JAMA Clinical Evidence Synopsis Echinacea for Preventing and Treating the Common Cold

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CLINICAL QUESTION Are *Echinacea* products associated with a reduced incidence and a shorter duration of common colds compared with placebo?

BOTTOM LINE Individual prophylaxis trials show no association with prevention of the common cold, but exploratory meta-analysis suggests that *Echinacea* products may be associated with a small reduction in cold incidence. In treatment trials, there was no association of *Echinacea* products with a shorter duration of colds.

Preparations of the plant *Echinacea* are widely used in North America and Europe for prevention and treatment of the common cold.¹ This JAMA Clinical Evidence Synopsis summarizes the results of a Cochrane review² regarding the association of *Echinacea* products with prevention and treatment of colds.

Summary of Findings

Using the Cochrane "risk of bias" tool, 10 trials had a low risk of bias, 8 trials had a high risk of bias, and 6 trials had an unclear risk of bias. Because the trials studied chemically diverse extracts of *Echinacea*, we made an a priori decision not to pool data in a meta-analysis. Nevertheless, to provide a crude estimate of overall results and to generate hypotheses for future testing, we pooled study findings when there was no indication of statistical heterogeneity.

Ten trials investigated the prevention of colds. Nine of these (with 12 comparisons of an *Echinacea* product and placebo) reported the number of patients with at least 1 cold. None of the individual trials found a significant association of *Echinacea* with a reduction of colds (**Figure**). As findings were highly consistent ($l^2 = 0\%$, $\tau^2 = 0.00$), we pooled all 9 trials, regardless of the product used. Among these trials, an exploratory meta-analysis showed that prophylactic treatment with *Echinacea* products was

Evidence Profile

No. of randomized clinical trials: 24 (33 comparisons of *Echinacea* and placebo; 14 treatment trials, 9 prevention trials, 1 trial examining both prevention and treatment); 1 trial included only children.

Study years: Conducted, 1990-2010; published, 1992-2012

No. of patients: 2809 in treatment trials, 1822 in prevention trials

Men: 1640 (42%) Women: 2294 (58%) (sex not reported in 5 trials) Race/ethnicity: Unavailable

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Age, mean: 27.9 years (age not reported in 5 trials) Settings: Variety of inpatient and outpatient settings

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Countries: United States, Sweden, Germany, Canada, United Kingdom, Australia

Comparison: Echinacea vs placebo

Primary outcome: Cold duration in days for treatment trials; number of participants with at least 1 cold for prevention trials

associated with a lower rate of colds, 285 of 666 patients in the *Echinacea* group vs 279 of 501 patients in the placebo group (risk ratio, 0.83 [95% CI, 0.75-0.92]; *P* < .001).

Fifteen trials were treatment trials. Six of these reported on duration of cold symptoms. One with a high risk of bias and 1 trial with unclear risk of bias reported large, statistically significant associations of *Echinacea* with shorter duration of cold symptoms compared with placebo, whereas the 4 trials with low risk of bias did not find any associations of *Echinacea* with duration of cold symptoms. Due to the strong heterogeneity of findings and methodological quality, we did not include exploratory meta-analyses across trials or products on efficacy.

Most common adverse effects reported were headache, nausea, and a bad taste. In treatment trials, *Echinacea* products were associated with a higher rate of adverse effects (323 of 946 patients receiving *Echinacea* vs 281 of 863 patients receiving placebo; odds ratio [OR], 1.28 [95% CI, 1.02-1.60]; P = .03). In prevention trials, there was no association of *Echinacea* with adverse effects (102 of 868 patients receiving *Echinacea* vs 65 of 757 patients receiving placebo; OR, 1.49 [95% CI, 0.95-2.35]; P = .09). The treatment trial among children found that *Echinacea* was associated with a higher prevalence of rashes.

Discussion

In individual trials, there was no association of *Echinacea* products with prevention of the common cold. However, exploratory metaanalysis across products suggests that *Echinacea* products may be associated with a small reduction in cold incidence and that individual trials seem to be underpowered. The overall evidence for clinically relevant treatment effects is weak. We updated the search in July 2014, but could not identify more recent randomized clinical trials on *Echinacea* for preventing or treating the common cold.

Limitations

A variety of products prepared from different *Echinacea* species, plant parts, and in different forms have been compared with placebo in randomized clinical trials. These preparations contain different amounts of bioactive components and hence are not biochemically similar. Preparations based on the aerial parts of *Echinacea purpurea* were investigated most often. Furthermore, trial approaches and methods for cold assessment were highly

	Echinacea		Placebo				
Study by Echinacea product	No. With Outcome	Total No. of Participants	No. With Outcome	Total No. of Participants	Risk Ratio (95% CI)	Favors Favors Echinacea Placebo	
E purpurea herb							
Hall, 2007	7	18	7	14	0.78 (0.36-1.70)		
E purpurea herb pressed juice							
Grimm, 1999	35	54	40	54	0.88 (0.68-1.13)	— — —	
Sperber, 2004	14	24	18	22	0.71 (0.48-1.05)		
E purpurea dried plant extract							
O'Neill, 2008	22	28	27	30	0.87 (0.70-1.10)		
E purpurea root extract							
Zhang, 2003	25	54	33	57	0.80 (0.56-1.15)		
E purpurea root alcoholic extract							
Melchart, 1998	29	99	33	90	0.80 (0.53-1.20)	_	
E purpurea root and E angustofolia root							
Tiralongo, 2012	37	85	49	85	0.76 (0.56-1.02)		
4% phenolic extract of E purpurea and E angust	ifolia						
Turner, 2000	11	55	14	46	0.66 (0.33-1.30)	_	
E angustifolia root alcoholic extract							
Melchart, 1998	32	100	33	90	0.87 (0.59-1.29)	-	
E angustifolia root extract with CO ₂							
Turner, 2005	25	45	58	103	0.99 (0.72-1.35)		
E angustifolia root extract with 60% ethanol							
Turner, 2005	24	52	58	103	0.82 (0.58-1.15)	_	
E angustifolia root extract with 20% ethanol							
Turner, 2005	24	52	58	103	0.82 (0.58-1.15)		
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						0.2 1.0	÷,
						Risk Ratio (95% CI)	

Figure. Risk Ratios for Participants With at Least One Cold Episode In the Prevention Trials

Source: Data have been adapted with permission from Wiley.² The analysis includes 9 trials with 12 comparisons of an *Echinacea* product and placebo with

a total of 1167 patients. Melchart, 1998, compared 2 distinct *Echinacea* products and Turner, 2005, 3 distinct *Echinacea* products with a placebo group.

variable between studies. Therefore, trial results could not be combined for our primary outcome analyses. Publication bias cannot be ruled out with certainty.

Comparison of Findings With Current Practice Guidelines

Treatment of the common cold with *Echinacea* products is not specifically addressed in current guidelines and recommendations.^{3,4} Recommendations by the British National Institute for Health and Care Excellence (NICE)⁵ generally advise against the use of complementary and alternative medicine in cold treatment, without mentioning *Echinacea* products particularly.

Areas in Need for Future Study

Future studies should use well-defined preparations and a standard set of outcome measures in randomized clinical trials. Trials investigating the prevention of colds need large sample sizes as the potential effects of *Echinacea* products are likely to be small.

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Submissions: We encourage authors to submit papers for consideration as a JAMA Clinical Evidence Synopsis. Please contact Dr McDermott at mdm608@northwestern.edu.

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