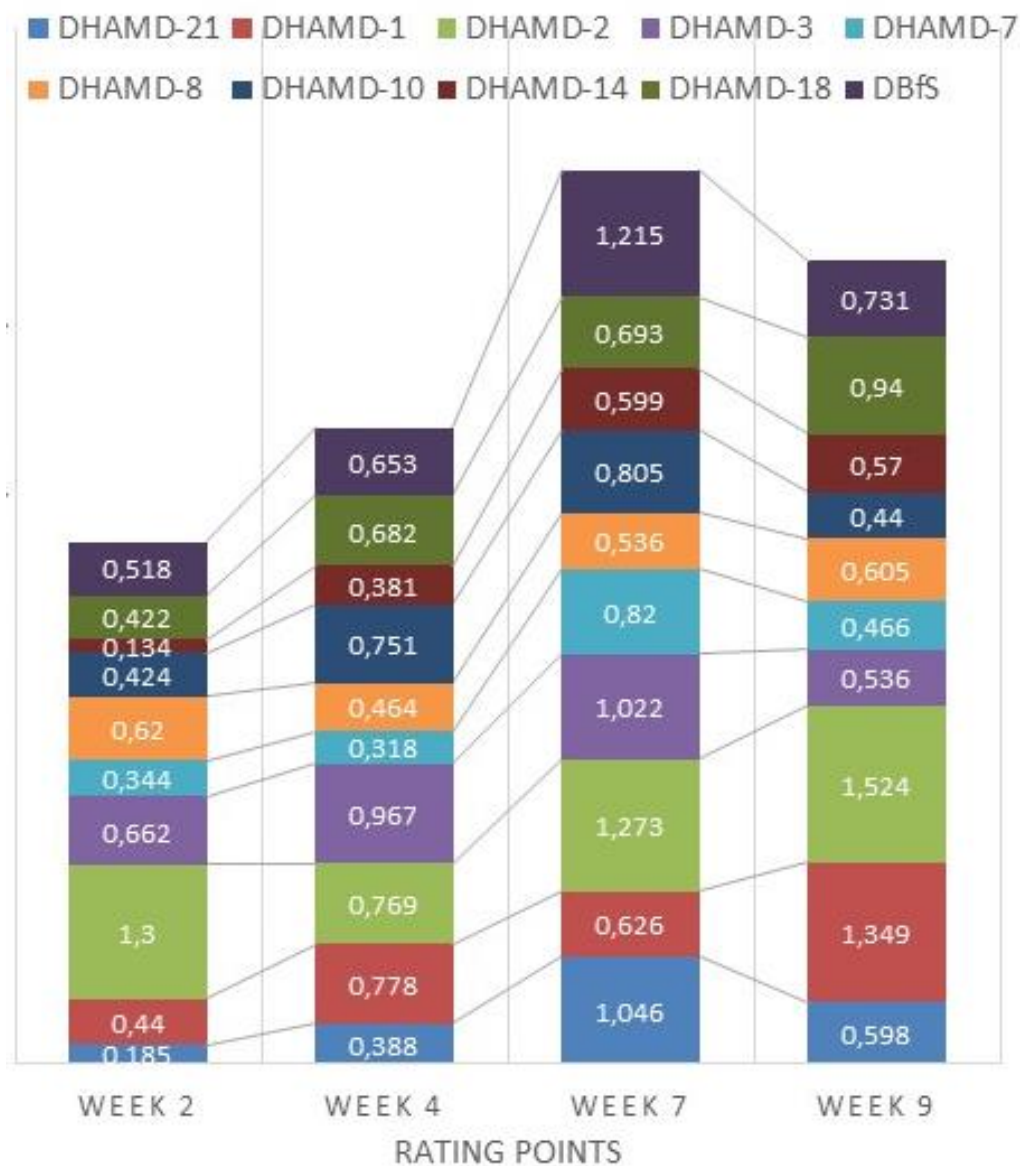


Additional file 4:

Figure S1. Cumulative effect size development of amantadine vs. placebo Groups (Cohen's d-values).



Cohen's d-values of clinical significance: values of 0.2, 0.5, and 0.8 were proposed to represent small, medium, and large effects, respectively.

D=Difference scores of

HAMD-21 Hamilton rating scale of depression; 21 items. Single items:

HAMD-1 "depressed mood"; **HAMD-2** "feeling of guilt"; **HAMD-3** "suicide"; **HAMD-7** "work and activities"; **HAMD-8** "retardation"; **HAMD-10** "anxiety (psychic)"; **HAMD-14** "sexual function"; **HAMD-18** "day-variations";

BfS "Befindlichkeitsskala nach von Zerssen (self-rating well-being scale)"