

FIG. 1

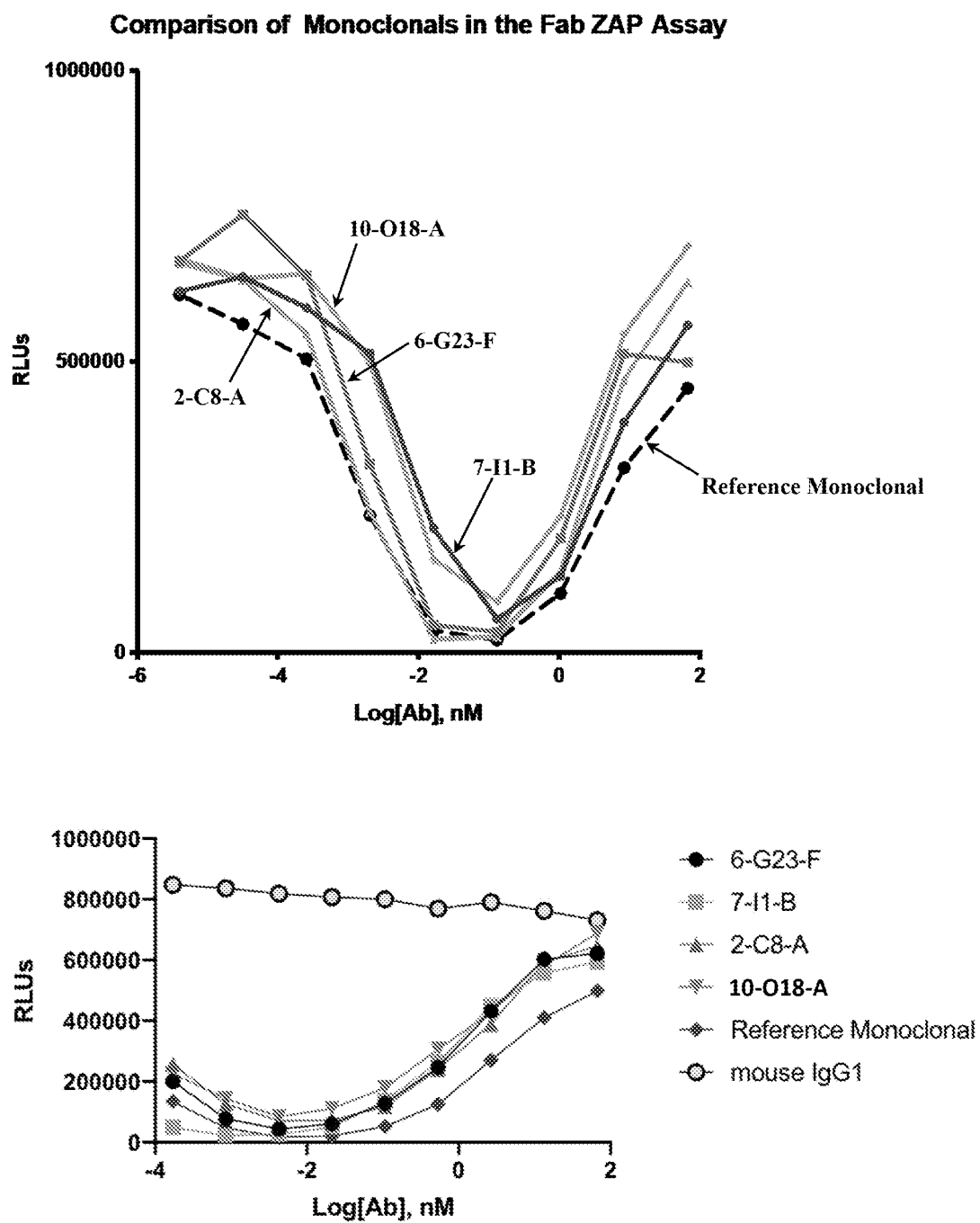


FIG. 2A

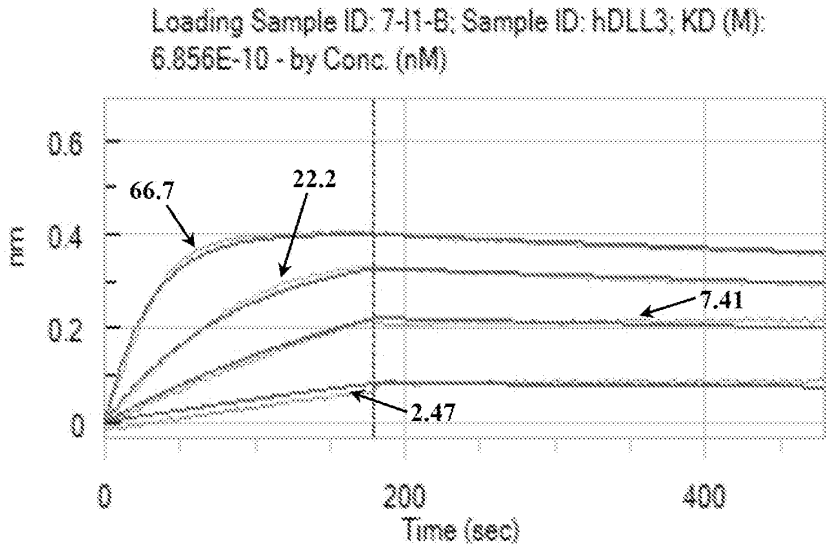


FIG. 2B

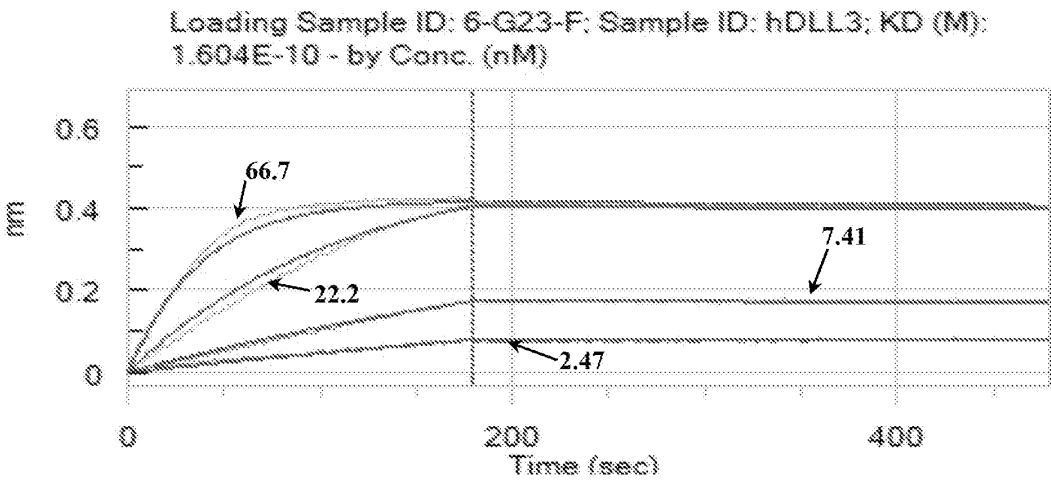


FIG. 2C

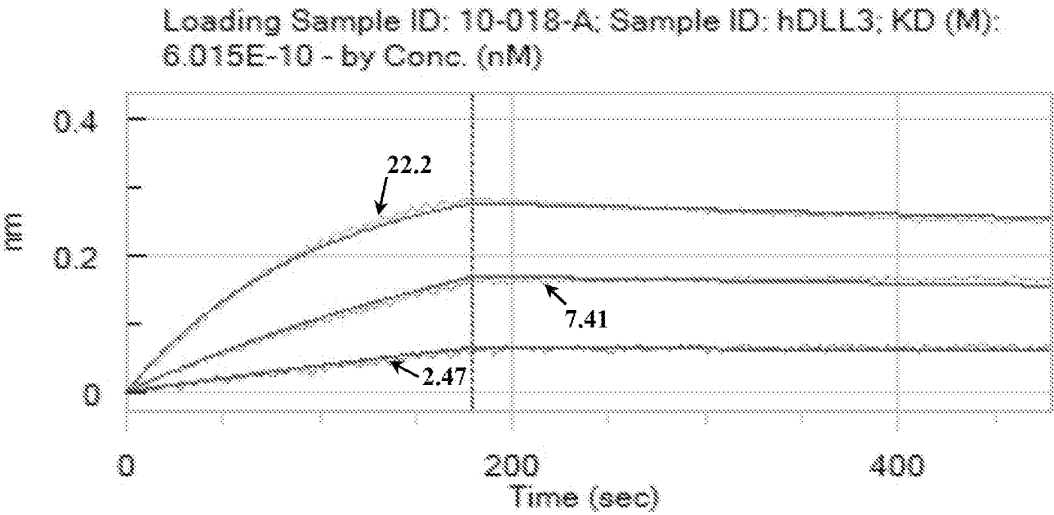
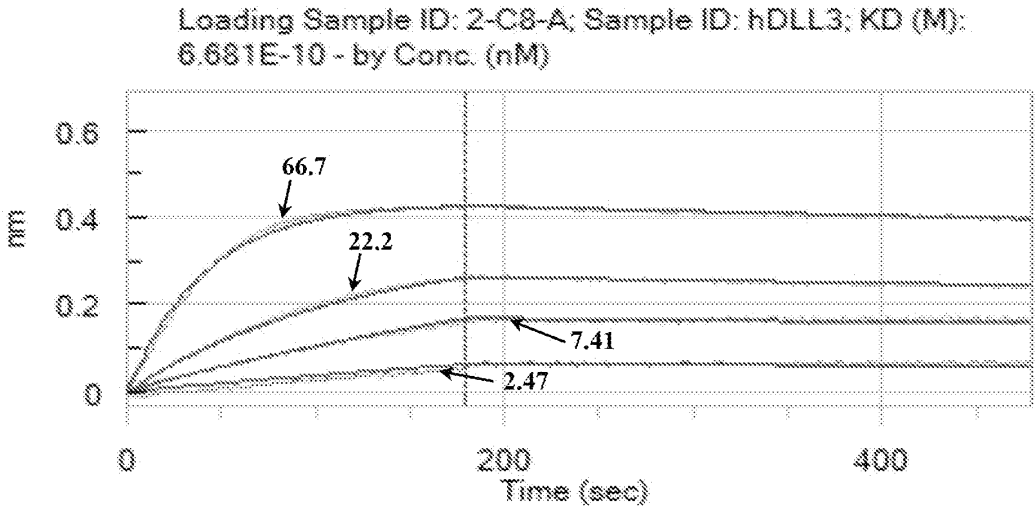


FIG. 2D



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FIG. 2E

	Kd nM	Kon (1/Ms)	Kdis(1/s)	FullX^2	FullR^2
7-I1-B	0.68	5.02E+05	3.44E-04	0.168	0.99
10-O18-A	0.62	4.49E+05	2.67E-04	0.038	0.97
6-G23-F	0.16	4.05E+05	6.49E-05	0.15	0.97
2-C8-A	0.668	3.63E+05	2.42E-04	0.073	0.99

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FIG. 3ANucleotide Sequence of Heavy Chain Variable Domain (V_H) of 7-I1-B (SEQ ID NO: 1)

GAGGTGCAGCTGGTGGAGTCTGGGGGGGGCTTGGTAAAGCCTGGGGGGTCCCTT
AGACTCTCCTGTGCAGCCTCTGGATTCACTTTCAGTAACACCTGGATGAGCTGGG
TCCGCCAGGCTCCAGGGAAGGGGCTGGAGTGGGTTGGCCGTATTAAAAGCAAAT
CTGATGGTGGGACAACAGACTACGCTGCACCCGTGAAAGGCAGATTACCATCT
CAAGAGATGATTCAAAAAACACGCTGTATCTGCAAATGAACAGCCTGAAAACCG
AGGACACAGCCGTGTATTACTGTACCCAGTATTATTGGAACCTCTTTGACTACTG
GGGCCAGGGAACCCTGGTCACCGTCTCCTCA

Amino Acid Sequence of Heavy Chain Variable Domain (V_H) of 7-I1-B (SEQ ID NO: 2)

EVQLVESGGGLVKPGGSLRLSCAAS**GF****TFS****NTW**MSWVRQAPGKGLEWVGRIKSKS
DGGTTDYAAPVKGRFTISRDDSKNTLYLQMNSLKTEDTAVYYCTQYYWNSFDYWGQ
GTLVTVSS

FIG. 3BNucleotide Sequence of Light Chain Variable Domain (V_L) of 7-I1-B (SEQ ID NO: 6)

GACATCCAGATGACCCAGTCTCCATCCTCCCTGTCTGCATCTGTAGGAGACAGAG
TCACCATCACTTGCCAGGCGAGTCAGGACATTAGCAACTATTTAAATTGGTATCA
GCAGAAACCAGGGAAAGCCCCTAAGCTCCTGATCTACGATGCATCCAATTTGGA
AACAGGGGTCCCATCAAGGTTCAGTGGAAGTGGATCTGGGACAGATTTTACTTTC
ACCATCAGCAGCCTGCAGCCTGAAGATATTGCAACATATTACTGTCAACAGTATG
ATAATCTCCCGCTCACTTTCGGCGGAGGGACCAAGGTGGAGATCAAA

Amino Acid Sequence of Light Chain Variable Domain (V_L) of 7-I1-B (SEQ ID NO: 7)

DIQMTQSPSSLSASVGDRVTITC**QASQ****DIS****NYL**NWYQQKPGKAPKLLIYDASNLETG
VPSRFSGSGSGTDFTFTISSLQPEDATYYCQYDNLPLTFGGGTKVEIK

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FIG. 4A

Nucleotide Sequence of Heavy Chain Variable Domain (V_H) of 2-C8-A (SEQ ID NO: 11).

GAGGTGCAGCTGGTGGAGTCTGGGGGAGGCTTGGTCCAGCCTGGGGGGTCCCAG
AGACTCTCCTGTGCAGCCTCTGGATTACCTTTAGTAGCTATTGGATGAACTGGG
TCCGCCAGGCTCCAGGGAAGGGGCTGGAGTGGGTGGCCAACATAAAGGAAGAT
GGAAGTGAGAAATACTATGTGGACTCTGTGAAGGGCCGATTACCATCTCCAGA
GACAACGCCAAGAAGCTACTGTATCTGCAATGAACAGCCTGAGAGCCGAGGAC
ACGGCTGTGTATTACTGTGCGAGAGATCCGGGCTGGGCTCCCTTTGACTACTGGG
GCCAGGGAACCCTGGTCACCGTCTCCTCA

Amino Acid Sequence of Heavy Chain Variable Domain (V_H) of 2-C8-A (SEQ ID NO: 12).

EVQLVESGGGLVQPGGSQLSCAAS**GF****TFSS**YWMNWVRQAPGKGLEWVANIKEDG
SEKYYVDSVKGRFTISRDNAKNSLYLQMNSLRAEDTAVYYCARDPGWAPFDYWGQ
GTLVTVSS

FIG. 4B

Nucleotide Sequence of Light Chain Variable Domain (V_L) of 2-C8-A (SEQ ID NO: 16).

GACATCCAGATGTCCCAGTCTCCATCCTCACTGTCTGCATCTGTAGGAGACAGAG
TCACCATCACTTGTCTGGGCGAGTCAGGGCATTAGCAATTATTTAGCCTGGTTTCA
GCAGAAACCAGGGAAAGCCCCTAAGTCCCTGATCTATGCTGCATCCAGTTTGCA
AAGTGGGGTCCCATCAAAGTTCAGCGGCAGTGGATCTGGGACAGATTTCACTCTC
GCCATCAGCAGCCTGCAGCCTGAAGATTTTGCAACTTATTACTGCCAACAGTATA
ATAGTTTCCCGTACACTTTTGGCCAGGGGACCACGCTGGAGATCAAA

Amino Acid Sequence of Light Chain Variable Domain (V_L) of 2-C8-A (SEQ ID NO: 17).

DIQMSQSPSSLSASVGDRVTITCRASQGISNYLAWFQQKPGKAPKSLIYAASSLQSGV
PSKFSGSGSGTDFTLAISLQPEDFATYYCQQYNSFPYTFGQGTLEIK

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FIG. 5A

Nucleotide Sequence of Heavy Chain Variable Domain (V_H) of 10-O18-A (SEQ ID NO: 21).

CAGGTGCAGCTGCAGGAGTCGGGCCCAGGACTGGTGAAGCCTTCGGAGACCCTG
TCCCTCACCTGCACTGTCTCTGGTGGCTCCATCAATAGTTACTACTGGAGCTGGA
TCCGGCAGCCCCCAGGGAAGGGACTGGAGTGGATTGGGTATATCTTTTACAGTG
GGATCACCAACTACAACCCCTCCCTCAAGAGTCGAGTCACCATATCATTAGACAC
GTCCAAGAACCAGTTCTCCCTGAAGCTGAGCTCTGTGACCGCTGCGGACACGGCC
GTGTATTACTGTGCGAGAATCGGCGTGGCTGGTTTTTACTTTGACTACTGGGGCC
AGGGAACCCTGGTCACCGTCTCCTCA

Amino Acid Sequence of Heavy Chain Variable Domain (V_H) of 10-O18-A (SEQ ID NO: 22).

QVQLQESGPGLVKPSSETLSLTCTV**SGGSINSYY**WSWIRQPPGKGLEWIGYIFYSGITN
YNPSLKS RVTISLDTSKNQFSLKLSSVTAADTAVYYCAR**IGVAGFYFDY**WGQGTTLVT
VSS

FIG. 5B

Nucleotide Sequence of Light Chain Variable Domain (V_L) of 10-O18-A (SEQ ID NO: 26).

GAAATTGTGTTGACGCAGTCTCCAGGCACCCTGTCTTTGTCTCCAGGGGAAAGAG
CCACCCTCTCCTGCAGGGCCAGTCAGAGTGTTAGCAGCAGCTACTTAGCCTGGTA
CCAGCAGAAACCTGGCCAGGCTCCAGGCTCCTCATCTATGGTGCATCCAGCAG
GGCCACTGGCATCCCAGACAGGTTCAAGTGGCAGTGGGTCTGGGACAGACTTCAC
TCTCACCATCAGCAGACTGGAGCCTGAAGATTTTGCAGTGTATTACTGTCAGCAG
TATGGTACCTCACCGCTCACTTTCGGCGGAGGGACCAAGGTGGAGATCAAA

Amino Acid Sequence of Light Chain Variable Domain (V_L) of 10-O18-A (SEQ ID NO: 27).

EIVLTQSPGTLSPGERATLSC**RASQSVSSSYL**AWYQQKPGQAPRLLIY**GASSRATG**
IPDRFSGSGSGTDFTLTISRLEPEDFAVYYC**QOYGTSP**LTFGGGTKVEIK

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FIG. 6A

Nucleotide Sequence of Heavy Chain Variable Domain (V_H) of 6-G23-F (SEQ ID NO: 31).

CAGGTGCAGCTGGTGCAGTCTGGGGCTGAGGTGAAGAAGCCTGGGGCCTCAGTG
AAGGTTTCCTGCAAGGCATCTGGATACACCTTCACCAGCTACTATATACACTGGG
TGCGACAGGCCCCCTGGACAAGGGCTTGAGTGGATGGGAATAATCGACCCAAGTG
ATGGTAGCACAACTACGCACAGAAAGTTCCAGGGCAGAGTCACCATGACCAGGG
ACACGTCCACGAGCACAGTCTACATGGAGCTGAGCAGCCTGAGATCTGAGGACA
CGGCCGTGTATTACTGTGCGAGAGATCGGGAATATAACTACTACGGTTTGGACGT
CTGGGGCCAAGGGACCACGGTCACCGTCTCCTCA

Amino Acid Sequence of Heavy Chain Variable Domain (V_H) of 6-G23-F (SEQ ID NO: 32).

QVQLVQSGAEVKKPGASVKVSCKASGYTFTSYIHWVRQAPGQGLEWMGHI~~DPSD~~
GSTNYAQKFQGRVTMTRDTSTSTVYMELSSLRSEDTAVYYCARDREYNYGLDIWG
QGTTVTVSS

FIG. 6B

Nucleotide Sequence of Light Chain Variable Domain (V_L) of 6-G23-F (SEQ ID NO: 36).

GATGTTGTGATGACTCAGTCTCCACTCTCCCTGCCCGTCACCCTTGGACAGCCGG
CCTCCATCTCCTGCAGGTCTAGTCAAAGCCTCGTATACCGTGATGGAAACACCTA
CTTGAATTGTTTTAGCAGAGGCCAGGCCAATCTCCAAGGCGCCTAATTTATAAG
GTTTCTAACCGGGACTCTGGGGTCCCAGACAGATTCCGCGGCAGTGGGTCAGGC
ACTGATTTCACTGAAAATCAGCCGGGTGGAGGCTGAGGATGTTGGGGTTTATT
ACTGCATGCAAGGTACACACTGGCCTCCGACGTTTCGGCCAAGGGACCAAGGTGG
AAATCAAA

Amino Acid Sequence of Light Chain Variable Domain (V_L) of 6-G23-F (SEQ ID NO: 37).

DVVMTQSPLSLPVTLGQPASISCRSSQSLVYRDGNTYLNWFQQRPGQSPRRLIYKVS
NRDSGVDPDRFRGSGSGTDFTLKISRVEAEDVGVYYCMOGTHWPPTFGQGTKVEIK

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FIG. 7

Monoclonal	DLL3	DLL4	DLL1
10-O18-A	1.361	0.056	0.126
2-C8-A	1.359	0.07	0.098
6-G23-F	1.459	0.05	0.206
7-I1-B	1.302	0.96	0.127
Background (anti-mouse IgG HRPO)	0.085	0.085	0.095

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FIG. 8A

>NM_016941.4 Homo sapiens delta like canonical Notch ligand 3 (DLL3), transcript variant 1, mRNA (SEQ ID NO: 55)

ACTCCCCGAGACCCCCACCAGAAGGCCATGGTCTCCCCACGGATGTCCGGGCTCCTCTCCCAGACTGTG
ATCCTAGCGCTCATTTTCTCCCCAGACACGGCCCGCTGGCGTCTTCGAGCTGCAGATCCACTCTTTTCG
GGCCGGGTCCAGGCCCTGGGGCCCCCGGCTCCCCCTGCAGCGCCCGGCTCCCCTGCCGCTCTTCTTCAG
AGTCTGCCTGAAGCCTGGGCTCTCAGAGGAGGCCGCCGAGTCCCCGTGCGCCCTGGGCGCGGCGCTGAGT
GCGCGCGGACCGGTCTACACCGAGCAGCCCGGAGCGCCCGCGCTGATCTCCCACTGCCCGACGGCCTCT
TGCAGGTGCCCTTCCGGGACGCTGGCCTGGCACCTTCTCTTTTCATCATCGAAACCTGGAGAGAGGAGTT
AGGAGACCAGATTGGAGGGCCCCGCTGGAGCCTGCTGGCGCGCTGGCTGGCAGGCGGCGCTTGGCAGCC
GGAGGCCCGTGGGCCCCGGGACATTACGCGCGCAGGCGCCTGGGAGCTGCGCTTCTCGTACCGCGCGCGCT
GCCAGCCGCTGCCGTCCGGACCGCGTGCACGCGCCTTGCCTGCCGCGCAGCGCCCCCTCGCGGTGCGG
TCCGGGACTGCCCCCTGCCACCGCTCGAGGACGAATGTGAGGCGCGCTGGTGTGCCGAGCAGGCTGC
AGCCCTGAGCATGGCTTCTGTGAACAGCCCGGTGAATGCCGATGCCCTAGAGGGCTGGACTGGACCCCTCT
GCACGGTCCCTGTCTCCACCGACGAGTGCCTCAGCCCCAGGGGCCCGTCTCTGCTACCAACGGGATGCCCT
TGTCCTGGGCTGGGCCCTGTGACGGGAACCCGTGTGCCAATGGAGGCAGCTGTAGTGAGACACCCAGG
TCCTTTGAATGCACCTGCCCGCGTGGGTCTACGGGCTGCGGTGTGAGGTGAGCGGGGTGACATGTGCAG
ATGGACCTGCTTCAACGGCGGCTTGTGTGTCGGGGGTGCAGACCTGACTCTGCCTACATCTGCCACTG
CCCACCCGTTTCCAAGGCTCCAACCTGTGAGAAGAGGGTGGACCGGTGCAGCCTGCAGCCATGCCGCAAT
GGCGGACTCTGCCTGGACCTGGGCCACGCCCTGCGCTGCGCTGCCGCGCGCGCTTCCGGGCTCCTCGCT
GCCAGCACGACTGGACGACTGCGCGGGCGCGCCTGCGCTAACGGCGGCACGTGTGTGGAGGGCGGCGG
CGCGCACCGCTGCTCCTGCGCGCTGGGCTTCCGGCGCGCGACTGCCGCGAGCGCGCGGACCCGTGCGCC
GCGCGCCCTGTGCTCACGGCGGCGCGCTGTACGCCCACTTCTCCGGCTCGTCTGCGCTTGGCTCCCG
GCTACATGGGAGCGCGGTGTGAGTTCCCACTGCACCCCGACGGCGCAAGCGCCTTCCCCGCGGCCCGCC
GGGCTCAGGCCCGGGGACCCCTCAGCGCTACCTTTTGCTCCGGCTCTGGGACTGCTCGTGGCCGCGGGC
GTGGCCGGCGCTGCGCTCTTGTGCTCCAGTGCAGCGCCCGTGGCCACTCCCAGGATGCTGGGTCTCGCT
TGCTGGCTGGGACCCCGAGCGCTCAGTCCACGCACTCCCGGATGCACTCAACAACCTAAGGACGCAAGGA
GGGTTCGGGGATGGTCCGAGCTCGTCCGTAGATTGGAATCGCCCTGAAGATGTAGACCCCTCAAGGGATT
TATGTCATATCTGCTCCTTCCATCTACGCTCGGGAGGTAGCGACGCCCTTTTCCCCCGCTACACACTG
GGCGCGCTGGGCGAGGCGACCTGCTTTTCCCTACCCCTTCTCGATTCTGTCCGTGAAATGAATTGG
GTAGAGTCTCTGGAAGGTTTTAAGCCATTTCAGTTCTAACTTACTTTTATCTTATTTGTCATCCCTCT
TATCGTTTTTGTAGCTACCTGCCATCTTCTTTGAAAAAOCATATGGGCTTGAGGAGGTACAGATGCCGACT
CCGCCAGAGCTTTTCCACTGATTGTACTCAGCGGGGAGGCAGGGGAGGCAGAGGGGCAGCCTCTCTAATG
CTTCTACTCATTTTGTCTTCTAGGCCTGACGCGTCTCTCCATCCGCACCTGGAGTCAGAGCGTGGATTT
TTGATTTTGTGCGGTGGTGGCCAGTCTCTGCCCCAGAGGCTTTGGAGTTCAATCTTGAAGGGGTGCTGG
GGGAACCTTACTGTTGCAAGTTGTAAATAATGGTTATTTATATCTATTTTTTCTCACCCCATCTCTCTA
GAAACACCTATAAAGGCTATTATTGTGATCA

>NP_058637.1 delta-like protein 3 isoform 1 precursor [Homo sapiens] (SEQ ID NO: 50)

MVSPRMSGLLSQTIVILALIFLPQTRPAGVFELQIHSEFGPGPGAPRSPCSARLPCRLFFRVCLKPGLSE
EAAESPCALGAALSARGPVYTEQPGAPAPDLPLPDGLLQVPPFRDAWPGTFSFIIETWREELGDQIGGPAW
SLLARVAGRRRLAAGGPWARDIQRAGAWELRFSYRARCPEPAVGTACTRLCRPRSAFRCGPGLRFPAPL
EDECEAPLVCRAGCSPEHGFCEQPGECRCLEGTGPLCTVPVSTSSCLSPRGPSATGCLVPGPFGCDG
NPCANGGSCSETPRSFECTCPRGFYGLRCEVSGVTCADGPCFNGGLCVGGADPDSAYICHCPPGFQGSNC
EKRVDRCSLQPCRNGGLCLDLGHALRCRCRAGFAGPRCEHDLDDCAGRACANGGTCVEGGGAHRCSCALG
FGGRDCRERADPCAARPCAAGGRCAHAFSGLVACAPGYMGARCEFPVHPDGASALPAAPPGLRPGDPQR
YLLFPALGLLVAGVAGAAALLLVHVRRRGHSQDAGSRLLAGTPEPSVHALPDALNNLRTQEGSGDGPSSS
VDWNRPEDVDPQGIYVISAPSIYAREVATPLFPPLHTGRAGQRQHLLFPYPSSILSVK

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FIG. 8B

>NM_203486.3 Homo sapiens delta like canonical Notch ligand 3 (DLL3), transcript variant 2, mRNA (SEQ ID NO: 56)

ACTCCCCGAGACCCCCACCAGAAGGCCATGGTCTCCCCACGGATGTCCGGGCTCCTCTCCCAGACTGTG
ATCCTAGCGCTCATTTTCCTCCCCAGACACGGCCCGCTGGCGTCTTCGAGCTGCAGATCCACTCTTTTCG
GGCCGGSTCCAGGCCCTGGGGCCCCGGTCCCCCTGCAGCGCCCGGCTCCCCCTGCCGCCCTCTTCTCAG
AGTCTGCCCTGAAGCCTGGGCTCTCAGAGGAGGCCGCCGAGTCCCCGTGCGCCCTGGGCGCGGCGCTGAGT
GCGCGCGGACCGGTCTACACCGAGCAGCCCGAGCGCCCGCGCTGATCTCCCACTGCCCGACGGCCTCT
TGCAGGTGCCCTTCCGGGACGCCTGGCCTGGCACCTTCTCTTTTCATCATCGAAACCTGGAGAGAGGAGTT
AGGAGACCAGATTGGAGGGCCCCGCCTGGAGCCTGCTGGCGCGCGTGGCTGGCAGGCGGCGCTTGGCAGCC
GGAGGCCCGTGGGCCCCGGACATTACAGCGCGCAGGCGCCTGGGAGCTGCGCTTCTCGTACCGCGCGCGCT
GGAGCCCGCTGCCGTGGGACCGCGTGCACGCGCCTCTGCCGTCCGCGCAGCGCCCCCTCGCGGTGCGG
TCCGGGACTGCGCCCCCTGCGCACCGCTCGAGGACGAATGTGAGGCGCCGCTGGTGTGCCGAGCAGGCTGC
AGCCCTGAGCATGGCTTCTGTGAACAGCCCGGTGAATGCCGATGCCTAGAGGGCTGGACTGGACCCCTCT
GCACGGTCCCTGTCTCCACCAGCAGTGCCTCAGCCCCAGGGGCCGCTCTCTGCTACCACCGGATGCCT
TGTCCCTGGGCTGGGCCCTGTGACGGGAACCCGCTGTCGAATGGAGGCAGCTGTAGTGAGACACCCAGG
TCCTTTGAATGCACCTGCCCGCGTGGGTTCTACGGGCTGCGGTGTGAGGTGAGCGGGGTGACATGTGCAG
ATGGACCCCTGCTTCAACGGCGGCTTGTGTGTCGGGGGTGCAGACCTGACTCTGCCTACATCTGCCACTG
CCCACCCGGTTTCCAAGGCTCCAACCTGTGAGAAGAGGGTGGACCGGTGCAGCCTGCAGCCATGCCGCAAT
GGCGGACTCTGCCTGGACCTGGGCCACGCCCTGCGCTGCGCGTGCAGCGCCGGCTTCGCGGGTCTCTCGCT
GCGAGCACGACCTGGACGACTGCGCGGGCCGCGCCTGCGCTAACGCGCGCACGTGTGTGGAGGGCGGCGG
CGCGCACCGCTGCTCCTGCGCGCTGGGCTTCGGCGGCGCGGACTGCCGCGAGCGCGCGGACCCGTGCGCC
GCGCGCCCCCTGTGCTCACGGCGGCGCTGCTACGCCCACTTCTCCGGCCTCGTCTGCGCTTGGCGCTCCCG
GCTACATGGGAGCGCGGTGTGAGTTCCAGTGCACCCGACGGCGCAAGCGCCTTGCCCGCGGCCCCCGCC
GGGCTCAGGCCCGGGGACCCCTCAGCGCTACCTTTTGCCTCCGGCTCTGGGACTGCTCGTGGCGCGGGG
GTGGCCGGCGCTGCGCTCTTGTGTGTCACGTGCGCCGCCGTGGCCACTCCCAGGATGCTGGGTCTCGCT
TGCTGGCTGGGACCCCGAGCCGTCAGTCCACGCACTCCCGGATGCACTCAACAACCTAAGGACGCGAGGA
GGGTTCGGGGATGGTCCGAGCTCGTCCGTAGATTGGAATCGCCCTGAAGATGTAGACCCFCAAGGGATT
TATGTATATCTGCTCCTTCCATCTACGCTCGGGAGGCCTGACGCGTCTCTCCATCCGCACCTGGAGTC
AGAGCGTGGATTTTTGTATTTGCTCGGTGGTGCCAGTCTCTGCCCGAGAGGCTTTGGAGTTCAATCTTG
AAGGGGTGCTGGGGGAACCTTTACTGTTGCAAGTTGTAAATAATGGTTATTTATATCTATTTTTTCTCA
CCCCATCTCTCTAGAAACACCTATAAAGGCTATTATTGTGATCA

>NP_982353.1 delta-like protein 3 isoform 2 precursor [Homo sapiens] (SEQ ID NO: 51)

MVSPRMSGLLSQTIVILALIFLPQTRPAGVFELQIHSFGPGPGAPRSPCSARLPCLRFFRVCLKPLGLSE
EAAESPCALGAALSARGPVYTEQPGAPAPDLPLPDGLLQVPFRDAWPGTFSFIETWREELGDQIGGPWW
SLLARVAGRRRLAAGGPWARDIQRAGAWELRFSYRARCPEPAVGTACTRLCRPRSAPSRCPGLRPCAPL
EDECEAPLVCRAGCSPEHGFCEQPGECRCLEGTGPLCTVPVSTSSCLSPRGPSSATTGCLVPGPGPCDG
NPCANGGSCSETPRSFECTCPRGFYGLRCEVSGVTCADGPCFNGGLCVGGADPDSAYICHCPPGFQGSNC
EKRVDRCSLQPCRNGLCLDLGHALRCRCRAGFAGPRCEHDLDDCAGRACANGGTCVEGGGAHRCSCALG
FGGRDRCRERADPCAARPCAHHGRCYAHFSGLVACAPGYMGARCEFPVHPDGASALPAAPPGLRPGDPQR
YLLPPALGLLVAAGVAGAALLLVHVRRRGHSQDAGSRLLAGTPEPSVHALPDALNNLRQTQEGSGDGPSSS
VDWNRPEDVDFQGIYVISAPSIYAREA