

## SUPPLEMENTARY MATERIAL

# Characterization of Chromosomal Translocation Breakpoint Sequences in Solid Tumours: “An *In Silico* Analysis”

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**Table S1. Nucleotide sequence of fusion partners retrieved from TICdb of major translocations.**

Sr. No	Translocations	TICdb reference ID	Fusion sequences(FS)	Length of FS in bps	Partner Chr. Having 100% identity	Partner Genes [5' / 3']	BLAT result of FS	UCSC location of FS	UCSC location for 500US /DS From Breakpoint
(1)	t(2;2)	17625570	atataatgtctaactcgggagactatgaaatattgtacTTGTACCGCCGGAAGCA CCAGGAGCTGCAAGCCATGCAGATGGAGCTGC	87	Chr2	5' EML4 (I)	1-39	42552656- 42552694	42552194-42553194
					Chr2	3' ALK (I)	40-87	29446347- 29446394	29445847-29446847
(2)	t(10;10)	10931090	cccgagtagctgtattacaggcacacggcaccacactggctgcttttgaTTATCTC CTCAGCTGAGATGACCTTCCGGAGGCCCGCCAGGCCTT CCCGGTCAGCTACTCCTCTCCGGTG	125	Chr10	5' CCDC6 (I)	1-55	61553383- 61553437	61552937-61553937
					Chr10	3' RET (I)	56-125	43610053- 43610122	43609553-43610553
(3)	t(12;15)	9949179	atactgatcagaacctgaagaagATGTGCAGCACATTAAGAGGAGA GAC	51	Chr12	5'ETV6(I)	1-25	12006471 12006495	12005995-12006995
					Chr15	3'NTRK3(I)	24-51	88483959 88483986	88483486-88484486
(4)	t(5;6)	18083107	aagcccactgacgctccaccgaaagATGATTTTTGGATAACCAGAAACA AGTTTCATACTT	60	Chr5	5'CD74(I)	1-25	149784243 149784267	149783743-149784743
					Chr6	3'ROS1(I)	24-60	117645544 117645580	117645080-117646080
(5a)	t(21;21)	16575875	cgctggcctactctggaagttcagAACTCTCTGATGAATGCAGTGT	47	Chr21	5'TMPRSS2(I)	1-24	42861433 42861456	42860933-42861933

Sr. No	Translocations	TICdb reference ID	Fusion sequences(FS)	Length of FS in bps	Partner Chr. Having 100% identity	Partner Genes [5' / 3']	BLAT result of FS	UCSC location of FS	UCSC location for 500US /DS From Breakpoint
					Chr21	3'ERG(I)	24-47	39795460 39795483	39794983-39795983
(5b)	t(21;21)	16575875	gcaagatggcttgaactcagAAGCCTTATCAGTTGTGAGTG	42	Chr21	5'TMPRSS2(I)	1-21	42870045 42870065	42869545-42870545
					Chr21	3'ERG(I)	21-42	39817523 39817544	39817044-39818044
(6)	t(15;7)	17671502	caactaacactgcggcttctcagTGCCTATGATCAGAAGCCACAAGT G	49	Chr15	5'HMG2P46 (I)	1-24	45806138 45806161	45805661-45806661
					Chr7	3'ETV1(I)	23-49	13975497 13975523	13975023-13976023
(7)	t(1;X)	9393982	gtggttccatgatgggaagtacatgCTGCCTGTGTCAGGGAATCTGCTT	50	Chr1	5'SFPQ(I)	1-26	35652602 35652627	35652102-35653102
					ChrX	3'TFE3(I)	26-50	48891743 48891767	48891267-48892267
(8)	t(5;8)	9525740	tctgcgactcattcatcgccctcagATTGGCCAAAATGGGAAGGATTG GA	51	Chr5	5'LIFR(I)	1-26	38595363 38595388	38594863-38595863
					Chr8	3'PLAG1(I)	25-51	57083724 57083750	57083250-57084250
(9)	t(12;3)	8988031	tttcacaggaatgacagcatctatgaggagCCACAACAAGTTGTTTCAGAAG AAGCTGCT	60	Chr12	3'HMGA2(I)	29-60	66345161 66345192	66344661-66345661
					Chr3	5'FHIT(I)	1-30	59908072 59908101	59907572-59908572
(10a)	t(12;9)	9484777	agttgttcagaagaagcctgctcagTCCTGGTACCTGGGCTAGCTTGGT TCCTTTCCAAG	60	Chr12	5'HMGA2(I)	1-25	66345171 66345195	66344695-66345695
					Chr9	3'NFIB(I)	23-60	14088291 14088328	14087828-14088828
(10b)	t(12;9)	9484777	CcaagaggcagacctaggaatggTCCTGGTACCTGGGCTAGCTTGGT TCCTTTCCAAGTG	61	Chr12	5'HMGA2(I)	1-24	66232326 66232349	66231849-66232849
					Chr9	3'NFIB(I)	24-61	14088289 14088326	14087826-14088826
(11)	t(9;15)	17934517	tatgtcaagtctgtttacagaaaaagcaaggaaccgttctCATCTGCATTGCCG GGACCGGATATGAGCATGAAACCTAGT	84	Chr9	5'BRD3(I)	1-43	136901154 136901196	136900654-136901654
					Chr15	3'NUTM1(I)	43-84	34640170 34640210	34639670-34640670
(12)	t(22;21)	S82335	ctattcctctacacagccgactagttatgatcatCAGATTCTTGGACCAACAA GTAGCCGCCTTGCAAATCCAG	74	Chr22	5'EWSR1(I)	1-33	29682912 29682944	29682444-29683444

Sr. No	Translocations	TICdb reference ID	Fusion sequences(FS)	Length of FS in bps	Partner Chr. Having 100% identity	Partner Genes [5' / 3']	BLAT result of FS	UCSC location of FS	UCSC location for 500US /DS From Breakpoint
					Chr21	3'ERG(I)	33-74	39762917 39762958	39762458-39763458
(13a)	t(22;11)	18602673	agcagcagctacgggcagcagaACCCTTCTTATGACTCAGTC	42	Chr22	5'EWSR1(I)	1-22	29683102 29683123	29682623-29683623
					Chr11	3'FLI1(I)	23-42	128675261 128675280	128674761-128675761
(13b)	t(22;11)	S62665	AGCCAACAGAGCAGCAGCTACGGGCAGCAGAGTCCTCCC CTTGGAGGGGCACAAACGATC	66	Chr22	5'EWSR1(I)	1-33	29683093 29683125	29682625-29683625
					Chr11	3'FLI1(I)	32-60	128677075 128677103	128676575-128677575

**Note:** Sequence in lower case represents 5' partner gene and upper case represents 3' partner gene. US: upstream, DS: downstream.

**Table S2. Analysis and characterization of various parameters for 1000bps (500 bp Upstream & 500 bp Downstream) sequence around breakpoint.**

Sr no.	Translocations	Chr. partners	Location of RSS	Location of Repeats	Partner Gene [5' / 3' ends] & Intronic(I)/ Exonic (E) Break	# of SIDD sites [Threshold cut off of G(x) <4]
(1)	t(2;2)	Chr2	U(435 – 462, 225 – 263, 250 – 288)	D (518-585) <sup>e</sup>	5' EML4 (I)	1
		Chr2	D(534 - 561,795 – 833)	D(802-1215) <sup>e</sup>	3' ALK (I)	5
(2)	t(10;10)	Chr10	U (75 - 113)	W(323-610) <sup>e</sup> /D(694-714) <sup>T</sup>	5' CCDC6 (I)	7
		Chr10	U(127 – 165) D(851 – 889)	U(112-282) <sup>y</sup>	3' RET (I)	1
(3)	t(12;15)	Chr12	U(244 - 282 ,288 - 326 ), D(839 - 866 ,959 - 986 )	U(1-128) <sup>e</sup> ,D(592-639) <sup>y</sup> , D(815-1000) <sup>D</sup>	5'ETV6(I)	3
		Chr15	D(551 - 578 )	D(702-790 , 983-1000) <sup>e</sup> , D(841-968) <sup>D</sup>	3'NTRK3(I)	2
(4)	t(5;6)	Chr5	U(28 - 55 , 291 - 329 )	U(2-37) <sup>e</sup>	5'CD74(I)	1
		Chr6	D(686 - 724 , 709 - 747)	D(639-718 , 952-1000) <sup>D</sup>	3'ROS1(I)	11
(5a)	t(21;21)	Chr21	U(105 - 132,105 - 143, 267 - 305), D(676 - 703 )	D(873-1000) <sup>e</sup>	5'TMPRSS2(I)	3

Sr no.	Translocations	Chr. partners	Location of RSS	Location of Repeats	Partner Gene [5' / 3' ends] & Intronic(I)/ Exonic (E) Break	# of SIDD sites [Threshold cut off of G(x) <4]
		Chr21	U(26 - 53, 32 - 59 ) , D(573 - 611,822 - 860 )	U(22-60) <sup>S</sup> , U(118-146) <sup>E</sup> , D(643-918) <sup>E</sup> , D(922-945) <sup>S</sup>	3'ERG(I)	4
(5b)	t(21;21)	Chr21	D(956 - 994)	—	5'TMPRSS2(I)	2
		Chr21	U(134 - 172 ) , D(936 - 974)	U(8-191) <sup>E</sup>	3'ERG(I)	4
(6)	t(15;7)	Chr15	U(271 - 309)	U(12-502) <sup>V</sup> , D(511-598,924-1000) <sup>V</sup> , D(600-923) <sup>E</sup>	5'HMGN2P46(I)	3
		Chr7	U(413 - 451 )	D(994-1000) <sup>E</sup>	3'ETV1(I)	8
(7)	t(1;X)	Chr1	D(951 - 989 )	U(409-435) <sup>S</sup> , D(948-1000) <sup>E</sup>	5'SFPQ(I)	7
		ChrX	—	—	3'TFE3(I)	1
(8)	t(5;8)	Chr5	U(132 - 170)	U(64-208) <sup>D</sup> , D(785-877, 972-1000) <sup>E</sup>	5'LIFR(I)	4
		Chr8	U(466 - 493 ) , D(837 - 864 )	D(832-857) <sup>E</sup>	3'PLAG1(I)	10
(9)	t(12;3)	Chr12	U(50 - 77)	U(52-357) <sup>E</sup> , D(763-791) <sup>E</sup>	3'HMGA2(I)	9
		Chr3	—	—	5'FHIT(I)	6
(10a)	t(12;9)	Chr12	U(16 - 43)	U(20-325) <sup>E</sup> , D(733-761) <sup>E</sup>	5'HMGA2(I)	10
		Chr9	—	U(348-375) <sup>E</sup>	3'NFIB(I)	10
(10b)	t(12;9)	Chr12	U(108 - 146 )	D(904-947) <sup>S</sup>	5'HMGA2(I)	8
		Chr9	—	U(348-375) <sup>E</sup>	3'NFIB(I)	12
(11)	t(9;15)	Chr9	U(364 - 402 )	—	5'BRD3(I)	1
		Chr15	U(49 - 87 ) , D(624 - 662 ,844 - 882 )	U(1-112 , 128-407) <sup>E</sup>	3'NUTM1(I)	3
(12b)	t(22;21)	Chr22	U(222 - 249,305 - 332, 356 - 383)	U(1-264) <sup>E</sup> , D(754-1000) <sup>E</sup>	5'EWSR1(I)	6
		Chr21	U(5 - 43 , 313 - 340 ) , D(874 - 912 )	U(1-255) <sup>E</sup>	3'ERG(I)	6
(13a)	t(22;11)	Chr22	U(43 - 70,126 - 153 ,	U(1-77) <sup>E</sup> ,	5'EWSR1(I)	4

Sr no.	Translocations	Chr. partners	Location of RSS	Location of Repeats	Partner Gene [5' / 3' ends] & Intronic(I)/ Exonic (E) Break	# of SIDD sites [Threshold cut off of G(x) <4]
			177 - 204), D(883 - 910 )	D(565-1000) <sup>€</sup>		
		Chr11	—	U(68-287) <sup>€</sup>	3'FLI1(I)	8
(13b)	t(22;11)	Chr22	U(41 - 68 , 124 - 151,175 - 202) ,D(881 - 908)	U(1-73) <sup>€</sup> , D(562-1000) <sup>€</sup>	5'EWSR1(I)	4
		Chr11	—	U(75-378) <sup>¥</sup> , D(733-929) <sup>€</sup>	3'FLI1(I)	7

Note: W: within, U: upstream, D: downstream of the breakpoint. €=Sine, ¥ = Line, £= Low complexity repeat, T=Tandem repeats, D = DNA Repeat, S= simple repeat

Table S3. Analysis of physico-chemical parameters/AT% within destabilized region (SIDD) & 1000 bps sequence around breakpoint.

Serial No.	Translocation	Chromosome Partner	Partner Gene [5' / 3' ends]	Base pair Location	Flexibility (kJ / mol A <sup>3</sup> )	Melting Temperature (°C)	Stacking Energy (kcal/mol)	AT (%)
					Expected Values			
					7.931	73.622	-8.080	
(1)	t(2;2)	Chr2	EML4	1..1001	8.602	66.943	-7.297	64.1
				495..600	9.001	55.519	-6.136	69.5
		Chr2	ALK	1..1001	7.766	73.333	-7.958	48.5
				663..803	9.458	62.461	-6.760	57.9
(2)	t(10;10)	Chr10	CCDC6	1..1001	8.544	69.180	-7.502	59.3
				654..813	9.433	62.831	-6.794	65.3
		Chr10	RET	1..1001	7.042	77.745	-8.459	38.7
				418..454	8.459	74.581	-8.344	74.1
(3)	t(12;15)	Chr12	5'ETV6	1..1001	8.189	71.104	-7.756	55.8
				488..623	9.121	67.538	-7.341	65.7
		Chr15	3'NTRK3	1..1001	7.925	72.920	-7.916	53.7
				425..599	7.924	74.195	-8.133	62.5
(4)	t(5;6)	Chr5	5'CD74	1..1001	7.542	75.690	-8.179	45.8
				523..622	7.888	72.598	-7.945	53.9
		Chr6	3'ROS1	1..1001	9.023	65.170	-7.046	71.4
				443..549	9.920	60.957	-6.593	82.5

Serial No.	Translocation	Chromosome Partner	Partner Gene [5' / 3' ends]	Base pair Location	Flexibility (kJ/mol A <sup>2</sup> )	Melting Temperature (°C)	Stacking Energy (kcal/mol)	AT (%)
					Expected Values			
					7.931	73.622	-8.080	
(5a)	t(21;21)	Chr21	5'TMPRSS2	1..1001	7.813	72.936	-7.929	35.2
				539..670	8.391	70.153	-7.687	48.4
		Chr21	3'ERG	1..1001	7.834	73.090	-7.981	54.3
				486..620	8.584	70.674	-7.745	65.3
(5b)	t(21;21)	Chr21	5'TMPRSS2	1..1001	7.829	73.131	-7.928	53.1
				440..672	9.037	66.783	-7.250	67.8
		Chr21	3'ERG	1..1001	8.327	71.554	-7.786	56.3
				490..710	9.129	67.834	-7.352	65.2
(6)	t(15;7)	Chr15	5'HMGN2P46	1..1001	8.345	70.454	-7.640	60.5
				451..597	8.505	69.878	-7.615	63.6
		Chr7	3'ETV1	1..1001	9.026	66.615	-7.199	65.8
				459..598	9.274	67.475	-7.248	76.3
(7)	t(1;X)	Chr1	5'SFPQ	1..1001	8.653	68.140	-7.388	63.6
				324..471	9.373	65.499	-7.033	69.3
		ChrX	3'TFE3	1..1001	7.863	73.596	-7.958	49.2
				519..596	8.008	73.803	-7.934	61.6
(8)	t(5;8)	Chr5	5'LIFR	1..1001	8.208	71.100	-7.734	55.6
				338..486	8.543	71.271	-7.652	66.1
		Chr8	3'PLAG1	1..1001	8.886	66.034	-7.165	66.2
				413..580	9.117	67.847	-7.343	73.4
(9)	t(12;3)	Chr12	3'HMGA2	1..1001	8.805	67.061	-7.265	64.1
				385..579	9.667	63.274	-6.804	74.3
		Chr3	5'FHIT	1..1001	8.518	68.835	-7.523	60.2
				472..626	8.655	69.445	-7.599	71.4
(10a)	t(12;9)	Chr12	5'HMGA2	1..1001	8.776	67.293	-7.293	64.4
				408..606	9.722	63.428	-6.807	73.7
		Chr9	3'NFIB	1..1001	9.166	65.848	-7.123	67.3
				300..546	9.319	68.225	-7.302	74.7

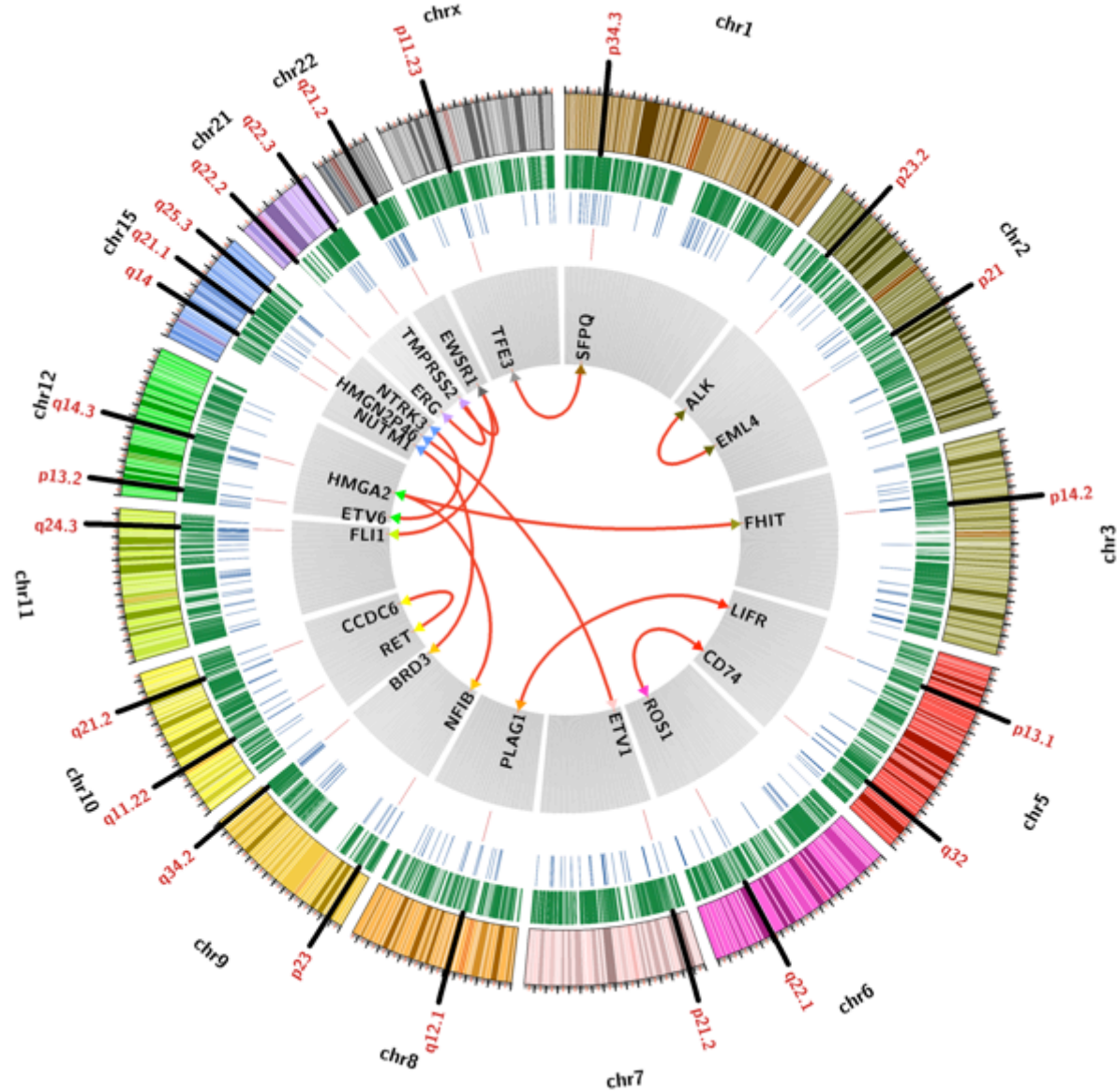
Serial No.	Translocation	Chromosome Partner	Partner Gene [5' / 3' ends]	Base pair Location	Flexibility (kJ/mol A <sup>2</sup> )	Melting Temperature (°C)	Stacking Energy (kcal/mol)	AT (%)
					Expected Values			
					7.931	73.622	-8.080	
(10b)	t(12;9)	Chr12	5'HMGA2	1..1001	9.379	65.759	-7.116	68.3
				414..548	9.606	66.236	-7.106	79.1
		Chr9	3'NFIB	1..1001	9.153	65.944	-7.135	67.1
				490..716	9.264	64.234	-6.959	70.3
(11)	t(9;15)	Chr9	5'BRD3	1..1001	7.420	76.818	-8.321	43.2
				493..600	8.882	69.527	-7.585	61.4
		Chr15	3'NUTM1	1..1001	7.908	73.850	-7.999	49.5
				385..578	9.219	68.324	-7.348	63.4
(12)	t(22;21)	Chr22	5'EWSR1	1..1001	8.200	71.633	-7.787	55.1
				389..514	9.382	67.398	-7.336	65.3
		Chr21	3'ERG	1..1001	8.331	70.033	-7.629	57.4
				419..541	8.950	68.334	-7.434	64.6
(13a)	t(22;11)	Chr22	5'EWSR1	1..1001	8.245	71.199	-7.736	55.2
				489..611	9.265	67.670	-7.308	66.7
		Chr11	3'FLI1	1..1001	8.765	68.196	-7.410	61.3
				472..601	8.863	67.694	-7.311	63.8
(13b)	t(22;11)	Chr22	5'EWSR1	1..1001	8.261	71.150	-7.729	55.1
				486..611	9.173	67.639	-7.305	64.5
		Chr11	3'FLI1	1..1001	8.560	69.460	-7.545	58.3
				659..817	9.312	65.801	-7.163	68.2

Table S4. Analysis &amp; characterization of various parameters of housekeeping gene GAPDH taken as control.

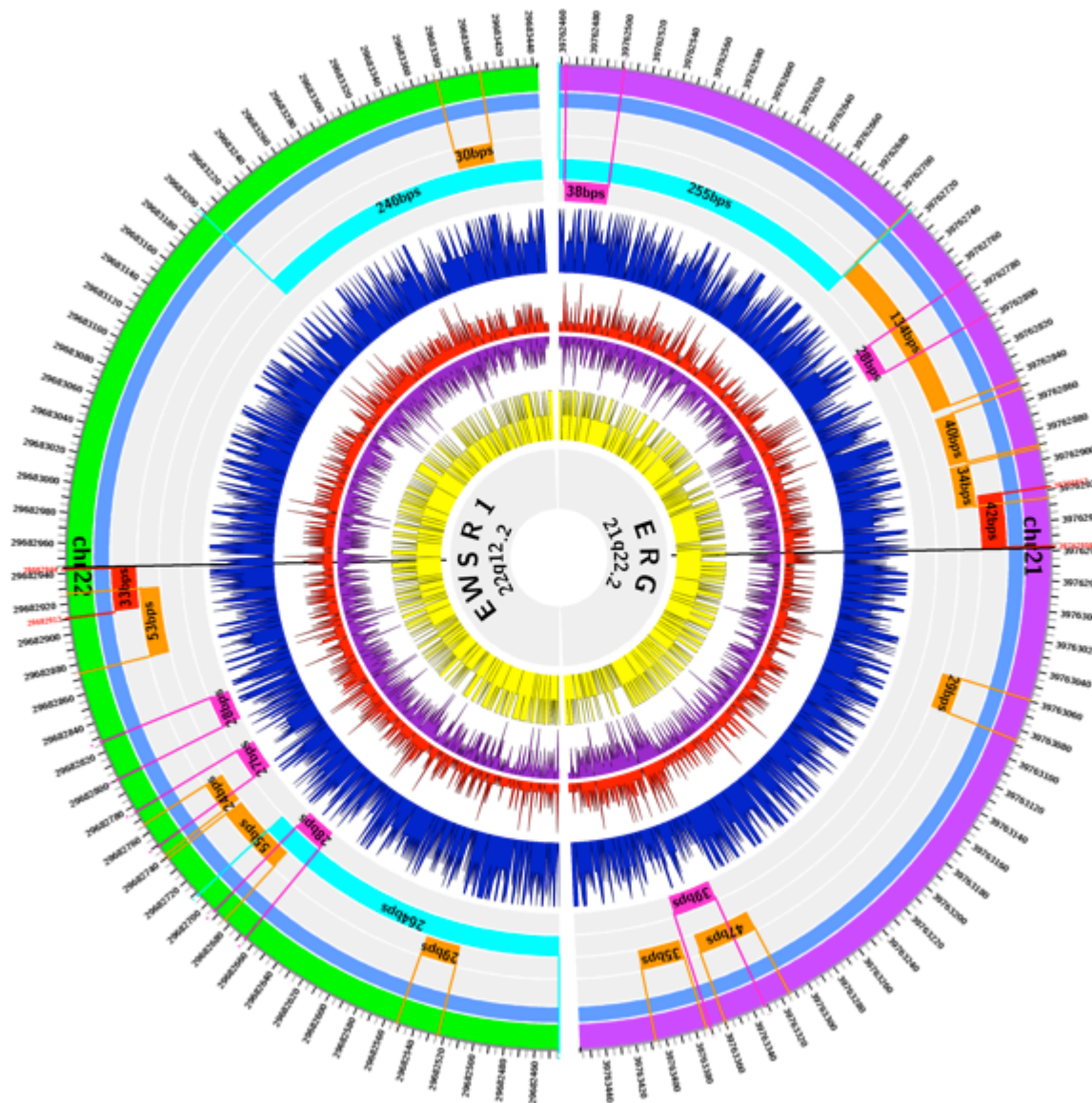
Sr No.	Genomic location of GAPDH Gene(12p13.31) (feb 2009/hg19)	Length of exonic sequences (Bps)	Genomic location of RSS (feb 2009/hg19)	Genomic Location of Repeats (feb 2009/hg19)	# of SIDD sites [Threshold cut off of G(x) <4]	Flexibility (kJ/mol A <sup>2</sup> )	Melting Temperature (°C)	Stacking Energy (kcl/mol)	AT (%)
						Cutoff Values (ROC)			
						7.9 Sp=100 Sn=96.7	73.6 Sp =100 Sn =100	-8.1 Sp=100 Sn=100	47.5 Sp=100 Sn=96.7
(1)	6643977 – 6644026 (E)	51	0	0	0	7.4	77.3	-8.5	40.0
(2)	6646087 – 6646176 (E)	90	0	0	0	7.3	77.2	-8.5	40.1
(3)	6646268 – 6646382 (E)	115	0	0	0	7.5	73.6	-8.0	47.3
(4)	6646476 – 6646556 (E)	81	0	0	0	7.3	74.2	-8.1	47.5
(5)	6646749 – 6647162 (E)	413	0	0	0	7.2	77.2	-8.4	40.7
(6)	6643927 – 6644026 (I/E)	100	0	0	0	7.1	80.4	-8.85	34.3
(7)	6646037 – 6646136 (I/E)	100	0	0	0	7.5	75.4	-8.15	41.9
(8)	6646218 – 6646317 (I/E)	100	0	0	0	7.3	75.9	-8.25	42.4
(9)	6646426 – 6646525 (I/E)	100	0	0	0	7.1	77.5	-8.4	39.8
(10)	6646701 – 6646800 (I/E)	100	0	0	0	7.3	76.7	-8.3	41.1

**Note:** E: Core Exonic sequence; I/E: Intron exon flanking sequence; Sp: Specificity; Sn: Sensitivity.





**Fig. (S1). Circos plot of chromosomal translocations (Oncogenic Fusion genes) in solid tumors.** Outer most track indicates the total length of the chromosomal coordinates in the clockwise direction (scale 1 = 1000000bps). The second track is denoting the complete set of genes of respective chromosomes. The next (blue bands) shows the total numbers of cancerous genes present within each of these chromosomes and the innermost track (red bands) depicts oncogenic fusion genes involved in the rearrangements which are shown by connecting lines and their cytobands (red) are shown on the outermost circle.



**Fig. (S2). Circos plot of various physico-chemical parameters at BpI for translocation t(21;22).** Sequential/ systematic depiction of all the parameters at or near the breakpoint region that are scrutinized in our study. The outermost trail demarcates the *BpI* sequence of 1000 bps of chr 21 (magenta) and 22 (fluorescent Green) in clockwise direction. While the inner trails demarcate various parameters as mentioned: Intronic region(blue); TICdb fusion sequence (red); SIDD regions(orange); Repeats (turquoise blue); Recombination signal sequence region (RSS-pink); graphical representation of Flexibility (dark blue); Stability(red); Stacking Energy (purple); Percentage of AT content(yellow); name of genes and their locations are mentioned in innermost circle. Horizontal black line indicates the exact breakpoint location and the respective features present at that particular point of break.