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Pharmacology

Topical steroids, keratolytics and Antipruritics

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Preparations of topical cutaneous steroids

1- Ointments

- ☐ The medical substance presents in a **greasy** vehicle.
- □ Ointments are better administered to thick hyperkeratotic and dry lesions (e.g. chronic eczema).
- □ It is the most potent vehicle (they are the most occlusive).
- ■Avoid ointments on <u>hair-bearing regions</u> because it may result in <u>folliculitis</u>.
- 2- Creams: less potent than ointment but cosmetically better (not greasy).
- 3- Lotions: less occlusive and less greasy; better in hair-bearing regions.
- 4- Gels like lotions, and more beneficial for the scalp as they do not cause greasy hair.
- 5- Foams highly effective for steroid delivery to the scalp but they are expensive. . 1←2←3←4←5 "צמוף" אל אליילאל—

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Mechanisms of action Glucocorticoid activity

- 1- Anti-inflammatory (inhibition of phospholipase A2).
- 2- Immunosuppressant (decreasing lymphocytes)
- 3- Antiallergic effects.
- 4- Antimtotic effects on epidermal cells.
- 5- Decrease Collagen cross linking.

Mineral ocorticoid activity

Vasoconstriction which leads to poor healing.

The <u>vasoconstrictor properties</u> determines the potency of steroid.

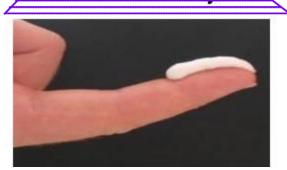
The European classification of topical steroids

- Weak (e.g. Hydrocortisone).
- Moderate (e.g. Clobetasone butyrate).
- Potent (e.g. Mometasone furoate & Betamethasone esters).
- Very potent (e.g. Clobetasol propionate). الما ف و العملات

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Pharmacokinetics

- ➤ Topical steroids are absorbed to the circulation when applied to thin epidermis (e.g. the eyelid, genital skin, and face) up to 300 folds compared to thicker regions of epidermis (like palms and soles).
- ➤ The penetration increases two- to ten-fold in diseased states (e.g. inflammation, scratches, and desquamation).
- ➤ The skin of children is more permeable to topical steroids.
- > Avoid High-potency steroids in face, gentalia, & intertriginous areas.
- ➤ The amount of the applied steroid used affects its absorption. One fingertip unit (FTU) is equal to 0.5 grams.
- ➤ Topical corticosteroids are recommended for once twice daily use.



Therapeutic uses

- ✓ Topical steroids are FDA-approved for treating:
- 1-Psoriasis 2-Eczema 3-Lichen planus
- 4-Discoid lupus erythematosus
- 5- Limited areas of vitiligo
- 6-Atopic dermatitis
- 7-Acute radiation dermatitis
- 8-Lichen simplex chronicus (neurodermatitis)
- 9-lichen sclerosis.
- ✓ Topical steroids are also effective for treating:
- 1- Hyper-proliferation disorders (e.g. hyper-proliferation disorders (e.g. hyperkeratosis & skin tumors).
- 2- Immunological disorders (e.g. alopecia areata, and bullous pemphegoid).
- 3- Inflammatory conditions (contact dermatitis & sun burns).
- 4- Allergic conditions (e.g. **urticaria** and some drug eruptions).
- 5- Topical steroids are also used for treating pityriasis rosea, and granulomas like sarcoidosis.

Adverse effects of topical corticosteroids

1-Local adverse effects (skin)

It occurs with **prolonged use**& depends on the **potency**, **vehicle**, and **application site of topical steroid**. They include:

- 1- Delayed wound healing.
- 2- Skin atrophy and striae. Collagen
- 3- Rosacea
- 5-Acne.
- 6-Spread of skin infections (viral, fungal, bacterial or parasitic)
- 7-Hypertrichosis.
- 8-Pigment alteration (<u>leukoderma</u> occurs with intralesional injection of **triamcinolone**).
- 4- Peri-oral dermatitis.
- 9- Contact allergy with hydrocortisone.



Steroid rosacea (erythema with or without pustules)



Skin atrophy is the most common adverse effect and occurs due to the anti-mitotic effect of topical corticosteroids &

The loss of connective tissue



Striae (stretch marks)

- ➤ Tachyphylaxis: the skin is developing tolerance to topical steroids, which leads to loss of vasoconstriction at the level of the capillaries.
- Addiction of topical steroid is a common clinical problem usually presented with rosacea, thin facial skin and irritation.
- ✓ Gradual withdrawal of topical steroids and <u>replacing with</u> topical calcineurin inhibitors help the improvement.

2- Systemic adverse effects

In Children

They are less likely to occur due to low cutaneous absorption
They occur on <u>prolonged</u> use of <u>high-potency</u> steroids on <u>thin</u>
<u>epidermal</u> regions or in <u>children</u>.

☐ The systemic adverse effects include: glaucoma, hypothalamicpituitary axis suppression, Cushing syndrome, hypertension, and hyperglycemia.

Contraindications of topical steroids

- 1-Bacterial infections (e.g. impetigo, cellulitis, erysipelas, etc).
- 2- In fungal infection including candida and dermatophytes.
- 3- Viral infections like herpes.
- 4- Wounds and ulcers.
- 5- Previous history of allergy to the preparation.

Topical calcineurin inhibitors
(tacrolimus & preclorimus) can be used alternative to topical steroids in different dermatological diseases (especially vitiligo, and atopic dermatitis).

Calcineurin inhibitors will not cause skin atrophy, srtia, rosecia, or tachyphylaxis.



Keratolytic agents

- Keratolytic agents break down the outer layer of the skin.
- They help skin to bind moisture.
- They can dissolve skin flakes and scales.
- ➤ Removing the scales & minimizes dandruff
- ➤ Help other topical medications to penetrate the skin better.

Side Effects of Keratolytic agents:

- Reddening of the skin____
- Reducting of the Sensation
 Burning or tingling sensation
 Sensitivity to UV light Apple
- Peeling of the skin.

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1- Lactic acid

It exfoliates the skin, stimulate skin cell renewal & keeps the skin hydrated

Over-the-counter creams and lotions that contain lactic acid help to relieve symptoms of psoriasis and keratosis pilaris "chicken skin".

Brittle = Unea Nails aeolly

2- Urea

very

Urea is an organic compound known to soften the skin.

Urea is very safe even in high conc.

It also has **hydrating and anti-itching properties**, and helps other medications to penetrate the skin.

Topical application of urea in a cream or gel has been shown to effectively treat **atopic dermatitis**, <u>eczema</u>, <u>xerosis</u> (dry skin), <u>psoriasis</u>, <u>keratosis</u> <u>pilaris</u> as well as **brittle nails**.

3- Salicylic acid

It is used to treat common conditions like <u>acne</u>, <u>psoriasis</u>, and <u>dandruff</u>. Salicylic acid works as a keratolytic by <u>breaking down connections</u> <u>between dead cells on the surface of the skin</u>.

It is usually applied to the skin surface as a **cream or lotion** of small concentrations (e.g. **0.5-1%)** for <u>acne and psoriasis</u>, or <u>higher conc</u> (up to 30%) to treat warts and corns or callus.

Other examples of keratolytic agents:

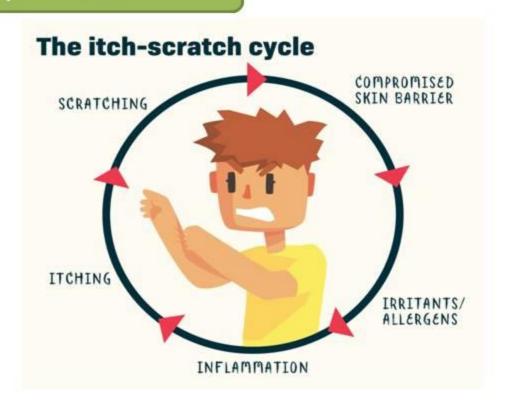
- 1- Topical tretinoin and oral acitretin,
- 2- Topical coal tar.
- 3- Glycolic acid,
- 4- Pyrithione zinc
- 5-All chemical peels

Antipruritics

Antipruritics, or anti-itch drugs, are medications that inhibit itching (pruritus).

Treatment of itching:

- Treat the cause.
- 2- General measures.
- 3- Topical medications.
- 4-Systemic medications.



General measures

Moisturization

Cool environment

Avoid irritants

Break itch-scratch cycle

Behavioural therapy, relaxation, stress reduction



Calamine

- □Calamine (means Zinc) lotion is an over-the-counter (OTC) medicine to treat mild itchiness (pruritus).
- □ Calamine lotion contains <u>zinc oxide</u> & 0.5% <u>ferric oxide</u> (the iron oxide gives calamine lotion its distinctive pink color).
- □It may contain purified water, Glycerin, Calcium hydroxide and Phenol.

Topical medications for itching

sycologecal Uses

- Topical corticosteroids especially in inflammatory skin diseases
- Topical calcineurin inhibitors.
- ➤ Topical Capsaicin is an active component of chili peppers, It is a chemical irritant & triggers heat sensation from skin.
- Topical Menthol (from mint) chemically trigger the cold-sensitive receptors in the skin (cooling sensation), Menthol's analgesic properties are due to activation of κ-opioid receptors and blocking of voltagesensitive sodium & calcium channels.
- ➤ <u>Topical local anesthetics like</u> <u>Pramoxine</u> (pramocaine), Prilocaine & Lidocaine.
- Doxepin is a medication belonging to the tricyclic antidepressant.





Systemic treatment

- To fo
- 1- Non-sedating antihistamines: Urticaria, insect bite reactions and drug reactions.
- 2- First-generation antihistamines (sedating) for Nocturnal itch.
- 3- μ-Opioid receptor antagonists for <u>Cholestatic</u> pruritus & atopic dermatitis.

4- κ-Opioid receptor agonists for uremic pruritus.

5- Antidepressants (paroxetine, and sertraline) palliative treatment in atopic dermatitis, lymphoma, solid carcinoma, uremic and cholestatic pruritus,

- 6-Doxepin in atopic dermatitis, HIV-related pruritus, allergic reactions, and urticaria.
- 7-Anticonvulsants (gabapentin & pregabalin) Uremic, Neuropathic pruritus.
- 8-Ursodeoxycholic acid for Intrahepatic cholestasis of pregnancy.
- 9- Oral <u>immunosuppressants</u> (cyclosporine, Azathioprine and Mycophenolate mofetil) in inflammatory dermatoses.
- 10 Systemic corticosteroids in inflammatory dermatoses.

