

Supplemental Material Table 1.

Primers used to resequence *NT5C3* and create expression constructs

| Primer Name | Primer Sequence |
|----------------------------------|---------------------------------------|
| F-5'-FR & Exon 1(NM_001002009.1) | GGTGCAA A ACTGGATTTCATAAAA |
| R-5'-FR & Exon 1(NM_001002009.1) | CTAGCGAGATCCCAGCCTGA |
| F-5'-FR & Exon 1(NM_016489.11) | GGACAGTAGAGAGAAAGCCGATA |
| R-5'-FR & Exon 1(NM_016489.11) | CCCTCAACCTGGAAAGAGACT |
| F-Exon 2 | TGGCATAACAGGAAGAACCTG |
| R-Exon 2 | TCTGAGAAGCACGACAATGTG |
| F-Exon R | ATGGTTGTATGTTGCACAGGAG |
| R-Exon R | ACCAATCCATTATGGTGAAAGC |
| F-Exon 3 | GCTTGGTGTCCCACACTTTT |
| R-Exon 3 | GGAAGCTGAGGAGGGAAAAC |
| F-Exon 4 | TATAAAGGAACCGCATGAGAAAA |
| R-Exon 4 | AAGAAAAAGCAGGTCTCCTCACT |
| F-Exon 5 & Exon 6 | TCTCAGTGATGTAAGCGAGCA |
| R-Exon 5 & Exon 6 | TCTGTTTGTGGCAATACAGGT |
| F-Exon 7 | GGAGGACTGGGAAACTCAGTAA |
| R-Exon 7 | AGGTGGCATATTTTGGATATGG |
| F-Exon 8 | ACAGCCCTCTTGGGCAGTAT |
| R-Exon 8R | GAGAAGGATGGAGTCCCACA |
| F-Exon 9 & Exon 10 | AGCTATGGTGAATGTGATGGACT |
| R-Exon 9 & Exon 10 | AGTGTGTTGAGAGAGGTGGAGAA |

Site-directed mutagenesis, expression construct

| | |
|-----------------|--|
| F-Exon 2 (9) | ATGACTAA <u>C</u> CAAGAGTCTGCCGTACATGTGAA |
| R-Exon 2 (9) | TTCACATGTACGGCAGACTCTT <u>G</u> TTAGTCAT |
| F-Exon 6 (276) | ACTAAAGGAAAAATA <u>T</u> TACGCTATTGAAGTTG |
| R-Exon 6 (276) | CAACTTCAATAGCGTA <u>A</u> TATTTTTCCCTTAGT |
| F-Exon 6 (306) | TGATCCTGTTCTTAC <u>C</u> GTAGAAGAGAAGTACC |
| R-Exon 6 (306) | GGTACTTCTCTTCTAC <u>G</u> GTAAGTTCAGGATCA |
| F-Exon 9 (959) | AGTGGCCAATGTTGA <u>A</u> CACATTCTGAAAATTG |
| R-Exon 9 (959) | CAATTTTCAGAATGTG <u>T</u> TCAACATTGGCCACT |
| F-Exon 10 (847) | ATTGTTTTAGTACA <u>A</u> CATGAATCATTAGAAGT |
| R-Exon 10 (847) | ACTTCTAATGATTCAT <u>G</u> TTGTTACTAAAACAAT |

Bold, underlined letters are mutated bases in site-directed mutagenesis primers.

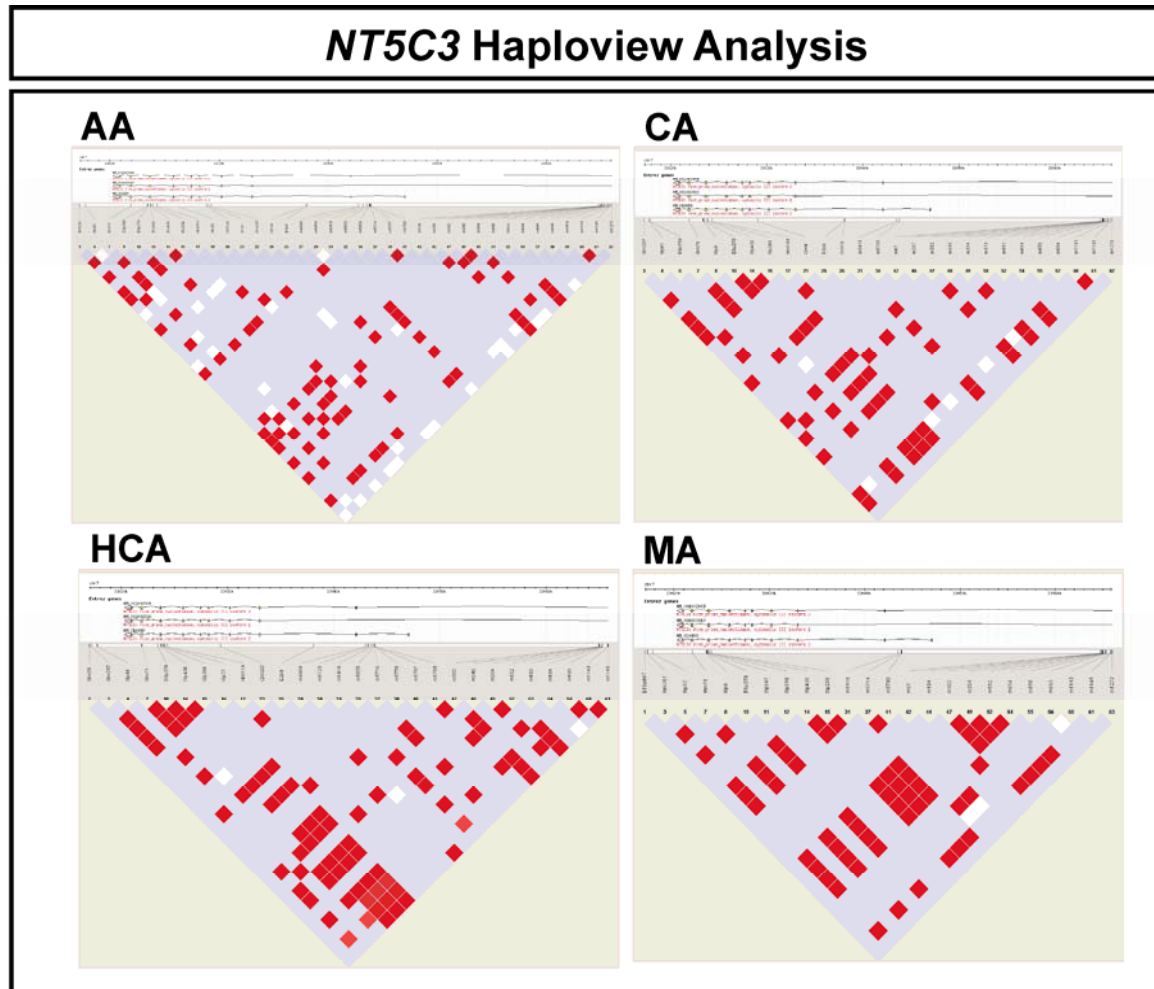
| Haplotype frequency | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------------|--------|-------|--------|--------|---------|---------|---------|---------|---------|---------|--------|--------|--------|--------|--------|--------|--------|--------|--------|--------|--------|--------|--------|-------|---------|---------|---------|---------|---------|---------|---|
| observed | allele | AA | CA | HCA | MA | (-1273) | (-1270) | (-1145) | (-1143) | (-1018) | (-944) | (-884) | (-693) | (-656) | (-654) | (-653) | (-552) | (-373) | (-354) | (-340) | (-302) | (-267) | (-258) | (-67) | (-5780) | (-5767) | (-5757) | (-5756) | (-5714) | (-5705) | |
| o | *1A | 0.200 | | | 0.167 | A | T | T | G | T | T | D | C | C | G | T | G | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| o | *1B | 0.158 | 0.289 | 0.192 | 0.433 | A | T | G | G | T | T | D | C | C | G | T | G | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| o | *1C | 0.150 | 0.292 | 0.300 | 0.133 | A | T | T | A | T | T | D | C | C | G | T | G | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| o | *1D | 0.083 | | | | A | T | T | G | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| o | *1E | 0.075 | | | | A | T | T | A | T | T | D | C | C | G | T | G | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1F | 0.058 | | | | A | T | T | G | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1G | 0.058 | | | | C | T | T | G | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1H | 0.050 | | | | A | T | T | A | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1I | 0.033 | 0.008 | | | A | T | T | G | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1J | 0.025 | | | | A | T | T | G | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1K | 0.017 | | | | A | T | G | G | T | T | D | C | C | A | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1L | 0.008 | | | | A | T | G | C | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1M | 0.008 | | | | A | T | T | G | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1N | 0.008 | | | | A | T | T | G | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1O | 0.008 | | | | C | T | T | G | T | C | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1P | 0.009 | | | | C | T | T | G | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1Q | 0.008 | | | | C | T | T | G | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| o | *1R | | 0.225 | 0.150* | 0.117 | A | T | T | G | T | T | D | C | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C |
| o | *1S | | 0.042* | 0.083 | 0.067 | A | T | T | G | T | T | D | C | C | A | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1T | | 0.036 | | | A | T | G | G | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1U | | 0.019 | | | A | T | G | G | T | T | D | C | C | G | T | G | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1V | | 0.017 | 0.008 | | A | T | T | G | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1W | | 0.017 | | | A | T | T | G | T | T | D | C | C | T | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C |
| i | *1X | | 0.008 | | | A | C | T | G | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1Y | | 0.008 | | | A | T | T | G | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1Z | | 0.008 | | | A | T | T | G | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1AA | | 0.008 | | | A | T | T | G | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1AB | | 0.008 | | | A | T | T | G | T | T | D | C | C | G | T | G | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| o | *1AC | | | 0.125 | 0.0167* | A | T | T | G | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| o | *1AD | | | 0.083 | | A | T | T | G | T | T | D | C | C | A | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1AE | | | 0.033 | | A | T | G | G | T | T | D | C | C | G | T | G | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| o | *1AF | | | 0.025 | | A | T | G | G | T | T | D | C | C | G | T | G | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1AG | | | 0.017 | | A | T | T | G | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1AH | | | 0.008 | | A | T | T | A | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1AJ | | | 0.008 | | A | T | T | G | T | T | D | C | C | A | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1AK | | | 0.008 | | A | T | T | G | T | T | D | C | C | G | C | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1AL | | | 0.008 | | A | T | T | G | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1AM | | | | 0.008 | A | T | T | G | T | T | D | C | C | A | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1AN | | | | 0.008 | A | T | T | G | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1AO | | | | 0.008 | A | T | T | G | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *1AP | | | | 0.008 | A | T | T | G | T | T | D | C | C | G | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |
| i | *2A | | | | 0.008 | A | T | T | G | T | T | D | C | C | A | T | C | C | T | T | A | D | D | T | C | A | A | T | A | C | |

| Haplotype frequency | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---------------------|--------|-------|--------|--------|---------|---------|---------|---------|---------|---------|---------|-------|----------|--------|--------|----------|--------|---------|---------|---------|---------|---------|-------|--------|---------|--------|--------|----------|---------|----------|
| observed | allele | AA | CA | HCA | MA | (-5635) | (-5633) | (-5418) | (-5059) | (-4900) | (-4645) | E2(9) | I2(-237) | I2(-4) | I4(35) | I4(-114) | I5(77) | I5(266) | I5(430) | I5(548) | E6(276) | E6(306) | I6(9) | I6(71) | E9(759) | I9(52) | I9(61) | I9(-267) | I9(-38) | E10(847) |
| o | *1A | 0.200 | | | 0.167 | G | A | G | G | G | T | G | T | C | G | C | G | G | A | C | T | A | C | G | T | A | A | C | G | |
| o | *1B | 0.158 | 0.289 | 0.192 | 0.433 | G | A | G | G | G | T | G | T | C | G | C | G | G | A | C | T | A | C | G | T | A | A | C | G | |
| o | *1C | 0.150 | 0.292 | 0.300 | 0.133 | G | A | G | G | G | T | G | T | C | G | C | G | G | A | C | T | A | C | G | T | A | A | C | G | |
| o | *1D | 0.083 | | | | G | A | A | G | G | T | G | T | C | G | C | A | A | A | T | T | A | C | G | T | A | A | C | G | |
| o | *1E | 0.075 | | | | G | A | A | G | G | T | G | T | C | G | C | G | G | A | A | C | C | A | C | G | T | A | A | C | G |
| i | *1F | 0.058 | | | | G | A | A | G | G | T | G | T | C | G | C | A | A | A | C | T | A | C | T | G | T | A | A | C | G |
| i | *1G | 0.058 | | | | G | A | A | G | G | T | G | T | C | G | C | G | G | A | A | C | T | A | C | G | T | A | A | C | G |
| i | *1H | 0.050 | | | | G | A | A | G | G | T | G | T | C | G | C | G | G | A | A | C | T | A | C | G | T | A | A | C | G |
| i | *1I | 0.033 | 0.008 | | | G | A | A | G | G | T | G | T | C | G | C | A | A | A | T | T | A | C | T | G | T | A | A | C | G |
| i | *1J | 0.025 | | | | G | A | A | G | G | C | G | T | C | A | C | A | A | A | T | T | A | C | G | T | A | A | C | G | G |
| i | *1K | 0.017 | | | | G | A | A | G | G | T | G | T | C | G | C | A | A | A | T | T | A | C | T | G | T | A | A | C | G |
| i | *1L | 0.008 | | | | G | A | A | G | G | T | G | T | C | G | C | A | A | A | T | T | A | C | G | T | A | A | C | G | G |
| i | *1M | 0.008 | | | | G | A | A | G | G | T | C | G | T | C | G | C | A | A | A | C | T | A | C | G | T | A | A | C | G |
| i | *1N | 0.008 | | | | G | A | A | G | G | T | G | T | C | G | C | A | A | A | C | T | A | C | G | T | A | A | C | G | G |
| i | *1O | 0.008 | | | | G | A | A | G | G | T | G | T | C | G | C | A | A | A | C | T | A | C | G | T | A | A | C | G | G |
| i | *1P | 0.009 | | | | G | A | A | G | G | T | G | T | C | G | C | A | A | A | T | T | A | C | G | T | A | A | C | G | G |
| o | *1R | | 0.225 | 0.150* | 0.117 | G | A | A | G | G | T | G | T | C | G | C | A | A | A | T | T | A | C | G | T | A | A | C | G | G |
| o | *1S | | 0.042* | 0.083 | 0.067 | G | A | A | G | G | T | G | T | C | G | C | A | A | A | T | T | A | C | G | T | A | A | C | G | G |
| i | *1T | | 0.036 | | | G | A | A | G | G | T | G | T | C | G | C | A | A | A | T | T | A | C | G | T | A | A | C | G | G |
| i | *1U | | 0.019 | | | G | A | A | G | G | T | G | T | C | G | C | A | A | A | T | T | A | C | G | T | A | A | C | G | G |
| i | *1V | | 0.017 | 0.008 | | G | A | A | G | G | T | G | T | C | G | C | A | A | A | C | T | A | C | G | T | A | A | C | G | G |
| i | *1W | | 0.017 | | | G | A | A | G | G | T | G | T | C | G | C | A | A | A | T | T | A | C | T | G | T | A | A | C | G |
| i | *1X | | 0.008 | | | G | A | A | G | G | T | G | T | C | G | C | A | A | A | T | T | A | C | T | G | T | A | A | C | G |
| i | *1Y | | 0.008 | | | G | A | A | G | G | T | G | T | C | G | C | A | A | A | T | T | A | C | T | G | T | A | A | C | G |
| i | *1Z | | 0.008 | | | G | A | A | G | G | T | G | T | C | G | C | A | A | A | T | T | A | C | T | G | T | A | A | C | G |
| i | *1AA | | 0.008 | | | G | A | A | G | G | T | G | T | C | G | C | A | A | A | T | T | A | C | T | G | T | A | A | C | G |
| i | *1AB | | 0.008 | | | G | A | A | G | G | T | G | T | C | G | C | A | A | A | C | T | A | C | A | T | A | A | C | G | G |
| o | *1AC | | | 0.125 | 0.0167* | G | A | A | G | G | T | G | T | C | G | C | A | A | A | T | T | A | C</ | | | | | | | |

| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|------|--|--|-------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| i | *1AN | | | 0.008 | G | A | A | G | G | G | T | G | T | C | G | C | A | A | A | T | T | A | T | G | T | A | G | C | G |
| i | *1AO | | | 0.008 | G | A | A | G | G | G | T | G | T | C | G | C | A | A | A | T | T | A | T | G | T | A | G | C | G |
| i | *1AP | | | 0.008 | G | A | A | G | G | G | T | G | T | C | G | C | A | A | A | T | T | A | T | G | T | A | G | C | G |
| i | *2A | | | 0.008 | G | A | A | G | G | G | T | G | T | C | G | C | A | A | A | T | T | A | T | G | T | A | G | C | C |

Supplementary Table 2. Human *NT5C3* haplotypes. Variant nucleotides compared with the "reference sequence" (i.e., the most common sequence in African American subjects) are highlighted as white on black. Initial haplotype designations (*1 and *2) were made on the basis of amino acid sequence, with the WT allozyme designated *1. The *2 designation was used for the sequence that encoded His283. Subsequent assignments/letter designations were made based on descending allele frequencies, starting with haplotypes present in African-American subjects. The symbols I and D at 5'-FR locations (-884), (-267) and (-258) represent insertions and deletions, respectively. Polymorphisms at positions 5'FR(-496), 5'UTR(-194), 5'UTR(-134), I1(-5694), I1(-5654), I1(-5125), I1(-19), I2(-307), I3(103), I5(533) and I5(647) were excluded from this table because they were not represented in any of the haplotypes listed. We used a 1% frequency cutoff for inclusion in the table, but haplotypes containing synonymous or nonsynonymous SNP with frequencies lower than 1% were also included. o = observed data; i = inferred data.

Supplementary Figure 1. Human *NT5C3* linkage disequilibrium in AA, CA, HCA and MA subjects. D' values were calculated for each polymorphism pair. All values shown in color were statistically significant ($p < 0.05$). Numbers identifying individual polymorphisms are those listed in **Table 1**.



Supplemental Figure 1