**Appendix Table D52. Adverse effects with topiramate vs. placebo in children (pooled with random effects results from randomized controlled clinical trials)**

| **Outcome** | **Dose** | **Reference,**  **Risk of bias** | **Events/**  **Randomized**  **[rate, %] with drug** | **Events/**  **Randomized**  **[rate, %] with placebo** | **Relative risk**  **(95% CI)** | **Weight** | **Absolute risk difference**  **(95% CI)** | **Weight** |
| --- | --- | --- | --- | --- | --- | --- | --- | --- |
| Abdominal pain | 50mg | Lewis, 20097 Low | 3/35 [8.6] | 3/33 [9.1] | 0.9(0.2 to 4.3) | 16.43 | -0.01(-0.14 to 0.13) | 22.37 |
| Abdominal pain | 50mg | Winner, 200615 Low | 0/12 [0.0] | 1/12 [8.3] | 0.3(0.0 to 7.5) | 3.97 | -0.08(-0.29 to 0.12) | 9.92 |
| Abdominal pain |  | Pooled | 3/47 [6.4] | 4/45 [8.9] | 0.8(0.2 to 3.0) | 20.4 | -0.03(-0.14 to 0.08) | 32.28 |
| Abdominal pain | 100mg | Lewis, 20097 Low | 5/35 [14.3] | 3/33 [9.1] | 1.6(0.4 to 6.1) | 21.05 | 0.05(-0.10 to 0.20) | 17.67 |
| Abdominal pain | 100mg | Winner, 200615 Low | 2/13 [15.4] | 1/12 [8.3] | 1.8(0.2 to 17.8) | 7.45 | 0.07(-0.18 to 0.32) | 6.48 |
| Abdominal pain |  | Pooled | 7/48 [14.6] | 4/45 [8.9] | 1.6(0.5 to 5.2) | 28.5 | 0.06(-0.07 to 0.19) | 24.15 |
| Abdominal pain | 200mg | Winner, 20058 Medium | 11/112 [9.8] | 6/50 [12.0] | 0.8(0.3 to 2.1) | 43.67 | -0.02(-0.13 to 0.08) | 36.55 |
| Abdominal pain | 200mg | Winner, 200615 Low | 2/14 [14.3] | 1/12 [8.3] | 1.7(0.2 to 16.6) | 7.42 | 0.06(-0.18 to 0.30) | 7.02 |
| Abdominal pain |  | Pooled | 13/126 [10.3] | 7/62 [11.3] | 0.9(0.4 to 2.2) | 51.1 | -0.01(-0.11 to 0.09) | 43.57 |
| Abdominal pain |  | Pooled | 23/221 [10.4] | 15/152 [9.9] | 1.0(0.6 to 1.9) | 100 | 0.00(-0.06 to 0.06) | 100 |
| Anorexia | 50mg | Lewis, 20097 Low | 3/35 [8.6] | 1/33 [3.0] | 2.8(0.3 to 25.9) | 11.32 | 0.06(-0.05 to 0.17) | 26.37 |
| Anorexia | 50mg | Winner, 200615 Low | 1/12 [8.3] | 1/12 [8.3] | 1.0(0.1 to 14.2) | 7.87 | 0.00(-0.22 to 0.22) | 6.48 |
| Anorexia |  | Pooled | 4/47 [8.5] | 2/45 [4.4] | 1.8(0.3 to 10.1) | 19.19 | 0.04(-0.05 to 0.14) | 32.85 |
| Anorexia | 100mg | Lewis, 20097 Low | 4/35 [11.4] | 1/33 [3.0] | 3.8(0.4 to 32.0) | 12.11 | 0.08(-0.04 to 0.21) | 21.81 |
| Anorexia | 100mg | Winner, 200615 Low | 1/13 [7.7] | 1/12 [8.3] | 0.9(0.1 to 13.2) | 7.84 | -0.01(-0.22 to 0.21) | 6.98 |
| Anorexia |  | Pooled | 5/48 [10.4] | 2/45 [4.4] | 2.2(0.4 to 11.5) | 19.95 | 0.06(-0.04 to 0.17) | 28.79 |
| Anorexia | 200mg | Winner, 20058 Medium | 15/112 [13.4] | 4/50 [8.0] | 1.7(0.6 to 4.8) | 50.14 | 0.05(-0.04 to 0.15) | 32.9 |
| Anorexia | 200mg | Winner, 200615 Low | 2/14 [14.3] | 1/12 [8.3] | 1.7(0.2 to 16.6) | 10.72 | 0.06(-0.18 to 0.30) | 5.46 |
| Anorexia |  | Pooled | 17/126 [13.5] | 5/62 [8.1] | 1.7(0.6 to 4.4) | 60.86 | 0.06(-0.04 to 0.15) | 38.36 |
| Anorexia |  | Pooled | 26/221 [11.8] | 9/152 [5.9] | 1.8(0.9 to 3.8) | 100 | 0.05(0.00 to 0.11) | 100 |
| Dizziness | 50mg | Lewis, 20097 Low | 2/35 [5.7] | 0/33 [0.0] | 4.7(0.2 to 94.8) | 19.56 | 0.06(-0.04 to 0.15) | 30.22 |
| Dizziness | 50mg | Winner, 200615 Low | 0/12 [0.0] | 2/12 [16.7] | 0.2(0.0 to 3.8) | 20.11 | -0.17(-0.41 to 0.07) | 13.56 |
| Dizziness |  | Pooled | 2/47 [4.3] | 2/45 [4.4] | 1.0(0.0 to 21.2) | 39.67 | -0.03(-0.24 to 0.19) | 43.78 |
| Dizziness | 100mg | Lewis, 20097 Low | 3/35 [8.6] | 0/353 [0.0] | 6.6(0.4 to 123.3) | 20.21 | 0.09(-0.02 to 0.19) | 28.43 |
| Dizziness | 100mg | Winner, 200615 Low | 0/13 [0.0] | 2/12 [16.7] | 0.2(0.0 to 3.5) | 20.07 | -0.17(-0.40 to 0.07) | 13.79 |
| Dizziness |  | Pooled | 3/48 [6.3] | 2/365 [4.4] | 1.1(0.0 to 36.8) | 40.28 | -0.02(-0.26 to 0.23) | 42.23 |
| Dizziness | 200mg | Winner, 200615 Low | 0/14 [0.0] | 2/12 [16.7] | 0.2(0.0 to 3.3) | 20.05 | -0.17(-0.40 to 0.07) | 13.99 |
| Dizziness |  | Pooled | 5/109 [4.6] | 6/422 [5.9] | 0.7(0.1 to 3.7) | 100 | -0.03(-0.14 to 0.08) | 100 |
| Fatigue | 50mg | Lewis, 20097 Low | 2/35 [5.7] | 2/33 [6.1] | 0.9(0.1 to 6.3) | 13.98 | 0.00(-0.12 to 0.11) | 25.91 |
| Fatigue | 50mg | Winner, 200615 Low | 0/12 [0.0] | 1/12 [8.3] | 0.3(0.0 to 7.5) | 5.23 | -0.08(-0.29 to 0.12) | 7.91 |
| Fatigue |  | Pooled | 2/47 [4.3] | 3/45 [6.7] | 0.7(0.1 to 3.6) | 19.21 | -0.02(-0.12 to 0.08) | 33.82 |
| Fatigue | 100mg | Lewis, 20097 Low | 3/35 [8.6] | 2/33 [6.1] | 1.4(0.3 to 7.9) | 16.98 | 0.03(-0.10 to 0.15) | 21.34 |
| Fatigue | 100mg | Winner, 200615 Low | 1/13 [7.7] | 1/12 [8.3] | 0.9(0.1 to 13.2) | 7.15 | -0.01(-0.22 to 0.21) | 7.15 |
| Fatigue |  | Pooled | 4/48 [8.3] | 3/45 [6.7] | 1.2(0.3 to 5.3) | 24.13 | 0.02(-0.09 to 0.12) | 28.49 |
| Fatigue | 200mg | Winner, 20058 Medium | 7/112 [6.3] | 6/50 [12.0] | 0.5(0.2 to 1.5) | 46.88 | -0.06(-0.16 to 0.04) | 32.1 |
| Fatigue | 200mg | Winner, 200615 Low | 2/14 [14.3] | 1/12 [8.3] | 1.7(0.2 to 16.6) | 9.78 | 0.06(-0.18 to 0.30) | 5.6 |
| Fatigue |  | Pooled | 9/126 [7.1] | 7/62 [11.3] | 0.6(0.2 to 1.6) | 56.66 | -0.04(-0.13 to 0.05) | 37.7 |
| Fatigue |  | Pooled | 15/221 [6.8] | 13/152 [8.6] | 0.8(0.4 to 1.6) | 100 | -0.02(-0.08 to 0.04) | 100 |
| Injury | 50mg | Lewis, 20097 Low | 3/35 [8.6] | 2/33 [6.1] | 1.4(0.3 to 7.9) | 15.79 | 0.03(-0.10 to 0.15) | 23.1 |
| Injury | 50mg | Winner, 200615 Low | 1/12 [8.3] | 1/12 [8.3] | 1.0(0.1 to 14.2) | 6.67 | 0.00(-0.22 to 0.22) | 7.19 |
| Injury |  | Pooled | 4/47 [8.5] | 3/45 [6.7] | 1.3(0.3 to 5.4) | 22.47 | 0.02(-0.09 to 0.13) | 30.3 |
| Injury | 100mg | Lewis, 20097 Low | 4/35 [11.4] | 2/33 [6.1] | 1.9(0.4 to 9.6) | 17.7 | 0.05(-0.08 to 0.19) | 19.84 |
| Injury | 100mg | Winner, 200615 Low | 1/13 [7.7] | 1/12 [8.3] | 0.9(0.1 to 13.2) | 6.65 | -0.01(-0.22 to 0.21) | 7.74 |
| Injury |  | Pooled | 5/48 [10.4] | 3/45 [6.7] | 1.6(0.4 to 6.2) | 24.35 | 0.04(-0.08 to 0.15) | 27.58 |
| Injury | 200mg | Winner, 20058 Medium | 8/112 [7.1] | 6/50 [12.0] | 0.6(0.2 to 1.6) | 46.56 | -0.05(-0.15 to 0.05) | 33.87 |
| Injury | 200mg | Winner, 200615 Low | 1/14 [7.1] | 1/12 [8.3] | 0.9(0.1 to 12.3) | 6.63 | -0.01(-0.22 to 0.20) | 8.25 |
| Injury |  | Pooled | 9/126 [7.1] | 7/62 [11.3] | 0.6(0.2 to 1.6) | 53.19 | -0.04(-0.13 to 0.05) | 42.12 |
| Injury |  | Pooled | 18/221 [8.1] | 13/152 [8.6] | 0.9(0.5 to 1.8) | 100 | 0.00(-0.06 to 0.06) | 100 |
| Mood problems | 50mg | Winner, 200615 Low | 0/12 [0.0] | 2/12 [16.7] | 0.2(0.0 to 3.8) | 18.56 | -0.17(-0.41 to 0.07) | 7.29 |
| Mood problems | 50mg | 20159428 Low | 1/35 [2.9] | 0/33 [0.0] | 2.8(0.1 to 67.2) | 15.97 | 0.03(-0.05 to 0.11) | 39.29 |
| Mood problems |  | Pooled | 1/47 [2.1] | 2/45 [4.4] | 0.7(0.1 to 9.4) | 34.53 | -0.04(-0.21 to 0.14) | 46.58 |
| Mood problems | 100mg | Winner, 200615 Low | 0/13 [0.0] | 2/12 [16.7] | 0.2(0.0 to 3.5) | 18.51 | -0.17(-0.40 to 0.07) | 7.47 |
| Mood problems | 100mg | 20159428 Low | 1/35 [2.9] | 0/33 [0.0] | 2.8(0.1 to 67.2) | 15.97 | 0.03(-0.05 to 0.11) | 39.29 |
| Mood problems |  | Pooled | 1/48 [2.1] | 2/45 [4.4] | 0.7(0.0 to 9.8) | 34.48 | -0.04(-0.22 to 0.14) | 46.75 |
| Mood problems | 200mg | Winner, 200615 Low | 1/14 [7.1] | 2/12 [16.7] | 0.4(0.0 to 4.2) | 30.98 | -0.10(-0.35 to 0.16) | 6.67 |
| Mood problems |  | Pooled | 3/109 [2.8] | 6/102 [5.9] | 0.6(0.2 to 2.1) | 100 | -0.01(-0.08 to 0.06) | 100 |
| Nausea | 50mg | Lewis, 20097 Low | 2/35 [5.7] | 2/33 [6.1] | 0.9(0.1 to 6.3) | 23.74 | 0.00(-0.12 to 0.11) | 25.71 |
| Nausea | 100mg | Lewis, 20097 Low | 3/35 [8.6] | 2/33 [6.1] | 1.4(0.3 to 7.9) | 28.84 | 0.03(-0.10 to 0.15) | 21.18 |
| Nausea | 200mg | Winner, 20058 Medium | 6/112 [5.4] | 3/50 [6.0] | 0.9(0.2 to 3.4) | 47.42 | -0.01(-0.08 to 0.07) | 53.11 |
| Nausea |  | Pooled | 11/182 [6.0] | 7/116 [6.0] | 1.0(0.4 to 2.6) | 100 | 0.00(-0.06 to 0.06) | 100 |
| Paresthesia | 50mg | Lewis, 20097 Low | 8/35 [22.9] | 1/33 [3.0] | 7.5(1.0 to 57.1) | 14.01 | 0.20(0.05 to 0.35) | 9.89 |
| Paresthesia | 50mg | Winner, 200615 Low | 1/12 [8.3] | 2/12 [16.7] | 0.5(0.1 to 4.8) | 11.2 | -0.08(-0.35 to 0.18) | 3.27 |
| Paresthesia |  | Pooled | 9/47 [19.1] | 3/45 [6.7] | 2.0(0.1 to 29.1) | 25.21 | 0.08(-0.19 to 0.35) | 13.16 |
| Paresthesia | 100mg | Lewis, 20097 Low | 4/35 [11.4] | 1/33 [3.0] | 3.8(0.4 to 32.0) | 12.54 | 0.08(-0.04 to 0.21) | 15.5 |
| Paresthesia | 100mg | Winner, 200615 Low | 5/13 [38.5] | 2/12 [16.7] | 2.3(0.5 to 9.7) | 27.67 | 0.22(-0.12 to 0.56) | 1.97 |
| Paresthesia |  | Pooled | 9/48 [18.8] | 3/45 [6.7] | 2.7(0.8 to 8.9) | 40.21 | 0.10(-0.01 to 0.21) | 17.47 |
| Paresthesia | 200mg | Winner, 20058 Medium | 9/112 [8.0] | 0/50 [0.0] | 8.6(0.5 to 144.5) | 7.19 | 0.08(0.02 to 0.14) | 67.27 |
| Paresthesia | 200mg | Winner, 200615 Low | 5/14 [35.7] | 2/12 [16.7] | 2.1(0.5 to 9.1) | 27.39 | 0.19(-0.14 to 0.52) | 2.1 |
| Paresthesia |  | Pooled | 14/126 [11.1] | 2/62 [3.2] | 2.9(0.8 to 10.4) | 34.58 | 0.08(0.03 to 0.14) | 69.37 |
| **Paresthesia** |  | **Pooled** | **32/221 [14.5]** | **8/152 [5.3]** | **2.6(1.2 to 5.6)** | **100** | **0.09(0.05 to 0.14)** | **100** |
| Sinusitis | 50mg | Lewis, 20097 Low | 2/35 [5.7] | 1/33 [3.0] | 1.9(0.2 to 19.8) | 10.17 | 0.03(-0.07 to 0.12) | 25.26 |
| Sinusitis | 50mg | Winner, 200615 Low | 1/12 [8.3] | 0/12 [0.0] | 3.0(0.1 to 67.1) | 5.83 | 0.08(-0.12 to 0.29) | 5.74 |
| Sinusitis |  | Pooled | 3/47 [6.4] | 1/45 [2.2] | 2.2(0.3 to 14.6) | 16.01 | 0.04(-0.05 to 0.13) | 30.99 |
| Sinusitis | 100mg | Lewis, 20097 Low | 1/35 [2.9] | 1/33 [3.0] | 0.9(0.1 to 14.5) | 7.55 | 0.00(-0.08 to 0.08) | 36.45 |
| Sinusitis | 100mg | Winner, 200615 Low | 1/13 [7.7] | 0/12 [0.0] | 2.8(0.1 to 62.5) | 5.82 | 0.08(-0.12 to 0.27) | 6.34 |
| Sinusitis |  | Pooled | 2/48 [4.2] | 1/45 [2.2] | 1.5(0.2 to 11.8) | 13.37 | 0.01(-0.06 to 0.08) | 42.8 |
| Sinusitis | 200mg | Winner, 20058 Medium | 11/112 [9.8] | 6/50 [12.0] | 0.8(0.3 to 2.1) | 64.12 | -0.02(-0.13 to 0.08) | 21.14 |
| Sinusitis | 200mg | Winner, 200615 Low | 2/14 [14.3] | 0/12 [0.0] | 4.3(0.2 to 82.3) | 6.5 | 0.14(-0.07 to 0.36) | 5.07 |
| Sinusitis |  | Pooled | 13/126 [10.3] | 6/62 [9.7] | 1.0(0.3 to 3.1) | 70.62 | 0.03(-0.12 to 0.18) | 26.21 |
| Sinusitis |  | Pooled | 18/221 [8.1] | 8/152 [5.3] | 1.2(0.5 to 2.5) | 100 | 0.02(-0.03 to 0.07) | 100 |
| Somnolence | 50mg | Winner, 200615 Low | 1/12 [8.3] | 1/12 [8.3] | 1.0(0.1 to 14.2) | 11.69 | 0.00(-0.22 to 0.22) | 6.34 |
| Somnolence | 100mg | Lewis, 20097 Low | 2/35 [5.7] | 0/33 [0.0] | 4.7(0.2 to 94.8) | 9.15 | 0.06(-0.04 to 0.15) | 36.34 |
| Somnolence | 100mg | Winner, 200615 Low | 1/13 [7.7] | 1/12 [8.3] | 0.9(0.1 to 13.2) | 11.65 | -0.01(-0.22 to 0.21) | 6.82 |
| Somnolence |  | Pooled | 3/48 [6.3] | 1/45 [2.2] | 1.9(0.3 to 13.8) | 20.8 | 0.05(-0.04 to 0.13) | 43.17 |
| Somnolence | 200mg | Winner, 20058 Medium | 9/112 [8.0] | 3/50 [6.0] | 1.3(0.4 to 4.7) | 51.58 | 0.02(-0.06 to 0.10) | 45.15 |
| Somnolence | 200mg | Winner, 200615 Low | 2/14 [14.3] | 1/12 [8.3] | 1.7(0.2 to 16.6) | 15.93 | 0.06(-0.18 to 0.30) | 5.34 |
| Somnolence |  | Pooled | 11/126 [8.7] | 4/62 [6.5] | 1.4(0.5 to 4.3) | 67.51 | 0.03(-0.05 to 0.10) | 50.49 |
| Somnolence |  | Pooled | 15/186 [8.1] | 6/119 [5.0] | 1.4(0.6 to 3.6) | 100 | 0.03(-0.02 to 0.09) | 100 |
| Upper respiratory tract infection | 50mg | Lewis, 20097 Low | 7/35 [20.0] | 3/33 [9.1] | 2.2(0.6 to 7.8) | 16.37 | 0.11(-0.06 to 0.27) | 21.42 |
| Upper respiratory tract infection | 50mg | Winner, 200615 Low | 5/12 [41.7] | 1/12 [8.3] | 5.0(0.7 to 36.7) | 6.61 | 0.33(0.01 to 0.65) | 5.69 |
| Upper respiratory tract infection |  | Pooled | 12/47 [25.5] | 4/45 [8.9] | 2.8(1.0 to 8.1) | 22.99 | 0.18(-0.03 to 0.38) | 27.11 |
| Upper respiratory tract infection | 100mg | Lewis, 20097 Low | 8/35 [22.9] | 3/33 [9.1] | 2.5(0.7 to 8.7) | 17.11 | 0.14(-0.03 to 0.31) | 20.09 |
| Upper respiratory tract infection | 100mg | Winner, 200615 Low | 3/13 [23.1] | 1/12 [8.3] | 2.8(0.3 to 23.1) | 5.83 | 0.15(-0.13 to 0.43) | 7.57 |
| Upper respiratory tract infection |  | Pooled | 11/48 [22.9] | 4/45 [8.9] | 2.6(0.9 to 7.5) | 22.93 | 0.14(-0.01 to 0.29) | 27.66 |
| Upper respiratory tract infection | 200mg | Winner, 20058 Medium | 22/112 [19.6] | 8/50 [16.0] | 1.2(0.6 to 2.6) | 48.28 | 0.04(-0.09 to 0.16) | 36.99 |
| Upper respiratory tract infection | 200mg | Winner, 200615 Low | 3/14 [21.4] | 1/12 [8.3] | 2.6(0.3 to 21.6) | 5.8 | 0.13(-0.14 to 0.40) | 8.24 |
| Upper respiratory tract infection |  | Pooled | 25/126 [19.8] | 9/62 [14.5] | 1.3(0.7 to 2.7) | 54.08 | 0.05(-0.06 to 0.17) | 45.23 |
| **Upper respiratory tract infection** |  | **Pooled** | **48/221 [21.7]** | **17/152 [11.2]** | **1.8(1.1 to 3.1)** | **100** | **0.11(0.03 to 0.18)** | **100** |
| Weight decrease | 50mg | Lewis, 20097 Low | 10/35 [28.6] | 7/33 [21.2] | 1.3(0.6 to 3.1) | 32.69 | 0.07(-0.13 to 0.28) | 9.37 |
| Weight decrease | 50mg | Winner, 200615 Low | 2/12 [16.7] | 1/12 [8.3] | 2.0(0.2 to 19.2) | 4.51 | 0.08(-0.18 to 0.35) | 5.69 |
| Weight decrease |  | Pooled | 12/47 [25.5] | 8/45 [17.8] | 1.4(0.6 to 3.1) | 37.2 | 0.08(-0.08 to 0.24) | 15.06 |
| Weight decrease | 100mg | Lewis, 20097 Low | 17/35 [48.6] | 7/33 [21.2] | 2.3(1.1 to 4.8) | 42.12 | 0.27(0.06 to 0.49) | 8.36 |
| Weight decrease | 100mg | Winner, 200615 Low | 2/13 [15.4] | 1/12 [8.3] | 1.8(0.2 to 17.8) | 4.49 | 0.07(-0.18 to 0.32) | 6.23 |
| Weight decrease |  | Pooled | 19/48 [39.6] | 8/45 [17.8] | 2.2(1.1 to 4.5) | 46.61 | 0.18(-0.02 to 0.38) | 14.6 |
| Weight decrease | 200mg | Winner, 20058 Medium | 11/112 [9.8] | 2/50 [4.0] | 2.5(0.6 to 10.7) | 10.7 | 0.06(-0.02 to 0.14) | 65.47 |
| Weight decrease | 200mg | Winner, 200615 Low | 4/14 [28.6] | 1/12 [8.3] | 3.4(0.4 to 26.7) | 5.49 | 0.20(-0.08 to 0.49) | 4.87 |
| Weight decrease |  | Pooled | 15/126 [11.9] | 3/62 [4.8] | 2.8(0.8 to 9.1) | 16.19 | 0.07(-0.01 to 0.14) | 70.35 |
| **Weight decrease** |  | **Pooled** | **46/221 [20.8]** | **19/152 [12.5]** | **2.0(1.2 to 3.2)** | **100** | **0.09(0.02 to 0.15)** | **100** |

| **Outcome** | **Dose** | **Reference** | **Events/**  **Randomized**  **[rate,%] with drug** | **Events/**  **Randomized**  **[rate,%] with placebo** | **Peto Odds ratio**  **(95% CI)** | **% Weight** | **Arcsine transformed risk difference**  **(95% CI)** | **% Weight** |
| --- | --- | --- | --- | --- | --- | --- | --- | --- |
| Abdominal pain | 50mg | Lewis, 20097 Low | 3/35 [8.6] | 3/33 [9.1] | 0.9(0.2 to 5.0) | 17.19 | -0.01(-0.25 to 0.23) | 19.47 |
| Abdominal pain | 50mg | Winner, 200615 Low | 0/12 [0.0] | 1/12 [8.3] | 0.1(0.0 to 6.8) | 3.1 | -0.29(-0.69 to 0.11) | 6.88 |
| Abdominal pain |  | Pooled | 3/47 [6.4] | 4/45 [8.9] | 0.7(0.2 to 3.2) | 20.29 | -0.10(-0.37 to 0.16) | 26.34 |
| Abdominal pain | 100mg | Lewis, 20097 Low | 5/35 [14.3] | 3/33 [9.1] | 1.6(0.4 to 7.1) | 22.19 | 0.08(-0.16 to 0.32) | 19.47 |
| Abdominal pain | 100mg | Winner, 200615 Low | 2/13 [15.4] | 1/12 [8.3] | 1.9(0.2 to 20.2) | 8.51 | 0.11(-0.28 to 0.50) | 7.15 |
| Abdominal pain |  | Pooled | 7/48 [14.6] | 4/45 [8.9] | 1.7(0.5 to 5.9) | 30.7 | 0.09(-0.11 to 0.29) | 26.61 |
| Abdominal pain | 200mg | Winner, 20058 Medium | 11/112 [9.8] | 6/50 [12.0] | 0.8(0.3 to 2.3) | 40.5 | -0.04(-0.20 to 0.13) | 39.64 |
| Abdominal pain | 200mg | Winner, 200615 Low | 2/14 [14.3] | 1/12 [8.3] | 1.8(0.2 to 18.7) | 8.5 | 0.10(-0.29 to 0.48) | 7.4 |
| Abdominal pain |  | Pooled | 13/126 [10.3] | 7/62 [11.3] | 0.9(0.3 to 2.4) | 49.01 | -0.02(-0.17 to 0.14) | 47.05 |
| Abdominal pain |  | Pooled | 23/221 [10.4] | 15/152 [9.9] | 1.0(0.5 to 2.1) | 100 | -0.01(-0.11 to 0.10) | 100 |
| Anorexia | 50mg | Lewis, 20097 Low | 3/35 [8.6] | 1/33 [3.0] | 2.7(0.4 to 19.9) | 12.95 | 0.12(-0.12 to 0.36) | 19.47 |
| Anorexia | 50mg | Winner, 200615 Low | 1/12 [8.3] | 1/12 [8.3] | 1.0(0.1 to 17.0) | 6.49 | 0.00(-0.40 to 0.40) | 6.88 |
| Anorexia |  | Pooled | 4/47 [8.5] | 2/45 [4.4] | 1.9(0.4 to 9.9) | 19.43 | 0.09(-0.11 to 0.30) | 26.34 |
| Anorexia | 100mg | Lewis, 20097 Low | 4/35 [11.4] | 1/33 [3.0] | 3.4(0.6 to 20.6) | 15.93 | 0.17(-0.07 to 0.41) | 19.47 |
| Anorexia | 100mg | Winner, 200615 Low | 1/13 [7.7] | 1/12 [8.3] | 0.9(0.1 to 15.6) | 6.49 | -0.01(-0.40 to 0.38) | 7.15 |
| Anorexia |  | Pooled | 5/48 [10.4] | 2/45 [4.4] | 2.3(0.5 to 10.6) | 22.42 | 0.12(-0.08 to 0.32) | 26.61 |
| Anorexia | 200mg | Winner, 20058 Medium | 15/112 [13.4] | 4/50 [8.0] | 1.7(0.6 to 4.7) | 48.84 | 0.09(-0.08 to 0.25) | 39.64 |
| Anorexia | 200mg | Winner, 200615 Low | 2/14 [14.3] | 1/12 [8.3] | 1.8(0.2 to 18.7) | 9.3 | 0.10(-0.29 to 0.48) | 7.4 |
| Anorexia |  | Pooled | 17/126 [13.5] | 5/62 [8.1] | 1.7(0.7 to 4.4) | 58.15 | 0.09(-0.06 to 0.24) | 47.05 |
| Anorexia |  | Pooled | 26/221 [11.8] | 9/152 [5.9] | 1.9(0.9 to 3.8) | 100 | 0.10(-0.01 to 0.20) | 100 |
| Dizziness | 50mg | Lewis, 20097 Low | 2/35 [5.7] | 0/33 [0.0] | 7.2(0.4 to 117.5) | 18.55 | 0.24(0.00 to 0.48) | 22.1 |
| Dizziness | 50mg | Winner, 200615 Low | 0/12 [0.0] | 2/12 [16.7] | 0.1(0.0 to 2.1) | 18.03 | -0.42(-0.82 to -0.02) | 18.43 |
| Dizziness |  | Pooled | 2/47 [4.3] | 2/45 [4.4] | 1.0(0.1 to 7.1) | 36.58 | -0.07(-0.72 to 0.58) | 40.52 |
| Dizziness | 100mg | Lewis, 20097 Low | 3/35 [8.6] | 0/353 [0.0] | 7.4(0.7 to 73.8) | 27.4 | 0.30(0.06 to 0.54) | 22.1 |
| Dizziness | 100mg | Winner, 200615 Low | 0/13 [0.0] | 2/12 [16.7] | 0.1(0.0 to 1.9) | 18.03 | -0.42(-0.81 to -0.03) | 18.61 |
| Dizziness |  | Pooled | 3/48 [6.3] | 2/365 [4.4] | 1.4(0.2 to 8.4) | 45.44 | -0.04(-0.75 to 0.66) | 40.71 |
| Dizziness | 200mg | Winner, 200615 Low | 0/14 [0.0] | 2/12 [16.7] | 0.1(0.0 to 1.8) | 17.99 | -0.42(-0.81 to -0.04) | 18.77 |
| Dizziness |  | Pooled | 5/109 [4.6] | 6/422 [5.9] | 0.8(0.2 to 2.6) | 100 | -0.12(-0.46 to 0.22) | 100 |
| Fatigue | 50mg | Lewis, 20097 Low | 2/35 [5.7] | 2/33 [6.1] | 0.9(0.1 to 7.0) | 15.62 | -0.01(-0.25 to 0.23) | 19.47 |
| Fatigue | 50mg | Winner, 200615 Low | 0/12 [0.0] | 1/12 [8.3] | 0.1(0.0 to 6.8) | 4.09 | -0.29(-0.69 to 0.11) | 6.88 |
| Fatigue |  | Pooled | 2/47 [4.3] | 3/45 [6.7] | 0.6(0.1 to 3.8) | 19.71 | -0.10(-0.37 to 0.16) | 26.34 |
| Fatigue | 100mg | Lewis, 20097 Low | 3/35 [8.6] | 2/33 [6.1] | 1.4(0.2 to 8.8) | 19.22 | 0.05(-0.19 to 0.29) | 19.47 |
| Fatigue | 100mg | Winner, 200615 Low | 1/13 [7.7] | 1/12 [8.3] | 0.9(0.1 to 15.6) | 7.83 | -0.01(-0.40 to 0.38) | 7.15 |
| Fatigue |  | Pooled | 4/48 [8.3] | 3/45 [6.7] | 1.3(0.3 to 5.8) | 27.05 | 0.03(-0.17 to 0.24) | 26.61 |
| Fatigue | 200mg | Winner, 20058 Medium | 7/112 [6.3] | 6/50 [12.0] | 0.5(0.1 to 1.6) | 42.01 | -0.10(-0.27 to 0.07) | 39.64 |
| Fatigue | 200mg | Winner, 200615 Low | 2/14 [14.3] | 1/12 [8.3] | 1.8(0.2 to 18.7) | 11.23 | 0.10(-0.29 to 0.48) | 7.4 |
| Fatigue |  | Pooled | 9/126 [7.1] | 7/62 [11.3] | 0.6(0.2 to 1.8) | 53.24 | -0.07(-0.22 to 0.08) | 47.05 |
| Fatigue |  | Pooled | 15/221 [6.8] | 13/152 [8.6] | 0.7(0.3 to 1.7) | 100 | -0.05(-0.15 to 0.06) | 100 |
| Injury | 50mg | Lewis, 20097 Low | 3/35 [8.6] | 2/33 [6.1] | 1.4(0.2 to 8.8) | 17.42 | 0.05(-0.19 to 0.29) | 19.47 |
| Injury | 50mg | Winner, 200615 Low | 1/12 [8.3] | 1/12 [8.3] | 1.0(0.1 to 17.0) | 7.09 | 0.00(-0.40 to 0.40) | 6.88 |
| Injury |  | Pooled | 4/47 [8.5] | 3/45 [6.7] | 1.3(0.3 to 5.9) | 24.52 | 0.04(-0.17 to 0.24) | 26.34 |
| Injury | 100mg | Lewis, 20097 Low | 4/35 [11.4] | 2/33 [6.1] | 1.9(0.4 to 10.2) | 20.57 | 0.10(-0.14 to 0.33) | 19.47 |
| Injury | 100mg | Winner, 200615 Low | 1/13 [7.7] | 1/12 [8.3] | 0.9(0.1 to 15.6) | 7.1 | -0.01(-0.40 to 0.38) | 7.15 |
| Injury |  | Pooled | 5/48 [10.4] | 3/45 [6.7] | 1.6(0.4 to 6.7) | 27.67 | 0.07(-0.14 to 0.27) | 26.61 |
| Injury | 200mg | Winner, 20058 Medium | 8/112 [7.1] | 6/50 [12.0] | 0.5(0.2 to 1.8) | 40.74 | -0.08(-0.25 to 0.08) | 39.64 |
| Injury | 200mg | Winner, 200615 Low | 1/14 [7.1] | 1/12 [8.3] | 0.9(0.1 to 14.5) | 7.08 | -0.02(-0.41 to 0.36) | 7.4 |
| Injury |  | Pooled | 9/126 [7.1] | 7/62 [11.3] | 0.6(0.2 to 1.7) | 47.82 | -0.07(-0.23 to 0.08) | 47.05 |
| Injury |  | Pooled | 18/221 [8.1] | 13/152 [8.6] | 0.9(0.4 to 2.0) | 100 | -0.01(-0.11 to 0.10) | 100 |
| Mood problems | 50mg | Winner, 200615 Low | 0/12 [0.0] | 2/12 [16.7] | 0.1(0.0 to 2.1) | 22.33 | -0.42(-0.82 to -0.02) | 17.27 |
| Mood problems | 50mg | 20159428 Low | 1/35 [2.9] | 0/33 [0.0] | 7.0(0.1 to 352.3) | 11.66 | 0.17(-0.07 to 0.41) | 23.68 |
| Mood problems |  | Pooled | 1/47 [2.1] | 2/45 [4.4] | 0.5(0.1 to 4.9) | 33.99 | -0.10(-0.68 to 0.47) | 40.96 |
| Mood problems | 100mg | Winner, 200615 Low | 0/13 [0.0] | 2/12 [16.7] | 0.1(0.0 to 1.9) | 22.33 | -0.42(-0.81 to -0.03) | 17.55 |
| Mood problems | 100mg | 20159428 Low | 1/35 [2.9] | 0/33 [0.0] | 7.0(0.1 to 352.3) | 11.66 | 0.17(-0.07 to 0.41) | 23.68 |
| Mood problems |  | Pooled | 1/48 [2.1] | 2/45 [4.4] | 0.5(0.0 to 4.6) | 33.99 | -0.10(-0.68 to 0.47) | 41.24 |
| Mood problems | 200mg | Winner, 200615 Low | 1/14 [7.1] | 2/12 [16.7] | 0.4(0.0 to 4.3) | 32.02 | -0.15(-0.54 to 0.24) | 17.8 |
| Mood problems |  | Pooled | 3/109 [2.8] | 6/102 [5.9] | 0.5(0.1 to 1.7) | 100 | -0.09(-0.35 to 0.16) | 100 |
| Nausea | 50mg | Lewis, 20097 Low | 2/35 [5.7] | 2/33 [6.1] | 0.9(0.1 to 7.0) | 24.14 | -0.01(-0.25 to 0.23) | 24.77 |
| Nausea | 100mg | Lewis, 20097 Low | 3/35 [8.6] | 2/33 [6.1] | 1.4(0.2 to 8.8) | 29.7 | 0.05(-0.19 to 0.29) | 24.77 |
| Nausea | 200mg | Winner, 20058 Medium | 6/112 [5.4] | 3/50 [6.0] | 0.9(0.2 to 3.8) | 46.16 | -0.01(-0.18 to 0.15) | 50.45 |
| Nausea |  | Pooled | 11/182 [6.0] | 7/116 [6.0] | 1.0(0.4 to 2.8) | 100 | 0.00(-0.12 to 0.12) | 100 |
| Paresthesia | 50mg | Lewis, 20097 Low | 8/35 [22.9] | 1/33 [3.0] | 5.5(1.4 to 22.1) | 23.86 | 0.32(0.09 to 0.56) | 19.47 |
| Paresthesia | 50mg | Winner, 200615 Low | 1/12 [8.3] | 2/12 [16.7] | 0.5(0.0 to 5.1) | 8.25 | -0.13(-0.53 to 0.27) | 6.88 |
| Paresthesia |  | Pooled | 9/47 [19.1] | 3/45 [6.7] | 2.9(0.9 to 9.7) | 32.12 | 0.13(-0.31 to 0.57) | 26.34 |
| Paresthesia | 100mg | Lewis, 20097 Low | 4/35 [11.4] | 1/33 [3.0] | 3.4(0.6 to 20.6) | 14.16 | 0.17(-0.07 to 0.41) | 19.47 |
| Paresthesia | 100mg | Winner, 200615 Low | 5/13 [38.5] | 2/12 [16.7] | 2.8(0.5 to 15.6) | 15.79 | 0.25(-0.14 to 0.64) | 7.15 |
| Paresthesia |  | Pooled | 9/48 [18.8] | 3/45 [6.7] | 3.1(0.9 to 10.6) | 29.95 | 0.19(-0.01 to 0.39) | 26.61 |
| Paresthesia | 200mg | Winner, 20058 Medium | 9/112 [8.0] | 0/50 [0.0] | 4.6(1.1 to 19.5) | 22 | 0.29(0.12 to 0.45) | 39.64 |
| Paresthesia | 200mg | Winner, 200615 Low | 5/14 [35.7] | 2/12 [16.7] | 2.5(0.5 to 14.0) | 15.94 | 0.22(-0.17 to 0.61) | 7.4 |
| **Paresthesia** |  | **Pooled** | **14/126 [11.1]** | **2/62 [3.2]** | **3.6(1.2 to 10.8)** | **37.93** | **0.28(0.12 to 0.43)** | **47.05** |
| **Paresthesia** |  | **Pooled** | **32/221 [14.5]** | **8/152 [5.3]** | **3.2(1.6 to 6.3)** | **100** | **0.24(0.13 to 0.34)** | **100** |
| Sinusitis | 50mg | Lewis, 20097 Low | 2/35 [5.7] | 1/33 [3.0] | 1.9(0.2 to 18.6) | 13.31 | 0.07(-0.17 to 0.30) | 20.37 |
| Sinusitis | 50mg | Winner, 200615 Low | 1/12 [8.3] | 0/12 [0.0] | 7.4(0.1 to 372.4) | 4.58 | 0.29(-0.11 to 0.69) | 9.07 |
| Sinusitis |  | Pooled | 3/47 [6.4] | 1/45 [2.2] | 2.7(0.4 to 19.3) | 17.88 | 0.13(-0.08 to 0.33) | 29.44 |
| Sinusitis | 100mg | Lewis, 20097 Low | 1/35 [2.9] | 1/33 [3.0] | 0.9(0.1 to 15.4) | 9.01 | -0.01(-0.24 to 0.23) | 20.37 |
| Sinusitis | 100mg | Winner, 200615 Low | 1/13 [7.7] | 0/12 [0.0] | 6.8(0.1 to 345.9) | 4.57 | 0.28(-0.11 to 0.67) | 9.38 |
| Sinusitis |  | Pooled | 2/48 [4.2] | 1/45 [2.2] | 1.8(0.2 to 17.9) | 13.58 | 0.09(-0.17 to 0.36) | 29.75 |
| Sinusitis | 200mg | Winner, 20058 Medium | 11/112 [9.8] | 6/50 [12.0] | 0.8(0.3 to 2.3) | 59.8 | -0.04(-0.20 to 0.13) | 31.15 |
| Sinusitis | 200mg | Winner, 200615 Low | 2/14 [14.3] | 0/12 [0.0] | 6.9(0.4 to 118.1) | 8.73 | 0.39(0.00 to 0.77) | 9.66 |
| Sinusitis |  | Pooled | 13/126 [10.3] | 6/62 [9.7] | 1.0(0.4 to 2.9) | 68.54 | 0.14(-0.27 to 0.55) | 40.81 |
| Sinusitis |  | Pooled | 18/221 [8.1] | 8/152 [5.3] | 1.3(0.6 to 3.1) | 100 | 0.09(-0.04 to 0.22) | 100 |
| Somnolence | 50mg | Winner, 200615 Low | 1/12 [8.3] | 1/12 [8.3] | 1.0(0.1 to 17.0) | 10.58 | 0.00(-0.40 to 0.40) | 8.54 |
| Somnolence | 100mg | Lewis, 20097 Low | 2/35 [5.7] | 0/33 [0.0] | 7.2(0.4 to 117.5) | 10.89 | 0.24(0.00 to 0.48) | 24.17 |
| Somnolence | 100mg | Winner, 200615 Low | 1/13 [7.7] | 1/12 [8.3] | 0.9(0.1 to 15.6) | 10.58 | -0.01(-0.40 to 0.38) | 8.87 |
| Somnolence |  | Pooled | 3/48 [6.3] | 1/45 [2.2] | 2.6(0.4 to 19.1) | 21.47 | 0.17(-0.06 to 0.39) | 33.05 |
| Somnolence | 200mg | Winner, 20058 Medium | 9/112 [8.0] | 3/50 [6.0] | 1.3(0.4 to 4.8) | 52.78 | 0.04(-0.13 to 0.21) | 49.23 |
| Somnolence | 200mg | Winner, 200615 Low | 2/14 [14.3] | 1/12 [8.3] | 1.8(0.2 to 18.7) | 15.17 | 0.10(-0.29 to 0.48) | 9.19 |
| Somnolence |  | Pooled | 11/126 [8.7] | 4/62 [6.5] | 1.4(0.5 to 4.4) | 67.95 | 0.05(-0.10 to 0.20) | 58.42 |
| Somnolence |  | Pooled | 15/186 [8.1] | 6/119 [5.0] | 1.6(0.6 to 3.9) | 100 | 0.09(-0.03 to 0.20) | 100 |
| Upper respiratory tract infection | 50mg | Lewis, 20097 Low | 7/35 [20.0] | 3/33 [9.1] | 2.4(0.6 to 8.9) | 17.07 | 0.16(-0.08 to 0.40) | 19.47 |
| Upper respiratory tract infection | 50mg | Winner, 200615 Low | 5/12 [41.7] | 1/12 [8.3] | 5.5(0.9 to 33.5) | 9.26 | 0.41(0.01 to 0.81) | 6.88 |
| **Upper respiratory tract infection** |  | **Pooled** | **12/47 [25.5]** | **4/45 [8.9]** | **3.2(1.1 to 9.3)** | **26.33** | **0.23(0.01 to 0.45)** | **26.34** |
| Upper respiratory tract infection | 100mg | Lewis, 20097 Low | 8/35 [22.9] | 3/33 [9.1] | 2.7(0.8 to 9.8) | 18.45 | 0.19(-0.05 to 0.43) | 19.47 |
| Upper respiratory tract infection | 100mg | Winner, 200615 Low | 3/13 [23.1] | 1/12 [8.3] | 2.9(0.4 to 23.3) | 6.89 | 0.21(-0.18 to 0.60) | 7.15 |
| Upper respiratory tract infection |  | Pooled | 11/48 [22.9] | 4/45 [8.9] | 2.8(0.9 to 8.2) | 25.34 | 0.20(-0.01 to 0.40) | 26.61 |
| Upper respiratory tract infection | 200mg | Winner, 20058 Medium | 22/112 [19.6] | 8/50 [16.0] | 1.3(0.5 to 3.0) | 41.42 | 0.05(-0.12 to 0.21) | 39.64 |
| Upper respiratory tract infection | 200mg | Winner, 200615 Low | 3/14 [21.4] | 1/12 [8.3] | 2.6(0.3 to 21.4) | 6.9 | 0.19(-0.20 to 0.57) | 7.4 |
| Upper respiratory tract infection |  | Pooled | 25/126 [19.8] | 9/62 [14.5] | 1.4(0.6 to 3.1) | 48.33 | 0.07(-0.08 to 0.22) | 47.05 |
| **Upper respiratory tract infection** |  | **Pooled** | **48/221 [21.7]** | **17/152 [11.2]** | **2.1(1.2 to 3.6)** | **100** | **0.14(0.04 to 0.25)** | **100** |
| Weight decrease | 50mg | Lewis, 20097 Low | 10/35 [28.6] | 7/33 [21.2] | 1.5(0.5 to 4.4) | 26.6 | 0.09(-0.15 to 0.32) | 19.47 |
| Weight decrease | 50mg | Winner, 200615 Low | 2/12 [16.7] | 1/12 [8.3] | 2.1(0.2 to 22.2) | 5.64 | 0.13(-0.27 to 0.53) | 6.88 |
| Weight decrease |  | Pooled | 12/47 [25.5] | 8/45 [17.8] | 1.6(0.6 to 4.2) | 32.24 | 0.10(-0.11 to 0.30) | 26.34 |
| Weight decrease | 100mg | Lewis, 20097 Low | 17/35 [48.6] | 7/33 [21.2] | 3.3(1.2 to 8.7) | 32.4 | 0.29(0.06 to 0.53) | 19.47 |
| Weight decrease | 100mg | Winner, 200615 Low | 2/13 [15.4] | 1/12 [8.3] | 1.9(0.2 to 20.2) | 5.65 | 0.11(-0.28 to 0.50) | 7.15 |
| **Weight decrease** |  | **Pooled** | **19/48 [39.6]** | **8/45 [17.8]** | **3.0(1.2 to 7.5)** | **38.05** | **0.24(0.04 to 0.45)** | **26.61** |
| Weight decrease | 200mg | Winner, 20058 Medium | 11/112 [9.8] | 2/50 [4.0] | 2.2(0.6 to 7.4) | 21.13 | 0.12(-0.05 to 0.28) | 39.64 |
| Weight decrease | 200mg | Winner, 200615 Low | 4/14 [28.6] | 1/12 [8.3] | 3.5(0.5 to 23.8) | 8.59 | 0.27(-0.11 to 0.66) | 7.4 |
| Weight decrease |  | Pooled | 15/126 [11.9] | 3/62 [4.8] | 2.5(0.9 to 7.0) | 29.72 | 0.14(-0.01 to 0.30) | 47.05 |
| **Weight decrease** |  | **Pooled** | **46/221 [20.8]** | **19/152 [12.5]** | **2.3(1.3 to 4.0)** | **100** | **0.16(0.05 to 0.26)** | **100** |

| **Outcome** | **Topiramate**  **dose** | **Degree of freedom** | **Effect Measure** | **P** | **I-squared** | **Effect**  **Measure** | **P** | **I-squared** |
| --- | --- | --- | --- | --- | --- | --- | --- | --- |
| Abdominal pain | 50 | 1 | Relative risk | 0.556 | 0.00% | Risk difference | 0.529 | 0.00% |
| Abdominal pain | 100 | 1 | Relative risk | 0.905 | 0.00% | Risk difference | 0.901 | 0.00% |
| Abdominal pain | 200 | 1 | Relative risk | 0.556 | 0.00% | Risk difference | 0.545 | 0.00% |
| Abdominal pain | Overall | 5 | Relative risk | 0.905 | 0.00% | Risk difference | 0.875 | 0.00% |
| Anorexia | 50 | 1 | Relative risk | 0.555 | 0.00% | Risk difference | 0.66 | 0.00% |
| Anorexia | 100 | 1 | Relative risk | 0.419 | 0.00% | Risk difference | 0.469 | 0.00% |
| Anorexia | 200 | 1 | Relative risk | 0.985 | 0.00% | Risk difference | 0.966 | 0.00% |
| Anorexia | Overall | 5 | Relative risk | 0.957 | 0.00% | Risk difference | 0.978 | 0.00% |
| Dizziness | 50 | 1 | Relative risk | 0.14 | 54.10% | Risk difference | 0.086 | 66.00% |
| Dizziness | 100 | 1 | Relative risk | 0.091 | 64.90% | Risk difference | 0.055 | 72.90% |
| Dizziness | Overall | 4 | Relative risk | 0.187 | 35.10% | Risk difference | 0.052 | 57.30% |
| Fatigue | 50 | 1 | Relative risk | 0.576 | 0.00% | Risk difference | 0.499 | 0.00% |
| Fatigue | 100 | 1 | Relative risk | 0.792 | 0.00% | Risk difference | 0.802 | 0.00% |
| Fatigue | 200 | 1 | Relative risk | 0.35 | 0.00% | Risk difference | 0.38 | 0.00% |
| Fatigue | Overall | 5 | Relative risk | 0.871 | 0.00% | Absolute risk difference | 0.858 | 0.00% |
| Injury | 50 | 1 | Relative risk | 0.83 | 0.00% | Risk difference | 0.846 | 0.00% |
| Injury | 100 | 1 | Relative risk | 0.653 | 0.00% | Risk difference | 0.639 | 0.00% |

Bold = significant at 95% confidence level when 95% CI of absolute risk difference do not include 0