The Properties of CTP (Copper Tripeptide-1)



OVERVIEW

Copper peptides are naturally occurring small protein fragments that have high affinity to copper ions. In human plasma, the level of GHK-Cu is about 200ng/ml at age 20. By the age of 60, the level drops to 80ng/ml.

Scientific studies conducted in different research laboratories have established in humans, tripeptide GHK-Cu can promote activation of wound healing, attraction of immune cells, antioxidant and anti-inflammatory effects, stimulation of collagen and glycosaminoglycan synthesis in skin fibroblasts and promotion of blood vessels growth. Recent studies indicate its role in stem cell biology and anti-tumor defense. Synthetic GHK-Cu is used in cosmetics as a reparative and anti-aging ingredient.

THE DEFINITION OF CTP

Copper peptide, GHK-Cu is a naturally occurring copper complex of a glycyl-Lhistidyl-L-lysine peptide. Since it has three amino acids it is called tripeptide. The GHK-Cu tripeptide has strong affinity for copper(II) and was first isolated from human plasma. It can be found also in saliva and urine.



COPPER COMPLEX COPPER TRIPEPTIDE-1

H N

O



0

0

 $\dot{N}H_2$

THE CHARTERISTICS OF CTP



INCI Name Other Name Cas No Purity (%)

Copper Tripeptide-1 GHK-Cu 89030-95-5 95% (min)



PROPOSED MECHANISM OF ACTION OF GHK-Cu¹



THE EFFECT OF COPPER TRIPEPTIDE-1

- 1. Calm irritated and reddened skin
- 2. Tighten loose skin and improve elasticity
- 3. Tighten the protective skin barrier proteins
- 4. Improve skin firmness
- 5. Reduce fine lines
- 6. Reduce the depth of deep wrinkles
- 7. Improve skin clarity and "glow"
- 8. Reduce spots, photodamage, and hyperpigmentation
- 9. Smooth rough skin
- 10. Improve overall appearance
- 11. Improve hair transplant success
- 12. Improve hair growth and thickness
- 13. Restore liver after toxic poisoning
- 14. Block stomach ulcer development
- 15. Heal intestinal ulcers
- 16. Accelerate wound healing
- 17. Act as powerful anti-inflammatories on skin
- 18. Cause the differentiation of stem cells needed for tissue repair

ACTIONS OF COPPER TRIPEPTIDE-1



SIMPLE OVERVIEW OF THE ACTION OF COPPER TRIPEPTIDE-1



COMPARISON THE FUNCTION OF CTP WITH RETINOID AND VITAMIN C

| Skin Effect | Copper peptide | Retinoid | Vitamin C |
|---|---|---|---|
| Reducing wrinkle | Yes | Yes | Minor improvements |
| Skin Color Tone | Very good tone | Often reddish, leathery appearance | Fair tone |
| New Collagen | Yes Significant in 70% of users in 30days | Yes Significant in 40% of users in 30days | Yes Significant in 50% of users in 30days |
| New Elastin | Yes | Yes | Minor improvement |
| Increase water– holding proteoglycans and glycosaminoglycans | Yes | Yes | No |
| Repair skin barrier | Yes | No Actually degrades skin barrier | Νο |

COMPARISON THE FUNCTION OF CTP WITH RETINOID AND VITAMIN C

| Skin Effect | Copper peptide | Retinoid | Vitamin C |
|--|---|--|--|
| Improve capillary circulation | Yes | Yes | No |
| Activate Metalloproteinases to remove damaged proteins | Yes | No | No |
| Causes irritation | No | Yes Strong irritant | No |
| Anti-inflammatory | Yes | No | No |
| Anti-oxidant Yes Activates superoxide No dismutase in skin | | No | Yes Vitamin C acts as anti-oxidant |
| Used for clinical skin repair? | Yes For wound healing and dermatological skin repair | Yes For skin renewal and anti–acne | No Effects are too minor |

EFFECT OF INJECTION FOR GHK-Cu INTO THE WOUND CHAMBER





3 Days after injection of GHK-Cu





14 Days after injection of GHK-Cu

EFFECT OF INJECTION FOR GHK-Cu INTO THE WOUND CHAMBER

At days 3

More inflammatory cells (polymorphonuclear cells and macrophages) were present in the treated chambers (B) than in the controls (A) and some neovascularization (v) was already visible.

At days 14

Large areas of well-organized fibrosis with elongated fibroblasts were seen in the GHK-Cu-injected chambers (D), whereas the control ones (C) were characterized by a cellular accumulation

EFFECT OF GHK-Cu (LSE)

Fig. 3 Histology of normal skin (a) and LSE (b LSE without copper–GHK, c LSE with copper–GHK). Following H&E staining, the shapes of basal cells were compared by microscopic examination. The basal layer keratinocytes appeared to be more cuboidal in the copper– GHK treated models (×200, *insert box* ×400)



Control





copper-GHK



Integrin α6

Integrin β1

EFFECT OF GHK-Cu (WESTERN BLOT ANALYSES)



Fig 5 Cultured keratinocytes were treated with or without 1 μ M copper-GHK for 3 days. The cells were then lysed and Western blot analyses were conducted on the lysate as described in the "Materials and methods". The results shown are representative of experiments that were conducted in triplicate



ANTI-AGING EFFECT OF GHK-Cu

Aged Epidermis:



Increased Skin Renewal & Repair:



GHK-Cu Effects on Skin Stem Cells

to basement membrane

Integrins and p63 increase Stemness revived stem cells

EFFECT OF GHK-Cu ON KERATINOCYTES



The long-term effect of Copper–GHK on the proliferation of Keratinocytes



LSE with or without GHK-Cu



IMPROVING SUNTANNING AND REDUCING UV DAMAGE TO SKIN CELLS

GHK blocks lethal ultraviolet radiation damage to cultured skin keratinocytes by bindind and inactivating reactive carbonyl species such as 4-hydroxylnoneal, acrolein, malondialdehyde and glyoxal.



GHK increasing the efficiency of melanin formation and reducing post-tanning peeling damage. Decreasing the time in sunlight and post-tanning peeling would reduce overall ultraviolet exposure.

COSMETICS ADVANTAGE OF CTP

- Skin matrix remodeling
- Penetrate stratum corneum membranes
- Anti-inflammatory and Antioxidant action
- Improves wound healing
- Increases hair follicle size
- Stimulation of hair growth
- Restores function in damaged cells





COSMETICS ADVANTAGE OF CTP





Skin Remodeling (Humans)

- Six placebo controlled studies on 270+ subjects
- Increases keratinocyte proliferation
 Improves appearance, firmness
- elasticity, and skin thickness
- Improves wrinkles, mottled hyperpigmentation & photodamage
- Increases skin collagen
- Tightens protective skin barrier proteins
- · Improves skin clarity



- Wound Healing
- Heals:
- Rats, mice, pigs, rabbits, humans, dogs and Guinea pigs
- Heals:
- · Surgical wounds
- · Burn wounds
- Ischemic wounds
- Wound chambers
- Punch biopsy
- · Dog paws
- Skin transplants
- Accelerates:
- Increased re-epithelialization, wound closure, wound strength, granulation tissue, collagen, elastin, proteoglycans, glycosaminoglycans, decorin, and subcutaneous fat cells



Tissue Remodeling Copper peptides Induce Regeneration of Diverse Organs



CTP IN MARKET

Skin renewal

Neutrogena Visibly Firm Night Cream® Neutrogena Visibly Firm Eye Cream® Blue Copper Firming Elasticity Repair® Climate Extreme Body Repair with Copper Peptide® Blue Razor Aftershave® Neova®Eye Therapy Neova®Ight Therapy Cream® Neova®Body Therapy Lotion® Neova®Cuticle Therapy Neova®Cleansing Bar Neova®Body Scruband BioPeptide-CL®

Tissue regeneration

lamin®group of wound products, BioHeal®for "At-Risk" skin in persons with conditions Graftcyte®products for hair transplantation, Folligen®Hair growth stimulating products Tricomin®Hair growth stimulating products, post-surgical skin healing Complex Cu3®products for use after I aser resurfacing Protect & Restore®skin care products, Neutrogena Visibly Firm Night Cream®skin care products, Neutrogena Visibly Firm Eye Cream®skin care products, Blue Copper®skin care products, NeovaNight Repair®skin care products, Protect & Restore SuntanningLotion®

Wound healing

IaminGel Wound Dressing® IaminImpregnated Gauze Dressing® Iamin Wet Dressing (copper-saline)® Iamin-2 Hydrating Gel® IaminWound Cleanser®

Stimulation of Hair Growth

- •Tricomin®Solution Follicle Therapy Spray,
- •Tricomin®Revitalizing Shampoo,
- Tricomin®Restructuring Conditioner
- Tricomin®Conditioning Shampoo

Post-Surgical Skin Healingafter laser resurfacing, dermabrasion, and chemical peels

•Complex Cu3®Intensive Tissue Repair Cream

•Complex Cu3®Hydrating Gel

Complex Cu3®Gentle Face Cleanser



CTP IN KOREAN MARKET

| 제품 분류 | 제조사 | 제품명 |
|-------|---------|----------|
| 샴푸 | 현대약품 | 미녹솔C |
| 샴푸 | DK코스메틱 | 코미인 |
| 샴푸 | 한국미용메디팜 | 테라피션 |
| 샴푸 | - | 고려황칠 |
| 샴푸 | - | 디아스포라 |
| 샴푸 | - | 프로메톡 |
| 샴푸 | - | 조아산파낙스 |
| 크림 | 프로유코스메틱 | 쿠퍼펩타이드크림 |
| 탈모치료제 | - | 온헤어 |
| 수용액 | DK코스메틱 | 코미인 |







THE FUNCTION OF CTP (SUMMARY)

Causes skin remodeling

- Rebuilds skin protective barrier
- Increases collagen and elastin more than vitamin C
- Increases water-holding proteoglycans
- Rebuilds blood microcirculation
- Activates removal of damaged skin proteins and scars

Present in wounds

- The human body's natural remodeling signal

Anti-inflammatory

- Potent anti-inflammatory, activates skin's main antioxidant protein--dismutase superoxide

THE FUNCTION OF CTP (SUMMARY)

Increases hair follicle(humans) size

- Also blocks hair loss if given before chemotherapy and accelerates hair growth after chemotherapy (rats)

Hair Growth

- Copper peptide (GHK-Cu) stimulate hair growth

- The efficiency of synthetic analog of GHK-Cu was similar to that of 5% minoxidil.



CTP

| INCI Name | Copper Tripeptide-1 |
|------------|---------------------|
| Other Name | GHK-Cu |
| Cas No | 89030-95-5 |
| Purity (%) | 95% (min) |

CTP (1,000 ppm)

| INCI Name | Water, Butylene Glycol, Copper Tripeptide– 1, Phenoxyethanol, Methylparaben, Ethylparaben, Butylparaben, Propylparaben, isobutylparaben |
|--------------------------------------|--|
| Appearance | Transparent Blue Solution |
| Density at 20℃ | 0.990 - 1.010 |
| Copper Tripeptide–1 Content (ppm) | 950 ~ 1050 |
| Recommended Dossage | 5% (min) |

FORMULATIONS

| Face and Eye Care – Treatm | nent |
|----------------------------|-------|
| Product/INCI Name | % w/w |
| <part a=""></part> | |
| Stearic acid | 1.00 |
| Cetanol | 1.50 |
| BP | 0.10 |
| Bees wax | 1.0 |
| Aracel 60 | 1.00 |
| Tween 60 | 1.00 |
| Aracel 165 | 1.50 |
| Squalane | 9.00 |
| LP #70 | 4.00 |
| Silicone oil(200F) | 2.00 |
| <part b=""></part> | |
| MP | 0.20 |
| EDTA-2Na | 0.02 |
| Allantoin | 0.05 |
| TEA | 0.50 |
| 1,3-BG | 10.00 |
| Glycerin | 2.00 |
| Carbopol 940 | 0.24 |
| B-Glucan | 1.00 |
| D.I water | 61.43 |
| <part c=""></part> | |
| CTP-1 | 0.10 |
| Perfume | 0.02 |

Methods

- 1. Heat Phases A to 75-80 ℃
- 2. Combine ingredients of Part B with mixing $% 10^{-1}$ and heat to 75-80 $^{\circ}\mathrm{C}$
- 3. Add part A to Part B with mixing and cool to 45 $^\circ\!\!C$

4. Add Part C of CTP-1 and perfume at 40 to 45 °C and stir until temp.



FORMULATIONS

Conditioning Shampoo

| Product/INCI Name | % w/v |
|--|----------|
| <part a=""></part> | |
| Deionised Water | To 100.0 |
| Sodium Laureth Sulfate, 3 EO(70%) | 14.50 |
| Cocamidopropyl Betain | 9.00 |
| Sodium Lauroyl Sarcosinate | 5.00 |
| Isostearamidopropyl Morpholine Lactate | 5.00 |
| Stearamidopropyl Dimethylamine Lactate | 5.00 |
| | |

<Part B> Polyquaternimu-10 **Deionised Water**

| <part c=""></part> |
|------------------------------------|
| CTP-1 |
| PEG-6 Carpric/Carprylic Glycerides |
| PHENOVA |
| Perfume |
| Total |

w/w 00.00

4.50

0.20

2.50

0.05 1.50 0.50 qs 100.00

Methods

- 1. Dissolve SLES in water
- 2. Add the rest of Part A ingredients and mix until homogeneous.
- 3. Combine part B ingredients and add into Part A heat to 60 °C and mix for 10min.
- 4. Cool to below 40 ℃
- 5. Add Part C ingredients and mixing.
- 6. Adjust pH to 7.0 if necessary and viscosity as desired.



FORMULATIONS

Anti-hair loss Hair Tonic

| Product/INCI Name | % w/w | |
|--------------------------------|-----------|----------------------|
| Deionised Water | To 100.00 | |
| Citric acid | 0.26 | |
| Trisodic acid | 1.20 | |
| Potassium Sorbate | 0.10 | Method |
| <part b=""> PHENOVA</part> | 0.50 | 1. Add r 2. Add I |
| PPG-5 Ceteth 20 | 2.00 | 5. Mix P Helix |
| <part c=""></part> | | |
| CTP-1 | 0.20 | |
| <part d=""></part> | | |
| Polysorbate 20 | 1.00 | |
| Perfume | qs | |
| Total | 100.00 | |



- ds
- part B to Part A with mix.
- Part C to Part A+B and mix
- Part D and add it to Part A+B+C with stirring.



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THANK YOU

