

cyspera[®]

PIGMENT CORRECTION

HOPE FOR
YOUR SKIN



INTENSIVE PIGMENT CORRECTION SYSTEM

« Pigmentary disorders can be psychologically devastating »



Brown patches are #1 aesthetic concern of Latin type women



Dark spots are #1 aesthetic concern of West Asian & African type women



Facial Dyschromia is #1 aesthetic concern of East Asian women

Cyspera Cysteamine 5% has become the first-line treatment, free of hydroquinone

Hydroquinone has been used for decades as the gold standard for stubborn discolouration concerns. Although still controversial, hydroquinone has a reputation as a mutagenic and carcinogenic compound. It is a melano-cytotoxic agent that is known to induce ochronosis. Non-hydroquinone compounds such as kojic acid, arbutin, azelaic acid, retinoids, hydroxy acids show lower efficacy.



Significant efficacy on melasma

2015, first double-blind, randomised clinical study on 50 epidermal melasma patients.² Other clinical studies have confirmed results.³
– **British Journal**

Significant response of Kligman's resistant patients

Melasma lesions almost disappearing and signs of skin atrophy reversing after treatment with topical cysteamine.⁴
– **Journal of Cosmetic Dermatology**

Greater depigmenting effects than HQ4% Comparison of Cyspera vs. HQ4% in a randomised controlled trial. Investigator concludes Cyspera as an alternative to avoid or rotate off HQ4%⁵
– **Australian Journal of Dermatology**

Better efficacy & tolerability than Kligman Formula Comparison of Cyspera vs. modified Kligman Formula in a randomised clinical study of 50 melasma patients⁶
– **Skin Research Technology**

As effective with less complication than TXA mesotherapy Comparison of Cyspera vs. Tranexamic acid mesotherapy in a randomised clinical study of 54 melasma patients⁷
– **Archive of Dermatological Research**

Scientis introduces the new Cysteamine Isobionic-Amide Complex™

The novel Cysteamine Isobionic-Amide Complex™ activates powerful synergistic effects.

Cysteamine + Isobionic-Amide

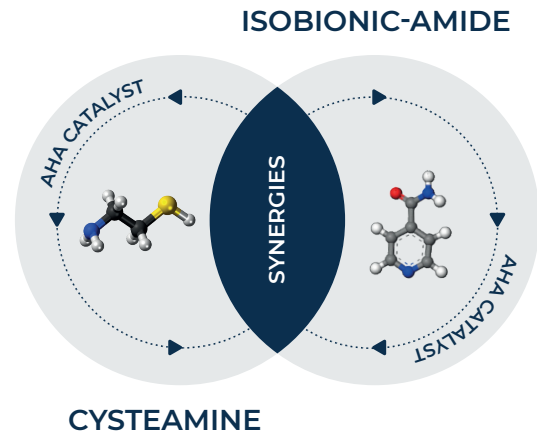
Isobionic-Amide multiplies the pigment correction efficacy of Cysteamine by inhibiting multiple steps in the melanogenesis pathway^{10,11,12}

Cysteamine + AHA

AHA stabilises Cysteamine, which enhances its permeability into the skin and reduces its smell on skin^{13,14,15}

Isobionic-Amide + AHA

AHA reacts immediately with Isobionic-Amide, which produces an instant pigment lightening¹⁶



Cyspera Intensive System™ is clinically proven to be fast and effective against hyperpigmentation without the concerns of corticosteroids, retinoic-acid and hydroquinone.



- ✓ Significant improvement in the appearance of discolouration: 18% more effective¹⁸
- ✓ Fast onset of action: 10% better evening of skin tone within 4 weeks¹⁷
- ✓ Well tolerated for long-term use: 4x better tolerated after 16 weeks¹⁸
- ✓ Powerful antioxidant activity for improved skin health
- ✓ Significant improvement in quality of life

Long term efficacy and tolerability

The new Cysteamine Isobionic-Amide Complex™ leverages the long-term efficacy from Scientis Cysteamine 5% and shows fast onset of action and better evening of skin tone within 4 weeks of use.



Before



Results after 16 weeks



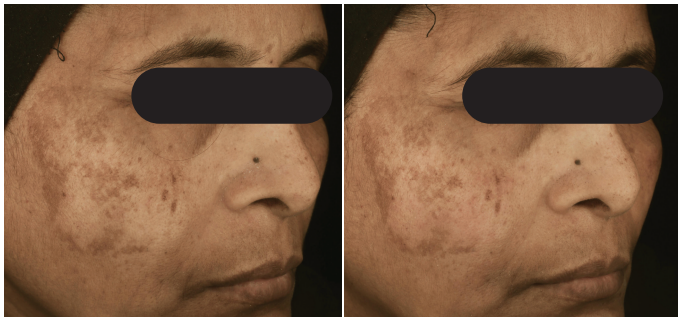
Results after 5 years

18% MORE EFFECTIVE, 4X BETTER TOLERATED AND SUITABLE FOR LONG-TERM MAINTENANCE¹⁸

Fast onset of action

Baseline

Week 4



Cyspera Intensive System™



Traditional "Gold-standard" Triple Combination

Clinical Study Observations at week 4¹⁷:

- Cyspera® acts rapidly on pigment correction with significant mMASI Score reduction, on par with Triple Combination in comparison to baseline already within 4 weeks
- Cyspera® is more effective at evening skin tone already within 4 weeks, with 10% greater reduction of contrast between lesional and non-lesional areas
- Cyspera® improves the quality of life of patients already within 4 weeks, with condition being significantly less bothersome after one month of regular test product usage

Clinical cases of pigmentation concerns

BROWN PATCHES



Results after 8 weeks. 15 minutes daily application of Cyspera Intensive System™.
Courtesy from Lesley Clark-Loeser, MD

DARK SPOTS



Results after 4 weeks. 15 minutes daily application of Cyspera Intensive System™.
Courtesy from Corey Hartman, MD

FACIAL DYSCHROMIA



Results after 8 weeks. 15 minutes daily application of Cyspera Intensive System™.
Courtesy from Lesley Clark-Loeser, MD

A 3-step approach for multiple effects

CYSPERA INTENSIVE™

Cysteamine Isobionic-Amid Complex™ delivers a powerful antioxidant activity to improve the appearance of persistent brown patches and dark spots.^{9,17}

1. **Cysteamine** reduces melanin production in the melanogenesis pathway at multiple locations
2. **Cysteamine with AHA**. AHA creates an acidic environment enhancing cysteamine permeability into the skin²¹
3. **Isobionic-Amide with AHA** immediate reaction leads to instant pigment lightening²²



Clinically-proven to be more effective and faster than the gold standard triple combination without the concerns of corticosteroids, retinoic-acid and hydroquinone

- ✓ 18% more effective and 4x better tolerated after 16 weeks²⁰
- ✓ Fast onset of action and 10% better evening of skin tone already within 4 weeks¹⁷
- ✓ Significantly improves the quality of life of patients already within 4 weeks¹⁷

CYSPERA NEUTRALIZE™

AHA-Amino Acid Complex neutralises the smell of cysteamine and re-balance the epidermis before application of Cyspera Boost™²²

1. **L-Arginine Complex** stops Isobionic-Amide / AHA reaction
2. **Mild-Surfactant medium** gently removes lingering residue, while helping soothe skin and promoting healthy skin barrier
3. **Lactobionic Acid** prepares the skin for Cyspera Boost™ by neutralising and balancing the skins pH



- ✓ **Neutralise smell of cysteamine:** 77% of the patients think it neutralises the smell of cysteamine on the skin²²
- ✓ **Re-balance:** 90% of the patients think it re-balances the skin after use of Intensive²²
- ✓ **Cleanse:** 90% of the patients think it deeply cleanses the skin²²
- ✓ **Pleasant to use:** 87% of the patients find it pleasant to use after Intensive²²

CYSPERA BOOST™

The Isobionic-Amide Complex™ works synergistically with Cyspera Intensive™ to even skin tone, improve complexion and deliver a natural and healthy glow.¹⁷

1. **Isobionic-Amide** multiplies the efficacy of Cysteamine by inhibiting multiple steps melanogenesis in the skin²⁴
2. **Retinol and an anti-oxidative complex** provides a visual pigment correction effect by increasing non-pigmented skin layers and shading²⁵
3. **Isobionic-Amide with Retinol** provides an anti-inflammatory effect to the skin after application of Cyspera Intensive™²⁶



- ✓ **Brightening:** 81% of the patients noticed improvement of skin complexion, and 71% of the patients find their skin more radiant and luminous²³
- ✓ **Skin Health:** 84% of the patients find their skin smoother, and 77% of the patients feel their skin is visibly healthier²³
- ✓ **Pleasant use:** 94% of the patients find the product easy to apply, not sticky and easy to combine with other products²³
- ✓ **Pleasant smell:** 90% of the patients find the smell pleasant²³

Mode of use - 15-minute short application

Intensive phase: Once per day, everyday, for 16 weeks or until the desired results are achieved.

Maintenance phase: Once per day, twice-weekly.



Always apply Cyspera Intensive System™ on rested skin.


Do not wash the skin before application. If necessary to wash, wait one hour before application.



15 mn

Apply one pump as a thin layer on the skin. Leave on for 15 minutes (a warming sensation or mild tingling may occur and last up to 30 min).


**Step 1
INTENSIVE**



2

Apply 2 pumps on unwashed face. Gently massage in a circular motion. Rinse thoroughly with water and pat dry.

**Step 2
NEUTRALIZE**



3

Apply a thin layer to face.

**Step 3
BOOST**

It is important for the skin to acclimate to Cyspera Intensive™ by limiting initial use to 5 minutes once per day. After one week, increase to 15 minutes exposure (or advised by your skincare professional).

In case of irritation, discontinue use for a few days until your skin recovers, then restart application of Cyspera Intensive™ for 5 minutes.

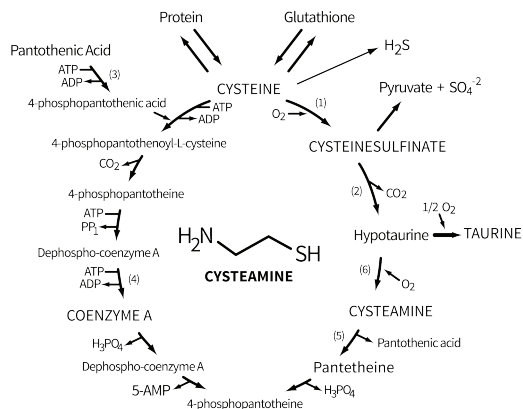
Use can be tailored to meet the needs of an individual patient and their routine. It can be applied in the morning or evening, offering patients true flexibility and personalised choice.

For optimal results, daily use of a broad-spectrum sunscreen with SPF 30 or higher is highly recommended.

Caution : For external use only. Do not use on damaged or irritated skin. Do not use in case of known allergy to a component. Discontinue use and ask a doctor if signs of irritation or allergy appear. Avoid contact with eyes. In case of eye contact, rinse off with clear water. Keep out of reach of children. Do not use for pregnant or lactating women. Use an adapted solar protection in case of sun exposure.

Cysteamine

50 years of proven efficacy

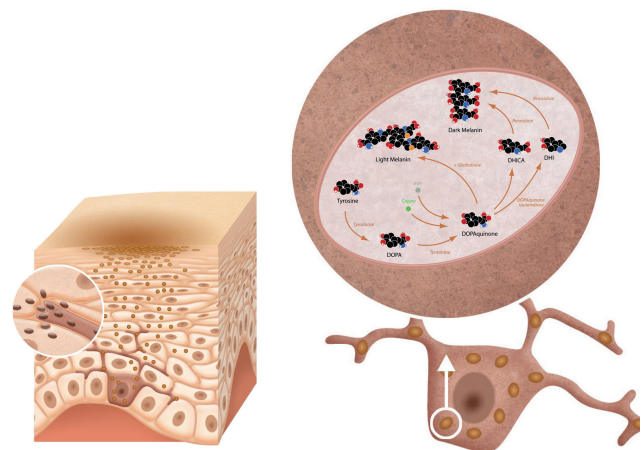


Cysteamine is biologically present in human tissue¹⁹

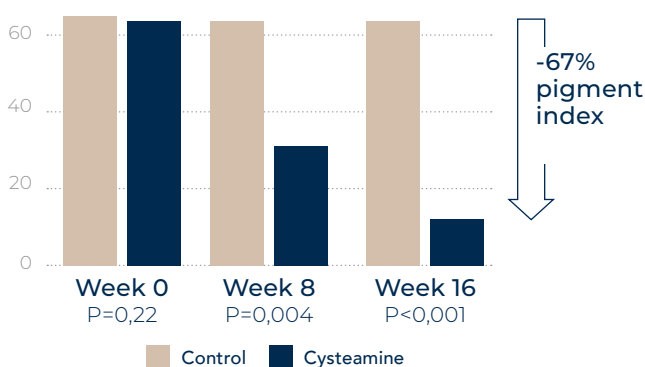
- ✓ The simplest aminothiols physiologically produced in human cells
- ✓ A high concentration of cysteamine is found in human milk
- ✓ Biocompatible and well tolerated

Cysteamine delivers powerful antioxidant activity in the skin¹

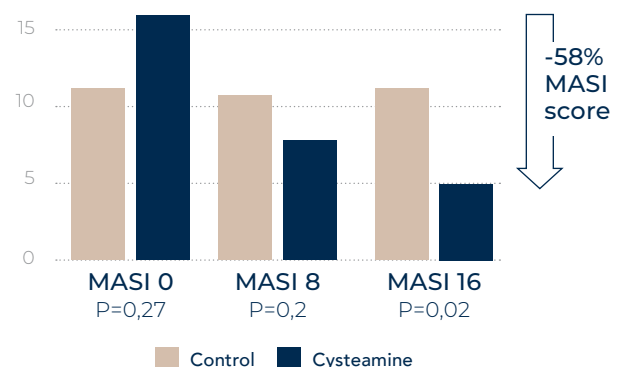
- ✓ Inhibition of tyrosinase
- ✓ Inhibition of peroxidase
- ✓ Increase of intra-cellular glutathione
- ✓ Reduction of dark melanin



67% pigment correction after 16 weeks measured by Mexameter²⁰



58% MASI score reduction after 16 weeks²⁰



Considerable efficacy

Hsu et al (2013) Journal of the American Academy of Dermatology 68 (4,1) AB189, P6024

Significant efficacy both by investigators' and patients' assessment

Mansouri et al (2015) British Journal of Dermatology 173 (1) 209-217

Higher pigment correction compared to all other agents

Kasraee (2017) 23rd International Pigment Cell Conference in Denver Co, USA

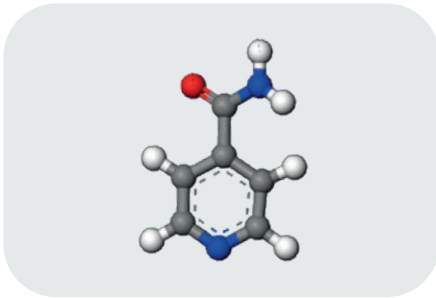
Superior benefit / risk ratio compared to all other agents

Goorochurn (2017) 26th Meeting of the EADV, Geneva, Switzerland

Significant efficacy in pigment correction

Farshi et al (2018) Journal of Dermatological Treatment, 29 (2) 182-189

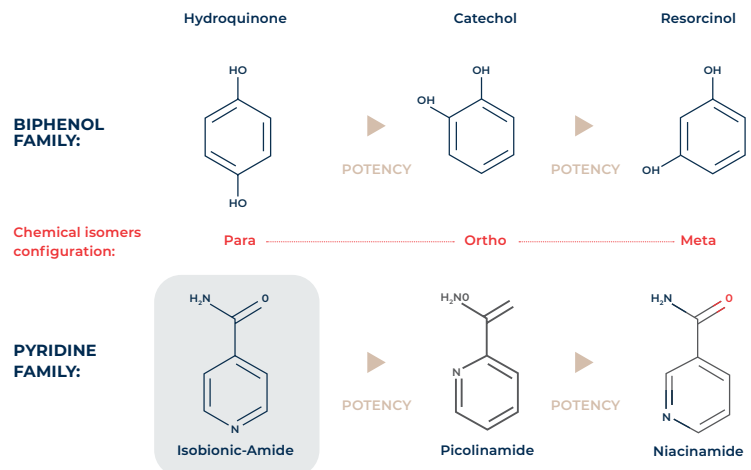
Isobionic-Amide a new lighthening agent technology



- Isobionic-Amide is a part of the vitamin B3 family, which are naturally occurring molecules.
- Isobionic-Amide has a pyridine structure in para-isomer configuration.
- Isobionic-Amide related INCI function is skin conditioning. It is as well use pharmacologically to increase the bioavailability of other molecules by formation of co-crystals

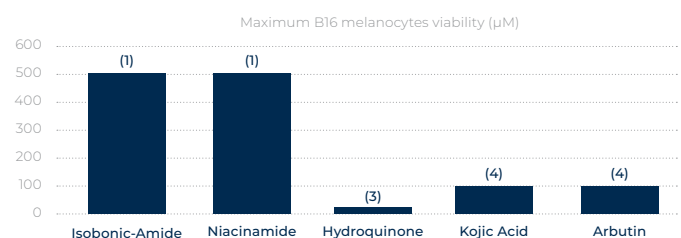
Increased potency

- Scientis discovered in 2011 the depigmenting effect of Isobionic-Amide
- The effectiveness of Isobionic Amide was first hypothesized by analysing the chemical isomer configuration of the pyridine family in comparison of the biphenol family
- Similarly to hydroquinone vs. resorcinol, the hypothesis was that a pyridine in para configuration would be more potent than in Meta configuration
- In-vitro evaluation confirmed the greater potency of Isobionic-Amide vs. niacinamide
- Scientis patented in 2013 the synergy between Cysteamine and Isobionic-Amide



Excellent safety profile

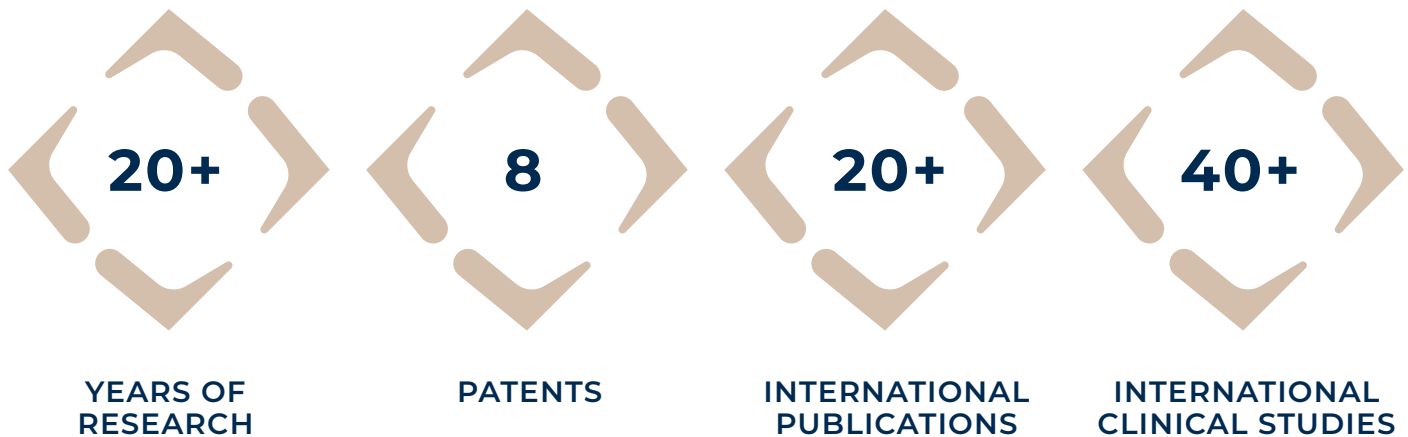
- Considered as hydrophilic with a partition coefficient: $\log P = -0.28^2$, which is **favorable to penetrate through corneal layer**.
- Considered low molecular weight at 122,125 g/mol, which is **favourable to diffuse in epidermis**.
- Considered readily **biodegradable**¹
- **Negligeable Acute Toxicity**
 - LD50 (oral, rat) is > 2.000 mg/kg
 - No death observed at oral dose of >875 mg/kg in rat³
- **Not considered teratogenic & mutagenic**
- **Non-Carcinogenic**
 - Not carcinogenic at 5g/kg bw/day (mice, 1% solutions in drinking water during their lifetime, equivalent to 5g/kg bw/day)⁶
- **Low Melanocytes & Keratinocytes cytotoxicity**
 - None up to very high concentration (500µM), as for Niacinamide.^{4,5}
 - Considerably less toxic to melanocytes compared to other depigmenting agents (25µM for hydroquinone and hydroquinone-acetate⁷; 100µM for kojic acid and arbutin⁸)



Scientis : The Pigmentation Authority

Scientis is a Swiss dermatology company dedicated to skin pigmentation.

We aspire at leading the transformation of the **Aesthetic Dermatology with the Skin of Colours in mind**. We work to offer highly differentiated molecules, technologies and products specifically addressing the unmet Medical and Aspirational needs of the multiple diversity of the People of Colour.



Recommended by physicians and patients

“I’ve have seen excellent efficacy. My patients really appreciate Cyspera®’s ability to not only help reduce the appearance of dark spots, but also blend their complexion to provide a more even skin tone.”
– Corey L. Hartman, MD

“We have seen promising results with Cyspera®, and I’m excited that there is a new, long-term solution to address patchy pigmentation and dark spots.”
– Seemal R. Desai, MD

“Was on hydroquinone on and off for several years and although I managed to lighten my dark patches, as soon as I stopped, the dark patches returned. Cyspera® is a game changer. I have used it for 4 weeks and I can tell that there is 30% improvement to my dark patches”
– Hilary (US Patient – phototype III)

“Best system for hyperpigmentation!! I’ve used everything made for hyperpigmentation for the last 2 years and this is the only product that has helped my dark pigment. I’ve used it for about 4 months total, and I would say my pigmentation is nearly gone”
– Mika V. (US Patient – phototype V)



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