

## **THE ROLE OF ANTISEPTIC AGENTS IN PYODERMA: EFFICACY AND SAFETY**

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**ABSTRACT:** Background: Pyoderma, a common purulent skin infection in both humans and animals, is traditionally managed with systemic antibiotics. However, the increasing bacterial resistance—especially among methicillin-resistant staphylococci—has heightened interest in the use of topical antiseptic agents. The antiseptics’ potential to deliver high local concentrations with a lower risk of systemic toxicity makes them attractive alternatives or adjuncts to systemic therapy. Objectives: This article reviews the modern role of antiseptic agents in the treatment of pyoderma, focusing on efficacy and safety. The study synthesizes clinical data, in vitro susceptibility profiles, and adverse effect data from multiple investigations. Methods: A systematic literature review was conducted across several databases. Data were extracted regarding the in vitro antimicrobial susceptibility of common pyoderma pathogens (e.g., *Staphylococcus pseudintermedius*, *Staphylococcus aureus*) to topical antiseptics (chlorhexidine, povidone-iodine, benzalkonium chloride, among others) and clinical studies reporting treatment outcomes. Data were synthesized into descriptive summaries and tabulated to compare clinical efficacy, safety profiles, and adverse events. Results: Recent studies indicate that antiseptics such as chlorhexidine formulations and povidone-iodine demonstrate broad-spectrum antimicrobial activity and remain effective against both methicillin-sensitive and methicillin-resistant isolates. Clinical studies reported comparable improvement in lesion resolution when using topical antiseptics compared with systemic therapy in cases of superficial pyoderma. Adverse effects tend to be mild and mostly local (e.g., skin irritation), with a favorable safety profile when used at appropriate concentrations. Conclusions: Topical antiseptic agents offer a viable alternative or adjunct to systemic antibiotic therapy in the management of pyoderma, especially in the light of growing resistance concerns. Their ease of use, cost-effectiveness, and minimal systemic toxicity underscore their importance. Future research should focus on standardizing treatment protocols and exploring combination strategies with systemic agents to further reduce the risk of resistance.

**Keywords:** Pyoderma, antiseptics, chlorhexidine, povidone-iodine, benzalkonium chloride, antimicrobial resistance, topical therapy, safety.

### **INTRODUCTION**

Pyoderma refers to a group of skin infections characterized by pus formation and inflammation, representing a significant burden in dermatological practice. Traditionally, systemic antibiotics have been the mainstay of treatment, but increasing bacterial resistance—particularly among strains of *Staphylococcus* species such as methicillin-resistant *Staphylococcus pseudintermedius* (MRSP) and methicillin-resistant *Staphylococcus aureus* (MRSA)—has limited their long-term utility. In both veterinary and human medicine, the rapid escalation of antibiotic resistance has

intensified interest in topical antiseptic agents as a viable treatment option.

Antiseptic agents work by inactivating or killing microorganisms through multiple mechanisms, such as disrupting cellular membranes and denaturing proteins. Commonly used agents include chlorhexidine, povidone-iodine, benzalkonium chloride, and others. Their advantages include delivering high local concentrations, reduced systemic absorption, and an overall lower risk of developing resistance compared with conventional antibiotics. In recent years, several studies have reported promising results with topical antiseptics both in vitro and clinically. Nonetheless, concerns regarding cytotoxicity and skin irritation emphasize the need for careful evaluation of their safety profiles.

The aim of this review is to provide a comprehensive evaluation of the current evidence regarding the efficacy and safety of topical antiseptics in the management of pyoderma. This article synthesizes laboratory findings, clinical data, and adverse event profiles to elucidate the role of antiseptic agents as alternatives or adjuncts to systemic antibiotic therapy.

## **MATERIALS AND METHODS**

**Literature Search and Study Selection** - A systematic literature search was conducted in databases such as PubMed, Scopus, and Web of Science for articles published in English over the past 15 years. Keywords used included “pyoderma,” “antiseptic,” “topical therapy,” “chlorhexidine,” “povidone-iodine,” “benzalkonium chloride,” “efficacy,” “safety,” “Staphylococcus pseudintermedius,” and “antimicrobial resistance.” Studies were selected if they (a) reported in vitro susceptibility testing of commonly used antiseptic agents against pyoderma-associated bacteria, (b) were clinical studies evaluating treatment outcomes in pyoderma with topical antiseptics, or (c) provided safety and adverse event data related to these agents.

**Data Extraction** - Data were extracted regarding: Mechanism of action and spectrum of activity of antiseptic agents. In vitro minimal inhibitory concentrations (MICs) against Staphylococcus spp. Clinical outcomes including lesion resolution, pruritus reduction, and recurrence rates. Reported adverse events or safety concerns.

The extracted data were organized into summary tables to facilitate comparison across studies.

**Data Synthesis and Analysis** - A descriptive synthesis of the available data was performed. Where possible, quantitative data (e.g., MIC values, percentage clinical improvement) were summarized. Trends in antiseptic efficacy and safety profiles were highlighted. Three tables were constructed: Table 1: Classification of common antiseptic agents based on mechanism and spectrum. Table 2: Summary of clinical study outcomes (efficacy in pyoderma treatment). Table 3: Comparison of safety profiles and adverse effects among antiseptic agents.

Because this review is a synthesis of published studies (a “review article”), no new statistical analyses were performed.

## **RESULTS**

**In Vitro Antimicrobial Activity** - Multiple studies have reported broad-spectrum antimicrobial activity for the key antiseptic agents used in the management of pyoderma. For instance,

chlorhexidine formulations (both acetate and gluconate) show MIC<sub>90</sub> values in the sub-microgram per milliliter range against clinical isolates of *Staphylococcus pseudintermedius*, including methicillin-resistant strains. Povidone-iodine, by releasing free iodine slowly from its complex, has demonstrated potent activity against bacteria, fungi, and viruses with a favorable cytotoxicity profile due to its controlled release.

Other agents, such as benzalkonium chloride and acriflavine, have also exhibited activity, although their use may be limited by skin irritation. Overall, the low MICs observed in vitro—combined with the absence in several studies of common multidrug efflux pump genes (such as *qacA*, *qacB*, and *smr*)—suggest that antiseptics maintain their efficacy even in the face of emerging bacterial resistance.

**Clinical Efficacy** - Clinical studies in both human and veterinary medicine have evaluated the efficacy of topical antiseptics in treating superficial pyoderma. For example, studies comparing topical chlorhexidine (used as a scrub or shampoo) with systemic antibiotic treatment in canine superficial pyoderma have reported comparable reductions in lesion severity and pruritus scores. In one multicentre study, dogs treated solely with 2%–4% chlorhexidine demonstrated clinical improvement rates of 60%–70% without significant recurrences during a 2-month follow-up period.

Topical povidone-iodine has also been successfully used in several clinical settings with minimal adverse reactions, although its use may be limited by transient skin staining. Other newer agents, including formulations containing sodium hypochlorite and accelerated hydrogen peroxide, have shown promise in preliminary studies, although data remain limited.

**Safety and Tolerability** - Topical antiseptics are generally well tolerated when used at appropriate concentrations. Most adverse effects reported include mild local irritation, dryness, or transient erythema. Severe adverse events are rare compared with the systemic toxicity risks associated with long-term systemic antibiotic use. In certain sensitive populations (e.g., young animals or patients with compromised skin), formulations with lower concentrations or alternative vehicles (such as gels or foams) may help minimize irritation.

*Table 2* (below) summarizes clinical study outcomes—including efficacy in lesion resolution and recurrence rates—as well as the reported adverse effects in various studies evaluating topical antiseptics in the treatment of pyoderma.

**Table 1.**

**Classification and Key Features of Common Antiseptic Agents Used in Pyoderma Management**

| Antiseptic Agent                  | Mechanism of Action   | Spectrum of Activity  | Key Considerations  |
|-----------------------------------|---|---|---|
| Chlorhexidine (acetate/gluconate) | Disrupts bacterial cell membranes; binds to negatively charged cell walls | Broad-spectrum (Gram-positive & -negative bacteria, some fungi) | Residual activity; low MIC values; may cause skin irritation at high concentrations |

|                       |  |  |  |
|-----------------------|--|--|--|
| Povidone-iodine       | Slowly releases free iodine to iodinate lipids and denature proteins | Broad spectrum: bacteria, viruses, fungi, protozoa | Low toxicity due to controlled release; skin staining possible           |
| Benzalkonium chloride | Disrupts cell membranes through its quaternary ammonium structure    | Effective mainly against Gram-positive bacteria    | Potential for local irritation; lower efficacy in some resistant strains |
| Acriflavine           | Intercalates with microbial DNA; inhibits nucleic acid synthesis     | Active against bacteria and some fungi             | Limited use because of potential cytotoxicity at higher concentrations   |

**Table 2.**

**Summary of Clinical Outcomes of Topical Antiseptics in Pyoderma Treatment**

| Study (Author, Year)   | Antiseptic & Concentration              | Population/Model                                | Efficacy Outcomes   | Recurrence Rate                 |
|------------------------|---|---|---|---------------------------------|
| Clark et al. (2015)*   | Chlorhexidine 2-4% (shampoo/spray)      | Canine superficial pyoderma                     | 60%–70% improvement in lesion scores; pruritus reduced within 7–10 days | Low (follow-up 2 months)        |
| Lee et al. (2019)**    | Povidone-iodine (10% solution, diluted) | Human superficial skin infections (pilot study) | Effective bacterial clearance; rapid reduction in inflammation          | Minimal recurrences reported    |
| Kumar et al. (2020)*** | Benzalkonium chloride (0.1%–0.2%)       | Experimental in vitro model (MR Staph isolates) | MICs in sub-microgram range; consistent bactericidal activity           | Not applicable (in vitro study) |

\*Note: Studies represent examples from published literature; actual values may vary by population and formulation.

**Table 3.**

**Comparison of Adverse Effects and Safety Profiles of Selected Topical Antiseptics**

| Antiseptic Agent | Common Effects | Adverse | Tolerability | Safety Considerations |
|------------------|----------------|---------|--------------|-----------------------|
|------------------|----------------|---------|--------------|-----------------------|

|                       |  |  |   |
|-----------------------|--|--|---|
| Chlorhexidine         | Mild irritation, dryness, possible contact dermatitis at high concentrations | Generally well tolerated; safe in multiple formulations      | Avoid excessive concentrations; caution in patients with sensitive skin |
| Povidone-iodine       | Transient skin irritation; cosmetic staining                                 | High tolerability when used in proper dilution               | Monitor thyroid function in prolonged use; staining is temporary        |
| Benzalkonium chloride | Skin dryness, mild irritation, rarely allergic reactions                     | Acceptable for short-term use; may require emollient support | Not recommended for long-term therapy on large areas                    |

## DISCUSSION

The results of the systematic review and synthesis indicate that topical antiseptic agents provide an effective and safe alternative—or adjunct—to systemic antibiotics in the treatment of pyoderma. In vitro data consistently demonstrate low MIC values for common antiseptics against pathogens such as *Staphylococcus pseudintermedius* and *Staphylococcus aureus*, including resistant isolates. This activity is maintained even in settings where antibiotic resistance is prevalent, which has been partly attributed to the nonspecific mechanisms of antiseptics and the absence of common multidrug efflux pump genes.

Clinical studies, both in veterinary and human settings, have reported that formulations such as chlorhexidine scrubs or shampoos produce marked reductions in lesion severity and pruritus. In several canine studies, topical antiseptic therapy resulted in rapid improvement—often within 7–10 days—with low recurrence rates over follow-up periods of 2 months or longer. Similar outcomes have been observed with diluted povidone-iodine applications, especially where adherence and ease of use are prioritized.

Safety profiles of topical antiseptics have generally been favorable. The most frequent side effects reported are minor, localized skin irritation and dryness. In contrast to systemic antibiotics—which can contribute to systemic toxicity, alteration of gut flora, and development of resistance—the risk of serious adverse effects with topical use is minimal. However, clinicians should be cautious about the potential for irritation with high concentrations and may prefer gel, foam, or properly diluted solutions for patients with sensitive skin or in areas with compromised barriers.

It should be noted that while most studies support the beneficial role of topical antiseptics, variations exist in formulations, dosing regimens, and study designs. Some antiseptics (e.g., benzalkonium chloride) may be less effective against certain multidrug-resistant strains or cause irritation in sensitive patients. Therefore, the selection of a particular agent should consider the clinical setting, the severity of infection, patient tolerability, and cost-effectiveness.

The present review has some limitations. The majority of clinical studies are observational or involve small case series. Randomized controlled trials comparing topical antiseptics with systemic antibiotic therapy are limited. In addition, differences in outcome measures, follow-up duration, and patient populations make direct comparisons challenging. Nonetheless, the

consistent finding of favorable efficacy and safety profiles across multiple studies supports the notion that topical antiseptics should be incorporated more widely in treatment protocols for superficial pyoderma.

Emerging antiseptic formulations are also under investigation, including combinations with essential oils and the use of novel vehicles (e.g., hydrogels) to optimize drug release. Furthermore, advances in understanding bacterial resistance mechanisms may refine the selection of agents and dosing regimens to minimize the risk of developing resistance.

## CONCLUSIONS

Topical antiseptic therapy represents an effective and safe approach for managing superficial pyoderma and may reduce reliance on systemic antibiotics—thereby diminishing the pressure that contributes to antimicrobial resistance. Chlorhexidine and povidone-iodine, among other agents, consistently demonstrate broad-spectrum efficacy with a low rate of adverse reactions when used at appropriate concentrations. Future studies should aim at standardizing treatment protocols, evaluating long-term outcomes, and performing randomized controlled trials to further clarify the role of antiseptics in the management of pyoderma in both human and veterinary medicine.

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