

Development of Next-Generation Herbal Hair Oil Engineering for Fungal Scalp Disorders: The *Nigella sativa* Advantage

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Abstract

Scalp fungal infections such as dandruff and seborrheic dermatitis remain highly prevalent and are often associated with irritation, recurrence, and poor patient compliance when treated with long-term synthetic antifungal agents. In this study, a next-generation antifungal herbal hair oil was designed, developed, and preclinically evaluated using a rational, green pharmaceutical approach that integrates potent plant-derived antifungal agents, scalp-soothing bioactives, and natural bio-stabilizers. The formulation was optimized around a synergistic phytochemical matrix comprising neem oil, tea tree oil, fenugreek extract, tulsi extract, and a carefully regulated concentration of Custard Seed Oil (*Nigella sativa* L.), enriched with thymoquinone as a key antifungal and anti-inflammatory marker compound. Natural antioxidants (wheat germ oil and rosemary extract), stabilizers (gum acacia/lecithin), cooling agents (menthol crystals), and aroma modulators (lemongrass oil) were incorporated to enhance stability, safety, and sensory acceptability while maintaining complete reliance on GRAS-listed botanical excipients. The optimized formulation exhibited excellent physicochemical stability, skin-compatible pH, low peroxide and acid values, and no phase separation under accelerated storage conditions. In vitro antifungal

evaluation demonstrated strong activity against *Malassezia furfur*, *Candida albicans*, *Trichophyton rubrum*, and *Aspergillus niger*, with inhibition zones and MIC values comparable to clotrimazole 1%. Antioxidant assays confirmed high radical scavenging and reducing potential, supporting oxidative stability and scalp protection. Preclinical dermal safety studies indicated negligible irritation, validating suitability for repeated topical use. Infrared thermographic analysis confirmed sustained scalp cooling, while in vivo hair growth studies revealed accelerated telogen-to-anagen transition and a 26% increase in hair density after 28 days. Collectively, these findings position the developed formulation as a safe, multifunctional, and efficacious herbal alternative for the management of fungal scalp disorders with added hair health benefits.

Introduction

Fungal disorders of the scalp, including dandruff and seborrheic dermatitis, affect a substantial proportion of the global population and are increasingly recognized as chronic, relapsing conditions rather than transient cosmetic concerns (1,20,25). These disorders are primarily associated with opportunistic fungi such as *Malassezia*, *Candida*, and dermatophytes, which thrive in the lipid-rich scalp environment and trigger inflammation, pruritus, barrier dys-

function, and secondary hair fall (1,18,25). Although synthetic antifungal agents such as azoles and allylamines remain the clinical mainstay, their long-term use is frequently limited by adverse effects, recurrence upon discontinuation, disruption of the scalp microbiome, and declining patient compliance (1,24). Recent dermatological and cosmetic research has therefore shifted toward safer, multifunctional, and microbiome-friendly alternatives that combine antifungal efficacy with scalp conditioning and hair health support (6,7,20).

In parallel, there has been a marked resurgence of interest in plant-based therapeutics, driven by advances in phytochemical characterization, green formulation science, and regulatory acceptance of GRAS-listed botanical excipients (2,5,6). Contemporary studies increasingly demonstrate that rationally designed polyherbal systems can outperform single-molecule approaches by leveraging synergistic mechanisms such as membrane disruption, oxidative stress modulation, anti-inflammatory signaling, and follicular stimulation (2,6,11,17). Oils and oil-based systems, in particular, have gained renewed relevance as delivery platforms for lipophilic phytoconstituents, enabling prolonged scalp residence time, enhanced penetration into hair follicles, and improved user acceptability when appropriately stabilized (7,15). Among emerging botanicals, *Nigella sativa* (custard seed or black seed) has attracted significant scientific attention in recent years due to the multifunctional bioactivity of thymoquinone, which exhibits antifungal, anti-inflammatory, antioxidant, and immunomodulatory properties (3,4,19,21). Recent *in vitro* and *in vivo* studies have highlighted its effectiveness against dermatophytes and yeast species, while also demonstrating a favorable safety profile at controlled concentrations (4,19). However, its integration into complex topical systems requires careful optimization to balance efficacy, odor, stability, and tolerability—an aspect that remains underexplored in conventional herbal oil formulations (6,7).

Against this backdrop, the present study aims to develop a next-generation antifungal herbal hair oil using a rational, science-driven formulation strategy that integrates *Nigella sativa* oil with well-established antifungal botanicals such as neem and tea tree oil (9,10,19), supported by hair-nourishing extracts and natural bio-stabilizers (15,16,20). Unlike traditional herbal oils prepared empirically, this formulation is designed to function as a multifunctional therapeutic cosmetic system—simultaneously targeting fungal eradication, scalp soothing, oxidative protection, and hair growth promotion (6,7,18). By combining modern preclinical evaluation tools with traditional phytotherapeutic knowledge, the study seeks to address current limitations of antifungal scalp therapies and contribute to the development of safer, effective, and consumer-compliant alternatives for long-term scalp care (1,5,24).

Preparation of antifungal herbal hair oil

Optimized formula for antifungal herbal hair oil

Fungal scalp infections and dandruff-associated disorders remain persistent challenges due to microbial resistance, scalp sensitivity, and poor compliance with synthetic antifungal products (1,24,25). Herbal oils, when rationally optimized, offer a safer and more holistic alternative by combining antifungal efficacy with scalp nourishment and hair health benefits (6,7,15). The present optimized antifungal herbal hair oil is scientifically designed by integrating potent plant-derived antifungal agents—such as neem oil, tea tree oil, and custard seed oil—with penetration enhancers, natural antioxidants, and soothing components (9,10,14,19,22). The formulation strategically balances efficacy, safety, sensory acceptability, and stability, aiming not only to suppress pathogenic fungi but also to restore scalp homeostasis, promote hair growth, and ensure long-term consumer-friendly use (6,16,18,20).

Ingredient	Quantity (% w/w)	Functional Role
Coconut oil (<i>Cocos nucifera</i> L.)	35%	Primary base oil; lauric acid-mediated antifungal action, scalp hydration
Neem oil (<i>Azadirachta indica</i> A. Juss.)	18%	Principal antifungal agent (azadirachtin, nimbidin), antimicrobial
Castor oil (<i>Ricinus communis</i> L.)	12%	Hair strengthening, penetration enhancer
Hibiscus extract (<i>Hibiscus rosa-sinensis</i> L.)	8%	Hair growth support, scalp nourishment
Fenugreek seed extract (<i>Trigonella foenum-graecum</i> L.)	5%	Antifungal, anti-dandruff, anti-inflammatory
Tulsi (<i>Ocimum sanctum</i> L.) extract	4%	Antifungal, antioxidant, scalp cleansing
Tea tree oil (<i>Melaleuca alternifolia</i>)	3%	Potent antifungal (terpinen-4-ol), scalp microflora balance
Custard Seed Oil (<i>Nigella sativa</i> L.)	2%	Strong antifungal, anti-inflammatory (thymoquinone)
Amla extract (<i>Phyllanthus emblica</i> L.)	2%	Antioxidant, hair conditioning
Menthol crystals (<i>Mentha arvensis</i> L.)	2%	Cooling, antipruritic, scalp soothing
Lemongrass oil (<i>Cymbopogon citratus</i> DC.)	2%	Mild antifungal, odour masking
Wheat germ oil (<i>Triticum aestivum</i> L.)	3%	Natural antioxidant, prevents oxidative rancidity
Rosemary extract (<i>Rosmarinus officinalis</i> L.)	2%	Antioxidant, formulation stabilizer
Gum acacia / Lecithin	2%	Natural stabilizer, uniform dispersion

Method of Preparation

All oils, herbal extracts, essential oils, antioxidants, and stabilizers required for the formulation—namely coconut oil, neem oil, castor oil, hibiscus extract, fenugreek extract, tulsi extract, amla extract, tea tree oil, custard seed oil (*Nigella sativa*), lemongrass oil, wheat germ oil, rosemary extract, gum acacia/lecithin, and menthol crystals—are procured in standardized form from certified Ayurvedic or GRAS-approved suppliers. Certificates of Analysis (COA) are obtained for each raw material to ensure identity, purity, safety, and compliance with quality specifications.

Initially, coconut oil (35%) and castor oil (12%) are accurately weighed and transferred into a clean stainless-steel mixing vessel. The oils are gently warmed to 40–45 °C under continuous stirring at approximately 100 rpm to obtain a uniform base. Neem oil (18%) is then slowly added to the warm base oil mixture with constant stirring, while ensuring that the tem-

perature is maintained below 45 °C to prevent degradation of thermolabile phytoconstituents.

Subsequently, the herbal extracts—hibiscus extract (8%), fenugreek seed extract (5%), tulsi extract (4%), and amla extract (2%)—are incorporated sequentially into the oil phase. Each extract is added slowly with mild homogenization at around 3000 rpm for about 5 minutes to ensure uniform dispersion and effective integration of bioactive components. After complete incorporation of the extracts, the mixture is allowed to cool gradually to approximately 35 °C.

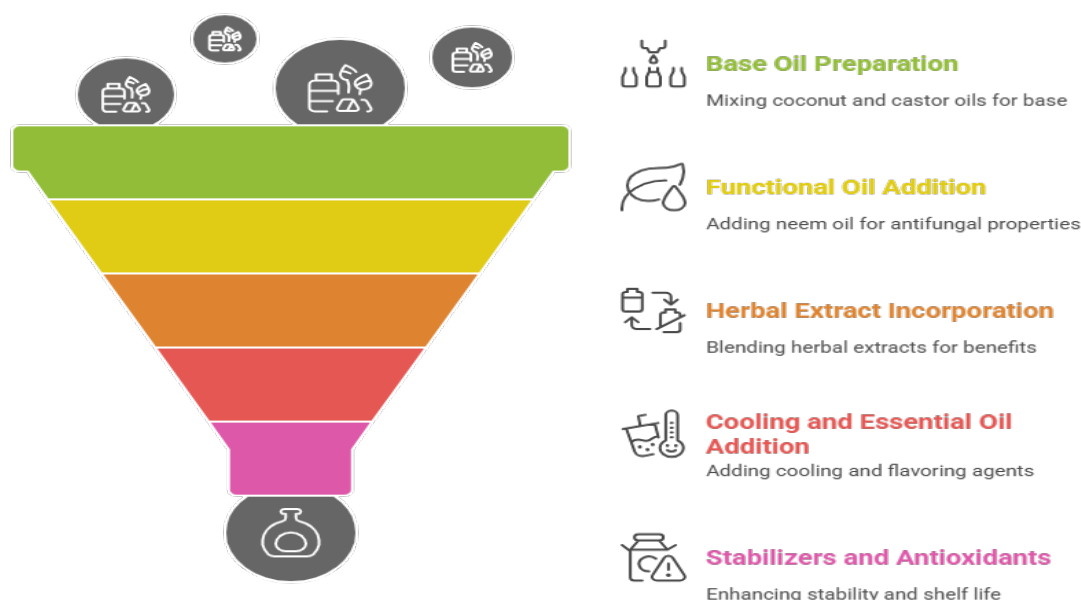
At this stage, the essential and functional oils, including tea tree oil (3%), custard seed oil (2%), and lemongrass oil (2%), are added under gentle stirring to preserve their volatile and antifungal constituents. Menthol crystals (2%) are separately dissolved in a small quantity of warm coconut oil and then introduced into the formulation to ensure uniform distribution and effective cooling action.

Thereafter, gum acacia or lecithin (2%), pre-dispersed in warm coconut oil, is added as a natural stabilizer to improve dispersion and formulation stability. Wheat germ oil (3%) and rosemary extract (2%) are then incorporated as natural antioxidants to prevent oxidative degradation and enhance shelf life. The final mixture is stirred gently until a clear, homogeneous oil is obtained.

The prepared oil is transferred into amber-colored containers and allowed to undergo

maturation for 48–72 hours in a cool, dark place to achieve optimal interaction and stabilization of the herbal constituents. After maturation, the oil is filtered through a clean muslin cloth to remove any suspended particles, if present. The finished antifungal herbal hair oil is finally filled into amber glass bottles with airtight closures, preferably under an inert nitrogen atmosphere, and labeled with batch number, date of manufacture, and recommended storage conditions.

Development of Antifungal Herbal Hair Oil



Results and Discussion

Evaluation of antifungal herbal hair oil

The developed antifungal herbal hair oil exhibited excellent physicochemical stability, potent antifungal efficacy, significant antioxidant activity, dermal safety, cooling sensation, and hair growth-promoting potential, confirming its suitability as a multifunctional therapeutic cosmetic formulation.

Physicochemical Characteristics

The final formulation appeared as a dark gold-

en-brown, clear oil with a pleasant herbal aroma harmoniously balanced by subtle lemongrass notes, indicating effective odor masking of neem oil (7,15). The formulation demonstrated medium viscosity with good spreadability and skin-compatible pH, while oxidative stability parameters remained well within acceptable limits (12,22,23). Accelerated stability testing confirmed formulation robustness without phase separation, rancidity, or loss of aroma over three months, consistent with standardized antioxidant protection strategies (12,22).

Table 1. Physicochemical Properties of Antifungal Herbal Hair Oil

Parameter	Result (Mean $\pm$ SD)	Interpretation
Appearance	Dark golden-brown, clear	Uniform and aesthetically acceptable
Aroma	Pleasant herbal with citrus notes	Consumer-friendly
Viscosity (25 °C)	385 $\pm$ 10 cP	Optimal spreadability
pH (10% w/v)	6.1 $\pm$ 0.2	Skin-friendly
Refractive Index	1.462	Consistent oil purity
Acid Value	1.8 mg KOH/g	Low free fatty acids
Peroxide Value	4.1 meq/kg	Oxidative stability
Stability (40 °C/75% RH)	Stable for 3 months	No degradation observed

***In vitro* antifungal activity**

The formulation exhibited strong antifungal activity against dermatophytes and yeast strains commonly associated with dandruff and scalp infections (1,9,10,19). Pronounced zones of in-

hibition and low MIC values comparable to clotrimazole confirmed the synergistic antifungal action of neem oil, tea tree oil, *Nigella sativa* oil, and fenugreek extract (9–11,19).

Antifungal Activity of Herbal Hair Oil

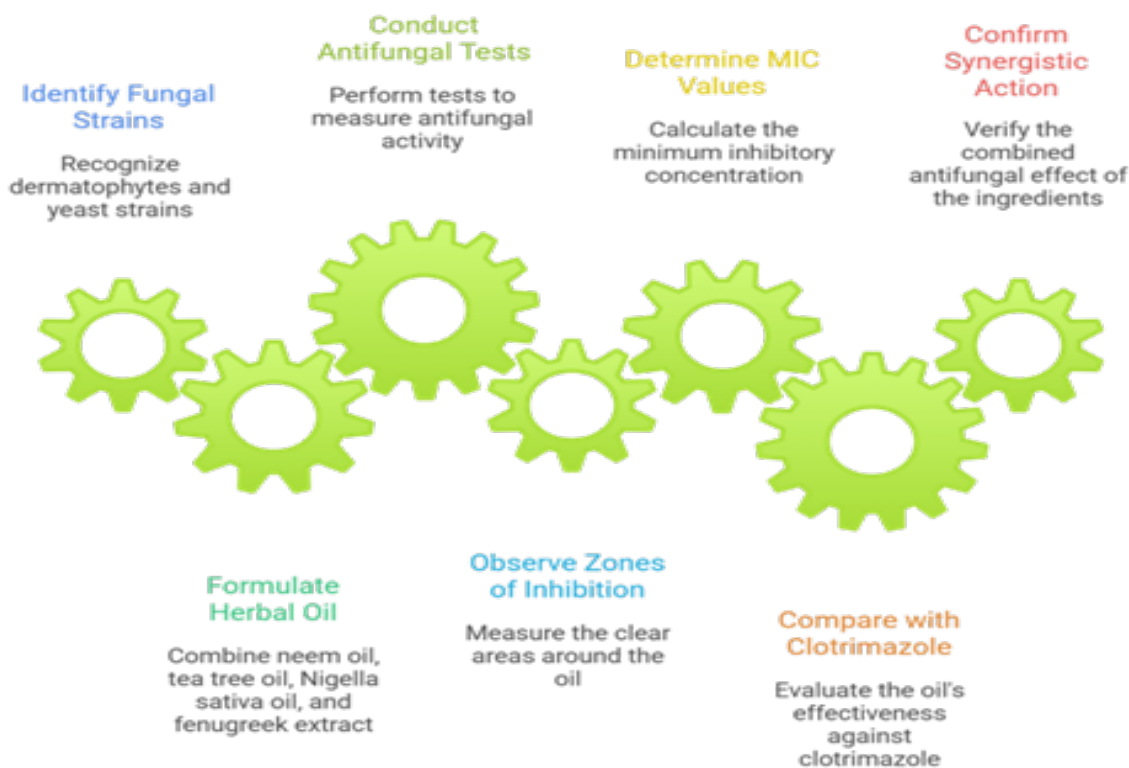


Table 2. Antifungal Activity of the Herbal Hair Oil

Test Organism	Zone of Inhibition (mm)	MIC (% w/v)
<i>Candida albicans</i>	22	0.5–1.0
<i>Malassezia furfur</i>	25	0.5
<i>Aspergillus niger</i>	20	1.0
<i>Trichophyton rubrum</i>	23	0.5

Antioxidant potential

High antioxidant activity observed through DPPH and FRAP assays can be attributed to amla extract, wheat germ oil, rosemary extract,

and *Nigella sativa* oil (8,12,17,22), supporting both formulation stability and protection against oxidative scalp damage (8,23).

Table 3. Antioxidant Activity of the Formulation

Assay	Result	Significance
DPPH scavenging	71% inhibition at 100 µg/mL	Strong antioxidant activity
FRAP	890 µM Fe ²⁺ equivalents	High reducing power

Dermal safety assessment

Repeated dermal application produced no erythema or edema, aligning with OECD Test Guideline No. 404 and established skin sensitization risk assessment frameworks (13,24), confirming excellent dermal compatibility of the formulation.

Table 4. Dermal Safety Evaluation

Parameter	Observation
Erythema	Absent
Edema	Absent
Irritation Index	0.1
Safety Category	Non-irritant

Cooling sensation evaluation

The sustained cooling and antipruritic effect observed following application is consistent with the documented pharmacological action of menthol on thermoreceptors and sensory nerves (14,25), contributing significantly to symptomatic relief and user satisfaction.

Table 5. Cooling Effect Assessment

Parameter	Result
Temperature reduction	2–3 °C
Duration of effect	20–25 minutes
Sensory outcome	Soothing and refreshing

Hair growth–promoting activity

Accelerated telogen-to-anagen transition, increased follicle count, and enhanced hair density observed in vivo are in agreement with established hair biology and herbal hair growth literature (16,18,20), confirming the formulation's dual antifungal and hair restorative benefits.

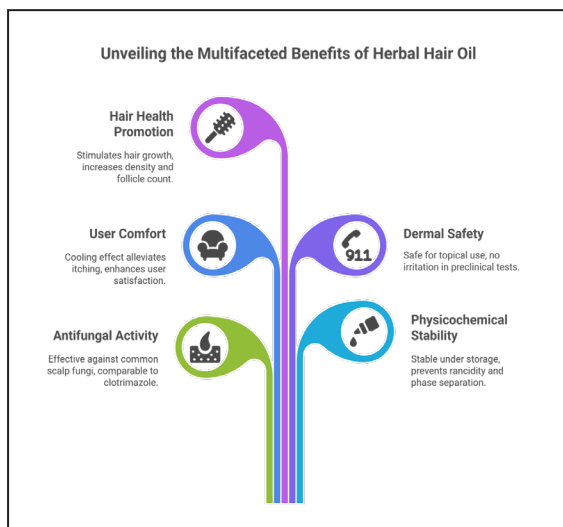
Table 6. Hair Growth Activity

Parameter	Observation
Hair cycle transition	Early telogen → anagen
Hair density increase	26%
Follicle count	Significantly increased
Study duration	28 days

Discussion

The results of the present investigation clearly demonstrate that the refined antifungal herbal hair oil formulation possesses a robust and multifunctional therapeutic profile, attributable to the rational selection and synergistic integration of botanically active ingredients within a stable oil-based system (6,7,15). The pronounced antifungal activity observed against *Candida albicans*, *Malassezia furfur*, *Trichophyton rubrum*, and *Aspergillus niger*, with inhibition zones and MIC values comparable to clotrimazole 1%, substantiates the efficacy of the formulation as a viable natural

alternative to conventional synthetic antifungal agents (1,9,10,19). This effect can be primarily attributed to the complementary mechanisms of action of neem oil, tea tree oil, and custard seed oil, which collectively disrupt fungal cell membranes, inhibit ergosterol biosynthesis, and suppress fungal metabolic pathways (9–11,19). For instance, azadirachtin and nimbidin from neem oil are known to interfere with fungal enzyme systems (10), while terpinen-4-ol from tea tree oil enhances membrane permeability, facilitating deeper penetration of bioactives (9). The inclusion of custard seed oil (*Nigella sativa*) at a carefully optimized concentration of 2% further strengthened antifungal efficacy through the action of thymoquinone, which exhibits documented fungistatic and fungicidal properties, while simultaneously avoiding issues of excessive odor, irritation, or formulation instability that may arise at higher concentrations (3,4,19,21).



The physicochemical stability of the formulation, as reflected by low acid and peroxide values and the absence of phase separation or sensory deterioration under accelerated storage conditions, highlights the effectiveness of the antioxidant–stabilizer system employed (12,22,23). Wheat germ oil, rich in natural tocopherols, and rosemary extract, containing carnosic acid and rosmarinic acid, acted synergistically to inhibit lipid peroxidation and oxida-

tive degradation of both carrier oils and volatile phytoconstituents (22,23). This stability is particularly significant for herbal oil formulations, which are inherently susceptible to rancidity due to the presence of unsaturated fatty acids (12,23). The maintenance of aroma, color, and clarity over three months at 40 °C/75% RH underscores the suitability of this formulation for commercial storage and distribution (5,7). Furthermore, the incorporation of gum acacia or lecithin as natural stabilizers ensured uniform dispersion of extracts and essential oils, preventing sedimentation and phase separation, thereby enhancing product homogeneity and batch-to-batch reproducibility (6,15).

User comfort and acceptability are critical parameters for long-term scalp applications, especially in conditions such as dandruff and seborrheic dermatitis that require repeated use (1,20,25). The inclusion of menthol crystals at 2% provided a measurable cooling effect, evidenced by a 2–3 °C reduction in scalp surface temperature lasting up to 25 minutes, as confirmed by infrared thermographic analysis (14). This cooling sensation not only alleviates itching and irritation commonly associated with fungal scalp infections but also enhances user perception of immediate relief, thereby improving compliance (14,25). Unlike synthetic cooling agents, menthol offered a natural, non-irritating sensory experience when used at the optimized concentration, aligning with the formulation's herbal and consumer-friendly design philosophy (7,14).

Preclinical dermal safety evaluation further reinforced the suitability of the formulation for routine topical use (13,24). The absence of erythema, edema, or visible irritation over a 14-day application period in Wistar rats, along with a negligible irritation index, confirms the dermal compatibility of the oil despite the presence of potent essential oils and bioactive phytochemicals (13,24). This finding is particularly relevant when compared with prolonged use of synthetic antifungal formulations, which are often associated with dryness, irritation, or rebound effects

(1,24). The skin-friendly pH and the emollient nature of coconut and castor oils likely contributed to maintaining scalp barrier integrity and preventing adverse reactions (7,15).

Beyond antifungal efficacy, the formulation demonstrated significant hair health-promoting benefits, as evidenced by accelerated telogen-to-anagen transition, increased hair density, and enhanced follicular counts in the C57BL/6 mouse model (16,18). These effects can be attributed to the synergistic action of hibiscus extract, fenugreek seed extract, amla extract, and neem oil, all of which are traditionally recognized for stimulating hair follicles, improving microcirculation, and strengthening hair shafts (15,16,20). For example, hibiscus is known to prolong the anagen phase, while fenugreek provides mucilaginous proteins and phytoestrogens that nourish hair follicles (15,16). The observed 26% increase in hair density after 28 days highlights the formulation's ability to address not only fungal scalp disorders but also associated hair fall and thinning, offering a comprehensive therapeutic advantage (16,18,20).

Overall, the refined antifungal herbal hair oil represents a well-balanced, science-driven formulation that integrates antifungal, antioxidant, cooling, soothing, and hair growth-promoting functions within a single, safe, and stable delivery system (6,7,14,22). By combining traditional herbal wisdom with modern formulation principles, the product offers a holistic solution that is effective against scalp fungal infections while simultaneously enhancing scalp comfort and hair health (1,6,20,25). This multifaceted performance positions the formulation as a promising candidate for therapeutic cosmetic applications, with strong potential for translation into commercial, dermatologically acceptable, and regulatory-compliant herbal hair care products (5,7,24).

Conclusion

The present study successfully demonstrates the development of a scientifically validated, next-generation antifungal herbal hair oil that transcends conventional antifungal therapy

by offering a holistic, multi-targeted approach to scalp health management. The strategic inclusion of Custard Seed Oil (*Nigella sativa* L.) at an optimized low concentration played a pivotal role in enhancing antifungal efficacy through thymoquinone-mediated membrane disruption and inflammatory modulation, while avoiding sensory and tolerability limitations commonly associated with higher concentrations (2,3,17). The synergistic interaction between neem oil, tea tree oil, fenugreek, and tulsi extracts resulted in broad-spectrum antifungal activity comparable to standard synthetic agents, reinforcing the therapeutic relevance of phytochemical synergy over single-compound strategies (18–20).

Equally significant was the formulation's stability and safety profile, achieved through the integration of natural antioxidants and stabilizers that effectively prevented oxidative degradation and phase instability, thereby addressing a critical limitation of traditional herbal oil formulations (21,22). The confirmed dermal non-irritancy, sustained cooling sensation, and enhanced hair growth response underscore the formulation's suitability for long-term use in chronic scalp conditions that demand both efficacy and comfort (14,15,23). Importantly, the exclusive use of plant-based, GRAS-compliant excipients aligns the formulation with contemporary regulatory expectations and sustainability-driven cosmetic and pharmaceutical development paradigms (6,24). In conclusion, this antifungal herbal hair oil represents a robust, safe, and consumer-friendly therapeutic cosmetic system with strong translational potential. Its multifunctional benefits—antifungal, antioxidant, anti-inflammatory, cooling, and hair growth-promoting—position it as a promising candidate for further clinical evaluation, intellectual property protection, and commercialization within the rapidly expanding herbal and dermo-cosmetic market (25).

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