



REVIEW

Microneedling and Its Use in Hair Loss Disorders: A Systematic Review

Robert S. English Jr. · Sophia Ruiz · Pedro DoAmaral

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ABSTRACT

Introduction: Microneedling (MN) is a minimally invasive procedure involving the induction of percutaneous wounds with medical-grade needles. In this literature review, we investigate clinical data on MN for the treatment of hair loss disorders.

Methods: A literature search was conducted through PubMed up to November 2021 to identify original articles evaluating the use of MN on hair loss disorders. The database was searched using the following keywords: “microneedling,” “micro needling,” “micro needle,” “microneedle,” “needle,” “dermaroller” and “alopecia,” “hair loss,” “alopecia,” “areata,” “cicatrical,” or “effluvium.”

Results: A total of 22 clinical studies featuring 1127 subjects met our criteria for inclusion. Jadad scores ranged from 1 to 3, with a mean of 2. As an adjunct therapy, MN improved hair parameters across genders and a range of hair loss types, severities, needling devices, needling depths of 0.50–2.50 mm, and session frequencies from once weekly to monthly. Across 17

investigations totaling 911 androgenic alopecia (AGA) subjects, MN improved hair parameters when paired with 5% minoxidil, growth factor solutions, and/or platelet-rich plasma (PRP) topicals, or when introduced to subjects whose hair count changes had plateaued for ≥ 6 months on other treatments. Across four investigations on 201 alopecia areata (AA) subjects, MN improved hair parameters as a standalone therapy versus cryotherapy, as an adjunct to 5-aminolevulinic acid and photodynamic therapy, and equivalently when paired with topical PRP versus carbon dioxide laser therapy with topical PRP. Across 657 subjects receiving MN, no serious adverse events were reported.

Conclusions: Clinical studies demonstrate generally favorable results for MN as an adjunct therapy for AGA and AA. However, data are of relatively low quality. Significant heterogeneity exists across interventions, comparators, and MN procedures. Large-scale randomized controlled trials are recommended to discern the effects of MN as a standalone and adjunct therapy, determine best practices, and establish long-term safety.

Supplementary Information The online version contains supplementary material available at <https://doi.org/10.1007/s13555-021-00653-2>.

Keywords: Microneedling; Alopecia; Hair loss

R. S. English Jr. (✉) · S. Ruiz · P. DoAmaral
Perfect Hair Health, 2021 Fillmore, Ste 98, San
Francisco, CA 94115, USA
e-mail: rob@perfecthairhealth.com

Key Summary Points

Why carry out this study?

There is growing interest in the use of microneedling as a standalone and adjunct therapy for hair loss disorders.

This literature reviews summarizes a body of clinical evidence on microneedling for hair loss disorders to evaluate hair loss outcomes, evidence quality, limitations in research, and areas of opportunity for future investigations.

What was learned from the study?

Microneedling improves hair loss parameters across a range of hair loss types, needling devices, needling depths, session frequencies, and combination therapies.

While evidence suggests that microneedling might improve hair loss, clinical data are of relatively low quality. With better study designs and efforts to standardize best practices, microneedling could become a staple adjuvant to US Food and Drug Administration (FDA)-approved hair loss treatments.

INTRODUCTION

Alopecia is a common cosmetic concern affecting over 50% of adults throughout a lifetime [1]. Hair loss disorders are typically categorized into scarring and nonscarring alopecias, with treatments dependent on the pathogenesis and diagnosis determined during dermatological evaluation [2]. While drug and nondrug interventions often help to improve many hair loss disorders, treatments for androgenic alopecia (AGA) are typically relegated to stopping the progression of the condition [3]. Moreover, treatments for alopecia areata (AA) and alopecia totalis (AT) remain limited, with recurrence

rates high [4]. Consequently, there remains demand for novel and effective hair loss treatments.

Microneedling (MN) is a minimally invasive procedure involving the induction of percutaneous wounds with 0.25–5.00 mm medical-grade needles. First described by Orentreich in 1995 for the use of wrinkles and atrophic scars, MN purportedly releases platelet-derived growth factor and vascular endothelial growth factor to promote wound-healing responses, improve angiogenesis, and attenuate or partially reverse fibrosis resulting from acute injury and skin aging [5, 6]. MN can be administered at-home or in-clinic, with devices ranging from needling stamps, manual rollers, and automated pens with or without fractional radiofrequency. Across a range of devices, needling depths, and session frequencies, MN has demonstrated clinical improvements as a standalone and/or adjunct therapy for patients with atrophic scars, actinic keratoses, and pigmentation disorders such as vitiligo and melasma [6, 7].

In the last decade, studies have demonstrated that MN may enhance transdermal delivery, promote anagen-initiating Wnt/ β -catenin signaling, and improve dermal papillae stem cell proliferation—thus potentiating therapeutic effects for a variety of hair loss disorders [8–10]. In 2013, the landmark study by Dhurat et al. on 100 AGA subjects found that over a 12-week period, once-weekly MN combined with twice-daily 5% minoxidil increased hair counts significantly versus minoxidil monotherapy [11]. Since then, investigators have continued to assess the effects of MN as both a standalone and adjunct therapy for hair loss.

In this systematic review, we investigate the use of MN as a standalone, adjunct, and comparator therapy on hair loss disorders. We evaluate patient populations, interventions, comparators, MN procedures, outcomes, and adverse events—as well as evidence quality using Jadad scoring. We discuss possible mechanisms by which MN may improve hair loss disorders as a monotherapy and an adjunct intervention. Finally, we identify limitations in

the current body of research and provide recommendations for future clinical trials.

METHODS

Literature Search

A broad literature search was conducted through PubMed up to November 2021 to identify original articles that evaluate the use of MN on hair loss disorders. The database was searched using combinations of the following keywords: “microneedling,” “micro needling,” “micro needle,” “microneedle,” “needle,” “dermaroller” and “alopecia,” “hair loss,” “alopecia,” “areata,” “cicatricial,” or “effluvium.” This article is based on previously conducted studies and does not contain any new studies with human participants or animals performed by any of the authors.

Inclusion and Exclusion Criteria

All search hits were screened and examined for relevant titles and abstracts. Full texts were reviewed to determine eligibility. Articles were included if they featured all of the following: (a) human subjects with scalp hair loss, (b) MN as a standalone or adjunct therapy, and (c) endpoint measurements related to scalp hair. Articles were excluded if they did not feature (a) original data, (b) human data, (c) endpoint measurements for hair parameters, and/or (d) designs that adequately evaluated the effects of MN on hair. This literature review is based on previously conducted studies and does not contain any new studies with human participants or animals performed by any of the authors. Full inclusion and exclusion criteria can be found in Table 1.

RE and SR each independently identified 367 records for screening. RE, SR, and PD each independently screened all 367 titles and abstracts to assess eligibility, and the 42 full texts to determine inclusion. RE, SR, and PD each independently assessed Jadad scores. Any disagreements in identifications, screenings,

Table 1 PICOS inclusion and exclusion criteria

Parameter	Inclusion criteria	Exclusion criteria
Patients	Patients of any age treated for scalp hair loss	
Intervention	MN as a standalone or adjunct therapy	MN devices with needle-releasing drugs, acupuncture needles
Comparator	How effective is MN at improving hair loss outcomes?	
Outcomes	<i>Primary endpoints:</i> phototrichogram, investigator, and/or patient assessments	Any study not designed to adequately test for the standalone or additive effect of MN
Study design	Prospective studies	Retrospective design, case series, literature reviews, or nonhuman subjects; studies with fewer than five patients; ongoing clinical trials; Jadad scores lower than 1

A table summarizing the inclusion and exclusion criteria in our systematic review for clinical studies investigating the use of MN for the treatment of hair loss disorders

selections, and/or Jadad scores were discussed by RE and PD and resolved by RE.

RESULTS

Of the 42 full texts accessed to assess eligibility, 20 were excluded on the basis of the wrong intervention ($n = 2$), outcome ($n = 4$), or study

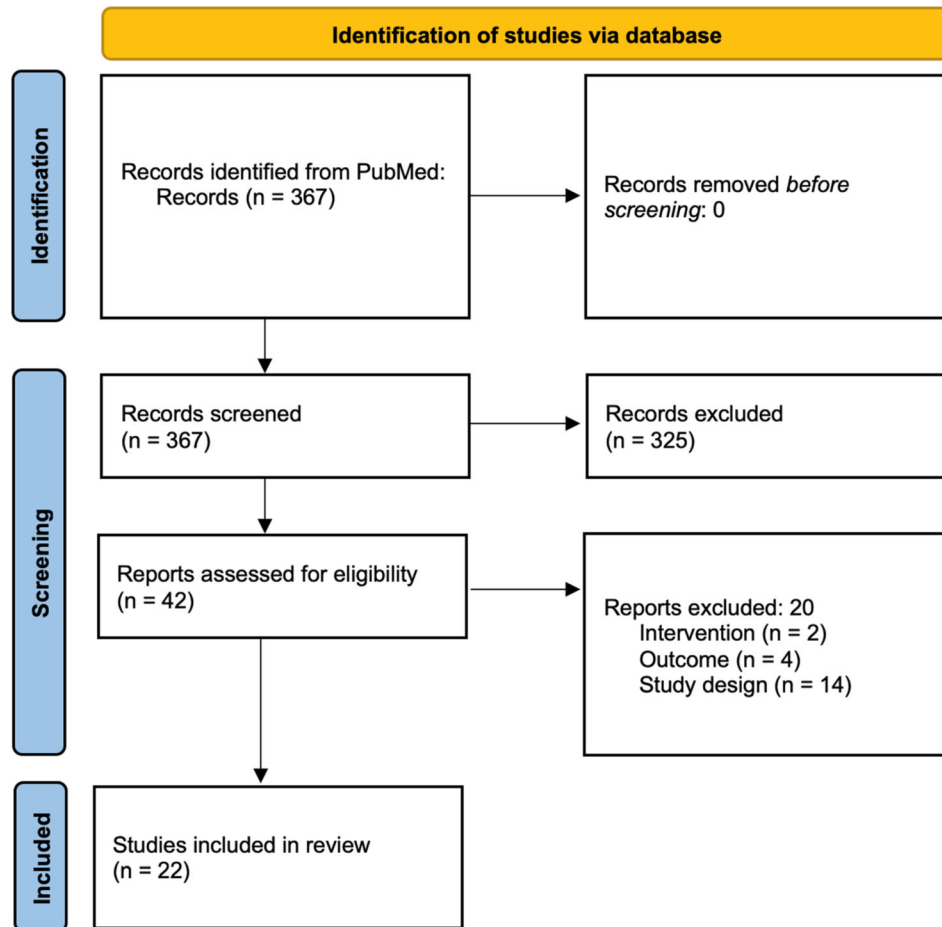


Fig. 1 PRISMA flowchart. A PRISMA flowchart detailing the process of eligibility for all records reviewed for the literature review, as well as the number of studies identified, screened, excluded, and included

design ($n = 14$). A total of 22 clinical studies met our criteria for inclusion: 17 trials with randomization and 5 nonrandomized prospective cohorts (Fig. 1). Jadad scores ranged from 1 to 3, with a mean score of 2 (Table S1).

Of the 22 studies, 16 were conducted on AGA subjects, 4 on alopecia areata subjects, 1 on alopecia totalis subjects, and 1 on both AGA and telogen effluvium (TE) subjects. A total of 1127 subjects (856 males and 269 females) were included featuring the following hair loss types: AGA ($n = 911$), AA ($n = 201$), AT ($n = 8$), and TE ($n = 7$) [11–32].

AGA

Patients

Within studies featuring AGA subjects, enrollment ages ranged from 18 to 70 years, with a subject-weighted average of 33.75 years. Of the 15 studies with male AGA subjects, 1 did not enroll subjects based on a classification system, while 14 included males with hair loss based on the Norwood–Hamilton scale: I (0.0%), II (46.7%), III (93.3%), IV (93.3%), V (60%), and VI (26.7%). Of the seven studies with female AGA subjects, one did not enroll subjects based on any classification system, one enrolled based on Sinclair scores, and five enrolled females with hair loss determined by the Ludwig scale: I (80.0%), II (60.0%), and III (60.0%) (Table 2).

Table 2 Parameter summaries for studies assessing the use of MN on AGA subjects

Androgenic alopecia (AGA)										
Author (year)	Total subjects (sex); alopecia type	Study type	Treatment regimen	MN procedure	No. of MN sessions	Treatment duration	Endpoints	Effectiveness	Adverse events	Jadad score
Ramadan et al. [12]	n = 126 (46 m, 80 f); AGA	Combination: PRP, 5% minoxidil, 2.50 mg finasteride (men), 200 mg spironolactone (women)	Group 1: medications + PRP injections Group 2: medications + MN + topical PRP Group 3: medications	MN automated pen, 2.00 mm needles, session endpoints marked as three passes followed by PRP application	6, once every month	6 months	Hair counts, hair diameters, photographic evaluation	Yes, in both sexes Hair density increased in group 2 versus group 1; larger effect for groups 1 and 2 versus group 3	No serious events reported Twenty-three subjects reported transient pain after PRP	2
Sohng et al. [13]	n = 29 (24 m, 5 f); AGA	Combination: 5% minoxidil	Group 1: home-use MN Group 2: home-use MN + 5% minoxidil Group 3: 5% minoxidil	MN stamp, 0.25 mm spiral needles, endpoint sessions marked as gentle tapping 20 times in target area	52, twice weekly	6 months	Hair counts, self assessments	No	No serious events reported; mild and transient pruritus noted in one subject	2
Burns et al. [14]	n = 11 (f); AGA	Combination: 5% minoxidil + MN added to ongoing hair loss treatments	5% minoxidil + MN twice per month added to ongoing hair loss treatments	MN automated pen, session endpoints marked as two passes across the frontal, crown, vertex, and upper-parietal scalp	6, twice every month	3 months	Photographic evaluation, self assessments	Yes Investigators noted that 11/11 subjects improved at least 1–1.5 Sinclair scores	No serious events reported	1
Gowda et al. [15]	n = 90 (m); AGA	Combination: 5% minoxidil, PRP	Group 1: 5% minoxidil Group 2: 5% minoxidil + MN Group 3: 5% minoxidil + PRP injections	MN roller, 1.50 mm needles, session endpoints marked as passes longitudinally, vertically, and diagonally until pinpoint bleeding noted	4, once every month	4 months	Hair counts, photographic evaluation	Yes Hair counts increased in groups 1, 2, and 3; investigators noted improvements in group 3	No serious events reported in MN group	1

Table 2 continued

Author (year)	Total subjects (sex); alopecia type	Study type	Treatment regimen	MN procedure	No. of MN sessions	Treatment duration	Endpoints	Effectiveness	Adverse events	Jadad score
Shome et al. [16]	<i>n</i> = 50 (25 m, 25 f); AGA	Combination: intradermal versus MN delivery of growth factor solution	Group 1: intradermal QR678 Neo(®) Group 2: MN + QR678 Neo(®) topical	MN roller, 1.50 mm needles, session endpoints marked as 4–5 passes longitudinally, vertically, and diagonally until light erythema noted	8, once every 3 weeks	12 months	Hair counts, hair diameters, photographic evaluation, self assessments	Yes Hair counts and hair diameters increased in groups 1 and 2; no difference in changes to hair counts and diameters in group 1 versus group 2	No serious events reported; transient scalp itchiness higher in group 2	2
Ozcan et al. [17]	<i>n</i> = 62 (m); AGA	Combination: PRP solution	Group 1: PRP injections Group 2: MN + PRP solution	MN automated pen, 1.50 mm needles	4, once every two weeks then once after 1 month	10 weeks	Hair counts, hair diameters, pull tests, photographic evaluation, self assessments	Yes Hair counts and hair diameters increased in groups 1 and 2; changes to anagen:telogen hairs increased in group 2 versus group 1	No events reported	2
Yu et al. [18]	<i>n</i> = 40 (m); AGA	Combination: 5% minoxidil, growth factors	Group 1: saline + MN Group 2: 5% minoxidil + MN Group 3: growth factors + MN Group 4: 5% minoxidil + growth factors + MN	MN roller	16, once weekly	16 weeks	Hair counts, hair diameters, photographic evaluation, self assessments	Yes Hair density increased in groups 2, 3, and 4	No serious events reported Three subjects developed mild erythema which alleviated after 24 hours	3

Table 2 continued

Androgenic alopecia (AGA)										
Author (year)	Total subjects (sex); alopecia type	Study type	Treatment regimen	MN procedure	No. of MN sessions	Treatment duration	Endpoints	Effectiveness	Adverse events	Jadad score
Bao et al. [19]	n = 75 (m); AGA	Combination: 5% minoxidil	Group 1: 5% minoxidil Group 2: MN Group 3: 5% minoxidil + MN	MN automated pen, 1.00–2.00 mm needles, session endpoints marked as hemorrhage and redness	8, once every 3 weeks	24 weeks	Hair counts, hair diameters, photographic evaluation	Yes Hair counts increased in groups 1, 2, and 3; hair density increased in groups 2 and 3; larger improvements for group 3 versus groups 1 and 2	Twelve events related to scalp irritation were noted; all resolved within 4 days; no differences in occurrence across groups	3
Aggarwal et al. [20]	n = 30 (m); AGA	Split scalp: MN, PRP	Side 1: MN Side 2: MN + PRP injections	MN roller, 1.50–2.00 mm needles, session endpoints marked as gentle rolling until pinpoint bleeding	4, once monthly	6 months	Hair counts, hair diameters, self assessments	Yes Hair counts and hair diameters increased in sides 1 and 2; no difference in change of hair counts and hair diameters in side 1 versus side 2	No serious events reported	3

Table 2 continued

Androgenic alopecia (AGA)		Study type		Treatment regimen		MN procedure		No. of MN sessions		Treatment duration		Endpoints		Effectiveness		Adverse events		Jadad score	
Author (year)	Total subjects (sex); alopecia type	Combination: minoxidil	Study type	Treatment regimen	Treatment regimen	MN procedure	MN procedure	No. of MN sessions	Treatment duration	Endpoints	Effectiveness	Adverse events	Jadad score						
Faghghi et al. [21]	<i>n</i> = 59 (29 m, 30 f); AGA	Combination: 5% minoxidil	Combination: minoxidil	Group 1: 5% minoxidil Group 2: 5% minoxidil + MN 1.2 mm Group 3: 5% minoxidil + MN 0.6 mm	Group 1: 5% minoxidil Group 2: 5% minoxidil + MN 1.2 mm Group 3: 5% minoxidil + MN 0.6 mm	MN automated pen, 1.20 or 0.60 mm needles, session endpoints marked as pinpoint bleeding	MN automated pen, 1.20 or 0.60 mm needles, session endpoints marked as pinpoint bleeding	6, once every 2 weeks	12 weeks	Hair counts, hair diameters, photographic evaluation, self assessments	Yes Hair counts and diameters increased in groups 1, 2, and 3; change in hair counts and diameters greater in group 3 versus group 1; change in hair diameters greater in group 2 versus group 1	More pain reported from MN in group 2 versus 3	2						
Starace et al. [22]	<i>n</i> = 50 (14 m, 36 f); AGA, TE	Combination: MN added to ongoing hair loss treatments	Combination: MN added to ongoing hair loss treatments	MN every 3 weeks added to ongoing hair loss treatments	MN every 3 weeks added to ongoing hair loss treatments	MN roller, 1.50 mm needles, 20–25 minute sessions, session endpoints marked as eight passes longitudinally, vertically, and diagonally in affected regions or until mild erythema and pinpoint bleeding noted	MN roller, 1.50 mm needles, 20–25 minute sessions, session endpoints marked as eight passes longitudinally, vertically, and diagonally in affected regions or until mild erythema and pinpoint bleeding noted	3, once every 4 weeks	6 months	Hair counts, hair diameters, photographic evaluation, self assessments	AGA: yes, in both sexes TE: yes Hair counts and hair diameters increased	No serious events reported	1						

Table 2 continued

Androgenic alopecia (AGA)										
Author (year)	Total subjects (sex); alopecia type	Study type	Treatment regimen	MN procedure	No. of MN sessions	Treatment duration	Endpoints	Effectiveness	Adverse events	Jadad score
Kumar et al. [23]	<i>n</i> = 60 (m); AGA	Combination: 5% minoxidil	Group 1: 5% minoxidil Group 2: 5% minoxidil + MN	MN roller, 1.50 mm needles, session endpoints marked as passes longitudinally, vertically, and diagonally in affected regions until pinpoint bleeding noted	8; four sessions (month 1), two sessions (month 2), two sessions (month 3)	12 weeks	Hair counts, photographic evaluation, self assessments	Yes Hair counts increased in group 2 versus group 1; higher self assessment scores in group 2 versus group 1	No serious events reported	2
Yu et al. [24]	<i>n</i> = 19 (m); AGA	Split scalp: 5% minoxidil	Side 1: 5% minoxidil Side 2: 5% minoxidil + fractional radiofrequency MN	Fractional radiofrequency MN, 1.50 mm needles	5, once every 4 weeks	5 months	Hair counts, hair diameters, photographic evaluation, self assessments	Yes Hair counts and hair diameters increased in sides 1 and 2; changes to hair counts and diameters greater in side 2 versus 1	No serious events reported	3
Bao et al. [25]	<i>n</i> = 60 (m); AGA	Combination: 5% minoxidil	Group 1: 5% minoxidil Group 2: MN Group 3: 5% minoxidil + MN	MN automated pen, 1.50–2.50 mm needles, session endpoints marked as 3–4 passes until redness and/or hemorrhaging noted	12, once every 2 weeks	24 weeks	Hair counts, hair diameters, photographic evaluation, self assessments	Yes Hair counts increased in groups 1, 2, and 3; hair counts, hair diameters, photographic evaluations, and self assessments better in group 3 versus groups 1 and 2	No serious events reported; no differences in adverse event reporting across groups	3

Table 2 continued

Androgenic alopecia (AGA)		Study type		Treatment regimen		MN procedure		No. of MN sessions		Treatment duration		Endpoints		Effectiveness		Adverse events		Jadad score	
Author (year)	Total subjects (sex); alopecia type	Study type	Treatment regimen	MN procedure	No. of MN sessions	Treatment duration	Endpoints	Effectiveness	Adverse events	Jadad score									
Shah et al. [26]	<i>n</i> = 50 (m); AGA	Combination: 5% minoxidil, PRP	Group 1: 5% minoxidil Group 2: 5% minoxidil + MN + PRP injections	MN roller, 1.50 mm needles, session endpoints marked as eight passes longitudinally, vertically, and diagonally in affected regions or until mild erythema	6, once monthly	6 months	Photographic evaluation, self assessments	Yes Photographic evaluations better in group 2 versus group 1	No events reported	2									
Dhurat et al. [11]	<i>n</i> = 100 (m); AGA	Combination: 5% minoxidil	Group 1: 5% minoxidil Group 2: 5% minoxidil + MN	MN roller, 1.50 mm needles, session endpoints marked as passes longitudinally, vertically, and diagonally in affected regions until mild erythema	12, once weekly	3 weeks	Hair counts, photographic evaluation, self assessments	Yes Hair counts increased in groups 1 and 2; hair counts increased in group 1 versus 2; higher photographic evaluations and self assessments in group 1 versus 2	No serious events reported	3									
Lee et al. [27]	<i>n</i> = 11 (f); AGA	Split scalp: growth factor solution	Side 1: growth factor solution + MN Side 2: saline + MN	MN automated pen, 0.50 mm needles	5, once weekly	5 weeks	Hair counts, self assessments	Yes Hair counts increased in side 1 versus side 2	No events reported	1									

A table summarizing all included studies assessing the use of MN on AGA subjects. Each study summary features parameters regarding author, year of publication, total subjects, gender, study type, treatment regimen, MN procedure, number of MN sessions, treatment duration, assessment endpoints, effectiveness, adverse events, and Jadad score

Interventions and Comparators

In total, 536 subjects received MN therapy, while 375 received other hair loss interventions. Across all study groups, MN was included as a standalone therapy in 6 groups ($n = 105$). As an adjunct therapy, MN was evaluated alongside topical minoxidil in 10 groups ($n = 234$), proprietary topicals and/or growth factor solutions in 3 groups ($n = 46$), PRP in 2 groups ($n = 61$), continued medication use in 1 group ($n = 50$), PRP and systemic medications in 1 group ($n = 42$), PRP with topical minoxidil in 1 group ($n = 25$), topical minoxidil and continued medications in 1 group ($n = 11$), and topical minoxidil alongside growth factors in 1 group ($n = 10$) (Table 2).

MN Procedures

MN devices tested included manual rollers ($n = 8$), automated pens ($n = 7$), manual stamps ($n = 1$), and automated fractional radiofrequency pens ($n = 1$). Needle lengths ranged from 0.25 to 2.50 mm, with a mean needle length of 1.39 mm.

The frequency of MN sessions ranged from twice weekly to once monthly, with a mean session frequency of once per 2.64 weeks. The number of MN sessions ranged from 3 to 52, with a mean of 9.53 MN sessions per study. Treatment durations averaged 20.01 weeks.

While four studies did not specify MN session endpoints, 13 studies standardized endpoints by a number of passes, directions, and/or taps ($n = 3$), mild erythema ($n = 3$), a number of passes and/or bleeding ($n = 3$), or passes until hemorrhage ($n = 4$) (Table 2).

Outcomes

In total, 15 of 17 studies assessed hair parameters through phototrichograms (i.e., hair counts, hair diameters, and/or hair densities). Of the 17 studies, 6 included MN-only groups, whereas all studies tested MN alongside other hair loss interventions.

Of the six MN monotherapy groups, two noted significant increases to total hair counts, one found significant increases to hair diameters and total hair density, and three showed no effect [13, 18–20, 25, 27].

Of the seven studies testing MN with 5% minoxidil, six found statistically significantly increased hair counts versus 5% minoxidil alone, and for a range of devices and needle lengths: rollers, automated pens, and fractional radiofrequency devices with depths from 0.60 to 2.50 mm [11, 15, 18, 19, 21, 23, 25]. However, Sohng et al. tested 5% minoxidil with a 0.25 mm needling stamp twice-weekly and found no effect on hair parameters [13]. When comparing 5% minoxidil with 0.60 mm or 1.20 mm needle lengths, Faghihi et al. found that 0.60 mm needle lengths led to significant hair count and diameter increases versus 5% minoxidil, whereas 1.20 mm needle lengths only saw hair count increases versus 5% minoxidil [21].

All three studies testing MN alongside proprietary topicals and/or growth factors noted increases to hair counts [16, 18, 27]. Lee et al. and Yu et al. demonstrated improved hair parameters but no differences across groups when comparing MN use with topical versus intradermal delivery of proprietary products and/or growth factors [18, 27]. Conversely, Ozcan et al. found that MN alongside either topical or injectable PRP significantly increased hair counts and diameters similarly across groups, but that subjects receiving MN alongside topical PRP saw greater improvements to anagen:telo-gen hairs [17]. Interestingly, in a split-scalp study, Aggarwal et al. showed that both MN and MN with PRP injections increased hair diameters and density equivalently—with no significant differences noted across groups [20].

Two studies tested the introduction of MN alongside ongoing hair loss medications [14, 22]. Burns et al. found that twice monthly MN combined with 5% minoxidil improved Ludwig scores for 11/11 females who had previously plateaued for ≥ 6 months on other hair loss treatments [14]. Starace et al. showed that the addition of MN improved hair counts in those already using hair loss treatments for > 1 year (Table 2) [22].

Adverse Events

Across 536 subjects receiving MN, no serious adverse events were reported. Of mild adverse

events, transient pain, scalp irritation, and mild erythema were most commonly reported. Withdrawal rates across MN groups were low and comparable to non-MN groups.

AA and AT

Patients

Of the five studies with AA and AT subjects, enrollment ages ranged from 16 to 45 years, with a subject-weighted average of 28.34 years. Three investigations enrolled subjects with hair loss gradients according to Severity Of Alopecia Tool (SALT) score, one study enrolled on the basis of severe AA, and one study enrolled on the basis of AT (Table 3).

Interventions and Comparators

In total, 114 subjects received MN therapy while 95 received other hair loss interventions. Of the five studies featuring AA and AT subjects, three compared treatments across patients ($n = 181$), while two compared treatments across lesions within the same patients ($n = 28$).

Across all study groups, MN was included as a standalone therapy in three groups ($n = 68$). As an adjunct therapy, MN was evaluated alongside a PRP topical in one group ($n = 20$), and with 5-aminolevulinic acid or methyl 5-aminolevulinic acid alongside photodynamic therapy in two groups ($n = 25$). As a comparator, MN was included as a control against cryotherapy in one group ($n = 40$), PRP injections in one group ($n = 20$), fractional CO₂ laser alongside a PRP topical in one group ($n = 20$), triamcinolone acetonide injections in one group ($n = 20$), 5% minoxidil injections in one group ($n = 20$), and 5-aminolevulinic acid or methyl 5-aminolevulinic acid alongside photodynamic therapy in two groups ($n = 23$) (Table 3).

MN Procedures

MN devices tested included manual rollers ($n = 3$) and automated pens ($n = 2$). Needle lengths ranged from 1.00 to 5.00 mm, with a mean needle length of 2.25 mm.

The frequency of MN sessions ranged from once every 2 weeks to once monthly, with a

mean session frequency of once every 3.46 weeks. The number of MN sessions ranged from three to six, with a mean of 4.40 MN sessions per study. Treatment durations averaged 14.20 weeks.

While one study did not specify MN session endpoints, four studies standardized endpoints by a number of passes ($n = 2$), a number of passes and/or mild erythema ($n = 1$), or minutes of passes in affected areas ($n = 1$) (Table 3).

Outcomes

In total, two of five studies assessed hair parameters through objective measurements (i.e., phototrichograms or 4 mm punch biopsies). Subjective measurements for the remaining three studies included Severity of Alopecia Tool (SALT) scores ($n = 2$), Lesional Area & Density (LAD) scores ($n = 1$), and/or a four-point scale ($n = 1$).

Of the three studies testing standalone MN groups, Aboeldahab et al. and Abdallah et al. showed that MN alone increased hair density and improved SALT scores, respectively [28, 30]. However, Giorgio et al. demonstrated no changes to hair parameters using a four-point scale to evaluate MN alone [31].

When comparing MN with cryotherapy, Aboeldahab et al. found significant increases to hair counts and hair densities across both interventions, with MN demonstrating greater changes to SALT scores versus cryotherapy [28]. Conversely, Abdallah et al. found that triamcinolone acetonide injections with and without 5% intradermal minoxidil led to greater improvements to SALT and LAD scores versus controls than MN alone [30].

As an adjunct therapy, Giorgio et al. showed that MN alongside 5-aminolevulinic acid and photodynamic therapy improved 94% of AA lesions versus 53% of lesions receiving only 5-aminolevulinic acid and photodynamic therapy [31]. However, Yoo et al. found that in AT subjects, methyl 5-aminolevulinic acid and photodynamic therapy with and without MN led to no hair parameter improvements according to 4 mm punch biopsies [32].

Finally, Ragab et al. demonstrated that, over a 3-month period, MN alongside topical PRP improves SALT scores similarly to fractional

Table 3 Parameter summaries for studies assessing the use of MN on AA and AT subjects

Author (year)	Total subjects (gender); alopecia type	Study type	Treatment regimen	MN procedure	No. of MN sessions	Treatment duration	Endpoints	Effectiveness	Adverse events	Jadad score
Aboeldahab et al. [28]	n = 80 (50 m, 30 f); alopecia areata (AA)	Comparison: cryotherapy versus MN	Group 1: cryotherapy Group 2: MN	MN automated pen, 1.00–2.00 mm needles, session endpoints marked as 4–5 passes longitudinally, vertically, and diagonally	6, once every 2 weeks	12 weeks	Hair counts, hair density, Severity of Alopecia Tool (SALT) scores, photographic evaluation, and self assessments	Yes Hair counts, hair density, SALT scores, photographic evaluation, and self assessments improved in groups 1 and 2; higher changes to SALT scores in group 2 versus 1	No adverse events in group 2	3
Ragab et al. [29]	n = 60 (48 m, 12 f); AA	Combination: PRP solution	Group 1: PRP injections Group 2: fractional CO ₂ laser + PRP topical Group 3: MN + PRP topical	MN roller, 1.50 mm needles, session endpoints marked as 4–5 passes longitudinally, vertically, and obliquely until mild erythema noted	3, once monthly	3 months	SALT	Yes SALT scores improved in groups 1, 2, and 3; no difference in SALT scores across groups	No serious events reported; pain higher in group 1 versus group 3	3
Abdallah et al. [30]	n = 20 (19 m, 1 f); AA	Inpatient comparison: triamcinolone acetamide injections, minoxidil 5% injections	Patch 1: triamcinolone acetamide injections Patch 2: 5% minoxidil injections Patch 3: triamcinolone acetamide injections + 5% minoxidil injections Patch 4: MN Patch 5: control	MN roller, session endpoints marked as 4–5 passes longitudinally, vertically, and diagonally	4, once every 4 weeks	16 weeks	SALT, Lesional Area & Density (LAD) scores	Yes SALT and LAD scores decreased in patches 1, 2, 3, and 4; largest differences seen in patches 1 and 3 versus patch 5	No serious events reported	2

Table 3 continued

Alopecia areata (AA), alopecia totalis (AT)										
Author (year)	Total subjects (gender); alopecia type	Study type	Treatment regimen	MN procedure	No. of MN sessions	Treatment duration	Endpoints	Effectiveness	Adverse events	Jadad score
Giorgio et al. [31]	<i>n</i> = 41 (17 m, 24 f); AA	Combination: 5-aminolevulinic acid photodynamic therapy	Group 1: MN Group 2: 5-aminolevulinic acid + photodynamic therapy Group 3: MN + 5-aminolevulinic acid + photodynamic therapy	MN automated pen, 1.00 mm needles, session endpoints marked as 5 minute of passes on AA-affected areas	6, once every 3 weeks	18 weeks	4-point scale	Yes No change in group 1; improvements noted in 53% of group 2 and 94% of group 3	No events reported	1
Yoo et al. [32]	<i>n</i> = 8 (2 m, 6 f); alopecia totalis (AT)	Split scalp: 5-aminolevulinic acid + photodynamic therapy	Side 1: MN + methyl 5-aminolevulinic acid + photodynamic therapy Side 2: methyl 5-aminolevulinic acid + photodynamic therapy	MN roller, 5.00 mm needles	3, once every 4 weeks	12 weeks	Histological assessments (4 mm punch biopsy)	No	No serious events reported	1

A table summarizing all included studies assessing the use of MN on AA and AT subjects. Each study summary features parameters regarding author, year of publication, total subjects, gender, study type, treatment regimen, MN procedure, number of MN sessions, treatment duration, assessment endpoints, effectiveness, adverse events, and Jadad score

CO₂ laser therapy alongside topical PRP (Table 3) [29].

Adverse Events

Among 114 subjects receiving MN, no serious adverse events were reported. Of mild adverse events, transient pain and mild erythema were most common. Of the five studies, no withdrawals were reported in MN or non-MN groups.

DISCUSSION

Clinical studies demonstrate generally favorable results for MN as an adjunct therapy for AGA and AA. However, data are of relatively low quality and should be interpreted with caution. Due to significant heterogeneity across interventions, comparators, and MN procedures (i.e., devices, needle lengths, session frequencies, and session endpoints), we could not conduct a meta-analysis. Here we discuss the proposed mechanisms of MN, limitations in the current body of research, and design considerations for future studies.

Mechanisms

AGA

In AGA-affected hair follicles, dihydrotestosterone dysregulates the Wnt/ β -catenin pathway, induces transforming growth factor β 1, and triggers apoptosis in dermal papillae and epithelial cells. This leads to a shortened anagen phase, reductions to dermal papillae cell cluster sizes with each re-entry into anagen and, consequently, microvascular degradation alongside progressive hair follicle miniaturization [33–36]. In mid-to-late stages of miniaturization, perifollicular fibrosis is often observed and may reduce the effectiveness of both systemic and topical AGA treatments [37].

As a monotherapy for AGA, data on MN are limited, and the mechanisms by which MN might improve AGA remain speculative. In a pooled linear regression across six subgroups, Gupta et al. found that MN significantly increased total hair counts, by more than 5%

topical minoxidil [38]. However, two of the subgroups were a part of split-scalp studies assessing MN versus PRP injections or topical growth factor solutions [20, 27]. Therefore, the possibility of percutaneous treatment diffusion across scalp zones cannot be discounted.

Animal models suggest that MN may promote anagen-initiating Wnt/ β -catenin signaling and dermal papillae stem cell proliferation. In particular, percutaneous wounds from MN appear to activate hair follicle stem cells, platelet-derived growth factor, and vascular endothelial growth factor—potentiating the initiation of angiogenesis, neocollagenesis, and a new anagen cycle [5–7, 9, 10]. Clinical data show that MN reduces scarring and improves the density and thickness of epidermal and dermal skin layers [39, 40]. In two randomized controlled clinical trials, Bao et al. found that MN alone increased terminal hair counts—with their latter study analyzing biopsies from a subset of AGA subjects showing that MN alone upregulated protein levels of both FZD3 and LEF-1 but not β -catenin [19, 25]. However, RT-PCR testing revealed no statistical increases to mRNA expression—with the authors postulating that the inconsistent results might be due to small sample sizes, infrequent needling sessions, shallow needling depths, and/or post-transcriptional modifications [25].

As an adjunct therapy, MN may improve AGA by enhancing transdermal delivery, and by improving sulfation and Wnt pathway expression when paired with topical minoxidil. Henry et al. demonstrated in vitro that 0.15 mm needles inserted into human skin for 10 seconds enhanced transdermal permeability of calcein by more than 1000-fold [41]. These effects may partly explain the equivalent and/or additive improvements to hair parameters from intradermal growth factors or PRP injections versus MN alongside their topical applications [16, 17]. Additionally, MN may enhance topical minoxidil activation. Topical minoxidil is a pro-drug that requires sulfation by sulfotransferase enzymes in the outer root sheath of hair follicles [8]. Goren et al. demonstrated that reduced sulfotransferase activity in hair follicles predicted topical minoxidil nonresponders [42]. More recently, Sharma et al. found that, over

21 days, once-weekly microneedling led to a median increase in sulfotransferase activity of 37.5% [8]. Finally, Bao et al. demonstrated that MN with 5% minoxidil upregulated the expression of FZD3, LEF-1, and β -catenin in mRNA and protein more than MN or 5% minoxidil monotherapy—suggesting that the addition of MN might amplify the effects of minoxidil on the Wnt pathway [19].

AA and AT

AA and AT are autoimmune forms of alopecia resulting from the collapse of immune privilege in affected hair follicles. In particular, peribulbar lymphocytic infiltrates appear to induce apoptosis in hair follicle keratinocytes—leading to inflammation, impaired hair shaft production, and sometimes hair shaft miniaturization within the same hair cycle. While the histologic features of AA and AT are well studied, their underlying pathogenesis remain poorly understood—with researchers speculating the involvement of immunological shifts related to genetic and environmental factors [4].

As a monotherapy for AA and AT, data on MN are unrobust. While Aboeldahab et al. and Abdallah et al. noted improvements to AA from MN alone, these results should be interpreted with caution due to relatively small sample sizes, as well as the 50% spontaneous recoveries observed in many AA studies with adequate controls [28, 30]. Moreover, Giorgio et al. found no effect from MN alone in AA subjects [31]. As an adjunct therapy, Giorgio et al. suggested that the release of growth factors from MN may induce immunosuppressive actions that amplify the effects of substances such as 5-aminolevulinic acid [31]. Ragab et al. found that MN alongside topical PRP improved hair parameters similarly to PRP injections, and suggested that MN may also enhance transdermal delivery for AA [29].

Limitations

Due to significant heterogeneity across MN studies regarding interventions, comparators, MN devices, needle lengths, session frequencies, and session endpoints, our systematic review

does not include a meta-analysis and cannot establish best practices for MN procedures.

Faghihi et al. found hair parameters improved more when pairing 5% minoxidil with fortnightly MN using an automated pen with needle lengths of 0.60 mm versus 1.20 mm [21]. Faghihi et al. postulated that puncture depths of 0.60 mm still generate enough of an inflammatory response for stem cell and growth factor recruitment, but without damaging the hair follicle bulge residing 1.00–1.80 mm from the skin surface [43]. Interestingly, Sasaki found that with MN automated pens, needling lengths matched penetration depths up to 1.50 mm [44]. Due to user pressure variability and needle entry angulation, Lima et al. estimated that a 3.00 mm MN manual roller only penetrates to skin depths 50–70% of its needle length [45]. Taken together, equivalent MN penetration depths of 0.60–0.80 mm may be achievable with MN automated pens and MN manual rollers set to needle lengths of 0.60–0.80 mm and 1.25–1.50 mm, respectively. Relatedly, Fernandes postulated that MN device preferences do not matter so long as the skin is penetrated to the same depths [46]. Regardless of standardizations for MN devices or needle lengths, additional methodological considerations—i.e., session durations, frequencies, and endpoints—still likely exert influence over the degree of inflammation induced, and thereby the magnitude of outcomes across a variety of hair parameters. As such, no procedural best practices can be ascertained with the current body of evidence.

While 8 of 22 clinical studies on MN included groups to evaluate MN alone ($n = 174$), only 1 study compared MN monotherapy against an untreated control patch for AA ($n = 20$). Moreover, 21 of 22 clinical studies on MN assessed hair parameter changes over periods of less than 52 weeks. Study durations of less than 52 weeks often do not allow investigators to separate the effects of any intervention against seasonal fluctuations to hair cycling—particularly in the absence of untreated control groups [47, 48].

Finally, 4 out of 22 clinical studies utilized split-scalp study designs to evaluate MN against or as an adjuvant to topicals and/or injectables.

Since MN is suspected to enhance transdermal drug delivery, split-scalp study designs leave open the possibility of percutaneous drug diffusion across scalp zones, thus limiting the interpretability of endpoint assessments.

Recommendations

Large-scale, randomized, placebo-controlled clinical trials assessing the use of MN for hair loss are needed. Future investigations should consider study durations of at least 12 months, include groups for MN as a monotherapy, and evaluate MN against a placebo (i.e., manual rollers with removed needles, automated pens with uninstalled needle cartridges, and/or an untreated control group). Split-scalp studies should be avoided, particularly when evaluating MN against or as an adjuvant to topicals and/or injectables. Finally, studies evaluating the use of MN across different procedural standards (i.e., shorter needle lengths and more frequent sessions versus longer needle lengths and less frequent sessions) will help toward establishing best practices.

CONCLUSION

Among 22 clinical studies featuring 1127 subjects, MN as an adjunct therapy improved hair parameters across genders as well as a range of hair loss types, hair loss severities, needling devices, needling depths of 0.50–2.50 mm, and session frequencies from once weekly to once monthly—with no serious adverse events reported. However, results should be interpreted with caution due to significant heterogeneity across study interventions, comparators, and MN procedures (i.e., devices, needle lengths, session frequencies, and session endpoints). Large-scale randomized controlled trials are needed to discern the effects of MN as a standalone and adjunct therapy, determine best practices for MN procedures, and establish long-term safety data. Study designs should consider 12-month durations, include groups using MN as a monotherapy, and evaluate MN against a placebo and/or untreated group.

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