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Figure 1A

SNPs genotyped in AVITA and associated with bevacizumab outcome

	SNP_ID	Rare diagnostic allele	SEQUENCE
SEQ ID NO:1	rs9554316	T	ATAGACACAAATTCTAAGAAATCGTAAGACATTCGATTTTTTTTCT[T/G]GAACTCAATG GAATTCTCAGCCCCCAGCCCTTGGTCATGGTCTCCAGCAA
SEQ ID NO:2	rs9582036	C	TGCTGCGATTACAGGCACGAGCCACTGTGCCCAGCAACAATAGCCTTCTT[C/A]CAAAATGT ATACTAAAGTATTTAGGAGTAAAGTGCTATTATGCCTTAAGC
SEQ ID NO:3	rs9513070	G	TTGCTTTTATACTATAATGCAAGCTTGCCCAACTTGTGGCCACGGGCT[G/A]CATGAATCC CAGGATGGCTTTGAGTGAGCCCAACACAACTCGTAAACT
SEQ ID NO:4	rs9554320	A	TTTCCGGGGCTGTGAGCAAGGTTCTGTGTGTAGCTGATCATTTCTCCCCC[A/C]GCACCTGCA AAGCCGCTGTTACAAACTGGCCTCCCACTCAAGGCACAC

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Figure 1B

SNPs belonging to the rs9554316 and rs9582036 haplotype block (within the locus) but not genotyped within AVITA

	SNP_ID	Rare diagnostic allele	SEQUENCE
SEQ ID NO:5	rs17619037	G	AGTGCAAAACATCCACAAGCCCCAGGAAAAAAGTGTCGAAGACAGGCA[A/G]GCAAAGTG CAATTGACACAGTGGAGAAAGCCTCACATGGACACCCGAAAAAG
SEQ ID NO:6	rs9579177	G	ACATTACTGGCGTTGGTGTGTTGGAAGGATCTGAAGAAGGGGTCTATCGG[G/A]GTGCACT AGGTACCTTAACCTTACGACTTACATCAACATGGAGGTGGCA
SEQ ID NO:7	rs9579176	G	CCCTTAGCCCTCAACCCACATCACAGGGCTTTAGAGAGAAGAGCTGG[G/A]AGACCTCT TGTTGGCCAGGCTATACTGGTAGCTTCTCCCATTTGATTG
SEQ ID NO:8	rs9554319	C	CTTCTCCAAAATGTATACTAAAGTATTTAGGAGTAAAGTGCTATTATGC[C/T]TTAAGCCTC CTTTAAATGGTTTGGCAAAAAGAAAAATCCGTAAACAAACA
SEQ ID NO:9	rs7996030	A	CTCCAGAGGGACCTACTCTGGTCCGTTGTGAAATCAGAGAATGCCAGAC[A/G]TAGGAGA GACTGGAGGCATCATCTCGTCTGAGTGCCCAATGGCACAGGAG
SEQ ID NO:10	rs9513075	G	GAGGACACCAAGCAGTGAACAAGAACAGGTCCCTCCCGTCCCTAC[G/A]CTCAGGC AGAGCTCCCCAGCCCTGCTGTGGGCCCTGAGTCTCCTGTCCAC
SEQ ID NO:11	rs7993418	G	TCTTCAAGGTTTGTATCTTCCAGGCTCATGAACCTTGAAGCATTTAC[G/A]TATCTAATGA AGAAACAGAAAGAATTATCAAGACAGGAAAGGCATCCAG
SEQ ID NO:12	rs9513071	G	TTACTTGTCTGGGAGATGGCGGTGCTGGGGAAAAACCTGAGAAACCC[T/G]AGTTTCAT ATCCTCCCTCTGACACTTCTTAGAGGTATGCCTTAGGGGGCG

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Figure 1C

SNPs 5' up or downstream of the haplotype block (outside the locus) and in LD ($D'/\text{LOD} > 80$) with rs9554316 or rs9582036 but not typed in AVITA

	SNP_ID	Rare diagnostic allele	SEQUENCE
SEQ ID NO:13	rs11620238	G	AGGAAGTCATGATCTTTTCTGACCTGGCCTCCAGCCATGCAGTGTCCC[T/G]TCTTTC ACATTGCATTGATCACAAGTGAGTCTCTAAAGCTAGCTCTGATTC
SEQ ID NO:14	rs17618631	G	CTTCAAGTCAATTTGCCTGAATGAATGATTTTATTCTTATGTTATGAAACC[C/G]ATAATAC TTTTTAAAGGATTTCAGATTCTTGAAGACAAAATGCCTCTTTC
SEQ ID NO:15	rs12429309	C	GCAGTGTATGAGGGTTCCAAATTTCTCCATATCCTCGCCACACTGACTTCA[T/C]CTAGCT ATCCTCATGGGCGTAAAGTGGTATCTTATGTTAAATTTCTTCCC
SEQ ID NO:16	rs7982251	C	GTGGCTGTCTGCAGGGACATGCTCTGCATTGAGAGAACACTGGGGAGA[C/T]GACAC GTTTCTCTTTTGTGAACATGCTCCCGTAGTGATTACTAAATGA
SEQ ID NO:17	rs9508016	A	AGGGAGGACATGGGAGGTCTCCACAAAAGTGAAAAATACACTATGTCT[A/G]AAAAAC AAGCAGGGAAGAGAAGCCAGTACGAGCCAGCGGGGAGATGCTAGG
SEQ ID NO:18	rs7982957	A	AGAACACTGGGGAGACGACACGTTTCTCTTTTGTGAAACATGCTCCCGT[A/T]GTGATT ACTAAATGACATTCTCCAGAGCATGCAACACACGGGCAGAGCTA
SEQ ID NO:19	rs3794400	T	AGAAGGACAGAAATACGTTTGAGGAGGGGAGAGGCAAAAGCCAGCCTCCC[T/C]GCCT CTTAACCAGACCCCTTGGGAGTAGAGCGGCTTCTGGTACTGTACACA

Linkage disequilibrium was defined here as having a $D'/\text{LOD} > 80$

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Figure 1D

SNPs linked with r^2 equal or larger than 0.12 to rs9554316 (SEQ ID NO. 1), rs9582036 (SEQ ID NO. 2), rs9513070 (SEQ ID NO. 3) or rs9554320 (SEQ ID NO. 4)

	SNP_ID	Rare diagnostic allele	SEQUENCE
SEQ ID NO: 40	rs45455097	C	TGTGTTTTTAAATTTTTTAAACCAATTTCTTAATACTTTACATTTT/CJATTATTAATATTT ATACTTCTAAGTATTTTCTTAAGGAAAAAGTACTTC
SEQ ID NO: 41	rs1886233	A	TACTGGGAGAAAAATTTATGGAGCATCAACTGGGTAAACTATAATACAAAC[A/G]AACGTTAAG CGTGAAGGGATAGAAACATAGGGCAGTGTGTGTAATAAT
SEQ ID NO: 42	rs9554309	C	TAAATAATTTCATAGAAGTTTCAGTATAGTACGTAATTTCTGTAGAGAGTAT[C/T]GTATGCAAGC TCTCTGTATGCCACCCACTGTTAGTTCAAATTGAGATTTT
SEQ ID NO: 43	rs11619230	T	TGTATGCCACCCACTGTTAGTTCAAATTGAGATTTTGTGTTTGTGTTT[C/T]TTAAGAACATA CCAAGAAATACCAGTGAATTGTTGGATATGAATTCCT
SEQ ID NO: 44	rs9554311	C	AAATTGAGATTTTGTGTTTGTGTTTCTTAAGAACATACCAAGAAAA[A/C/T]CAGTGAATTG TTGGATATGAATTCCTGTTAGTTGATTAAATCCTTTCT
SEQ ID NO: 45	rs4771233	T	GAGCCAGAGTTTGAACAGGCTGGGCAACATAGTGAGACCTTGTCTC[T/C]ACAAAAAG TAAATCAGCCAGGATGGTGTACGCACCTGTAGCCCCAGC
SEQ ID NO: 46	rs6491274	C	GGAGACCTTCACATGGCTCCATGTGGCTTAGGTTTCTCCAAGTATGGC[T/C]GCTGGGTC CCTGGCGGAACTGTTGAAGAGTGAGTGACCAAGACTCAGG
SEQ ID NO: 47	rs7982639	G	TTCAAAAGTCAGGAAAGAGGTAACCAACGCAAGCAAGACAGAGTAGCCT[A/G]TGAGATGG GAAGAAAGAAAGTATCATGCTCTGGAACCAACTGAAGTTGTT
SEQ ID NO: 48	rs12877718	C	ATAAGTGTGTTCTTTAAAGCACITTTAAATAGTTGCTGGCACATATTGGG[T/C]ACTCAAATGA TAGCAATGATAGCAATTACTAGCTATCAGTCTCCTCCCAA
SEQ ID NO: 49	rs10507382	C	TGGGAAAAAGAGGATAATATAGGTATGTTTCAAAACAAATATATTTAG[T/C]CTGAGATTAA GCCCTTCCACCTTTTAAATGGGTTTCTCCTATGAAAAG
SEQ ID NO: 50	rs57354941	T	AACATTAGTAGCATCTGGTTGGATTGGGTTTTTTTTTCCCTCCCT[C/T]AAGTGTCTT AGCTCGCAGATAGAGAATGGCCATCCCTGTGTGGAT
SEQ ID NO: 51	rs17086497	A	AGTTCCAGAGCCATGAGAACATTTCTAACTGACGTGACATTGATTCTT[C/A]AAGCATTCTT CAAGTCATTTGCCTGAATGAATGATTTTATCTTATGTTA
SEQ ID NO: 52	rs3794395	T	TGGCATGGCTACACGCTAAATATTTAATGGCATAATCGAGTGCTAG[C/T]GCCAGGCTT GTATTTTGTGTGTGTGATTACAGATTGGGAAATTGG
SEQ ID NO: 53	rs9554317	T	CCATCTCGGCCTCCAGAGTGTGGGATTACAGGGCTGAGCCACTGTGCC[C/T]GGCCGCAA GCAGTATTTCTGACTGCTGCAGCTTCTTCCCATATCTTCCC
SEQ ID NO: 54	rs9513073	G	TATGTAAGACTTGAAGGTTTGAGAAAGTCCAAAGCAGTTTGGTCCAA[A/G]GCTGATGTG ACTTTGTACCTCTGAGTTATTTGGGACCACAGCCTGGGTAT

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SNPs linked with r^2 equal or larger than 0.12 to rs9554316 (SEQ ID NO. 1), rs9582036 (SEQ ID NO. 2), rs9513070 (SEQ ID NO. 3) or rs9554320 (SEQ ID NO. 4)

	SNP_ID	Rare diagnostic allele	SEQUENCE
SEQ ID NO: 55	rs9513074	T	ATGCTTAAATAGTGGCACTTATAAGCACCAATAACTATTTAATCAGTA[C/T]TATTCAATAC TGTTATAATAAACTAAATTCAGGCCAGGCATGGAGGCT
SEQ ID NO: 56	rs9508015	T	TAGTGGCACTTATAAGCACCAATAACTATTTAATCAGTACTATTCAATA[C/T]TGTATAATAA AAACTAATTCAGGCCAGGCATGGAGGCTCAGGCTGTGA
SEQ ID NO: 57	rs2011950	G	CTGCAATCTCCGCTTCATAGGCTCAAGTGATTCTCATGCCCTCAGCCTCCC[A/G]AGTAGCTGC CACCAGGCCGGCTAACTTTGTATTTTAGTAGAGCGG
SEQ ID NO: 58	rs9513110	G	AACAGATCTTTGTTTAGAAAGAAAGAAAGACATCCAGAAACCAACCAGG[T/G]GGGACTCTAG ACACCCATGGAGGAGGTGATCAGAGCTCATGAGAGTACCAG
SEQ ID NO: 59	rs9513112	A	TAGCTCTATAGTAGGAGAAATATCCTTTAAGTTTCATTAAAGGTTCTCAG[G/A]TCTTATTTAA AGAAATGTACCCCAATGCTAGCAATGATTACAACTTGCA
SEQ ID NO: 60	rs9513113	T	CGTTACACGTTTTGTTGCTAAGTATTCCATTTCCATGGCAGAGAAATGT[C/T]GATGAAAAAC AACACTAGCTTGAGGCTGATGCATTAATCTTTCATACATA
SEQ ID NO: 61	rs9551471	G	ACAATCCCTTTCTCCACCTAGAAATCATGCAATAGAAATACCAGTCTAC[A/G]CTCTTCAAAA CAAAACCCAGACATCAAAAGACAGTATCTATGCATATC
SEQ ID NO: 62	rs2296285	A	CCAGTTGAAAAGGGGAAAGAAATCAAAATTACCTTCCCCAACAGATTAT[T/A]AAAGGAAAT ACCAAACAGCCACAAAGAGCTCGAAACCTTTTCTATCGCC
SEQ ID NO: 63	rs9513116	A	CTCTAAAAGCCAAATACCAATTCAGCACTGGGTCCACTGAAATTTCTATCC[G/A]AAGATATCA GAACCTCAGTGATTGTGTCCTCCAAACACTATGCAAAAC
SEQ ID NO: 64	rs9551473	G	ACATATAAAAGAATTATATACAATGCCAAGTGGGATTTTTTCAGGAA[T/G]GCAAGATTG GTTTAACATCCAAAACCAATGTAATTACTATATCATGTCA
SEQ ID NO: 65	rs7330109	T	TAAAATGGCAGAAGAGGATTTGACAAAATCCAAAGCTCCTTTCGTGATG[A/T]AAACACTCA GGAAAATGGGAATTGCACTGAATTTCTCAACCTAATAAAT
SEQ ID NO: 66	rs9508037	G	AGAAATTAAAGAAGATCTACTAAGTGGGAAACATCTCATGTTTCATGGATT[A/G]GAAAGACTTA ATATTGTTAAGATGTTAATGTCCCCCAAATTTGATCTACAGA
SEQ ID NO: 67	rs1924981	T	ATACATGAAAGAACCCTCAAAATATATGCTGAGTGAAGAAATTAGTCA[C/T]AAAGACTA CAAATTATGTATCTATTATATAAAATATCCATGAGACAA
SEQ ID NO: 68	rs34140996	G	ATAAGGGGTGATAGCTAAAGGCTATGGAGTTTATTTTGAGACAATGACAT[A/G]TCTTAAAT TGACCGTGATGATGATGGTTGCATATATCTGCAATATGCT
SEQ ID NO: 69	rs7985584	A	TGTCATATGCTACAAACCACTGAATCATCATTTTAAATGGGTGGTAT[G/A]TAAATTATAT CTCAATAAACTTATTTTGAATTAACACAGAGGTTTAGTC
SEQ ID NO: 70	rs7992940	G	AATAAGAAAGTGCCTCTTCTAAAAAGAGCTCTTTCCTACTACCTGCCT[T/G]TCCCTCAGAA GAAAGACATTGCTTTTGTGTTTGTGTTTGTGTTT

	SNP_ID	Rare diagnostic allele	SEQUENCE
SEQ ID NO: 71	rs718273	T	CCTTGAAGCTATTACTGACCAATCAGAATACTATCATCTCTCTTC/TGGCCTCCTTCC TCCCCTCATCCCTCAGTCTCTCTCTCTCTCTCCTTC

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Figure 2

	SNP ID	2nd-PCR PRIMER SEQ	
SEQ ID NO:2	rs9582036	ACGTTGGATGACTGTGCCCCAGCAACAATAG	SEQ ID NO:20
SEQ ID NO:1	rs9554316	ACGTTGGATGATCGTAAAGACATCATTCG	SEQ ID NO:25
SEQ ID NO:4	rs9554320	ACGTTGGATGTTGGGAGGCCAGTTTGTAAAC	SEQ ID NO:30
SEQ ID NO:3	rs9513070	ACGTTGGATGGCAAGCTTGCCCCAACTTGTG	SEQ ID NO:35

	SNP ID	1st-PCR PRIMER SEQ	
SEQ ID NO:2	rs9582036	ACGTTGGATGGCATAATAGCATTACTCC	SEQ ID NO:21
SEQ ID NO:1	rs9554316	ACGTTGGATGTGCTGGAGACCAATGACCAAG	SEQ ID NO:26
SEQ ID NO:4	rs9554320	ACGTTGGATGAGCAAGGTTCTGTGTAG	SEQ ID NO:31
SEQ ID NO:3	rs9513070	ACGTTGGATGGTTGTGTGGGCTGCACTC	SEQ ID NO:36

	SNP ID	AL	MT	UEP MASS	UEP SEQ	
SEQ ID NO:2	rs9582036	91	46,9	5442,6	AGCAACAATAGCCTTCTT	SEQ ID NO:22
SEQ ID NO:1	rs9554316	100	48,2	7282,8	AAGACATCATTCGATTTTCTTCT	SEQ ID NO:27
SEQ ID NO:4	rs9554320	93	98,4	5515,6	ACAGCGGCTTTGCAGTGC	SEQ ID NO:32
SEQ ID NO:3	rs9513070	93	97	5203,4	CGTGTGGCCACACGGGCT	SEQ ID NO:37

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Figure 2 (continued)

	SNP ID	EXT1 CALL	EXT1 MASS	EXT1 SEQ	
SEQ ID NO:2	rs9582036	C	5689,8	AGCAACAATAGCCTTCTTC	SEQ ID NO:23
SEQ ID NO:1	rs9554316	G	7570	AAGACATCATTCGATTTTCTG	SEQ ID NO:28
SEQ ID NO:4	rs9554320	C	5802,8	ACAGCGGCTTGCAGTGG	SEQ ID NO:33
SEQ ID NO:3	rs9513070	A	5474,6	CGTGTGGCCACGGGCTA	SEQ ID NO:38

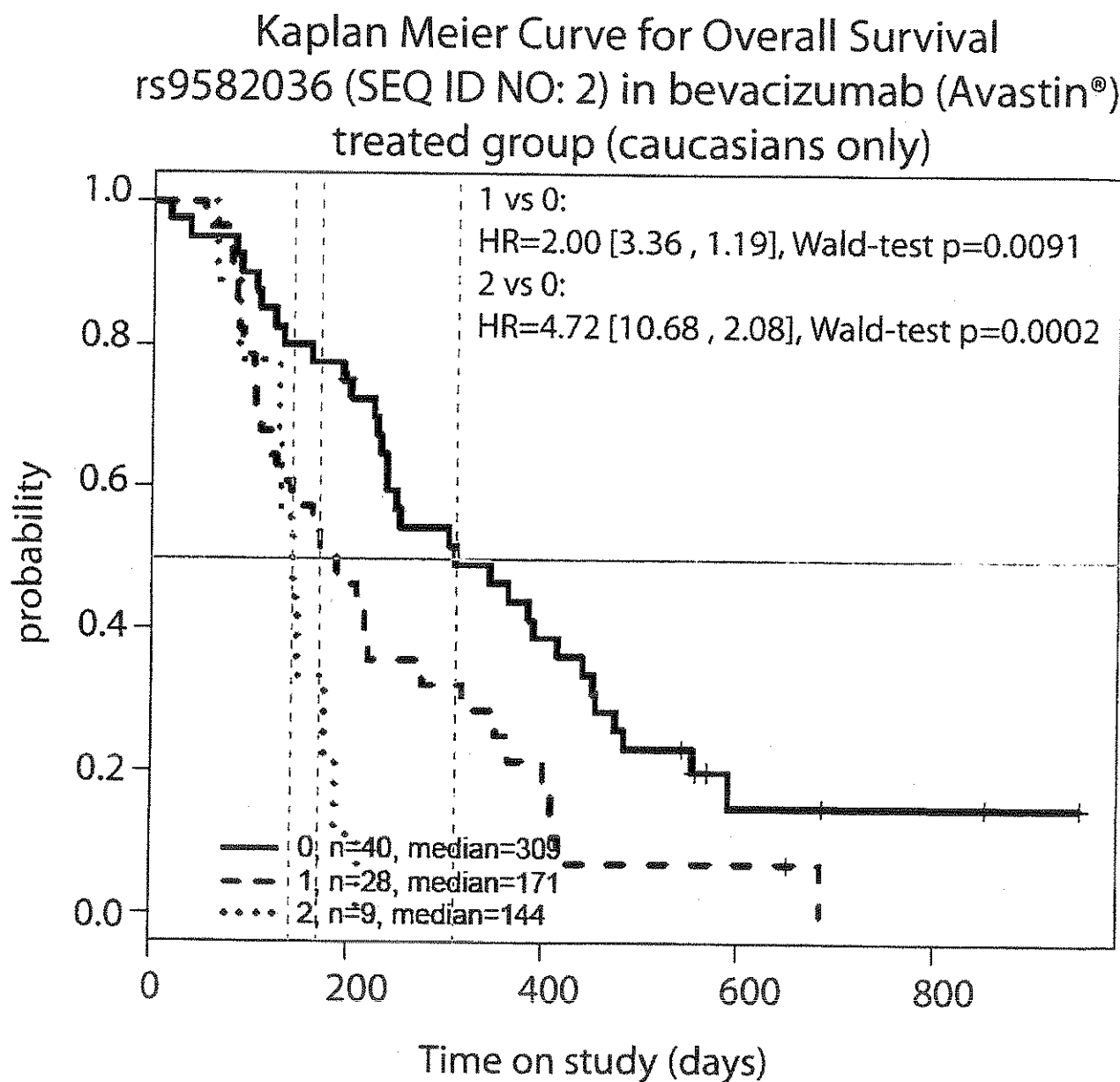
	SNP ID	EXT2 CALL	EXT2 MASS	EXT2 SEQ	
SEQ ID NO:2	rs9582036	A	5713,8	AGCAACAATAGCCTTCTTA	SEQ ID NO:24
SEQ ID NO:1	rs9554316	T	7609,9	AAGACATCATTCGATTTTCTT	SEQ ID NO:29
SEQ ID NO:4	rs9554320	A	5842,7	ACAGCGGCTTGCAGTGG	SEQ ID NO:34
SEQ ID NO:3	rs9513070	G	5490,6	CGTGTGGCCACGGGCTG	SEQ ID NO:39

Abbreviations:

SEQ	Sequence
AL	AMPLICON LENGTH
MT	Melting Temperature
UEP	Unextended primer mass
EXT1	Extended product 1
EXT2	Extended product 2

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Figure 3



In the legend 0, 1 and 2 refer to genotypes:

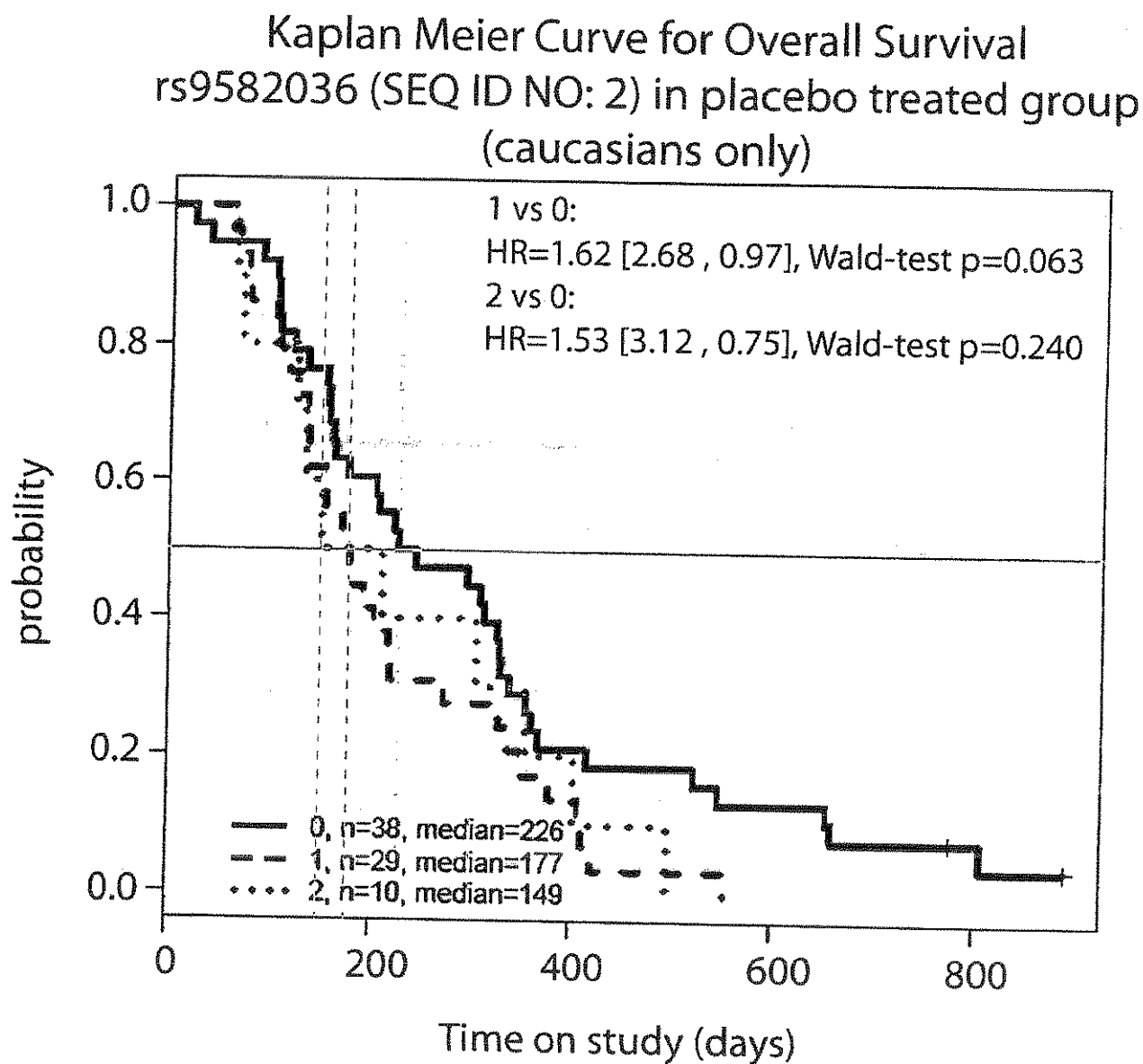
0: AA (n=40, median=309)

1: AC (n=28, median=171)

2: CC (n=9, median=144)

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Figure 4



In the legend 0, 1 and 2 refer to genotypes:

0: AA (n=38, median=226)

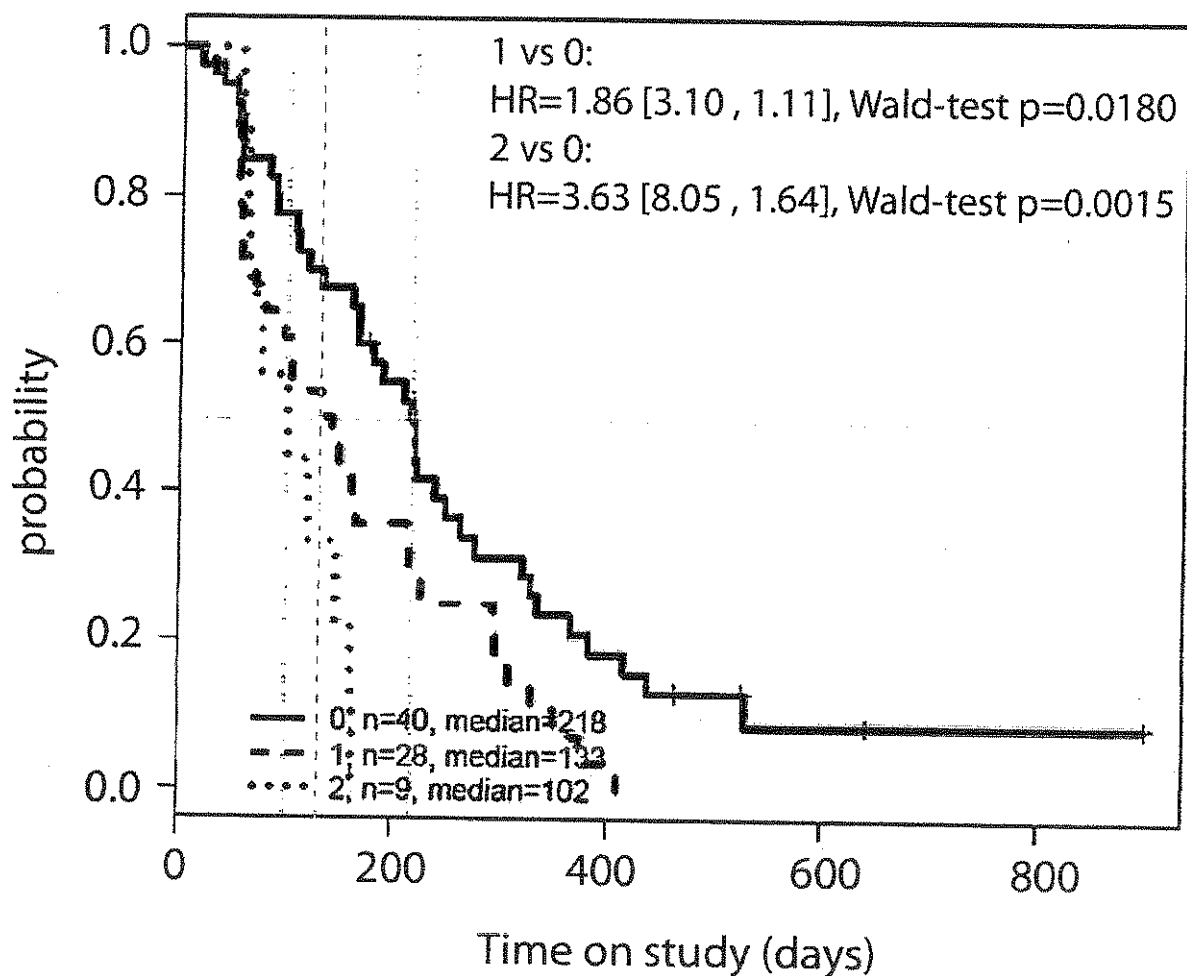
1: AC (n=29, median=177)

2: CC (n=10, median=149)

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Figure 5

Kaplan Meier Curve for Progression Free Survival
rs9582036 (SEQ ID NO: 2) in bevacizumab (Avastin®)
treated group (caucasians only)



In the legend 0, 1 and 2 refer to genotypes:

0: AA (n=40, median=218)

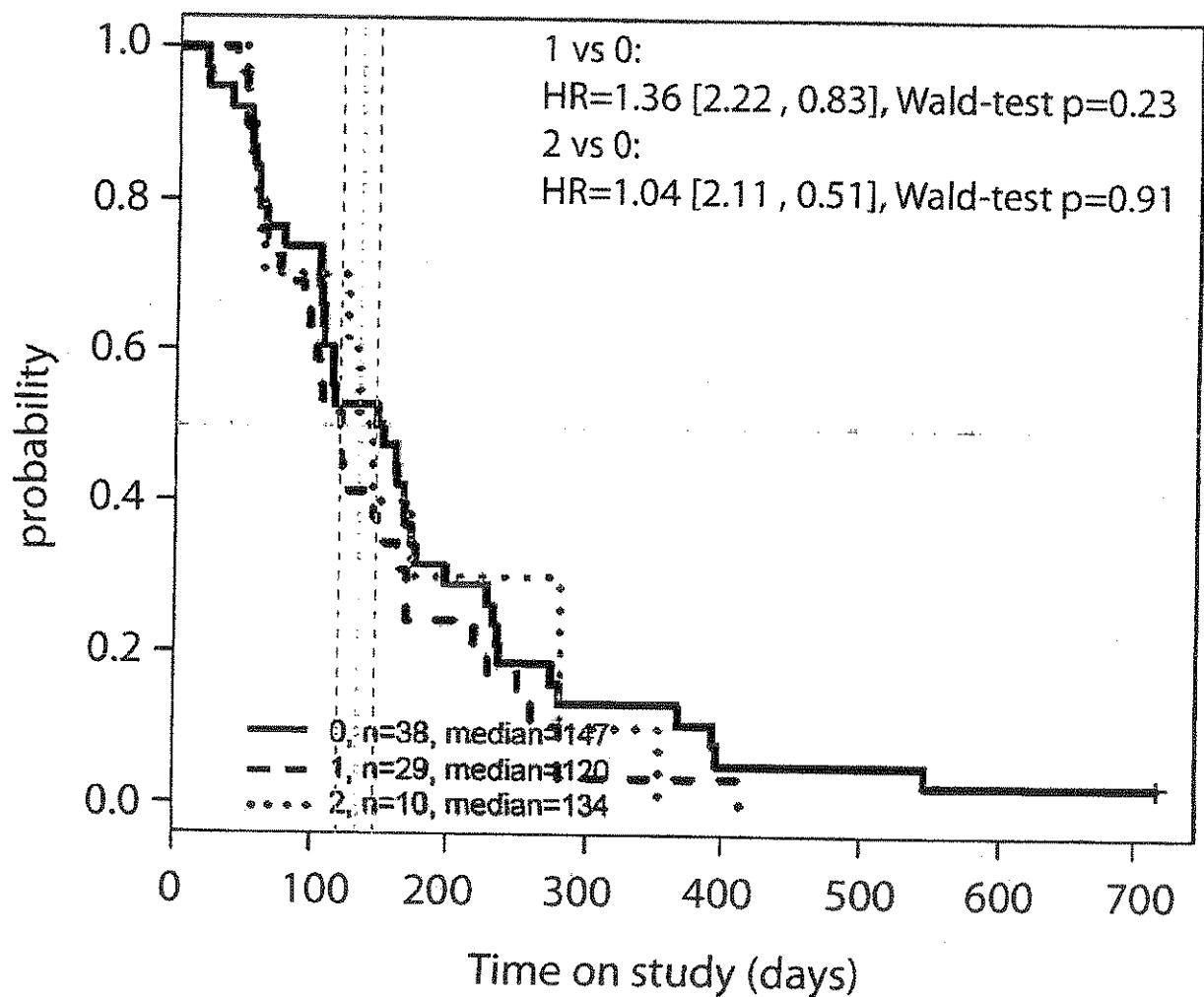
1: AC (n=28, median=133)

2: CC (n=9, median=102)

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Figure 6

Kaplan Meier Curve for Progression Free Survival
rs9582036 (SEQ ID NO: 2) in placebo treated group
(caucasians only)



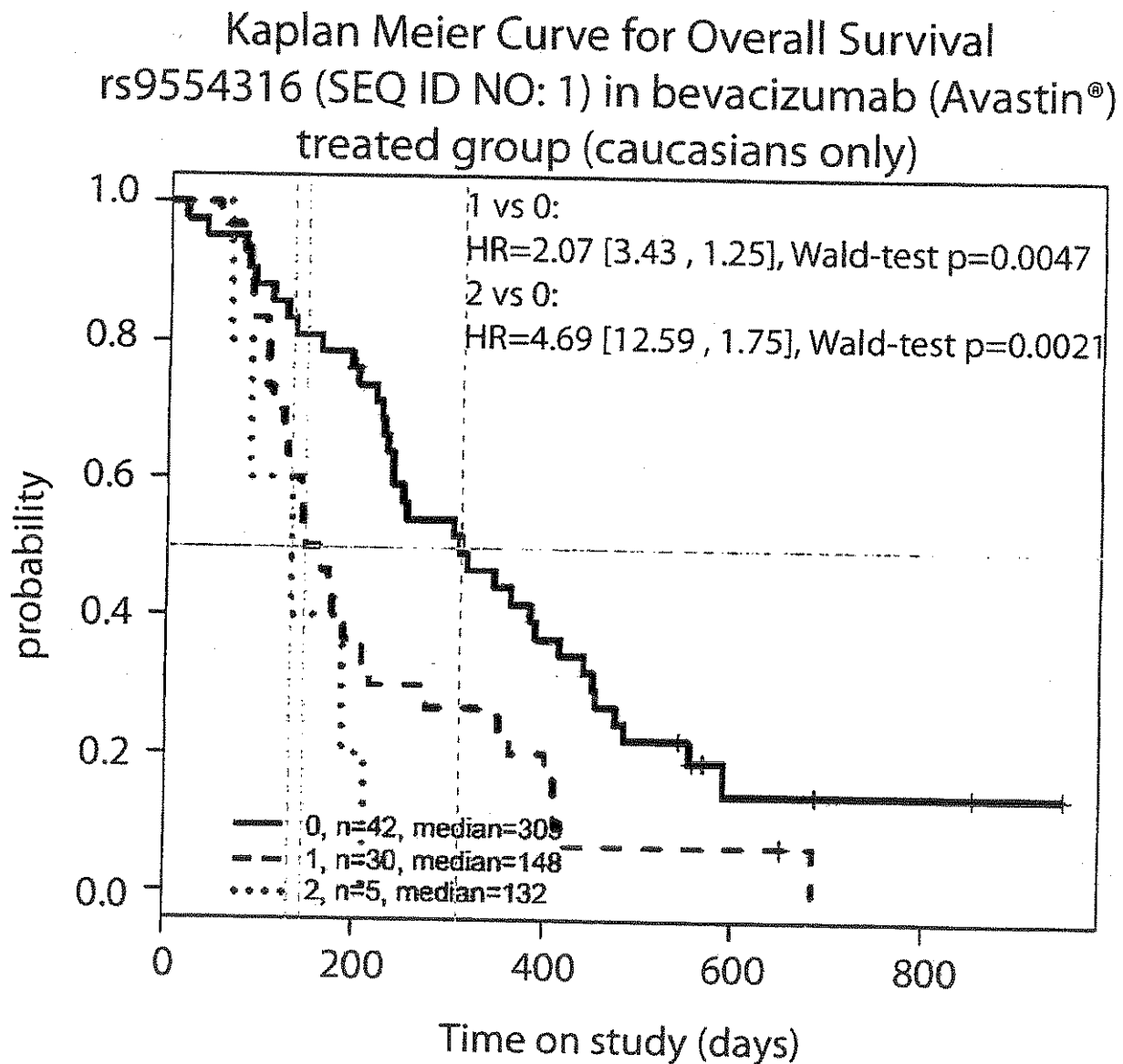
In the legend 0, 1 and 2 refer to genotypes:

0: AA (n=38, median=147)

1: AC (n=29, median=120)

2: CC (n=10, median=134)

Figure 7



In the legend 0, 1 and 2 refer to genotypes:

0: GG (n=42, median=309)

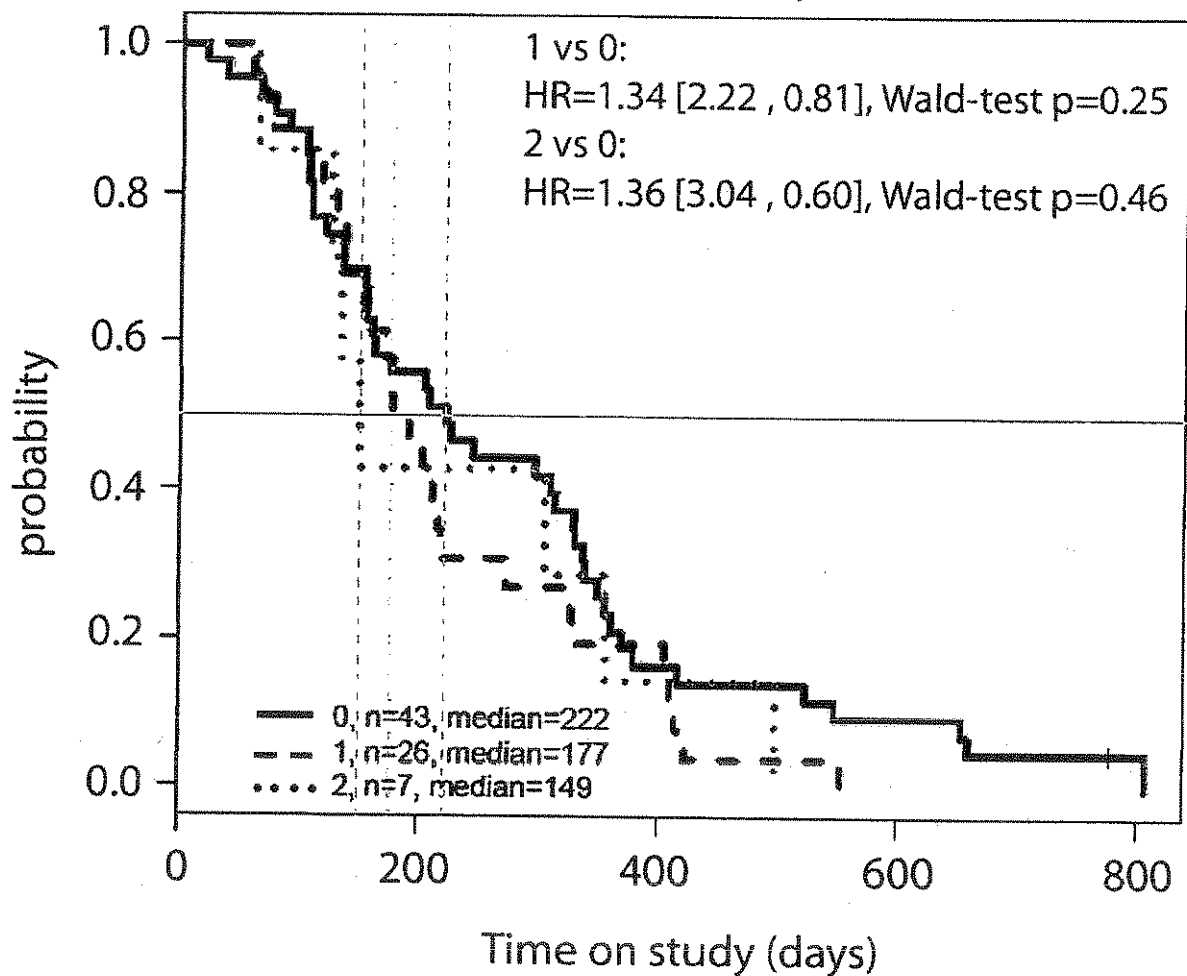
1: TG (n=30, median=148)

2: TT (n=5, median=132)

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Figure 8

Kaplan Meier Curve for Overall Survival
rs9554316 (SEQ ID NO: 1) in placebo treated group
(caucasians only)



In the legend 0, 1 and 2 refer to genotypes:

0: GG (n=43, median=222)

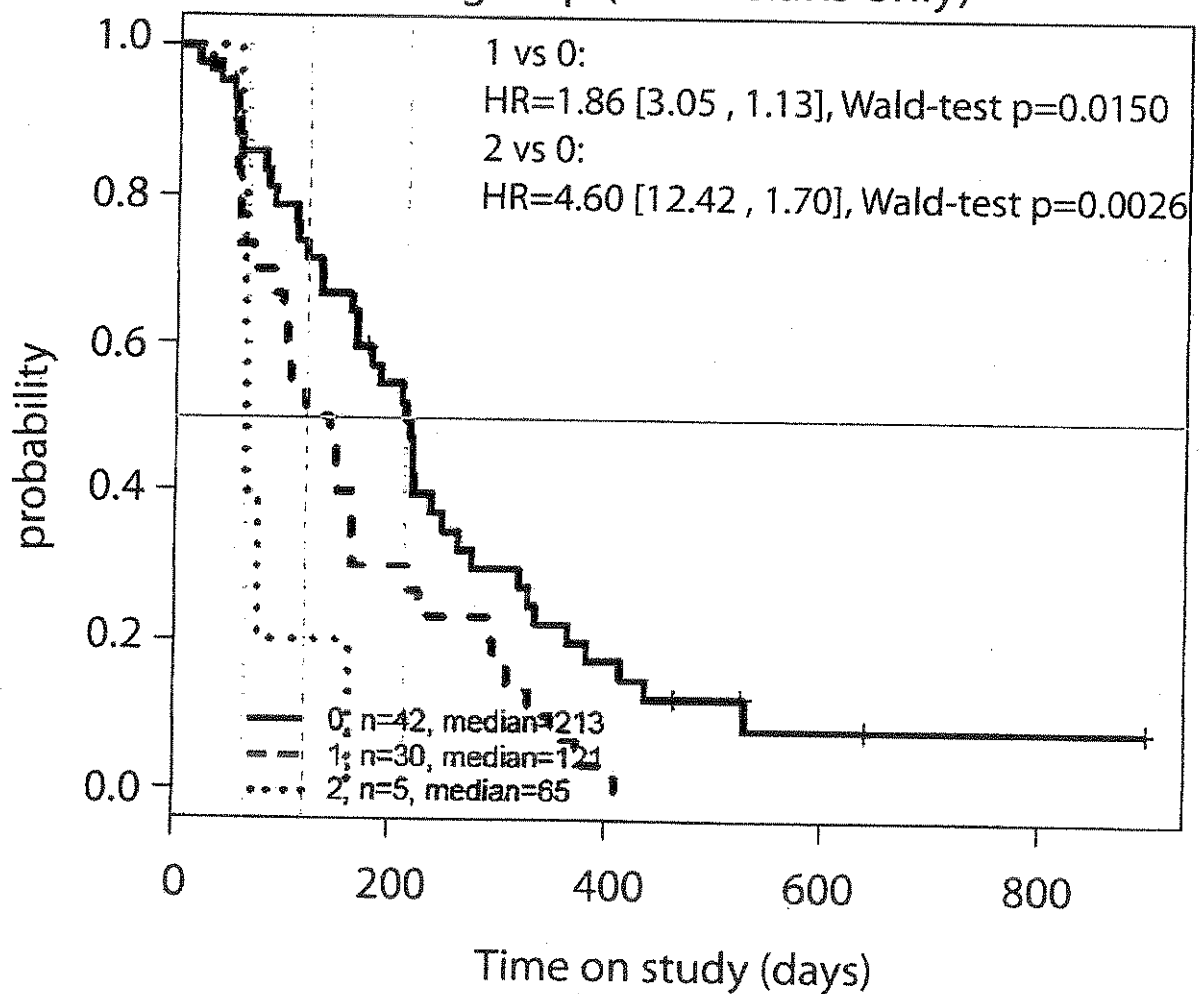
1: TG (n=26, median=177)

2: TT (n=7, median=149)

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Figure 9

Kaplan Meier Curve for Progression Free Survival
rs9554316 (SEQ ID NO: 1) in bevacizumab (Avastin®)
treated group (caucasians only)



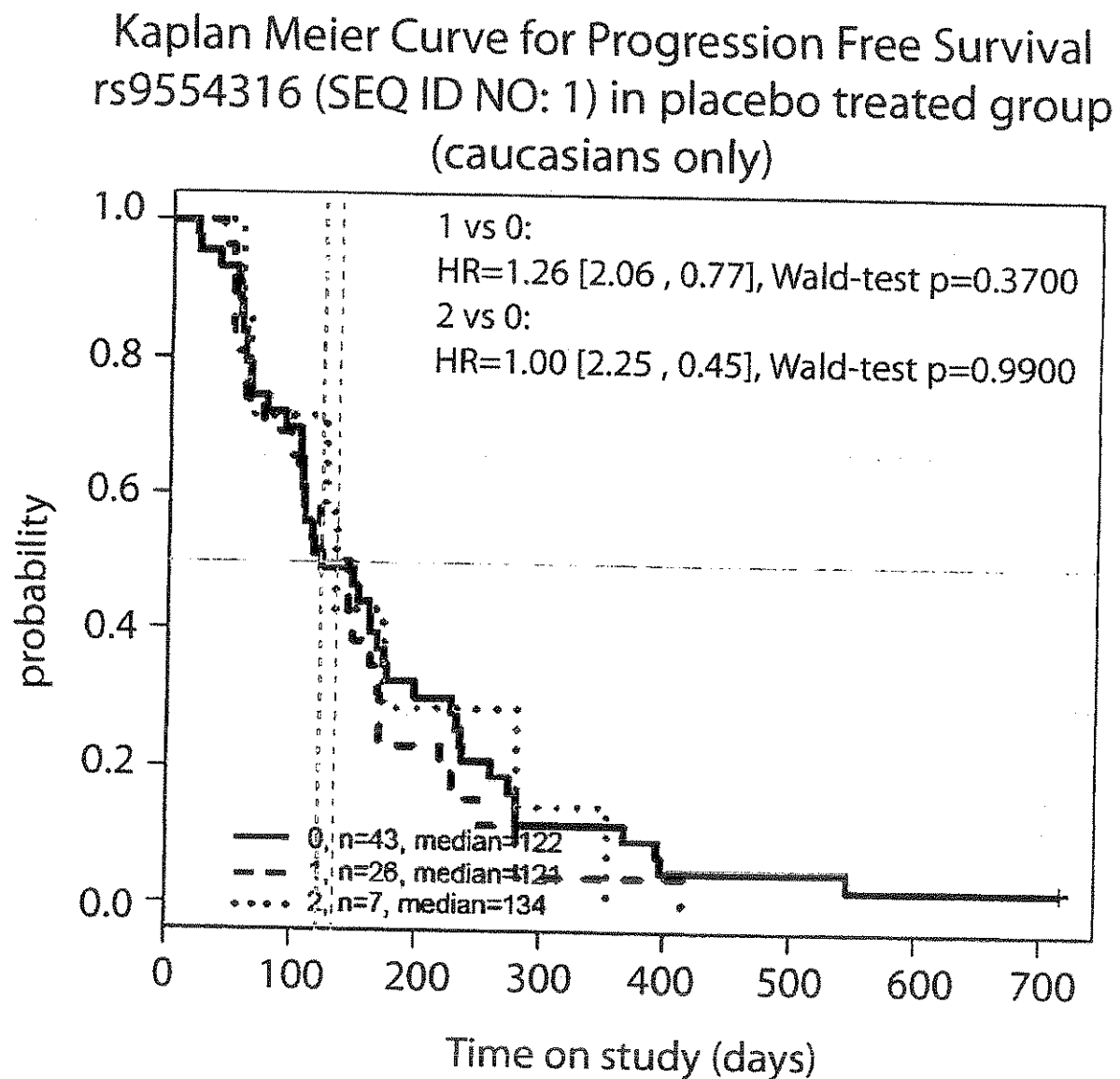
In the legend 0, 1 and 2 refer to genotypes:

0: GG (n=42, median=213)

1: TG (n=30, median=121)

2: TT (n=5, median=65)

Figure 10



In the legend 0, 1 and 2 refer to genotypes:

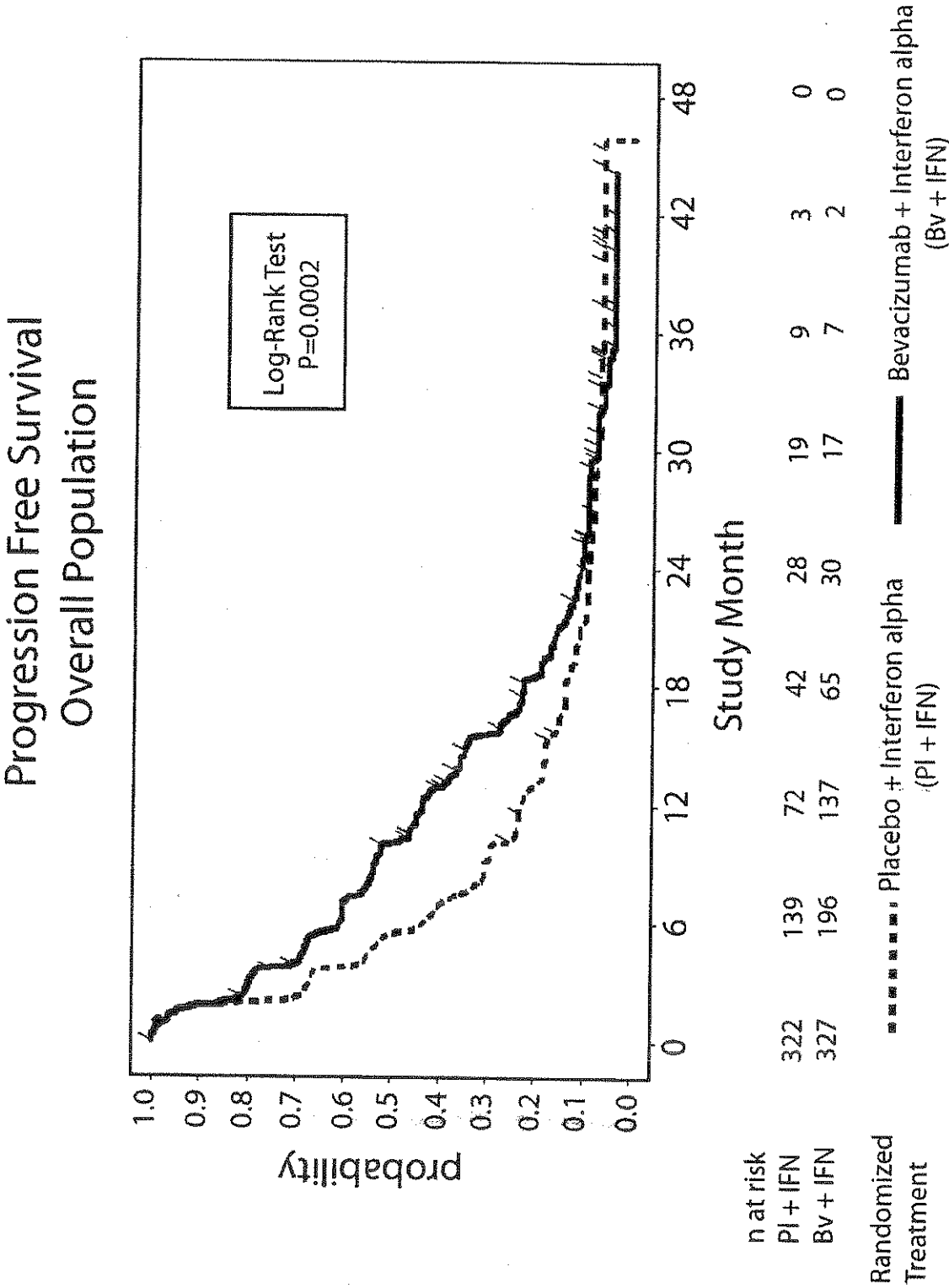
0: GG (n=43, median=122)

1: TG (n=26, median=121)

2: TT (n=7, median=134)

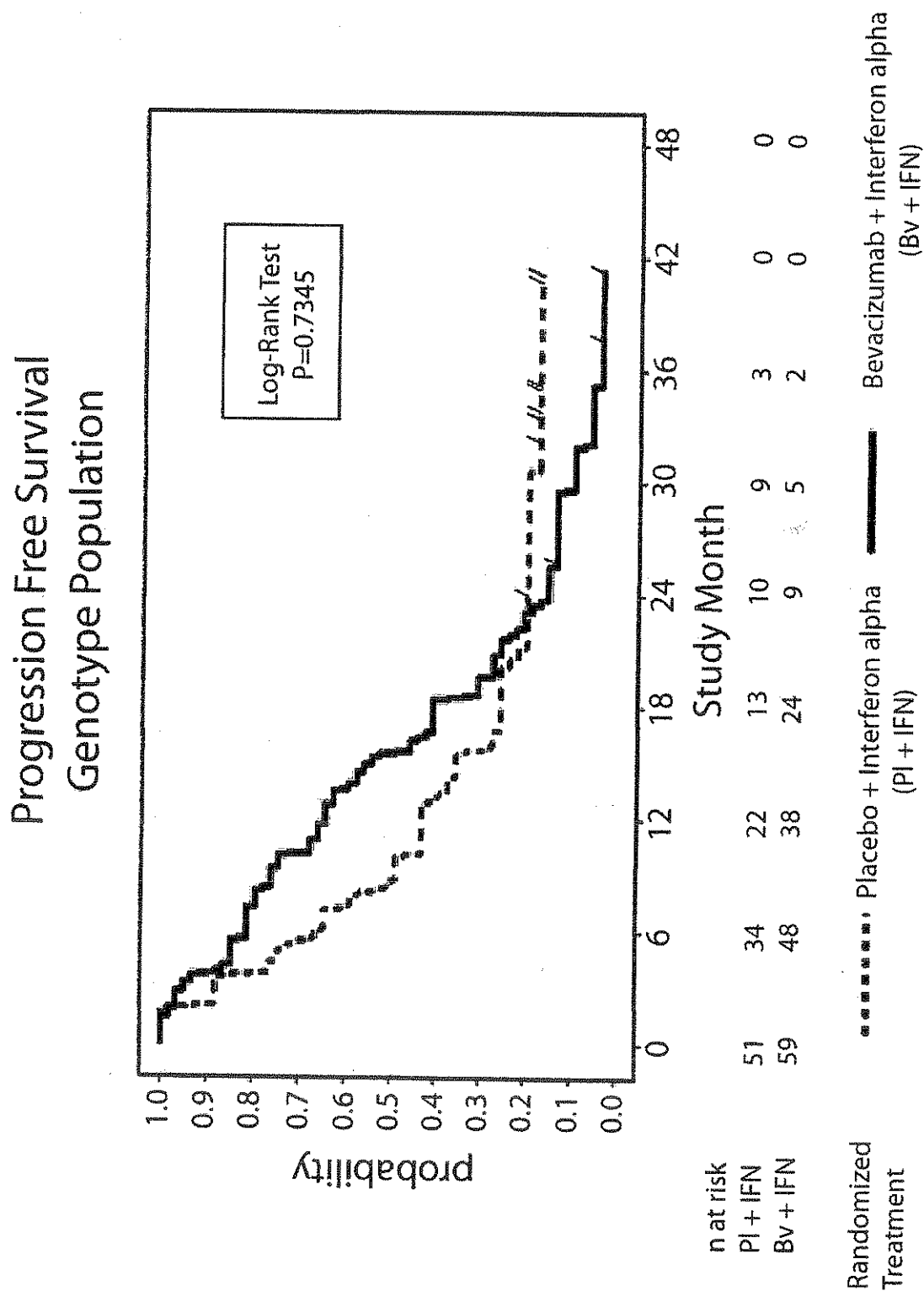
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Figure 11



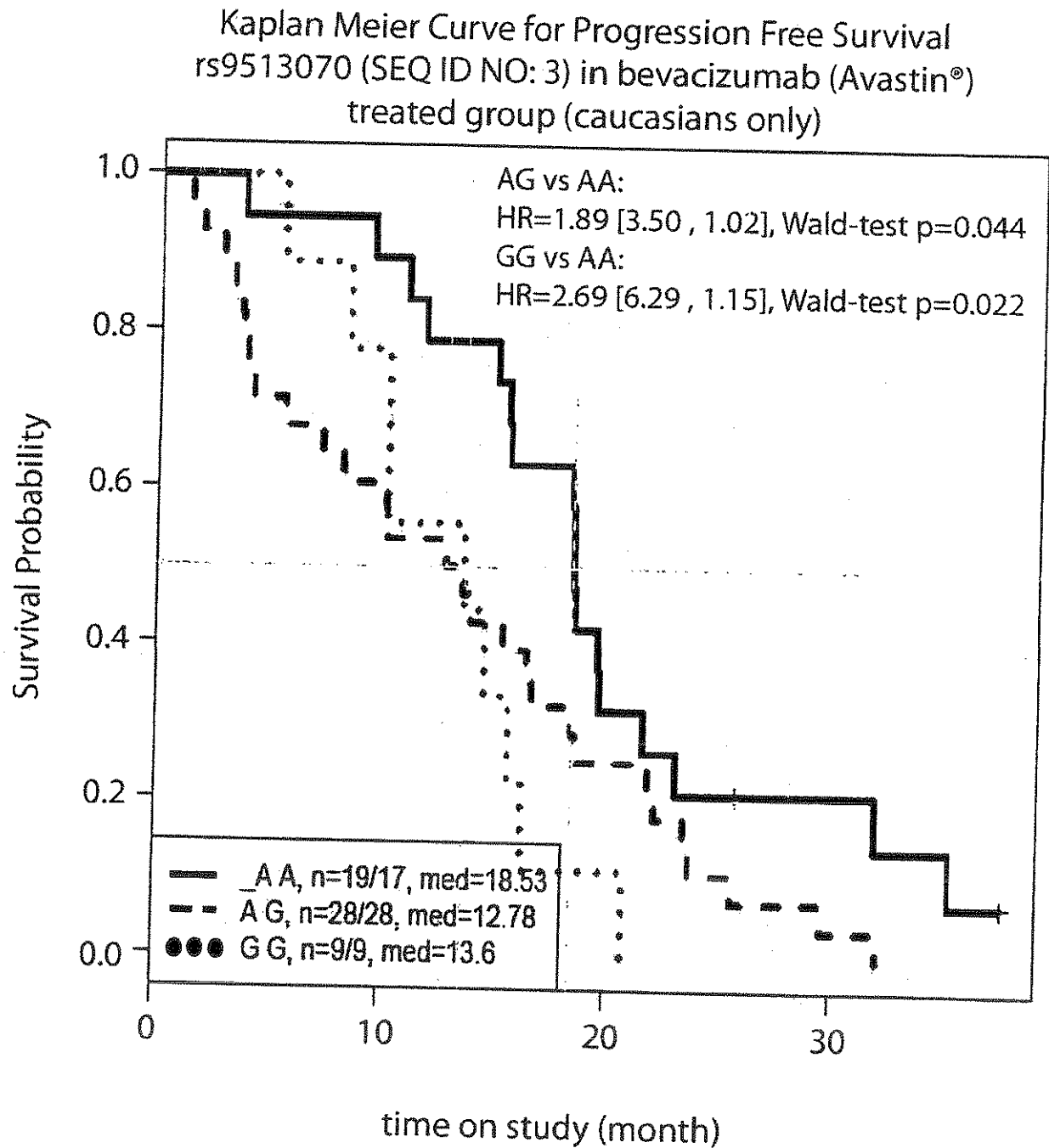
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Figure 12



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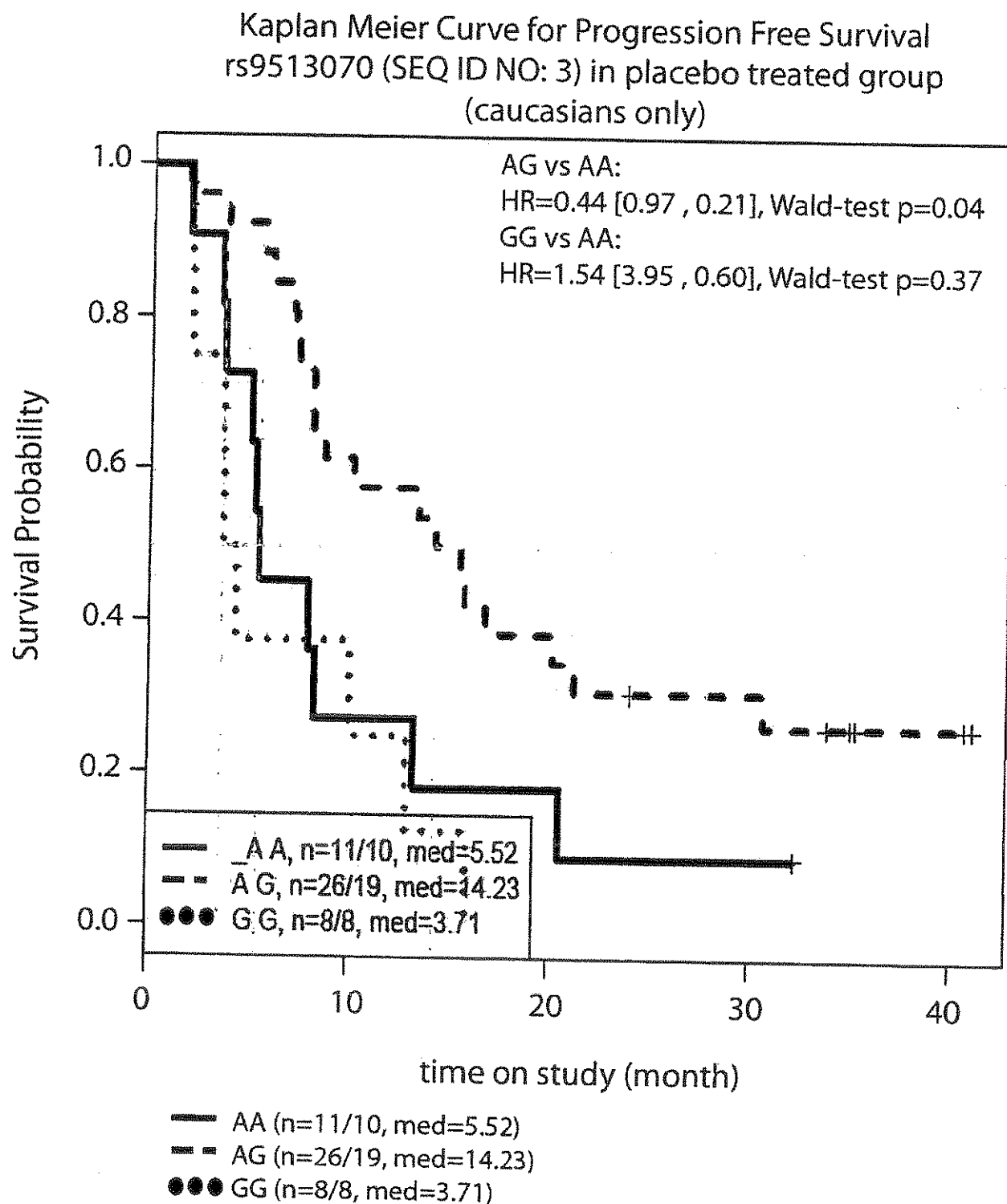
Figure 13



— AA (n=19/17, med=18.53)
-- AG (n=28/28, med=12.78)
●●● GG (n=9/9, med=13.6)

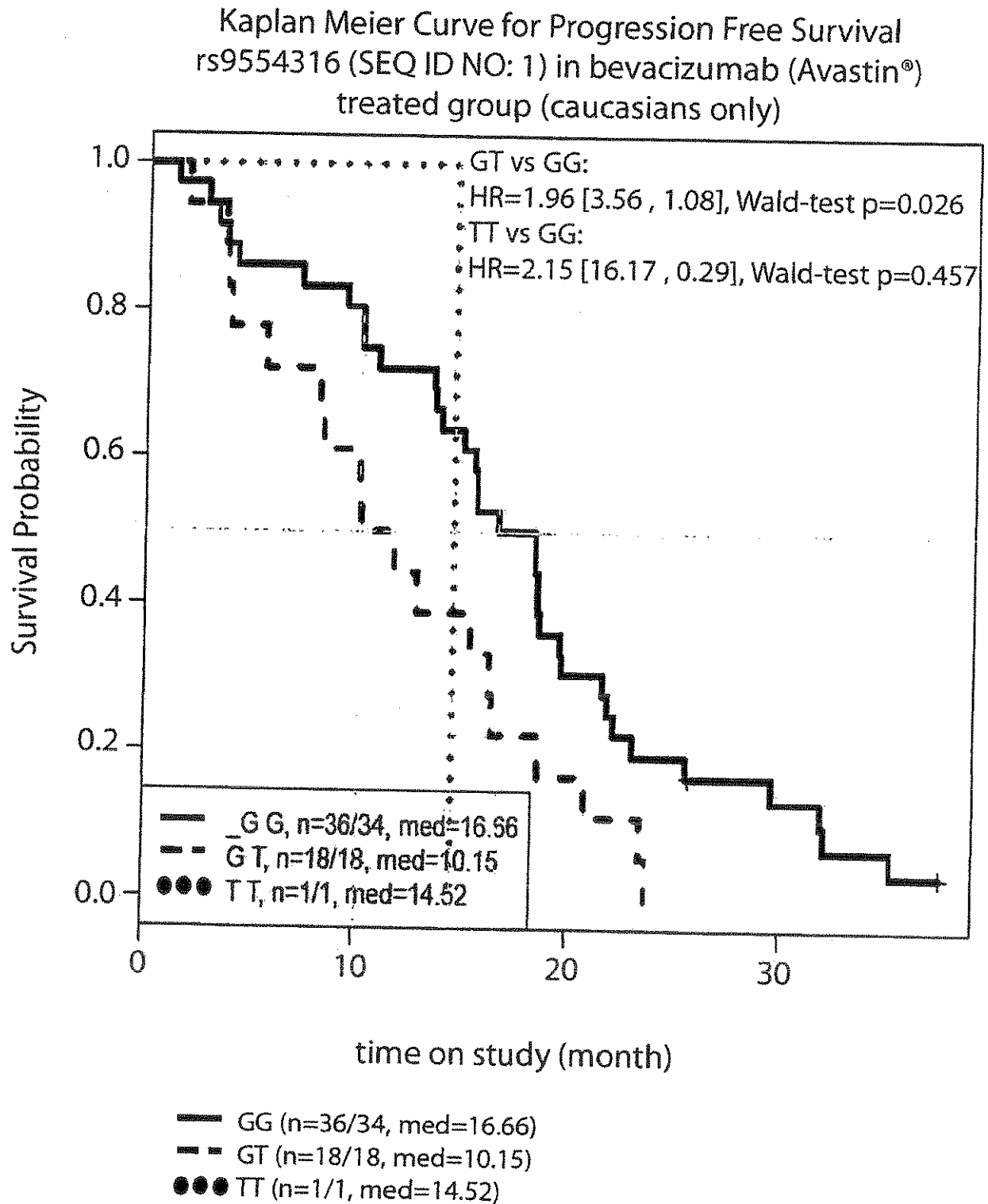
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Figure 14



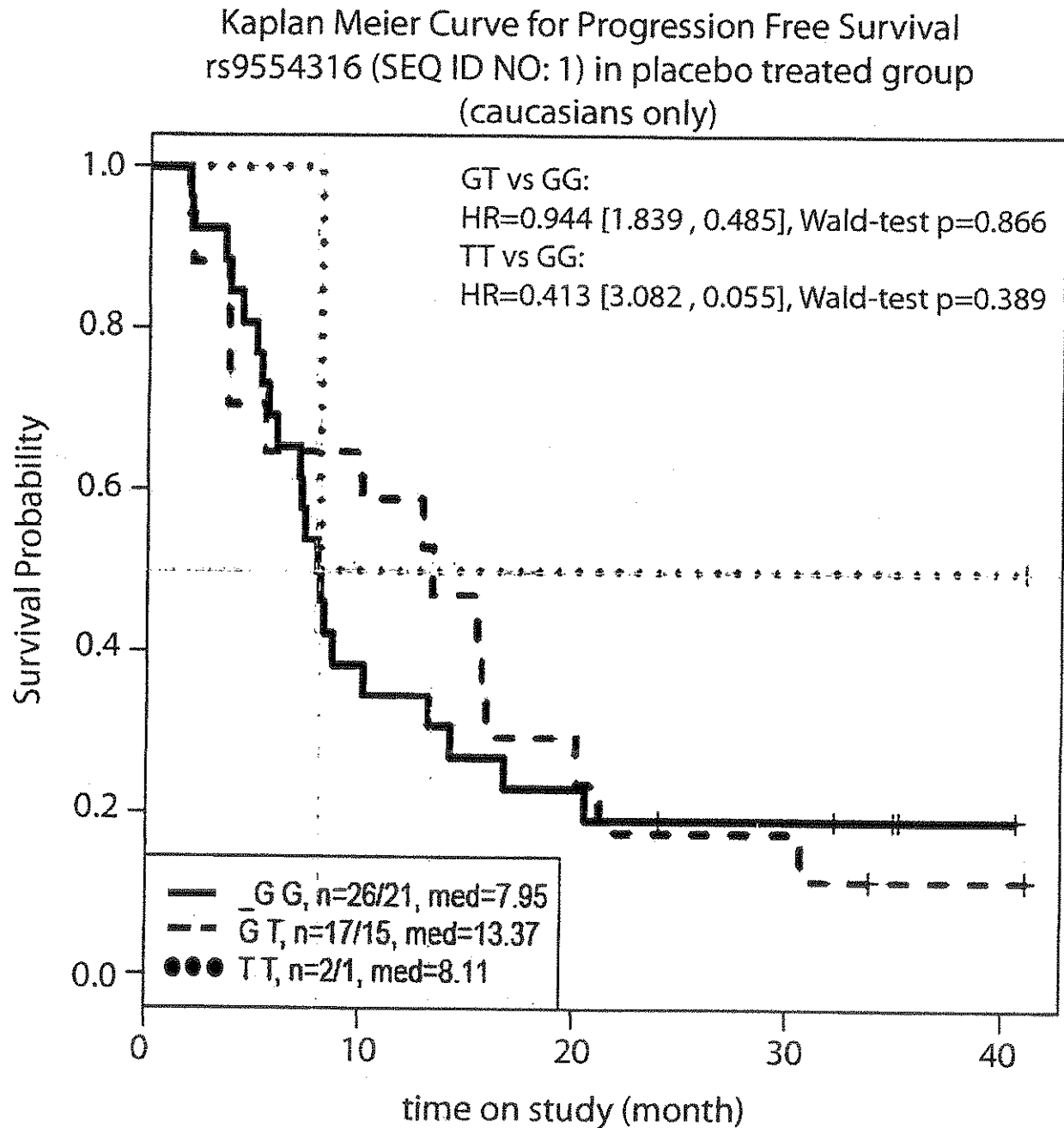
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Figure 15



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Figure 16



— GG (n=26/21, med=7.95)
-- GT (n=17/15, med=13.37)
... TT (n=2/1, med=8.11)