



Literature Review on Korean Medicine Treatment for Alopecia

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This study aimed to analyze the use of Korean medicine treatments for alopecia in among clinical studies. We identified and analyzed 22 studies from Korean databases; Oriental Medicine Advanced Searching Integrated System (OASIS), Science ON, Korean Studies Information Service System (KISS), and Research Information Sharing Service (RISS) and international database; PubMed. We analyzed the Korean medical treatment in each case and determined the tendency to use each intervention. We analyzed 1,464 patients from 22 selected studies. Herbal medicine, acupuncture, external medicine or products, pharmacopuncture, and phototherapy were used for alopecia treatment. The herbal medicines mainly used to treat alopecia were Gagam Cheongyoung-tang, Gagam Hwajung-hwan, and Yukmijihwang-tang-hwan. The acupoints primarily used were GV20, EX-HN1, GB5, KI3, PC6, ST36, GV22, and A-shi. The most commonly used pharmacopuncture therapies were Hwangryunhaedoktang (HH), Carthami Fructus (CF), Bee Venom (BV), and Hominis placenta (HP). The Korean medical treatment for alopecia improved the condition of patients. However, seven studies reported the occurrence of side effects such as pruritus, dazed, drowsiness, headache, pain, and diarrhea. This study shows the potential of Korean medicine for the treatment of alopecia. Further studies with a large sample size and long-term follow-up are warranted to establish the primary treatment guidelines and objective outcome measures for alopecia.

Keywords: alopecia, hair loss, korean medicine, alopecia areata, androgenetic alopecia

INTRODUCTION

Alopecia, a disease that results in hair loss in areas where hair normally exists [1], is associated with a negative change in the appearance of the scalp due to gradual decrease in hair density [2]. In Korea, the number of patients with alopecia has increased in the last 5 years (from 208,534 in 2016 to 210,408 in 2020) [3]. In addition, alopecia has been increasingly affected by not only aging or genetic factors but also acquired factors such as environmental pollution, stress, and hormonal secretion abnormalities caused by dietary changes [4].

Alopecia can be divided into cicatricial alopecia and noncicatricial alopecia. Cicatricial alopecia refers to permanent hair loss, whereas non-cicatricial alopecia refers to recoverable hair loss. Non-cicatricial alopecia includes alopecia areata, androgenetic alopecia, and telogen effluvium [5].

Androgenetic alopecia is the most common type of hair loss in both men and women. Androgenic and genetic predispositions appear to be important factors in androgenetic alopecia [6]. Alopecia areata is the formation of round or oval patches of alopecia of various sizes. It is divided into patchy alopecia areata (partial loss of scalp hair), alopecia totalis (total loss of scalp hair), and alopecia universalis (total loss of body and scalp hair) [7].

Currently, the most common treatments for alopecia are oral Finasteride, a Food and Drug Administration-approved drug,

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and topical Minoxidil [8]. However, Finasteride treatment takes 6-12 months to reduce hair loss or increase hair growth and recurrence of alopecia is observed within 2 months of discontinuation. In addition, the use of Finasteride is restricted because of side effects such as reduced sexual desire, decreased sexual functions such as erectile dysfunction, and the risk of birth defects in women. Further, the use of Minoxidil has limitations such as it should be applied twice a day consistently and it works only if the medication stays on the scalp for 3-4 hours [9]. In particular, the use of Minoxidil can cause side effects such as dizziness, dry scalp, itching, skin irritation, and erythema [10]. Treatments such as hair transplant have been proposed; however, these treatments have side effects such as skin atrophy, blood vessel enlargement, hormonal imbalance, and growth inhibition because of long-term drug application [11].

The use of Western medical treatments and hair transplantation has limitations such as side effects; hence, it is important to develop a safe and effective treatment method in Korean medicine.

In Korean medicine, alopecia is generally diagnosed into four syndrome—blood heat syndrome, qi stagnation and blood stasis syndrome, qi and blood deficiency syndrome, and liver and kidney deficiency syndrome. Based on the Deficiency-Excess syndrome classification, deficiency syndromes are classified into blood deficiency syndrome, qi and blood deficiency syndrome, and liver and kidney deficiency syndrome, and excess syndromes are classified into blood heat syndrome, windheat syndrome, qi stagnation, and blood stasis syndrome [12, 13]. The various treatment methods used for alopecia include herbal medicine [14], which involve diagnosis and treatment based on an overall analysis of symptoms and signs, and external treatments such as acupuncture [15], pharmacopuncture [16], and phototherapy [17]. Recently, various case reports and experimental studies have reported the use of microneedling [19] and needle embedding [20] for the treatment of alopecia.

Various Korean medical treatments have been developed and used owing to the growing demand for hair loss treatment; however, the consistency and direction of treatment methods have not been established sufficiently. We conducted this study to establish future treatment directions and effective treatments for alopecia by analyzing case reports and clinical studies on Korean medical treatments for alopecia reported in Korean database and PubMed, a foreign database.

MATERIALS AND METHODS

1. Data sources

May 7, 2021, the study was conducted using four Korean databases—Oriental Medicine Advanced Searching Integrated System (OASIS), Science ON, Korean Studies Information Service System (KISS), and Research Information Sharing Service (RISS)—and one international database—PubMed.

We used "Alopecia" as the search term in OASIS and "(Alopecia) AND (Herbal medicine OR Pharmacopuncture OR Electroacupuncture OR Acupuncture OR Moxibustion OR Cupping Therapy)" in Science ON, KISS, and RISS to search for papers on Korean medical treatment for improving alopecia. We searched PubMed using the following search strategy: "Acupuncture" [MeSH] OR "Electroacupuncture" [MeSH] OR "Cupping Therapy" [MeSH] OR "Dry Needling" [MeSH] OR "Moxibustion" [MeSH] OR "Plants, Medicinal" [MeSH] OR "Phytotherapy" [MeSH] OR "Pharmacognosy" [MeSH] OR "Ethnobotany" [MeSH] OR "Ethnopharmacology" [MeSH]) AND ("Alopecia" [MeSH]).

2. Inclusion and exclusion criteria of studies

In this study, we reviewed papers published in Korean and international journals on Korean medicine treatments to improve alopecia. This study analyzed the use of several treatments methods such as herbal medicine, acupuncture, electropuncture, pharmacopuncture, moxibustion, and cupping therapy.

We excluded non-human experimental studies, reviews, protocol studies, concise reports, and studies unrelated to alopecia treatment. In addition, we excluded studies with interventions unrelated to the direction of the study, studies without original text, and studies with symptoms or diseases unrelated to alopecia.

3. Data extraction

In total, 22 articles were included. The following data were collected: publication year, study type, number of cases. In addition, we analyzed the type of alopecia, Korean medical treatment used, outcome measurements, treatment results, and side effects.

RESULTS

1. Search results

We found 144 articles using Korean databases and 77 articles using the international database. A total of 174 articles were obtained after all articles were ordered in Excel and duplicates based on author names, publication date, and title were removed. Titles and abstracts of 174 articles were reviewed, and 29 non-human experimental studies, 36 reviews, one concise report, one protocol study, and 56 articles unrelated to alopecia treatment were excluded. Finally, we reviewed the full text of the remaining 51 articles. A total of 22 articles were included in analysis after excluding two articles that were not found, 21 articles unrelated to intervention, and six articles on alopecia with other diseases (Fig. 1, Table 1, 2).

2. Number of patients

A total of 1,464 patients received alopecia treatment in 22 studies on Korean medical treatments for alopecia. Among the 1,464 patients, 980 were men and 484 were women.

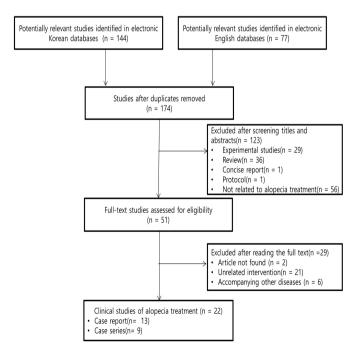


Figure 1. Flow chart.

3. Classification by alopecia types

Six types of alopecia were found in the selected 22 studies. The frequency of each type of alopecia was as follows: alopecia areata, 1,286; androgenetic alopecia, 139; febrile alopecia, 30; telogen effluvium, 7; and alopecia totalis and alopecia universalis, 1 (Table 3).

4. Classification by treatments

The treatments used in 22 studies were herbal medicine, acupuncture, pharmacopuncture, external medicine or product, phototherapy, cupping therapy, life management, microneedling, needle-embedding, blood-letting therapy, moxibustion (Zhuang medicine) and galvanic therapy. The frequency of use of each treatment was as follows (papers): herbal medicine, 17; acupuncture, 15; pharmacopuncture, 9; external medicine or product, 7; phototherapy, 4; cupping therapy, life management, microneedling, and needle embedding, 2; and blood-letting therapy, moxibustion (Zhuang medicine), and galvanic therapy, 1 (Table 4).

In subsection, small-scale case reports (<six) accounted for all of those cases, and large-scale case reports (>six) were calculated by treating them as one. In addition, if different types of treatments were used in one case, each treatment was considered to have been used once.

5. Herbal medicine

In the case of combined prescriptions such as Samul-tang plus Yukmijihwang-tang, each prescription was considered to have been used once. Gamibang and Gagambang were considered to be of the same type as the original prescription. For example, Hyeongbangjihwang-tang gamibang was considered to be the same type as Hyeongbangjihwang-tang. Hwan was treated as Tang.

Herbal medicine was used in 25 of 30 cases. The names of the prescribed herbal medicines were available in 23 cases. A total of 18 types of herbal medicines were prescribed. In the order of frequency, Gagam Cheongyoung-tang and Gagam Hwajung-hwan were prescribed six times; Yukmijihwang-tang·hwan was prescribed three times; and Samul-tang, Soshiho-tang, and Hyungbangjihwang-tang gamibang were prescribed two times. In addition to Yangdokbeakho-tang gamibang, Shihogayong-golmoryeo-tang, Baekhogainsam-tang, Galgeun-tang, Banha-

Author (Year)	Type of alopecia	Patient (Sex/Age)	Intervention (Sort / Site)	Outcome measurement	Result	Adverse events
Yun (2001) [21]	AA	1 (M/37)	1. HM: Hyeongbangjiwhang-tang + Rehmanniae Radix 2. Atx: Kidney tonification (腎正格) 3. Phx: (1) CF:JSD=7:3 / Alopecia region (2) HH / GB21 (3) CC / BL23 4. Cupping therapy: Dry cupping / Shoulder region 5. Phototherapy: IR / Head area 6. Blood-letting therapy: Alopecia region	 Subjective symptoms change 	 Hair loss (6 points) → Improved (4 points), Slightly improved (2 points) 	œ. خ
Kim (2004) [22]	AA	1 (M/30)	 Atx: GV20, GB20, A-shi point Phx: Phx: BV (10000:1)→(4000:1)→(2000:1) / Center&border of scalp CF / Center&border of scalp 	 Visual evaluation&Pictures EEAA 	1. Improved 2. $0 \rightarrow 3$	Slightly pruritus (BV)
Ha (2004) [<mark>15</mark>]	AT	1 (M/8)	 HM: Yangdokbeakho-tang Gamibang → Samul-tang + Yukmijihwang-tang → Samulgamri-hwan Atx: Kidney tonification(腎正格), Friction acupuncture/ Scalp 	1. Visual evaluation&Pictures	1. Improved	œ Ż
Lee (2008) [16]	AU	1 (M/38)	 Phx: BV / Alopecia starting point, Parietal Region feeling stiff tension Life management: lightly shampooing once a day, wiping hair with a towel and drying it after shampoo 	Visual evaluation&Pictures VAS (Alopecia, Tension of scalp) SALT score Stress index	1. Improved 2. (1) Alopecia: $7 \rightarrow 3$ (2) Tension: $5 \rightarrow 1$ 3. S2 (39.8% \rightarrow 43.0%) 4. $37 \rightarrow 38$	Dazed (1 h)
Lee (2009) [23]	AA	1 (F/19)	 HM: not mentioned Phx: CF / Alopecia region Atx: GV20, ST8, GV23, GV21, BL6, EX-HN1, GB20 Life management: only shampooing once a day, reduce eating processed food 	 Visual evaluation&Pictures VAS (Patient's awareness about hair loss progress) Stress index EEAA Response of treatment 	1. Improved 2. 9 → 2 3. 48 → 43 4. 0 → 3 5. No → Good	œ. Z
Hwang (2012) [24]	AA	1 (F/5)	 HM: Shihogayonggolmoryeo-tang → Baekhogainsam-tang 	 Visual evaluation&Pictures 	 Hair loss patch diameter (cm): 2.5 → 0.5 	N. R.

Table 1. Continued	ontinued					
Author (Year)	Type of alopecia	Patient (Sex/Age)	Intervention (Sort / Site)	Outcome measurement	Result	Adverse events
Ju (2013) [18]	AA	1 (F/13)	 HM: Galgeun-tang → Banhabackchulchunma-tang → Galgeuntang → Soshiho-tang + Rhinocerotis Cornu→ Soshiho-tang + Rhinocerotis Cornu→ Soshiho-tang + Rhinocerotis Cornu, Daehamhyung-hwan Atx: A-shi point (Hair loss area), CV4 Phx: (1) HP / 2 cm outward of lesion to inward (2) JsD / Induratum of upper trapezius, GB21, Intersecting points (Medial line of scapula&T3-T7 height), LU1 (3) BUM / BL32, BL34, Inner point of 12 ribs end, Inner point of ASIS External medicine or product: Scalp mist, Shampoo 5. Phototherapy: (1) High frequency wave therapy / Trapezius (2) Low frequency wave therapy / Shoulder, Back (3) IR 6. Cupping therapy: Dry cupping / Shoulder region 7. Microneedling: GL / Alopecia region 8. Galvanic 	1. Visual evaluation&Pictures	1. Improved	-Drowsiness after meals -Headache
Yoon (2014) [19]	AGA	1 (F/42)	HM: Soshiho-tang Atx: GV20, EX-HN1, ST8, Ll4, Kl3, BL2, TE23, GB1, ST1, A-shi point (Scalp, Circumbulbar) Shx: HH / Whole Parietal region Needle embedding: Scalp	Visual evaluation&Pictures VAS (Cephalic fever and perspiration) Ludwig classification	1. Improved 2. $10 \rightarrow 1$ 3. Grade $2 \rightarrow 1$	-Pain -Aching (1 d)
Ko (2016) [25]	AGA	1 (M/27)	 HM: Hyeongbangjihwang-tang Gamibang Atx: CV12, ST36, LI4, EX-HN5, LR3 Phx: V / Frontal region, CV1 + HA / GB21, GB20, BL31 → V / Frontal region + HA / Frontal region 	 Visual evaluation&Pictures Subjective evaluation (Pruritus&Seborrhea) 	1. Improved 2. Improved	Diarrhea (degeneration)
Kim (2016) [17]	AA (by CA)	1 (M/19)	 HM: Daeyeon&jeon Atx: A-shi point (Alopecia region) Phototherapy: Hani-maehwa Laser / Alopecia region 	1. Visual evaluation&Pictures	1. Improved	N.S.
Kim (2016) (2) [26]	¥	1 (F/42)	 HM: Shipjeondaebo-tang Gamibang Atx: A-shi point (Alopecia region), ST36, Ll4 Ptx: HH / Scalp External medicine or product: Herbal spray (Houttuyniae Herba, Perillae Folium, Betulae Cortex) / Scalp 	Visual evaluation&Pictures VAS (subjective symptoms changed)	1. Improved 2. $10 \rightarrow 1$	ĸ.

5. Phototheraphy: IR 6. Needle-embedding: Scalp

Table 1. (Table 1. Continued					
Author (Year)	Type of alopecia	Patient (Sex/Age)	Intervention (Sort / Site)	Outcome measurement	Result	Adverse events
Yoon (2017) [27]	AA	4 (M/39, M/35, M/26, M/25)*	 HM: Gagam Cheongyoung-tang Atx: GV20, GV22, BL7, PC6, KI3, LR2 External medicine or product: Scalp soothing ointment (Rehmanniae Radix, Osterici Radix, Liriopes Radix, Saposhnikovia Radix, Bupleuri Radix, Scutellariae Radix, Laminariae Thallus, Ecliptae Herba, menthol) / Crown of the head (Arounding GV20) 	1. Visual evaluation&Pictures	1. Improved	None
Choi (2017) [14]	AA	5 (M/46, M/30, M/27, F/53, F/22) [†]	 HM: Gagam Hwajung-hwan Atx: GV20, EX-HN1, GB5 etc. (According to the symptoms) 	Wisual evaluation&Pictures Magnifying glass SALT score	1. Improved 2. Improved 3. $(M/46)$: $S1 \rightarrow S0$, $(M/30)$: $S2 \rightarrow S0$, $(F/53)$: $S2 \rightarrow S0$, $(M/27)$: $S2 \rightarrow S0$, $(F/22)$: $S1 \rightarrow S0$	K.
Choi (2019) [28]	AA	1 (F/4)	 HM: Gagam Cheongyoung-tang→ Gagam Palmul-tang→ Gagam Cheongyoung-tang 	 Visual evaluation&Pictures Magnifying glass SALT score 	 Improved Improved S1 → S0 	Diarrhea
Zhang (2019) [29]	AA	1 (M/36)	1. Medicated thread moxibustion (Zhuang medicine): ST36, SP10, GV20, LR3, Kuihua acupoint	Visual evaluation&Pictures Pull test	1. Improved 2. Improved	S. S.
Yun (2020) [30]	AGA	2 (M/70, M/60)	 HM: Yukmijihwang-hwan External medicine or product: Hair-growth tonic (Acori Calami Rhizoma, Glycyrrhizae Radix, Centellae herba, Sophorae Radix, Capsici Fructus, Camelliae 	 Visual evaluation&Pictures HNS 	1. Improved 2. No change	Z Z

Folium)

(2) Hair-growth shampoo (Ligustici Rhizoma Acori Calami Rhizoma, Coicis Semen, Mori Radicis Cortex, Centellae herba, Camelliae

Herbal medicine; Phx, Pharmacopuncture; CF, Carthami Fructus; JsD, Juglandis Semen; HH, Hwangryunhaedoktang; CC, Cervi pantotrichum Cornu; BV, Bee Venom; EEAA, Evaluation of the effect on alopecia areata; VAS, Visual Analog Scale; SALT, Severity of alopecia tool; HP, Hominis placenta; BUM, Calculus bovis Fel ursi Moschus; GL, Ganoderma lucidum; HNS, Hamilton-AA, Alopecia areata; AU, Alopecia universalis; AT, Alopecia totalis; AGA, Androgenetic alopecia; FA, Febrile alopecia; TE, Telogen effluvium; CA, Cicatricial alopecia; Atx, Acupuncture; HM, Norwood scale.

^{*}Patients were already taking Finasteride over 1 year before treatment and during treatment accompanied Finasteride.

[†]Patients who were resulted S2 in SALT scale.

Table 2. Analyzation of clinical treatments and results (patients more than six)

Adverse events	None	Ä.	х ж.	Ä.
Result	1. Improved 2. Improved 3. Improved 4. Improved	1. Improved 2. Improved 3. Improved	1. Improved (56.2%) (p = 0.003) 2. 13.14 \pm 4.13 \rightarrow 13.82 \pm 4.12 (p < 0.001) 3. Improved [§]	1. (1) 5.35 \pm 2.82 \rightarrow 4.21 \pm 2.29 (p = 0.082) (2) 4.5 \pm 1.28 \rightarrow 1.6 \pm 1.10 (p < 0.001) 2. 1.43 \pm 0.27 \rightarrow 1.61 \pm 0.24 (p < 0.001)
Outcome measurement	1. Magnifying glass 2. Questionnaire (Effect on the herb medicine prescription) 3. Hair loss during shampooing 4. Hair regrowth change	Visual evaluation&Pictures Questionnaire (Patient self-assessment) Investigator assessment	 Visual evaluation&Pictures Magnifying glass Questionnaire (Parietal Fever, Sebum, Keratin, Pruritus, Hair elasticity, Change of hair loss volume) 	1. Questionnaire(1) Cold-heat questionnaire(2) Hair and scalp questionnaire2. Magnifying glass
Intervention (Sort / Site)	1. HM : Honey pill (Biotae Cacumen, Mori Folium, Phyllostachydis Folium, Ginkgo Folium, Pini Polium, Glycyrrhizae Radix, Ginseng Radix, Pulsatillae Radix, Kochiae Fructus, Moutan Cortex, Lycii Fructus, Rehmanniae Radix Preparat, Poria (Hoelen), Corni Fructus, Sesami Semen Nigrum, Prunellae Spica, Acanthopanacis Cortex, Alismatis Rhizoma, Taraxci Herba, Laminariae Thallus, Biotae Semen) 2. External medicine or products 3. External medicine or products 4. Herbal spray (Biotae Cacumen, Mori Folium, Phyllostachydis Folium, Ginkgo Folium, Pini Polium, Glycyrrhizae Radix, Ginseng Radix, Pulsatillae Radix, Kochiae Fructus, Houttuyniae Herba, Lycii Fructus, Glycinis Semen Nigra, Sesami Semen Nigrum, Prunellae Spica, Acanthopanacis Cortex, Cannabis Folium, Laminariae Thallus extract)	 Atx: Oh-Shu acupoint (According to the symptoms) Phx: HP, CF etc. / ST8, GV23, GV21, GV20, BL6, EX-HN1 External medicine or products: Shampoo (Blue spring from Dr. hoo Corp) 	 HM: Unknown Atx: GV20, EX-HN1, GB5, ST8, PC6, HT8, ST36, BL66 etc. (According to the symptoms) External medicine or product: Oriental hair care products (Balmers M shampoo, M tonic, 701 serum) 	1. HM: Gagam Chengyoung-tang 2. Atx: GV20, GV21, GV22, GV23, ST8, HT8, ST36
Patient N (M, F) Age [Mean]	6 (2, 4) [49.6]	82 (82, 0) [30 ± 5.5]	1,050 [‡] (714, 336) [30.67 ± 9.03]	30 (30, 0) [31.73 ± 6.34]
Type of alopecia	AGA	AGA	AA A	₹
Author (Year)	Kim (2004) (2) [31]	Yi (2006) [32]	Lee (2013) [33]	Hong (2013) [34]

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'1,050 people conducted an initial survey, but 153 people conducted a retrospective survey in the actual paper.

SALT score was calculated according to these categories (Age and Sex Distribution, Clinical Type of AA, Extent of AA, Duration before Visit, Age of onset, Age of first visit, The period until Regrowth of Terminal Hair, The Period until Cured, Associated immunological disease, Previous treatments, Family history, Relapse) but the p-value of Age and Sex Distribution, Age of onset, Improved patient rate (%): Parietal Fever (37.3%), Sebum (19.6%), Keratin (23.5%), Pruritus (30.7%), Hair elasticity (83%), Change of hair loss volume (31.4%). Age of first visit, Associated immunological disease, Previous treatments, Family history is over 0.005.

Table 3. Types of alopecia reported in alopecia clinical studies

Type of alopecia	Number of patients n (%)
Alopecia areata	1,286 (87.8%)
Androgenetic alopecia	139 (9.5%)
Febrile alopecia	30 (2.0%)
Telogen effluvium	7 (0.5%)
Alopecia totalis, Alopecia universalis	1 (0.1%)
Total	1,464

Table 4. Interventions reported in alopecia clinical studies

Intervention	Number of papers n (%)	Number of cases n (%)
Herbal medicine	17 (77.3%)	25 (83.3%)
Acupuncture	15 (68.2%)	22 (73.3%)
Pharmacopuncture	9 (40.9%)	9 (30.0%)
External medicine or product	7 (31.8%)	11 (36.7%)
Phototherapy	4 (18.2%)	4 (13.3%)
Cupping therapy, Life management, Microneedling, Needle-embedding	2 (9.1%)	2 (6.7%)
Blood-letting therapy, Moxibustion (Zhuang medicine), Galvanic	1 (4.5%)	1 (3.3%)
Total	22	30

Table 5. Herbal medicines reported in alopecia clinical studies

Herbal medicine	Number of cases n (%)
Gagam-cheongyoung-tang (加減淸營湯), Gagam-hwajung-hwan (加減和中丸)	6 (18.2%)
Yukmijihwang-tang·huan (六味地黃湯·丸)	3 (9.1%)
Samul-tang (四物湯) (including Gami) Soshiho-tang (小柴胡湯) (including Gami) Hyeongbangjihwang-tang gamibang (荊防地黃湯加味方)	2 (6.1%)
Yangdokbeakho-tang (陽毒白虎湯) Shihogayonggolmoryeo-tang (柴胡加龍骨牡蠣湯) Baekhogainsam-tang (白虎加人蔘湯) Galgeun-tang (葛根湯) Banhabackchulchunma-tang (半夏白朮天麻湯) Daehamhyung-hwan (大陷胸丸) Daeyeong-jeon (大營煎) Gagam Palmul-tang (加減八物湯) Gagam Hyeongbangpaedok-san (加減荊防敗毒散) Gagam Jaeumbaekbo-hwan (加減滋陰百補丸) Samulgamri-hwan (四物坎离丸) Honey pill (蜜丸)	1 (3.0%)
Total	33

backchulchunma-tang, Daehamhyung-hwan, Daeyeong-jeon, Gagam Palmul-tang, Gagam Hyeongbangpaedok-san, Gagam Jaeumbaekbo-hwan, and Samulgamri-hwan, Honey pills were used once (Table 5).

6. Acupuncture

Acupuncture was used in 22 of the 30 cases analyzed. In order of frequency, the following acupuncture points were used: GV20, 15 times; EX-HN1, nine times; GB5, seven times; KI3 and PC6, six times; ST36, GV22, and A-shi point, five times; BL7, LR2, and ST8, four times; LI4, three times; and Kidney tonification, GV23, GV21, GB20, and HT8, two times. In addition, BL6, CV4, BL2, TE23, GB1, ST1, CV12, EX-HN5, LR3, BL66, TE5 and Oh-shu acupoint were used once (Table 6).

7. Pharmacopuncture

Pharmacopuncture treatment was used in nine of the 30 cases analyzed. Hwangryunhaedoktang (HH) and Carthami Fructus (CF) pharmacopuncture were the most commonly

Table 6. Acupoints reported in alopecia clinical studies

Acupoint	Number of cases n (%)
GV20	15 (15.8%)
EX-HN1	9 (9.5%)
GB5	7 (7.4%)
KI3, PC6	6 (6.3%)
ST36, GV22, A-shi point	5 (5.3%)
BL7, LR2, ST8	4 (4.2%)
LI4	3 (3.2%)
Kidney tonification, GV23, GV21, GB20, HT8	2 (2.1%)
BL6, CV4, BL2, TE23, GB1, ST1, CV12, EX-HN5, LR3, BL66, TE5, Oh-shu acupoint	1 (1.1%)
Total	95

Table 7. Pharmacopunctures reported in alopecia clinical studies

Pharmacopuncture	Number of cases n (%)
HH, CF	3 (18.8%)
BV, HP	2 (12.5%)
JsD, CC, BUM, V, HA, CF · JsD (Mixed)	1 (6.3%)
Total	16

used pharmacopuncture treatments (three times), followed by Bee Venom (BV) and Hominis placenta (HP) (two times), and Juglandis Semen (JsD), Cervi pantotrichum Cornu, Calculus bovis Fel ursi Moschus, V, HA and CF plus JsD (one time) (Table 7).

V pharmacopuncture consists of the cystic stone of Bos taurus domesticus Gmelin, the gallbladder of Ursus thibetanus G. Cuvier, the gland secretion of Moschus moschiferus L, etc. HA pharmacopuncture consists of the flower of Lonicera japonica Thunb, the fruit of Eriobotrya japonica Lindl, etc.

8. External medicine or product

External medicine or product was used in 11 of the 30 cases analyzed. The most commonly used external medicine or product was shampoo (five times), followed by ointment (four times), tonic (three times), herbal spray (two times), and serum and mist (one time) (Table 8).

9. Phototherapy

Phototherapy was used in four of the 30 cases analyzed. Infrared radiation was the most commonly used phototherapy. High-frequency, low-frequency and laser were used once each. Hani-maehwa Laser, a low-power laser, was used (Table 9).

Table 8. External medicine or products reported in clinical studies

External medicine or product	Number of papers n (%)
Shampoo	5 (31.3%)
Ointment	4 (25.0%)
Tonic	3 (18.8%)
Herbal spray	2 (12.5%)
Serum, Mist	1 (6.3%)
Total	16

Table 9. Phototherapies reported in clinical studies

Phototherapy	Number of cases n (%)
IR (Infrared ray)	3 (50.0%)
High frequency, Low frequency, Laser	1 (16.7%)
Total	6

10. Other interventions

The other types of treatments used were cupping therapy, life management, microneedling, needle embedding, blood-letting therapy, moxibustion (Zhuang medicine) and galvanic therapy. Cupping therapy, life management, microneedling, and needle embedding were used twice, and blood-letting therapy, moxibustion (Zhuang medicine) and galvanic therapy were used once (Table 10).

11. Classification by outcome measurement

The treatment can change for each case in the same study; however, the outcome measurements are always the same. Therefore, we assessed the outcome by the number of papers and not by the number of cases. In total, 18 types of outcome measurements were used in 22 alopecia treatment studies. Eight types of evaluation tools were used in two or more papers. visual observation and images, 16 studies; magnifying glass, 6 studies; Visual Analog Scale (VAS), Severity of alopecia tool (SALT), and questionnaire, 4 studies; and stress index, Evaluation of the effect on alopecia areata (EEAA), and pull test, 2 studies. In ad-

Table 10. Other interventions reported in clinical studies

Other intervention	Number of cases n (%)
Cupping therapy, Life management, Microneedling, Needle embedding	2 (18.2%)
Blood-letting therapy, Moxibustion (Zhuang medicine), Galvanic	1 (9.1%)
Total	11

Table 11. Outcome measurements reported in clinical studies

Outcome measurement	Number of papers n (%)
Visual evaluation&Pictures	16 (72.7%)
Magnifying glass	6 (27.3%)
VAS, SALT, Questionnaire	4 (18.2%)
Stress index, EEAA, Pull test	2 (9.1%)
Subjective symptoms change, subjective evaluation, response of treatment, Ludwig Classification, HNS, hair condition change, hair loss during shampooing, hair regrowth change, investigator assessment, iconography	1 (4.5%)
Total	22

dition to the following outcome measurements were used once; subjective symptoms change, subjective evaluation, response to treatment, Ludwig classification, Hamilton-Norwood scale (HNS), hair condition change, hair loss during shampooing, hair regrowth change, investigator assessment, and iconography (Table 11).

The aforementioned outcome measurements can be largely divided into visual, subjective, and objective measurements. Visual measurements (visual evaluation and pictures, magnifying glass, severity on the alopecia tool, evaluation of the effect on alopecia areata, Ludwig classification, Hamilton-Norwood scale, hair condition change, and iconography) were used 32 times. Subjective measurements (VAS, questionnaire, stress index, subjective symptoms change, subjective evaluation, response of treatment, and investigator assessment) were used 14 times. Objective measurements (pull test, hair loss during shampooing, and hair regrowth change) were used four times (Table 12).

12. Side effects

In 12 studies, side effects were not mentioned. Three studies reported no side effects. In the remaining seven studies, the side effects were described. Choi (2019) [20] and Choi (2019) (2) [28] reported diarrhea, Ko (2016) [25] reported worsening of diarrhea symptoms in patients with diarrhea, Kim (2004) [22] reported slight pruritus during treatment with BV, Lee (2008) [16] reported dizziness for an hour, Ju (2013) [18] reported drowsi-

Table 12. Outcome measurements reported in clinical studies (divided into 3 groups)

Туре	Outcome measurement	Number of total outcome measurement n (%)
Visual	Visual evaluation&Pictures, Magnifying glass, SALT, EEAA, Ludwig Classification, HNS, hair condition change, iconography	32 (64%)
Subjective	VAS, Questionnaire, Stress index, subjective symptoms change, subjective evaluation, response of treatment, investigator assessment	14 (28%)
Objective	Pull test, hair loss during shampooing, hair regrowth change	4 (8%)
Total		50

ness after meals and headache, and Yoon (2014) [19] reported pain and aching for a day.

DISCUSSION

Recently, the incidence of alopecia, a hair loss disease, has been increasing [1, 3]. Currently, Finasteride and Minoxidil are mainly used in Western medicine for the treatment of alopecia [8]. However, they can cause problems such as decrease sexual function and increase the risk of birth defects [9]. Hence, we investigated treatments for alopecia in Korean medicine, which has relatively few side effects.

A total of 22 studies were selected for analysis. These studies have been published in the past 20 years (one in 2001, three in 2004, 1 in 2006, one in 2008, one in 2009, one in 2012, three in 2013, one in 2014, three in 2016, two in 2017, four in 2019, and one in 2020). This suggests that studies on treatment for alopecia in Korean medicine are steadily underway.

In 22 studies, 1,464 patients (980 men and 484 women) were treated for alopecia using Korean medical treatments. A total of six types of alopecia—alopecia areata, androgenetic alopecia, febrile alopecia, telogen effluvium, alopecia totalis, and alopecia universalis—were identified in 1,464 patients. Most patients (n = 1,286) had alopecia areata, while 139 patients had androgenetic alopecia. This gap is believed to have been particularly large because a case study [33] included 1050 patients with alopecia areata.

The treatments for alopecia in Korean medicine include herbal medicine, acupuncture, pharmacopuncture, external medicine or product, cupping therapy, microneedling, moxibustion (Zhuang medicine) and blood-letting therapy. In addition, phototherapy, life management, and galvanic therapy are used as a supplement therapy. However, Yoon (2017) [27] used Finasteride along with Korean medication to treat alopecia. To determine the frequency of each Korean medical treatment, small-scale case reports (<6 cases) were considered each, and large-scale case reports (>6 cases) were calculated by treating the number of cases as one.

Herbal medicine was used in 25 of 30 cases. The most commonly used prescriptions were Gagam Cheongyoung-tang and Gagam Hwajung-hwan, six times each. Cheongyoung-tang, a typical prescription for clearing heat and cooling blood, nourishes the liver and kidney to cool the heat. Gagam Cheongyoung-tang is a basic addition to Cheongyoung-tang [36]. As mentioned in the introduction, the cause of alopecia is bloodheat syndrome, wind-heat syndrome, and dampness-heat syndrome [12], thus, the use of Gagam Cheongyoung-tang which has the effect of clearing heat can control alopecia. Gagam Hwajung-hwan is used in patients with stress heat syndrome accompanied by digestive congestion. Currently, alopecia is considered to be affected by acquired factors such as stress [23]; therefore, it is believed that alopecia can be controlled by Gagam Hwajung-hwan.

Acupuncture was used in 22 of 30 cases. The most commonly used acupoint was GV20, which was used 15 times. This acupoint is located on the scalp and is believed to have been used to relieve scalp fever, a major cause of alopecia. EX-HN1, GB5, GV22, and A-shi point are all local acupuncture points used more than five times. Local acupuncture is considered to induce new hair by increasing hair cells, dermal papilla cells, and capillary blood flow circulation, which play an important role in hair development [21]. The distal acupuncture points were KI3, PC6, and ST36, which were used more than five times. ST36 is usually used for alopecia caused by qi and blood deficiency syndrome [26].

Pharmacopuncture was used in 9 of 30 cases. The most commonly used pharmacopuncture was HH and CF. HH pharmacopuncture was used to control the heat in the parietal region by clearing away heat toxin and improve alopecia by stimulating the scalp [37]. CF pharmacopuncture is effective in activating blood circulation and detoxification; hence, it is likely to be effective in a syndrome in which the hair coat cannot be nourished due to body fluid deficiency, blood deficiency, and poor communication between blood circulation [23]. In addition, BV and HP pharmacopuncture were used. Ko (2016) [25] reported the use of HN pharmacopuncture; however, on analyzing the details, HA acupuncture, not HN acupuncture, was reported. Hence, we determined and analyzed it as HA acupuncture.

External medicine or product was used in 11 of 30 cases. The most commonly used external medicine or product was shampoo (five times), which is thought to be versatile because it is closely involved in scalp health and alopecia. Ointment was the second most commonly used product to soothe the scalp, rather than treat alopecia directly. Yoon (2017) [27] reported the use of a self-made sedation ointment of the root of Rhemannia glutinosa (Gaertner) Liboschitz, the root of Ostericum koreanum Kitagawa, and the tuberous root of Liriope platyphylla Wang et Tang to be applied on the head.

Phototherapy was used in 4 of 30 cases. The most commonly used phototherapy was infrared radiation, which is believed

to have been used to aid the supply of nutrients to hair by promoting blood circulation and repair the scalp damaged by cell activation [38]. A high frequency, low frequency, and laser were used once each. High-frequency laser was used to administer heat to the deep parts of the body, and low-frequency laser was used for systemic stimulation of nerves and muscles [18]. The Hani-maehwa laser was used to induce hair growth and reduce the side effects [17].

The other treatments included cupping therapy, microneedling, needle embedding, moxibustion (Zhuang medicine), blood-letting therapy, life management, and galvanic therapy. Regarding cupping, dry cupping was performed in the shoulder region. Microneedling facilitates the absorption of pharmacopuncture solution into the skin and treatment of alopecia through scalp irritation [18]. Needle-embedding therapy was performed using a 29Gx30mm Miracu in the parietal region of the scalp. This treatment induces the body's defense system by introducing harmless foreign substances into the bloodstream, which is believed to facilitate alopecia treatment by stimulating the parietal region [26].

Regarding life management, Lee (2008) [16] indicated that hair should be shampooed lightly once a day, wiped lightly with a towel, and dried with a dryer. Lee (2009) [23] set a standard for shampooing once a day and reducing the consumption of processed foods. Yun (2001) [21] indicated that blood circulation should be increased in the alopecia area through bloodletting therapy, and Ju (2013) [18] reported scalp care before acupuncture treatment with galvanic.

In 22 studies, 18 types of outcome measurements (50 times) were used. Outcome measurements were largely divided into visual, subjective, and objective measurements, Visual measurements were used the most frequently (32/50 [64%]), with a high proportion of visual evaluations and pictures. This is because alopecia is a visible disease, and visual evaluation methods are simple and accurate. SALT, one of the popular hair loss assessment tools, consists of visual indicators. This is related to the use of more subjective tools than objective evaluation tools.

This study has limitations. To assess the trend of foreign research, we used PubMed; however, the PubMed search yielded only two studies that were in accordance with the eligibility criteria. In addition, the assessment of objective outcome measurements for alopecia is essential in the future, given that most of the outcome measurements used in the studies were far from objective, forcing the authors to intervene in subjective interpretation. Most selected studies were case studies; hence, in many studies, interventions changed as the treatment progressed. For example, in Ju's study (2013) [18], the herbal medicine administered to a patient was changed to Galgeun-tang, Banhabaekchulcheonma-tang, and Soshiho-tang plus the horn of Cervus elaphus L., Daehamhyung-hwan four times. Thus, further studies are warranted to determine the therapeutic effectiveness of each intervention.

Compared with a previous review in 2019 [39], we analyzed six more studies, including those from foreign databases. Moreover, we corrected the incorrect data in the previous review. We hope that this study will be used as a reference to develop guidelines for alopecia for actual clinical use.

CONCLUSION

This study analyzed 22 studies on Korean medicine treatments for alopecia in using four Korean databases and a international database. The main findings of this study are as follows:

- 1. A total of 1,464 patients were included in 22 studies. Patients with alopecia areata accounted for 87.8% of all patients. The treatments used for alopecia, in order of frequency (cases), were as follows: herbal medicine, acupuncture, external medicine or product, pharmacopuncture, phototherapy, cupping therapy, life management, microneedling, needle embedding, blood letting therapy, moxibustion (Zhuang medicine) and galvanic therapy.
- 2. In herbal medicine, Gagam Cheongyoung-tang and Gagam Hwajung-hwan, Yukmijihwang-tang-hwan, Samul-tang, Soshiho-tang, and Hyungbangjihwang-tang gamibang were mainly used.
- 3. The most commonly used acupuncture point was GV20, followed by EX-HN1, GB5, KI3, PC6, ST36, GV22, and A-shi. In addition, both local and distal acupuncture treatments were performed; however, the proportion of local acupuncture treatments was higher, with the top eight most commonly used acupoints. HH and CF were mainly used in pharmacopuncture.
- 4. The main external medicine or product used was shampoo, followed by ointment. The phototherapy mainly used was infrared radiation, followed by high-frequency, low-frequency and laser.
- 5. The outcome measurements were largely divided into visual, subjective, and objective measurements. Visual measurements (such as visual evaluation and pictures and magnifying glass) were the most commonly used. The other outcome mea-

sures used were VAS, SALT, questionnaire, and stress index.

CONFLICTS OF INTEREST

The authors declare no conflict of interest.

AUTHORS' CONTRIBUTION

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