

## NKR 44: calcineurin inhibitor vs. local steroid

### Review information

#### Authors

Danish Health Authority<sup>1</sup>

<sup>1</sup>[Empty affiliation]

Citation example: DHA. NKR 44: calcineurin inhibitor vs. local steroid. Cochrane Database of Systematic Reviews [Year], Issue [Issue].

### Characteristics of studies

#### Characteristics of included studies

##### Bauer 2012

<b>Methods</b>	<b>Study design:</b> Randomized controlled trial <b>Study grouping:</b> Parallel group <b>Open Label:</b> <b>Cluster RCT:</b>
<b>Participants</b>	<b>Baseline Characteristics</b> intervention <ul style="list-style-type: none"> <li>● <i>Age: mean(SD):</i> 34.8(12.47)</li> <li>● <i>Sex (%male):</i> 35</li> </ul> control <ul style="list-style-type: none"> <li>● <i>Age: mean(SD):</i> 30.87(8.08)</li> <li>● <i>Sex (%male):</i> 43.8</li> </ul> <p><b>Included criteria:</b> Patients suffering from moderate to very severe chronic relapsing atopic hand dermatitis (IGA 3) aged 18 years, who had responded to treatment with mometasone fuorate 0.1 % once daily over 1–3 weeks (IGA 2), were included after having obtained written informed consent.</p> <p><b>Excluded criteria:</b> Atopic dermatitis covering &gt;20 % of body surface area, use of phototherapy, systemic prednisolone or systemic immunosuppressive agents 4 weeks before screening visit, topical tar, pimecrolimus and tacrolimus 7 days prior to screening visit, hypersensitivity to ingredients of the study medication and/or the vehicle, women without adequate contraception or pregnancy or lactation. Further exclusion criteria were malignant disease within the last five years, concomitant skin disease or infections on the hands.</p> <p><b>Pretreatment:</b> There were no significant differences between the patients randomized to pimecrolimus (n = 20) and vehicle (n = 16) concerning age, sex, body weight, height and ethnicity (Table 2). No significant differences were seen in duration of treatment in patients randomized to pimecrolimus (37.3 ± 19.5 days) and vehicle (33.2 ± 21.7 days, p = 0.60) or use of study medication (pimecrolimus 82.9 ± 59.0 g; vehicle 79.0 ± 48.7 g, p = 0.97). Moreover, there were no significant differences in the numbers of discontinuations per group (pimecrolimus n = 10, 50.0 %; vehicle n = 9, 56.3 %; p = 0.749).</p>
<b>Interventions</b>	<b>Intervention Characteristics</b> intervention <ul style="list-style-type: none"> <li>● <i>treatment:</i> pimecrolimus 1 % cream twice a day after clinical response (IGA 2) to a 1–3 week treatment with mometasone fuorate 0.1 %.</li> <li>● <i>dose:</i> twice a day</li> <li>● <i>duration:</i> 8 weeks</li> </ul> control <ul style="list-style-type: none"> <li>● <i>treatment:</i> pimecrolimus Vehicle twice a day after clinical response (IGA 2) to a 1–3 week treatment with mometasone fuorate 0.1 %.</li> <li>● <i>dose:</i> twice a day</li> <li>● <i>duration:</i> 8 weeks</li> </ul>
<b>Outcomes</b>	<p><i>Sværhedsgrad af eksemet</i></p> <ul style="list-style-type: none"> <li>● <b>Outcome type:</b> Continuous Outcome</li> <li>● <b>Reporting:</b> Fully reported</li> <li>● <b>Scale:</b> HECSI</li> <li>● <b>Range:</b> 0–360</li> <li>● <b>Direction:</b> Lower is better</li> <li>● <b>Data value:</b> Change from baseline</li> </ul> <p><i>Livskvalitet</i></p> <ul style="list-style-type: none"> <li>● <b>Outcome type:</b> Continuous Outcome</li> <li>● <b>Reporting:</b> Fully reported</li> <li>● <b>Scale:</b> DLQI</li> <li>● <b>Range:</b> 0–30</li> <li>● <b>Direction:</b> Lower is better</li> <li>● <b>Data value:</b> Change from baseline</li> </ul> <p><i>Hudatrofi</i></p> <ul style="list-style-type: none"> <li>● <b>Outcome type:</b> Continuous Outcome</li> <li>● <b>Reporting:</b> Fully reported</li> <li>● <b>Scale:</b> TEWL</li> <li>● <b>Data value:</b> Endpoint</li> </ul>

<b>Identification</b>	<p><b>Sponsorship source:</b> AB: Although not directly related to this study AB served as a paid lecturer for Novartis and other companies. DL: none. UM: none. MM: Although not directly related to this study MM served as a paid lecturer for Novartis and other companies. MB is employee of Novartis Pharma GmbH, the manufacturer of pimecrolimus cream. TLD: Although not directly related to this study TLD served as a paid lecturer for Novartis and other companies.</p> <p><b>Country:</b> Germany</p> <p><b>Setting:</b> Outpatient clinics</p> <p><b>Comments:</b> -</p> <p><b>Authors name:</b> Andrea Bauer</p> <p><b>Institution:</b> Klinik und Poliklinik für Dermatologie, Medizinische Fakultät Carl Gustav Carus, Technische Universität Dresden</p> <p><b>Email:</b> andrea.bauer@uniklinikum-dresden.de</p> <p><b>Address:</b> Department of Dermatology University Hospital Carl Gustav Carus Technical University Dresden Fetscherstrasse 74D-01307 Dresden</p>
<b>Notes</b>	

## Risk of bias table

Bias	Authors' judgement	Support for judgement
Sequence Generation	Low risk	Quote: "The allocation sequence was generated by use of a permuted- block randomisation list in blocks' of 4 with equal allocation to pimecrolimus and vehicle."
Allocation concealment	Low risk	Quote: "Group [31]. Randomisation and blinding <b>Randomization was performed by allocation of the consecutive patients to the lowest available number from the randomisation list.</b> The allocation sequence was generated"
Blinding of participants and personnel	Low risk	Quote: "Verum and vehicle creams were prepared from the commercial product and blinded labelled for this study by Novartis Pharma GmbH. Patients and investigators were blinded to assignment of patients during the entire study period until the closing of the data bank."
Blinding of outcome assessors	Low risk	
Incomplete outcome data	Low risk	Judgement Comment: 1/16 dropout in the control group, no dropouts in intervention group (n=20)
Selective outcome reporting	Low risk	
Other sources of bias	Low risk	

## Hordinsky 2010

<b>Methods</b>	<p><b>Study design:</b> Randomized controlled trial</p> <p><b>Study grouping:</b> Parallel group</p> <p><b>Open Label:</b></p> <p><b>Cluster RCT:</b></p>
<b>Participants</b>	<p><b>Baseline Characteristics</b></p> <p>intervention</p> <ul style="list-style-type: none"> <li>● Age: mean(SD): 43.9 (14.4)</li> <li>● Sex (%male): 40</li> </ul> <p>control</p> <ul style="list-style-type: none"> <li>● Age: mean(SD): 44.1 (15.1)</li> <li>● Sex (%male): 35.5</li> </ul> <p><b>Included criteria:</b> Patients of either sex were eligible for inclusion in the study if they were at least 18 years of age and had a history of chronic hand dermatitis for at least 90 days. Patients were required to have mild-to-moderate hand dermatitis of the target hand, as defined by an Investigator's Global Assessment (IGA) score of 2 (mild) or 3 (moderate). When both hands were affected, the target hand was defined as the more severely affected.</p> <p><b>Excluded criteria:</b> The principal exclusion criteria were dyshidrotic dermatitis, psoriasis affecting the hand, flares of atopic dermatitis, and medication or concomitant conditions that could interfere with the conduct of the study or interpretation of the results. Immunocompromised patients, and those with a history of malignant disease, were also excluded</p> <p><b>Pretreatment:</b> There were no clinically relevant differences in baseline demographic characteristics or disease history between the pimecrolimus cream 1% and vehicle groups</p>
<b>Interventions</b>	<p><b>Intervention Characteristics</b></p> <p>intervention</p> <ul style="list-style-type: none"> <li>● treatment: pimecrolimus cream 1%</li> <li>● dose: twice daily</li> <li>● duration: up to 6 weeks</li> </ul> <p>control</p> <ul style="list-style-type: none"> <li>● treatment: vehicle cream</li> <li>● dose: twice daily</li> <li>● duration: up to 6 weeks</li> </ul>
<b>Outcomes</b>	<p><i>Sværhedsgrad af eksemet</i></p> <ul style="list-style-type: none"> <li>● Outcome type: Dichotomous Outcome</li> <li>● Reporting: Fully reported</li> <li>● Scale: number of patients with IGA score 0-1</li> <li>● Direction: Lower is better</li> </ul>

	<ul style="list-style-type: none"> <li>● <b>Data value:</b> Endpoint</li> </ul> <i>Livskvalitet</i> <ul style="list-style-type: none"> <li>● <b>Outcome type:</b> ContinuousOutcome</li> </ul> <i>Hudatrofi</i> <ul style="list-style-type: none"> <li>● <b>Outcome type:</b> ContinuousOutcome</li> </ul>
<b>Identification</b>	<p><b>Sponsorship source:</b> M. Hordinsky received research and/or consulting support from Novartis. A. Fleischer: advisory board – Allergan, Amgen, Astellas, Galderma, GSK, Ortho Dermatologics, Stiefel; consultant – Allergan, Astellas, Asubio, Combe, Galderma, Gerson Lehrman, Intendis, Kikaku America International, Merz, Novartis, Serentis, Taisho; investigator – 3M, Abbott, Amgen, Astellas, Asubio, Beiersdorf, Biogen, BioCryst, Dow, Centocor, Coria, Galderma, GSK, Genentech, Intendis, Mediciis, Novartis, Ortho Dermatologics, Pfizer, Serentis, Stiefel, Taisho; speaker bureau – Amgen, Astellas, Galderma, Intendis, Mediciis, Novartis, Stiefel, Upsher-Smith. J.K. Rivers has received research support and honoraria for speaking from Graceway Pharmaceuticals, GlaxoSmithKline, Merck, Solta Medical, Mediciis and Novartis. Y. Poulin has received research grants and/or honoraria from Abbott, Advitech, Amgen, Astellas-Biogen, Boehringer, Celgene, Centocor, Dermik, Galderma, GlaxoSmithKline, Isotechnika, LEO Pharma, Merck Serono, Novartis and Schering. D. Belsito is a consultant and/or received honoraria from Novartis, Astellas, Allderm, Basilea, Galderma, Sanofi-Aventis, Stiefel, Warner-Chilcott and Amgen/Wyeth. T. Hultsch was an employee of Novartis Pharma AG at the time of this study.</p> <p><b>Country:</b> Austria, Canada, Denmark, Hungary, Italy, Norway and the USA</p> <p><b>Setting:</b> -</p> <p><b>Comments:</b> -</p> <p><b>Authors name:</b> Maria Hordinsky</p> <p><b>Institution:</b> University of Minnesota School of Medicine, Minneapolis, Minn.</p> <p><b>Email:</b> hordi001 @ umn.edu</p> <p><b>Address:</b> Dr. M. Hordinsky University of Minnesota, Department of Dermatology 420 Delaware St, MMC 98 Minneapolis, MN 55455 (USA)</p>
<b>Notes</b>	

## Risk of bias table

Bias	Authors' judgement	Support for judgement
Sequence Generation	Low risk	Judgement Comment: Randomization was performed using a validated automated system and was stratified by baseline IGA score at each center.
Allocation concealment	Unclear risk	no comments
Blinding of participants and personnel	Low risk	Quote: "The study, which was randomized, double-blind and vehicle- controlled,"
Blinding of outcome assessors	Low risk	
Incomplete outcome data	Low risk	Judgement Comment: Withdrawal was equal between groups: 48/325 in the intervention group and 49/327 in the control group.
Selective outcome reporting	Low risk	
Other sources of bias	Low risk	

## Katsarou 2012

<b>Methods</b>	<p><b>Study design:</b> Randomized controlled trial</p> <p><b>Study grouping:</b> Parallel group</p> <p><b>Open Label:</b></p> <p><b>Cluster RCT:</b></p>
<b>Participants</b>	<p><b>Baseline Characteristics</b></p> <p>intervention</p> <ul style="list-style-type: none"> <li>● <i>Age: mean(SD):</i> -</li> <li>● <i>Sex (%male):</i> -</li> </ul> <p>control</p> <ul style="list-style-type: none"> <li>● <i>Age: mean(SD):</i> -</li> <li>● <i>Sex (%male):</i> -</li> </ul> <p><b>Included criteria:</b> age <math>\geq</math> 18 years. chronic hand eczema present at least 6 months before referral. pos patch test reactions relevant to patients records, absence of atopy documented with negative personal and family atopic history and total serum IgE concentration <math>\leq</math> 100 IU/ml, avoidance of topical and systemic corticosteroids and/or immunosuppressants for preceding two weeks before study onset.</p> <p><b>Excluded criteria:</b> -</p> <p><b>Pretreatment:</b> none</p>
<b>Interventions</b>	<p><b>Intervention Characteristics</b></p> <p>intervention</p> <ul style="list-style-type: none"> <li>● <i>treatment:</i> clobetasol propionate for 3 days. Tacrolimus ointment 0.1% x 2 daily for 30 days then x 1 daily for 2 months</li> <li>● <i>dose:</i> 2 times daily for 30 days then 1 time daily for 2 months</li> <li>● <i>duration:</i> 90th days</li> </ul> <p>control</p> <ul style="list-style-type: none"> <li>● <i>treatment:</i> Clobetasol propionate for 3 days. Mometasone furoate 0.1%</li> </ul>

	<ul style="list-style-type: none"> <li>● <b>dose:</b> 2 times daily for 1 week, then 1 time daily for 2 weeks, then 2 times a week up</li> <li>● <b>duration:</b> 90th days</li> </ul>
<b>Outcomes</b>	<p><i>Sværhedsgrad af eksemet</i></p> <ul style="list-style-type: none"> <li>● <b>Outcome type:</b> ContinuousOutcome</li> </ul> <p><i>Livskvalitet</i></p> <ul style="list-style-type: none"> <li>● <b>Outcome type:</b> ContinuousOutcome</li> </ul> <p><i>Hudatrofi</i></p> <ul style="list-style-type: none"> <li>● <b>Outcome type:</b> ContinuousOutcome</li> </ul>
<b>Identification</b>	<p><b>Sponsorship source:</b> None</p> <p><b>Country:</b> Greece</p> <p><b>Setting:</b> hospital</p> <p><b>Comments:</b> -</p> <p><b>Authors name:</b> Alexandra Katsarou</p> <p><b>Institution:</b> Dept. of dermatology, A. Sygros Hospital</p> <p><b>Email:</b> alkats,duoa@yahoo.gr</p> <p><b>Address:</b> Dept. of Dermatology, A. Sygros Hospital</p>
<b>Notes</b>	

## Risk of bias table

Bias	Authors' judgement	Support for judgement
Sequence Generation	Low risk	Judgement Comment: Random numbers in a computerized way
Allocation concealment	High risk	Judgement Comment: an investigator's assistant enrolled an assigned the treatment of the participants
Blinding of participants and personnel	Unclear risk	not reported
Blinding of outcome assessors	Unclear risk	not reported
Incomplete outcome data	Unclear risk	Judgement Comment: No baseline data available
Selective outcome reporting	High risk	
Other sources of bias	Unclear risk	not reported

## Krejci Manwaring 2008

<b>Methods</b>	<p><b>Study design:</b> Randomized controlled trial</p> <p><b>Study grouping:</b> Parallel group</p> <p><b>Open Label:</b></p> <p><b>Cluster RCT:</b></p>
<b>Participants</b>	<p><b>Baseline Characteristics</b></p> <p>intervention</p> <ul style="list-style-type: none"> <li>● <b>Age: mean(SD):</b> 46</li> <li>● <b>Sex (%male):</b> 19</li> </ul> <p>control</p> <ul style="list-style-type: none"> <li>● <b>Age: mean(SD):</b> 44</li> <li>● <b>Sex (%male):</b> 36</li> </ul> <p><b>Included criteria:</b> Diagnosed atopic/allergic/irritant dermatitis of the hands with a combined symptom severity score of 5-16 (score 0-16), adults aged 18 or over (female neg. pregnancy test), not using topical tacrolimus 28 days prior to the study, not using topical corticosteroids, nonsteroid immunosuppressants or light treatments 1 week prior to the study.</p> <p><b>Excluded criteria:</b> -</p> <p><b>Pretreatment:</b> no significant</p>
<b>Interventions</b>	<p><b>Intervention Characteristics</b></p> <p>intervention</p> <ul style="list-style-type: none"> <li>● <b>treatment:</b> First week prednisone 30mg/day combined with tacrolimus ointment 1%. Tacrolimus ointment 1% for the rest of the study.</li> <li>● <b>dose:</b> 2 times a day</li> <li>● <b>duration:</b> 12 weeks</li> </ul> <p>control</p> <ul style="list-style-type: none"> <li>● <b>treatment:</b> First week prednisone 30mg/day combined with vechile. Vechile ointment for the rest of the study.</li> <li>● <b>dose:</b> 2 times a day</li> <li>● <b>duration:</b> 12 weeks</li> </ul>
<b>Outcomes</b>	<p><i>Sværhedsgrad af eksemet</i></p> <ul style="list-style-type: none"> <li>● <b>Outcome type:</b> DichotomousOutcome</li> <li>● <b>Unit of measure:</b> %change in IGA</li> <li>● <b>Direction:</b> Higher is better</li> <li>● <b>Data value:</b> Change from baseline</li> </ul> <p><i>Livskvalitet</i></p> <ul style="list-style-type: none"> <li>● <b>Outcome type:</b> ContinuousOutcome</li> </ul> <p><i>Hudatrofi</i></p> <ul style="list-style-type: none"> <li>● <b>Outcome type:</b> ContinuousOutcome</li> </ul>

	<i>Flare</i> <ul style="list-style-type: none"> <li>● <b>Outcome type:</b> DichotomousOutcome</li> <li>● <b>Reporting:</b> Fully reported</li> <li>● <b>Direction:</b> Lower is better</li> <li>● <b>Data value:</b> Endpoint</li> </ul>
<b>Identification</b>	<b>Sponsorship source:</b> Astekka Pharma inc, Fujisawa Healthcare Inc, Connetics, Galderma, Amgen, Biogen, Genentich, novartis Pharmaceuticals Corp., Bioform Medical. <b>Country:</b> USA <b>Setting:</b> - <b>Comments:</b> - <b>Authors name:</b> Jennifer Krejci-Manwaring <b>Institution:</b> Dept. of Dermatology, Wake Forest university School of Medicine, Winston.Salem, NC <b>Email:</b> sfeldman@wfubmc.edu <b>Address:</b> Dept. of Dermatology, Wake Forrest University School of Medicine Winston-Salem, NC
<b>Notes</b>	

## Risk of bias table

Bias	Authors' judgement	Support for judgement
Sequence Generation	Low risk	Judgement Comment: The randomization code list, correlating the kit numbers with the content of each kit, was kept on file at the Fujisawa Healthcare Medical Information Dept. until the time of analysis.
Allocation concealment	Low risk	Judgement Comment: Both the investigators and the patients were blinded to treatment assignment
Blinding of participants and personnel	Low risk	Judgement Comment: Each patient was assigned a subject number with corresponding prelabeled tubes of study medication. The vehicle and tacrolimus ointments were packaged in identical containers labeled with the subject number, so neither the subject, coordinator, nor the investigator knew which treatment the patient received.
Blinding of outcome assessors	Low risk	Judgement Comment: Each patient was assigned a subject number with corresponding prelabeled tubes of study medication. The vehicle and tacrolimus ointments were packaged in identical containers labeled with the subject number, so neither the subject, coordinator, nor the investigator knew which treatment the patient received.
Incomplete outcome data	Low risk	Judgement Comment: 5/22 in the intervention group and 1/11 in the control were lost to follow up.
Selective outcome reporting	Low risk	
Other sources of bias	Low risk	

## Schnopp 2002

<b>Methods</b>	<b>Study design:</b> Randomized controlled trial <b>Study grouping:</b> Parallel group <b>Open Label:</b> <b>Cluster RCT:</b>
<b>Participants</b>	<b>Baseline Characteristics</b> Tacrolimus (FK506 0.1%) <ul style="list-style-type: none"> <li>● <i>Age (mean):</i> 43</li> <li>● <i>Palm affected:</i> 12</li> <li>● <i>Female (%):</i> 15 (94)</li> </ul> Mometasone furoate 0.1% <ul style="list-style-type: none"> <li>● <i>Age (mean):</i> 43</li> <li>● <i>Palm affected:</i> 12</li> <li>● <i>Female (%):</i> 15 (94)</li> </ul> <b>Included criteria:</b> moderate to severe chronic, relapsing dyshidrotic eczema <b>Excluded criteria:</b> use of topical glucocorticoids or any systemic treatment with possible influence on the course of the disease (eg. steroids, antibiotics, antihistamines, nonsteroidal anti-inflammatory medications). <b>Pretreatment:</b> Each person was his/her own control (right vs left hand)
<b>Interventions</b>	<b>Intervention Characteristics</b> Tacrolimus (FK506 0.1%) <ul style="list-style-type: none"> <li>● <i>Description:</i> tacrolimus</li> <li>● <i>Dose:</i> twice daily</li> <li>● <i>Duration:</i> 4 weeks</li> </ul> Mometasone furoate 0.1% <ul style="list-style-type: none"> <li>● <i>Description:</i> Mometasone ointment</li> <li>● <i>Dose:</i> twice daily</li> <li>● <i>Duration:</i> 4 weeks</li> </ul>
<b>Outcomes</b>	<i>Sværhedsgrad af eksemet</i> <ul style="list-style-type: none"> <li>● <b>Outcome type:</b> ContinuousOutcome</li> <li>● <b>Reporting:</b> Fully reported</li> <li>● <b>Scale:</b> DASI</li> <li>● <b>Direction:</b> Lower is better</li> </ul>

	● Data value: Endpoint
<b>Identification</b>	<p><b>Sponsorship source:</b> Funding sources: none</p> <p><b>Country:</b> Germany</p> <p><b>Setting:</b> Hospital</p> <p><b>Comments:</b></p> <p><b>Authors name:</b> Christina Schnopp</p> <p><b>Institution:</b> Dept. of Dermatology and Allergy Biederstein, Technical University Munich</p> <p><b>Email:</b> nina.schnopp@lrz.tu-muenchen.de</p> <p><b>Address:</b> Dept. of Dermatology and Allergy Biederstein, Technical University Munich, Biedersteiner Str. 29, D-80802 Munchen, Germany</p>
<b>Notes</b>	<p>Louise Klokke Madsen on 06/04/2016 02:52</p> <p><b>Outcomes</b></p> <p>mean værdier aflæst fra figur 3, SD taget fra teksten (2 ugers målingen). 16 deltagere randomiseret, antageligt 8 i hver gruppe (ikke opgivet)</p> <p>Nkr44 HåNdeksem on 14/04/2016 21:14</p> <p><b>Outcomes</b></p> <p>SD taget fra uge 2.tæller hænder personer. n=16 hænder i hver gruppe</p>

## Risk of bias table

Bias	Authors' judgement	Support for judgement
Sequence Generation	Unclear risk	not reported
Allocation concealment	Unclear risk	not reported
Blinding of participants and personnel	High risk	Judgement Comment: not reported (but it says: 'observer-blinded', not double blind.
Blinding of outcome assessors	Low risk	Judgement Comment: 'observer-blinded'
Incomplete outcome data	Low risk	Judgement Comment: no dropouts during treatment period (4 dropped out during the washout period)
Selective outcome reporting	Low risk	
Other sources of bias	Low risk	

## Footnotes

## Characteristics of excluded studies

**Abramovits 2010**

Reason for exclusion	Wrong study design
----------------------	--------------------

**Bissonnette 2010**

Reason for exclusion	Wrong study design
----------------------	--------------------

**Day 2008**

Reason for exclusion	Wrong study design
----------------------	--------------------

**Diepgen 2007**

Reason for exclusion	Wrong study design
----------------------	--------------------

**Gelmetti 2010**

Reason for exclusion	Wrong study design
----------------------	--------------------

**Luger 2007**

Reason for exclusion	Wrong study design
----------------------	--------------------

**Madan 2007**

Reason for exclusion	Wrong study design
----------------------	--------------------

**Mensing 2008**

Reason for exclusion	Wrong study design
----------------------	--------------------

**Pacor 2006**

Reason for exclusion	Wrong comparator
----------------------	------------------

## Ricci 2007

Reason for exclusion	Wrong study design
----------------------	--------------------

## Schliemann 2008

Reason for exclusion	Wrong study design
----------------------	--------------------

## Van 2012

Reason for exclusion	Wrong intervention
----------------------	--------------------

## Wollina 2006

Reason for exclusion	Wrong study design
----------------------	--------------------

## Footnotes

## Characteristics of studies awaiting classification

## Footnotes

## Characteristics of ongoing studies

## Footnotes

## Summary of findings tables

## Additional tables

## References to studies

## Included studies

### Bauer 2012

Bauer,Andrea; Lange,Nora; Mattered,Uwe; Meurer,Michael; Braeutigam,Matthias; Diepgen,Thomas L.. Efficacy of pimecrolimus 1% cream in the long term management of atopic hand dermatitis. A double-blind RCT. Journal der Deutschen Dermatologischen Gesellschaft = Journal of the German Society of Dermatology : JDDG 2012;10(6):426-33. [DOI: ]

### Hordinsky 2010

Hordinsky,Maria; Fleischer,Alan; Rivers,Jason K.; Poulin,Yves; Belsito,Donald; Hultsch,Thomas. Efficacy and safety of pimecrolimus cream 1% in mild-to-moderate chronic hand dermatitis: a randomized, double-blind trial. Dermatology (Basel, Switzerland) 2010;221(1):71-7. [DOI: ]

### Katsarou 2012

Katsarou,Alexandra; Makris,Manolis; Papagiannaki,Konstantina; Lagogianni,Eirini; Tagka,Anna; Kalogeromitros,Dimitrios. Tacrolimus 0.1% vs mometasone furoate topical treatment in allergic contact hand eczema: a prospective randomized clinical study. European journal of dermatology : EJD 2012;22(2):192-6. [DOI: ]

### Krejci Manwaring 2008

Krejci-Manwaring,Jennifer; McCarty,Martha Ann; Camacho,Fabian; Manuel,Janeen; Hartle,Jennifer; Fleischer,Alan,Jr; Feldman,Steven R.. Topical tacrolimus 0.1% improves symptoms of hand dermatitis in patients treated with a prednisone taper. Journal of drugs in dermatology : JDD 2008;7(7):643-6. [DOI: ]

### Schnopp 2002

Schnopp,C.; Remling,R.; Mohrenschlager,M.; Weigl,L.; Ring,J.; Abeck,D.. Topical tacrolimus (FK506) and mometasone furoate in treatment of dyshidrotic palmar eczema: a randomized, observer-blinded trial. Journal of the American Academy of Dermatology 2002;46(1):73-77. [DOI: S0190962202206787 [pii]]

## Excluded studies

### Abramovits 2010

Abramovits W.; Granowski P.. Innovative management of severe hand dermatitis. Dermatologic clinics 2010;28(3):453-465. [DOI: ]

### Bissonnette 2010

Bissonnette R.; Diepgen T.L.; Elsner P.; English J.; Graham-Brown R.; Homey B.; Luger T.; Lynde C.; Maars J.; Maibach H.I.. Redefining treatment options in chronic hand eczema (CHE). Journal of the European Academy of Dermatology and Venereology 2010;24(Web Page):1-20. [DOI: ]

### Day 2008

Day,Isaiah; Lin,Andrew N.. Use of pimecrolimus cream in disorders other than atopic dermatitis. Journal of cutaneous medicine and surgery 2008;12(1):17-26. [DOI: ]

### Diepgen 2007

Diepgen T.L.; Agner T.; Aberer W.; Berth-Jones J.; Cambazard F.; Elsner P.; McFadden J.; Coenraads P.J.. Management of chronic hand eczema. Contact dermatitis 2007;57(4):203-210. [DOI: ]

**Gelmetti 2010**

Gelmetti,Carlo; Frasin,Adina; Restano,Lucia. Innovative therapeutics in pediatric dermatology. *Dermatologic clinics* 2010;28(3):619-29. [DOI: ]

**Luger 2007**

Luger,Thomas; Paul,Carle. Potential new indications of topical calcineurin inhibitors. *Dermatology (Basel, Switzerland)* 2007;215 Suppl 1(Journal Article):45-54. [DOI: ]

**Madan 2007**

Madan,V.; Griffiths,C. E. M.. Systemic ciclosporin and tacrolimus in dermatology. *Dermatologic therapy* 2007;20(4):239-50. [DOI: ]

**Mensing 2008**

Mensing,C. O.; Mensing,C. H.; Mensing,H.. Treatment with pimecrolimus cream 1% clears irritant dermatitis of the periocular region, face and neck. *International journal of dermatology* 2008;47(9):960-4. [DOI: ]

**Pacor 2006**

Pacor,Maria L.; Di Lorenzo,Gabriele; Martinelli,Nicola; Mansueto,Pasquale; Friso,Simonetta; Pellitteri,Maria Esposito; Di Fede,Gaetana; Rini,Giovambattista; Corrocher,Roberto. Tacrolimus ointment in nickel sulphate-induced steroid-resistant allergic contact dermatitis. *Allergy and Asthma Proceedings : The Official Journal of Regional and State Allergy Societies* 2006;27(6):527-31. [DOI: ]

**Ricci 2007**

Ricci G.; Dondi A.; Patrizi A.. Role of topical calcineurin inhibitors on atopic dermatitis of children. *Current medicinal chemistry* 2007;14(14):1579-1591. [DOI: ]

**Schliemann 2008**

Schliemann,Sibylle; Kelterer,Daniela; Bauer,Andrea; John,Swen M.; Skudlik,Christoph; Schindera,Ingo; Wehrmann,Wolfgang; Elsner,Peter. Tacrolimus ointment in the treatment of occupationally induced chronic hand dermatitis. *Contact dermatitis* 2008;58(5):299-306. [DOI: ]

**Van 2012**

Van,Gils R.; Boot C.R.L.; Knol D.L.; Rustemeyer T.; Van,Mechelen W.; Van Der,Valk P.; Anema J.R.. The effectiveness of integrated care for patients with hand eczema: Results of a randomized, controlled trial. *Contact dermatitis* 2012;66(4):197-204. [DOI: ]

**Wollina 2006**

Wollina U.; Hansel G.; Koch A.; Abdel-Naser M.B.. Topical pimecrolimus for skin disease other than atopic dermatitis. Expert opinion on pharmacotherapy 2006;7(14):1967-1975. [DOI: ]

**Studies awaiting classification****Ongoing studies****Other references****Additional references****Other published versions of this review****Classification pending references****Data and analyses****1 Pimecrolimus vs control**

Outcome or Subgroup	Studies	Participants	Statistical Method	Effect Estimate
1.1 Sværhedsgrad af eksem (ptt. med moderat/svær eksem), EOT	1	36	Mean Difference (IV, Random, 95% CI)	-10.75 [-51.98, 30.48]
1.1.1 HECSI (lower=better)	1	36	Mean Difference (IV, Random, 95% CI)	-10.75 [-51.98, 30.48]
1.2 Sværhedsgrad af eksem (ptt. med mild/moderat eksem), EOT	1		Risk Ratio (IV, Random, 95% CI)	Subtotals only
1.2.1 patients with IGA 0-1	1	652	Risk Ratio (IV, Random, 95% CI)	1.28 [0.99, 1.66]
1.3 Livskvalitet (ptt. med moderat/svær eksem), EOT	1	36	Mean Difference (IV, Random, 95% CI)	-1.07 [-4.92, 2.78]
1.3.1 DLQI (lower=better)	1	36	Mean Difference (IV, Random, 95% CI)	-1.07 [-4.92, 2.78]
1.4 Hudatrofi (ptt. med moderat/svær eksem), EOT	1	36	Mean Difference (IV, Random, 95% CI)	1.45 [-9.54, 12.44]
1.4.1 TEWL (lower=better)	1	36	Mean Difference (IV, Random, 95% CI)	1.45 [-9.54, 12.44]
1.5 Cancer	0	0	Odds Ratio (M-H, Fixed, 95% CI)	Not estimable
1.6 Flare	0		Risk Ratio (IV, Fixed, 95% CI)	No totals

**2 Tacrolimus vs control**

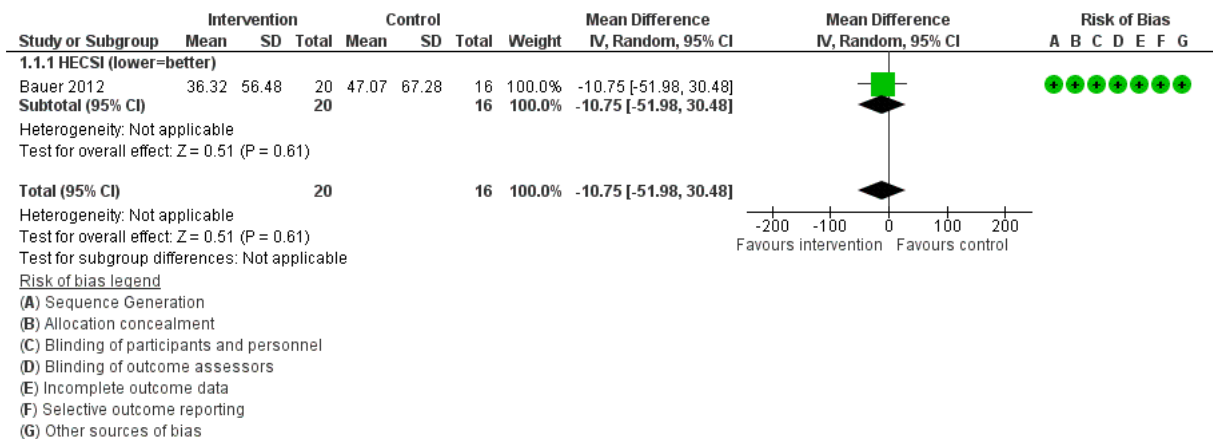
Outcome or Subgroup	Studies	Participants	Statistical Method	Effect Estimate
---------------------	---------	--------------	--------------------	-----------------



2.1 Sværhedsgrad af eksemet (ptt. med moderat/svær eksem), længste follow up (4 uger)	1	32	Mean Difference (IV, Fixed, 95% CI)	1.50 [-3.34, 6.34]
2.1.1 DASI (lower=better)	1	32	Mean Difference (IV, Fixed, 95% CI)	1.50 [-3.34, 6.34]
2.2 Sværhedsgrad af eksemet (ptt. med moderat/svær eksem), EOT	1	20	Mean Difference (IV, Random, 95% CI)	-0.30 [-2.21, 1.61]
2.2.2 % change in IGA (higher=better)	1	20	Mean Difference (IV, Random, 95% CI)	-0.30 [-2.21, 1.61]
2.3 Cancer	0	0	Odds Ratio (M-H, Fixed, 95% CI)	Not estimable
2.4 Flare	0		Risk Ratio (IV, Fixed, 95% CI)	No totals

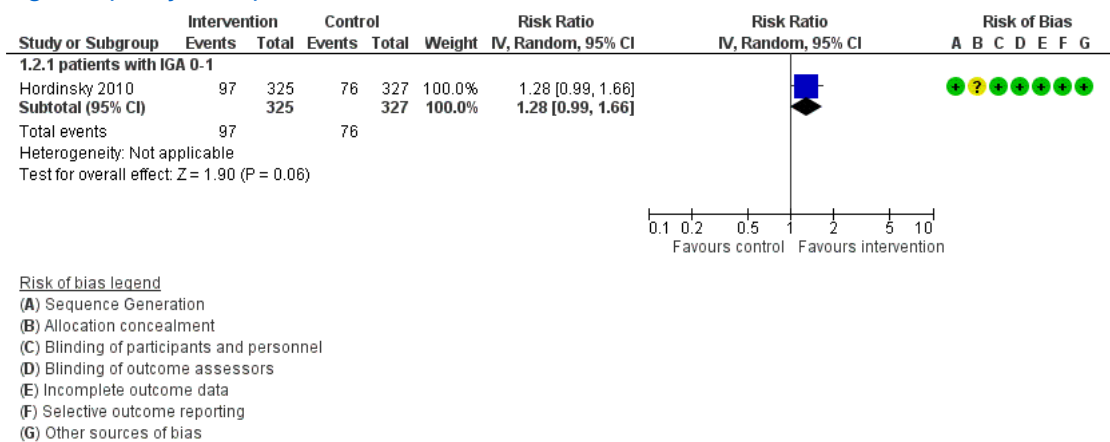
## Figures

### Figure 1 (Analysis 1.1)



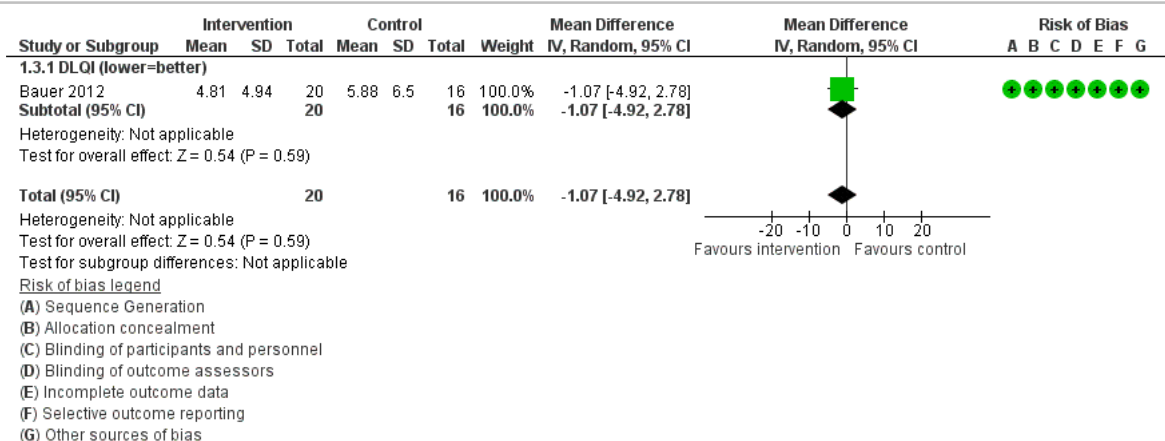
Forest plot of comparison: 1 Pimecrolimus vs control, outcome: 1.1 Sværhedsgrad af eksemet (ptt. med moderat/svær eksem), EOT.

### Figure 2 (Analysis 1.2)



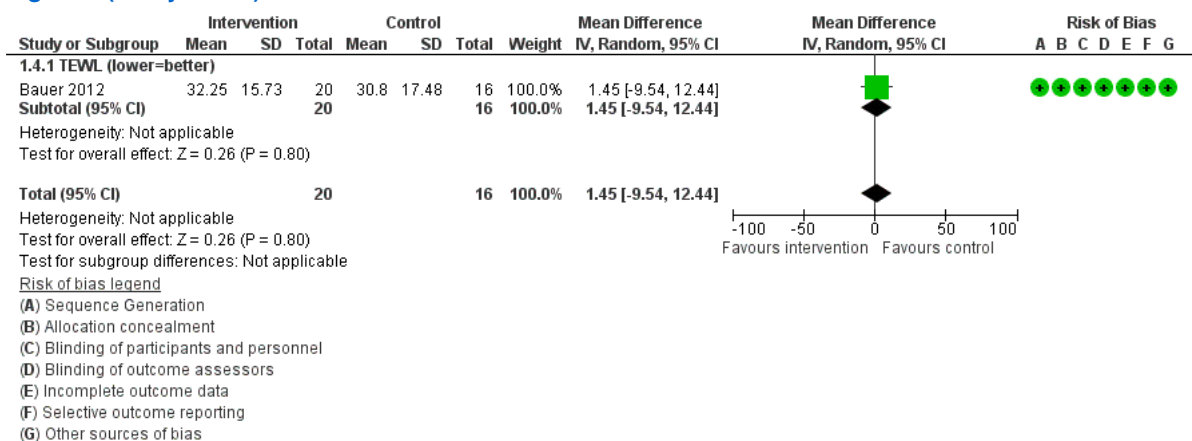
Forest plot of comparison: 1 Pimecrolimus vs control, outcome: 1.2 Sværhedsgrad af eksemet (ptt. med mild/moderat eksem), EOT.

### Figure 3 (Analysis 1.3)



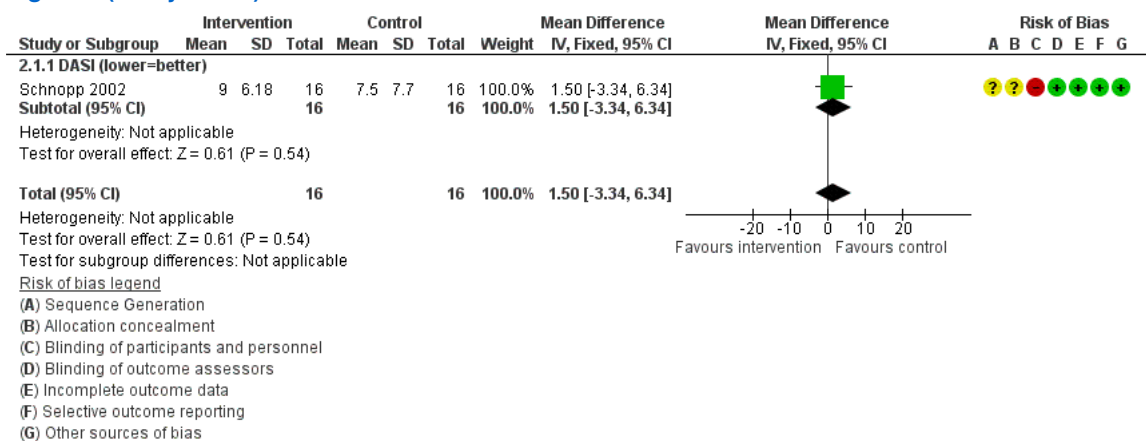
Forest plot of comparison: 1 Pimecrolimus vs control, outcome: 1.3 Livskvalitet (ptt. med moderat/svær eksem), EOT.

Figure 4 (Analysis 1.4)



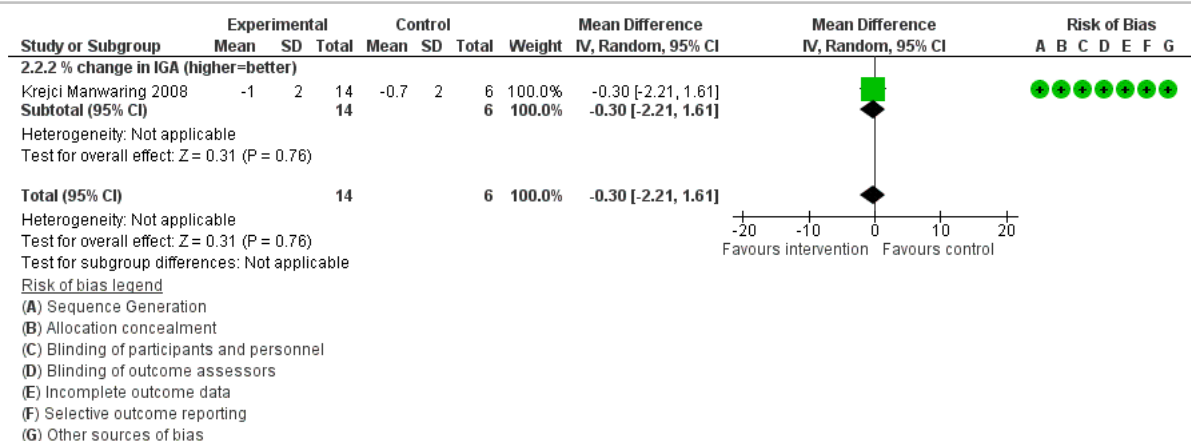
Forest plot of comparison: 1 Pimecrolimus vs control, outcome: 1.4 Hudatrofi (ptt. med moderat/svær eksem), EOT.

Figure 5 (Analysis 2.1)



Forest plot of comparison: 2 Tacrolimus vs control, outcome: 2.1 Sværhedsgrad af eksemet (ptt. med moderat/svær eksem), længste follow up (4 uger).

Figure 6 (Analysis 2.2)



Forest plot of comparison: 2 Tacrolimus vs control, outcome: 2.2 Sværhedsgrad af eksemet (ptt. med moderat/svær eksem), EOT.