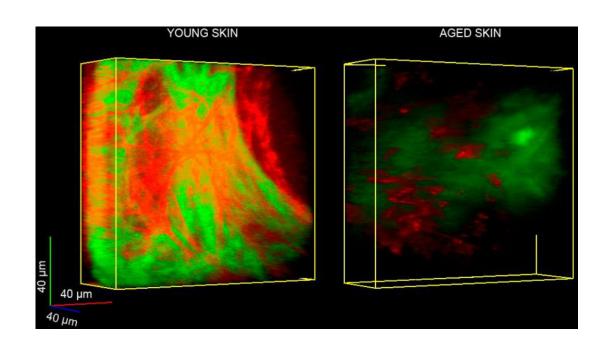
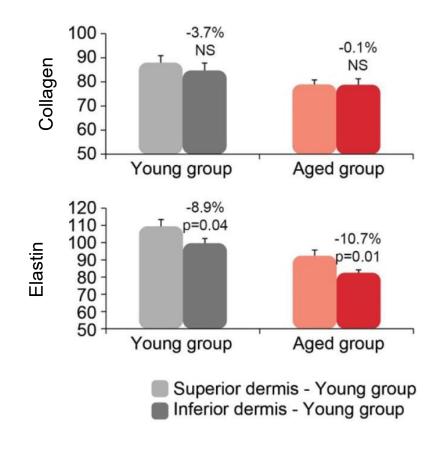


## Collagen and elastin reduction in aging skin

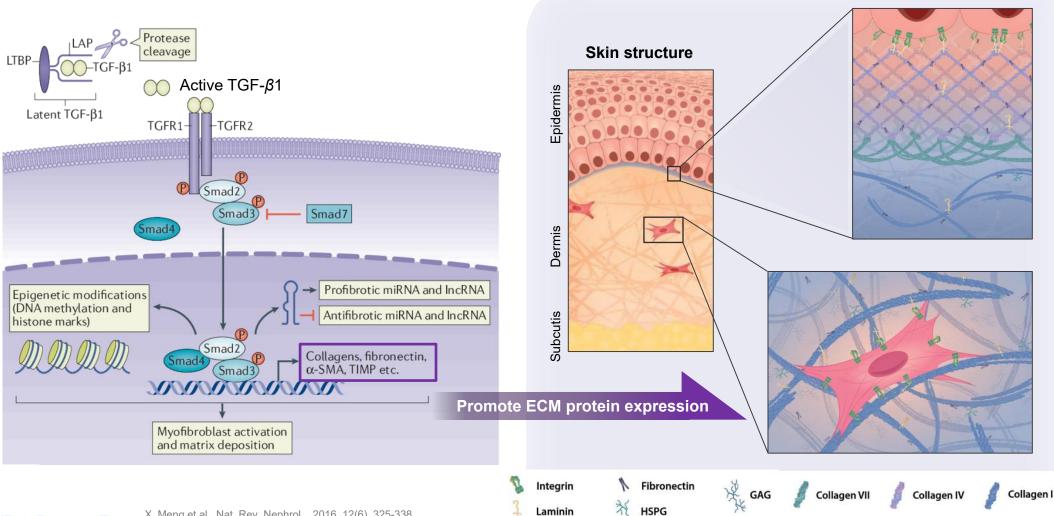


Collagen: SHG-signal in green; Elastin: autofluorescence in red



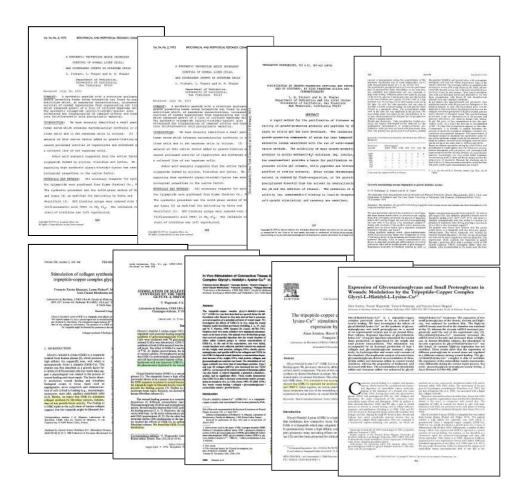


## The role of TGF- $\beta$ in ECM protein synthesis





### Key milestones in tripeptide-1 (GHK) discovery and research



#### ◆ 1973 Discovery:

L. Pickart identified a tripeptide (Gly-His-Lys) in human serum that promotes mouse liver cell proliferation.

#### ♦ 1973 Synthesis:

The tripeptide Gly-His-Lys was synthesized and shown to stimulate liver cell growth in mice.

#### 1975 Purification:

A method for purifying this growth factor using silica gel chromatography was developed.

#### 1977 Confirmation:

D. H. Schlesinger confirmed the growth factor in human serum is **Gly-His-Lys**.



### **Natural source of tripeptide-1: Promotes** collagen Enzymatic fragment of SPARC protein synthesis via TGF-β **Antioxidant** effect $-NH_2$ HM NH **Accelerates** wound healing $H_2N$ Increases **SPARC** protein **GHK**



skin density and thickness

### Palmitoyl modification enhances transdermal delivery

#### **Methods of promoting osmolality**

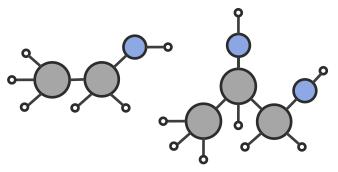
# Chemical modification (Palmitoyl, acetyl)



#### Complexation



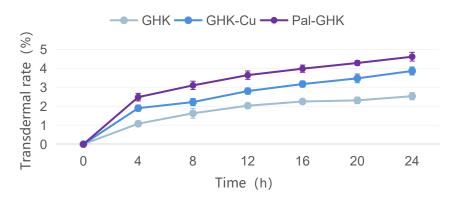
#### **Chemical osmolality promoter**



# READLINE

#### **Variation in transdermal delivery**

## Comparison of transdermal properties of modified GHK



- Concentration: GHK, GHK-Cu and Pal-GHK are at 10 mg/mL.
- 4-hour transmittance: GHK-Cu shows 1.76x, and Pal-GHK shows
  2.3x the transmittance of GHK.
- ◆ 24-hour transmittance: GHK-Cu reaches 1.53x, and **Pal-GHK** reaches 1.82x the transmittance of GHK.
- Transdermal rate: Pal-GHK > GHK-Cu > GHK



### WrinFix-P3P1:

A potent collagen-stimulating peptide

INCI Name: Palmitoyl tripeptide-1

■ CAS No.: 147732-56-7

■ Molecular Formula:  $C_{30}H_{54}N_6O_5$ 

■ Molecular Weight: 578.8 g mol<sup>-1</sup>

■ Efficacy: Anti-aging, anti-wrinkle

■ **Dosage:** 10-1000 ppm





## Versatile market application



#### **CLARINS**

Precious La Lotion Age-Defying Treatment Essence

Anti-wrinkle, firming, repairing

\$160/150 mL



#### **SKIN CEUTICALS**

A.G.E. Eye Complex for Dark Circles

Anti-aging, remove dark circles

\$132/15 g



#### **Drunk Elephant**

Protini™ Polypeptide Cream

Anti-aging, repairing

\$69/50 mL



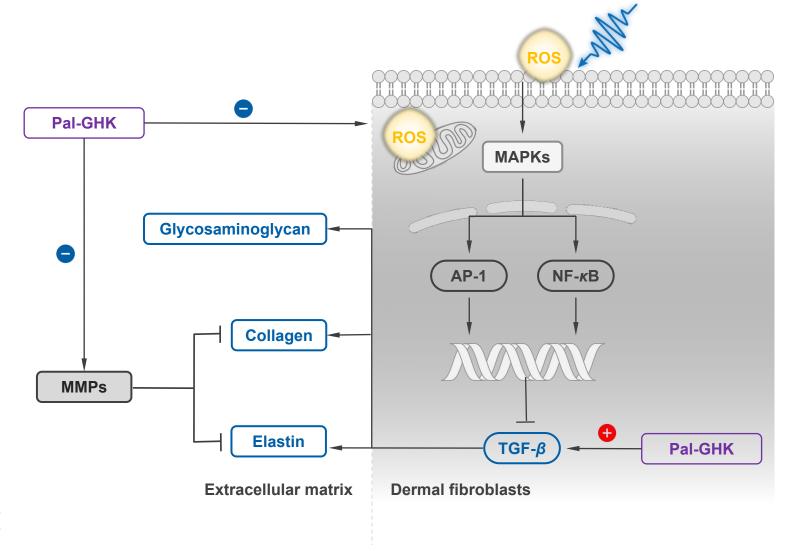
Product

Efficacy

Price

### WrinFix-P3P1:

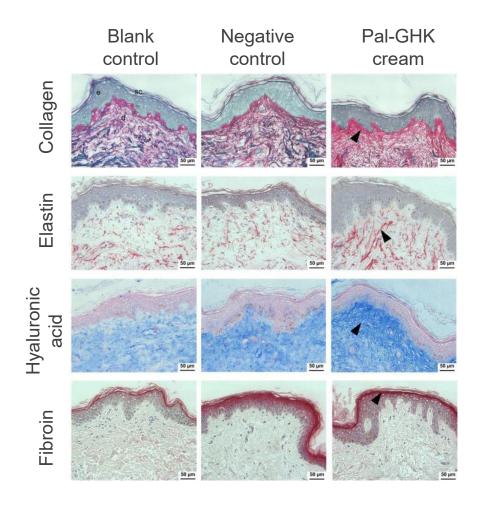
Activates TGF- $\beta$  and promotes collagen production



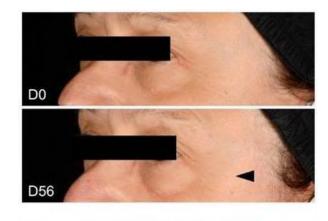


### WrinFix-P3P1:

### Promotes collagen, elastin and reduces wrinkles



Comparison before and after using 100 ppm palmitoyl tripeptide-1 cream

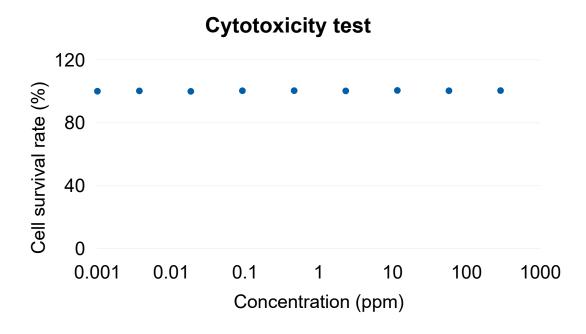








## WrinFix-P3P1: No cytotoxicity within 1000 ppm



➤ Cytotoxicity was not observed in the concentration range of 1 – 1000 ppm WrinFix-P3P1.



## WrinFix-P3P1: Safety for use

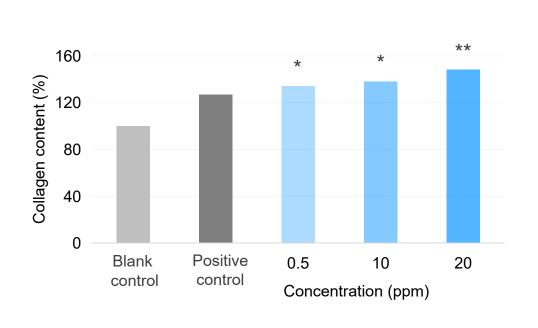
**Patch Test** 

Group	Number of subjects	observe time	Patch test number of different skin reaction levels				
			0	1	2	3	4
Readline WrinFix-P3P1	30	0.5 h	30	0	0	0	0
		24 h	30	0	0	0	0
		48 h	30	0	0	0	0
Negative control	I 30	0.5 h	30	0	0	0	0
		24 h	30	0	0	0	0
		48 h	30	0	0	0	0

**Result**: Human skin patch test showed **200 ppm WrinFix-P3P1**, and 0 of 30 subjects had skin adverse reactions.



## WrinFix-P3P1 can promote the collagen sysnthesis



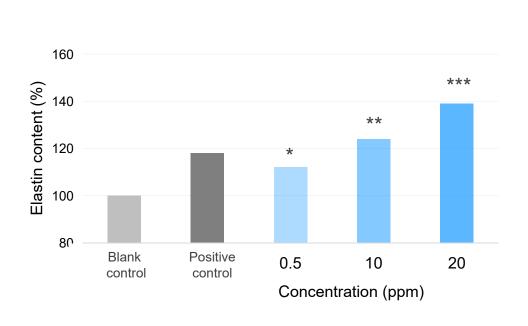
In vitro	Human fibroblasts
Blank control	Cell culture medium
Negative control	UVB(200 mJ/cm²)
Positive control	100 ng/mL TGF <i>β</i> -1
Test group	WrinFix-P3P1
Test method	Detect absorbance at 450 nm
	Readline Laboratory



20 ppm WrinFix-P3P1 increases collagen synthesis by 48.2%.



## WrinFix-P3P1 can promote the elastin synthesis



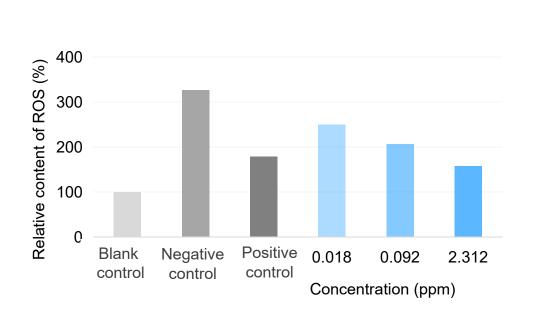
In vitro	Human fibroblasts
Blank control	Cell culture medium
Negative control	UVB (200 mJ/cm²)
Positive control	100 ng/mL TGF <i>β</i> -1
Test group	WrinFix-P3P1
Test method	Detect absorbance at 450 nm
	Readline Laboratory



20 ppm WrinFix-P3P1 increases elastin synthesis by 39.1%.



### WrinFix-P3P1 can inhibit the generation of intracellular ROS



In vitro	Human fibroblasts	
Blank control	Cell culture medium	
Negative control	UV	
Positive control	UV + Vitamin E	
Test group	WrinFix-P3P1	
Test method	Detect absorbance at 450 nm	
	Readline Laboratory	



WrinFix-P3P1 can significantly reduce intracellular ROS.



## WrinFix-P3P1: COSMOS approved







## WrinFix-P3P1: Application information





### **Application Guide**

• **Purity:** ≥ 98.0%

Solubility: Partly soluble in water (< 1000 ppm)</li>

• Condition: Protect from light, seal tightly, and store at 2-8 ° C

• Suggested dosage: 10-1000 ppm

Incompatibility: Avoid with metal complexes



⊠ SPEC	⊠ COA	⊠ MSDS
⊠ TDS	⊠Material safety information	⊠ Leaflet
⊠ Product PPT		
⊠ HPLC spectrum	☐ Stability study	☐ Structure confirmation report
☐ Literature	⊠ In vitro study	☐ Human efficacy report
☐ Permeability report	⊠ Flow chart	□ Declaration of ingredients
□ COSMOS certificate	⊠ HALAL	☐ Formulation guide
	☐ Natural origin index	<ul><li>☑ Declaration of origin</li></ul>
⊠ Patent	⊠ Non-GMO	☐ Others





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