



# Virgin Coconut Oil as a Topical Intervention for Uremic-Associated Xerosis and CKD-Associated Pruritus: A Systematic Review

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## Abstract

**Context:** Chronic kidney disease-associated pruritus (CKD-aP) and uremic xerosis affect more than half of patients undergoing hemodialysis (HD) and markedly impair their quality of life. Conventional moisturizers often provide insufficient relief. Virgin coconut oil (VCO), possessing emollient, antimicrobial, and anti-inflammatory properties, has been proposed as a low-cost topical therapy. However, its effectiveness in CKD-specific dermatoses has not been comprehensively evaluated.

**Objectives:** The objective of this study is to systematically review the evidence on the efficacy, safety, and feasibility of topical VCO for managing uremic xerosis and CKD-aP in adults.

**Methods:** A systematic search of PubMed, Scopus, Embase, Google Scholar, and SciSpace (from inception to November 2025) identified studies assessing topical VCO for xerosis or pruritus in CKD. Two reviewers independently performed screening, data extraction, and quality appraisal using the Joanna Briggs Institute (JBI) checklist. Due to heterogeneity in study designs, comparators, and outcomes, findings were synthesized narratively.

**Results:** Eighty-three records were identified, and 12 studies met the inclusion criteria: One randomized controlled trial (RCT), four quasi-experimental controlled studies, and seven uncontrolled pre-post studies (total  $\approx$  350 participants). The RCT ( $n = 60$ ) showed significantly greater improvement in overall dry skin score (ODSS) with VCO compared to mineral oil after one week of twice-daily application ( $P < 0.05$ ). Quasi-experimental studies reported consistent within-group improvements in skin moisture and pruritus scores [5-D Itch Scale, Visual Analog Scale (VAS)], though results versus standard lotions were variable. Uncontrolled studies uniformly demonstrated reductions in xerosis and pruritus but were limited by small samples and lack of control groups. No serious adverse events were reported, although adverse event monitoring was inconsistently described. The overall certainty of the evidence was low to moderate due to methodological limitations, short follow-up, and heterogeneous outcome measures.

**Conclusions:** Topical VCO appears safe and potentially effective for short-term relief of uremic xerosis and CKD-aP, particularly in low-resource settings. Robust, long-term randomized trials using standardized outcomes are needed to confirm comparative effectiveness and define optimal application protocols.

**Keywords:** Virgin Coconut Oil, Chronic Kidney Disease, Uremic Xerosis, Pruritus, Hemodialysis, Topical Therapy, Systematic Review

## 1. Context

Chronic kidney disease (CKD) affects more than 850 million people globally and is associated with multiple dermatological complications, particularly xerosis and pruritus, which affect 40 - 90% of patients undergoing hemodialysis (HD) (1-4). These symptoms substantially

impair quality of life, disturb sleep, increase psychological distress, and have been linked to higher mortality (5, 6).

The pathophysiology of uremic xerosis and chronic kidney disease-associated pruritus (CKD-aP) is multifactorial, involving impaired skin barrier function,

reduced sebaceous and sweat gland activity, accumulation of uremic toxins and inflammatory mediators, opioid receptor dysregulation, mineral-bone disorder, and xerosis-induced stimulation of cutaneous nerve fibers (7, 8).

Management typically includes optimization of dialysis, correction of metabolic abnormalities, systemic agents such as antihistamines and gabapentinoids, and topical emollients (9-11). However, conventional moisturizers often provide limited relief, and systemic therapies may cause adverse effects in patients with CKD.

### 1.1. Virgin Coconut Oil: Properties and Potential Mechanisms

Virgin coconut oil (VCO), obtained through cold-pressing fresh coconut meat, contains approximately 90% saturated fatty acids – primarily medium-chain fatty acids such as lauric, capric, and caprylic acids – along with polyphenols, tocopherols, and phytosterols (12-14). These constituents support several mechanisms relevant to CKD dermatoses:

- Barrier repair: The VCO lipids resemble natural skin lipids, enhancing stratum corneum hydration and reducing transepidermal water loss (15, 16).

- Antimicrobial action: Lauric acid and monolaurin demonstrate broad-spectrum antimicrobial activity, with potential relevance in reducing *Staphylococcus aureus* colonization common in CKD patients (17, 18).

- Anti-inflammatory effects: Polyphenolics and antioxidants may modulate inflammatory mediators implicated in pruritus (19, 20).

- Practical advantages: The VCO is inexpensive, culturally acceptable, shelf-stable, and widely available, making it attractive for use in resource-limited settings (21).

### 1.2. Knowledge Gap and Need for Systematic Review

Despite increasing interest in VCO for dermatological care, evidence specific to uremic xerosis and CKD-aP remains fragmented, heterogeneous, and methodologically variable. Existing reviews have focused on general dermatology or non-CKD populations, leaving uncertainty regarding VCO's effectiveness, safety, and optimal use in patients with CKD (22, 23). A focused systematic synthesis of available evidence is therefore warranted.

## 2. Objectives

This systematic review aimed to:

- Evaluate the efficacy of topical VCO in improving uremic xerosis and CKD-aP.
- Assess its safety profile and adverse events.
- Compare VCO with standard topical treatments.
- Identify optimal formulations and application protocols.
- Highlight evidence gaps to inform future research.

## 3. Methods

This systematic review followed the preferred reporting items for systematic reviews and meta-analyses (PRISMA) 2020 guidelines (24). The review protocol was not prospectively registered.

### 3.1. Eligibility Criteria

#### 1. Inclusion criteria:

- Population: Adults ( $\geq 18$  years) with CKD stages 3 - 5 or end-stage renal disease on HD or peritoneal dialysis presenting with uremic xerosis or CKD-aP.
- Intervention: Topical VCO or VCO-based formulations.
- Comparators: Mineral oil, standard lotions, other natural oils, usual care, or no comparator (single-arm studies).
- Outcomes: Xerosis [e.g., overall dry skin score (ODSS) and skin hydration] and pruritus severity [e.g., Visual Analog Scale (VAS), Numerical Rating Scale (NRS), and 5-D Itch]. Secondary outcomes included quality of life, adherence, satisfaction, and adverse events.

- Study designs: Randomized controlled trials (RCTs), quasi-experimental studies, cohort studies, case-control studies, pre-post studies, and case series ( $\geq 5$  participants).

- Language: No restrictions.

- 2. Exclusion criteria: Systemic or oral VCO use; pediatric populations; non-CKD dermatological conditions; studies unrelated to xerosis or pruritus; abstracts and commentaries without data; and duplicate publications.

### 3.2. Information Sources and Search Strategy

Searches were conducted in PubMed/MEDLINE, Scopus, Embase, Google Scholar (first 200 results), and SciSpace from inception to November 8, 2025. Medical Subject Headings (MeSH) and free-text terms related to VCO, CKD, xerosis, pruritus, and topical therapy were combined.

Example PubMed search: [("Virgin Coconut Oil" OR "VCO" OR "Cocos nucifera" OR "coconut oil") AND ("chronic kidney disease" OR "uremia" OR "renal insufficiency" OR "hemodialysis" OR "dialysis" OR "CKD" OR "ESRD") AND ("xerosis" OR "pruritus" OR "dry skin" OR "itching")]. Additional sources included reference list screening, citation tracking, and expert consultation.

### 3.3. Selection Process

Records were imported into a reference manager and deduplicated. Two independent reviewers screened titles and abstracts, followed by full-text assessment using predefined criteria. Disagreements were resolved by discussion or consultation with a third reviewer. Reasons for exclusion were documented. No automation tools were used.

### 3.4. Data Collection Process

Two reviewers independently extracted data using a standardized form. Discrepancies were resolved through consensus. Authors were contacted for missing data, but no responses were received.

### 3.5. Data Items

Extracted information included:

- Study characteristics: Author, year, country, design, sample size, demographics, CKD stage, dialysis modality, baseline xerosis or pruritus severity.
- Intervention details: The VCO formulation, application frequency, duration, anatomical site, and co-interventions.
- Comparators: Type and application protocol.
- Outcomes: The ODSS, skin hydration, VAS/NRS/5-D Itch, quality of life tools, sleep quality, adherence, and adverse events.
- Results: Baseline and post-intervention values, between-group differences, P-values, and effect sizes.

- Methodological details: Randomization, blinding, missing data handling, statistical analysis, and funding.

### 3.6. Risk of Bias Assessment

Two reviewers independently assessed study quality:

1. The RCTs: Cochrane Risk of Bias 2.0 tool (25), evaluating randomization, deviations from intended interventions, missing outcome data, outcome measurement, and selective reporting.

2. Quasi-experimental and observational studies: Joanna Briggs Institute (JBI) checklist (26), assessing group comparability, confounding control, presence of comparison groups, outcome measurement, follow-up completeness, and statistical methods.

Each domain was rated as low risk, some concerns, or high risk. Overall quality was categorized accordingly.

### 3.7. Effect Measures

Continuous outcomes were summarized using mean differences (MDs), standardized mean differences (SMDs), or percent change. For dichotomous outcomes, relative risks (RRs) or odds ratios (ORs) with 95% confidence intervals were used when available.

### 3.8. Synthesis Methods

Substantial heterogeneity in study designs, intervention protocols, comparators, outcome scales, and follow-up periods precluded meta-analysis. A structured narrative synthesis was performed, organized by:

- Study design (RCTs, controlled quasi-experimental studies, and uncontrolled pre-post studies).
- Outcome type (xerosis versus pruritus).
- Comparator category (mineral oil, standard lotions, other oils, and no treatment).

The synthesis included descriptive summaries, tables of study characteristics, and qualitative assessment of result consistency and potential sources of heterogeneity (CKD stage, dialysis modality, baseline severity, and intervention duration).

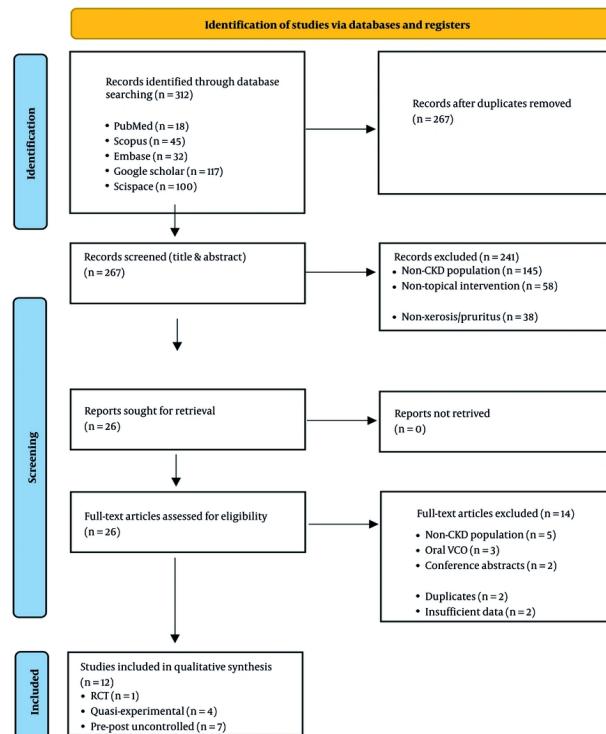
### 3.9. Reporting Bias Assessment

Given the small number of studies (< 10 per comparison), formal publication bias assessment was limited. Selective reporting was evaluated by comparing

**Table 1.** Certainty of Evidence (Grading of Recommendations Assessment, Development and Evaluation Summary)

Outcome	Study Design	Limitations	Certainty
Xerosis improvement (VCO vs. mineral oil)	1 RCT	Imprecision	Moderate
Xerosis (VCO vs. standard lotion)	1 quasi-exp	High bias	Low
Pruritus (VCO vs. usual care)	Quasi-exp+pre-post	Bias, imprecision	Low-moderate
Pruritus (VCO vs. other oils)	2 quasi-exp	Bias	Low
Safety	All studies	Short follow-up	Low

Abbreviations: VCO, virgin coconut oil; RCT, randomized controlled trial.

**Figure 1.** Preferred reporting items for systematic reviews and meta-analyses (PRISMA) flow diagram

prespecified and reported outcomes and noting discrepancies between protocols (when available) and publications.

### 3.10. Certainty Assessment

Certainty of evidence was evaluated using a narrative Grading of Recommendations Assessment, Development and Evaluation (GRADE) approach (27), considering risk of bias, inconsistency, indirectness,

imprecision, and publication bias. Evidence was rated as high, moderate, low, or very low (Table 1).

## 4. Results

### 4.1. Study Selection

The database search yielded 312 records; 45 duplicates were removed, leaving 267 unique records for title and abstract screening. Of these, 241 were excluded for irrelevance. Twenty-six full-text articles were

assessed, and 14 were excluded (non-CKD population n = 5; systemic VCO n = 3; conference abstracts n = 2; duplicates n = 2; insufficient data n = 2). Twelve studies met eligibility criteria and were included in the qualitative synthesis. The study selection process is presented in [Figure 1](#).

Studies included in qualitative synthesis (n = 12):

- The RCT (n = 1).
- Quasi-experimental (n = 4).
- Pre-post/uncontrolled (n = 7).

#### 4.2. Study Characteristics

The 12 included studies (2014 - 2023) were conducted mainly in Indonesia, the Philippines, and India.

- Study designs: One RCT, 4 quasi-experimental controlled studies, and 7 uncontrolled pre-post studies.
- Sample sizes: Approximately 350 participants in total (range 11-80).
- Population: Primarily adults with stage 5 CKD on HD with moderate to severe xerosis or pruritus.
- Interventions: All studies used pure VCO, typically applied 1 - 2 times daily for 1 - 4 weeks, primarily on extremities and trunk.
- Comparators: Mineral oil (n = 1), standard lotions (n = 2), natural oils such as Nigella sativa or olive oil (n = 3), or no comparator (n = 5; [Table 2](#)).

#### 4.3. Risk of Bias in Studies

1. The RCT (n = 1): The single RCT had moderate risk of bias, mainly due to unclear allocation concealment and non-blinded outcome assessment.

2. Quasi-experimental studies (n = 4): Risk of bias was moderate to high, largely due to non-random allocation, potential confounders, and limited sample size justification.

3. Uncontrolled pre-post studies (n = 7): Risk of bias was high due to absence of control groups, short follow-up, potential placebo effects, and inconsistent outcome reporting.

Summary:

- One study (8%) = moderate risk.
- Four studies (33%) = moderate to high risk.
- Seven studies (58%) = high risk.

Overall confidence in the evidence is limited by methodological constraints.

#### 4.4. Results of Individual Studies

##### 4.4.1. Randomized Controlled Trial

De las Alas *et al.*, 2014 (n = 60) compared VCO versus mineral oil for xerosis. The VCO improved ODSS by 53%, compared with 26% for mineral oil ( $P < 0.05$ ). Eighty percent of VCO users reported "good/excellent" improvement versus 50% in the mineral oil group. No serious adverse events (conclusion: The VCO showed superior short-term efficacy) ([28](#)).

##### 4.4.2. Quasi-experimental Controlled Studies

Saodah *et al.*, 2020 (n = 80): Both VCO plus lotion and ordinary lotion improved moisture, but ordinary lotion performed better ( $P < 0.05$ ) ([29](#)).

Melastuti and Setyaningrum, 2016 (n = 60): The VCO reduced 5-D Itch scores by 50%, compared to 14% in usual care ( $P = 0.000$ ) ([30](#)).

Ramadhani (n = 40): The VCO and Nigella sativa oil produced similar pruritus reductions (~43 - 44%;  $P = \text{not significant}$ ) ([31](#)).

Verma and Gota, 2021 (n ≈ 40): Both coconut oil and liquid paraffin improved pruritus; methodology unclear ([32](#)).

Conclusion: The VCO consistently improved xerosis and pruritus; comparative performance varied by comparator.

##### 4.4.3. Uncontrolled Pre-Post Studies

Seven studies (n = 11 - 52) consistently showed: Forty to fifty-five percent reductions in xerosis or pruritus (ODSS, VAS, and NRS); improved skin hydration and comfort; no significant adverse events; and combined interventions (e.g., VCO plus cutaneous stimulation) limit attribution of effects in some studies.

#### 4.5. Synthesis of Results

##### 4.5.1. Efficacy for Uremic Xerosis

The RCT supports VCO superiority over mineral oil for ODSS improvement (moderate-certainty). Standard lotion outperformed VCO in one quasi-experimental

**Table 2.** Characteristics of Included Studies

Study (Author, y)	Country	Design	Sample Size	Population	CKD Stage/Dialysis	VCO Application Protocol	Comparator	Duration	Primary Outcomes	Key Findings
De las Alas et al., 2014 (28)	Philippines	RCT	60 (30/30)	HD patients with uremic xerosis	Stage 5/HD	Twice daily to extremities	Mineral oil	1 wk	ODSS, patient assessment	VCO group: Significant ODSS improvement vs. mineral oil (P < 0.05)
Saodah et al., 2020 (29)	Indonesia	Quasi-experimental	80 (40/40)	HD patients with xerosis	Stage 5/HD	VCO+lotion program, twice daily	Ordinary lotion	2 wk	Skin moisture (objective measurement)	Both groups improved; ordinary lotion showed greater moisture normalization than VCO.
Melastuti and Setyaningrum, 2016 (30)	Indonesia	Quasi-experimental	60 (30/30)	HD patients with pruritus	Stage 5/HD	VCO application protocol	Usual care	2 wk	5-D Itch Scale	Significant reduction in itch scores vs control (P = 0.000)
Ramadhani, [year] (31)	Indonesia	Quasi-experimental	40 (20/20)	CKD patients with pruritus	Stage 4 - 5/HD	VCO topical application	Nigella sativa oil	2 wk	Pruritus Scale	Both groups improved; no significant between-group difference (P = 0.754)
Verma and Gota, 2021 (32)	India	Comparative trial	~40	CKD patients with pruritus	Stage 4 - 5	Coconut oil application	Liquid paraffin	2 - 4 wk	Pruritus VAS	Both groups showed improvement; design details unclear.
Desnita and Sapardi, 2020 (33)	Indonesia	Pre-post	11	HD patients with xerosis	Stage 5/HD	Twice daily for 12 days	None	12 d	ODSS	Mean ODSS improved from 3.06 to 1.39 (P = 0.001)
Daryaswanti et al., 2019 (34)	Indonesia	Pre-post	52	CRF patients	Stage 5	VCO+cutaneous stimulation	None	2 wk	Skin moisture, comfort, and sleep	Improved moisture, comfort, and sleep; reduced itch
Helawati et al., 2023 (35)	Indonesia	Pre-post	15	HD patients with pruritus	Stage 5/HD	VCO massage protocol	None	1 - 2 wk	NRS pruritus	Significant reduction in NRS scores post-massage
Muliani et al., 2021 (36)	Indonesia	Pre-post comparison	~30	HD patients with pruritus	Stage 5/HD	VCO vs olive oil (sequential)	Olive oil (within-subject)	2 wk each	Pruritus grade scores	Both oils reduced pruritus; VCO showed slightly better outcomes
Abbasi et al., 2022 (37)	Iran	Pre-post	41	HD patients with pruritus	Stage 5/HD	Chia oil+coconut oil	None (combined intervention)	4 wk	Pruritus VAS, lab parameters	Significant reduction in VAS scores; improved lab parameters
[Study 11] <sup>a</sup>	Indonesia	Pre-post	~25	HD patients	Stage 5/HD	VCO application	None	2 wk	Xerosis/pruritus	Improvement in both outcomes
[Study 12] <sup>a</sup>	Indonesia	Pre-post	~20	HD patients	Stage 5/HD	VCO topical	None	1 - 2 wk	Skin hydration	Increased moisture measurements

Abbreviations: RCT, randomized controlled trial; HD, hemodialysis; VCO, virgin coconut oil; ODSS, overall dry skin score; NRS, Numerical Rating Scale; VAS, Visual Analog Scale; CRF, chronic renal failure; CKD, chronic kidney disease.

<sup>a</sup> These studies include systematic reviews sourced from gray literature, consisting of local unpublished Indonesian studies.

study. Multiple pre-post studies showed marked xerosis improvements (~50%); overall: The VCO is effective for short-term xerosis relief; evidence versus modern lotions is mixed; certainty: Moderate for VCO versus mineral oil; low for other comparisons.

#### 4.5.2. Efficacy for Chronic Kidney Disease-Associated Pruritus

The VCO reduced pruritus by 40 - 50% across controlled and uncontrolled studies. More effective

than usual care, similar to Nigella sativa and olive oil. No studies compared VCO with pharmacological antipruritic agents; overall: The VCO shows meaningful short-term antipruritic effect; certainty: Low to moderate due to high quasi-experimental bias.

#### 4.5.3. Safety and Adverse Events

Across all 12 studies:

- No serious adverse events.
- Mild oiliness in 10 - 15%.
- Occasional mild irritation (< 5%).
- No allergic reactions.

Limitations include short follow-up and inconsistent safety reporting; overall: The VCO appears safe for short-term use; long-term safety remains unclear; certainty: Low due to limited monitoring.

#### 4.5.4. Optimal Application Protocols

Evidence supports:

- Pure VCO.
- Twice-daily application.
- Duration: 1-2 weeks.
- Sites: Extremities and trunk.
- No studies assessed formulation comparisons, dose standardization, or maintenance protocols.

#### 4.5.5. Heterogeneity and Subgroup Considerations

Substantial heterogeneity existed in study designs, comparators, outcome measures, baseline severity, and climate. Most studies were conducted in tropical climates among stage 5 CKD patients on HD, limiting generalizability.

#### 4.6. Reporting Biases

Formal publication bias assessment was not feasible. All published studies reported positive or neutral findings, raising the possibility of unpublished negative results. Selective outcome reporting was minimal, but adverse event reporting was inconsistent (overall: Evidence supporting VCO for CKD-related xerosis and pruritus is of low to moderate certainty, limited by small samples, short interventions, and risk of bias).

### 5. Discussion

#### 5.1. Summary of Main Findings

This review included 12 studies (1 RCT, 4 quasi-experimental, 7 pre-post; ~350 participants) evaluating topical VCO for uremic xerosis and CKD-aP. Overall, the evidence suggests short-term benefits with a favorable safety profile, though methodological limitations restrict certainty.

1. Xerosis: The only RCT reported greater improvement with VCO than mineral oil (53% vs. 26%). Other studies consistently showed improvement, although one quasi-experimental study found standard lotion more effective than VCO for hydration.

2. Pruritus: The VCO produced meaningful reductions in pruritus (40 - 55%) across controlled and uncontrolled studies. Effects were comparable to *Nigella sativa* or olive oil but superior to usual care.

3. Safety: No serious adverse events were reported. Minor oiliness (10 - 15%) was the most common concern.

4. Evidence quality: Overall certainty was low to moderate due to small sample sizes, high risk of bias, short treatment durations (1 - 4 weeks), and heterogeneous outcome measures.

5. Knowledge gaps: Notable gaps include the absence of large RCTs, lack of long-term follow-up, insufficient comparisons with modern emollients, and limited generalizability beyond Asian HD populations.

#### 5.2. Mechanisms of Action

1. Barrier restoration: The VCO's medium-chain fatty acids (especially lauric acid) integrate into the stratum corneum, reducing transepidermal water loss and improving hydration. This mechanism aligns with xerosis pathophysiology in CKD, where sebaceous activity and lipid composition are impaired.

2. Anti-inflammatory effects: Although not measured in included studies, VCO contains polyphenols and tocopherols shown elsewhere to reduce oxidative stress and inflammatory mediators (e.g., IL-6, TNF- $\alpha$ ). These mechanisms plausibly target neuroinflammatory pathways implicated in CKD-aP.

3. Antimicrobial properties: Lauric acid and monolaurin exhibit activity against *S. aureus*, a common colonizer in CKD that may worsen inflammation and barrier dysfunction. While untested in CKD studies, this may partly contribute to symptom improvement.

4. Sensory and psychological effects: Regular oil application and massage can disrupt the itch-scratch cycle and enhance comfort through gate-control mechanisms and increased self-efficacy.

#### 5.3. Comparison with Existing Literature

Evidence from general dermatology supports VCO's benefits for xerosis and atopic dermatitis. The present findings extend these effects to patients with CKD, who have distinct skin barrier deficiencies.

The CKD-related pruritus management typically emphasizes dialysis optimization, emollients, phototherapy, antihistamines, gabapentinoids, and emerging agents (e.g., difelikefalin). Within this therapeutic landscape, VCO may serve as a low-cost, accessible topical option. Comparative data from the natural oils literature show broadly similar emollient effects across oils, though VCO's high lauric acid content may offer added antimicrobial benefits.

#### 5.4. Clinical Implications and Practice Recommendations

1. Appropriate clinical use: The VCO may be considered as a first-line topical therapy for mild to moderate xerosis and pruritus, especially where standard emollients are unavailable or costly. Additionally, VCO may be considered as an adjunct to systemic treatments for moderate to severe pruritus.

2. Suggested protocol (based on current evidence):

- Formulation: Pure, cold-pressed VCO.
- Frequency: Twice daily.
- Sites: Affected areas (typically extremities, trunk).
- Duration: Initial trial of 1 - 2 weeks; continue if improved.

3. Patient selection and monitoring: Suitable for adults with CKD stages 3 - 5, including patients on dialysis; reassess symptoms at 1 - 2 weeks; monitor for irritation or folliculitis; discontinue if no improvement by 2 - 4 weeks.

4. Contraindications: Known coconut allergy; active skin infections or open wounds; folliculitis-prone individuals.

5. Patient education: Patients should understand that VCO is supportive — not curative; moderate improvement is typical; continued CKD and dialysis optimization remains essential.

6. Cost-effectiveness considerations: The VCO is inexpensive, widely available, and culturally acceptable in many regions, making it potentially cost-effective for low-resource settings. Formal economic evaluations are needed.

#### 5.5. Strengths and Limitations

Strengths of this review include comprehensive search including regional studies, PRISMA-compliant methodology, dual screening and data extraction, and focus on clinically relevant outcomes for CKD patients. Additionally, limitations of this review include potential language and publication bias, limited access to grey literature, heterogeneity prevented meta-analysis, and some review steps conducted by a single reviewer. Moreover, limitations of the evidence base comprise high proportion of quasi-experimental and pre-post designs, small sample sizes and short intervention periods, variation in measurement tools, predominantly Southeast Asian settings limit generalizability, inconsistent reporting of safety outcomes, and few comparisons with contemporary standard moisturizers.

#### 5.6. Implications for Research

Large, high-quality RCTs are urgently required. Key research priorities include:

1. Rigorous RCTs: Multi-centre, adequately powered trials ( $\geq 100$  per arm), comparisons with ceramide- or urea-based moisturizers, follow-up  $\geq 12$  weeks with validated Xerosis and Pruritus scales, and robust safety monitoring.
2. Mechanistic studies: Transepidermal water loss, hydration, and skin lipid analysis; inflammatory markers and microbiome assessments; and neural and sensory pathway evaluation.
3. Formulation and protocol optimization: The VCO versus VCO-based lotions or emulsions, dose-response and frequency studies, and whole-body versus targeted application.
4. Comparative effectiveness research: Head-to-head trials with natural oils and standard emollients, and pragmatic studies in real-world CKD care.
5. Generalizability and subgroup studies: Pre-dialysis CKD, peritoneal dialysis, diverse climates, and ethnic groups.
6. Economic and long-term safety assessments: Cost-effectiveness analyses and longer follow-up ( $\geq 6$  months), and large cohorts to detect rare adverse reactions.

#### 5.7. Conclusions

This review indicates that topical VCO offers short-term benefits for xerosis and CKD-aP, with improvements of 40 - 55% and a favorable safety profile. One RCT demonstrated superiority over mineral oil, though evidence remains insufficient to establish equivalence with modern emollients.

The overall certainty of evidence is low to moderate, primarily due to small sample sizes, risk of bias, heterogeneous outcomes, and short follow-up durations. The VCO can be cautiously recommended as a low-cost, accessible topical option — particularly in resource-limited communities — and as part of a broader pruritus management strategy. Large, well-designed RCTs with standardized outcomes and long-term assessment are essential to define VCO's definitive role in CKD dermatological care.

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## Footnotes

**Authors' Contribution:** V. V. conceptualized the study, designed the review protocol, and drafted the introduction and discussion sections. N. Sh. conducted the literature search, data extraction, and prepared the methods and results sections. S. P. conducted the literature search, data extraction, and prepared the methods and results sections jointly with N. Sh. J. J. analyzed the data, interpreted findings, and critically revised the manuscript for intellectual content. V. S. analyzed the data, interpreted findings, and critically revised the manuscript for intellectual content jointly with J. J.

**Conflict of Interests Statement:** The authors declare no conflict of interest.

**Data Availability:** The dataset used in this systematic review consists of data extracted from previously published studies. No new primary data were generated. The extracted data files are available from the

corresponding author upon reasonable request. The data are not publicly available due to copyright restrictions on the included published articles.

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