC1A4				
						TMTC1	SMARCC1				
						EN1	SLC7A1				
						NA	METAP2				
						NA	ANKFY1				
						MAP3K2	EPRS				
						ELL2	CLCN3				
						SLC45A3	SNX4				
						HRG	MBTPS1				
						DENND5B	CSNK1A1				
						MDFIC	ALG13				
						TGM2	GPR56				
						NA	FOXJ3				
						GPD2	MFNG				
						NA	TRIO				

						NA	AP1S1				
						MFGE8	DCTN4				
						NA	EDEM3				
						HBEGF	HSP90B1				
						AKR1C2	TSN				
						GLDN	SLC7A1				
						OCIAD2	TRA2B				
						LONRF3	RIPK1				
						ELL2	SERINC3				
						MAP9	GM2A				
						NA	TAGLN2				
						NA	SMG1				
						LOC283454	OSBP				
						NA	QPCT				
						PSME4	MAN1A1				
						NA	SH3GLB2				
						NTRK2	FUCA1				
						SLC6A12	APBA1				
						IL18RAP	GSPT1				
						PLEKHA5	MCM4				
						NA	H2AFY				
						NT5DC4	RANBP2				
						CPD	HADHA				
						NA	PTPRF				
						SLC8A1	TLR4				
						ANKRD29	DAPK3				
						PCMTD1	ELP3				
						FAM178B	NA				

						ZFP36L1	SETD8				
						SLFN5	ARFIP1				
						SMURF2	ZFY				
						SIGLEC8	C19orf6				
						LOC284454	TGFBR2				
						LOC401097	TANK				
						C17orf91	BSG				
						NLRP12	NA				
						USP53	NETO2				
						AZI2	SERINC3				
						NA	ENG				
						C19orf59	MAPK14				
						NPR1	ADD1				
						IFIT3	FAS				
						NA	CD46				
						FMNL3	ARF3				
						NA	HMGA1				
						SLC11A1	AGFG1				
						NA	HNRNPC				
						SLC41A1	MYO9B				
						KCNA3	YAP1				
						CACNA2D1	DDX3X				
						ARID3B	PCK2				
						GAPT	PRPF40A				
						ALOX5	SEL1L				
						BLOC1S2	RAB6A				
						IGF2BP1	GLIPR1				
						NA	ACLY				

						NA	NOLC1				
						CADM1	MAPRE1				
						NA	CDV3				
						PCNX	SKAP2				
						NA	ACTR2				
						SCD	ZC3H15				
						NA	NA				
						NA	MAPK1				
						CCDC157	ZMYND11				
						NA	ITPR2				
						BLMH	DAZAP2				
						NA	EDC3				
						SLC5A7	PCYOX1				
						MFI2	MAP2K2				
						NA	TFG				
						FKBP15	KCTD20				
						CPD	C19orf6				
						TRMT1	DOCK9				
						NA	MTMR1				
						FCGR3B	CAPRIN1				
						NA	SMARCC1				
						NA	PIP5K1A				
						TEX14	HNRNPH1				
						NA	BCLAF1				
						OK/SW-CL.36	C14orf101				
						ABCC3	TMED2				
						COX15	CD46				
						MEX3D	CLN5				

						MERTK	KIAA0494				
						LTA4H	SEL1L				
						NA	GRLF1				
						AKR1C1	MPZL1				
						CNIH4	ROD1				
						NPTXR	SH3BP4				
						NA	CFLAR				
						PPARD	SKAP2				
						TP53INP2	BCLAF1				
						C12orf59	ACTR2				
						KLHL29	AP2B1				
						LARGE	PRKDC				
						NA	SEC23IP				
						MMP2	RRN3P1				
						NA	NUMA1				
						NA	GDI2				
						NA	ME2				
						KCNC3	CYB5R4				
						NAPSB	DCLRE1C				
						ACADVL	NAA15				
						NA	NNT				
						C1orf126	TRIM27				
						NA	DLG1				
						NA	IGF2R				
						CYP19A1	GTF2F1				
						NA	TUBGCP2				
						TNNT2	ERC1				
						NRK	COPA				

						MSTN	NA				
						CYLD	DLAT				
						RGS8	DHX9				
						NA	SCP2				
						HDGFRP3	MAGOH				
						SLC16A10	SRP72				
						HS3ST5	TLE4				
						LOC255167	MAT2A				
						KCNIP2	UBE4B				
						IL17RB	FAM120A				
						ANG	SOCS2				
						FLJ13197	CANX				
						TPSB2	SSR1				
						NA	FAS				
						FURIN	AASDHPPT				
						ABCC3	NA				
						TGFBR1	API5				
						CLDN11	ROD1				
						NEK6	SLC30A1				
						SLC11A1	GGA2				
						CD58	VAC14				
						PYROXD1	KPNA3				
						ALK	TTC37				
						GOLT1A	SSB				
						CNKSIR3	C1orf103				
						CHD9	KIAA0494				
						RHOH	ARF1				
						L3MBTL4	AP1S1				

						GLDN	06-Mar				
						NA	RELN				
						HAL	ARHGDIA				
						NA	USP46				
						S100A8	TNFRSF10B				
						NLRC4	PVR				
						PTGER3	C20orf30				
						NA	FAS				
						CYP19A1	PNO1				
						CLP1	ADD3				
						SLC35F4	CANX				
						IL27RA	PSME4				
						PARVA	MPZL1				
						CARD11	TTC3				
						NA	TFRC				
						MTMR11	PRPF4B				
						PDK4	RAB1A				
						FSTL5	RHOBTB3				
						NA	ZNF192				
						CSN1S1	CHD4				
						ANK1	ATP1B1				
						BCAS4	NA				
						CNIH4	EIF1AX				
						VASH1	DEDD				
						PPIH	MOBKL1B				
						NA	TP53				
						TP53I11	TMBIM6				
						NA	API5				

						NA	DDX3X				
						SLC11A1	HPCAL1				
						DIXDC1	ANAPC5				
						ALOX5	STX16				
						VASH1	EIF4E				
						FOS	TGOLN2				
						CCDC40	TMED2				
						MIA2	NA				
						FOXQ1	MBTPS2				
						MNDA	COMMD10				
						NA	CLCN3				
						NA	ERLIN1				
						SLC1A2	SRSF10				
						CNIH4	PSG3				
						KIAA1274	FKBP15				
						TGM2	GLYR1				
						NA	STS				
						NA	TIA1				
						NA	ANP32A				
						COQ2	SYNJ1				
						WLS	KCTD5				
						MSRA	NFYC				
						NR4A3	LARP4B				
						APOC2	KDM6A				
						GSN	ARHGDIA				
						TFAP2B	POLR2E				
						CCNA1	WDR1				
						LRRC1	CCT2				

							PRDM14	STX3				
							NA	AMD1				
							SMAD6	PDE8A				
							NA	U2AF2				
							NA	BRCC3				
							NA	SERBP1				
							NA	KIF5B				
							ST8SIA2	HSPD1				
							MAP3K2	POLDIP3				
							PHGDH	PNO1				
							NA	YKT6				
							TMEM111	UBE3B				
							STX6	SEC63				
							BEST1	G3BP1				
							PSEN1	SLC1A4				
							NA	ADD3				
							HIPK2	ILF3				
							MSR1	CYB5B				
							ARPP21	ROCK2				
							SBNO1	PTPN11				
							SP100	MOBKL1B				
							NEDD9	PTBP1				
							PKD2L1	RHOBTB3				
							FBP1	KIAA0776				
							CCDC40	ITGA4				
							BEST1	AGA				
							TTC12	GATC				
							FAM71F2	GPR137				

							TOPORS	GFPT1				
							STX6	STAM2				
							B3GNT5	DLG1				
							GPR25	ACTR2				
							NA	CDYL				
							NA	SMPD1				
							EWSR1	MARCKS				
							ADC	UBE2G2				
							FBXO9	EPB41L2				
							NA	AKIRIN1				
							DYX1C1	PSG6				
							CELF6	PREPL				
							TMEM38B	MCM3AP				
							NA	GART				
							TMED5	ESF1				
							NA	NA				
							DENND5B	CHD4				
							ZNF527	CLINT1				
							SPTBN1	DLG1				
							NA	SRPR				
							MEX3C	SEL1L3				
							NA	SLC35A2				
							LRRC27	SDHC				
							GCHFR	MMP1				
							PGLYRP2	AGPS				
							CPD	PLXNA2				
							TRIO	HNRNPR				
							LMX1A	SLC1A4				

						NA	GRAP				
						NA	SRRT				
						KCP	KIAA0182				
						GPR34	C6orf62				
						CADM4	ORC5L				
						ADAM8	NUDT21				
						AZI2	LSS				
						RIT2	CTSS				
						UTY	PDCD4				
						NA	TBX1				
						NA	RNF14				
						NA	DIMT1L				
						ARFGEF2	PLEKHB2				
						CDK5RAP2	SRSF1				
						NUP62CL	CSTF2T				
						PDCD1	HS2ST1				
						PTPDC1	CSNK2A1				
						PWRN2	GM2A				
						NA	TOP1				
						NA	PTPRB				
						PTPRZ1	LSM12				
						CDK3	USP1				
						RBP4	ZFPL1				
						NA	TSPAN12				
						NA	SLC35A3				
						PLAC1L	DLAT				
						C12orf49	UBE2K				
						LOC339807	CCND2				

						DYNC1H1	DSCR3				
						LRRC6	GCLM				
						NA	PAPSS2				
						CCDC40	SEMA3C				
						EFCAB5	SRPK1				
						SLC46A2	SEL1L3				
						NA	CALM1				
						NCR3	GOT1				
						TCEB3B	MOBK1B				
						SDCCAG8	ACOT7				
						TUBE1	CSNK2A1				
						SKIL	DCAF7				
						FCRL1	IDH3A				
						NA	GALNT2				
						GPR37	PRPF6				
						NA	CCND2				
						NA	YBX1				
						RALA	C6orf62				
						SYTL1	UBE2N				
						NA	PPIF				
						NKTR	HSPA9				
						CES4	LARP4				
						CAMP	CYP20A1				
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						EPB41L5	TFAM				
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						NA					

						GOLGA8IP					
						FLJ33996					
						NA					
						GGT7					
						NA					
						NA					
						ECE1					
						SLC35D2					
						TCTN2					
						NA					
						MYO6					
						SLC44A5					
						ATP13A3					
						LIPG					
						DENR					
						ZNF540					
						TPTE2P1					
						NA					
						POU3F1					
						B3GNT5					
						OSM					
						BOLL					
						NA					
						A2ML1					
						S100A8					
						SH2D4A					
						KLHL6					
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						SDC1					
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						GPR110					
						NA					
						TPSAB1					
						SSH2					
						NA					
						C14orf109					
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						ELL2					
						COL8A2					
						DNAL1					
						TMEM38B					
						NDP					
						FGFRL1					
						PDE4DIP					
						XPO1					
						NA					
						MKI67IP					
						SLC7A1					
						NA					
						ALDH1A1					
						NA					
						PDXK					
						RDH12					

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							SYT2					
							GLUL					
							NA					
							ZNF540					
							CLIP3					
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							NA					
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							STOX2					
							SPDEF					
							AGAP3					
							TPSAB1					
							NA					
							CCDC123					
							NA					
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							KLF12					
							NA					
							SPINT1					
							NA					
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						LOC158696					
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						NA					
						CACNA1G					
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						LCK					
						IL1A					
						SLC16A3					
						NA					
						NA					
						STX6					
						MXD1					
						REPS2					
						C11orf75					
						MLPH					

						CCDC63					
						ENO3					
						RABGAP1L					
						SORL1					
						PHACTR1					
						ERBB3					
						NA					
						C11orf63					
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						CCDC114					
						CHERP					
						SKI					
						ROCK1					
						NA					
						CPEB3					
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						NA					
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						TYRO3					
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						NA					
						NA					
						ESR2					
						ISM1					
						NA					
						FNDC3B					
						GLUL					

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						BLID					
						WNT4					
						PTPN22					
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						SEMA7A					
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						DSTN					
						STOM					
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						CHD9					
						TACC2					
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							RRAD					
							NA					
							HAPLN1					
							SSH2					
							LTBR					
							NA					
							NA					
							ARL5A					
							SYT14					
							NA					
							MBD4					
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							PTPDC1					
							TPSAB1					
							CCDC68					
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							NA					
							NA					
							AK1					
							CALHM3					
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							MID2					

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						NA					
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						NA					
						SCGN					
						CNTNAP3					
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						NA					
						NA					
						AGPAT2					
						GNRHR					
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						AGRP					
						GPD2					
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						HES1					
						MTMR11					
						DMD					
						MTMR3					
						EML4					
						DNASE1					
						PNKD					
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						ACER3					

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							PSEN2					
							HIPK2					
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							OR51E1					
							MIINK1					
							ZNF589					
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							OLIG2					
							KLHL6					
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							GAD2					
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							KCNK13					
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							NA					
							KLK2					
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							TRAIP					
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							NA					
							TSPAN13					
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							FMNL3					
							CACNA1E					
							FAM71B					
							BPY2					
							DBF4B					
							SYTL3					
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							CYP2F1					
							OR1J4					
							GBP6					
							PLCG1					
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							NA					
							NA					
							FBXO9					
							LIPJ					
							SCD					
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							NA					
							TPD52					

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							IL1R2					
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							MAPKAPK2					
							REPS2					
							TMEM207					
							TANC2					
							NA					
							GPR157					
							TFEB					
							TRIM36					
							NA					
							TPSAB1					
							SLCO2B1					
							PRR5L					
							PPM1H					
							PEX14					
							CLDN23					
							TRPM3					
							ERMP1					
							AMBRA1					
							REPS2					
							IL17RB					
							SLC31A1					

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							TRIO					
							CYP19A1					
							ASB15					
							NA					
							NA					
							NA					
							NKG7					
							C20orf7					
							C12orf48					
							C18orf8					
							SNX7					
							LHX6					
							USP22					
							TLR3					
							NA					
							TASP1					
							NA					
							SNX10					
							SLC4A11					
							NA					
							MBD4					
							RAPGEF3					
							RNFT1					
							NA					
							ST3GAL6					
							TFCP2L1					

							APOC4					
							NA					
							HOXB8					
							NEB					
							QPCT					
							SGCB					
							DUSP16					
							ICOSLG					
							NEK6					
							ACER3					
							CAPN5					
							EHF					
							NA					
							ENOPH1					
							NA					
							NA					
							SCD					
							DGAT2					
							TIAM2					
							INADL					
							N4BP2L1					
							LOC149773					
							TFR2					
							HN1					
							SLC23A3					
							LOC646576					
							CYP27A1					
							CYB5A					

							NA					
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							HPS4					
							SKIL					
							PNPLA7					
							NEK6					
							CKLF					
							GPR125					
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							NA					
							EPB41L5					
							BUB1					
							PTH2R					
							NA					
							ESR1					
							PAQR5					
							ENG					
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							DNAJC6					
							AACSL					
							OR4D2					
							ATP2B2					
							PARP10					
							SLC24A1					
							LFNG					
							TFEB					

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						ABCG1					
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						NA					
						EPB41					
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						PKD1L2					
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						RHEB					
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						CLDN6					
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							NA					
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							NSUN7					
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							KIAA2026					
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							FANCC					
							RSPO2					
							CCR7					
							SCCPDH					
							PSG7					
							C14orf142					
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						SORL1					
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						ZAN					
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						NA					
						SPATA2					
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						FBXL17					
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						NA					
						SVEP1					
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						PRAM1					
						SKIL					
						NA					

						OGFOD1					
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						CEACAM21					
						CIT					
						TYRO3					
						BMX					
						EDNRB					
						TAGAP					
						C10orf35					
						TBC1D16					
						SH3BP4					
						NA					
						LOC339751					
						NA					
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						IL7R					
						MAPKAP1					
						KRT7					
						SAMD14					
						SPAG1					
						PDPN					
						GPRIN3					
						TLE3					
						CHDH					
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						NAPSB					
						TGM2					
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						NA					
						DENND1B					
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						ABCG1					
						WIP1					
						BIN2					
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							SLC9A5					
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							SARDH					
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							CD1D					
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							GRAMD4					
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						ERN2					
						ST14					
						CNR1					
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						NA					
						TGM4					
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						ACER3					
						CHDH					
						ALDOB					
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						NA					
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						STRBP					
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						C22orf45					
						FAM164A					
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						IL4					
						SYDE1					
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						NUSAP1					
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						C17orf99					
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						C18orf62					
						EDNRB					
						ACER3					
						NA					
						WLS					
						VDR					
						TRIM2					
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						PGP					
						ATP6V0A2					
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						DGAT2					
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						ERLIN1					
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						TRIM58					
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						LOC151121					
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						KIF23					
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						IL18R1					
						PDE4B					
						GKAP1					
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						VPS45					
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						GPRIN3					
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						MYOZ2					
						NA					
						FLJ37453					
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						ERGIC1					
						SERPINA1					
						NA					
						ATP8B1					
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						KRT222					
						IL1B					
						NA					
						NA					
						FAM195A					
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						VRK3					
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						CNTNAP2					
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						EDN3					
						HNF4A					
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						DAB2IP					
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						DKFZp566F0947					
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						MAOA					
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							TBC1D16					
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							MGST1					
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							MGST1					
							FANCI					
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							ATP9B					
							CHDH					
							ADAM3A					
							HOXA3					
							GNA14					
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							MBNL1					
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						C11orf93					
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						SLC34A1					
						TMEM185A					
						NA					
						C6orf182					
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						PAPLN					
						SEMA6A					
						HS6ST3					
						MAP1LC3A					
						C5orf56					
						BRSK1					
						IL1B					
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						ID1					
						SV2B					
						GALNT2					
						LOC652993					
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						ZNF552					
						TRAC					
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						NA					
						EPB41L5					
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						PSPH					
						GABRA1					
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						GLTPD2					
						SAMD11					
						MAOA					
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						FGFRL1					
						MAPK13					
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						TCF21					
						APBB1IP					
						CXCL14					
						CCDC67					
						HESS					
						NUSAP1					
						BLM					
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							SEMA3C					
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						WNT5A					
						NA					
						ARHGEF19					
						GPD1					
						NA					
						SOX6					
						PTTG1					
						ACVR1B					
						SAE1					
						MAEA					
						PPIF					
						ALDH2					
						NA					
						AGMAT					
						FAM46C					
						WNT5A					
						PAPOLB					
						NA					
						NA					

								ASRGL1					
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Table S15: Significant differential expression of miRNA between each RECIST response group comparison

Group comparison	# of significant miRNA	mirna list
SD (N=5) vs ARES (N=4)	14	<i>miR-9-5p</i> , miR-184, miR-1269a, <i>miR-204-5p</i> , miR-504-5p, miR-7641, miR-150-5p, miR-23b-5p, miR-27b-5p, miR342-3p, <i>miR-365a-5p</i> , <i>miR6510-3p</i> , miR-455-5p, <i>miR-934</i>
SD (n=5) vs IRES (n=4)	14	<i>miR-9-5p</i> , miR-9-3p, <i>miR-204-5p</i> , miR-1269a, miR-499a-5p, miR-200b-5p, <i>miR-365a-5p</i> , miR-128-3p, <i>miR-6510-3p</i> , miR-4664-3p, miR-144-5p, miR-486-5p, <i>miR-934</i> , miR-206
IRES (N=4) vs ARES (n=4)	1	miR-184