

Table SII. Controlled studies of the use of wet wrap therapy (WWT) with emollients only or in combination with (diluted) topical corticosteroids (TCS).

Study	Objective	Design	Study population	AD definition and outcome measurement	Duration	Treatment	Outcome	Side-effects	Follow-up after end of treatment.	Comments
Jannohamed et al. 2014 (26)	WWT with diluted corticosteroids in comparison with emollients alone.	Randomized, controlled, double-blind study	39 children, 0.5–10 years	Severe AD (AD defined by H&R)	4 weeks: (24 h) whole body	TCS group: 1:3 MF 0.1% Face: 1:19 MF 0.1% or 1:1 pimecrolimus (as open therapy) Conv. group: emollients only beneath wet wraps Inner layer re-wetted every 2–3 h (not night)	4 dropouts in the emollient group Significant greater decrease in SCORAD in TCS group	Folliculitis: 10 in TCS group vs. 2 in emollient group Sec. infection in eczema: 2 in emollient group Wounding from facemask: 3 in emollient groups vs. 2 in TCS Temporary suppression of HPA axis: 3 in TCS group	No follow-up	
Schnopp et al. 2002 (30)	Comparing the effect of a steroid (MF) and a steroid-free (vehicle) preparation of WWT	Randomized, controlled, double-blinded, left–right study	20 children, 2–17 years	Exacerbated AD (AD defined by H&R)	5 days.	5 days with WWT TD with either MF 0.1% (not diluted) or emollient No treatment during night	The improvement in the MF group was significantly better than emollients only ($p < 0.01$) TEWL measurements improved in both study arms without reaching statistical significance	No information on side effects	No follow-up.	No systemic treatment with corticosteroids or AB, or TCS 7 days prior to start No information on re-wetting. Various AD definitions used No reports when and why WWT was stopped No long-term results
Devillers et al. 2002 (31)	To evaluate a standardized treatment, using wet-wrap dressings with diluted corticosteroids, in patients with refractory AD	Retrospective, controlled left–right study week 1 After week 1: Uncontrolled study	14 children (6 months–10 years) and 12 adults (18–61 years)	Refractory AD (AD defined by H&R, UK and Sampson criteria)	Average 17 weeks (11 to 41 weeks)	Week 1: Left–right study with WWT OD with 1:10 vs. 1:4 diluted 0.05% FP in adults and 1:20 vs. 1:10 diluted 0.05% FP in children. Whole body application Week 2: 4 days with OD 'diluted' FP on lesions only. Followed by 3 days of emollients only (no WWT) in objective SCORAD (uncontrolled). Week 3 ->: WWT of a maximum of 5 days weekly WWT: Minimum 12 h daily treatment. Rewetted every 2–3 h	Outcome left–right study (week 1): On day 4/5 a side difference was observed in 1 child and 4 adults in favor of the less diluted suppressed HPA-axis' and striae, 9 had FP. No comments whether it was infections. a significant difference At day 6–9 significant decrease in objective SCORAD No results from follow-up period (after week 1).	During follow-up period (after week 1): 5 had exacerbation of AD, 3 had temporary decrease in serum-cortisone, 1 had 'prolonged' suppressed HPA-axis' and striae, 9 had infections.	No follow-up	
Wolkenstorfer et al. 2000 (33)	To investigate the influence of different CS dilutions on the efficacy and HPA-suppression in children with severe refractory AD having WWT	Group 1: Uncontrolled study Group 2: Controlled left–right study Group 3: Controlled study	31 children, 5 months–13 years Group 1: 18 children Group 2: 5 children Group 3: 8 children	Severe AD (AD defined by Sampson and UK criteria) AD severity measured by objective SCORAD	2 weeks	Group 1: 1:2 diluted FP+WWT OD during 2 weeks Group 2: Left–right study of 1:10, 1:4 or 1:2 dilutions of FP OD+WWT for 1 week, 2 nd week OD 1:10 dilution only Group 3: 8 children, 2 children respectively with emollient only, 1:20, 1:10 or 1:4 dilution of FP on the whole body +WWT OD for 2 weeks WWT 24 h daily and rewetted every second hour	Group 1: After 2 weeks significant decrease in SCORAD in 17/18 children. Group 2: No significant difference between the different dilutions after week 1 (1:10 vs. 1:4, 1:4 vs. 1:2 etc.). Group 3: Week 1: Only minor improvements when emollient only, great improvements regardless FP dilutions, tendency to dose-response relationship.	Group 1: serum cortisol not significant decreased after 2 weeks, 3/18 had temporary lower serum cortisol levels after 2 weeks Group 2: No HPA-suppression Group 3: Increased HPA-suppression, measured by s-cortisol, but not urinary cortisol/creatinine ratio, with greater amount of applied TCS. Other side-effects reported: Folliculitis, upper resp. tract infections, herpes simplex infection, itching, diarrhea, abdominal pain, balanitis, furunculosis	No follow-up	

AD: atopic dermatitis, conv.: conventional treatment; AB: antibiotics; SCORAD: scoring atopic dermatitis severity score (51); H&R: Hanifin and Rajka AD criteria (58); HC: hydrocortisone; MF: mometasone furoate; FP: flucanone propionate; OD: once daily; TD: twice daily; HPA: hypothalamic-pituitary-adrenocortical; TEWL: transepidermal water loss; UK criteria: UK working party AD criteria (59); Sampson criteria: Sampson AD criteria (60).