

## REFERENCES USED IN ALGORITHMS FOR THE TREATMENT OF PERSONS WITH DEPRESSION

1. American Psychological Association: Clinical practice guideline for the treatment of depression across three age cohorts. 2019.
2. Quality AfHRA: Second-Generation Antidepressants in the Pharmacologic Treatment of Adult Depression: An Update of the 2007 Comparative Effectiveness Review. In: Effective Health Care Program. Rockville, MD: AHRQ; 2019.
3. Kennedy SH, Lam RW, McIntyre RS, Tourjman SV, Bhat V, Blier P, Hasnain M, Jollant F, Levitt AJ, MacQueen GM et al: Canadian Network for Mood and Anxiety Treatments (CANMAT) 2016 Clinical Guidelines for the Management of Adults with Major Depressive Disorder: Section 3. Pharmacological Treatments. *Can J Psychiatry* 2016, 61(9):540-560.
4. Wagner G, Schultes MT, Titscher V, Teufer B, Klerings I, Gartlehner G: Efficacy and safety of levomilnacipran, vilazodone and vortioxetine compared with other second-generation antidepressants for major depressive disorder in adults: A systematic review and network meta-analysis. *J Affect Disord* 2018, 228:1-12.
5. Bauer M, Bschor T, Pfennig A, Whybrow PC, Angst J, Versiani M, Moller HJ: World Federation of Societies of Biological Psychiatry (WFSBP) Guidelines for Biological Treatment of Unipolar Depressive Disorders in Primary Care. *World J Biol Psychiatry* 2007, 8(2):67-104.
6. Wenzel-Seifert K, Wittmann M, Haen E: QTc prolongation by psychotropic drugs and the risk of Torsade de Pointes. *Dtsch Arztebl Int* 2011, 108(41):687-693.
7. Henssler J, Heinz A, Brandt L, Bschor T: Antidepressant Withdrawal and Rebound Phenomena. *Dtsch Arztebl Int* 2019, 116(20):355-361.
8. National Institute for Health and Care Excellence: Guideline on the Treatment and Management of Depression in Adults [NICE Guideline 90]. 2009.
9. American Psychiatric Association: Practice Guideline for the Treatment of Patients with Major Depressive Disorder. In: APA Guideline. Third Edition edn: American Psychiatric Association; 2010.
10. Guidelines B: Major Depressive Issues in Adults: Appendix D. 2013.
11. VA/DoD The Management of Major Depressive Disorder Working Group: VA/DoD Clinical Practice Guideline for the Management of Major Depressive Disorder. Online: Department of Veterans Affairs, Department of Defense; 2016.
12. Faquih AE, Memon RI, Hafeez H, Zeshan M, Naveed S: A Review of Novel Antidepressants: A Guide for Clinicians. *Cureus* 2019, 11(3):e4185.
13. Cipriani A, Furukawa TA, Salanti G, Chaimani A, Atkinson LZ, Ogawa Y, Leucht S, Ruhe HG, Turner EH, Higgins JPT et al: Comparative efficacy and acceptability of 21 antidepressant drugs for the acute treatment of adults with major depressive disorder: a systematic review and network meta-analysis. *Lancet* 2018, 391(10128):1357-1366.

14. McIntyre R, Gommoll C, Chen C, Ruth A: The efficacy of levomilnacipran ER across symptoms of major depressive disorder: a post hoc analysis of 5 randomized, double-blind, placebo-controlled trials. *CNS spectrums* 2016, -1:1-8.
15. McIntyre RS: The role of new antidepressants in clinical practice in Canada: a brief review of vortioxetine, levomilnacipran ER, and vilazodone. *Neuropsychiatr Dis Treat* 2017, 13:2913-2919.
16. Serretti A: The Present and Future of Precision Medicine in Psychiatry: Focus on Clinical Psychopharmacology of Antidepressants. *Clinical Psychopharmacology and Neuroscience* 2018, 16:1-6.
17. Malhi GS, Bassett D, Boyce P, Bryant R, Fitzgerald PB, Fritz K, Hopwood M, Lyndon B, Mulder R, Murray G et al: Royal Australian and New Zealand College of Psychiatrists clinical practice guidelines for mood disorders. *Aust N Z J Psychiatry* 2015, 49(12):1087-1206.
18. Fasipe OJ: The emergence of new antidepressants for clinical use: Agomelatine paradox versus other novel agents. *IBRO Rep* 2019, 6:95-110.
19. Wang SM, Han C, Lee SJ, Patkar AA, Masand PS, Pae CU: Vilazodone for the Treatment of Depression: An Update. *Chonnam Med J* 2016, 52(2):91-100.
20. Mathews M, Gommoll C, Chen D, Nunez R, Khan A: Efficacy and safety of vilazodone 20 and 40 mg in major depressive disorder: a randomized, double-blind, placebo-controlled trial. *Int Clin Psychopharmacol* 2015, 30(2):67-74.
21. Koesters M, Ostuzzi G, Guaiana G, Breilmann J, Barbui C: Vortioxetine for depression in adults. *Cochrane Database Syst Rev* 2017, 7:Cd011520.
22. Long JD: Vortioxetine for Depression in Adults. *Issues in Mental Health Nursing* 2019, 40(9):819-820.
23. Young AH, Evitt L, Brignone M, Diamand F, Atsou K, Campbell R, Cure S, Danchenko N: Cost-utility evaluation of vortioxetine in patients with Major Depressive Disorder experiencing inadequate response to alternative antidepressants in the United Kingdom. *J Affect Disord* 2017, 218:291-298.
24. Thase ME, Danchenko N, Brignone M, Florea I, Diamand F, Jacobsen PL, Vieta E: Comparative evaluation of vortioxetine as a switch therapy in patients with major depressive disorder. *European Neuropsychopharmacology* 2017, 27(8):773-781.
25. Cleare A, Pariante C, Young A, Anderson I, Christmas D, Cowen P, Dickens C, Ferrier IN, Geddes J, Gilbody S et al: Evidence-based guidelines for treating depressive disorders with antidepressants: A revision of the 2008 British Association for Psychopharmacology guidelines (BAP). *Journal of psychopharmacology (Oxford, England)* 2015, 29.
26. Lam RW, Kennedy SH, Grigoriadis S, McIntyre RS, Milev R, Ramasubbu R, Parikh SV, Patten SB, Ravindran AV, Canadian Network for M et al: Canadian Network for Mood and Anxiety Treatments (CANMAT) clinical guidelines for the management of major depressive disorder in adults. III. Pharmacotherapy. *J Affect Disord* 2009, 117 Suppl 1:S26-43.
27. Papakostas G, Fava M: A Metaanalysis of Clinical Trials Comparing Moclobemide with Selective Serotonin Reuptake Inhibitors for the Treatment of Major Depressive Disorder. *Canadian journal of psychiatry Revue canadienne de psychiatrie* 2006, 51:783-790.

28. Lotufo-Neto F, Trivedi M, Thase ME: Meta-analysis of the reversible inhibitors of monoamine oxidase type A moclobemide and brofaromine for the treatment of depression. *Neuropsychopharmacology* 1999, 20(3):226-247.
29. Linde K, Kriston L, Rucker G, Jamil S, Schumann I, Meissner K, Sigterman K, Schneider A: Efficacy and acceptability of pharmacological treatments for depressive disorders in primary care: systematic review and network meta-analysis. *Ann Fam Med* 2015, 13(1):69-79.
30. Cristancho MA, Thase ME: Critical appraisal of selegiline transdermal system for major depressive disorder. *Expert Opin Drug Deliv* 2016, 13(5):659-665.
31. National Institute for Health and Care Excellence: Depression in adults: recognition and management (summary) [NICE Guideline 90]. In: NICE Clinical Guideline. 2009 (2019 update).
32. Kasper S, Pail G: Milnacipran: a unique antidepressant? *Neuropsychiatr Dis Treat* 2010, 6:23-31.
33. Papakostas GI, Fava M: A meta-analysis of clinical trials comparing milnacipran, a serotonin--norepinephrine reuptake inhibitor, with a selective serotonin reuptake inhibitor for the treatment of major depressive disorder. *Eur Neuropsychopharmacol* 2007, 17(1):32-36.
34. Nakagawa A, Watanabe N, Omori IM, Barbui C, Cipriani A, McGuire H, Churchill R, Furukawa TA: Milnacipran versus other antidepressive agents for depression. *Cochrane Database Syst Rev* 2009(3):CD006529-CD006529.
35. Important Safety Information Regarding the Discontinuation of Sales of Nefazodone in Canada [<https://www.healthycanadians.gc.ca/recall-alert-rappel-avis/hc-sc/2003/14224a-eng.php>]
36. Bennabi D, Charpeaud T, Yroni A, Genty JB, Destouches S, Lancrenon S, Alaili N, Bellivier F, Bougerol T, Camus V et al: Clinical guidelines for the management of treatment-resistant depression: French recommendations from experts, the French Association for Biological Psychiatry and Neuropsychopharmacology and the fondation FondaMental. *BMC Psychiatry* 2019, 19(1):262.
37. Zhou X, Ravindran AV, Qin B, Del Giovane C, Li Q, Bauer M, Liu Y, Fang Y, da Silva T, Zhang Y et al: Comparative efficacy, acceptability, and tolerability of augmentation agents in treatment-resistant depression: systematic review and network meta-analysis. *J Clin Psychiatry* 2015, 76(4):e487-498.
38. Strawbridge R, Carter B, Marwood L, Bandelow B, Tsapekos D, Nikolova VL, Taylor R, Mantingh T, de Angel V, Patrick F et al: Augmentation therapies for treatment-resistant depression: systematic review and meta-analysis. *Br J Psychiatry* 2019, 214(1):42-51.
39. Zhou X, Keitner GI, Qin B, Ravindran AV, Bauer M, Del Giovane C, Zhao J, Liu Y, Fang Y, Zhang Y et al: Atypical Antipsychotic Augmentation for Treatment-Resistant Depression: A Systematic Review and Network Meta-Analysis. *Int J Neuropsychopharmacol* 2015, 18(11):pyv060.
40. Komossa K, Depping AM, Gaudchau A, Kissling W, Leucht S: Second-generation antipsychotics for major depressive disorder and dysthymia. *Cochrane Database Syst Rev* 2010(12):Cd008121.

41. Spielmans GI, Berman MI, Linardatos E, Rosenlicht NZ, Perry A, Tsai AC: Adjunctive atypical antipsychotic treatment for major depressive disorder: a meta-analysis of depression, quality of life, and safety outcomes. *PLoS Med* 2013, 10(3):e1001403.
42. Citrome L: Adjunctive aripiprazole, olanzapine, or quetiapine for major depressive disorder: an analysis of number needed to treat, number needed to harm, and likelihood to be helped or harmed. *Postgrad Med* 2010, 122(4):39-48.
43. Nelson JC, Papakostas GI: Atypical antipsychotic augmentation in major depressive disorder: a meta-analysis of placebo-controlled randomized trials. *Am J Psychiatry* 2009, 166(9):980-991.
44. Mohamed S, Johnson GR, Chen P, Hicks PB, Davis LL, Yoon J, Gleason TC, Vertrees JE, Weingart K, Tal I et al: Effect of Antidepressant Switching vs Augmentation on Remission Among Patients With Major Depressive Disorder Unresponsive to Antidepressant Treatment: The VAST-D Randomized Clinical Trial. *Jama* 2017, 318(2):132-145.
45. Hobart M, Skuban A, Zhang P, Josiassen MK, Hefting N, Augustine C, Brewer C, Sanchez R, McQuade RD: Efficacy and safety of flexibly dosed brexpiprazole for the adjunctive treatment of major depressive disorder: a randomized, active-referenced, placebo-controlled study. *Curr Med Res Opin* 2018, 34(4):633-642.
46. Papadimitropoulou K, Vossen C, Karabis A, Donatti C, Kubitz N: Comparative efficacy and tolerability of pharmacological and somatic interventions in adult patients with treatment-resistant depression: a systematic review and network meta-analysis. *Curr Med Res Opin* 2017, 33(4):701-711.
47. Bauer M, Dell'osso L, Kasper S, Pitchot W, Dencker Vansvik E, Kohler J, Jorgensen L, Montgomery SA: Extended-release quetiapine fumarate (quetiapine XR) monotherapy and quetiapine XR or lithium as add-on to antidepressants in patients with treatment-resistant major depressive disorder. *Journal of affective disorders* 2013, 151(1):209-219.
48. Fang Y, Yuan C, Xu Y, Chen J, Wu Z, Cao L, Yi Z, Hong W, Wang Y, Jiang K et al: A pilot study of the efficacy and safety of paroxetine augmented with risperidone, valproate, buspirone, trazodone, or thyroid hormone in adult Chinese patients with treatment-resistant major depression. *J Clin Psychopharmacol* 2011, 31(5):638-642.
49. Edwards SJ, Hamilton V, Nherera L, Trevor N: Lithium or an atypical antipsychotic drug in the management of treatment-resistant depression: a systematic review and economic evaluation. *Health Technol Assess* 2013, 17(54):1-190.
50. Turner P, Kantaria R, Young AH: A systematic review and meta-analysis of the evidence base for add-on treatment for patients with major depressive disorder who have not responded to antidepressant treatment: a European perspective. *J Psychopharmacol* 2014, 28(2):85-98.
51. Yoshimura R, Hori H, Umene-Nakano W, Ikenouchi-Sugita A, Katsuki A, Atake K, Nakamura J: Comparison of lithium, aripiprazole and olanzapine as augmentation to paroxetine for inpatients with major depressive disorder. *Ther Adv Psychopharmacol* 2014, 4(3):123-129.
52. Cheon EJ, Lee KH, Park YW, Lee JH, Koo BH, Lee SJ, Sung HM: Comparison of the Efficacy and Safety of Aripiprazole Versus Bupropion Augmentation in Patients With Major Depressive Disorder Unresponsive to

Selective Serotonin Reuptake Inhibitors: A Randomized, Prospective, Open-Label Study. *J Clin Psychopharmacol* 2017, 37(2):193-199.

53. Doree JP, Des Rosiers J, Lew V, Gendron A, Elie R, Stip E, Tourjman SV: Quetiapine augmentation of treatment-resistant depression: a comparison with lithium. *Curr Med Res Opin* 2007, 23(2):333-341.
54. Barbee JG, Conrad EJ, Jamhour NJ: The effectiveness of olanzapine, risperidone, quetiapine, and ziprasidone as augmentation agents in treatment-resistant major depressive disorder. *J Clin Psychiatry* 2004, 65(7):975-981.
55. Nierenberg AA, Fava M, Trivedi MH, Wisniewski SR, Thase ME, McGrath PJ, Alpert JE, Warden D, Luther JF, Niederehe G et al: A comparison of lithium and T(3) augmentation following two failed medication treatments for depression: a STAR\*D report. *Am J Psychiatry* 2006, 163(9):1519-1530; quiz 1665.
56. Joffe RT, Singer W, Levitt AJ, MacDonald C: A placebo-controlled comparison of lithium and triiodothyronine augmentation of tricyclic antidepressants in unipolar refractory depression. *Arch Gen Psychiatry* 1993, 50(5):387-393.
57. Joffe RT, Sokolov ST, Levitt AJ: Lithium and triiodothyronine augmentation of antidepressants. *Can J Psychiatry* 2006, 51(12):791-793.
58. Schindler F, Angheliescu IG: Lithium versus lamotrigine augmentation in treatment resistant unipolar depression: a randomized, open-label study. *Int Clin Psychopharmacol* 2007, 22(3):179-182.
59. Trivedi MH, Fava M, Wisniewski SR, Thase ME, Quitkin F, Warden D, Ritz L, Nierenberg AA, Lebowitz BD, Biggs MM et al: Medication augmentation after the failure of SSRIs for depression. *N Engl J Med* 2006, 354(12):1243-1252.
60. Wang HR, Woo YS, Ahn HS, Ahn IM, Kim HJ, Bahk WM: Can Atypical Antipsychotic Augmentation Reduce Subsequent Treatment Failure More Effectively Among Depressed Patients with a Higher Degree of Treatment Resistance? A Meta-Analysis of Randomized Controlled Trials. *Int J Neuropsychopharmacol* 2015, 18(8).
61. Luan S, Wan H, Zhang L, Zhao H: Efficacy, acceptability, and safety of adjunctive aripiprazole in treatment-resistant depression: a meta-analysis of randomized controlled trials. *Neuropsychiatr Dis Treat* 2018, 14:467-477.
62. Berman RM, Marcus RN, Swanink R, McQuade RD, Carson WH, Corey-Lisle PK, Khan A: The efficacy and safety of aripiprazole as adjunctive therapy in major depressive disorder: a multicenter, randomized, double-blind, placebo-controlled study. *J Clin Psychiatry* 2007, 68(6):843-853.
63. Berman RM, Fava M, Thase ME, Trivedi MH, Swanink R, McQuade RD, Carson WH, Adson D, Taylor L, Hazel J et al: Aripiprazole augmentation in major depressive disorder: a double-blind, placebo-controlled study in patients with inadequate response to antidepressants. *CNS Spectr* 2009, 14(4):197-206.
64. Marcus RN, McQuade RD, Carson WH, Hennicken D, Fava M, Simon JS, Trivedi MH, Thase ME, Berman RM: The efficacy and safety of aripiprazole as adjunctive therapy in major depressive disorder: a second multicenter, randomized, double-blind, placebo-controlled study. *J Clin Psychopharmacol* 2008, 28(2):156-165.

65. Nelson JC, Thase ME, Trivedi MH, Fava M, Han J, Van Tran Q, Pikalov A, Qi Y, Carlson BX, Marcus RN et al: Safety and Tolerability of Adjunctive Aripiprazole in Major Depressive Disorder: A Pooled Post Hoc Analysis (studies CN138-139 and CN138-163). *Prim Care Companion J Clin Psychiatry* 2009, 11(6):344-352.
66. Fava M, Mischoulon D, Iosifescu D, Witte J, Pencina M, Flynn M, Harper L, Levy M, Rickels K, Pollack M: A double-blind, placebo-controlled study of aripiprazole adjunctive to antidepressant therapy among depressed outpatients with inadequate response to prior antidepressant therapy (ADAPT-A Study). *Psychother Psychosom* 2012, 81(2):87-97.
67. Kamijima K, Higuchi T, Ishigooka J, Ohmori T, Ozaki N, Kanba S, Kinoshita T, Koyama T: Aripiprazole augmentation to antidepressant therapy in Japanese patients with major depressive disorder: a randomized, double-blind, placebo-controlled study (ADMIRE study). *J Affect Disord* 2013, 151(3):899-905.
68. Ozaki N, Otsubo T, Kato M, Higuchi T, Ono H, Kamijima K: Efficacy of aripiprazole augmentation in Japanese patients with major depressive disorder: a subgroup analysis and Montgomery-Asberg Depression Rating Scale and Hamilton Rating Scale for Depression item analyses of the Aripiprazole Depression Multicenter Efficacy study. *Psychiatry Clin Neurosci* 2015, 69(1):34-42.
69. Kamijima K, Yasuda M, Yamamura K, Fukuta Y: Real-world effectiveness and safety of aripiprazole augmentation therapy in patients with major depressive disorder. *Curr Med Res Opin* 2018, 34(12):2105-2112.
70. Clayton AH, Baker RA, Sheehan JJ, Cain ZJ, Forbes RA, Marler SV, Marcus R, Berman RM, Thase ME: Comparison of adjunctive use of aripiprazole with bupropion or selective serotonin reuptake inhibitors/serotonin-norepinephrine reuptake inhibitors: analysis of patients beginning adjunctive treatment in a 52-week, open-label study. *BMC Res Notes* 2014, 7:459.
71. Nelson JC, Rahman Z, Laubmeier KK, Eudicone JM, McQuade RD, Berman RM, Marcus RN, Baker RA, Sheehan JJ: Efficacy of adjunctive aripiprazole in patients with major depressive disorder whose symptoms worsened with antidepressant monotherapy. *CNS Spectr* 2014, 19(6):528-534.
72. Muzina DJ, Chambers JS, Camacho TA, Eudicone JM, Forbes RA, Berman RM, Baker RA: Adjunctive aripiprazole for depression: predictive value of early assessment. *Am J Manag Care* 2011, 17(12):793-801.
73. Mischoulon D, Witte J, Levy M, Papakostas GI, Pet LR, Hsieh WH, Pencina MJ, Ward S, Pollack MH, Fava M: Efficacy of dose increase among nonresponders to low-dose aripiprazole augmentation in patients with inadequate response to antidepressant treatment: a randomized, double-blind, placebo-controlled, efficacy trial. *J Clin Psychiatry* 2012, 73(3):353-357.
74. Stahl SM: Mechanism of action of brexpiprazole: comparison with aripiprazole. *CNS Spectr* 2016, 21(1):1-6.
75. Yoon S, Jeon SW, Ko YH, Patkar AA, Masand PS, Pae CU, Han C: Adjunctive Brexpiprazole as a Novel Effective Strategy for Treating Major Depressive Disorder: A Systematic Review and Meta-Analysis. *J Clin Psychopharmacol* 2017, 37(1):46-53.
76. Citrome L: Brexpiprazole for schizophrenia and as adjunct for major depressive disorder: a systematic review of the efficacy and safety profile for this newly approved antipsychotic - what is the number

needed to treat, number needed to harm and likelihood to be helped or harmed? *Int J Clin Pract* 2015, 69(9):978-997.

77. Thase M, Youakim J, Skuban A, Hobart M, Augustine C, Zhang P, McQuade R, Carson W, Nyilas M, Sanchez R et al: Efficacy and Safety of Adjunctive Brexpiprazole 2 mg in Major Depressive Disorder: A Phase 3, Randomized, Placebo-Controlled Study in Patients With Inadequate Response to Antidepressants. *The Journal of clinical psychiatry* 2015, 76.
78. Thase M, Youakim J, Skuban A, Hobart M, Zhang P, McQuade R, Nyilas M, Carson W, Sanchez R, Eriksson H: Adjunctive Brexpiprazole 1 and 3 mg for Patients With Major Depressive Disorder Following Inadequate Response to Antidepressants. *The Journal of clinical psychiatry* 2015, 76.
79. Hobart M, Skuban A, Zhang P, Augustine C, Brewer C, Hefting N, Sanchez R, McQuade RD: A Randomized, Placebo-Controlled Study of the Efficacy and Safety of Fixed-Dose Brexpiprazole 2 mg/d as Adjunctive Treatment of Adults With Major Depressive Disorder. *J Clin Psychiatry* 2018, 79(4).
80. Hobart M, Zhang P, Skuban A, Brewer C, Hefting N, Sanchez R, McQuade RD: A Long-Term, Open-Label Study to Evaluate the Safety and Tolerability of Brexpiprazole as Adjunctive Therapy in Adults With Major Depressive Disorder. *J Clin Psychopharmacol* 2019, 39(3):203-209.
81. Bauer M, Hefting N, Lindsten A, Josiassen MK, Hobart M: A randomised, placebo-controlled 24-week study evaluating adjunctive brexpiprazole in patients with major depressive disorder. *Acta Neuropsychiatr* 2019, 31(1):27-35.
82. Bobo WV, Shelton RC: Olanzapine and fluoxetine combination therapy for treatment-resistant depression: review of efficacy, safety, and study design issues. *Neuropsychiatr Dis Treat* 2009, 5:369-383.
83. Shelton R, Tollefson G, Tohen M, Stahl S, Gannon K, Jacobs T, Buras W, Bymaster F, Zhang W, Spencer K et al: A Novel Augmentation Strategy for Treating Resistant Major Depression. *The American journal of psychiatry* 2001, 158:131-134.
84. Shelton RC, Williamson DJ, Corya SA, Sanger TM, Van Campen LE, Case M, Briggs SD, Tollefson GD: Olanzapine/fluoxetine combination for treatment-resistant depression: a controlled study of SSRI and nortriptyline resistance. *J Clin Psychiatry* 2005, 66(10):1289-1297.
85. Thase ME, Corya SA, Osuntokun O, Case M, Henley DB, Sanger TM, Watson SB, Dube S: A randomized, double-blind comparison of olanzapine/fluoxetine combination, olanzapine, and fluoxetine in treatment-resistant major depressive disorder. *J Clin Psychiatry* 2007, 68(2):224-236.
86. Parker G, Brotchie H, Parker K: Is combination olanzapine and antidepressant medication associated with a more rapid response trajectory than antidepressant alone? *Am J Psychiatry* 2005, 162(4):796-798.
87. Corya SA, Andersen SW, Detke HC, Kelly LS, Van Campen LE, Sanger TM, Williamson DJ, Dube S: Long-term antidepressant efficacy and safety of olanzapine/fluoxetine combination: a 76-week open-label study. *J Clin Psychiatry* 2003, 64(11):1349-1356.
88. Bobo WV, Shelton RC: Fluoxetine and olanzapine combination therapy in treatment-resistant major depression: review of efficacy and safety data. *Expert Opin Pharmacother* 2009, 10(13):2145-2159.

89. Boku S, Inoue T, Honma H, Honma H, Nakagawa S, Koyama T: Olanzapine augmentation of milnacipran for stage 2 treatment-resistant major depression: an open study. *Hum Psychopharmacol* 2011, 26(3):237-241.
90. Bauer M, Pretorius HW, Constant EL, Earley WR, Szamosi J, Brecher M: Extended-release quetiapine as adjunct to an antidepressant in patients with major depressive disorder: results of a randomized, placebo-controlled, double-blind study. *J Clin Psychiatry* 2009, 70(4):540-549.
91. El-Khalili N, Joyce M, Atkinson S, Buynak RJ, Datto C, Lindgren P, Eriksson H: Extended-release quetiapine fumarate (quetiapine XR) as adjunctive therapy in major depressive disorder (MDD) in patients with an inadequate response to ongoing antidepressant treatment: a multicentre, randomized, double-blind, placebo-controlled study. *Int J Neuropsychopharmacol* 2010, 13(7):917-932.
92. Garakani A, Martinez JM, Marcus S, Weaver J, Rickels K, Fava M, Hirschowitz J: A randomized, double-blind, and placebo-controlled trial of quetiapine augmentation of fluoxetine in major depressive disorder. *Int Clin Psychopharmacol* 2008, 23(5):269-275.
93. Bauer M, Demyttenaere K, El-Khalili N, Thase ME, Papakostas GI, Szamosi J, Earley WR, Eriksson H: Pooled analysis of adjunct extended-release quetiapine fumarate in patients with major depressive disorder according to ongoing SSRI or SNRI treatment. *Int Clin Psychopharmacol* 2014, 29(1):16-25.
94. Mahmoud RA, Pandina GJ, Turkoz I, Kosik-Gonzalez C, Canuso CM, Kujawa MJ, Gharabawi-Garibaldi GM: Risperidone for treatment-refractory major depressive disorder: a randomized trial. *Annals of internal medicine* 2007, 147(9):593-602.
95. Keitner GI, Garlow SJ, Ryan CE, Ninan PT, Solomon DA, Nemeroff CB, Keller MB: A randomized, placebo-controlled trial of risperidone augmentation for patients with difficult-to-treat unipolar, non-psychotic major depression. *J Psychiatr Res* 2009, 43(3):205-214.
96. Rapaport MH, Gharabawi GM, Canuso CM, Mahmoud RA, Keller MB, Bossie CA, Turkoz I, Lasser RA, Loeschler A, Bouhours P et al: Effects of risperidone augmentation in patients with treatment-resistant depression: Results of open-label treatment followed by double-blind continuation. *Neuropsychopharmacology* 2006, 31(11):2505-2513.
97. Owenby RK, Brown LT, Brown JN: Use of risperidone as augmentation treatment for major depressive disorder. *Ann Pharmacother* 2011, 45(1):95-100.
98. Bauer M, Adli M, Ricken R, Severus E, Pilhatsch M: Role of lithium augmentation in the management of major depressive disorder. *CNS Drugs* 2014, 28(4):331-342.
99. Crossley NA, Bauer M: Acceleration and augmentation of antidepressants with lithium for depressive disorders: two meta-analyses of randomized, placebo-controlled trials. *J Clin Psychiatry* 2007, 68(6):935-940.
100. Nelson JC, Baumann P, Delucchi K, Joffe R, Katona C: A systematic review and meta-analysis of lithium augmentation of tricyclic and second generation antidepressants in major depression. *J Affect Disord* 2014, 168:269-275.
101. Alevizos B, Alevizos E, Leonardou A, Zervas I: Low dosage lithium augmentation in venlafaxine resistant depression: an open-label study. *Psychiatriki* 2012, 23(2):143-148.



102. Sugawara H, Sakamoto K, Harada T, Ishigooka J: Predictors of efficacy in lithium augmentation for treatment-resistant depression. *J Affect Disord* 2010, 125(1-3):165-168.
103. Nierenberg AA, Papakostas GI, Petersen T, Montoya HD, Worthington JJ, Tedlow J, Alpert JE, Fava M: Lithium augmentation of nortriptyline for subjects resistant to multiple antidepressants. *J Clin Psychopharmacol* 2003, 23(1):92-95.
104. Valerio MP, Martino DJ: Differential response to lithium between melancholic and non-melancholic unipolar depression. *Psychiatry Res* 2018, 269:183-184.
105. Harter M, Klesse C, Bermejo I, Schneider F, Berger M: Unipolar depression: diagnostic and therapeutic recommendations from the current S3/National Clinical Practice Guideline. *Dtsch Arztebl Int* 2010, 107(40):700-708.
106. Cooper-Kazaz R, Lerer B: Efficacy and safety of triiodothyronine supplementation in patients with major depressive disorder treated with specific serotonin reuptake inhibitors. *Int J Neuropsychopharmacol* 2008, 11(5):685-699.
107. Abraham G, Milev R, Stuart Lawson J: T3 augmentation of SSRI resistant depression. *J Affect Disord* 2006, 91(2-3):211-215.
108. Aronson R, Offman HJ, Joffe RT, Naylor CD: Triiodothyronine augmentation in the treatment of refractory depression. A meta-analysis. *Arch Gen Psychiatry* 1996, 53(9):842-848.
109. Sokolov ST, Levitt AJ, Joffe RT: Thyroid hormone levels before unsuccessful antidepressant therapy are associated with later response to T3 augmentation. *Psychiatry Res* 1997, 69(2-3):203-206.
110. Garlow SJ, Dunlop BW, Ninan PT, Nemeroff CB: The combination of triiodothyronine (T3) and sertraline is not superior to sertraline monotherapy in the treatment of major depressive disorder. *J Psychiatr Res* 2012, 46(11):1406-1413.
111. Posternak M, Novak S, Stern R, Hennessey J, Joffe R, Prange A, Jr., Zimmerman M: A pilot effectiveness study: placebo-controlled trial of adjunctive L-triiodothyronine (T3) used to accelerate and potentiate the antidepressant response. *Int J Neuropsychopharmacol* 2008, 11(1):15-25.
112. Kelly TF, Lieberman DZ: Long term augmentation with T3 in refractory major depression. *J Affect Disord* 2009, 115(1-2):230-233.
113. Cooke RG, Joffe RT, Levitt AJ: T3 augmentation of antidepressant treatment in T4-replaced thyroid patients. *J Clin Psychiatry* 1992, 53(1):16-18.
114. Rosenthal LJ, Goldner WS, O'Reardon JP: T3 augmentation in major depressive disorder: safety considerations. *Am J Psychiatry* 2011, 168(10):1035-1040.
115. Henssler J, Bschor T, Baethge C: Combining Antidepressants in Acute Treatment of Depression: A Meta-Analysis of 38 Studies Including 4511 Patients. *Canadian journal of psychiatry Revue canadienne de psychiatrie* 2016, 61(1):29-43.

116. Keks NA, Burrows GD, Copolov DL, Newton R, Paoletti N, Schweitzer I, Tiller J: Beyond the evidence: is there a place for antidepressant combinations in the pharmacotherapy of depression? *Med J Aust* 2007, 186(3):142-144.
117. Blier P, Ward HE, Tremblay P, Laberge L, Hebert C, Bergeron R: Combination of antidepressant medications from treatment initiation for major depressive disorder: a double-blind randomized study. *Am J Psychiatry* 2010, 167(3):281-288.
118. Jha MK, Malchow AL, Grannemann BD, Rush AJ, Trivedi MH: Do baseline sub-threshold hypomanic symptoms affect acute-phase antidepressant outcome in outpatients with major depressive disorder? Preliminary findings from the randomized CO-MED trial. *Neuropsychopharmacology* 2018, 43(11):2197-2203.
119. Rush AJ, Trivedi MH, Stewart JW, Nierenberg AA, Fava M, Kurian BT, Warden D, Morris DW, Luther JF, Husain MM et al: Combining medications to enhance depression outcomes (CO-MED): acute and long-term outcomes of a single-blind randomized study. *Am J Psychiatry* 2011, 168(7):689-701.
120. Rocha FL, Fuzikawa C, Riera R, Hara C: Combination of antidepressants in the treatment of major depressive disorder: a systematic review and meta-analysis. *J Clin Psychopharmacol* 2012, 32(2):278-281.
121. Patel K, Allen S, Haque MN, Angelescu I, Baumeister D, Tracy DK: Bupropion: a systematic review and meta-analysis of effectiveness as an antidepressant. *Ther Adv Psychopharmacol* 2016, 6(2):99-144.
122. Gulrez G, Badyal DK, Deswal RS, Sharma A: Bupropion as an augmenting agent in patients of depression with partial response. *Basic Clin Pharmacol Toxicol* 2012, 110(3):227-230.
123. Fornaro M, Martino M, Mattei C, Prestia D, Vinciguerra V, De Berardis D, De Pasquale C, Iasevoli F, Mungo S, Fornaro P: Duloxetine-bupropion combination for treatment-resistant atypical depression: a double-blind, randomized, placebo-controlled trial. *Eur Neuropsychopharmacol* 2014, 24(8):1269-1278.
124. Leuchter AF, Lesser IM, Trivedi MH, Rush AJ, Morris DW, Warden D, Fava M, Wisniewski SR, Luther JF, Perales M et al: An open pilot study of the combination of escitalopram and bupropion-SR for outpatients with major depressive disorder. *J Psychiatr Pract* 2008, 14(5):271-280.
125. DeBattista C, Solvason HB, Poirier J, Kendrick E, Schatzberg AF: A prospective trial of bupropion SR augmentation of partial and non-responders to serotonergic antidepressants. *J Clin Psychopharmacol* 2003, 23(1):27-30.
126. Spier SA: Use of bupropion with SRIs and venlafaxine. *Depress Anxiety* 1998, 7(2):73-75.
127. Papakostas GI, Worthington JJ, 3rd, Iosifescu DV, Kinrys G, Burns AM, Fisher LB, Homberger CH, Mischoulon D, Fava M: The combination of duloxetine and bupropion for treatment-resistant major depressive disorder. *Depress Anxiety* 2006, 23(3):178-181.
128. Nasr S, Wendt B, Popli A, Crayton J: Comparing outcomes of adjunctive treatment in depression: aripiprazole versus bupropion. *J Affect Disord* 2014, 162:50-54.
129. Bodkin JA, Lasser RA, Wines JD, Jr., Gardner DM, Baldessarini RJ: Combining serotonin reuptake inhibitors and bupropion in partial responders to antidepressant monotherapy. *J Clin Psychiatry* 1997, 58(4):137-145.

130. Lam RW, Hossie H, Solomons K, Yatham LN: Citalopram and bupropion-SR: combining versus switching in patients with treatment-resistant depression. *J Clin Psychiatry* 2004, 65(3):337-340.
131. Stewart JW, McGrath PJ, Blondeau C, Deliyannides DA, Hellerstein D, Norris S, Amat J, Pilowsky DJ, Tessier P, Laberge L et al: Combination antidepressant therapy for major depressive disorder: Speed and probability of remission. *Journal of psychiatric research* 2014, 52:7-14.
132. Taylor MJ, Rudkin L, Bullemor-Day P, Lubin J, Chukwujekwu C, Hawton K: Strategies for managing sexual dysfunction induced by antidepressant medication. *Cochrane Database Syst Rev* 2013(5):Cd003382.
133. SwitchRx [<https://www.switchrx.com/antidepressants.php/switch>]
134. Blier P, Gobbi G, Turcotte J, Montigny C, Boucher N, Hébert C, Debonnel G: Mirtazapine and paroxetine in major depression: A comparison of monotherapy versus their combination from treatment initiation. *European neuropsychopharmacology : the journal of the European College of Neuropsychopharmacology* 2009, 19:457-465.
135. Kessler D, Burns A, Tallon D, Lewis G, MacNeill S, Round J, Hollingworth W, Chew-Graham C, Anderson I, Campbell J et al: Combining mirtazapine with SSRIs or SNRIs for treatment-resistant depression: the MIR RCT. *Health Technol Assess* 2018, 22(63):1-136.
136. Carpenter LL, Yasmin S, Price LH: A double-blind, placebo-controlled study of antidepressant augmentation with mirtazapine. *Biol Psychiatry* 2002, 51(2):183-188.
137. Goss AJ, Kaser M, Costafreda SG, Sahakian BJ, Fu CH: Modafinil augmentation therapy in unipolar and bipolar depression: a systematic review and meta-analysis of randomized controlled trials. *J Clin Psychiatry* 2013, 74(11):1101-1107.
138. Candy B, Jones L, Williams R, Tookman A, King M: Psychostimulants for depression. *Cochrane Database of Systematic Reviews* 2008(2).
139. Fava M, Thase ME, DeBattista C: A multicenter, placebo-controlled study of modafinil augmentation in partial responders to selective serotonin reuptake inhibitors with persistent fatigue and sleepiness. *J Clin Psychiatry* 2005, 66(1):85-93.
140. The Management of Major Depressive Disorder Work Group: VA/DoD Clinical Practice Guideline for the Management of Major Depressive Disorder. Online: Department of Veterans Affairs, Department of Defense; 2022.
141. Papakostas GI, Salloum NC, Hock RS, Jha MK, Murrough JW, Mathew SJ, Iosifescu DV, Fava M: Efficacy of Esketamine Augmentation in Major Depressive Disorder: A Meta-Analysis. *J Clin Psychiatry* 2020, 81(4).
142. Citrome L, DiBernardo A, Singh J: Appraising esketamine nasal spray for the management of treatment-resistant depression in adults: Number needed to treat, number needed to harm, and likelihood to be helped or harmed. *J Affect Disord* 2020, 271:228-238.
143. Zheng W, Cai DB, Xiang YQ, Zheng W, Jiang WL, Sim K, Ungvari GS, Huang X, Huang XX, Ning YP et al: Adjunctive intranasal esketamine for major depressive disorder: A systematic review of randomized double-blind controlled-placebo studies. *J Affect Disord* 2020, 265:63-70.

144. McIntyre RS, Carvalho IP, Lui LMW, Majeed A, Masand PS, Gill H, Rodrigues NB, Lipsitz O, Coles AC, Lee Y et al: The effect of intravenous, intranasal, and oral ketamine in mood disorders: A meta-analysis. *J Affect Disord* 2020, 276:576-584.
145. Esketamine for treatment-resistant depression. *Drug and Therapeutics Bulletin* 2020, 58(12):183-188.
146. Wajs E, Aluisio L, Holder R, Daly EJ, Lane R, Lim P, George JE, Morrison RL, Sanacora G, Young AH et al: Esketamine Nasal Spray Plus Oral Antidepressant in Patients With Treatment-Resistant Depression: Assessment of Long-Term Safety in a Phase 3, Open-Label Study (SUSTAIN-2). *J Clin Psychiatry* 2020, 81(3).
147. Ochs-Ross R, Daly EJ, Zhang Y, Lane R, Lim P, Morrison RL, Hough D, Manji H, Drevets WC, Sanacora G et al: Efficacy and Safety of Esketamine Nasal Spray Plus an Oral Antidepressant in Elderly Patients With Treatment-Resistant Depression-TRANSFORM-3. *Am J Geriatr Psychiatry* 2020, 28(2):121-141.
148. Solmi M, Veronese N, Zaninotto L, Loos M, Gao K, Schaffer A, Reis C, Normann C, Angheliescu I, Correll C: Lamotrigine compared to placebo and other agents with antidepressant activity in patients with unipolar and bipolar depression: A comprehensive meta-analysis of efficacy and safety outcomes in short-term trials. *CNS spectrums* 2016, 21:403-418.
149. Joffe RT, Singer W: A comparison of triiodothyronine and thyroxine in the potentiation of tricyclic antidepressants. *Psychiatry Res* 1990, 32(3):241-251.
150. Papakostas GI, Fava M, Baer L, Swee MB, Jaeger A, Bobo WV, Shelton RC: Ziprasidone Augmentation of Escitalopram for Major Depressive Disorder: Efficacy Results From a Randomized, Double-Blind, Placebo-Controlled Study. *Am J Psychiatry* 2015, 172(12):1251-1258.
151. Fava M, Rosenbaum JF, McGrath PJ, Stewart JW, Amsterdam JD, Quitkin FM: Lithium and tricyclic augmentation of fluoxetine treatment for resistant major depression: a double-blind, controlled study. *Am J Psychiatry* 1994, 151(9):1372-1374.
152. Fava M, Alpert J, Nierenberg A, Lagomasino I, Sonawalla S, Tedlow J, Worthington J, Baer L, Rosenbaum JF: Double-blind study of high-dose fluoxetine versus lithium or desipramine augmentation of fluoxetine in partial responders and nonresponders to fluoxetine. *J Clin Psychopharmacol* 2002, 22(4):379-387.
153. Coppen A, Bailey J: Enhancement of the antidepressant action of fluoxetine by folic acid: a randomised, placebo controlled trial. *J Affect Disord* 2000, 60(2):121-130.
154. Kelly CB, McDonnell AP, Johnston TG, Mulholland C, Cooper SJ, McMaster D, Evans A, Whitehead AS: The MTHFR C677T polymorphism is associated with depressive episodes in patients from Northern Ireland. *J Psychopharmacol* 2004, 18(4):567-571.
155. Papakostas GI, Shelton RC, Zajecka JM, Etemad B, Rickels K, Clain A, Baer L, Dalton ED, Sacco GR, Schoenfeld D et al: L-methylfolate as adjunctive therapy for SSRI-resistant major depression: results of two randomized, double-blind, parallel-sequential trials. *Am J Psychiatry* 2012, 169(12):1267-1274.
156. Shelton RC, Sloan Manning J, Barrentine LW, Tipton EV: Assessing Effects of L-Methylfolate in Depression Management: Results of a Real-World Patient Experience Trial. *Prim Care Companion CNS Disord* 2013, 15(4).

157. Passeri M, Cucinotta D, Abate G, Senin U, Ventura A, Stramba Badiale M, Diana R, La Greca P, Le Grazie C: Oral 5'-methyltetrahydrofolic acid in senile organic mental disorders with depression: results of a double-blind multicenter study. *Aging (Milano)* 1993, 5(1):63-71.
158. Godfrey PS, Toone BK, Carney MW, Flynn TG, Bottiglieri T, Laundry M, Chanarin I, Reynolds EH: Enhancement of recovery from psychiatric illness by methylfolate. *Lancet* 1990, 336(8712):392-395.
159. Guaraldi GP, Fava M, Mazzi F, la Greca P: An open trial of methyltetrahydrofolate in elderly depressed patients. *Ann Clin Psychiatry* 1993, 5(2):101-105.
160. Paton C, Ferrier IN: SSRIs and gastrointestinal bleeding. *BMJ* 2005, 331(7516):529-530.
161. Meijer WE, Heerdink ER, Nolen WA, Herings RM, Leufkens HG, Egberts AC: Association of risk of abnormal bleeding with degree of serotonin reuptake inhibition by antidepressants. *Arch Intern Med* 2004, 164(21):2367-2370.
162. Cheng YL, Hu HY, Lin XH, Luo JC, Peng YL, Hou MC, Lin HC, Lee FY: Use of SSRI, But Not SNRI, Increased Upper and Lower Gastrointestinal Bleeding: A Nationwide Population-Based Cohort Study in Taiwan. *Medicine (Baltimore)* 2015, 94(46):e2022.
163. Wang SM, Han C, Bahk WM, Lee SJ, Patkar AA, Masand PS, Pae CU: Addressing the Side Effects of Contemporary Antidepressant Drugs: A Comprehensive Review. *Chonnam Med J* 2018, 54(2):101-112.
164. Sanis Health Inc: Trazodone. In: Health Canada Monograph. 2017.
165. Allergan: Vilazodone. In: CA Product Monograph. 2019.
166. Lundbeck: Vortioxetine. In: CA Product Monograph. 2019.
167. Dalton SO, Johansen C, Mellekjaer L, Norgard B, Sorensen HT, Olsen JH: Use of selective serotonin reuptake inhibitors and risk of upper gastrointestinal tract bleeding: a population-based cohort study. *Arch Intern Med* 2003, 163(1):59-64.
168. Carvalho AF, Sharma MS, Brunoni AR, Vieta E, Fava GA: The Safety, Tolerability and Risks Associated with the Use of Newer Generation Antidepressant Drugs: A Critical Review of the Literature. *Psychother Psychosom* 2016, 85(5):270-288.
169. Na KS, Jung HY, Cho SJ, Cho SE: Can we recommend mirtazapine and bupropion for patients at risk for bleeding?: A systematic review and meta-analysis. *J Affect Disord* 2018, 225:221-226.
170. Park LT, Zarate CA, Jr.: Depression in the Primary Care Setting. *The New England Journal of Medicine* 2019, 380(23):2279-2280.
171. Henssler J, Kurschus M, Franklin J, Bschor T, Baethge C: Trajectories of Acute Antidepressant Efficacy: How Long to Wait for Response? A Systematic Review and Meta-Analysis of Long-Term, Placebo-Controlled Acute Treatment Trials. *J Clin Psychiatry* 2018, 79(3).
172. Lewis G, Duffy L, Ades A, Amos R, Araya R, Brabyn S, Button KS, Churchill R, Derrick C, Dowrick C et al: The clinical effectiveness of sertraline in primary care and the role of depression severity and duration

- (PANDA): a pragmatic, double-blind, placebo-controlled randomised trial. *The Lancet Psychiatry* 2019, 6(11):903-914.
173. McCarron RM, Shapiro B, Rawles J, Luo J: Depression. *Annals of internal medicine* 2021, 174(5):itc65-itc80.
174. Park LT, Zarate CA, Jr.: Depression in the Primary Care Setting. *N Engl J Med* 2019, 380(6):559-568.
175. Arnold MJ, Fulleborn S, Farrell J: Medications for Treatment-Resistant Depression in Adults. *Am Fam Physician* 2021, 103(1):16-18.
176. Jensen B RL: Antidepressants: Comparison Chart. May 2019 edn. RxFiles.ca: RxFiles Academic Detailing; 2019.
177. Psychiatry.net [<http://wiki.psychiatry.net.nl/index.php/SwitchAntidepressants>]
178. Jensen B RL: Antipsychotics: Comparison Chart. Nov 2019 edn. RxFiles.ca: RxFiles Academic Detailing; 2019.
179. Jibson M: Second-generation antipsychotic medications: Pharmacology, administration, and side effects. In: *UpToDate*. Edited by Marder S; 2019.
180. Muench J, Hamer AM: Adverse effects of antipsychotic medications. *Am Fam Physician* 2010, 81(5):617-622.
181. Arnow B, Blasey C, Williams L, Palmer D, Rekshan W, Schatzberg A, Etkin A, Kulkarni J, Luther J, Rush A: Depression Subtypes in Predicting Antidepressant Response: A Report From the iSPOT-D Trial. *American Journal of Psychiatry* 2015, 172.
182. Jha MK, Minhajuddin A, Gadad BS, Greer T, Grannemann B, Soyombo A, Mayes TL, Rush AJ, Trivedi MH: Can C-reactive protein inform antidepressant medication selection in depressed outpatients? Findings from the CO-MED trial. *Psychoneuroendocrinology* 2017, 78:105-113.
183. Arteaga-Henríquez G, Simon MS, Burger B, Weidinger E, Wijkhuijs A, Arolt V, Birkenhager TK, Musil R, Müller N, Drexhage HA: Low-Grade Inflammation as a Predictor of Antidepressant and Anti-Inflammatory Therapy Response in MDD Patients: A Systematic Review of the Literature in Combination With an Analysis of Experimental Data Collected in the EU-MOODINFLAME Consortium. *Front Psychiatry* 2019, 10:458-458.
184. Keks N, Hope J, Keogh S: Switching and stopping antidepressants. *Aust Prescr* 2016, 39(3):76-83.
185. Henssler J, Heinz A, Brandt L, Bschor T: Antidepressant Withdrawal and Rebound Phenomena. *Dtsch Arztebl Int* 2019, 116(20):355-361.
186. Zafra-Tanaka JH, Goicochea-Lugo S, Villarreal-Zegarra D, Taype-Rondan A: Characteristics and quality of clinical practice guidelines for depression in adults: a scoping review. *BMC Psychiatry* 2019, 19(1):76.
187. Bahr R, Lopez A, Rey JA: Intranasal Esketamine (Spravato(TM)) for Use in Treatment-Resistant Depression In Conjunction With an Oral Antidepressant. *P t* 2019, 44(6):340-375.

188. Andrade C: Relative Efficacy and Acceptability of Antidepressant Drugs in Adults With Major Depressive Disorder: Commentary on a Network Meta-Analysis. *J Clin Psychiatry* 2018, 79(2).
189. Gabriel FC, de Melo DO, Fráguas R, Leite-Santos NC, Mantovani da Silva RA, Ribeiro E: Pharmacological treatment of depression: A systematic review comparing clinical practice guideline recommendations. *PLoS One* 2020, 15(4):e0231700.
190. Powell JG, Garland S, Preston K, Piszczatoski C: Brexanolone (Zulresso): Finally, an FDA-Approved Treatment for Postpartum Depression. *Ann Pharmacother* 2020, 54(2):157-163.
191. Athreya A, Brückl T, Binder E, Rush A, Biernacka J, Frye M, Neavin D, Skime M, Monrad D, Iyer R et al: Prediction of short-term antidepressant response using probabilistic graphical models with replication across multiple drugs and treatment settings. *Neuropsychopharmacology* 2021:1-11.
192. Tomlinson A, Furukawa TA, Efthimiou O, Salanti G, De Crescenzo F, Singh I, Cipriani A: Personalise antidepressant treatment for unipolar depression combining individual choices, risks and big data (PETRUSHKA): rationale and protocol. *Evid Based Ment Health* 2020, 23(2):52-56.
193. Kraus C, Kadriu B, Lanzenberger R, Zarate CA, Jr., Kasper S: Prognosis and improved outcomes in major depression: a review. *Transl Psychiatry* 2019, 9(1):127.
194. Schatzberg AF, Blier P, Delgado PL, Fava M, Haddad PM, Shelton RC: Antidepressant discontinuation syndrome: consensus panel recommendations for clinical management and additional research. *J Clin Psychiatry* 2006, 67 Suppl 4:27-30.
195. National Institute for Health and Care Excellence: Depression in Adults with a Chronic Physical Health Problem: Treatment and Management. 2010.
196. Baldwin DS, Montgomery SA, Nil R, Lader M: Discontinuation symptoms in depression and anxiety disorders. *Int J Neuropsychopharmacol* 2007, 10(1):73-84.
197. Fava M: Prospective studies of adverse events related to antidepressant discontinuation. *J Clin Psychiatry* 2006, 67 Suppl 4:14-21.
198. Taylor D, Stewart S, Connolly A: Antidepressant withdrawal symptoms-telephone calls to a national medication helpline. *J Affect Disord* 2006, 95(1-3):129-133.
199. American Psychiatric Association: Practice Guideline for the Treatment of Patients with Major Depressive Disorder. Third Edition edn: American Psychiatric Association; 2010.
200. Perahia DG, Pritchett YL, Kajdasz DK, Bauer M, Jain R, Russell JM, Walker DJ, Spencer KA, Froud DM, Raskin J et al: A randomized, double-blind comparison of duloxetine and venlafaxine in the treatment of patients with major depressive disorder. *J Psychiatr Res* 2008, 42(1):22-34.
201. Gartlehner G, Hansen RA, Morgan LC, Thaler K, Lux L, Van Noord M, Mager U, Thieda P, Gaynes BN, Wilkins T et al: Comparative benefits and harms of second-generation antidepressants for treating major depressive disorder: an updated meta-analysis. *Annals of internal medicine* 2011, 155(11):772-785.
202. National Institute for Health and Care Excellence: Depression: Evidence Update. 2012.

203. Schueler YB, Koesters M, Wieseler B, Grouven U, Kromp M, Kerekes MF, Kreis J, Kaiser T, Becker T, Weinmann S: A systematic review of duloxetine and venlafaxine in major depression, including unpublished data. *Acta psychiatrica Scandinavica* 2011, 123(4):247-265.
204. Hicks JK, Bishop JR, Sangkuhl K, Muller DJ, Ji Y, Leckband SG, Leeder JS, Graham RL, Chiulli DL, A LL et al: Clinical Pharmacogenetics Implementation Consortium (CPIC) Guideline for CYP2D6 and CYP2C19 Genotypes and Dosing of Selective Serotonin Reuptake Inhibitors. *Clinical pharmacology and therapeutics* 2015, 98(2):127-134.
205. Nierenberg AA, Farabaugh AH, Alpert JE, Gordon J, Worthington JJ, Rosenbaum JF, Fava M: Timing of onset of antidepressant response with fluoxetine treatment. *Am J Psychiatry* 2000, 157(9):1423-1428.
206. Nierenberg AA, McLean NE, Alpert JE, Worthington JJ, Rosenbaum JF, Fava M: Early nonresponse to fluoxetine as a predictor of poor 8-week outcome. *Am J Psychiatry* 1995, 152(10):1500-1503.
207. Szegedi A, Jansen WT, van Willigenburg AP, van der Meulen E, Stassen HH, Thase ME: Early improvement in the first 2 weeks as a predictor of treatment outcome in patients with major depressive disorder: a meta-analysis including 6562 patients. *J Clin Psychiatry* 2009, 70(3):344-353.
208. Szegedi A, Muller MJ, Angheliescu I, Klawe C, Kohlen R, Benkert O: Early improvement under mirtazapine and paroxetine predicts later stable response and remission with high sensitivity in patients with major depression. *J Clin Psychiatry* 2003, 64(4):413-420.
209. Dreimuller N, Wagner S, Engel A, Braus DF, Roll SC, Elsner S, Tadic A, Lieb K: Predictors of the effectiveness of an early medication change strategy in patients with major depressive disorder. *BMC Psychiatry* 2019, 19(1):24.
210. Bobo WV, Chen H, Trivedi MH, Stewart JW, Nierenberg AA, Fava M, Kurian BT, Warden D, Morris DW, Luther JF et al: Randomized comparison of selective serotonin reuptake inhibitor (escitalopram) monotherapy and antidepressant combination pharmacotherapy for major depressive disorder with melancholic features: a CO-MED report. *J Affect Disord* 2011, 133(3):467-476.
211. Kunugi H, Hori H, Ogawa S: Biochemical markers subtyping major depressive disorder. *Psychiatry Clin Neurosci* 2015, 69(10):597-608.
212. Keller MB: Issues in treatment-resistant depression. *J Clin Psychiatry* 2005, 66 Suppl 8:5-12.
213. Berlim MT, Turecki G: What is the meaning of treatment resistant/refractory major depression (TRD)? A systematic review of current randomized trials. *Eur Neuropsychopharmacol* 2007, 17(11):696-707.
214. MacQueen G, Santaguida P, Keshavarz H, Jaworska N, Levine M, Beyene J, Raina P: Systematic Review of Clinical Practice Guidelines for Failed Antidepressant Treatment Response in Major Depressive Disorder, Dysthymia, and Subthreshold Depression in Adults. *Canadian journal of psychiatry Revue canadienne de psychiatrie* 2017, 62(1):11-23.
215. Rush AJ, Fava M, Wisniewski SR, Lavori PW, Trivedi MH, Sackeim HA, Thase ME, Nierenberg AA, Quitkin FM, Kashner TM et al: Sequenced treatment alternatives to relieve depression (STAR\*D): rationale and design. *Control Clin Trials* 2004, 25(1):119-142.



216. Gaynes BN, Dusetzina SB, Ellis AR, Hansen RA, Farley JF, Miller WC, Sturmer T: Treating depression after initial treatment failure: directly comparing switch and augmenting strategies in STAR\*D. *J Clin Psychopharmacol* 2012, 32(1):114-119.
217. Bauer M, Dopfmer S: Lithium augmentation in treatment-resistant depression: meta-analysis of placebo-controlled studies. *J Clin Psychopharmacol* 1999, 19(5):427-434.
218. Adli M, Wiethoff K, Baghai TC, Fisher R, Seemuller F, Laakmann G, Brieger P, Cordes J, Malevani J, Laux G et al: How Effective Is Algorithm-Guided Treatment for Depressed Inpatients? Results from the Randomized Controlled Multicenter German Algorithm Project 3 Trial. *Int J Neuropsychopharmacol* 2017, 20(9):721-730.
219. Bschor T, Baethge C: No evidence for switching the antidepressant: systematic review and meta-analysis of RCTs of a common therapeutic strategy. *Acta psychiatrica Scandinavica* 2010, 121(3):174-179.
220. Thase ME: No evidence for switching antidepressants in treatment-resistant depressed patients. *Evid Based Med* 2010, 15(4):120-121.
221. Kato T, Furukawa TA, Mantani A, Kurata K, Kubouchi H, Hirota S, Sato H, Sugishita K, Chino B, Itoh K et al: Optimising first- and second-line treatment strategies for untreated major depressive disorder - the SUND study: a pragmatic, multi-centre, assessor-blinded randomised controlled trial. *BMC Med* 2018, 16(1):103.
222. Bose A, Tsai J, Li D: Early non-response in patients with severe depression: escitalopram up-titration versus switch to duloxetine. *Clin Drug Investig* 2012, 32(6):373-385.
223. Irene R, Luis MA, Helena DC, David P, Ramon DJ, Inmaculada G: Switching to duloxetine from selective serotonin reuptake inhibitors in non- or partial responders: Results from a Spanish sample. *Int J Psychiatry Clin Pract* 2009, 13(2):100-108.
224. Simon G: Unipolar major depression in adults: Choosing initial treatment. In: *UpToDate*. Edited by P R-B; 2019.
225. NHS WGotCPGotMoDiAfs: Clinical Practice Guideline on the Management of Depression in Adults. In: *Spanish NHS Guideline*. [guiasalud.es](http://guiasalud.es): Ministry of Health, Social Services and Equality; 2015.
226. Ramanuj P, Ferenchick EK, Pincus HA: Depression in primary care: part 2-management. *Bmj* 2019, 365:l835.
227. Chokka PR, Hankey JR: Assessment and management of sexual dysfunction in the context of depression. *Ther Adv Psychopharmacol* 2018, 8(1):13-23.
228. Solmi M, Murru A, Pacchiarotti I, Undurraga J, Veronese N, Fornaro M, Stubbs B, Monaco F, Vieta E, Seeman MV et al: Safety, tolerability, and risks associated with first- and second-generation antipsychotics: a state-of-the-art clinical review. *Ther Clin Risk Manag* 2017, 13:757-777.
229. Warner CH, Bobo W, Warner C, Reid S, Rachal J: Antidepressant discontinuation syndrome. *Am Fam Physician* 2006, 74(3):449-456.
230. Bhat V, Kennedy SH: Recognition and management of antidepressant discontinuation syndrome. *J Psychiatry Neurosci* 2017, 42(4):E7-e8.

231. Berigan TR, Harazin JS: Bupropion-Associated Withdrawal Symptoms: A Case Report. *Prim Care Companion J Clin Psychiatry* 1999, 1(2):50-51.
232. Baldwin DS, Chrones L, Florea I, Nielsen R, Nomikos GG, Palo W, Reines E: The safety and tolerability of vortioxetine: Analysis of data from randomized placebo-controlled trials and open-label extension studies. *Journal of psychopharmacology (Oxford, England)* 2016, 30(3):242-252.
233. Beach SR, Celano CM, Sugrue AM, Adams C, Ackerman MJ, Noseworthy PA, Huffman JC: QT Prolongation, Torsades de Pointes, and Psychotropic Medications: A 5-Year Update. *Psychosomatics* 2018, 59(2):105-122.
234. Crepeau-Gendron G, Brown HK, Shorey C, Madan R, Szabuniewicz C, Koh S, Veinich S, Mah L: Association between citalopram, escitalopram and QTc prolongation in a real-world geriatric setting. *J Affect Disord* 2019, 250:341-345.
235. Petry N, Lupu R, Gohar A, Larson EA, Peterson C, Williams V, Zhao J, Wilke RA, Hines LJ: CYP2C19 genotype, physician prescribing pattern, and risk for long QT on serotonin selective reuptake inhibitors. *Pharmacogenomics* 2019, 20(5):343-351.
236. Perahia DG, Quail D, Desai D, Corruble E, Fava M: Switching to duloxetine from selective serotonin reuptake inhibitor antidepressants: a multicenter trial comparing 2 switching techniques. *J Clin Psychiatry* 2008, 69(1):95-105.
237. Romera I, Perez V, Menchon JM, Schacht A, Papen R, Neuhauser D, Abbar M, Picard H, Gilaberte I: Early vs. conventional switching of antidepressants in patients with MDD and moderate to severe pain: a double-blind randomized study. *J Affect Disord* 2012, 143(1-3):47-55.
238. Nakajima S, Uchida H, Suzuki T, Watanabe K, Hirano J, Yagihashi T, Takeuchi H, Abe T, Kashima H, Mimura M: Is switching antidepressants following early nonresponse more beneficial in acute-phase treatment of depression?: a randomized open-label trial. *Prog Neuropsychopharmacol Biol Psychiatry* 2011, 35(8):1983-1989.
239. Angita Pharma Inc: AG-Amitriptyline. In: *CA Product Monograph*. Boucherville, QC; 2021.
240. Upsher-Smith Laboratories: Amitriptyline hydrochloride. In: *US Product Monograph*. Maple Grove, MN; 2021.
241. Apotex Inc: Apo-amitriptyline. In: *CA Product Monograph*. Toronto, ON; 2019.
242. Mylan Pharmaceuticals: Amitriptyline. In: *US Product Monograph*. Morgantown, WV; 2016.
243. Odan Laboratories Ltd.: Odan-Bupropion SR. In: *CA Product Monograph*. Pointe Claire, QC; 2021.
244. Sun Pharma Canada Inc.: Taro-Bupropion XL. In: *CA Product Monograph*. Brampton, ON; 2021.
245. Solco Healthcare US LLC: Bupropion hydrochloride tablet extended release (SR). In: *US Product Monograph*. Somerset, NJ; 2022.

246. Graviti Pharmaceuticals Private Limited: Bupropion hydrochloride extended-release tablet (XL). In: US Product Monograph. Wilmington, DE; 2022.
247. Angita Pharma Inc.: Citalopram. In: CA Product Monograph. Boucherville, QC; 2021.
248. Aurobindo Pharma Limited: Citalopram hydrobromide tablet. In: US Product Monograph. East Windsor, NJ; 2022.
249. JAMP Pharma Corporation: Desvenlafaxine Extended Release Tablets. In: CA Product Monograph. Boucherville, QC; 2022.
250. Greenstone LLC: Desvenlafaxine succinate tablet, extended release. In: US Product Monograph. Peapack, NJ; 2022.
251. Sivem Pharmaceuticals ULC: Duloxetine Delayed-Release Capsules. In: CA Product Monograph. Saint-Laurent, QC; 2022.
252. Lupin Pharmaceuticals Inc.: Duloxetine capsule delayed release. In: US Product Monograph. Baltimore, MD; 2022.
253. Lundbeck Canada Inc.: CipraleX (escitalopram). In: CA Product Monograph. St-Laurent, QC; 2022.
254. Lupin Pharmaceuticals Inc.: Escitalopram tablet. In: US Product Monograph. Baltimore, MD; 2022.
255. Accord Healthcare Inc.: ACH-Fluoxetine. In: CA Product Monograph. Kirkland, QC; 2022.
256. Modavar Pharmaceuticals LLC: Fluoxetine. In: US Product Monograph. Washington, DC; 2022.
257. Pro Doc Ltée: Fluvoxamine. In: CA Product Monograph. Laval, QC; 2022.
258. Allergan Inc.: Fetzima (Levomilnacipran Extended Release Capsule). In: CA Product Monograph. Markham, ON; 2021.
259. Allergan Inc.: Fetzima (levomilnacipran hydrochloride capsule extended release). In: US Product Monograph. Madison, NJ; 2021.
260. Apotex Inc.: Apo-mirtazapine. In: CA Product Monograph. Toronto, ON; 2022.
261. Apotex Corp.: Mirtazapine. In: US Product Monograph. Weston, FL; 2022.
262. Bausch health CI: Manerix (moclobemide). In: CA Product Monograph. Laval, QC; 2021.
263. GlaxoSmithKline Inc.: Paxil (paroxetine). In: CA Product Monograph. Mississauga, ON; 2021.
264. GlaxoSmithKline Inc.: Paxil CR (paroxetine controlled release). In: CA Product Monograph. Mississauga, ON; 2021.
265. Apotex Corp.: Paroxetine. In: US Product Monograph. Weston, FL; 2021.
266. Sebelá Pharmaceuticals Inc.: Pexeva (paroxetine mesylate). In: US Product Monograph. Roswell, GA; 2022.

267. Apotex Corp: Paroxetine extended release. In: US Product Monograph. Weston, FL; 2021.
268. Erfa Canada Inc: Nardil (phenelzine). In: CA Product Monograph. Montréal, QC; 2015.
269. Parke-Davis Div of Pfizer Inc: Nardil (phenelzine). In: US Product Monograph. New York, NY; 2020.
270. Mylan Specialty L.P.: Emsam (selegiline patch). In: US Product Monograph. Morgantown, WV; 2020.
271. Pharmascience Inc.: pms-Sertraline. In: CA Product Monograph. Montreal, QC; 2022.
272. Greenstone LLC: Sertraline hydrochloride tablet. In: US Product Monograph. Peapack, NJ; 2022.
273. GlaxoSmithKline: Tranylcypromine. In: CA Product Monograph. 2019.
274. Par Pharmaceutical Inc.: Tranylcypromine. In: US Product Monograph. Chestnut Ridge, NY; 2022.
275. Pro Doc Ltée: Trazodone. In: CA Product Monograph. Laval, QC; 2021.
276. Torrent Pharmaceuticals Limited: Trazodone. In: US Product Monograph. Levittown, PA; 2022.
277. Upjohn Canada ULC: Effexor XR (Venlafaxine Hydrochloride Extended Release Capsules). In: CA Product Monograph. Kirkland, QC; 2022.
278. Acetris Health LLC: Venlafaxine hydrochloride capsule extended release. In: US Product Monograph. Saddle Brook, NJ; 2022.
279. Allergan Inc.: Viibryd (vilazodone hydrochloride). In: CA Product Monograph. Markham, ON; 2021.
280. Allergan Inc.: Viibryd (vilazodone hydrochloride). In: US Product Monograph. Irvine, CA; 2021.
281. Lundbeck Canada Inc.: Trintellix (vortioxetine). In: CA Product Monograph. St-Laurent, QC; 2021.
282. Takeda Pharmaceuticals America Inc.: Trintellix (vortioxetine). In: US Product Monograph. Lexington, MA; 2021.
283. Otsuka Canada Pharmaceutical: Abilify (aripiprazole). In: CA Product Monograph. Saint-Laurent, QC; 2021.
284. Otsuka America Pharmaceutical: Abilify (aripiprazole). In: US Product Monograph. Rockville, MD; 2021.
285. Otsuka Canada Pharmaceutical Inc: Rexulti (brexpiprazole). In: CA Product Monograph. Saint-Laurent, QC; 2020.
286. Otsuka America Pharmaceutical Inc: Rexulti (brexpiprazole). In: US Product Monograph. Rockville, MD; 2021.
287. Janssen Inc: Spravato (esketamine nasal spray). In: CA Product Monograph. Toronto, ON; 2021.
288. Janssen Pharmaceuticals Inc.: Spravato (esketamine hydrochloride solution). In: US Product Monograph. Titusville, NJ; 2020.

289. Bausch Health Canada Inc: Carbolith (lithium). In: CA Product Monograph. Laval, Quebec; 2019.
290. AA Pharma Inc: Lithmax (lithium). In: CA Product Monograph. Vaughan, Ontario; 2013.
291. ANI Pharmaceuticals Inc: Lithium Carbonate extended-release tablet. In: US Product Monograph. Baudette, MN; 2022.
292. Pharmaceuticals C: Lithium Carbonate. In: US Product Monograph. Piscataway, NJ; 2021.
293. Teva Canada Limited: Alertec (modafinil). In: CA Product Monograph. Toronto, ON; 2021.
294. Sanis Health Inc: Olanzapine. In: CA Product Monograph. Brampton, ON; 2021.
295. Eli Lilly and Company: Zyprexa (olanzapine). In: US Product Monograph. Indianapolis, IN; 2022.
296. AstraZeneca Canada Inc: Seroquel (quetiapine). In: CA Product Monograph. Mississauga, ON; 2021.
297. Accord Healthcare: ACH-Quetiapine Fumarate XR. In: CA Product Monograph. Kirkland, QC; 2022.
298. AstraZeneca Pharmaceuticals LP: Seroquel (quetiapine). In: US Product Monograph. Wilmington, DE; 2022.
299. AstraZeneca Pharmaceuticals LP: Seroquel XR - quetiapine extended release. In: US Product Monograph. Wilmington, DE; 2022.
300. Marcan Pharmaceuticals Inc: Mar-risperidone. In: CA Product Monograph. Ottawa, ON; 2022.
301. PD-Rx Pharmaceuticals Inc: Risperdal (risperidone tablet). In: US Product Monograph. Oklahoma City, OK; 2022.
302. Teva Canada Limited: Teva-liothyronine. In: CA Product Monograph. Toronto, ON; 2019.
303. Sun Pharmaceutical Industries Inc: Liothyronine. In: US Product Monograph. Cranbury, NJ; 2021.
304. Hicks JK, Sangkuhl K, Swen JJ, Ellingrod VL, Muller DJ, Shimoda K, Bishop JR, Kharasch ED, Skaar TC, Gaedigk A et al: Clinical pharmacogenetics implementation consortium guideline (CPIC) for CYP2D6 and CYP2C19 genotypes and dosing of tricyclic antidepressants: 2016 update. *Clin Pharmacol Ther* 2017, 102(1):37-44.
305. Dutch Guidelines. [<https://www.knmp.nl/downloads/pharmacogenetic-recommendations-august-2019.pdf>]
306. Swen JJ, Nijenhuis M, de Boer A, Grandia L, Maitland-van der Zee AH, Mulder H, Rongen GA, van Schaik RH, Schalekamp T, Touw DJ et al: Pharmacogenetics: from bench to byte--an update of guidelines. *Clin Pharmacol Ther* 2011, 89(5):662-673.
307. Dutch guidelines May 2020 update [[https://api.pharmgkb.org/v1/download/file/attachment/DPWG\\_May\\_2020.pdf](https://api.pharmgkb.org/v1/download/file/attachment/DPWG_May_2020.pdf)]

308. Ryu S, Park S, Lee JH, Kim YR, Na HS, Lim HS, Choi HY, Hwang IY, Lee JG, Park ZW et al: A Study on CYP2C19 and CYP2D6 Polymorphic Effects on Pharmacokinetics and Pharmacodynamics of Amitriptyline in Healthy Koreans. *Clin Transl Sci* 2017, 10(2):93-101.
309. Atasayar G, Eryilmaz IE, Karli N, Egeli U, Zarifoglu M, Cecener G, Taskapilioglu O, Tunca B, Yildirim O, Ak S et al: Association of MDR1, CYP2D6, and CYP2C19 gene polymorphisms with prophylactic migraine treatment response. *J Neurol Sci* 2016, 366:149-154.
310. de Vos A, van der Weide J, Looovers HM: Association between CYP2C19\*17 and metabolism of amitriptyline, citalopram and clomipramine in Dutch hospitalized patients. *Pharmacogenomics J* 2011, 11(5):359-367.
311. Bijl MJ, Visser LE, Hofman A, Vulto AG, van Gelder T, Stricker BH, van Schaik RH: Influence of the CYP2D6\*4 polymorphism on dose, switching and discontinuation of antidepressants. *Br J Clin Pharmacol* 2008, 65(4):558-564.
312. Steimer W, Zopf K, von Amelunxen S, Pfeiffer H, Bachofer J, Popp J, Messner B, Kissling W, Leucht S: Amitriptyline or not, that is the question: pharmacogenetic testing of CYP2D6 and CYP2C19 identifies patients with low or high risk for side effects in amitriptyline therapy. *Clin Chem* 2005, 51(2):376-385.
313. Ishigooka J, Iwashita S, Higashi K, Liew EL, Tadori Y: Pharmacokinetics and Safety of Brexpiprazole Following Multiple-Dose Administration to Japanese Patients With Schizophrenia. *J Clin Pharmacol* 2018, 58(1):74-80.
314. Kirchheiner J, Meineke I, Muller G, Roots I, Brockmoller J: Contributions of CYP2D6, CYP2C9 and CYP2C19 to the biotransformation of E- and Z-doxepin in healthy volunteers. *Pharmacogenetics* 2002, 12(7):571-580.
315. Fabbri C, Tansey KE, Perlis RH, Hauser J, Henigsberg N, Maier W, Mors O, Placentino A, Rietschel M, Souery D et al: Effect of cytochrome CYP2C19 metabolizing activity on antidepressant response and side effects: Meta-analysis of data from genome-wide association studies. *Eur Neuropsychopharmacol* 2018, 28(8):945-954.
316. Demirbugen Oz M, Uckun Z, Yuce-Artun N, Baskak B, Ozdemir H, Kizil Ozel T, Devrimci Ozguven H, Suzen HS: The relationship between the serotonin 2A receptor gene -1438A/G and 102T/C polymorphisms and citalopram/sertraline-induced nausea in major depressed patients. *Hum Psychopharmacol* 2018, 33(5):e2673.
317. Mrazek DA, Biernacka JM, O'Kane DJ, Black JL, Cunningham JM, Drews MS, Snyder KA, Stevens SR, Rush AJ, Weinshilboum RM: CYP2C19 variation and citalopram response. *Pharmacogenet Genomics* 2011, 21(1):1-9.
318. Kumar Y, Kung S, Shinozaki G: CYP2C19 variation, not citalopram dose nor serum level, is associated with QTc prolongation. *J Psychopharmacol* 2014, 28(12):1143-1148.
319. Chang M, Tybring G, Dahl ML, Lindh JD: Impact of cytochrome P450 2C19 polymorphisms on citalopram/escitalopram exposure: a systematic review and meta-analysis. *Clin Pharmacokinet* 2014, 53(9):801-811.

320. Shiroma PR, Drews MS, Geske JR, Mrazek DA: SLC6A4 polymorphisms and age of onset in late-life depression on treatment outcomes with citalopram: a Sequenced Treatment Alternatives to Relieve Depression (STAR\*D) report. *The American journal of geriatric psychiatry : official journal of the American Association for Geriatric Psychiatry* 2014, 22(11):1140-1148.
321. Rotberg B, Kronenberg S, Carmel M, Frisch A, Brent D, Zalsman G, Apter A, Weizman A: Additive effects of 5-HTTLPR (serotonin transporter) and tryptophan hydroxylase 2 G-703T gene polymorphisms on the clinical response to citalopram among children and adolescents with depression and anxiety disorders. *J Child Adolesc Psychopharmacol* 2013, 23(2):117-122.
322. Illi A, Poutanen O, Setälä-Soikkeli E, Kampman O, Viikki M, Huhtala H, Mononen N, Haraldsson S, Koivisto PA, Leinonen E et al: Is 5-HTTLPR linked to the response of selective serotonin reuptake inhibitors in MDD? *Eur Arch Psychiatry Clin Neurosci* 2011, 261(2):95-102.
323. Min W, Li T, Ma X, Li Z, Yu T, Gao D, Zhang B, Yun Y, Sun X: Monoamine transporter gene polymorphisms affect susceptibility to depression and predict antidepressant response. *Psychopharmacology (Berl)* 2009, 205(3):409-417.
324. Mrazek DA, Rush AJ, Biernacka JM, O'Kane DJ, Cunningham JM, Wieben ED, Schaid DJ, Drews MS, Courson VL, Snyder KA et al: SLC6A4 variation and citalopram response. *Am J Med Genet B Neuropsychiatr Genet* 2009, 150B(3):341-351.
325. Hu XZ, Rush AJ, Charney D, Wilson AF, Sorant AJ, Papanicolaou GJ, Fava M, Trivedi MH, Wisniewski SR, Laje G et al: Association between a functional serotonin transporter promoter polymorphism and citalopram treatment in adult outpatients with major depression. *Arch Gen Psychiatry* 2007, 64(7):783-792.
326. Arias B, Catalan R, Gasto C, Gutierrez B, Fananas L: 5-HTTLPR polymorphism of the serotonin transporter gene predicts non-remission in major depression patients treated with citalopram in a 12-weeks follow up study. *J Clin Psychopharmacol* 2003, 23(6):563-567.
327. Poland RE, Lesser IM, Wan YJ, Gertsik L, Yao J, Raffle LJ, Lin KM, Myers HF: Response to citalopram is not associated with SLC6A4 genotype in African-Americans and Caucasians with major depression. *Life Sci* 2013, 92(20-21):967-970.
328. Veldic M, Ahmed AT, Blacker CJ, Geske JR, Biernacka JM, Borreggine KL, Moore KM, Prieto ML, Vande Voort JL, Croarkin PE et al: Cytochrome P450 2C19 Poor Metabolizer Phenotype in Treatment Resistant Depression: Treatment and Diagnostic Implications. *Front Pharmacol* 2019, 10:83.
329. Fischer AG, Endrass T, Goebel I, Reuter M, Montag C, Kubisch C, Ullsperger M: Interactive effects of citalopram and serotonin transporter genotype on neural correlates of response inhibition and attentional orienting. *Neuroimage* 2015, 116:59-67.
330. Outhred T, Das P, Dobson-Stone C, Felmingham KL, Bryant RA, Nathan PJ, Malhi GS, Kemp AH: Impact of 5-HTTLPR on SSRI serotonin transporter blockade during emotion regulation: A preliminary fMRI study. *J Affect Disord* 2016, 196:11-19.
331. Tatham EL, Hall GB, Clark D, Foster J, Ramasubbu R: The 5-HTTLPR and BDNF polymorphisms moderate the association between uncinate fasciculus connectivity and antidepressants treatment response in major depression. *Eur Arch Psychiatry Clin Neurosci* 2017, 267(2):135-147.

332. Lee EJ, Oh MS, Kim JS, Chang DI, Park JH, Cha JK, Heo JH, Sohn SI, Kim DE, Kim HY et al: Serotonin transporter gene polymorphisms may be associated with poststroke neurological recovery after escitalopram use. *J Neurol Neurosurg Psychiatry* 2018, 89(3):271-276.
333. Basu A, Chadda RK, Sood M, Kaur H, Kukreti R: Association of serotonin transporter (SLC6A4) and receptor (5HT<sub>1A</sub>, 5HT<sub>2A</sub>) polymorphisms with response to treatment with escitalopram in patients with major depressive disorder: A preliminary study. *Indian J Med Res* 2015, 142(1):40-45.
334. Serretti A, Kato M, De Ronchi D, Kinoshita T: Meta-analysis of serotonin transporter gene promoter polymorphism (5-HTTLPR) association with selective serotonin reuptake inhibitor efficacy in depressed patients. *Mol Psychiatry* 2007, 12(3):247-257.
335. Kato M, Serretti A: Review and meta-analysis of antidepressant pharmacogenetic findings in major depressive disorder. *Mol Psychiatry* 2010, 15(5):473-500.
336. Porcelli S, Fabbri C, Serretti A: Meta-analysis of serotonin transporter gene promoter polymorphism (5-HTTLPR) association with antidepressant efficacy. *Eur Neuropsychopharmacol* 2012, 22(4):239-258.
337. Kawaguchi DM, Glatt SJ: GRIK4 polymorphism and its association with antidepressant response in depressed patients: a meta-analysis. *Pharmacogenomics* 2014, 15(11):1451-1459.
338. Pu M, Zhang Z, Xu Z, Shi Y, Geng L, Yuan Y, Zhang X, Reynolds GP: Influence of genetic polymorphisms in the glutamatergic and GABAergic systems and their interactions with environmental stressors on antidepressant response. *Pharmacogenomics* 2013, 14(3):277-288.
339. Horstmann S, Lucae S, Menke A, Hennings JM, Ising M, Roeske D, Muller-Myhsok B, Holsboer F, Binder EB: Polymorphisms in GRIK4, HTR2A, and FKBP5 show interactive effects in predicting remission to antidepressant treatment. *Neuropsychopharmacology* 2010, 35(3):727-740.
340. Paddock S, Laje G, Charney D, Rush AJ, Wilson AF, Sorant AJ, Lipsky R, Wisniewski SR, Manji H, McMahon FJ: Association of GRIK4 with outcome of antidepressant treatment in the STAR\*D cohort. *The American journal of psychiatry* 2007, 164(8):1181-1188.
341. Zou YF, Wang F, Feng XL, Li WF, Tao JH, Pan FM, Huang F, Su H: Meta-analysis of FKBP5 gene polymorphisms association with treatment response in patients with mood disorders. *Neurosci Lett* 2010, 484(1):56-61.
342. Lekman M, Laje G, Charney D, Rush AJ, Wilson AF, Sorant AJ, Lipsky R, Wisniewski SR, Manji H, McMahon FJ et al: The FKBP5-gene in depression and treatment response--an association study in the Sequenced Treatment Alternatives to Relieve Depression (STAR\*D) Cohort. *Biol Psychiatry* 2008, 63(12):1103-1110.
343. Binder EB, Salyakina D, Lichtner P, Wochnik GM, Ising M, Putz B, Papiol S, Seaman S, Lucae S, Kohli MA et al: Polymorphisms in FKBP5 are associated with increased recurrence of depressive episodes and rapid response to antidepressant treatment. *Nat Genet* 2004, 36(12):1319-1325.
344. Niitsu T, Fabbri C, Bentini F, Serretti A: Pharmacogenetics in major depression: a comprehensive meta-analysis. *Prog Neuropsychopharmacol Biol Psychiatry* 2013, 45:183-194.



345. Noro M, Antonijevic I, Forray C, Kasper S, Kocabas NA, Lecrubier Y, Linotte S, Mendlewicz J, Montgomery S, Snyder L et al: 5HT1A and 5HT2A receptor genes in treatment response phenotypes in major depressive disorder. *Int Clin Psychopharmacol* 2010, 25(4):228-231.
346. Viikki M, Huuhka K, Leinonen E, Illi A, Setala-Soikkeli E, Huuhka M, Mononen N, Lehtimaki T, Kampman O: Interaction between two HTR2A polymorphisms and gender is associated with treatment response in MDD. *Neurosci Lett* 2011, 501(1):20-24.
347. Kishi T, Yoshimura R, Kitajima T, Okochi T, Okumura T, Tsunoka T, Yamanouchi Y, Kinoshita Y, Kawashima K, Naitoh H et al: HTR2A is associated with SSRI response in major depressive disorder in a Japanese cohort. *Neuromolecular Med* 2010, 12(3):237-242.
348. Peters EJ, Slager SL, Jenkins GD, Reinalda MS, Garriock HA, Shyn SI, Kraft JB, McGrath PJ, Hamilton SP: Resequencing of serotonin-related genes and association of tagging SNPs to citalopram response. *Pharmacogenet Genomics* 2009, 19(1):1-10.
349. McMahon FJ, Buervenich S, Charney D, Lipsky R, Rush AJ, Wilson AF, Sorant AJ, Papanicolaou GJ, Laje G, Fava M et al: Variation in the gene encoding the serotonin 2A receptor is associated with outcome of antidepressant treatment. *Am J Hum Genet* 2006, 78(5):804-814.
350. GENDEP Investigators; MARS Investigators SDI: Common genetic variation and antidepressant efficacy in major depressive disorder: a meta-analysis of three genome-wide pharmacogenetic studies. *Am J Psychiatry* 2013, 170(2):207-217.
351. Hodgson K, Tansey K, Dernovsek MZ, Hauser J, Henigsberg N, Maier W, Mors O, Placentino A, Rietschel M, Souery D et al: Genetic differences in cytochrome P450 enzymes and antidepressant treatment response. *J Psychopharmacol* 2014, 28(2):133-141.
352. Waade RB, Hermann M, Moe HL, Molden E: Impact of age on serum concentrations of venlafaxine and escitalopram in different CYP2D6 and CYP2C19 genotype subgroups. *Eur J Clin Pharmacol* 2014, 70(8):933-940.
353. Jukic MM, Haslemo T, Molden E, Ingelman-Sundberg M: Impact of CYP2C19 Genotype on Escitalopram Exposure and Therapeutic Failure: A Retrospective Study Based on 2,087 Patients. *Am J Psychiatry* 2018, 175(5):463-470.
354. Aldrich SL, Poweleit EA, Prows CA, Martin LJ, Strawn JR, Ramsey LB: Influence of CYP2C19 Metabolizer Status on Escitalopram/Citalopram Tolerability and Response in Youth With Anxiety and Depressive Disorders. *Front Pharmacol* 2019, 10:99.
355. Ng C, Sarris J, Singh A, Bousman C, Byron K, Peh LH, Smith DJ, Tan CH, Schweitzer I: Pharmacogenetic polymorphisms and response to escitalopram and venlafaxine over 8 weeks in major depression. *Hum Psychopharmacol* 2013, 28(5):516-522.
356. Strohmaier J, Wust S, Uher R, Henigsberg N, Mors O, Hauser J, Souery D, Zobel A, Dernovsek MZ, Streit F et al: Sexual dysfunction during treatment with serotonergic and noradrenergic antidepressants: clinical description and the role of the 5-HTTLPR. *World J Biol Psychiatry* 2011, 12(7):528-538.

357. Keers R, Uher R, Huezo-Diaz P, Smith R, Jaffee S, Rietschel M, Henigsberg N, Kozel D, Mors O, Maier W et al: Interaction between serotonin transporter gene variants and life events predicts response to antidepressants in the GENDEP project. *Pharmacogenomics J* 2011, 11(2):138-145.
358. Maron E, Tammiste A, Kallassalu K, Eller T, Vasar V, Nutt DJ, Metspalu A: Serotonin transporter promoter region polymorphisms do not influence treatment response to escitalopram in patients with major depression. *Eur Neuropsychopharmacol* 2009, 19(6):451-456.
359. Alexopoulos GS, Murphy CF, Gunning-Dixon FM, Glatt CE, Latoussakis V, Kelly RE, Jr., Kanellopoulos D, Klimstra S, Lim KO, Young RC et al: Serotonin transporter polymorphisms, microstructural white matter abnormalities and remission of geriatric depression. *J Affect Disord* 2009, 119(1-3):132-141.
360. Huezo-Diaz P, Uher R, Smith R, Rietschel M, Henigsberg N, Marusic A, Mors O, Maier W, Hauser J, Souery D et al: Moderation of antidepressant response by the serotonin transporter gene. *Br J Psychiatry* 2009, 195(1):30-38.
361. Poweleit EA, Aldrich SL, Martin LJ, Hahn D, Strawn JR, Ramsey LB: Pharmacogenetics of Sertraline Tolerability and Response in Pediatric Anxiety and Depressive Disorders. *J Child Adolesc Psychopharmacol* 2019, 29(5):348-361.
362. Brandl EJ, Tiwari AK, Zhou X, Deluce J, Kennedy JL, Muller DJ, Richter MA: Influence of CYP2D6 and CYP2C19 gene variants on antidepressant response in obsessive-compulsive disorder. *Pharmacogenomics J* 2014, 14(2):176-181.
363. Rudberg I, Hermann M, Refsum H, Molden E: Serum concentrations of sertraline and N-desmethyl sertraline in relation to CYP2C19 genotype in psychiatric patients. *Eur J Clin Pharmacol* 2008, 64(12):1181-1188.
364. Grasmader K, Verwohlt PL, Rietschel M, Dragicevic A, Muller M, Hiemke C, Freymann N, Zobel A, Maier W, Rao ML: Impact of polymorphisms of cytochrome-P450 isoenzymes 2C9, 2C19 and 2D6 on plasma concentrations and clinical effects of antidepressants in a naturalistic clinical setting. *Eur J Clin Pharmacol* 2004, 60(5):329-336.
365. Wang JH, Liu ZQ, Wang W, Chen XP, Shu Y, He N, Zhou HH: Pharmacokinetics of sertraline in relation to genetic polymorphism of CYP2C19. *Clin Pharmacol Ther* 2001, 70(1):42-47.
366. Zastrozhin MS, Grishina EA, Denisenko NP, Skryabin VY, Markov DD, Savchenko LM, Bryun EA, Sychev DA: Effects of CYP2D6 genetic polymorphisms on the efficacy and safety of fluvoxamine in patients with depressive disorder and comorbid alcohol use disorder. *Pharmacogenomics Pers Med* 2018, 11:113-119.
367. Muller DJ, Brandl EJ, Hwang R, Tiwari AK, Sturgess JE, Zai CC, Lieberman JA, Kennedy JL, Richter MA: The AmpliChip(R) CYP450 test and response to treatment in schizophrenia and obsessive compulsive disorder: a pilot study and focus on cases with abnormal CYP2D6 drug metabolism. *Genet Test Mol Biomarkers* 2012, 16(8):897-903.
368. Suzuki Y, Sugai T, Fukui N, Watanabe J, Ono S, Inoue Y, Ozdemir V, Someya T: CYP2D6 genotype and smoking influence fluvoxamine steady-state concentration in Japanese psychiatric patients: lessons for genotype-phenotype association study design in translational pharmacogenetics. *J Psychopharmacol* 2011, 25(7):908-914.

369. Katoh Y, Uchida S, Kawai M, Takei N, Mori N, Kawakami J, Kagawa Y, Yamada S, Namiki N, Hashimoto H: Effects of cigarette smoking and cytochrome P450 2D6 genotype on fluvoxamine concentration in plasma of Japanese patients. *Biol Pharm Bull* 2010, 33(2):285-288.
370. Watanabe J, Suzuki Y, Fukui N, Sugai T, Ono S, Inoue Y, Someya T: Dose-dependent effect of the CYP2D6 genotype on the steady-state fluvoxamine concentration. *Ther Drug Monit* 2008, 30(6):705-708.
371. Suzuki Y, Sawamura K, Someya T: Polymorphisms in the 5-hydroxytryptamine 2A receptor and CytochromeP4502D6 genes synergistically predict fluvoxamine-induced side effects in Japanese depressed patients. *Neuropsychopharmacology* 2006, 31(4):825-831.
372. Kawanishi C, Lundgren S, Agren H, Bertilsson L: Increased incidence of CYP2D6 gene duplication in patients with persistent mood disorders: ultrarapid metabolism of antidepressants as a cause of nonresponse. A pilot study. *Eur J Clin Pharmacol* 2004, 59(11):803-807.
373. Gerstenberg G, Aoshima T, Fukasawa T, Yoshida K, Takahashi H, Higuchi H, Murata Y, Shimoyama R, Ohkubo T, Shimizu T et al: Effects of the CYP 2D6 genotype and cigarette smoking on the steady-state plasma concentrations of fluvoxamine and its major metabolite fluvoxamino acid in Japanese depressed patients. *Ther Drug Monit* 2003, 25(4):463-468.
374. Gerstenberg G, Aoshima T, Fukasawa T, Yoshida K, Takahashi H, Higuchi H, Murata Y, Shimoyama R, Ohkubo T, Shimizu T et al: Relationship between clinical effects of fluvoxamine and the steady-state plasma concentrations of fluvoxamine and its major metabolite fluvoxamino acid in Japanese depressed patients. *Psychopharmacology (Berl)* 2003, 167(4):443-448.
375. Ohara K, Tanabu S, Ishibashi K, Ikemoto K, Yoshida K, Shibuya H: CYP2D6\*10 alleles do not determine plasma fluvoxamine concentration/dose ratio in Japanese subjects. *European journal of clinical pharmacology* 2003, 58(10):659-661.
376. Spigset O, Axelsson S, Norstrom A, Hagg S, Dahlqvist R: The major fluvoxamine metabolite in urine is formed by CYP2D6. *Eur J Clin Pharmacol* 2001, 57(9):653-658.
377. Hartter S, Grozinger M, Weigmann H, Roschke J, Hiemke C: Increased bioavailability of oral melatonin after fluvoxamine coadministration. *Clin Pharmacol Ther* 2000, 67(1):1-6.
378. Nishimura M, Ueda M, Saruwatari J, Nakashima H, Ogusu N, Aoki A, Tsuchimine S, Matsuda K, Iwashita K, Ono T et al: Influence of the cytochrome P450 2D6 \*10/\*10 genotype on the pharmacokinetics of paroxetine in Japanese patients with major depressive disorder: a population pharmacokinetic analysis. *Pharmacogenet Genomics* 2016, 26(9):403-413.
379. Ishiguro S, Watanabe T, Ueda M, Saeki Y, Hayashi Y, Akiyama K, Saito A, Kato K, Inoue Y, Shimoda K: Determinants of pharmacodynamic trajectory of the therapeutic response to paroxetine in Japanese patients with panic disorder. *Eur J Clin Pharmacol* 2011, 67(12):1213-1221.
380. Yevtushenko OO, Oros MM, Reynolds GP: Early response to selective serotonin reuptake inhibitors in panic disorder is associated with a functional 5-HT1A receptor gene polymorphism. *J Affect Disord* 2010, 123(1-3):308-311.

381. Illi A, Setala-Soikkeli E, Viikki M, Poutanen O, Huhtala H, Mononen N, Lehtimäki T, Leinonen E, Kampman O: 5-HTR1A, 5-HTR2A, 5-HTR6, TPH1 and TPH2 polymorphisms and major depression. *Neuroreport* 2009, 20(12):1125-1128.
382. Haslemo T, Eliasson E, Jukic MM, Ingelman-Sundberg M, Molden E: Significantly lower CYP2D6 metabolism measured as the O/N-desmethylvenlafaxine metabolic ratio in carriers of CYP2D6\*41 versus CYP2D6\*9 or CYP2D6\*10: a study on therapeutic drug monitoring data from 1003 genotyped Scandinavian patients. *Br J Clin Pharmacol* 2019, 85(1):194-201.
383. Lin XQ, Wang P, Cai WK, Xu GL, Yang M, Zhou MD, Sun M, He F, He GH: The Associations Between CYP2D6 Metabolizer Status and Pharmacokinetics and Clinical Outcomes of Venlafaxine: A Systematic Review and Meta-Analysis. *Pharmacopsychiatry* 2018.
384. Lobello KW, Preskorn SH, Guico-Pabia CJ, Jiang Q, Paul J, Nichols AI, Patroneva A, Ninan PT: Cytochrome P450 2D6 phenotype predicts antidepressant efficacy of venlafaxine: a secondary analysis of 4 studies in major depressive disorder. *J Clin Psychiatry* 2010, 71(11):1482-1487.
385. McAlpine DE, O'Kane DJ, Black JL, Mrazek DA: Cytochrome P450 2D6 genotype variation and venlafaxine dosage. *Mayo Clin Proc* 2007, 82(9):1065-1068.
386. Lessard E, Yessine MA, Hamelin BA, O'Hara G, LeBlanc J, Turgeon J: Influence of CYP2D6 activity on the disposition and cardiovascular toxicity of the antidepressant agent venlafaxine in humans. *Pharmacogenetics* 1999, 9(4):435-443.
387. Van Nieuwerburgh FC, Denys DA, Westenberg HG, Deforce DL: Response to serotonin reuptake inhibitors in OCD is not influenced by common CYP2D6 polymorphisms. *Int J Psychiatry Clin Pract* 2009, 13(1):345-348.
388. Hermann M, Hendset M, Fosaas K, Hjerpset M, Refsum H: Serum concentrations of venlafaxine and its metabolites O-desmethylvenlafaxine and N-desmethylvenlafaxine in heterozygous carriers of the CYP2D6\*3, \*4 or \*5 allele. *European journal of clinical pharmacology* 2008, 64(5):483-487.
389. Shams ME, Arneith B, Hiemke C, Dragicevic A, Muller MJ, Kaiser R, Lackner K, Hartter S: CYP2D6 polymorphism and clinical effect of the antidepressant venlafaxine. *J Clin Pharm Ther* 2006, 31(5):493-502.
390. van der Weide J, van Baalen-Benedek EH, Kootstra-Ros JE: Metabolic ratios of psychotropics as indication of cytochrome P450 2D6/2C19 genotype. *Ther Drug Monit* 2005, 27(4):478-483.
391. Eap CB, Lessard E, Baumann P, Brawand-Amey M, Yessine MA, O'Hara G, Turgeon J: Role of CYP2D6 in the stereoselective disposition of venlafaxine in humans. *Pharmacogenetics* 2003, 13(1):39-47.
392. Fukuda T, Nishida Y, Zhou Q, Yamamoto I, Kondo S, Azuma J: The impact of the CYP2D6 and CYP2C19 genotypes on venlafaxine pharmacokinetics in a Japanese population. *Eur J Clin Pharmacol* 2000, 56(2):175-180.
393. Veefkind AH, Haffmans PM, Hoencamp E: Venlafaxine serum levels and CYP2D6 genotype. *Ther Drug Monit* 2000, 22(2):202-208.

394. Fukuda T, Yamamoto I, Nishida Y, Zhou Q, Ohno M, Takada K, Azuma J: Effect of the CYP2D6\*10 genotype on venlafaxine pharmacokinetics in healthy adult volunteers. *British journal of clinical pharmacology* 1999, 47(4):450-453.
395. Preskorn S, Patroneva A, Silman H, Jiang Q, Isler JA, Burczynski ME, Ahmed S, Paul J, Nichols AI: Comparison of the pharmacokinetics of venlafaxine extended release and desvenlafaxine in extensive and poor cytochrome P450 2D6 metabolizers. *J Clin Psychopharmacol* 2009, 29(1):39-43.
396. Jaquenoud Sirot E, Harenberg S, Vandel P, Lima CA, Perrenoud P, Kemmerling K, Zullino DF, Hilleret H, Crettol S, Jonzier-Perey M et al: Multicenter study on the clinical effectiveness, pharmacokinetics, and pharmacogenetics of mirtazapine in depression. *J Clin Psychopharmacol* 2012, 32(5):622-629.
397. Borobia AM, Novalbos J, Guerra-Lopez P, Lopez-Rodriguez R, Tabares B, Rodriguez V, Abad-Santos F, Carcas AJ: Influence of sex and CYP2D6 genotype on mirtazapine disposition, evaluated in Spanish healthy volunteers. *Pharmacol Res* 2009, 59(6):393-398.
398. Lind AB, Reis M, Bengtsson F, Jonzier-Perey M, Powell Golay K, Ahlner J, Baumann P, Dahl ML: Steady-state concentrations of mirtazapine, N-desmethyilmirtazapine, 8-hydroxymirtazapine and their enantiomers in relation to cytochrome P450 2D6 genotype, age and smoking behaviour. *Clin Pharmacokinet* 2009, 48(1):63-70.
399. Brockmoller J, Meineke I, Kirchheiner J: Pharmacokinetics of mirtazapine: enantioselective effects of the CYP2D6 ultra rapid metabolizer genotype and correlation with adverse effects. *Clin Pharmacol Ther* 2007, 81(5):699-707.
400. Kirchheiner J, Henckel HB, Meineke I, Roots I, Brockmoller J: Impact of the CYP2D6 ultrarapid metabolizer genotype on mirtazapine pharmacokinetics and adverse events in healthy volunteers. *J Clin Psychopharmacol* 2004, 24(6):647-652.
401. Murphy GM, Jr., Kremer C, Rodrigues HE, Schatzberg AF: Pharmacogenetics of antidepressant medication intolerance. *Am J Psychiatry* 2003, 160(10):1830-1835.
402. Saiz-Rodriguez M, Belmonte C, Derqui-Fernandez N, Cabaleiro T, Roman M, Ochoa D, Talegon M, Ovejero-Benito MC, Abad-Santos F: Pharmacogenetics of trazodone in healthy volunteers: association with pharmacokinetics, pharmacodynamics and safety. *Pharmacogenomics* 2017, 18(16):1491-1502.
403. Saiz-Rodriguez M, Belmonte C, Roman M, Ochoa D, Jiang-Zheng C, Koller D, Mejia G, Zubiaur P, Wojnicz A, Abad-Santos F: Effect of ABCB1 C3435T Polymorphism on Pharmacokinetics of Antipsychotics and Antidepressants. *Basic Clin Pharmacol Toxicol* 2018, 123(4):474-485.
404. Pharmacogenomic Testing for Psychotropic Medication Selection: A Systematic Review of the Assurex GeneSight Psychotropic Test. *Ont Health Technol Assess Ser* 2017, 17(4):1-39.
405. Areberg J, Petersen KB, Chen G, Naik H: Population pharmacokinetic meta-analysis of vortioxetine in healthy individuals. *Basic Clin Pharmacol Toxicol* 2014, 115(6):552-559.
406. David SP, Strong DR, Leventhal AM, Lancaster MA, McGeary JE, Munafo MR, Bergen AW, Swan GE, Benowitz NL, Tyndale RF et al: Influence of a dopamine pathway additive genetic efficacy score on smoking cessation: results from two randomized clinical trials of bupropion. *Addiction* 2013, 108(12):2202-2211.

407. Breitling LP, Twardella D, Hoffmann MM, Witt SH, Treutlein J, Brenner H: Prospective association of dopamine-related polymorphisms with smoking cessation in general care. *Pharmacogenomics* 2010, 11(4):527-536.
408. Lerman C, Shields PG, Wileyto EP, Audrain J, Hawk LH, Jr., Pinto A, Kucharski S, Krishnan S, Niaura R, Epstein LH: Effects of dopamine transporter and receptor polymorphisms on smoking cessation in a bupropion clinical trial. *Health Psychol* 2003, 22(5):541-548.
409. Tomaz PR, Santos JR, Issa JS, Abe TO, Gaya PV, Krieger JE, Pereira AC, Santos PC: CYP2B6 rs2279343 polymorphism is associated with smoking cessation success in bupropion therapy. *Eur J Clin Pharmacol* 2015, 71(9):1067-1073.
410. Hubacek JA, Pankova A, Stepankova L, Zvolaska K, Adamkova V, Lanska V, Kralikova E: SNPs within CHRNA5-A3-B4 and CYP2A6/B6 are associated with smoking dependence but not with tobacco dependence treatment outcomes in the Czech population. *Gene* 2017, 606:35-38.
411. Lee AM, Jepson C, Hoffmann E, Epstein L, Hawk LW, Lerman C, Tyndale RF: CYP2B6 genotype alters abstinence rates in a bupropion smoking cessation trial. *Biol Psychiatry* 2007, 62(6):635-641.
412. Lerman C, Shields PG, Wileyto EP, Audrain J, Pinto A, Hawk L, Krishnan S, Niaura R, Epstein L: Pharmacogenetic investigation of smoking cessation treatment. *Pharmacogenetics* 2002, 12(8):627-634.
413. David SP, Strong DR, Munafo MR, Brown RA, Lloyd-Richardson EE, Wileyto PE, Evins EA, Shields PG, Lerman C, Niaura R: Bupropion efficacy for smoking cessation is influenced by the DRD2 Taq1A polymorphism: analysis of pooled data from two clinical trials. *Nicotine Tob Res* 2007, 9(12):1251-1257.
414. David SP, Brown RA, Papandonatos GD, Kahler CW, Lloyd-Richardson EE, Munafo MR, Shields PG, Lerman C, Strong D, McCaffery J et al: Pharmacogenetic clinical trial of sustained-release bupropion for smoking cessation. *Nicotine Tob Res* 2007, 9(8):821-833.
415. Swan GE, Valdes AM, Ring HZ, Khroyan TV, Jack LM, Ton CC, Curry SJ, McAfee T: Dopamine receptor DRD2 genotype and smoking cessation outcome following treatment with bupropion SR. *Pharmacogenomics J* 2005, 5(1):21-29.
416. Zhang JP, Lencz T, Zhang RX, Nitta M, Maayan L, John M, Robinson DG, Fleischhacker WW, Kahn RS, Ophoff RA et al: Pharmacogenetic Associations of Antipsychotic Drug-Related Weight Gain: A Systematic Review and Meta-analysis. *Schizophr Bull* 2016, 42(6):1418-1437.
417. Zhang JP, Lencz T, Malhotra AK: D2 receptor genetic variation and clinical response to antipsychotic drug treatment: a meta-analysis. *Am J Psychiatry* 2010, 167(7):763-772.
418. Alenius M, Wadelius M, Dahl ML, Hartvig P, Lindstrom L, Hammarlund-Udenaes M: Gene polymorphism influencing treatment response in psychotic patients in a naturalistic setting. *J Psychiatr Res* 2008, 42(11):884-893.
419. Bakker PR, van Harten PN, van Os J: Antipsychotic-induced tardive dyskinesia and polymorphic variations in COMT, DRD2, CYP1A2 and MnsOD genes: a meta-analysis of pharmacogenetic interactions. *Mol Psychiatry* 2008, 13(5):544-556.

420. Zai CC, De Luca V, Hwang RW, Voineskos A, Muller DJ, Remington G, Kennedy JL: Meta-analysis of two dopamine D2 receptor gene polymorphisms with tardive dyskinesia in schizophrenia patients. *Mol Psychiatry* 2007, 12(9):794-795.
421. Muller DJ, Zai CC, Sicard M, Remington E, Souza RP, Tiwari AK, Hwang R, Likhodi O, Shaikh S, Freeman N et al: Systematic analysis of dopamine receptor genes (DRD1-DRD5) in antipsychotic-induced weight gain. *Pharmacogenomics J* 2012, 12(2):156-164.
422. Ohara K, Nagai M, Tani K, Nakamura Y, Ino A, Ohara K: Functional polymorphism of -141C Ins/Del in the dopamine D2 receptor gene promoter and schizophrenia. *Psychiatry Research* 1998, 81(2):117-123.
423. Zhang J-P, Lencz T, Malhotra AK: D2 Receptor Genetic Variation and Clinical Response to Antipsychotic Drug Treatment: A Meta-Analysis. *American Journal of Psychiatry* 2010, 167(7):763-772.
424. Mulder H, Cohen D, Scheffer H, Gispens-de Wied C, Arends J, Wilmink FW, Franke B, Egberts AC: HTR2C gene polymorphisms and the metabolic syndrome in patients with schizophrenia: a replication study. *J Clin Psychopharmacol* 2009, 29(1):16-20.
425. Mulder H, Franke B, van der-Beek van der AA, Arends J, Wilmink FW, Scheffer H, Egberts AC: The association between HTR2C gene polymorphisms and the metabolic syndrome in patients with schizophrenia. *J Clin Psychopharmacol* 2007, 27(4):338-343.
426. Risselada AJ, Mulder H, Heerdink ER, Gonera RK, Egberts TC: Association between the HTR2C rs1414334 and ADRA2A 1291 C/G polymorphisms and lipid levels in obese patients. In: *Genetic determinants for metabolic abnormalities*. edn.; 2012: 107.
427. Czerwensky F, Leucht S, Steimer W: MC4R rs489693: a clinical risk factor for second generation antipsychotic-related weight gain? *Int J Neuropsychopharmacol* 2013, 16(9):2103-2109.
428. Malhotra AK, Correll CU, Chowdhury NI, Muller DJ, Gregersen PK, Lee AT, Tiwari AK, Kane JM, Fleischhacker WW, Kahn RS et al: Association between common variants near the melanocortin 4 receptor gene and severe antipsychotic drug-induced weight gain. *Arch Gen Psychiatry* 2012, 69(9):904-912.
429. Ma X, Maimaitirexiati T, Zhang R, Gui X, Zhang W, Xu G, Hu G: HTR2C polymorphisms, olanzapine-induced weight gain and antipsychotic-induced metabolic syndrome in schizophrenia patients: a meta-analysis. *Int J Psychiatry Clin Pract* 2014, 18(4):229-242.
430. Hiemke C, Baumann P, Bergemann N, Conca A, Dietmaier O, Egberts K, Fric M, Gerlach M, Greiner C, Grunder G et al: AGNP Consensus Guidelines for Therapeutic Drug Monitoring in Psychiatry: Update 2011. *Pharmacopsychiatry* 2011, 44(6):195-235.
431. Xing Q, Qian X, Li H, Wong S, Wu S, Feng G, Duan S, Xu M, Gao R, Qin W: The relationship between the therapeutic response to risperidone and the dopamine D2 receptor polymorphism in Chinese schizophrenia patients. *International Journal of Neuropsychopharmacology* 2007, 10(5):631-637.
432. Ikeda M, Yamanouchi Y, Kinoshita Y, Kitajima T, Yoshimura R, Hashimoto S, O'Donovan MC, Nakamura J, Ozaki N, Iwata N: Variants of dopamine and serotonin candidate genes as predictors of response to risperidone treatment in first-episode schizophrenia. 2008.

433. Lencz T, Robinson DG, Xu K, Ekholm J, Sevy S, Gunduz-Bruce H, Woerner MG, Kane JM, Goldman D, Malhotra AK: DRD2 promoter region variation as a predictor of sustained response to antipsychotic medication in first-episode schizophrenia patients. *American Journal of Psychiatry* 2006, 163(3):529-531. DPWG. Dutch Guidelines. <https://www.knmp.nl/downloads/pharmacogenetic-recommendations-august-2019.pdf>. Published 2019. Accessed October 22, 2019.