Supplementary Table 1. Sociodemographic and clinical characteristics of non-remitters and remitters in CATIE.

| | All sample ^a | Non-remission | Remission N = 239 | |
|--------------------------------|-------------------------|----------------|----------------------|-----------------------------|
| | N = 765 | N=372 | | |
| | N/Mean (%/SD) | N/Mean (%/SD) | N/Mean (%/SD) | X ² /F (p-value) |
| Age | 40.93 (11.03) | 41.52 (11.32) | 40.14 (11.09) | 0.022 (0.883) |
| Total | 765 (100%) | 372 (100%) | 239 (100%) | |
| Missing | - | - | - | |
| Gender (F/tot) | 206/765 (26.93%) | 89/372 (23.9%) | 68/239 (28.5%) | 1.562 (0.211) |
| Total | 765 (100%) | 372 (100%) | 239 (100%) | |
| Missing | - | - | - | |
| Employment (Full time/tot) | 48/761 (6.31%) | 15/371 (4.0%) | 21/237 (8.9%) | 6.026 (0.014) |
| Total | 761 (99.48%) | 371 (99.73%) | 237 (99.16%) | |
| Missing | 4 (0.52%) | 1 (0.27%) | 2 (0.84%) | |
| Social withdrawal at baseline | 2.83 (1.74) | 2.52 (1.59) | 3.35 (1.92) | 25.117 (<0.001) |
| (QOLS 8) | | | | |
| Total | 762 (99.61%) | 371 (99.73%) | 237 (99.16%) | |
| Missing | 3 (0.39%) | 1 (0.27%) | 2 (0.84%) | |
| Psychopathological severity at | 74.20 (17.72) | 81.31 (15.40) | 62.21 (14.82) | 0.232 (0.630) |
| baseline (PANSS) | | | | |
| Total | 765 (100%) | 372 (100%) | 239 (100%) | |
| Missing | - | - | - | |
| Psychopathological severity at | 66.11 (17.61) | 77.18 (12.201) | 48.88 (8.553) | 24.427 (<0.001) |
| week 12 (PANSS) | | | | |
| Total | 611 (79.87%) | 372 (100%) | 239 (100%) | |
| Missing | 154 (20.13%) | - | - | |

Abbreviations: CATIE, Clinical Antipsychotic Trials of Intervention Effectiveness; SD, Standard deviations; F, Female; QUOLS 8, Quality-of-Life Scale item 8; PANSS, Positive and Negative Syndrome Scale.

^a including 154 individuals with missing information on remission status

Supplementary Table 2. Sociodemographic and clinical characteristics of non-remitters and remitters in CO-MED.

| | All sample ^a N = 482 | Non-remission N = 202 N/Mean (%/SD) | Remission | |
|-------------------------------|------------------------------------|---|-----------------|-----------------------------|
| | | | N = 150 | |
| | N/Mean (%/SD) | | N/Mean (%/SD) | X ² /F (p-value) |
| Age | 43.17 (12.46) | 44.69 (11.89) | 43.39 (12.71) | 1.127 (0.289) |
| Total | 482 (100%) | 202 (100%) | 150 (100%) | |
| Missing | - | - | - | |
| Gender (F/tot) | 338/482 (70.1%) | 146/202 (%) | 105/150 (%) | 0.218 (0.640) |
| Total | 482 (100%) | 202 (100%) | 150 (100%) | |
| Missing | - | - | - | |
| Employment (Employed/tot) | 237/482 (49.2%) | 98/202 (51.75%) | 83/150 (50.81%) | 1.602 (0.206) |
| Total | 482 (100%) | 202 (100%) | 150 (100%) | |
| Missing | - | - | - | |
| Social withdrawal at baseline | 11.70 (3.76) | 11.87 (3.67) | 11.18 (3.85) | 1.540 (0.215) |
| (WSAS 03 + WSAS 05) | | | | |
| Total | 482 (100%) | 202 (100%) | 150 (100%) | |
| Missing | - | - | - | |
| Psychopathological severity | 20.63 (6.25) | 20.90 (6.01) | 19.91 (6.69) | 2.864 (0.091) |
| at baseline (QIDS-C) | | | | |
| Total | 482 (100%) | 202 (100%) | 150 (100%) | |
| Missing | - | - | - | |
| Psychopathological severity | 8.12 (6.32) | 12.06 (5.60) | 2.81 (1.58) | 104.934 (<0.001) |
| at week 12 (QIDS-C) | | | | |
| Total | 352 (73.03%) | 202 (100%) | 150 (100%) | |
| Missing | 130 (26.97%) | - | - | |

Abbreviations: CO-MED, Combining Medications to Enhance Depression Outcomes; SD, Standard deviations; F, Female; WSAS 03, Work and Social Adjustment Scale item 3; WSAS 05, Work and Social Adjustment Scale item 5; QUIDS-C, Quick Inventory of Depressive Symptomatology-Clinician rated.

^a including 130 individuals with missing information on remission status

Supplementary Table 3. Sociodemographic and clinical characteristics of non-remitters and remitters in GSRD.

| | All sample | Non-remission N = 1197 | Remission | |
|--------------------------------|------------------|---------------------------|------------------|-----------------------------|
| | N = 1397 | | N = 200 | |
| | N/Mean (%/SD) | N/Mean (%/SD) | N/Mean (%/SD) | X ² /F (p-value) |
| Age | 56.06 (84.82) | 55.84 (86.50) | 57.39 (74.17) | 0.160 (0.690) |
| Total | 1396 (99.93%) | 1197 (100%) | 200 (100%) | |
| Missing | 1 (0.07%) | - | - | |
| Gender (F/tot) | 926/1395 (66.4%) | 788/1195 (65.94%) | 138/200 (69.00%) | 0.718 (0.397) |
| Total | 1395 (99.86%) | 1195 (99.83%) | 200 (100%) | |
| Missing | 2 (0.14%) | 2 (0.17%) | - | |
| Employment (Employed/tot) | 636/1361 (49.7%) | 539/1172 (45.99%) | 97/189 (51.32%) | 1.859 (0.173) |
| Total | 1361 (97.42%) | 1172 (97.91%) | 189 (94.50%) | |
| Missing | 36 (2.58%) | 25 (2.08%) | 11 (5.50%) | |
| Social withdrawal at baseline | 3.59 (1.17) | 3.65 (1.13) | 3.24 (1.30) | 3.697 (0.055) |
| (MADRS item 8) | | | | |
| Total | 1394 (99.78%) | 1196 (99.92%) | 198 (0.99%) | |
| Missing | 3 (0.22%) | 1 (0.08%) | 2 (0.01%) | |
| Psychopathological severity at | 24.71 (11.28) | 27.99 (8.47) | 5.07 (2.76) | 134.162 (<0.001) |
| study entry (MADRS) | | | | |
| Total | 1397 (100%) | 1197 (100%) | 200 (100%) | |
| Missing | - | - | - | |
| Psychopathological severity, | 34.13 (7.52) | 34.51 (7.38) | 31.89 (7.94) | 3.665 (0.056) |
| retrospective (rMADRS) | | | | |
| Total | 1397(100%) | 1197 (100%) | 200 (100%) | |
| Missing | - | - | - | |

Abbreviations: GSRD, European Group for the Study of Resistant Depression; SD, Standard deviations; F, Female; MADRS, Montgomery and Asberg Depression Rating Scale; rMADRS, retrospective Montgomery and Asberg Depression Rating Scale.

Supplementary Table 4. Sociodemographic and clinical characteristics of non-remitters and remitters in STAR*D.

| | All sample ^a N = 1953 | Non-remission N = 937 | Remission | |
|-----------------------------|-------------------------------------|--------------------------|------------------|-----------------------------|
| | | | N = 509 | |
| | N/Mean (%/SD) | N/Mean (%/SD) | N/Mean (%/SD) | X ² /F (p-value) |
| Age | 43.15 (13.37) | 44.45 (12.85) | 42.71 (13.33) | 1.660 (0.198) |
| Total | 1953 (100%) | 937 (100%) | 509 (100%) | |
| Missing | - | - | - | |
| Gender (F/tot) | 1205/1953 (61.70%) | 580/937 (61.90%) | 328/509 (64.44%) | 0.911 (0.340) |
| Total | 1953 (100%) | 937 (100%) | 509 (100%) | |
| Missing | - | - | - | |
| Employment (Full time/tot) | 738/1953 (37.79%) | 315/937 (33.62%) | 240/509 (47.15%) | 25.543 (<0.001) |
| Total | 1953 (100%) | 937 (100%) | 509 (100%) | |
| Missing | - | - | - | |
| Social withdrawal at | 9.96 (4.08) | 10.44 (3.91) | 9.03 (4.04) | 1.421 (0.234) |
| baseline (WSAS 03+WSAS | | | | |
| 05) | | | | |
| Total | 1876 (96.06%) | 924 (98.61%) | 498 (97.84%) | |
| Missing | 77 (3.94%) | 13 (1.39%) | 9 (2.16%) | |
| Psychopathological severity | 15.42 (5.10) | 16.35 (4.61) | 13.82 (5.24) | 11.95 (0.001) |
| at baseline (QIDS-C) | | | | |
| Total | 1876 (96.06%) | 924 (98.61%) | 498 (97.84%) | |
| Missing | 77 (3.94%) | 13 (1.39%) | 9 (2.16) | |
| Psychopathological severity | 9.35 (6.37) | 12.85 (5.13) | 2.89 (1.58) | 523.237 (<0.001) |
| at week 12 (QIDS-C) | | | | |
| Total | 1446 (74.04%) | 937 (100%) | 509 (%) | |
| Missing | 507 (25.86%) | - | - | |

Abbreviations: STAR*D, Sequenced Treatment Alternatives to Relieve Depression; SD, Standard deviations; F, Female; WSAS 03, Work and Social Adjustment Scale item 3; WSAS 05, Work and Social Adjustment Scale item 5; QUIDS-C, Quick Inventory of Depressive Symptomatology-Clinician Rated.

^a including 507 individuals with missing information on remission status

Supplementary Table 5. Sociodemographic and clinical characteristics of non-remitters and remitters in STEP-BD.

| | All sample a | Non-remission | Remission | |
|--------------------------------|------------------|------------------|-----------------|-----------------------------|
| | N = 1056 | N = 519 | N = 136 | |
| | N/Mean (%/SD) | N/Mean (%/SD) | N/Mean (%/SD) | X ² /F (p-value) |
| Age | 39.86 (11.46) | 41.10 (10.76) | 40.38 (12.06) | 4.840 (0.028) |
| Total | 1056 (100%) | 519 (100%) | 136 (100%) | |
| Missing | - | - | - | |
| Gender (F/tot) | 623/1055 (59.1%) | 308/518 (59.46%) | 79/136 (58.09%) | 0.084 (0.772) |
| Total | 1055 (100%) | 518 (100%) | 136 (100%) | |
| Missing | 1 (transgender) | 1 (transgender) | - | |
| Employment (Full time/tot) | 685/738 (92.8%) | 361/377 (95.75%) | 69/80 (86.25%) | 10.728 (0.001) |
| Total | 738 (69.89%) | 377 (72.64%) | 80 (58.82%) | |
| Missing | 318 (30.11%) | 142 (27.36%) | 56 (41.18%) | |
| Social withdrawal at the | 5.91 (2.47) | 6.02 (2.44) | 5.52 (2.54) | 1.150 (0.284) |
| baseline (LRIFT 2c + LRIFT | | | | |
| 2d) | | | | |
| Total | 1037 (98.20%) | 513 (98.84%) | 135 (99.26%) | |
| Missing | 19 (1.80%) | 6 (1.16%) | 1 (0.74%) | |
| Psychopathological severity at | 29.01 (5.49) | 29.40 (5.48) | 27.67 (5.14) | 0.798 (0.372) |
| baseline (MADRS) | | | | |
| Total | 1056 (100%) | 519 (100%) | 136 (100%) | |
| Missing | - | - | - | |
| Psychopathological severity at | 19.71 (10.80) | 23.60 (8.50) | 4.90 (2.97) | 111.632 (<0.001) |
| week 12 (MADRS) | | | | |
| Total | 655 (62.03%) | 519 (100%) | 136 (100%) | |
| Missing | 401 (37.97%) | - | - | |
| Symptoms of mania at | 7.32 (4.94) | 7.40 (4.91) | 6.28 (4.68) | 2.318 (0.128) |
| baseline (YMRS) | | | | |
| Total | 1045 (98.95%) | 516 (99.42%) | 135 (99.26%) | |
| Missing | 11 (1.05%) | 3 (0.58%) | 1 (0.74%) | |
| Symptoms of mania at week | 6.69 (6.18) | 7.16 (6.20) | 4.93 (5.87) | 1.486 (0.223) |
| 12 (YMRS) | | | | |
| Total | 660 (62.50%) | 513 (98.84%) | 136 (100%) | |
| Missing | 396 (37.50%) | 6 (1.16%) | - | |

Abbreviations: STEP-BD, Systematic Treatment Enhancement Program for Bipolar Disorder; SD, Standard deviations; F, Female; LRIFT 2c, Life-Range of Impaired Functioning Tool item 2c; LRIFT 2d, Life-Range of Impaired Functioning Tool item 2d; MADRS, Montgomery and Asberg Depression Rating Scale; YMRS, Young Mania Rating Scale.

^a including 401 individuals with missing information on remission status