Abstract: Many ingredients found within nail cosmetic products are capable of sensitizing patients’ immune systems and causing contact dermatitis (CD). These include but are not limited to tosylamide, (meth)acrylates, and formaldehyde. A clear temporal relationship between nail cosmetic procedures and an eczematous outbreak on the hands, face, or other ectopic body regions can be a key indicator of CD secondary to nail cosmetic exposure. Once an inciting allergen is identified through patch testing, elimination and avoidance becomes a mainstay of treatment alongside the use of emollients and topical anti-inflammatory therapies. Patients should be counselled to approach future nail cosmetic products and procedures with caution and careful attention to ingredients, regardless of whether or not it has a “hypoallergenic” label.

Keywords: allergic contact dermatitis; irritant contact dermatitis; nail cosmetics

1. Introduction

Nail cosmetics have been a mainstay of grooming and beauty since the 17th century, with the earliest findings of nail coloring and enhancements dating back to ancient civilizations in Egypt and China [1]. Today, millions of people worldwide utilize nail cosmetic products in order to achieve aesthetically pleasing nails, with the nail cosmetic industry growing from a USD 3 billion industry in 2007 to a USD 45 billion industry by 2012 [2]. In the USA alone, spending on nail salon services exceeds USD 8.5 billion yearly [3]. While many of the products used to achieve this look have been shown to be relatively safe, there are several complications associated with their use. These complications include mechanical and traumatic complications, infection, exposure to carcinogenic ultraviolet (UV) radiation (used to cure and set gel nail polish), and contact dermatitis (CD) [4]. The latter complication will be the focus of this narrative review article.

CD, which can be further divided into allergic (ACD; 20% of cases) and irritant CD (ICD; 80% of cases), is an immune reaction that occurs in response to external compounds. This causative relationship between nail cosmetic products and CD has been well documented for over 50 years [5–7]. Specifically, a 1987 study showed that 4.2% of CD cases was due to the use of a cosmetic product, with approximately 8% of those cases attributed to nail cosmetics [6]. Since then, the number of patients diagnosed with CD secondary to use of a nail cosmetic product has grown along with the nail cosmetics industry, likely due to a combination of increased use and sensitization to the included ingredients [8–11]. In fact, the prevalence of contact allergies to one or more (meth)acrylates was shown to have significantly increased by more than 20% between 2009–2013 and 2014–2019 among nail technicians and product users [12]. A retrospective analysis of 38,775 patients who had previously undergone patch testing found that 769 (2%) had an allergic or irritant patch test reaction to a common ingredient in nail cosmetics [9]. This study was consistent with prior estimates that 1–3% of the population is sensitized to at least one ingredient found in nail cosmetics [5]. CD secondary to nail cosmetics is most prevalent among women aged 26–46 and nail technicians within their first year of work; however, it can develop within any demographic if nail products are used.
In order to consolidate the current body of literature on this growing problem and provide suggestions for improved diagnosis, management, and education, PubMed searches were conducted using the key phrases “nail contact dermatitis”, “nail contact allergy”, “nail cosmetics allergy”, and “acylate allergy”. In the current narrative review article, we will discuss the most common causative ingredients, clinical presentation, evaluation, and management of contact dermatitis secondary to nail cosmetics.

2. Common Nail Cosmetic Products and Associated Allergens

According to modern beauty standards, aesthetically pleasing nails have a glossy, smooth surface, no overhanging or ragged cuticles, a tip extending beyond the nail plate, an oval contour to the nail plate, and a gentle curve when visualized from the side [1]. A variety of products can be used to achieve these looks, including liquid nail polish, nail wraps, gel nail polish, dipping powders, and acrylics (often referred to as “tips”) [4]. Other nail care products include nail strengtheners, cuticle softeners, and nail hydrating polishes, oils, and serums. More detailed descriptions of these products can be found in Table 1. Regular manicures using nail polish and gel manicures are the two most popular nail cosmetic procedures in the USA [3].

Table 1. Description of common nail cosmetic products.

| Product (Synonyms) | Purpose | Application Process | Approximate Duration |
|-------------------|---------|---------------------|----------------------|
| Nail Polish (lacquer, varnish) | Gives nails color and shine | Painted directly onto nail plate or on top of acrylic or gel nails, with or without base or top coats. Can be applied at home or in salons | <1 week |
| Base Coat | Creates an even surface for the color coat and facilitates adhesion | Applied directly to nail plate prior to application of colored nail polish | N/A |
| Top Coat | Gives a glossy finish and protects nail enamel | Applied on top of colored nail polish | N/A |
| Nail Polish Remover (acetone) | Dissolves and removes nail polish and gel polish | Nail polish: wiped onto nails using a cotton applicator (such as a cotton ball or swab) Gel polish: poured into a bowl in which nails are soaked for several min or tightly applied to the nails using a cotton applicator and aluminum foil for several min | N/A |
| Nail Wraps (silk nails, preformed nails) | Reinforce weak, brittle, or breaking nails | Sheets applied directly to nail plate with cyanoacrylate-based nail glue or adhesive backing, then warmed and trimmed to accommodate the shape of the underlying nail | 1–2 weeks |
| Press-on Nails (fake nails, stick-ons) | Immediate color/design with durable shine and shape Quick application that requires little to no skill | Plastic nail applied directly to nail plate with cyanoacrylate-based glue or adhesive backing | 1–2 weeks |
| Gel Nail Polish (long-lasting, semipermanent) | Long-lasting color Less drying time compared to liquid nail polish | Liquid polish made of acrylic monomers is painted directly onto the nail plate or on top of acrylic nails. Each applied layer (3+) is cured with a UV or LED light (30 s–2 min) to allow for polymerization of acrylic monomers | 2+ weeks |
| Powder Nails (dipped nails, dipping powders) | Long-lasting color with a hard, durable finish Appeals to many concerned about UV exposure with gel polish | Nail prepared with cyanoacrylate-based nail glue is dipped into polymer powder. A water-based activator is then applied to the nails, hardening the polish | 2–4 weeks |
| Acrylic Nails (tips, nail extensions, porcelain nails) | Adds length and shape to existing nail | Powder polymers mixed with liquid acrylic monomers and then sculpted onto the nail plate | 2–3 weeks |

While the products discussed above are certainly useful in enhancing nail appearance, various ingredients within these products can produce both ACD and ICD. Table 2 displays the most common allergens and irritants in nail cosmetic products and which products/procedures they are found in. While the list of these ingredients is extensive, a recent retrospective analysis of patch testing results performed in those diagnosed with CD secondary to nail cosmetic ingredients most frequently revealed positive reactions in
response to hydroxyethyl methacrylate (56.6%), tosylamide (36.2%), methylmethacrylate (27.8%), ethyl acrylate (25.2%), and ethyl-2-cyanoacrylate (6.9%) [9]. We discuss these and other common ingredients below:

- **Tosylamide/formaldehyde resin (TSFR)** has long been known to be the most common cause of ACD related to nail polish, with data from the North American Contact Dermatitis Group (NACDG) suggesting that 4% of all positive patch tests involved sensitivity to TSFR [13]. It has been found to sensitize both nail components as well as produce ectopic ACD in areas of touching, scratching, or rubbing [14]. Because of this, many nail polish brands have switched formulation to include tosylamide epoxy resin instead; however, this has also been shown to sensitize both locally and ectopically [15].

- **Methacrylates** (powder and liquid) are mixed to form a flexible polymer for acrylic manicures. These ingredients can also be found in smaller quantities in both regular and gel nail polish. The three most common allergenic forms of these ingredients are 2-hydroxyethyl methacrylate (2-HEMA), 2-hydroxypropyl methacrylate (2-HPMA), and ethylene glycol dimethacrylate (EGDMA) [16,17]. These ingredients are most sensitizing in their liquid form (during application) and rarely cause reactions once hardened and cured. In addition, methacrylates are extremely cross-reactive with one another, so people are often allergic to multiple [15].

- **Acrylates** are mainly used in gel manicure systems. These include 2-hydroxyethyl acrylate (2-HEA), 2-ethylhexyl acrylate (2-EHA), 2-hydroxypropyl acrylate (2-HPA), ethyl acrylate (EA), and triethylene glycol diacrylate (TREGDA) [18]. With the increasing popularity of and consumer access to “at-home” gel manicure kits, sensitization to acrylates has increased, and those that use such kits are also more likely to be sensitized [19,20].

- **Dibutyl phthalate (DBP)** is considered a “plasticizer” that increases nail polish flexibility [21]. It has not only been seen to cause ACD, but it has also been shown to interfere with male reproductive development in animal models [22].

- **Benzophenone** is an additive to regular and gel nail polishes that protects cosmetic products by absorbing UV light and preventing product degradation prior to use. Cases report both ACD and photocontact dermatitis as a result of contact with benzophenone [23].

- **Formaldehyde** (also listed as formalin or methylene glycol) is the main ingredient in nail strengthening products and has been shown to cause ACD [14].

- Solvents such as ethyl acetate and isopropyl alcohol have been shown to cause ICD and ACD, albeit rarely. However, more recent studies have led many to believe that reactions to isopropyl alcohol are more common than once thought [24]. These ingredients can be found in both nail polishes as well as nail dehydrators.

- **Ethyl Cyanoacrylate (ECA)** is an ingredient in nail glue that has been associated with ACD, paronychia, and nail dystrophy [25,26]. While similar in name to (meth)acrylates (see below), evidence shows that there is no cross-sensitization between the two ingredients.

- **Methacrylic Acid (MAA)** is an ingredient in acidic nail primer that is known to be an extremely corrosive chemical. It can cause ACD if it accidentally comes in contact with the skin or cuticle on application [10]. Non-acid primers, those that do not contain MAA, have become more popular in order to avoid this adverse reaction; however, these still contain other allergens and irritants.
### Table 2. Common allergens and irritants in nail cosmetic procedures.

| Components                        | Common Allergens (ACD)                                                                 | Common Irritants (ICD)                        | Procedures That Use This Product |
|-----------------------------------|--------------------------------------------------------------------------------------|----------------------------------------------|---------------------------------|
|                                   | (Meth)acrylates                                                                     | Acetone                                      | Regular Manicure   Gel    Acrylic |
| Regular Color Polishing (includes base and top coat) | Benzophenone                                                                         | Ethyl Acetate                                | Yes    No    Likely         |
|                                   | Camphor                                                                              | Toluene                                      |                   |
|                                   | Dibutyl phthalate (DBP)                                                             |                                              |                   |
|                                   | Formaldehyde                                                                         |                                              |                   |
|                                   | Tosylamide epoxy resin                                                             |                                              |                   |
|                                   | Ethyl acetate (rare)                                                                |                                              |                   |
|                                   | Isopropyl alcohol (rare)                                                            |                                              |                   |
| Gel Color Polishing (includes base and top coat) | (Meth)acrylates                                                                     | Acetates                                     | No     Yes   Possible    |
|                                   | Camphor                                                                              |                                               |                   |
|                                   | Dibutyl phthalate (DBP)                                                             |                                               |                   |
|                                   | Photoinitiators                                                                     |                                               |                   |
|                                   | Formaldehyde                                                                         |                                               |                   |
|                                   | Acetates (rare)                                                                     |                                               |                   |
|                                   | Isopropyl alcohol (rare)                                                            |                                               |                   |
| Nail Strengthener (optional)      | Formaldehyde                                                                        | Possible                                     | Possible   Possible   Possible |
| Nail Polish Remover               | Acetone (rare)                                                                       | Ethyl Acetate                                | Yes     Yes    Yes      |
|                                   | Ethyl acetate (rare)                                                                |                                              |                   |
| Nail Cleanser                      | Isopropyl alcohol                                                                   | Possible                                     | Possible   Yes    Yes      |
| Nail Dehydrator (optional)        | Ethyl acetate (rare)                                                                | Ethyl Acetate (rare)                         | No      Yes    Yes      |
|                                   | Isopropyl alcohol (rare)                                                            |                                              |                   |
| Nail Primer (acid)                | (Meth)acrylates                                                                      | No                                           | Yes     Yes    Yes      |
|                                   | Methacrylate acid                                                                   |                                              |                   |
| Nail Primer (non-acid)             | Ethyl acetate (rare)                                                                | Ethyl Acetate (rare)                         | No      Yes    Yes      |
| Acrylic Preparation (liquid)      | (Meth)acrylates                                                                      | N/A                                          | N/A      Yes    N/A      |
|                                   | Hydroquinnone (occasional)                                                         |                                              |                   |
| Acrylic Preparation (powder)      | (Meth)acrylates                                                                      | N/A                                          | N/A      N/A   Yes      |
|                                   | Benzoyl peroxide (occasional)                                                      |                                              |                   |

### 3. Clinical Presentation

The classic clinical presentation of CD associated with nail cosmetic use (typically ACD) includes proximal nail fold, hyponychial or paronychial tenderness, edema, erythema, fissuring, and paresthesia [3]. The exact location of the reaction within the nail complex depends on where the aggravating product was applied. More severe cases of ACD, typically associated with gel and acrylic manicures, may also include paronychia, thickened and dry nail plates, and onycholysis, including hemorrhagic onycholysis (Figure 1) with total nail detachment. Nail technicians who are more frequently exposed to these ingredients can present with eczematous hand dermatitis [17].

In addition to the nails, CD may develop on ectopic body sites that the nails frequently come in contact with through touching, itching, or rubbing. This includes the face, neck, eyes and presents similarly to ACD and ICD caused by other triggers—erythematous, indurated, scaly plaques with possible vesiculation, fissuring, and edema. Some ingredients, such as acrylates, are also capable of becoming airborne, producing a more generalized, symmetrical airborne contact dermatitis [27].
A recent NACDG retrospective analysis of data from 2001 to 2016 found that the face was actually more commonly involved in nail-cosmetic-related CD compared to the hands and nails themselves (43% vs. 27.6% of cases, respectively) [9]. Tosylamide allergy alone was more associated with face involvement, whereas (meth)acrylate allergy was more highly associated with hand involvement. In addition, 14.4% of cases involved the eyelids, and 12.1% of cases were considered to have scattered/generalized distribution [9]. A separate retrospective review of European data from 2013 to 2015 found greater involvement of the hands (88.9%) than the face (36.8%, including eyelids, lips, and cheeks) [8].

4. Evaluation and Workup

The diagnosis of CD secondary to nail cosmetic use can often be made through a thorough patient history and physical exam, with special attention paid to the temporal relationship of nail cosmetic use/procedures to the onset of skin findings [28]. ACD will typically present 7–10 days after first/current exposure to the allergen, whereas ICD will present much sooner, often immediately or within a few days. Additionally, ICD tends to occur more on the dorsal plane of the hands, compared to the palmar aspect in ACD, and less frequently presents with vesicles and/or pain [29]. The differential diagnosis may also include atopic dermatitis and dyshidrotic eczema. The diagnosis of ACD can be confirmed by patch testing, although treatment and counselling may begin in the interim with a strong clinical suspicion of CD.

Patch testing is the standard diagnostic and confirmatory test for determining causative allergens in ACD and has both a sensitivity and specificity around 70–80% [30]. Up to 10% of all patients with ACD have a positive result for one or more ingredients in nail cosmetics [31]. However, as many commercially available patch testing kits do not include many of the antigens found in nail cosmetic products, additional customized testing using diluted versions of these ingredients should be performed for the most thorough results [3]. In fact, close to one-fifth of nail care product-associated allergens may be missed without additional screenings beyond the NACDG screening [9]. Suggested patch testing protocols for these ingredients can be found in Table 3. While broad-spectrum testing is useful for both identifying the current causative agent and future planning, suspected causative agents can be narrowed based on the procedure experienced and its included products (Table 2).
Table 3. Patch testing protocols for nail cosmetic ingredients [3].

| Allergen                                      | Recommended Patch Testing Protocol | Commercially Available Patch Test Series Including This Allergen |
|------------------------------------------------|------------------------------------|----------------------------------------------------------------|
| (Tosylamide) epoxy resin                      | 1% in petrolatum                   | T.R.U.E. Test, ACDS, NACDG                                     |
| Acetone                                       | 10% in olive oil [32]              | None                                                          |
| Acrylate Monomers (2-HEA, 2-EHA, 2-HPA, EA,   | 0.1–5% in petrolatum oil [33]      | ACDS, NACDG                                                   |
| TREGDA)                                       |                                    |                                                                |
| Benzophenone                                  | 10% in petrolatum [34]             | ACDS, NACDG                                                   |
| Benzoyl Peroxide                              | 10% in petrolatum [35]             | None                                                          |
| DBP                                            | 5% in petrolatum [36]              | None                                                          |
| Ethacrynic Acid                               | 10% in petrolatum [33]             | ACDS                                                          |
| Ethyl and Butyl Acetate                       | 10% in petrolatum [32]             | None                                                          |
| Formaldehyde                                  | 1% in aqueous solution             | T.R.U.E. Test, ACDS, NACDG                                    |
| Hydroquinone                                  | 2% in petrolatum [35]              | None                                                          |
| Methacrylate Monomers (2-HEMA, 2-HPMA, EGDMA, | 1–5% in petrolatum or olive oil    | ACDS, NACDG                                                   |
| MMA, THFMA)                                   | [33]                               |                                                                |
| Potassium or Sodium Hydroxide                 | 0.5% in aqueous solution [37]      | None                                                          |
| TSFR                                          | 10% in petrolatum [5]              | ACDS, NACDG                                                   |

T.R.U.E. (thin-layer rapid use epicutaneous) test contains 36 allergens, and the ACDS (American Contact Dermatitis Society) core series contains 80 allergens.

5. Treatment

5.1. Behavioral Interventions

Causative substance identification, removal, and avoidance is a mainstay of treatment for CD of any etiology. Ingredients that are more specific to certain products and/or nail procedures are likely easier to eliminate and, if desired, substitute. However, caution must still be taken. Patients must be aware that most polishes labeled “hypoallergenic” do not contain TSFR; however, these polishes typically still contain (meth)acrylates [38]. Additionally, some allergic to (meth)acrylates may find regular nail polish (that contains (meth)acrylates in much lower concentrations) to be a suitable substitution, while others continue to react. Some brands, such as CND Shellac and BRISA, claim to have created “acrylate-free” gel polishes, which have anecdotally been nonreactive in some with proven acrylate allergies. However, since cosmetics do not have ingredient regulating bodies, anyone with a history of CD secondary to nail cosmetic use should try new products with extreme caution [16].

For nail technicians and beauticians who occupationally come in contact with these allergens, latex gloves should be avoided as (meth)acrylates have been shown to penetrate through this material [39]. Rather, nitrile gloves should be used and switched every hour, as after this time frame, protection can diminish. Trilaminated polyethylene gloves are the most effective at protecting underlying skin from (meth)acrylates and other ingredients, but these are typically impractical [40].

5.2. Topical Anti-Inflammatory Treatment

For immediate treatment of CD skin lesions, topical steroids can be prescribed. Caution should be taken to prescribe a steroid of appropriate potency for the affected region, such as higher potency steroids for hand lesions and lower potency steroids for the more delicate the face [41]. Ointments may be chosen over creams as they are less likely to contain allergenic preservatives [28]. Alternatively, topical calcineurin inhibitors, such as pimecrolimus, tacrolimus, and cyclosporine, can be useful for ACD of the eyelids [35].
5.3. Emollients and Barrier Creams

Emollients and barrier creams, such as petrolatum, dimethicone, and paraffin, should be recommended in both treatment and prevention of CD by providing an extra layer of protection from allergens above the stratum corneum [42]. These creams have also been shown to have some anti-inflammatory properties as well as the ability to restore the skin barrier [41]. Hyaluronic acids can enhance intracellular lipid production, while ceramides help to retain moisture [33]. Occlusive dressings with emollients placed over affected areas (when practical) may help emollient adherence to skin and decrease subsequent exposure to allergens prior to healing.

6. Conclusions

As the popularity of nail cosmetic products and procedures continues to increase, nail-cosmetic-related CD has increasingly become a significant burden to patients, clinicians, and nail technicians. The high number of potentially allergenic ingredients in combination with the lack of a cosmetic regulatory body makes identification and elimination of triggering substances particularly challenging. As such, careful diagnosis, causative agent identification, treatment, and counseling are pertinent for positive outcomes.

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