### NIZORAL: AN EFFECTIVE TREATMENT OF SEBORRHEIC DERMATITIS AND DANDRUFF

A ray of hope for hair loss associated with these disorders?



## Seborrheic dermatitis and dandruff: Two interconnected skin/scalp disorders

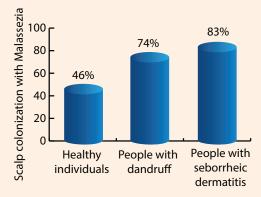
- Dandruff and seborrheic dermatitis are two related manifestations of the same dermatological disorder that affects seborrheic areas of the body (areas rich in sebaceous glands)<sup>1</sup>
- Dandruff involves itchy, flaking skin of the scalp without visible inflammation (erythema); seborrheic dermatitis, on the other hand, is associated with flaking or scaling of the skin, erythema (inflammation) and itching in the scalp and other seborrheic areas (behind the ear, face and upper chest)<sup>1</sup>
- Both seborrheic dermatitis and dandruff are associated with white to yellow flakes that can be dry or moist<sup>1</sup>
- Seborrheic dermatitis and dandruff affect 50% of the adult population worldwide<sup>1</sup>
- Dandruff is regarded as a milder and non-inflammatory form of seborrheic dermatitis; the latter often has a chronic recurrent disease course<sup>2</sup>





### The link between Malassezia and seborrheic dermatitis/dandruff

- Malassezia, a commensal fungus, is among the normal inhabitants of a healthy scalp<sup>3,4</sup>
- However, Malassezia is also regarded as one of the main causative factors in the development of seborrheic dermatitis and dandruff<sup>3</sup>
- Overgrowth of Malassezia on the scalp is reported in patients with seborrheic dermatitis and dandruff<sup>4</sup>



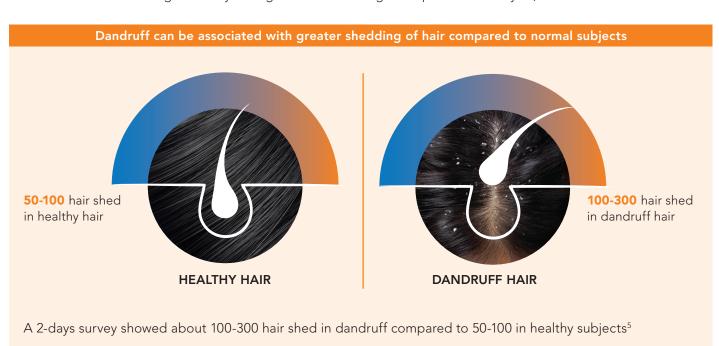
In a study, Malassezia colonization of the scalp was noted in 46% of healthy individuals, 74% in patients with dandruff and 83% in cases of seborrheic dermatitis<sup>4</sup>

### Seborrheic dermatitis and dandruff - Malassezia colonization and premature hair fall<sup>3</sup> Malassezia colonization of the scalp in seborrheic dermatitis and dandruff Fungal metabolite, free fatty acids accumulate Alteration of antioxidants in seborrheic dermatitis and dandruff Inflammation and oxidative stress Oxidative stress may accelerate aging process of dermal papilla cells and hair follicles; aged hair follicle stem cells show longer resting and shorter growth phases, may be slow in proliferation, and may be slow in tissue-regeneration Oxidative stress can increase lipid oxidation; oxidized lipids negatively influences hair growth Oxidative stress also adversely impact pre-emergent hair (inside the skin); anchoring forces of the hair fiber to follicles is altered

### Seborrheic dermatitis and dandruff – Other mechanisms contributing to hair fall

Premature hair fall

- Dandruff can also alter the exogen phase of the hair cycle (hair shedding from the scalp)<sup>5</sup>
- Co-existing dandruff may therefore increase hair shedding in androgenetic alopecia (male- and female-pattern baldness)<sup>5</sup>
- Additionally, seborrheic dermatitis and dandruff may also lead to prominent hair eclipse phase (latency period between hair shedding and early emergence of the next growth phase of hair cycle)<sup>5,6</sup>



## Targeting seborrheic dermatitis and dandruff with NIZORAL (ketoconazole) shampoo



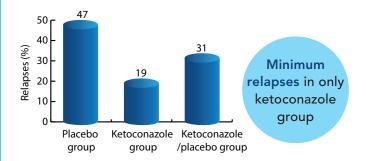
- Ketoconazole is an imidazole antifungal agent with activity against Malassezia spp along with direct anti-inflammatory effect<sup>7</sup>
- Ketoconazole has been shown to have high in vitro and in vivo efficacy against Malassezia<sup>8</sup>
- Ketoconazole 2% shampoo is advised to be used twice weekly over four weeks to improve symptoms of seborrheic dermatitis<sup>7</sup>
- Intermittent once weekly use of ketoconazole 2% shampoo is effective in preventing relapse of seborrheic dermatitis<sup>7</sup>
- Safety of ketoconazole 2% shampoo
  - » Good safety profile
  - » Low incidence of irritation and contact sensitivity<sup>7</sup>

# Efficacy of ketoconazole 2% shampoo in managing seborrheic dermatitis and dandruff and preventing relapse<sup>8</sup>

Efficacy of ketoconazole 2% shampoo vs placebo evaluated in patients with seborrheic dermatitis and dandruff



Responding patients assigned to prophylaxis phase for 6 months. Three groups (only ketoconazole 2% shampoo weekly, ketoconazole 2% and placebo alternate week, and only placebo weekly)



## By effectively treating seborrheic dermatitis and dandruff, can NIZORAL (Ketoconazole 2%) shampoo reduce hair fall?

- Ketoconazole 2% shampoo is approved for treating dandruff and seborrheic dermatitis of the scalp. It is not approved for treating hair loss or alopecia<sup>9</sup>
- However, there is some evidence to show that by improving scalp health (rebalancing scalp microbiome), it may reduce hair loss associated or exacerbated by seborrheic dermatitis and alopecia (indirect benefits)<sup>9,10</sup>

### Ketoconazole - Indirect mechanisms on reducing hair fall

- Scalp colonization with lipophillic flora, such as Malassezia, can trigger
  inflammation/oxidative stress leading to hair loss; since ketoconazole is a potent
  antifungal with activity against Malassezia spp., it may support hair growth by
  reducing scalp inflammation due to this fungus<sup>9,10</sup>
- Topical applications of ketoconazole may therefore be presumed to be useful for treating dandruff/seborrheic dermatitis-associated male pattern hair loss<sup>10</sup>
- Dandruff is associated with reduced proportion of anagen hair (active growth phase) and thin hair shaft diameter. Ketoconazole shampoos appear to increase the anagen hair percentage in subjects with dandruff, thereby it may reduce hair shedding associated with dandruff.<sup>11</sup>



### Indirect benefits of Nizoral on seborrheic dermatitis- and dandruffassociated hair fall





Being a potent antifungal with activity against Malassezia spp., it may reduce scalp inflammation and oxidative stress due to this fungus thereby reducing hair fall



Appear to increase anagen hair (hair in active growth phase) in subjects with dandruff, thereby reducing hair shedding





Indirect benefits on hair fall reduction by improving scalp health, treating seborrheic dermatitis/dandruff effectively and rebalancing scalp microbiome.

Daily application of Nizoral Expert (Crinipan PMC) can maintain healthy scalp balance and reduce relapse of seborrheic dermatitis/dandruff.

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