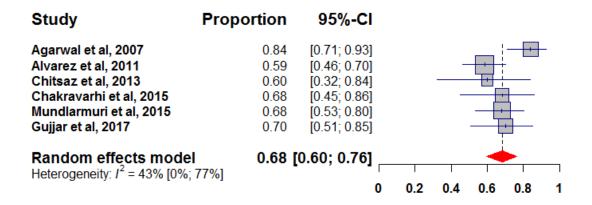
ONLY PROSPECTIVE STUDIES

Considering only prospective studies showed that the most effective non-BZD AED was PB with a SS of 0.79 (95% CI: 0.68-0.87) followed by VPA 0.71 (95% CI: 0.6-0.8), PHT 0.68 (95% CI: 0.6-0.76), LAC 0.66 (95% CI: 0.51-0.79), and the least effective non-BZD AED was LEV 0.64 (95% CI: 0.51-0.76).

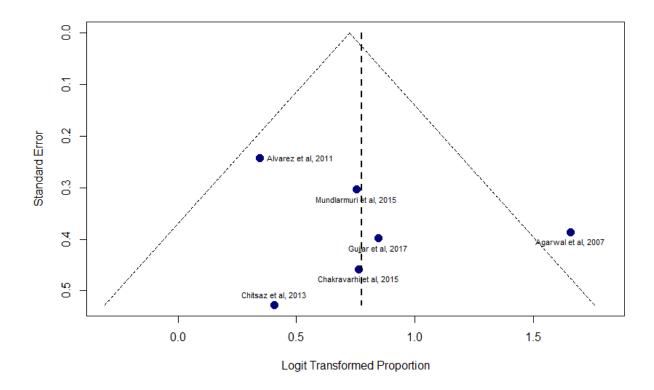
Study	Proportion	95%-CI	
PHT Agarwal et al, 2007 Alvarez et al, 2011 Chitsaz et al, 2013 Chakravarhi et al, 2015 Mundlarmuri et al, 2015 Gujjar et al, 2017 Random effects model Heterogeneity: I ² = 43% [0%	0.84 0.59 0.60 0.68 0.68 0.70 0.68	[0.71; 0.93] [0.46; 0.70] [0.32; 0.84] [0.45; 0.86] [0.53; 0.80] [0.51; 0.85] [0.60; 0.76]	——————————————————————————————————————
PB Malamiri et al, 2012 Su et al, 2016 Random effects model Heterogeneity: I ² = 0%	0.77 0.81 0.79	[0.58; 0.90] [0.65; 0.92] [0.68; 0.87]	
VPA Agarwal et al, 2007 Olsen et al, 2007 Alvarez et al, 2011 Chen et al, 2011 Malamiri et al, 2012 Chitsaz et al, 2013 Mundlarmuri et al, 2015 Su et al, 2016 Misra et al, 2017 Random effects model Heterogeneity: /² = 72% [44%]	0.88 0.73 0.75 0.50 0.90 0.73 0.68 0.44 0.70 0.71	[0.76; 0.95] [0.56; 0.86] [0.62; 0.85] [0.31; 0.69] [0.73; 0.98] [0.45; 0.92] [0.53; 0.80] [0.28; 0.62] [0.51; 0.84] [0.60; 0.80]	——————————————————————————————————————
LEV Eue et al, 2009 Alvarez et al, 2011 Atmaca et al, 2015 Chakravarhi et al, 2015 Mundlarmuri et al, 2015 Gujjar et al, 2017 Random effects model Heterogeneity: /² = 64% [13%]		[0.31; 0.66] [0.38; 0.65] [0.35; 0.93] [0.36; 0.79] [0.64; 0.88] [0.60; 0.95] [0.51; 0.76]	
LAC d'Orsi et al, 2016 Misra et al, 2017 Random effects model Heterogeneity: $I^2 = 0\%$	0.78 0.64 0.66	[0.40; 0.97] [0.45; 0.80] [0.51; 0.79]	0.2 0.4 0.6 0.8 1

ADJUSTMENT FOR PUBLICATION BIAS IN ONLY PROSPECTIVE STUDIES

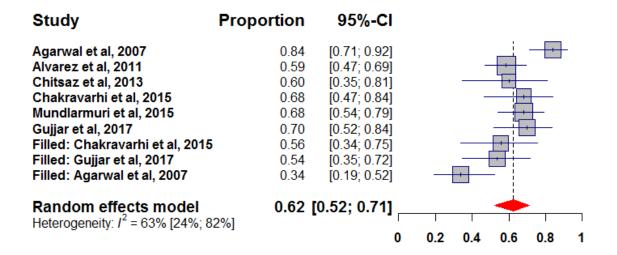
Phenytoin (PHT). The distribution of prospective studies evaluating the effectiveness of PHT showed evidence of publication bias and, therefore, adjustment for publication bias with Duval and Tweedie's trim and fill method resulted in a different estimation of SS than the original one.



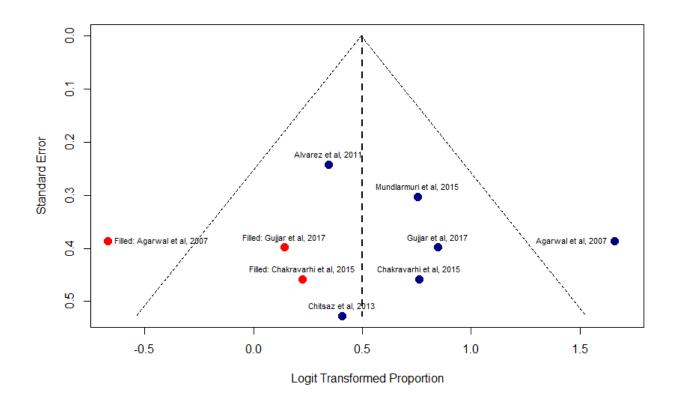
Original forest plot of the effectiveness of PHT. The probability of stopping seizures (SS) is 0.68 (95% CI: 0.6-0.76).



Funnel plot showing evidence of publication bias.

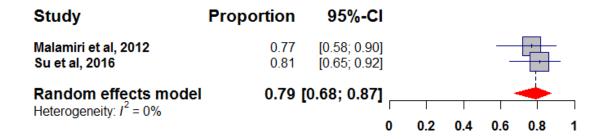


Forest plot of the effectiveness of PHT after applying Duval and Tweedie's trim and fill method. The probability of stopping seizures (SS) is 0.62 (95% CI: 0.52-0.71).

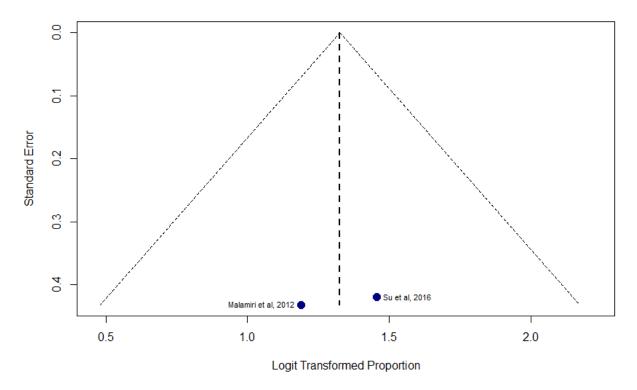


Funnel plot showing the imputed study after applying Duval and Tweedie's trim and fill method, adjusting for publication bias.

<u>Phenobarbital (PB).</u> The distribution of prospective studies evaluating the effectiveness of PB showed no evidence of publication bias and, correction for publication bias with Duval and Tweedie's trim and fill method is not possible with only two studies.

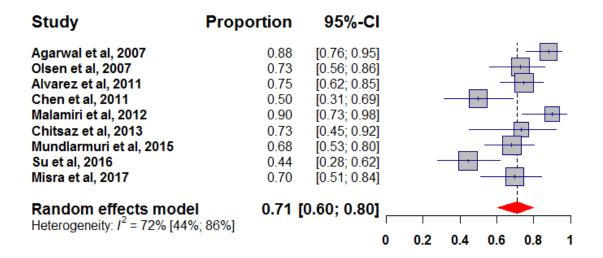


Original forest plot of the effectiveness of PB. The probability of stopping seizures (SS) is 0.79 (95% CI: 0.68-0.87).

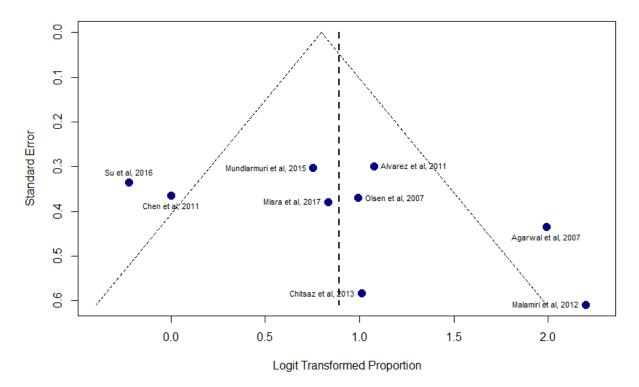


Funnel plot showing no evidence of publication bias.

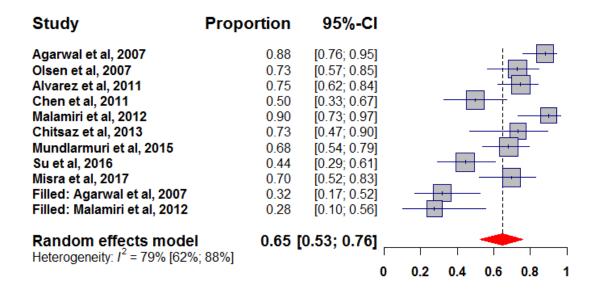
<u>Valproate (VPA)</u>. The distribution of prospective studies evaluating the effectiveness of VPA showed evidence of publication bias and, therefore, correction for publication bias with Duval and Tweedie's trim and fill method resulted in a different estimation of SS than the original one.



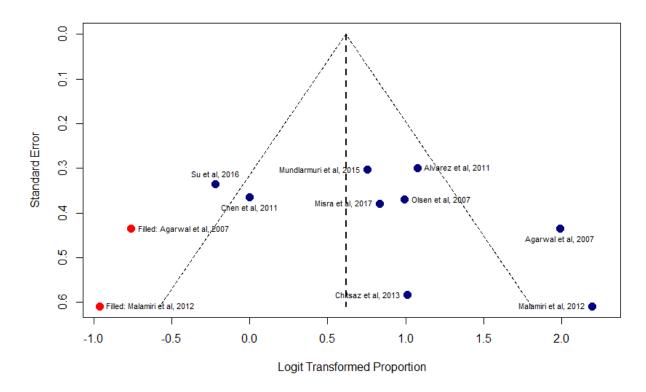
Original forest plot of the effectiveness of VPA. The probability of stopping seizures (SS) is 0.71 (95% CI: 0.6-0.8).



Funnel plot showing evidence of publication bias.

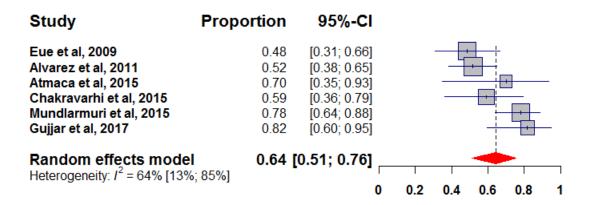


Forest plot of the effectiveness of VPA after applying Duval and Tweedie's trim and fill method. The probability of stopping seizures (SS) is 0.65 (95% CI: 0.53-0.76).

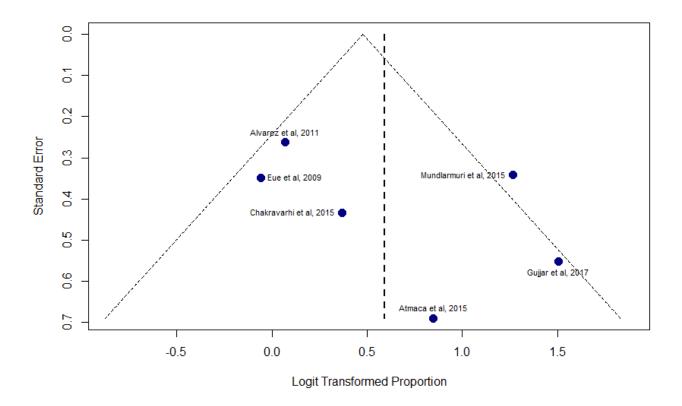


Funnel plot showing the imputed studies after applying Duval and Tweedie's trim and fill method, adjusting for publication bias.

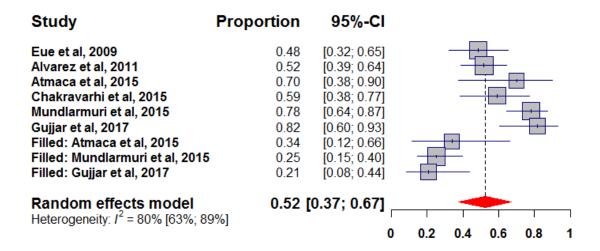
<u>Levetiracetam (LEV).</u> The distribution of prospective studies evaluating the effectiveness of LEV showed evidence of publication bias and, therefore, correction for publication bias with Duval and Tweedie's trim and fill method resulted in a different estimation of SS than the original one.



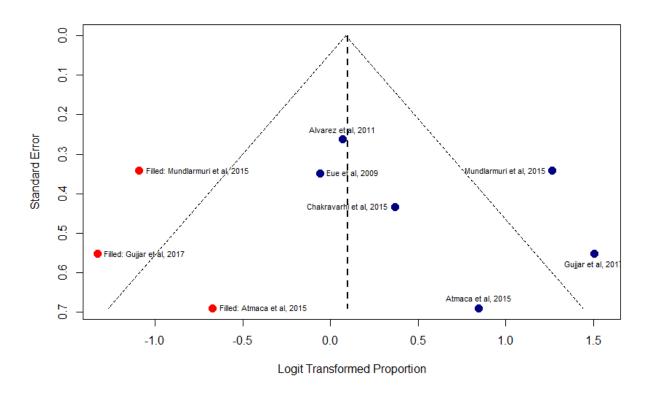
Original forest plot of the effectiveness of LEV. The probability of stopping seizures (SS) is 0.64 (95% CI: 0.51-0.76).



Funnel plot showing evidence of publication bias.

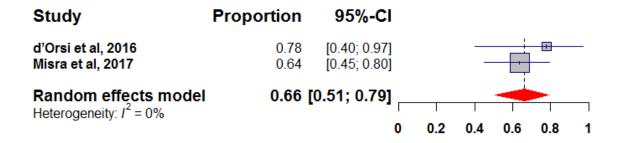


Forest plot of the effectiveness of LEV after applying Duval and Tweedie's trim and fill method. The probability of stopping seizures (SS) is 0.52 (95% CI: 0.37-0.67).

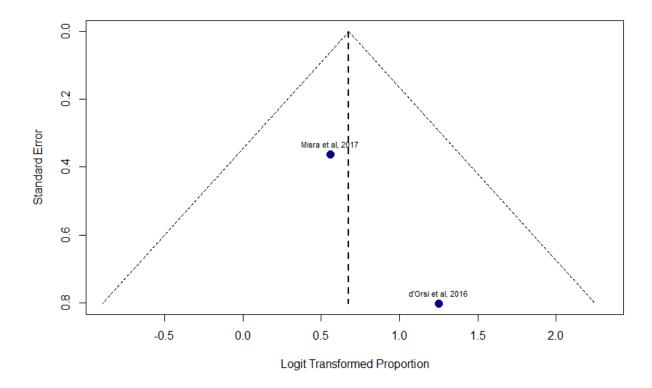


Funnel plot showing the imputed studies after applying Duval and Tweedie's trim and fill method, adjusting for publication bias.

<u>Lacosamide (LAC)</u>. The distribution of prospective studies evaluating the effectiveness of LAC showed no evidence of publication bias and, correction for publication bias with Duval and Tweedie's trim and fill method is not possible with only two studies.



Original forest plot of the effectiveness of LAC. The probability of stopping seizures (SS) is 0.66 (95% CI: 0.51-0.79).



Funnel plot showing no evidence of publication bias.

In summary, considering only prospective studies and after correcting for publication bias the most effective non-BZD AED was PB with a SS of 0.79 (95% CI: 0.68-0.87) followed by LAC 0.66 (95% CI: 0.51-0.79), VPA 0.65 (95% CI: 0.53-0.76), PHT 0.62 (95% CI: 0.52-0.71), and the least effective non-BZD AED was LEV 0.52 (95% CI: 0.37-0.67).