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# **ProductInformation**

CERAMIDES Non-Hydroxy Fatty Acid Sigma Prod. No. C2137

SYNONYMS: N-acyl-sphingosine

#### PHYSICAL DESCRIPTION:

Appearance: White powder

Molecular weight: Approximately 607 assuming sphingosine (trans-D-erythro-2-amino-4-octadecene-1,3-diol) is the base with N-acyl fatty acids of predominantly stearic (octadecanoic) and nervonic (cis-15-tetracosenoic) acids. Molecular formula:  $C_{39}H_{76}NO_3$  assuming sphingosine with predominantly stearic and nervonic acids.



R = Mixture of fatty acids, primarily stearic acid ( $C_{18}$ :0) and nervonic acid ( $C_{24}$ :1,cis-15)

# METHOD OF PREPARATION:

Prepared by the action of phospholipase C on pure bovine brain sphingomyelin.<sup>1</sup> Purification to approx. 99% as measured by thin-layer chromatography is achieved using crystallization and partition techniques.

#### STABILITY / STORAGE AS SUPPLIED:

Store at -20° C

# SOLUBILITY / SOLUTION STABILITY:

Soluble in chloroform at 10 mg per ml. Stable in chloroform for up to 3 months stored in freezer (-OEC).

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# **USAGE / APPLICATIONS:**

Ceramide has a role as a signal transduction effector molecule for bioactive regulation of metabolic systems<sup>2-12</sup> and is available as an analytical standard for chromatography.

Ceramide is an endogenous lipid component of a novel biochemical pathway termed the sphingomyelin cycle.<sup>23,4</sup> The sphingomyelin cycle was discovered in human leukemia HL-60 cells<sup>2,3</sup> which are activated during differentiation induced with 1-alpha-25-dihydroxyvitamin D3. Ceramide is generated by hydrolysis of membrane sphingomyelin by a novel magnesium-independent, neutral, cytosolic sphingomyelinase.<sup>5</sup> The use of cell permeable synthetic ceramide (C-2 ceramide, N-acetyl-D-sphingosine), has been shown to produce a similar dose dependant induction of differentiation of HL-60 cells.<sup>5</sup> Other synthetic analogs such as C-6 ceramide (N-hexanoyl-D-sphingosine) have been found useful to study cell responses.<sup>6</sup> Ceramide is generated in response to cellular stimulation by hormones, cytokines and antigens.<sup>6,7,8,9,10,11</sup> Mechanisms for Ceramide action involve regulation of protein phosphorylation via stimulation of a serine/threonine protein phosphatase, a proline-directed kinase and possibly other direct and/or indirect targets.<sup>8</sup> Ceramide metabolites such as sphingosine and sphingosine-1-phosphate have potent biological activities of their own.<sup>7</sup> Other analogs such as D-erythro-dihydroceramide do not exhibit bioactive effects in certain biological systems.<sup>4,6,10</sup>

Ceramide appears to have a role in mediating biological responses in a wide variety of cell types.<sup>8</sup> Ceramide is emerging as an intracellular messenger that mediates effects on terminal differentiation and cell proliferation as well as apoptosis or cell death and cell-cycle arrest.<sup>10,12</sup> The interrelationships of ceramide actions with other bioactive lipids and systems represents an ongoing active research area.

#### **GENERAL NOTES:**

Ceramide has been available for many years as a representative member of this (non-hydroxy fatty acid) class of lipids. In recent years it has been defined as an important regulator of metabolic systems. A significant part of current research has centered around its involvement in apoptosis systems.

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#### GENERAL NOTES: (continued)

Several of the following references (3,4,6,7,8,9,10, and 12) are review articles and cite the original research articles. They offer additional information on the role of ceramide in cellular events.

#### **REFERENCES:**

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