Table S1. Characteristics of included studies for *MTHFR* 677 and 1298 loci distributions (82 papers).

| First Author | Year | Ethnicity -Country | MTHFR 677  | MTHFR 1298 | Quality Score  |
| --- | --- | --- | --- | --- | --- |
| Cases, n (%) | Controls, source (s), n (%) | Cases n (%) | Controls n (%) |
| CC | CT | TT | s | CC | CT | TT | HWE | AA | AC | CC | AA | AC | CC |  HWE |
| **Australia** |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| Beetstra | 2008 | Caucasian-Australia | 9 (47.4) | 6(31.6) | 4 (21) | 1 | 29 (72.5) | 6(15) | 5(12.5) | No | 4(19) | 17(81) | No data | 15(36.6) | 26(63.4) | 0(0) | No | 17(5, 6, 6) |
| **Europe**  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| Weiner | 2010 | Caucasian-Russia | 399(47.7) | 364(43.5) | 74(8.8) | 1 | 386(49.6) | 326(41.9) | 66(8.5) | Yes | 398(47.9) | 353(42.5) | 80(9.6) | 379(48.3) | 330(42) | 76(9.7) | Yes | 15(7, 3, 5) |
| Ericson | 2009 | Caucasian-Sweden | 255(47.2) | 235(43.5) | 50(9.3) | 1 | 531(49.4) | 452(42.1) | 91(8.5) | Yes | 242(44.7) | 242(44.7) | 57(10.6) | 487(45.4) | 480(44.8) | 105(9.8) | Yes | 22(7, 9, 6) |
| Forsti | 2004 | Caucasian-Finland, Poland | 134(60.1) | 81(36.3) | 8(3.6) | 1 | 181(60.7) | 104(34.9) | 13(4.4) | Yes | 94(42.2) | 102(45.7) | 27(12.1) | 133(44.6) | 127(42.6) | 38(12.8) | Yes | 9 (2, 3, 4) |
| Jakubowska | 2008 | Caucasian-Poland | 154(49.1) | 126(40.1) | 34(10.8) | 1 | 134(46.2) | 138(47.6) | 18(6.2) | No | 151(47.3) | 134(42) | 34(10.7) | 117(40.3) | 144(49.7) | 29(10) | Yes | 21(6, 9, 6) |
| Lissowska | 2007 | Caucasian-Poland | 982(49.7) | 815(41.3) | 177(9) | 1 | 1132(49.6) | 915(40.1) | 235(10.3) | No # | 892(44.9) | 874(44) | 220(11.1) | 1086(47.7) | 941(41.3) | 251(11) | No # | 21(8, 7, 6) |
| Ozen | 2013 | Caucasian-Turkey | 28(54.9) | 18(35.3) | 5(9.8) | 1 | 76(71.7) | 3028.3 | 0(0) | Yes | 17(33.3) | 29(56.9) | 5(9.8 | 71(67) | 35(33) | 0(0) | No # | 22(7, 9, 6) |
| Cam | 2009 | Caucasian-Turkey | 48(43.6) | 49(44.6) | 13(11.8) | 1 | 47(49.5) | 42(44.2) | 6(6.3) | Yes | NA | 9(3, 4, 2) |
| Hekim | 2007 | Caucasian-Turkey | 22(55) | 16(40) | 2(5) | 1 | 38(55.9) | 26(38.2) | 4(5.9) | Yes | NA | 11(6, 2, 3) |
| Deligezer | 2005 | Caucasian-Turkey | 98(51.8) | 68(36) | 23(12.2) | 1 | 128(57.4) | 83(37.2) | 12(5.4) | Yes | NA | 14(7, 2, 5) |
| Ergul | 2003 | Caucasian-Turkey | 60(50.8) | 41(34.8) | 17(14.4) | 1 | 94(48.7) | 87(45.1) | 12(6.2) | Yes | 50(42.4) | 48(40.7) | 20(16.9) | 90(46.6) | 85(44.1) | 18(9.3) | Yes | 16(7, 5, 4) |
| Kakkoura | 2015 | Caucasian-Cyprus | 361(33.9) | 516(48.4) | 188(17.7) | 1 | 437(37.8) | 526(45.5) | 194(16.7) | Yes | 138(12.9) | 465(43.4) | 468(43.7) | 150(13.2) | 501(44.1) | 486(42.7) | Yes | 19(6, 7, 6) |
| Papandreou | 2012 | Caucasian-Greece | 105(35) | 150(50) | 45(15) | 1 | 99(35) | 161(56.9) | 23(8.1) | No # | 129(43) | 135(45) | 36(12) | 136(48) | 116(41) | 31(11) | Yes | 15(6, 3, 6) |
| Kalemi | 2005 | Caucasian-Greece | 19(45.2) | 16(38.1) | 7(16.7) | 1 | 23(45.1) | 20(39.2) | 8(15.7) | Yes | NA | 15(7, 2, 6) |
| Langsenlehner | 2008 | Caucasian-Austria | 40(38.1) | 48(45.7) | 17(16.2) | 1 | 51(48.6) | 43(40.9) | 11(10.5) | Yes | NA | 7(5, 0, 2) |
| Grieu | 2004 | Caucasian-Austria | 166(49.7) | 141(42.2) | 27(8.1) | 1 | 242(43.9) | 259(47) | 50(9.1) | Yes | NA | 17(6, 7, 4) |
| Langsenlehner | 2003 | Caucasian-Austria | 208(42.1) | 222(44.9) | 64(13) | 1 | 215(43.4) | 215(43.4) | 65(13.2) | Yes | NA | 17(6, 6, 5) |
| Cerne | 2011 | Caucasian-Slovenia | 222(42.5) | 238(45.6) | 62(11.9) | 1 | 108(40.1) | 124(46.1) | 37(13.8) | Yes | 258(49.2) | 219(41.8) | 47(9) | 131(48.7) | 117(43.5) | 21(7.8) | Yes | 24(9, 9, 6) |
| Reljic | 2007 | Caucasian-Croatia | 40(43) | 44(47.3) | 9(9.7) | 1 | 27(41.5) | 34(52.3) | 4(6.2) | Yes | NA | 20(6, 9, 5) |
| Justenhoven | 2005 | Caucasian-Germany | 249(42.6) | 274(47) | 61(10.4) | 1 | 261(41.2) | 279(44.1) | 93(14.7) | Yes | 273 (46.9) | 256(44) | 53(9.1) | 295(46.5) | 266(42) | 73(11.5) | Yes | 17(8, 3, 6) |
| Jakubowska | 2012 | Caucasian-Great Britain | 2032(42.5) | 2166(45.3) | 580(12.2) | 2 | 1447(43.2) | 1481(44.2) | 422(12.6) | Yes | NA | 12(5, 1, 6) |
| Campbell | 2002 | Caucasian-UK | 140(41.8) | 162(48.4) | 33(9.8) | 1 | 118(50.6) | 92(39.5) | 23(9.9) | Yes | NA | 15(7, 4, 4) |
| Sharp | 2002 | Caucasian-UK | 30(55.6) | 19(35.2) | 5(9.2) | 1 | 25(43.9) | 21(36.8) | 11(19.3) | Yes | 27(49.1) | 25(45.5) | 3(5.4) | 24(40) | 25(41.7) | 11(18.3) | Yes | 19(8, 6, 5) |
| Ferroni | 2009 | Caucasian-Italy | 7(38.9) | 7(38.9) | 4(22.2) | 1 | 18(38.3) | 25(53.2) | 4(8.5) | Yes | 9(50) | 8(44.4) | 1(5.6) | 31(66) | 14(29.8) | 2(4.2) | Yes | 19(7, 6, 6) |
| Macis | 2007 | Caucasian-Italy | 14(30.4) | 20(43.5) | 12(26.1) | 1 | 28(35) | 41(51.2) | 11(13.8) | Yes | NA | 20(7, 7, 6) |
| Henriquez-Hernandez | 2009 | Caucasian-Spain | 52(38.5) | 65(48.2) | 18(13.3) | 1 | 107(36.6) | 138(47.3) | 47(16.1) | Yes | NA | 21(8, 8, 5) |
| Guillem | 2007 | Caucasian-Spain | 53(37.3) | 61(43) | 28(19.7) | 1 | 56(38.9) | 67(46.5) | 21(14.6) | Yes | 66(46.5) | 60(42.2) | 16(11.3) | 69(48.2) | 61(42.7) | 13(9.1) | Yes | 17(6, 5, 6) |
| **North America** |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| Kotsopoulos | 2008 | Caucasian-Canada | 383(40.6) | 421(44.6) | 140(14.8) | 1 | 252(37.1) | 341(50.1) | 87(12.8) | Yes # | 466(49.5) | 390(41.5) | 85(9) | 398(51) | 309(39.6) | 73(9.4) | Yes # | 22(7, 9, 6) |
| Bentley | 2010 | Mixed-US | 346(36.8) | 402(42.8) | 191(20.3) | 1 | 429(35) | 592(48.3) | 205(16.7) | Yes | NA | 14(3, 5, 6) |
| Platek | 2009 | Mixed-US | 429(43.2) | 446(44.9) | 119(11.9) | 1 | 788(43.7) | 795(44.1) | 219(12.2) | Yes | 443(47.7) | 402(43.3) | 83(9) | 842(47.3) | 758(42.6) | 181(10.1) | Yes | 25(9, 10, 6) |
| Maruti | 2009 | Mixed-US | 133(41.8) | 139(43.7) | 46(14.5) | 1 | 301(46.5) | 284(43.9) | 62(9.6) | Yes | NA | 22(6, 10, 6) |
| Tao | 2009 | Mixed-US | 969(45.1) | 915(42.5) | 267(12.4) | 1 | 813(43.9) | 816(44.1) | 223(12) | Yes | 927(46.7) | 861(43.4) | 198(9.9) | 864(47.2) | 779(42.5) | 188(10.3) | Yes | 22(10, 6, 6) |
| Stevens | 2007 | Mixed-US | 208(42.1) | 224(45.3) | 62(12.6) | 1 | 236(47.8) | 193(39) | 65(13.2) | No | 224(45.3) | 228(46.2) | 42(8.5) | 252(51.1) | 201(40.8) | 40(8.1) | Yes # | 21(6, 9, 6) |
| Chen | 2005 | Mixed-US | 398(37.4) | 476(44.8) | 189(17.8) | 1 | 440(39.9) | 509(46.1) | 155(14) | Yes # | 558(52.5) | 417(39.3) | 87(8.2) | 536(48.6) | 457(41.4) | 110(10) | Yes # | 24(8, 10, 6) |
| Le Marchand | 2004 | Mixed-US | 573(48.2) | 479(40.3) | 137(11.5) | 1 | 1211(50.2) | 920(38.1) | 283(11.7) | No | 741(62.3) | 371(31.2) | 77(6.5) | 1493(61.8) | 801(33.2) | 120(5) | Yes # | 24(7, 11, 6) |
| Semenza | 2003 | Mixed-US | 42(40) | 58(55.2) | 5(4.8) | 3 | 112(45.3) | 111(44.9) | 24(9.8) | Yes | NA | 20(8, 6, 6) |
| Ramos-Silva | 2015 | Hispanic-Mexico | 216(43) | 178(36) | 103(21) | 1 | 167(49) | 140(41) | 32(10) | Yes | NA | 16(4, 6, 6) |
| **South America** |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| Barbosa | 2012 | Mixed-Brazil | 65(45.8) | 66(46.5) | 11(7.7) | 1 | 74(52.1) | 53(37.3) | 15(10.6) | Yes | 55(38.7) | 71(50) | 16(11.3) | 62(43.7) | 72(50.7) | 8(5.6) | No # | 21(5, 10, 6) |
| Carvalho | 2012 | Mixed-Brazil | 76(43.2) | 83(47.2) | 17(9.6) | 1 | 87(49.4) | 70(39.8) | 19(10.8) | Yes | 68(41.2) | 80(48.5) | 17(10.3) | 72(43.6) | 84(50.9) | 9(5.5) | No # | 22(9, 7, 6) |
| Batschauer | 2011 | Mixed-Brazil | 27(39.7) | 34(50) | 7(10.3) | 1 | 42(49.4) | 34(40) | 9(10.6) | Yes | NA | 17(8, 3, 6) |
| Ma | 2009 | Mixed-Brazil | 225(49.1) | 188(41.1) | 45(9.8) | 1 | 222(48.5) | 187(40.8) | 49(10.7) | Yes | 269(58.7) | 168(36.7) | 21(4.6) | 279(60.9) | 157(34.3) | 22(4.8) | Yes | 24(8, 10, 6) |
| Lopez-Cortes | 2015 | Mixed-Ecuador | 25(21.9) | 75(65.8) | 14(12.3) | 1 | 69(35.4) | 113(57.9) | 13(6.7) | No | 110(96.5) | 3(2.6) | 1(0.9) | 191(98) | 3(1.5) | 1(0.5) | No | 21(8, 7, 6) |
| **Asia** |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| Ma | 2009 | Asian-Japan | 124(32) | 183(47.1) | 81(20.9) | 1 | 115(29.7) | 188(48.6) | 84(21.7) | Yes | 254(65.5) | 119(30.6) | 15(3.9) | 256(66.1) | 116(30) | 15(3.9) | Yes | 23(8, 9, 6) |
| Suzuki | 2008 | Asia-Japan | 150(33) | 220(48.5) | 84(18.5) | 1 | 338(37.2) | 425(46.8) | 146(16) | Yes | NA | 25(10, 9, 6) |
| Lee | 2004 | Asia-Korea | 58(31.2) | 96(51.6) | 32(17.2) | 1 | 50(34) | 80(54.4) | 17(11.6) | Yes | NA | 22(8, 8, 6) |
| Huang | 2014 | Asia-Taiwan | 596(48.4) | 533(43.3) | 103(8.3) | 1 | 538(43.7) | 519(42.1) | 175(14.2) | No | 796(64.6) | 391(31.7) | 45(3.7) | 787(63.9) | 386(31.3) | 59(4.8) | Yes # | 13(4, 3, 6) |
| Cheng | 2008 | Asia-Taiwan | 185(53) | 133(38.1) | 31(8.9) | 1 | 268(50.6) | 221(41.7) | 41(7.7) | Yes | 207(59) | 125(35.6) | 19(5.4) | 310(58) | 207(38.8) | 17(3.2) | No | 24(9, 9, 6) |
| Yu | 2007 | Asia-Taiwan | 56(51.4) | 44(40.4) | 9(8.2) | 1 | 225(53.6) | 170(40.5) | 25(5.9) | Yes | NA | 24(8, 10, 6) |
| Chou | 2006 | Asia-Taiwan | 73(51.4) | 51(35.9) | 18(12.7) | 1 | 132(46.3) | 120(42.1) | 33(11.6) | Yes | 104(73.3) | 30(21.1) | 8(5.6) | 172(60.4) | 95(33.3) | 18(6.3) | Yes | 24(8, 11, 5) |
| Lin | 2004 | Asia-Taiwan | 43(48.9) | 38(43.2) | 7(7.9) | 1 | 173(50.6) | 145(42.4) | 24(7) | Yes | NA | 24(8, 10, 6) |
| Lin | 2015 | Asia-China | 142(61.4) | 70(30) | 20(8.6) | 1 | 131(55.3) | 82(34.6) | 24(10.1) | No# | NA | 22(5, 11, 6) |
| Lu | 2015 | Asia-China | 170(30.4) | 288(51.4) | 102(18.2) | 1 | 226(40.4) | 250(44.6) | 84(15) | Yes | 369(65.9) | 172(30.7) | 19(3.4) | 352(62.9) | 185(33) | 23(4.1) | Yes | 23(8, 9, 6) |
| He | 2014 | Asia-China | 159(51.3) | 97(31.3) | 54(17.4) | 1 | 220(57.7) | 117(30.7) | 44(11.6) | No | 138(44.5) | 132(42.6) | 40(12.9) | 173(45.4) | 155(40.7) | 53(13.9) | Yes # | 18(5, 8, 5) |
| Xi | 2014 | Asia-China | 489(59.8) | 279(34.1) | 50(6.1) | 1 | 497(58.6) | 298(35.1) | 53(6.3) | Yes# | NA | 24(7,11,6) |
| Jiang-Hua | 2014 | Asian-China | 241(45) | 189(35.4) | 105(19.6) | 1 | 365(54.3) | 226(33.5) | 82(12.2) | No# | 258(48.3) | 235(43.9) | 42(7.8) | 351(52.2) | 280(41.5) | 42(6.3) | Yes | 21(6, 9, 6) |
| Wang | 2014 | Asian-China | 250(57.5) | 153(35.2) | 32(7.3) | 1 | 255(58.6) | 150(34.5) | 30(6.9) | Yes | 206(47.4) | 176(40.5) | 53(12.1) | 214(49.3) | 172(39.5) | 49(11.2) | Yes | 19(7, 7, 5) |
| Weiwei | 2014 | Asian-China | 156(52.5) | 97(32.7) | 44(14.8) | 1 | 185(60.4) | 93(30.4) | 28(9.2) | No# | 135(45.6) | 129(43.6) | 32(10.8) | 151(49.3) | 130(42.5) | 25(8.2) | Yes | 19(4, 10, 5) |
| Liu | 2013 | Asian-China | 250(57.5) | 153(35.2) | 32(7.3) | 1 | 255(58.6) | 150(34.5) | 30(6.9) | Yes | 206(47.4) | 176(40.4) | 53(12.2) | 214(49.2) | 172(39.5) | 49(11.3) | Yes | 21(6, 10, 5) |
| Wu | 2012 | Asian-China | 32(42.7) | 30(40) | 13(17.3) | 1 | 37(49.3) | 32(42.7) | 6(8) | Yes | 37(49.3) | 32(42.7) | 6(8) | 42(56) | 28(37.3) | 5(6.7) | Yes | 17(6, 6, 5) |
| Hua | 2011 | Asian-China | 65(68.4) | 21(22.1) | 9(9.5) | 1 | 52(57.8) | 27(30) | 11(12.2) | No | 50(52.6) | 42(44.2) | 3(3.2) | 55(61.1) | 32(35.6) | 3(3.3) | Yes | 13(4, 5, 4) |
| Gao | 2009 | Asian-China | 202(32.4) | 305(48.9) | 117(18.7) | 1 | 235(37.7) | 301(48.2) | 88(14.1) | Yes | 446(71.5) | 169(27.1) | 9(1.4) | 425(68.1) | 188(30.1) | 11(1.8) | Yes | 21(9, 6, 6) |
| Li | 2009 | Asian-China | 38(58.5) | 17(26.1) | 10(15.4) | 1 | 90(62.9) | 50(35) | 3(2.1) | Yes | NA | 18(6, 6, 6) |
| Yuan | 2009 | Asian-China | 16(20) | 35(43.8) | 29(36.2) | 1 | 32(40) | 35(43.8) | 13(16.2) | Yes | NA | 18(5, 9, 4) |
| Jin | 2009 | Asian-China | 92(37.2) | 115(46.6) | 40(16.2) | 1 | 49(49) | 41(41) | 10(10) | Yes | NA | 24(9, 10, 5) |
| Kan | 2007 | Asian-China | 74(59.2) | 29(23.2) | 22(17.6) | 1 | 65(63.1) | 29(28.2) | 9(8.7) | No | 70(56) | 41(32.8) | 14(11.2) | 61(60.4) | 32(31.7) | 8(7.9) | Yes | 8(3, 2, 3) |
| Qi | 2004 | Asian-China | 42(19.4) | 104(47.9) | 71(32.7 | 1 | 59(27) | 105(48.2) | 54(24.8) | Yes | 155(71.4) | 58(26.7) | 4(1.9) | 144(66) | 71(32.6) | 3(1.4) | Yes | 20(6, 10, 4) |
| Shrubsole | 2004 | Asian-China | 374(33.6) | 555(49.9) | 183(16.5) | 1 | 387(33.4) | 577(49.7) | 196(16.9) | Yes | 768(68.5) | 311(27.7) | 42(3.8) | 824(68.2) | 344(28.5) | 40(3.3) | Yes | 21(6, 9, 6) |
| Inoue | 2008 | Asian-Singapore | 239(62.9) | 120(31.6) | 21(5.5) | 1 | 393(59.4) | 226(34.1) | 43(6.5) | Yes | 225(59.2) | 139(36.6) | 16(4.2) | 387(58.5) | 234(35.3) | 41(6.2) | Yes | 19(6, 7, 6) |
| Sangrajrang | 2010 | Asian-Thailand | 410(72.8) | 144(25.6) | 9(1.6) | 1 | 366(75.2) | 110(22.6) | 11(2.2) | Yes | 302(53.6) | 223(39.6) | 38(6.8) | 258(53) | 206(42.3) | 23(4.7) | No | 23(7, 10, 6) |
| Pooja | 2015 | Asian-India | 437(74.3) | 134(22.8) | 17(2.9) | 1 | 386(75.9) | 111(21.9) | 11(2.2) | Yes | NA | 17(8, 3, 6) |
| Prasad | 2011 | Asian-India | 124(95.4) | 5(3.8) | 1(0.8) | 1 | 116(92.8) | 8(6.4) | 1(0.8) | Yes | NA | 12(3, 5, 4) |
| Naushad | 2010 | Asian-India | 185(75.8) | 56(23) | 3(1.2) | 1 | 205(84) | 39(16) | 0(0) | Yes | NA | 23(7, 10, 6) |
| Mohammad | 2010 | Asian-India | 168(75.7) | 53(23.9) | 1(0.4) | 1 | 198(84.3) | 37(15.7) | 0(0) | Yes | NA | 18(6, 6, 6) |
| Mir | 2008 | Asian-India | 29(82.8) | 6(17.2) | 0(0) | 1 | 19(57.6) | 12(36.4) | 2(6) | Yes | 15(42.8) | 19(54.3) | 1(2.9) | 11(33.3) | 22(66.7) | 0(0) | No | 15(6, 5, 4) |
| Kalyankumar | 2006 | Asian-India | 45(51.1) | 37(42.1) | 6(6.8) | 1 | 61(64.2) | 31(32.6) | 3(3.2) | Yes | 49(55.7) | 33(37.5) | 6(6.8) | 65(68.4) | 26(27.4) | 4(4.2) | Yes | 13(7, 3, 3) |
| **Middle East Asia** |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| Akilzhanova | 2013 | Mid-Eastern -Kazakhstan | 181(57.5) | 109(34.6) | 25(7.9) | 1 | 287(47.5) | 269(44.5) | 48(8) | Yes | 138(43.8) | 142(45.1) | 35(11.1) | 318(52.6) | 242(40.1) | 44(7.3) | Yes | 21(9, 7, 5) |
| Akram | 2012 | Mid-Eastern-Pakistan | 65(59.1) | 25(22.7) | 20(18.2) | 1 | 55(50) | 45(40.9) | 10(9.1) | Yes | 35(31.8) | 55(50) | 20(18.2) | 30(27.3) | 75(68.2) | 5(4.5) | No | 12(3, 6, 3) |
| Hosseini | 2011 | Mid-Eastern -Iran | 168(57.1) | 84(28.6) | 42(14.3) | 1 | 150(50) | 90(30) | 60(20) | No | 162(55.1) | 96(32.7) | 36(12.2) | 105(35) | 135(45) | 60(20) | Yes | 13(5, 5, 3) |
| Lajin | 2012 | Mid-Eastern -Syria | 60(50.4) | 47(39.5) | 12(10.1) | 1 | 58(46) | 58(46) | 10(8) | Yes | 44(37) | 52(43.7) | 23(19.3) | 65(51.6) | 48(38.1) | 13(10.3) | Yes | 13(6, 3, 4) |
| Awwad | 2015 | Mid-Eastern -Jordan | 66(44) | 69(46) | 15(10) | 1 | 79(54.1) | 51(34.9) | 16(11) | Yes | 68(46.6) | 61(41.8) | 17(11.6) | 58(43) | 64(47.4) | 13(9.6) | Yes | 18(5, 8, 5) |
| Alshatwi | 2010 | Mid-Eastern-Saudi Arabia | 336(32.8) | 516(50.4) | 172(16.8) | 1 | 351(33.2) | 532(50.3) | 174(16.5) | Yes | NA | 22(8, 9, 5) |
| **Africa** |  |  |  |  |  |  |  |  |  |  |
| Diakite | 2012 | African-Morocco | 39(40.6) | 51(53.1) | 6(6.3) | 1 | 69(59) | 41(35) | 7(6) | Yes | NA | 19(7, 6, 6) |

HWE: Hardy Weinberg Equilibrium

#: HWE data updated (noted different from reports of original studies) based on available online calculator, retrieved at

http://www.tufts.edu/~mcourt01/Documents/Court%20lab%20-%20HW%20calculator.xls

Sources of controls: 1 = healthy adults, 2 = BRCA 1 or 2 carriers, 3 = clinic-based with breast diseases

Quality score ranges: 0 - 29 (external validity, 0 – 11; Internal Validity, 0 – 12; report quality, 0 - 6)

NA: Not available

UK: United Kingdom

US: United States of America

**Reference List of Studies included in the Meta-analysis**

19 Meta-analysis Papers

1. Xie SZ, Liu ZZ, Yu JH, Liu L, Wang W, Xie DL, Qin JB (2015). Association between the MTHFR C677T polymorphism and risk of cancer: Evidence from 446 case-control studies. *Tumour Biol,* 36(11):8953-72. doi: 10.1007/s13277-015-3648-z PMID: 26081619.
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