

Supplementary Table S1. Stratified and total pooled random-effect estimates of the proportion of drop-outs due to any adverse event and stratification by diagnosis. % W is the percentage weight of each study

Drop-outs due to any adverse event	Events (n)	Patients (n)	Proportion	95%-CI	%W (random)
Unipolar					
Fawcett et al. 2016 [85]	5	24	20.8%	7.1% - 42.2%	6.3
Hori et al., 2012 [49]	0	12	0.0%	0.0% - 26.5%	15.2
Inoue et al., 2010 [81]	2	10	20.0%	2.5% - 55.6%	2.7
Sporn et al., 2000 [84]	1	20	5.0%	0.1% - 24.9%	18.2
Tundo et al., 2022 [86]	5	79	6.3%	2.1% - 14.2%	57.6
Random-effect			6.4%	2.3% - 10.5%	
I^2 , %			30.0		
Bipolar					
Fawcett et al. 2016 [85]	3	18	16.7%	3.6% - 41.4%	16.9
Hori et al., 2012 [49]	0	5	0.0%	0.0% - 52.2%	10.2
Sporn et al., 2000 [84]	3	12	25.0%	5.5% - 57.2%	8.3
Tundo et al., 2022 [86]	3	37	8.1%	1.7% - 21.9%	64.6
Random-effect			10.1%	3.1% - 17.2%	
I^2 , %			0.0		
Total					
Cassano et al., 2004 [80]	5	23	21.7%	7.5% - 43.7%	9.4
Fawcett et al. 2016 [85]	8	42	19.1%	8.6% - 34.1%	14.1
Hori et al., 2012 [49]	0	17	0.0%	0.0% - 19.5%	19.7
Inoue et al., 2010 [81]	2	10	20.0%	2.5% - 55.6%	5.4
Lattanzi et al., 2002 [82]	5	31	16.1%	5.5% - 33.7%	12.9
Sporn et al., 2000 [84]	4	32	12.5%	3.5% - 29.0%	14.6
Tundo et al., 2022 [86]	8	116	6.9%	3.0% - 13.1%	23.9
Overall random-effects			11.4%	5.0% - 17.8%	
I^2 , %			54.9		

Supplementary Table S2. Stratified and total pooled random-effect estimates of the proportion of drop-outs due to any reasons and stratification by diagnosis.

Drop-outs due to any reasons	Events (n)	Patients (n)	Proportion	95%-CI	%W (random)
Unipolar					
Hori et al., 2012 [49]	2	12	16.7%	2.1% - 48.4%	22.8
Inoue et al., 2010 [81]	2	10	20.0%	2.5% - 55.6%	19.9
Sporn et al., 2000 [84]	9	20	45.0%	23.1% - 68.5%	22.2
Tundo et al., 2022 [86]	6	79	7.6%	2.8% - 15.8%	35.2
Random-effect			20.4%	4.1% - 36.7%	
<i>I², %</i>			73.8		
Bipolar					
Hori et al., 2012 [49]	0	5	0.0%	0.0% - 52.2%	17.1
Sporn et al., 2000 [84]	3	12	25.0%	5.5% - 57.2%	13.9
Tundo et al., 2022 [86]	5	37	13.5%	4.5% - 28.8%	69.0
Random-effect			12.8%	3.7% - 22.0%	
<i>I², %</i>			11.3		
Total					
Cassano et al., 2004 [80]	14	23	60.9%	38.5% - 80.3%	13.0
Fawcett et al. 2016 [85]	15	42	35.7%	21.6% - 52.0%	15.0
Hori et al., 2012 [49]	2	17	11.8%	1.5% - 36.4%	14.7
Inoue et al., 2010 [81]	2	10	20.0%	2.5% - 55.6%	11.3
Lattanzi et al., 2002 [82]	12	31	38.7%	21.9% - 57.8%	14.1
Sporn et al., 2000 [84]	12	32	37.5%	21.1% - 56.3%	14.2
Tundo et al., 2022 [86]	11	116	9.5%	4.8% - 16.3%	17.6
Overall random-effects			29.7%	16.2% - 43.3%	
<i>I², %</i>			86.6		

Supplementary Table S3. Stratified and total pooled random-effect estimates of the proportion of (hypo)mania switch and stratification by diagnosis.

(Hypo)mania switch	Events (n)	Patients (n)	Proportion	95%-CI	%W (random)
Unipolar					
Cassano et al., 2004 [80]	1	11	9.1%	0.2% 41.3%	0.8
Fawcett et al. 2016 [85]	0	24	0.0%	0.0% 14.3%	7.8
Hori et al., 2012 [49]	0	12	0.0%	0.0% 26.5%	2.2
Inoue et al., 2010 [81]	0	10	0.0%	0.0% 30.9%	1.6
Lattanzi et al., 2002 [82]	0	14	0.0%	0.0% 23.2%	2.9
Sporn et al., 2000 [84]	0	20	0.0%	0.0% 16.8%	5.6
Tundo et al., 2022 [86]	0	79	0.0%	0.0% 4.6%	79.2
Random-effect			0.1%	0.0%	1.6%
<i>I</i> ² , %			0.0		
Bipolar					
Cassano et al., 2004 [80]	2	12	16.7%	2.1% 48.4%	3.1
Fawcett et al. 2016 [85]	0	18	0.0%	0.0% 18.5%	27.0
Hori et al., 2012 [49]	0	5	0.0%	0.0% 52.2%	2.9
Lattanzi et al., 2002 [82]	2	17	11.8%	1.5% 36.4%	6.0
Perugi et al., 2001 [83]	1	10	10.0%	0.3% 44.5%	4.0
Sporn et al., 2000 [84]	1	12	8.3%	0.2% 38.5%	5.7
Tundo et al., 2022 [86]	1	37	2.7%	0.1% 14.2%	51.2
Random-effect			3.5%	0.0%	7.2%
<i>I</i> ² , %			0.0		
Total					
Cassano et al., 2004 [80]	3	23	13.0%	2.8% 33.6%	1.0
Fawcett et al. 2016 [85]	0	42	0.0%	0.0% 8.4%	18.6
Hori et al., 2012 [49]	0	17	0.0%	0.0% 19.5%	3.3
Inoue et al., 2010 [81]	0	10	0.0%	0.0% 30.9%	1.3
Lattanzi et al., 2002 [82]	2	31	6.5%	0.8% 21.4%	2.6
Perugi et al., 2001 [83]	1	10	10.0%	0.3% 44.5%	0.6
Sporn et al., 2000 [84]	1	32	3.1%	0.1% 16.2%	5.3
Tundo et al., 2022 [86]	1	116	0.9%	0.0% 4.7%	67.5
Overall random-effects			1.1%	0.0%	2.5%
<i>I</i> ² , %			0.0		