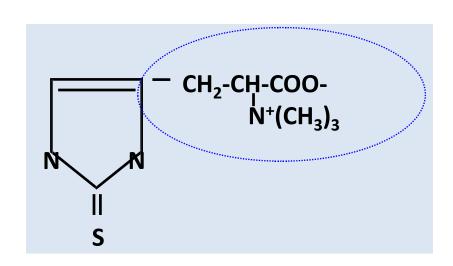




# THIOTAINE®



#### The Desirable Anti-oxidant

- Neutralizes Ozone
- Colorless Odorless

#### **CONCEPT**

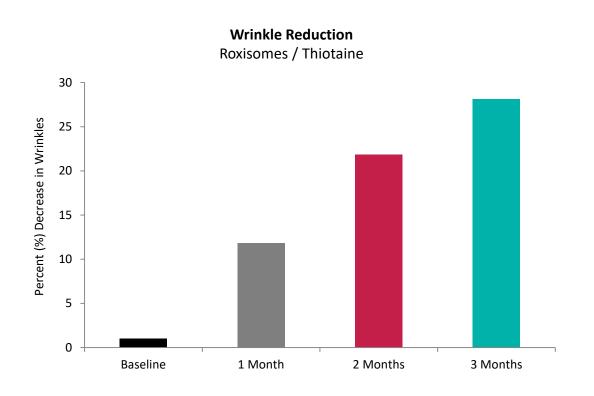
Cells need energy in the form of ATP. To create ATP mitochondria need fatty acids. With energy production come free radicals.

A car driving full speed without cooling fluid does not go far. Similarly, sparkles (in the form of free radicals) damage the mitochondrial membrane during ATP production. Cells need energy while quenching free radicals.

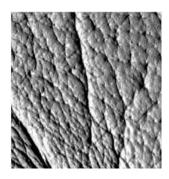
The solution is Thiotaine (Ergothioneine), a molecule with two functional groups: (1) to transfer fatty acids in the mitochondria, and (2) to act as a strong anti-oxidant with two extra characteristics.

- Thiotaine is odorless and colorless
- Thiotaine is desirable by the cells which have specific gates for this molecule.

## Effect of Roxisomes and Thiotaine on reducing skin wrinkles



Baseline



2 Months Treatment



#### Protocol:

- 21 subjects (female), Age 30-65
  - Subjects contained visible wrinkles in their outer canthus area (crows feet around the eyes)
- Formula containing 1% Roxisomes and 1% Thiotaine
- Formula used on full face, twice a day
- 3 months
- Images of the outer canthus area captured using imaging system.
- Photographs analyzed for fine lines and wrinkles using image analysis software

## THIOTAINE (ERGOTHIONEINE)

#### Two functional groups

HS—C CH Transfert group of fatty acids (similar to Carnitine)
$$C - CH_2 - CH - COO^{-1}$$

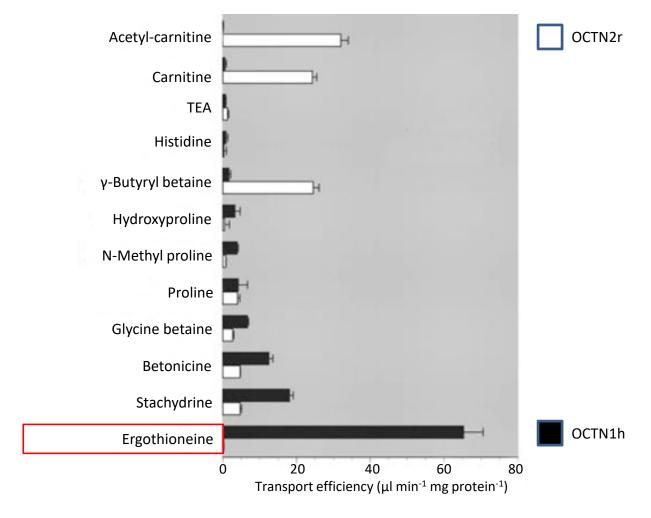
$$+N(CH_3)_3$$

Ergothioneine is an amino acid which we eat from dietary botanicals such as grains or vegetables. It will concentrate in the eye, and in red blood cells to protect hemoglobin (the carrier of oxygen). Ergothioneine is unmetabolized and slowly excreted.

Resistant – Colorless - Odorless

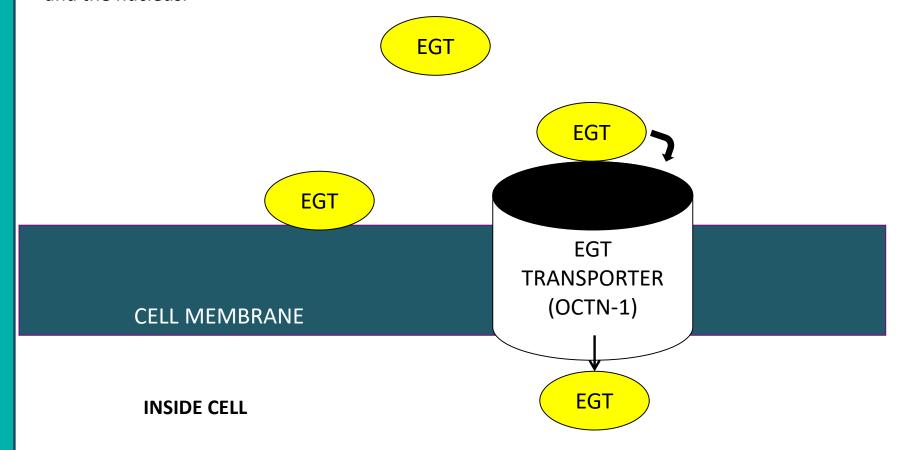
#### ERGOTHIONEINE IS DESIRABLE TO HUMAN SKIN CELLS

There are transporters in the cell membranes called OCTN. Different types of OCTN transport different molecules. OCTN1 works for Ergothioneine.



## OCTN1: THIOTAINE TRANSPORTER

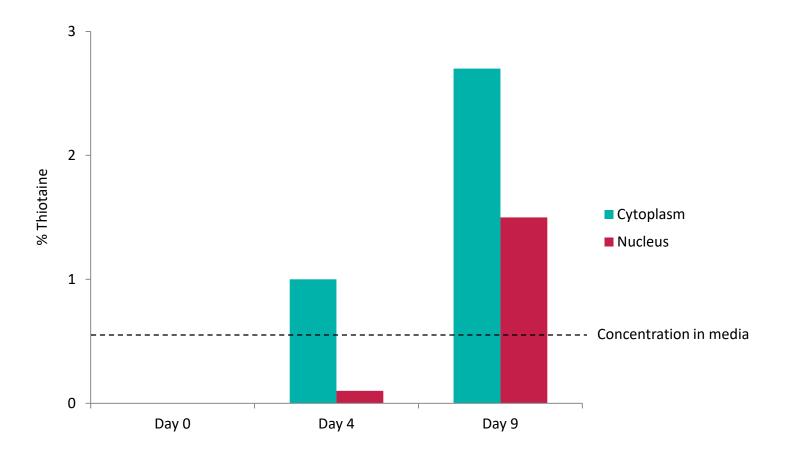
OCTN1 is in the membranes of fibroblasts, keratinocytes, melanocytes and in the mitochondria and the nucleus.



OCTN1 is the newly discovered transporter of EGT.

#### **ACCUMULATION IN CELLS**

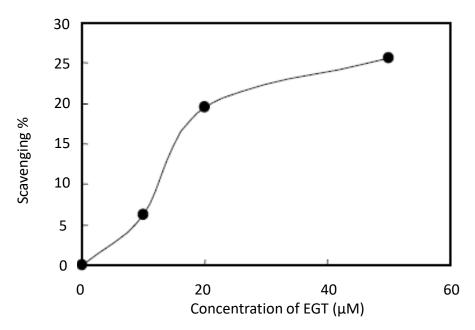
Normal human fibroblasts supplemented with 0.5% Thiotaine. Cells collected, separated into nucleus and cytoplasm and assayed for concentration.



The chart shows the assimilation of Thiotaine by the cells. EGT can also accumulate in keratinocytes and melanocytes using OCTN1.

#### THIOTAINE = SUPER ANTI-OXIDANT

## Superoxide Anion From hypoxanthine-xanthine oxidase

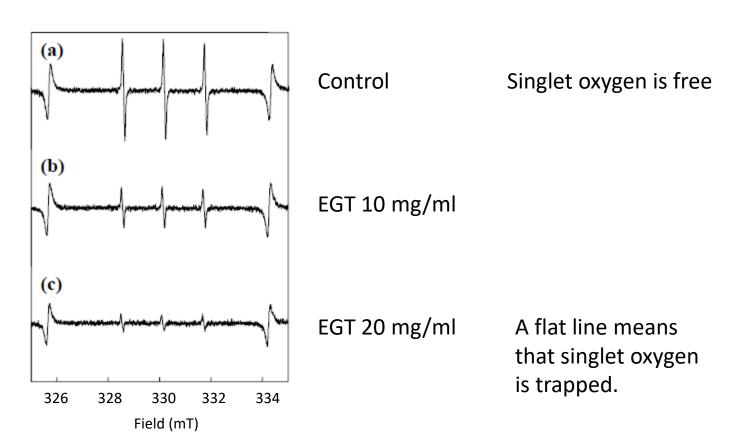


Obayashi et al., J. Cosmet. Sci. 56:17, 2005

With increasing concentration, EGT is increasingly efficient at catching the superoxide anion.

#### THIOTAINE = SUPER ANTI-OXIDANT

## Singlet Oxygen<br/>Spin Trapping



Obayashi et al., J. Cosmet. Sci. 56:17, 2005

#### THIOTAINE = SUPER ANTI-OXIDANT

## Stronger than:

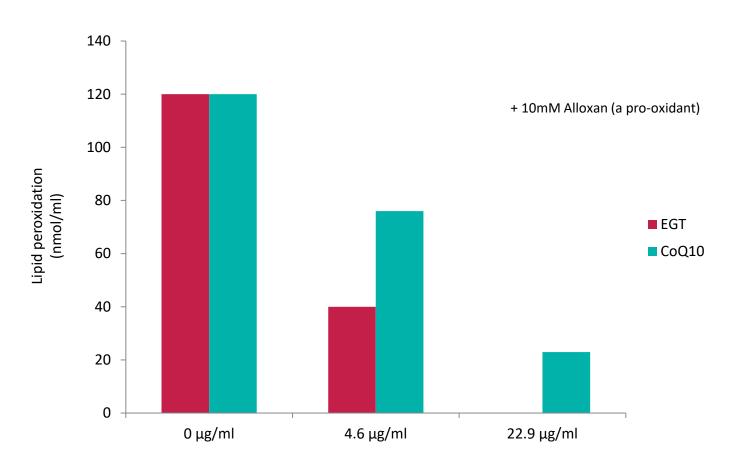
Coffeeberry – ORAC score 4X higher

(61,000 v 15,000) Brunswick Laboratories

- ➤ Idebenone and CoQ10 Dong et al., J. Cosmet. Dermatol. 6:183, 2007
- Vitamin C
  He et al., Skin Pharma Physiol. 18:183, 2004; Obayashi et al., J. Cosmet. Sci 56:17, 2005
- > Trolox (Vitamin E) Franzoni et al., Biomed. Pharmacother. 60:453, 2006
- Figure 1., Biochem. Biophys. Res. Comm. 231:389, 1997

## THIOTAINE: BETTER THAN CoQ10

#### Peroxidation of empty liposomes with Alloxan

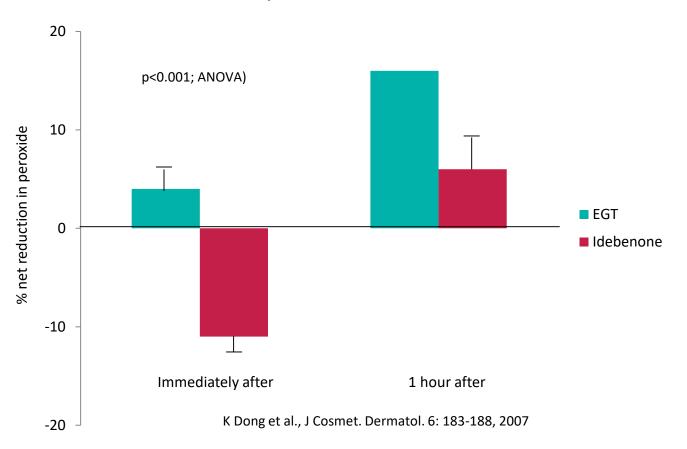


EGT at 22.9  $\mu$ g/ml stops peroxidation. CoQ10 does not.

#### THIOTAINE: BETTER THAN IDEBENONE

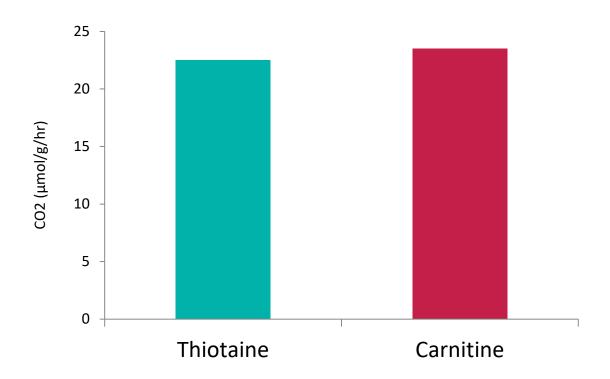
Oxidation of Rhodamine 123 in fibroblasts irradiated with Solar simulating UV (100 Kj/CM<sup>2</sup>).

10 µm of EGT or Idebenone



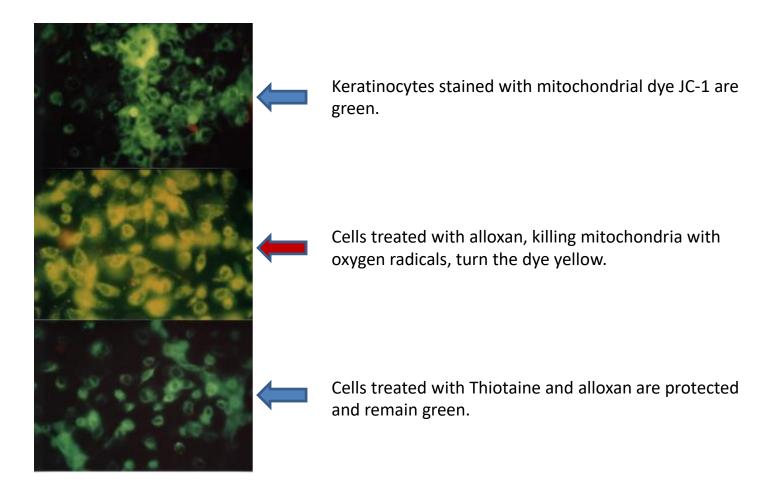
The chart shows the superiority of EGT over Idebenone for UV protection.

## SKIN ENERGY SUPPLEMENT: "CARNITINE-LIKE"



Both Carnitine and Thiotaine can supply fatty acids into mitochondria. The fatty acids are burned by oxygen to release CO<sub>2</sub> and ATP. This graph shows that they are equivalent. However, only Thiotaine controls the free radicals and protects the mitochondria.

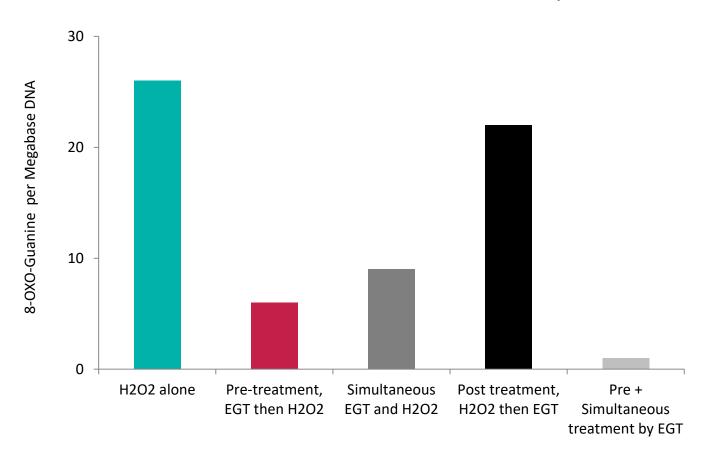
## THIOTAINE PROTECTS THE MITOCHONDRIA



Thiotaine, an energizing molecule, protects the membrane integrity of mitochondria against free radicals induced by alloxan.

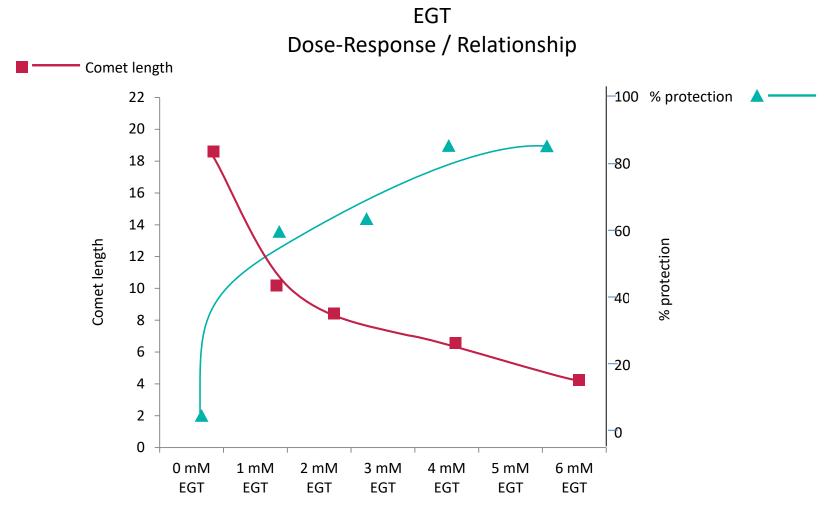
#### THIOTAINE PROTECTS AGAINST OXIDATIVE DNA DAMAGE

#### Thiotaine treatment of human keratinocytes



8oG is a damage on DNA, increased by pollution.

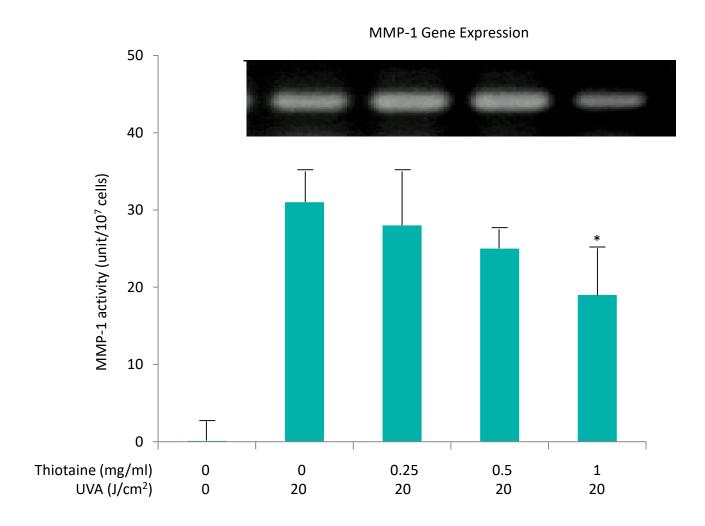
## THIOTAINE PROTECTS DNA AGAINST UV-A



Graph inspired by Decome et al., J. Photochem. Photobiol. 79:101, 2005

The more EGT is used, the more the DNA is protected.

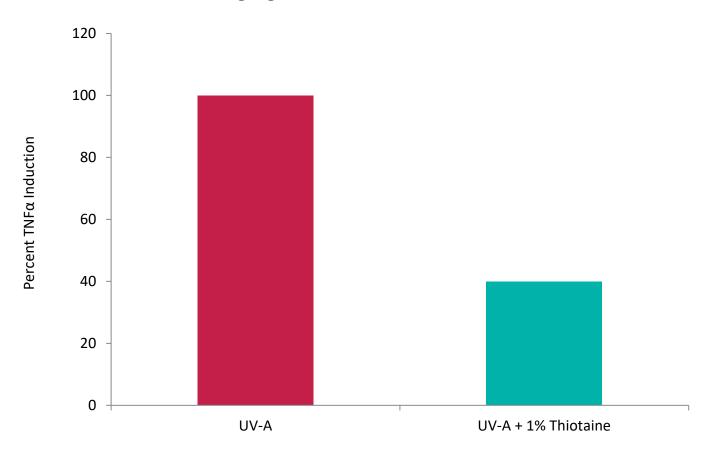
## THIOTAINE REDUCES MMP-1 EXPRESSION INDUCED BY UV-A



Thiotaine reduces MMP-1 expression in a dose dependent manner.

## THIOTAINE REDUCES STRESS SIGNALS

Photoaging: UV  $\rightarrow$  TNF $\alpha \rightarrow$  MMP-1  $\rightarrow$  Wrinkles



Obayashi et al., J. Cosmet. Sci. 56:17, 2005

Thiotaine reduces the release of the stress signal TNF- $\alpha$ .

#### **CONCLUSION**

#### Thiotaine:

- > Is colorless and odorless
- ➤ Is a super anti-oxidant
- Protects mitochondria
- > Shields against ozone
- Protects against oxidative stress and pollution
- Protects DNA against UV-A
- ➤ Reduces MMP-1
- ➤ Reduces stress signals



# THIOTAINE®

**INCI Name:** Water (and) Ergothioneine

**Preservative:** Phenoxyethanol

Suggested Use Level: 1.00%

Formulation Guidelines: Thiotaine is a colorless, clear liquid with a characteristic odor. Thiotaine is water

soluble and should be added to the cooling phase. The recommended pH range

is 4.0-7.0.

## Global Compliance and Product Features:

COUNTRY
COMPLIANCE

AUSTRALIA
Exempt

CANADA
Listed rICL

CHINA
Listed IECIC

EU
<1 MT Exempt

For additional information please contact technical@barnetproducts.com.

