

# **Cell Culture Tested Water-Soluble Complexes**

# **CHLORAMPHENICOL - WATER SOLUBLE**

#### Product Number: C3175

With approx. 100 mg chloramphenicol per gram; balance 2 - hydroxypropyl -  $\beta$ - cyclodextrin. Sold on the basis of mg of chloramphenicol.

#### **CHOLESTEROL - WATER SOLUBLE**

#### Product Number: C4951

With approx. 40 mg of cholesterol per gram; balance methyl -  $\beta$  - cyclodextrin.

Sold on the basis of mg of cholesterol.

# **DEXAMETHASONE - WATER SOLUBLE**

#### Product Number: D2915

(Cyclodextrin encapsulated dexamethasone)

With approx. 65 mg dexamethasone per gram; balance 2 - hydroxypropyl -  $\beta$  - cyclodextrin.

Sold on the basis of mg of dexamethasone.

# **ERGOCALCIFEROL - WATER SOLUBLE**

#### Product Number: E8014

#### (Vitamin D2)

With approx. 7 mg ergocalciferol per gram; balance methyl -  $\beta$  - cyclodextrin. Sold on the basis of mg of ergocalciferol.

# $\boldsymbol{\beta}$ - ESTRADIOL - WATER SOLUBLE

#### Product Number: E4389

(Cyclodextrin - encapsulated  $17\beta$  - estradiol)

With approx. 45 mg estradiol per gram; balance 2 - hydroxypropyl -  $\beta$  - cyclodextrin.

Sold on the basis of mg of estradiol.

# HYDROCORTISONE - WATER SOLUBLE

#### Product Number: H0396

(Cyclodextrin - encapsulated hydrocortisone)

# **ProductInformation**

With approx. 100 mg hydrocortisone per gram; balance 2 - hydroxypropyl -  $\beta$  - cyclodextrin.

Sold on the basis of mg of hydrocortisone

# LINOLEIC ACID - WATER SOLUBLE

#### Product Number: L5900

With approx. 30 mg of linoleic acid per gram; balance methyl -  $\beta$  - cyclodextrin.

#### **OLEIC ACID - WATER SOLUBLE**

#### Product Number: 01257

With approx. 30 mg oleic acid per gram; balance methyl -  $\beta$  - cyclodextrin

#### **PROGESTERONE - WATER SOLUBLE**

#### Product Number: P7556

(Cyclodextrin - encapsulated progesterone)

With approx. 70 mg progesterone per gram; balance 2 - hydroxypropyl -  $\beta$  - cyclodextrin.

Sold on the basis of mg of progesterone.

#### **RETINOL ACETATE - WATER SOLUBLE**

#### Product Number: R0635

(Vitamin A)

With a minimum of 5 mg retinol acetate per gram; balance methyl -  $\beta$  - cyclodextrin. Sold on the basis of mg of retinol acetate.

# **TESTOSTERONE - WATER SOLUBLE**

#### Product Number: T5035

(Cyclodextrin - encapsulated testosterone)

With approx. 109 mg testosterone per gram; balance 2 - hydroxypropyl -  $\beta$  - cyclodextrin.

Sold on the basis of mg of testosterone.

DEA CLASS III

# **PRODUCT SOLUBILITY**

Water-soluble complexes are soluble in water; difficulty may be encountered when attempting to solubilize in salt solutions or buffers. Typical stock solutions can be prepared at concentrations of 50 - 500 mg solid (powder) in 1 ml water. Solutions can be sterile filtered using a 0.2 µm membrane; the powders are not sterile. Solutions should be stored in working aliquots at -20°C. When calculating molar (e.g., µM, mM, etc.) solutions, the molecular weight of the cyclodextrin component of the complex is typically disregarded and the solution is prepared considering the amount of the hormone in the preparation and its respective molecular weight (printed in the alphabetical listings of the Sigma catalog).

# CYCLODEXTRINS - WHAT ARE THEY?

Many metabolically important compounds, such as, fatsoluble vitamins and hormones have very low solubilities in aqueous solutions. Various approaches have been taken to utilize these compounds in tissue and cell culture applications. Two most frequently used approaches are: (1) predissolving the compounds in organic solvents and (2) using "carrier" molecules to facilitate the dissolution of these compounds. One such class of "carrier" molecules is the **cyclodextrins** or cycloamyloses.

Cyclodextrins are cyclic oligosaccharides consisting of 6, 7, or 8 glucopyranose units, usually referred to as  $\alpha$ -,  $\beta$ -, or  $\gamma$ -cyclodextrins, respectively. These naturally occurring compounds have relatively rigid doughnut-shaped structures, and have attracted interest as possible natural complexing agents. The unique structures of these compounds owe their stability to intramolecular hydrogen bonding between the C2- and C3-hydroxyl groups of neighboring glucopyranose units. The molecule takes on the shape of a torus with the C2- and C3-hydroxyls located around the larger opening and the more reactive C6-hydroxyl aligned around the smaller opening. The arrangement of C6-hydroxyls opposite the hydrogen bonded C2- and C3-hydroxyls forces the oxygen bonds into close proximity within the cavity, leading to an electron rich, hydrophobic interior. The size of this hydrophobic cavity is a function of the number of glucopyranose units forming the cyclodextrin.

The solubility of natural cyclodextrins is very poor and initially this prevented cyclodextrins from becoming effective complexing agents. In the late 1960's, it was discovered that chemical substitutions at the 2,3, and 6 hydroxyl sites would greatly increase solubility. The degree of chemical substitution, as well as, the nature of the groups used for substitution, determine the final maximum concentration of cyclodextrin in an aqueous medium. Most chemically modified cyclodextrins are able to achieve a 50% (w/v) concentration in water.

Cavity size is the major determinant as to which cyclodextrin is used in complexation. "Fit" is critical to achieving good incorporation of cyclodextrins. Sixglucopyranose unit compounds or  $\alpha$ -cyclodextrins have small cavities which are not capable of accepting many molecules. Eight-glucopyranose unit compounds or  $\gamma$ -cyclodextrins have much larger cavities than many molecules to be incorporated and cyclodextrin hydrophobic charges can't effectively interact to facilitate complexation. The cavity diameter of b-cyclodextrins or  $\beta$ -glucopyranose unit compounds is well-suited for use with molecules the size of hormones, vitamins and many compounds frequently used in tissue and cell culture applications. For this reason,  $\beta$ -cyclodextrin is most commonly used as a complexing agent.

Hydrophobic molecules are incorporated into the cavity of cyclodextrins by displacing water. This reaction is favored by the repulsion of the molecule by water. This effectively encapsulates the molecule of interest within the cyclodextrin, rendering the molecule water soluble. When the water soluble complex is diluted in a much larger volume of aqueous solvent, the process is reversed, thereby releasing the molecule of interest into the solution.

SIGMA'S product line of water-soluble complexes includes host cyclodextrins and cyclodextrinencapsulated forms of many components commonly used in tissue and cell culture applications. For product listings, see the **REAGENTS** section of this catalog.

#### REFERENCES

1. The Source. (1991). Water-Soluble Complexes, Part 1: Cyclodextrins—What are they? Vol. 7 No. 3.

2. The Source. (1992). Water-Soluble Complexes, Part 2: Cyclodextrins and Cell Culture. Vol. 8 No. 1.



# CYCLODEXTRINS

# α- CYCLODEXTRIN

**Product Number: C4680** (Schardinger α - Dextrin; Cyclohexaamylose) Crystalline Molecular Formula: C36 H60 O30 Formula Weight: 972.87 CAS Number: 10016-20-3 <u>Solubility</u>: 50 mg/ ml in water

# $\beta$ - CYCLODEXTRIN

**Product Number: C4805** (Schardinger β- Dextrin; Cycloheptaamylose) Crystalline Molecular Formula: C42 H70 O35 Formula Weight: 1135.0 CAS Number: 7585-39-9 <u>Solubility</u>: 50 mg/ml in 1N NaOH

# $\gamma$ - CYCLODEXTRIN

Product Number: C4930 (Schardinger  $\gamma$ - Dextrin; Cyclooctaamylose) Molecular Formula: C48 H80 O40 Formula Weight: 1297.1 CAS Number: 17465-86-0 Solubility: 50 mg/ml in 1N NaOH

# 2 - HYDROXYPROPYL - $\beta$ - CYCLODEXTRIN Product Number: C0926 Mean degree of substitution: 4 - 10 determined by NMR

Crystalline Formula Weight: Approx. 1576 CAS Number: 128446-35-5 Solubility: 50 mg/ ml in water

#### METHYL - β- CYCLODEXTRIN Product Number: C4555

Mean degree of substitution: 10.5 - 14.7 Crystalline Formula Weight: Approx. 1320 CAS Number: 128446-36-6 <u>Solubility</u>: 50 mg/ ml in water