

# **Products for DNA Research**

2019 Catalog of Modification and Labeling





# TABLE OF CONTENTS

TRODUCTION	2
ABOUT US CATALOG	2
IODIFIERS	4
TERMINUS MODIFIERS SEQUENCE MODIFIERS 3'-MODIFIERS 3'-MODIFIERS CHEMICAL PHOSPHORYLATION ALDEHYDE MODIFICATION SPACER MODIFIERS DENDRIMERS BRANCHING PHOSPHORAMIDITE PHOTOCLEAVABLE MONOMERS CONJUGATION USING CLICK CHEMISTRY OLIGO-CLICK KITS COPPER-FREE CLICK CHEMISTRY	4 7 9 12 13 14 15 15 16 17 20 21
ABELING	24
SERINOL REAGENTS FOR MODIFICATION AND LABELING COT SERINOL PHOSPHORAMIDITE DABCYL LABELING BIOTIN LABELING FLUORESCEIN LABELING (SIMA) CYANINE LABELING ELITECHGROUP DYES AND QUENCHER BLACK HOLE QUENCHER DYES BLACKBERRY® QUENCHER (BBQ-650®) RHODAMINE (TAMRA) LABELING ACRIDINE LABELING DNP LABELING TOCOPHEROL LABELING TOCOPHEROL LABELING STEARYL LABELING N-ACETYLGALACTOSAMINE (GaINAC) LABELING CDDI] MGB™ LABELING PSORALEN LABELING FERROCENE LABELING EDTA LABELING FERROCENE LABELING DTA LABELING EDTA LABELING EDTA LABELING PORALEN LABELING EDTA LABELING OMETHYLENE BLUE LABELING LABELING WITH METAL CHELATES LABELING WITH METAL CHELATES LABELING WITH POLYAROMATIC HYDROCARBONS PUROMYCIN CPG QUENCHED AUTOLIGATION (QUAL) PROBES LABELING WITH ULTRAFAST PHOTO CROSS-LINKER	24 27 28 29 32 35 36 38 40 42 43 44 45 45 45 45 45 45 45 50 50 51 51 52 53
IDEX	54
ENERAL INFORMATION	58
ORDERING DISCOUNTS TERMS AND CONDITIONS OF SALE PATENTS	58 58 58 58

**G**len Research develops, manufactures and markets reagents for oligonucleotide synthesis, modification, labeling and purification. The company serves customers worldwide involved in basic research, diagnostics and therapeutics. Although Glen Research's original mission was to provide state-of-the-art reagents to researchers, the company also began offering standard reagents for oligonucleotide synthesis but with the innovation that every batch was accompanied by a Certificate of Analysis. The analytical techniques and quality criteria used for the evaluation and acceptance of these reagents were to become an industry standard years later. The company is headquartered in Sterling, Virginia. A privately held company, Glen Research was acquired by Maravai LifeSciences in December 2017.

# **OVER 30 YEARS OF ASSURED QUALITY FOR OLIGO SYNTHESIS**

# 1987

Glen Research was incorporated in the Commonwealth of Virginia

# 1993

Glen Research introduced the Sterling line of products, a new standard of quality for oligonucleotide synthesis

# 1996

Company negotiated an exclusive license with Gilead Sciences to supply C5-propynyl pyrimidine nucleosides and G-Clamp phosphoramidites

# 1999

Company awarded patents for a chemical phosphorylation reagent compatible with DMT-ON purification

# 2003

Glen Research negotiated an agreement with GE Healthcare Biosciences Corp. to supply Cyanine Dyes to the research market

# 2006

In collaboration with Berry & Associates, Inc., Glen Research awarded patents for pyrrolo-C analogues (fluorescent C analogues).

# 2013

In collaboration with Nelson Biotechnologies, Inc., company awarded patent for serinol phosphoramidites and supports

# 1991

Company awarded SBIR grant for the investigation of large scale oligonucleotide synthesis using H-phosphonate chemistry

# 1995

Glen Research negotiated an exclusive agreement to supply 5'-biotin phosphoramidite worldwide

# 1997

Glen Research moves into a custom built building in Sterling, Virginia

# 2002

Company made an agreement with Epoch Biosciences, Inc. to supply their proprietary dyes and nucleosides to the research market

# 2004

Company awarded patents for a truly universal support for oligonucleotide synthesis - US III.

# 2008

Glen Research obtained a license for the sale of Glen UnySupport from Ionis Pharmaceuticals

# 2017

Glen Research is acquired by Maravai LifeSciences

# OTHER INSTRUMENT TYPES

All minor bases, RNA products and modifiers are packaged in septum-capped vials suitable for ABI and other instruments. If you would like another type of vial/column add the following to the end of the catalog number.

Monomers For Instrument type	Add
Expedite MerMade	E M
Columns For Instrument type	Add

(Please inquire for availability of vials and columns for other instrument types.)

Applied Biosystems 3900

MerMade

# **CATALOG**

Welcome to the Glen Research Catalog containing the most complete selection of products for DNA and RNA research. The Table of Contents at the beginning and the Index at the end of the Catalog are the most comprehensive we have produced. There are always limitations to printed catalogs in a fast-moving technology sector and a complete and up-to-date catalog is also maintained on our web site.

All minor bases, modifiers and RNA products are packaged for Applied Biosystems instruments. We can provide vials and columns for a wide variety of other instruments. As shown in the table to the left, we can accommodate catalog numbers for unusual products to fit all popular instruments. The table to the left is reproduced on all relevant spreads of this catalog.

We are unique in conducting a QC test for supports to show the length of oligo that can be prepared before a drop-off in coupling due to steric effects begins to occur. The drop-off point is recorded in the Certificate of Analysis or Analytical Report. Unless otherwise specified, our minor base and modification supports are 1000Å CPG, which results in improved performance and the ability to make much longer oligos. Polystyrene supports are also available for some of our most popular items.

For reasons of quality assurance, we do not transfer powders or oils from stock Applied Biosystems vials to vials for other instruments. Powders may be hygroscopic and electrostatic, making transfer difficult, and oils have to be dissolved and the solvent evaporated. For best performance, it is preferable for the customer to dissolve the product and immediately transfer the solution to the correct instrument vial. Consequently, the product will be delivered in an industry-standard septum-capped vial along with a clean dry vial for the appropriate instrument.

Glen Research's distributors cover a very significant percentage of countries where oligonucleotide synthesis is commonly practiced. Our vast selection of unusual products is really only comprehensively stocked here in Virginia and some of our web viewers have asked us to set up a direct shipping channel. For them, we offer the eGlen program which is described in the following web link: <a href="http://www.glenresearch.com/Reference/eGlen.html">http://www.glenresearch.com/Reference/eGlen.html</a>.

Authorized distributors for Glen Research products are listed below. Other countries not listed are covered by direct sales from our Sterling, USA office.

# UK and Ireland

# Cambio Ltd

Telephone Number: +44 (0) 1954 210200 Fax Number: +44 (0) 1954 210300 e-mail addresses: <a href="mailto:support@cambio.co.uk">support@cambio.co.uk</a> and orders@cambio.co.uk

Website: http://www.cambio.co.uk/

# China

Beijing LeBo Biotech Co.,Ltd Telephone Number: +86-10-52405563 Fax Number: +86-10-58850899 email address: <u>info@lab-bio.com</u> Website:<u>http://www.lab-bio.com/</u>

# Netherlands

Eurogentec b.v.
Telephone Number: +31 43 352 06 98
Fax Number: +31 43 354 19 65
e-mail address: info@eurogentec.com

# Nordic and Baltic Countries

BioNordika AS
Telephone Number: +47 23 03 58 00
Fax Number: +47 23 03 58 01
e-mail address: info@bionordika.no
Website: http://www.bionordika.no/

# Belgium

Eurogentec S.A.
Telephone Number: +32 4 372 74 00
Fax Number: +32 4 372 75 00
e-mail address: info@eurogentec.com/
Website: http://www.eurogentec.com/

# Germany

Eurogentec GmbH
Telephone Number: +49 221 258 94 55
Fax Number: +49 221 258 94 54
e-mail address: info@eurogentec.com

# Republic of Korea

Bosung Scientific Co., Ltd.
Telephone Number: +82-02-6105-5630
Fax Number: +82-02-6105-5680
email address: info@bosungsci.com
Website: https://bosungsci.com/

# Japan

Nihon Techno Service Co., Ltd.
Telephone Number: +81 29 886 6811
Fax Number: +81 29 870 0210
e-mail address: info@ntsbio.com
Website: http://www.ntsbio.com/

# Israel

Eisenberg Bros. Ltd.
Telephone Number: 972-3-9777000
Fax Number: 972-3-9777001
e-mail address: nicoles@eb1.co.il
Website: http://www.eisenbros.co.il/

# France

Eurogentec s.a.
Telephone Number: +33 2 41 73 33 73
Fax Number: +33 2 41 73 10 26
e-mail address: info@eurogentec.com

INTELLECTUAL PROPERTY

Berger), Patent pending.

the end of the catalog number.

PDA amino-modifiers were eveloped by

Stefan Pitsch and ReseaChem GmbH (S.

# **INTELLECTUAL PROPERTY**

5'-Carboxy-Modifier C10 is offered for sale under license from TriLink BioTechnologies, Inc. It is intended for research and development purposes only, and may not be used for commercial, clinical, diagnostic or any other use. It is covered under US Patent No. 6,320,041.

# SEE ALSO

PC modifiers on page 16

# **ABBREVIATIONS**

CNEt = Cyanoethyl CPG = Controlled Pore Glass DMT = 4,4'-Dimethoxytrityl Fmoc = Fluorenylmethoxycarbonyl iPr = Isopropyl MMT = 4-Monomethoxytrityl T = Trityl TFA = Trifluroacetyl

# **TERMINUS MODIFIERS**

Glen Research 5'-Modifiers are designed for use in DNA synthesizers to functionalize the 5'-terminus of the target oligonucleotide. The 5'-Amino-Modifiers are available with a variety of chain lengths to fit exactly the desired application.

The DMS(O)MT-protected amino group is easier to deprotect compared to the MMT-protected one. The sulfoxy derivative survives conditions of oligonucleotide synthesis and can either be cleaved with standard deblock solution, or left intact for HPLC purification. At the same time, the DMS(O)MT group is fully compatible with cartridge purification. When detritylation on a cartridge is carried out, the DMS(O)MT+, which is more stable than MMT+, does not reattach itself to an amine. We now offer 5'-DMS(O)MT-Amino-Modifier C6 utilizing this new trityl based protecting group.

5'-Amino-Modifier TEG, a hydrophilic triethylene glycol ethylamine derivative, is 12 atoms in length and fully soluble in aqueous media.

Item	Catalog No.	Pack	Price (\$)
5'-Amino-Modifier C3-TFA	10-1923-90	100 μmole	50.00
	10-1923-02	0.25g	175.00
5'-Amino-Modifier C6	10-1906-90	100 μmole	60.00
	10-1906-02	0.25g	200.00
5'-Amino-Modifier C6-TFA	10-1916-90	100 μmole	30.00
	10-1916-02	0.25g	100.00
5'-Amino-Modifier C12	10-1912-90	100 μmole	90.00
	10-1912-02	0.25g	300.00
5'-Amino-Modifier 5	10-1905-90	100 μmole	60.00
	10-1905-02	0.25g	200.00
5'-DMS(O)MT-Amino-Modifier C6	10-1907-90	100 μmole	60.00
	10-1907-02	0.25g	200.00
5'-Amino-Modifier TEG	10-1917-90	100 μmole	115.00
	10-1917-02	0.25g	500.00
Methacrylate C6 Phosphoramidite	10-1891-90	100 μmole	110.00
	10-1891-02	0.25g	650.00
TFANH 0-P-N(Pr) <sub>2</sub> 0-CNEt	O-P-N(Pr) <sub>2</sub> O-CNEt	TFANH	O-P-N(Pr) <sub>2</sub> O-CNEt
5'-Amino-Modifier C3-TFA 5'-Amino-M	odifier C6	5'-Amino-Modi	fier C6-TFA
MMTNH O-P-N(Pr) <sub>2</sub> 5'-Amino-Modifier C12	MMTNH	O-P-N(Pr) <sub>2</sub> O-CNEt mino-Modifier 5	
F <sub>3</sub> C O O O O O P N(iPr) <sub>2</sub> O CNEt	0,	H	
0-P-N(iPr) <sub>2</sub>	H₃C S—		O—P—N(Pr) <sub>2</sub>     O—CNEt
O O—CNEt		OCH <sub>3</sub>	

Methacrylate C6 Phosphoramidite

5'-DMS(O)MT-Amino-Modifier C6

# **TERMINUS MODIFIERS (CONT.)**

Our more recent 5'-amino modifiers, protected by a novel phthalic acid diamide (PDA) protecting group, are stable solids. In contrast to the TFA protected amino modifiers, which are viscous oils, the analogous PDA protected compounds are granular powders. This important property of these compounds allows straightforward handling, storage and aliquoting and leads to a significant increase in stability.

Deprotection with methylamine in gas phase or aqueous solution or AMA leads to fast and complete removal of the PDA protecting group. However, ammonium hydroxide will not drive the equilibrium reaction to completion and only partial deprotection occurs - overnight deprotection with ammonium hydroxide will yield around 80% active amine.

We are offering three PDA Amino-Modifiers:

- 5'-Amino-Modifier C6-PDA
- Hydrophobic 5'-Amino-Modifier C12-PDA
- Hydrophilic 5'-Amino-Modifier-TEG-PDA

Item	Catalog No.	Pack	Price (\$)
5'-Amino-Modifier C6-PDA	10-1947-90	100 μmole	30.00
	10-1947-02	0.25g	100.00
5'-Amino-Modifier C12-PDA	10-1948-90	100 μmole	65.00
	10-1948-02	0.25g	240.00
5'-Amino-Modifier-TEG-PDA	10-1949-90	100 μmole	105.00
	10-1949-02	0.25g	420.00

5'-Amino-Modifier C6-PDA

5'-Amino-Modifier C12-PDA

5'-Amino-Modifier-TEG-PDA

Monomers For Instrument type	Add
Expedite MerMade	E M
Columns For Instrument type	Add
Expedite Applied Biosystems 3900 MerMade	E A M
(Please inquire for availabi	lity of vials

# INTELLECTUAL PROPERTY

and columns for other instrument types.)

5'-Maleimide Modifier Phosphoramidite is protected by a patent application and is offered by Glen Research under a non-exclusive license agreement from the University of Barcelona.

# **TERMINUS MODIFIERS (CONT.)**

5'-AminoOxy-Modifier 11

The disulfide thiol modifier may be used for introducing 3'- or 5'-thiol linkages. Dithiol Serinol, produced from lipoic acid and our patented serinol backbone, allows easy connection of multiply dithiol-labeled oligos to gold surfaces. 5'-Carboxy-Modifier C10 is a unique linker designed to be added at the terminus of an oligonucleotide synthesis. It generates an activated carboxylic acid N-hydroxysuccinimide (NHS) ester suitable for immediate conjugation on the synthesis column with molecules containing a primary amine, resulting in a stable amide linkage. An alternative carboxylate protecting group is the 2-chlorotrityl group, which is simply removed using the standard deblock cycle to generate a free carboxyl group on an otherwise fully protected oligonucleotide. The 2-chlorotrityl group is also removed during oligo deprotection with ammonium hydroxide or AMA and is incompatible with RP purification techniques. PC Amino-Modifier is a photocleavable C6 amino-modifier, part of our line of photocleavable (PC) modifiers. 5'-AminoOxy-Modifier 11 is based on a tetraethylene glycol linkage for improved solubility and for reducing the potential negative impact on hybridization of the oligo. The oxime formed from the reaction of alkyloxyamines with aldehydes creates a stable covalent bond. In comparison, the imine formed by the conjugation of primary amines with aldehydes is not stable to acidic or basic conditions and requires subsequent reduction with borohydride to form stable amine conjugates. 5'-Maleimide Modifier Phosphoramidite, developed at the University of Barcelona, incorporates a maleimide cycloadduct that is stable to ammonium hydroxide at room temperature. This phosphoramidite can be incorporated into DNA and RNA with both phosphate and phosphorothioate linkages. A retro-Diels-Alder reaction deprotects the maleimide immediately prior to conjugation.

Item	Catalog No.	Pack	Price (\$)
5'-Thiol-Modifier C6	10-1926-90	100 μmole	60.00
	10-1926-02	0.25g	200.00
Thiol-Modifier C6 S-S	10-1936-90	100 μmole	150.00
	10-1936-02	0.25g	360.00
Dithiol Serinol Phosphoramidite	10-1991-95	50 μmole	120.00
	10-1991-90	100 μmole	215.00
	10-1991-02	0.25g	585.00
PC Amino-Modifier Phosphoramidite	10-4906-90	100 μmole	135.00
	10-4906-02	0.25g	395.00
5'-Carboxy-Modifier C10	10-1935-90	100 μmole	65.00
	10-1935-02	0.25g	265.00
5'-Carboxy-Modifier C5	10-1945-90	100 μmole	95.00
	10-1945-02	0.25g	330.00
5'-AminoOxy-Modifier 11	10-1919-95	50 μmole	140.00
	10-1919-90	100 μmole	265.00
	10-1919-02	0.25g	895.00
5'-Maleimide-Modifier Phosphoramidite	10-1938-90	100 μmole	70.00
	10-1938-02	0.25g	335.00
TSO_P_N(Pr)2 DMTO	S S O P N(iPi O CNEt	, in the second	ODMT  O—P—N(iPr
5'-Thiol-Modifier C6  Thio  H <sub>3</sub> C  O-P-N(Pr) <sub>2</sub> O-CNEt  PC Amino-Modifier	N-CH <sub>C</sub> CH <sub>O</sub> -P-N(IPI) <sub>2</sub>	Et	O-CNEI  O-P-N(iPr  O-CNEI
DMT—N 0 0 0 0 — P—N(iPr)2 0—CNEt	H O O-CNEt	5'-Carboxy-I	Modifier C5

5'-Maleimide-Modifier

# **SEQUENCE MODIFIERS**

Sequence Modifiers are designed for use in automated synthesis. The carboxy-dT is hydrolyzed during deprotection and can be coupled directly to a molecule containing a primary amino group by a standard peptide coupling or via the intermediate N-hydroxysuccinimide (NHS) ester. Amino-Modifier dA, Amino-Modifier dC, N2-Amino-Modifier dG and both Amino-Modifier dT products can be added in place of a dA, dC, dG and dT residue, respectively, during oligonucleotide synthesis. Corresponding Amino-Modifier supports can replace their respective deoxynucleoside supports. After deprotection, the primary amine on the C6 analogues is separated from the oligonucleotide by a spacer arm with a total of 7 -10 atoms and can be labeled or attached to an enzyme. The C2 analogue is more suitable for the attachment of molecules designed to react with the oligonucleotide.

Item	Catalog No.	Pack	Price (\$)
Amino-Modifier C6 dA	10-1089-90	100 μmole	205.00
	10-1089-02	0.25g	455.00
Amino-Modifier C6 dC	10-1019-90	100 μmole	225.00
	10-1019-02	0.25g	450.00
N2-Amino-Modifier C6 dG	10-1529-95	50 μmole	240.00
	10-1529-90	100 μmole	480.00
	10-1529-02	0.25g	1100.00
Carboxy-dT	10-1035-90	100 μmole	180.00
	10-1035-02	0.25g	360.00
Amino-Modifier C2 dT	10-1037-90	100 μmole	180.00
	10-1037-02	0.25g	360.00
	10-1037-05	0.5g	720.00
Amino-Modifier C6 dT	10-1039-90	100 μmole	180.00
	10-1039-02	0.25g	360.00
	10-1039-05	0.5g	720.00

Amino-Modifier C6 dT

Amino-Modifier C2 dT

Carboxy-dT

# SEE ALSO

Amino-Modifier supports on page 9

# **OTHER INSTRUMENT TYPES**

All minor bases, RNA products and modifiers are packaged in septum-capped vials suitable for ABI and other instruments. If you would like another type of vial/column add the following to the end of the catalog number.

Monomers For Instrument type	Add
Expedite	E
MerMade	M
Columns For Instrument type	Add
Expedite	E
Applied Biosystems 3900	A
MerMade	M

(Please inquire for availability of vials and columns for other instrument types.)

Our repertoire of NHS ester derivatives has been expanded to include the NHS-Carboxy-dT-CE Phosphoramidite. By making a dT analog of the Carboxy-Modifier C10, it is possible to label one or multiple sites within an oligonucleotide. This opens up the possibility to label any number of different dyes or molecules within an oligonucleotide when the phosphoramidite is unavailable. Doing so is straightforward and may be done manually off the synthesizer or even in a fully-automated manner on the DNA synthesizer.

We have never found conditions which allow the TFA group to be removed from an amino-modifier while the oligonucleotide remains attached to the support. We are able to solve this problem by using a 9-fluorenylmethoxycarbonyl (Fmoc) protecting group. The Fmoc group is removed using a two step procedure, the first to remove the cyanoethyl protection groups and flush the formed acrylonitrile from the synthesis column using 1% diisopropylamine in acetonitrile, and the second to remove the Fmoc group using 10% piperidine in DMF. The amino group so formed on the column can be reacted with a variety of activated esters. We offer Fmoc-Amino-Modifier C6 dT Phosphoramidite as a nucleosidic option and Amino-Modifier Serinol Phosphoramidite as a non-nucleosidic alternative. We also offer S-Bz-Thiol-Modifier C6-dT to join the ranks of thiol-modifiers for oligonucleotide synthesis. Thiol-Modifier C6-dT can be added as usual at the desired locations within a sequence.

Item	Catalog No.	Pack	Price (\$)
NHS-Carboxy-dT	10-1535-90	100 μmole	210.00
	10-1535-02	0.25g	550.00
Fmoc-Amino-Modifier C6 dT	10-1536-90	100 μmole	180.00
	10-1536-02	0.25g	360.00
S-Bz-Thiol-Modifier C6-dT	10-1538-95	50 μmole	130.00
	10-1538-90	100 μmole	245.00
	10-1538-02	0.25g	550.00
Amino-Modifier Serinol Phosphoramidite	10-1997-95	50 μmole	125.00
	10-1997-90	100 μmole	225.00
	10-1997-02	0.25g	595.00

NHS-Carboxy-dT

Fmoc-Amino-Modifier C6 dT

# 3'-MODIFIERS

**3**′-Amino-Modifier CPGs, containing amino groups protected with the base-labile Fmoc group, are designed to functionalize the 3′-terminus of the target oligonucleotide by the introduction of a primary amine. In an alternative approach, the nitrogen destined to become the 3′-amino group is included in a phthalimide (PT) group which is attached to the support through an amide group attached to the aromatic ring. This simple linkage is very stable to all conditions of oligonucleotide synthesis and contains no chiral center. Using an extended ammonium hydroxide treatment (55°C for 17 hours), the cleavage of the amine from the phthalimide is accomplished along with the deprotection of the oligonucleotide. ABI-style columns are supplied unless otherwise requested.

Item	Cat. No.	Pack	Price (\$)
3'-Amino-Modifier C7 CPG 1000	20-2958-01	0.1g	95.00
	20-2958-10	1.0g	675.00
1 μmole columns	20-2958-41	Pack of 4	140.00
0.2 μmole columns	20-2958-42	Pack of 4	85.00
10 μmole column (ABI)	20-2958-13	Pack of 1	250.00
15 μmole column (Expedite)	20-2958-14	Pack of 1	375.00
3'-Amino-Modifier Serinol CPG	20-2997-01	0.1g	95.00
	20-2997-10	1.0g	675.00
0.2 μmole columns	20-2997-42	Pack of 4	85.00
1 μmole columns	20-2997-41	Pack of 4	140.00
10 μmole column (ABI)	20-2997-13	Pack of 1	250.00
15 μmole column (Expedite)	20-2997-14	Pack of 1	375.00
3'-PT-Amino-Modifier C3 CPG	20-2954-01	0.1g	95.00
	20-2954-10	1.0g	675.00
1 μmole columns	20-2954-41	Pack of 4	140.00
0.2 μmole columns	20-2954-42	Pack of 4	85.00
10 μmole column (ABI)	20-2954-13	Pack of 1	250.00
15 μmole column (Expedite)	20-2954-14	Pack of 1	375.00
3'-PT-Amino-Modifier C6 CPG	20-2956-01	0.1g	95.00
	20-2956-10	1.0g	675.00
1 μmole columns	20-2956-41	Pack of 4	140.00
0.2 μmole columns	20-2956-42	Pack of 4	85.00
10 μmole column (ABI)	20-2956-13	Pack of 1	250.00
15 μmole column (Expedite)	20-2956-14	Pack of 1	375.00
3'-PT-Amino-Modifier C6 PS	26-2956-01	0.1g	125.00
	26-2956-10	1.0g	1025.00
200 nmole columns (AB 3900)	26-2956-52	Pack of 10	220.00
40 nmole columns (AB 3900)	26-2956-55	Pack of 10	220.00

3'-Amino-Modifier C7 CPG

3'-PT Amino-Modifier C3 CPG

3'-PT Amino-Modifier C6 CPG

3'-Amino-Modifier Serinol CPG

9

( R

The 3'-Thiol-Modifier S-S CPG supports are used to introduce 3'-thiol linkages with three and six atom spacers into oligonucleotides. 3'-Dithiol Serinol CPG is used to introduce a dithiol group at the 3'-terminus. In conjunction with Dithiol Serinol Phosphoramidite, it is simple to produce oligonucleotides with multiple thiol groups at the 3' terminus, which is ideal for conjugation to gold surfaces. With Glyceryl CPG the 3'-terminus of an oligonucleotide is readily oxidized by sodium periodate to form a 3'-phosphoglycaldehyde. The aldehyde may be further oxidized to the corresponding carboxylic acid. Either the aldehyde or the carboxylate may be used for subsequent conjugation to amine-containing products.

Item	Cat. No.	Pack	Price (\$)
3'-Thiol-Modifier C3 S-S CPG	20-2933-01	0.1g	85.00
	20-2933-10	1.0g	600.00
1 μmole columns	20-2933-41	Pack of 4	125.00
0.2 μmole columns	20-2933-42	Pack of 4	75.00
10 μmole column (ABI)	20-2933-13	Pack of 1	225.00
15 μmole column (Expedite)	20-2933-14	Pack of 1	350.00
3'-Thiol-Modifier 6 S-S CPG	20-2938-01	0.1g	85.00
	20-2938-10	1.0g	600.00
0.2 μmole columns	20-2938-42	Pack of 4	75.00
1 μmole columns	20-2938-41	Pack of 4	125.00
10 μmole column (ABI)	20-2938-13	Pack of 1	225.00
15 μmole column (Expedite)	20-2938-14	Pack of 1	350.00
3'-Dithiol Serinol CPG	20-2991-01	0.1g	120.00
	20-2991-10	1.0g	995.00
0.2 μmole columns	20-2991-42	Pack of 4	120.00
1 μmole columns	20-2991-41	Pack of 4	200.00
10 μmole column (ABI)	20-2991-13	Pack of 1	300.00
15 μmole column (Expedite)	20-2991-14	Pack of 1	450.00
3'-Glyceryl CPG	20-2902-01	0.1g	85.00
	20-2902-10	1.0g	600.00
1 μmole columns	20-2902-41	Pack of 4	125.00
0.2 μmole columns	20-2902-42	Pack of 4	75.00
10 μmole column (ABI)	20-2902-13	Pack of 1	225.00
15 μmole column (Expedite)	20-2902-14	Pack of 1	350.00

All minor bases, RNA products and modifiers are packaged in septumcapped vials suitable for ABI and other instruments. If you would like another type of vial/column add the following to the end of the catalog number.

OTHER INSTRUMENT TYPES

SEE ALSO

Dithiol Serinol on page 6

Monomers For Instrument type	Add
Expedite	E
MerMade	M
Columns For Instrument type	Add
Expedite	E
Applied Biosystems 3900	A
MerMade	M

(Please inquire for availability of vials and columns for other instrument types.)

> S S O-succinyl-lcaa-CPG 3'-Thiol-Modifier 6 S-S CPG 3'-Thiol-Modifier C3 S-S CPG

3'-Glyceryl CPG

# 3'-MODIFIERS (CONT.)

3'-Amino-Modifier C6 dC CPG and 3'-Amino-Modifier C6 dT CPG replace a dC and T, respectively, at the 3'-terminus. These products allow convenient labeling at the 3' without blocking the terminus from desired enzymatic activity.

Item	Cat. No.	Pack	Price (\$)
3'-Amino-Modifier C6 dC CPG	20-2019-01	0.1g	120.00
	20-2019-10	1.0g	995.00
1 μmole columns	20-2019-41	Pack of 4	200.00
0.2 μmole columns	20-2019-42	Pack of 4	120.00
10 μmole column (ABI)	20-2019-13	Pack of 1	300.00
15 μmole column (Expedite)	20-2019-14	Pack of 1	450.00
3'-Amino-Modifier C6 dT CPG	20-2039-01	0.1g	96.00
	20-2039-10	1.0g	800.00
1 μmole columns	20-2039-41	Pack of 4	160.00
0.2 μmole columns	20-2039-42	Pack of 4	96.00
10 μmole column (ABI)	20-2039-13	Pack of 1	240.00
15 μmole column (Expedite)	20-2039-14	Pack of 1	360.00

Amino-Modifier C6 dC CPG

Amino-Modifier C6 dT CPG

Chemical Phosphorylation Reagent is most commonly used to phosphorylate the 5'-terminus of an oligonucleotide. Although this product is also successful in 3'-phosphorylation, 3'-Phosphate CPG allows direct preparation of oligonucleotides with a 3'-phosphate group. Chemical Phosphorylation Reagent II contains a DMT group on a side chain which is stable to base cleavage and can be left on the oligonucleotide for use in RP purification. The DMT group is later removed with aqueous acid and the side chain is eliminated after brief treatment with aqueous ammonium hydroxide to yield the 5'-phosphate.1 Solid CPR II is similar in performance to CPR II but it is easier to prepare aliquots since it is a powder. Many researchers treat synthesis supports with a hindered base (e.g., diethylamine, diisopropylethylamine, or DBU) post-synthesis to eliminate and remove the cyanoethyl phosphate groups. In this way, the acrylonitrile formed in situ is removed from the support and is not available to alkylate dT residues at the N3 position in the oligos. Since the sulfonylethyl group in 3'-Phosphate CPG is also susceptible to ß-elimination leading to oligo cleavage, this technique is not compatible with 3'-phosphate CPG. Using CPR II CPG, which is base labile but does not support ß-elimination, the cyanoethyl groups can be removed from

the oligo prior to cleavage and base deprotection. ABI-style vials and columns are supplied unless otherwise requested.

Item	Cat. No.	Pack
Chemical Phosphorylation Reagent	10-1900-90 10-1900-02	100 μmole 0.25g
3'-Phosphate CPG	20-2900-01 20-2900-10	0.1g 1.0g
1 μmole columns	20-2900-41	Pack of 4

Pack of 4 0.2 µmole columns 20-2900-42 Pack of 4 10 μmole column (ABI) 20-2900-13 Pack of 1 15 µmole column (Expedite) 20-2900-14 Pack of 1 3'-Phosphate PS 26-2900-01 0.1g 26-2900-10 1.0g 200 nmole columns (AB 3900) 26-2900-52 Pack of 10 40 nmole columns (AB 3900) 26-2900-55 Pack of 10 3'-Phosphate CPG 25-2900-01 0.1g (High Load) 25-2900-10 1.0g

Chemical Phosphorylation Reagent II 10-1901-90 100 umole 60.00 (CPR II) 10-1901-02 0.25g 200.00 Solid Chemical Phosphorylation Reagent II 10-1902-90 100 umole 60.00 (Solid CPR II) 10-1902-02 200.00 0.25g 3'-CPR II CPG 20-2903-01 0.1g 70.00 20-2903-10 1.0g 480.00 0.2 umole columns 20-2903-42 Pack of 4 60.00 1 umole columns 20-2903-41 Pack of 4 100.00

20-2903-13

20-2903-14

25-2900-46

O-succinyl-lcaa-CPG

2.5 µmole columns

10 µmole column (ABI)

15 µmole column (Expedite)

.0-P-N(Pr) O-CNEt

MeHNOC CONHMe .O --- succinvl-CPG

Price (\$)

50.00

160.00

70.00

480.00

100.00

60.00

180.00

280.00

75.00

510.00

150.00

150.00

85.00

600.00

120.00

180.00

280.00

Pack of 4

Pack of 1

Pack of 1

Chemical Phosphorylation Reagent II Solid Chemical Phosphorylation Reagent II

**ALDEHYDE MODIFICATION** 

**MODIFIERS** 

Aldehyde modifiers would be attractive electrophilic substitutions in oligonucleotides since they are able to react with amino groups to form a Schiff's base, with hydrazino groups to form hydrazones, and with semicarbazides to form semicarbazones. The Schiff's base is unstable and must be reduced with sodium borohydride to form a stable linkage but hydrazones and semicarbazides are very stable linkages.

Our collaboration with ELITechGroup, formerly Epoch Biosciences, has allowed us to offer 5'-Aldehyde-Modifier C2 Phosphoramidite. The acetal protecting group is sufficiently hydrophobic for use in RP HPLC and cartridge purification and is readily removed after oligonucleotide synthesis under standard oligonucleotide detritylation conditions with 80% acetic acid / 20% water or 2% aqueous trifluoroacetic acid during cartridge purification.

A formylindole nucleoside analogue has been used to introduce aldehyde groups within an oligonucleotide or at the 5' terminus. This product has no protecting group on the aldehyde, which means that deprotection of the modified oligonucleotide can be done without changing preferred conditions.

Item	Cat. No.	Pack	Price (\$)
5'-Aldehyde-Modifier C2 Phosphoramidite	10-1933-90 10-1933-02	100 μmole 0.25g	85.00 325.00
Formylindole CE Phosphoramidite	10-1934-90 10-1934-02	100 μmole 0.25g	85.00 325.00

# INTELLECTUAL PROPERTY

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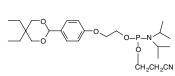
A simple agreement must be signed before end-users and custom oligo services may purchase these products for use as defined above. http://www.glenresearch.com/ Reference/ELITechGroupProducts.pdf

# **OTHER INSTRUMENT TYPES**

All minor bases, RNA products and modifiers are packaged in septumcapped vials suitable for ABI and other instruments. If you would like another type of vial/column add the following to the end of the catalog number.

Monomers For Instrument type	Add
Expedite MerMade	E M
Columns For Instrument type	Add
Expedite Applied Biosystems 3900 MerMade	E A M

(Please inquire for availability of vials and columns for other instrument types.)



O-CNEt

Formylindole

O-CNEt Chemical Phospl

\_O-P-N(IPr)2

**INTELLECTUAL PROPERTY** 

Solid Chemical Phosphorylation

Reagent II and related supports

are covered by European Patent:

1) A. Guzaev, H.Salo, A. Azhayev, and

High load supports on page 4

H. Lonnberg, Tetrahedron, 1995, 51,

EP0816368.

9375-9384.

SEE ALSO

	Α		

PC Modifiers on page 16 Pyrrolidine on page 4

he spacer phosphoramidites C3, 9, C12	2 and 18 are used to insert a spacer arm in	an oligonucleotide. Th	e compounds may	
be added in multiple additions when a longer spacer is required. 3'-Spacer C3 CPG may also act as a blocker of exonuclease				
and polymerase activity at the 3'-terminus. dSpacer is used to introduce a stable abasic site within an oligonucleotide. PC				
Spacer is a photocleavable C3 spacer modifier, part of our line of photocleavable (PC) modifiers.				
Item	Cat. No.	Pack	Price (\$)	

Item	Cat. No.	Pack	Price (\$)
Spacer Phosphoramidite 9	10-1909-90	100 μmole	75.00
	10-1909-02	0.25g	240.00
Spacer Phosphoramidite C3	10-1913-90	100 μmole	75.00
	10-1913-02	0.25g	240.00
dSpacer CE Phosphoramidite	10-1914-90	100 μmole	85.00
	10-1914-02	0.25g	295.00
Spacer Phosphoramidite 18	10-1918-90	100 μmole	95.00
	10-1918-02	0.25g	240.00
Spacer C12 CE Phosphoramidite	10-1928-90	100 μmole	95.00
	10-1928-02	0.25g	240.00
3'-Spacer C3 CPG	20-2913-01	0.1g	70.00
	20-2913-10	1.0g	480.00
1 μmole columns	20-2913-41	Pack of 4	100.00
0.2 μmole columns	20-2913-42	Pack of 4	60.00
10 μmole column (ABI)	20-2913-13	Pack of 1	180.00
15 μmole column (Expedite)	20-2913-14	Pack of 1	280.00
PC Spacer Phosphoramidite	10-4913-90	100 μmole	135.00
	10-4913-02	0.25g	395.00

# O-CNEt O-CNEt dSpacer Spacer C3 O-P-N(Pr)2 DMTO O-CNEt PC Spacer Spacer C12

# **DENDRIMERS**

**D**endrimers are discrete, highly branched, monodispersed polymers that possess patterns reminiscent of the branching of trees. Plain and mixed oligonucleotide dendrimers can be synthesized using novel doubling and trebling phosphoramidite synthons. <sup>1,2</sup> Dendrimers offer the following advantages. Incorporation of label using  $\gamma$ -32P-ATP and polynucleotide kinase increases in proportion to the number of 5'-ends. Fluorescent signal also increases in proportion to the number of 5'-ends, if spacers are incorporated between the labels and the ends of the branches. When using a dendrimeric oligonucleotide as a PCR primer, the strand bearing the dendrimer is resistant to degradation by T7 Gene 6 exonuclease making it easy to convert the double-stranded product of the PCR to a multiply labeled, single-stranded probe. Enhanced stability of DNA dendrimers makes them useful as building blocks for the 'bottom up' approach to nano-assembly. These features also suggest applications in DNA chip technology when higher temperatures are required, for example, to melt secondary structure in the target.

Item	Catalog No.	Pack	Price(\$)
Symmetric Doubler Phosphoramidite	10-1920-90	100 μmole	150.00
	10-1920-02	0.25g	240.00
Asymmetric Doubler (LEV) Phosphoramidite	10-1981-90	100 μmole	105.00
	10-1981-02	0.25g	250.00
Trebler Phosphoramidite	10-1922-90	100 μmole	180.00
	10-1922-02	0.25g	300.00
Long Trebler Phosphoramidite	10-1925-90	100 μmole	200.00
	10-1925-02	0.25g	300.00

# **BRANCHING PHOSPHORAMIDITE**

A branching monomer is required to construct comb-like oligonucleotide probes. The developers of the comb system from Chiron Corporation evaluated<sup>3</sup> several protecting groups for the branch point and chose levulinyl (LEV), which is specifically removed using a reagent containing hydrazine hydrate, acetic acid and pyridine.

Item	Catalog No.	Pack	Price(\$)
5-Me-dC Brancher Phosphoramidite	10-1018-90 10-1018-02	100 μmole 0.25g	205.00 505.00
DMTO  Symmetric Doubler  DMTO  O—P—N(Pr) <sub>2</sub> O—CNEt  DMTO  O—P—N(Pr) <sub>2</sub> O—CNEt  DMTO  O—P—N(Pr) <sub>2</sub> O—CNEt	NH O—P—N(iPr) <sub>2</sub> O—CNEt NH O orier (LEV)	DMTO O DMTO Trebler  DMTO O O O O O O O O O O O O O O O O O O	O-P-N(Pr) <sub>2</sub> O-CNEt
Long Trebler		5-Me-dC Brancher	

# OTHER INSTRUMENT TYPES

All minor bases, RNA products and modifiers are packaged in septumcapped vials suitable for ABI and other instruments. If you would like another type of vial/column add the following to the end of the catalog number.

Monomers For Instrument type	Add
Expedite MerMade	E M
Columns For Instrument type	Add
Expedite Applied Biosystems 3900 MerMade	E A M

(Please inquire for availability of vials and columns for other instrument types.)

# REFERENCES

- (1) M.S. Shchepinov, I.A. Udalova, A.J. Bridgman, and E.M. Southern, Nucleic Acids Res, 1997, 25, 4447-4454.
- (2) M.S. Shchepinov, K.U. Mir, J.K. Elder, M.D. Frank-Kamenetskii, and E.M. Southern, Nucleic Acids Res, 1999, 27, 3035-41.
- (3) T. Horn, C.A. Chang, and M.S. Urdea, Nucleic Acids Res, 1997, 25, 4842-4849.

# **OTHER INSTRUMENT TYPES**

All minor bases, RNA products and modifiers are packaged in septum-capped vials suitable for ABI and other instruments. If you would like another type of vial/column add the following to the end of the catalog number.

# lonomers

For Instrument type	Add			
Expedite MerMade	E M			
Columns For Instrument type	Add			
Expedite Applied Biosystems 3900 MerMade	E A M			
(Please inquire for availability of vials				

and columns for other instrument types.)

# INTELLECTUAL PROPERTY

Glen Research offers PC Biotin, PC Amino-Modifier and PC Spacer products in association with AmberGen, Inc. and Link Technologies, Ltd. For a commercial application license, please contact AmberGen, Inc., +617-923-9990, (sales@ambergen.com), http:// www.ambergen.com/.

PC Linker phosphoramidite is available from Glen Research in association with Link Technologies Ltd (Scotland).

# **SEE ALSO**

5'-Biotin on page 29

# REFERENCES

- P. Ordoukhanian and J-S. Taylor, J. Am. Chem. Soc., 117, 9570-9571, 1995.
   F. Hausch and A. Jäschke. Nucleic Acids
- Research, 2000, **28**, e35.

  (2b) F. Hausch and A. Jäschke, *Tetrahedron*,
- 2001, **57**, 1261-1268.
  (3) T. Wenzel, T. Elssner, K. Fahr, J. Bimmler, S. Richter, I. Thomas, and M. Kostrzewa, *Nucleosides, Nucleotides & Nucleic Acids*, 2003, **22**, 1579-1581.

# **PHOTOCLEAVABLE MONOMERS**

PC Biotin Phosphoramidite can be used to prepare 5'-biotinylated oligonucleotides suitable for capture by streptavidin in a mode similar to our popular 5' Biotin Phosphoramidite. Amino- and thiol-modified oligonucleotides have proven to be very useful for the attachment of a variety of haptens and fluorophores, as well as for the tethering of the oligonucleotides to a diversity of beads and surfaces. PC Amino-Modifier Phosphoramidite is used to prepare 5'-amino-modified oligonucleotides suitable for subsequent photocleavage. PC Spacer Phosphoramidite can be used as an intermediary to attach any modification reagent, available as a phosphoramidite, to the terminus of oligonucleotides. After photocleavage, a 5'-phosphate is generated on the DNA, rendering it suitable for further biological transformations, such as gene construction and cloning after ligation.

A versatile photocleavable DNA building block has been described by researchers in Washington University, Missouri and used in phototriggered hybridization.<sup>1</sup> This reagent has also been used in the design of multifunctional DNA and RNA conjugates<sup>2</sup> for the in vitro selection of new molecules catalyzing biomolecular reactions. Researchers at Bruker Daltonik in Germany have also developed genoSNIP, a method for single-nucleotide polymorphism (SNP) genotyping by MALDI-TOF mass spectrometry.<sup>3</sup> This method uses size reduction of primer extension products by incorporation of the photocleavable linker for phototriggering strand breaks near to the 3' end of the extension primer. PC Linker can be incorporated into oligonucleotides at any position by standard automated DNA synthesis methodology. PC Linker Phosphoramidite has the added advantage in that photocleavage results in monophosphate fragments at both the 3'- and 5'-termini of the oligonucleotide fragments.

Item	Catalog No.	Pack	Price(\$)
PC Biotin Phosphoramidite	10-4950-95	50 μmole	145.00
	10-4950-90	100 μmole	280.00
	10-4950-02	0.25g	675.00
PC Amino-Modifier Phosphoramidite	10-4906-90	100 μmole	135.00
	10-4906-02	0.25g	395.00
PC Spacer Phosphoramidite	10-4913-90	100 μmole	135.00
	10-4913-02	0.25g	395.00
PC Linker Phosphoramidite	10-4920-90	100 μmole	255.00
	10-4920-02	0.25g	795.00

$$\begin{array}{c|c} O & H_3C \\ \hline O - P - N(Pr)_2 \\ \hline O - CNEt \\ \hline \end{array}$$

# CONJUGATION USING CLICK CHEMISTRY

The copper(I)-catalyzed azide-alkyne cycloaddition (CuAAC) reaction between azides and alkynes to form 1,2,3-triazoles, as reported¹ by Sharpless, was found to be so exquisitely regioselective and efficient at even the most mild conditions that Sharpless coined the term 'Click Chemistry' to describe it. The use of this method for DNA modification has been somewhat delayed by the fact that copper ions damage DNA, typically yielding strand breaks.² As these problems have now been overcome by the use of copper(I)-stabilizing ligands (e.g., tris(benzyltriazolylmethyl)amine, TBTA³), Carell et al. and Seela et al. discovered that the CuAAC reaction can be used to functionalize alkyne-modified DNA nucleobases with extremely high efficiency.⁴

Oligonucleotides bearing a single nucleosidic alkyne group can be prepared using a C8-Alkyne-dC or dT-CE Phosphoramidite. Purified oligonucleotides are usually modified with 2-5 equivalents of the corresponding marker-azide (e.g., fluorescent-dye azides). After the addition of precomplexed Cu(I), complete conversion to the labeled oligo is observed in a time span between 30 min and 4 hours. After a simple precipitation step, labeled oligonucleotides can be recovered in near quantitative yields. Using a combination of C8-Alkyne, C8-TIPS-Alkyne and C8-TMS-Alkyne, it is possible to label oligonucleotides in up to three separate click reactions. The alkyne groups on the last two monomers are protected, respectively, with triisopropylsilyl (TIPS) and trimethylsilyl (TMS) protecting groups. The first click reaction on solid phase on a C8-Alkyne yields the singly modified oligonucleotide with full retention of the TIPS and/or TMS protecting group. For double click, a C8-TIPS-Alkyne is used as the second nucleoside and the TIPS protecting group is cleaved with tetrabutylammonium fluoride (TBAF) without causing any damage to the DNA. The second click reaction in solution yields the doubly modified oligonucleotide in excellent yield. For the introduction of three different labels, all three nucleosides are introduced into oligonucleotides. The first click reaction is performed directly on the resin. The singly modified oligonucleotide is subsequently cleaved from the support with concomitant cleavage of the TMS group and retention of the TIPS protecting group. The second click reaction is performed in solution. Precipitation of the doubly modified oligonucleotide, cleavage of the TIPS group with TBAF, and a subsequent third click reaction in solution furnishes the desired triply modified oligonucleotide in excellent overall yield.

Item		Catalog No.	Pack	Price(\$
C8-Alkyne-dT-CE Ph	osphoramidite	10-1540-95	50 μmole	165.0
		10-1540-90	100 μmole	315.0
		10-1540-02	0.25g	900.0
C8-TIPS-Alkyne-dC-	CE Phosphoramidite	10-1541-95	50 μmole	295.0
		10-1541-90	100 μmole	575.0
		10-1541-02	0.25g	1275.0
C8-TMS-Alkyne-dC-	CE Phosphoramidite	10-1542-95	50 μmole	270.0
,	·	10-1542-90	100 μmole	525.0
		10-1542-02	0.25g	1275.0
C8-Alkyne-dC-CE Ph	nosphoramidite	10-1543-95	50 μmole	225.0
		10-1543-90	100 μmole	435.0
		10-1543-02	0.25g	1125.0
	4.	TIPS	TMS	
HN	NHBz N N	NHBz N	NHBz	// <u>/</u> /
	DMTO	DMTO	DMTO	
O—P—N(iPr) <sub>2</sub> I O—CNEt	O—P—N(iPr)₂     O—CNEt	O—P—N(iPr) <sub>2</sub>	0—P—N(iPr)₂       O—CNEt	
C8-Alkyne-dT	C8-TIPS-Alkyne-dC	Ö—CNEt C8-TMS-Alkyne-dC	C8-Alkyne-dC	

# REFERENCES

- C.W. Tornoe, C. Christensen, M. Meldal, J. Org. Chem. 2002, 67, 3057-3064; V.
   V. Rostovtsev, L. G. Green, V. V. Fokin, K.
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- [2] C. J. Burrows, J. G. Muller, *Chem. Rev.* 1998, **98**, 1109 1151.
- [3] T. R. Chan, R. Hilgraf, K. B. Sharpless, V. V. Fokin, *Org. Lett.* 2004, **6**, 2853 – 2855
- [4] J. Gierlich, G. A. Burley, P. M. E. Gramlich, D. M. Hammond, T. Carell, Org. Lett. 2006, 8, 3639-3642. F. Seela, V. R. Sirivolu, Chem. Biodiversity 2006, 3, 509-514.
- [5] P. M. E. Gramlich, S. Warncke, J. Gierlich, T. Carell, Angew. Chem. 2008, 120, 3491–3493; Angew. Chem. Int. Ed. 2008, 47, 3442–3444.
- [6] P. M. E. Gramlich, C. T. Wirges, A. Manetto, T. Carell, *Angew. Chem. Int.* Ed. 2008, 47, 8350-8358.

# INTELLECTUAL PROPERTY

baseclick GmbH has been granted the following patents (1-3) besides its further patent applications (4-5).

- WO 2006/117161
   (New labeling strategies for the sensitive detection of analytes)
- WO 2008/952775 (Click chemistry for the production of reporter molecules)
   WO 2010/115957 (Click Chemistry on
- heterogeneous catalysts)
  4. PCT/EP 2013/064610 (Anandamide-
- PCT/EP 2013/064610 (Anandamide-modified nucleic molecules)
   PCT/EP 2015/056007 (Self-assembly of DNA Origami: a diagnostic tool)
- baseclick GmbH holds a worldwide exclusive license for granted patent application WO 03/101972 (Copper-catalysed ligation of azides and acetylenes for the nucleic acid field) in the area of diagnostics and research.

As Glen Research and baseclick are partners, Glen Research is now able to help in sublicensing this outstanding technology.

O-CNEt

C8-TIPS-Alkyne-dT

O-CNEt

C8-TMS-Alkyne-dT

5-Ethynyl-dU offers convenient click conjugation with an azide to generate a label rigidly attached to one of the oligonucleotide bases. 5-Ethynyl-dU is subject to base-catalyzed hydration during cleavage and deprotection, especially when using a strong base or heat. Hydration of an ethynyl group forms a methyl ketone which subsequently blocks potential click reactions. Mild deprotection conditions are necessary when using 5-Ethynyl-dU-CE Phosphoramidite to prevent this side reaction. TIPS-5-Ethynyl-dU-CE Phosphoramidite, containing a protected alkyne, offers broader compatibility with oligonucleotide synthesis and deprotection. Protecting the 5-ethynyl group with a triisopropylsilyl (TIPS) protecting group prevents acid or base catalyzed hydration during oligonucleotide synthesis and workup. A quick treatment with TBAF removes the TIPS protecting group.

Item	Catalog No.	Pack	Price(\$
C8-TIPS-Alkyne-dT-CE Phosphoramidite	10-1544-95	50 μmole	220.0
oo m o mayne ar oz moophoramate	10-1544-90	100 μmole	425.0
	10-1544-02	0.25g	1020.0
C8-TMS-Alkyne-dT-CE Phosphoramidite	10-1545-95	50 μmole	205.0
,	10-1545-90	100 μmole	395.0
	10-1545-02	0.25g	1050.0
5-Ethynyl-dU-CE Phosphoramidite	10-1554-95	50 μmole	130.0
, ,	10-1554-90	100 μmole	245.0
	10-1554-02	0.25g	775.0
TIPS-5-Ethynyl-dU-CE Phosphoramidite	10-1555-95	50 μmole	195.0
	10-1555-90	100 μmole	370.0
	10-1555-02	0.25g	975.0
THPTA Ligand	50-1004-92	25 μmole	50.0
(Water soluble)	50-1004-90	100 μmole	180.0
Click-Solution (DMSO/t-BuOH)	50-1002-11	10 x 1.0mL	185.0

# OTHER INSTRUMENT TYPES

3'-Propargyl-5-Me-dC CPG on

All minor bases, RNA products and modifiers are packaged in septum-capped vials suitable for ABI and other instruments. If you would like another type of vial/column add the following to the end of the catalog number.

# Monomers

Applied Biosystems 3900

MerMade

SEE ALSO

Expedite MerMade	E M
Columns For Instrument type	Add
Evenedite	_

(Please inquire for availability of vials and columns for other instrument types.)

O-CNEt

TIPS-5-Ethynyl-dU

5-Ethynyl-dU

# CONJUGATION USING CLICK CHEMISTRY (CONT.)

Oligonucleotides prepared using 5'-Hexynyl Phosphoramidite are stable to standard deprotection conditions and exhibit a slightly increased retention time on RP HPLC. Azides are not compatible with oligonucleotide synthesis using phosphoramidites so a post-synthesis reaction is required. Azidobutyrate NHS Ester is used¹ for azido-modification of amines at either the 3'-end or the 5'-end of an oligo and it can even be used for internal modification on an Amino-Modifier-C6 dX residue within the sequence. Specific to the 5'-terminus, 5'-Bromohexyl Phosphoramidite is added in the last cycle. This modifier can then be easily transformed into a 5'-azido group by displacement of bromide using sodium azide.² Alkyne NHS ester allows the functionalization of an amino moiety in a variety of molecules, including DNA and RNA oligonucleotides as well as peptides or proteins. We also offer two products for use in Click Chemistry based upon our 1,3-diol product portfolio with the serinol backbone - a phosphoramidite for adding an alkyne group at the 5' terminus or within the sequence, and a synthesis support for labeling the 3' terminus of oligonucleotides with an alkyne group.

Catalog No.	Pack	Price(\$)
10-1908-90	100 μmole	60.00
10-1908-02	0.25g	200.00
50-1904-23	2.3mg	60.00
50-1904-24	23mg	300.00
10-1946-90	100 μmole	60.00
10-1946-02	0.25g	200.00
50-1905-23	2.3mg	60.00
50-1905-24	23mg	300.00
10-1992-95	50 μmole	100.00
10-1992-90	100 μmole	185.00
10-1992-02	0.25g	575.00
20-2992-01	0.1g	105.00
20-2992-10	1.0g	800.00
20-2992-42 20-2992-41 20-2992-13 20-2992-14	Pack of 4 Pack of 4 Pack of 1 Pack of 1	100.00 175.00 260.00 390.00
	10-1908-90 10-1908-02 50-1904-23 50-1904-24 10-1946-90 10-1946-02 50-1905-23 50-1905-24 10-1992-95 10-1992-90 10-1992-02 20-2992-01 20-2992-10 20-2992-42 20-2992-41 20-2992-13	10-1908-90 100 μmole 10-1908-02 0.25g  50-1904-23 2.3mg 50-1904-24 23mg  10-1946-90 100 μmole 10-1946-02 0.25g  50-1905-23 2.3mg 50-1905-24 23mg  10-1992-95 50 μmole 10-1992-90 100 μmole 10-1992-02 0.25g  20-2992-01 0.1g 20-2992-10 1.0g 20-2992-42 Pack of 4 20-2992-13 Pack of 1

# REFERENCES

- (1) R. Kumar, et al., Journal of the American Chemical Society, 2007, **129**, 6859-6864
- (2) J. Lietard, A. Meyer, J.J. Vasseur, and F. Morvan, *Tetrahedron Letters*, 2007, 48, 8795-8798.

# SEE ALSO

Serinol Products on page 24

Alkyne-NHS Ester

Alkyne-Modifier Serinol Phosphoramidite

3'-Alkyne-Modifier Serinol CPG

18

SEE ALSO

# STABILITY NOTES

Oligonucleotides containing a 5'-iodo group are prepared conventionally with the exception that deprotection is carried out in ammonium hydroxide at room temperature for 24 hours. Under these conditions, degradation of the iodo group was less than 2%.

# **CONJUGATION USING CLICK CHEMISTRY (CONT.)**

**1**-Ethynyl-dSpacer CE Phosphoramidite can be used in any position within an oligonucleotide while still retaining the high efficiency of click chemistry. The modifier is efficiently incorporated into oligonucleotides using standard phosphoramidite chemistry, is stable to common deprotection conditions, and is compatible with Glen-Pak™ purification. 1-Ethynyl-dSpacer generates a substituted 1,2,3-triazole pseudo-nucleobase after click chemistry conjugation with an azide The 1-ethynyl-dSpacer modification exhibits similar duplex stability to the standard dSpacer (10-1914) and destabilizes the duplex when internally incorporated. Upon cycloaddition, the duplex stability is moderated by the resulting structure of the modification. Simple 1,2,3-triazoles were destabilizing, as were modifications that incorporated TEG linkers (6-FAM-TEG and Amino-TEG). Modifications that incorporated aromatic functional groups restored duplex stability to varying degrees with coumarin and psoralen significantly restoring stability. A 5′-iodo-modified oligonucleotide (prepared using 5′-lodo-dT) can be quantitatively converted to the corresponding 5′-azide.

Item	Catalog No.	Pack	Price(\$)
1-Ethynyl-dSpacer CE Phosphoramidite	10-1910-95	50 μmole	180.00
	10-1910-90	100 µmole	340.00
	10-1910-02	0.25g	1250.00
5'-I-dT-CE Phosphoramidite	10-1931-90	100 μmole	85.00
	10-1931-02	0.25g	295.00

# **OLIGO-CLICK KITS**

Oligo-Click Kits contain an air-stable, insoluble Cu(I) source in pellet form in a pre-loaded and ready-to-use vial. Within the kit, the TBTA ligand is replaced by an activator which is compatible with both aqueous and organic solvents. This innovative combination of catalyst and ligand/activator results in a much easier labeling work-flow of only three simple steps. The preparation of the oligonucleotide labeling via CuAAC now requires only a minimal hands-on time of a few minutes or even less and can be carried out in air without any additional precautions. Glen Research is offering the following kits in collaboration with baseclick GmbH.

- Oligo-Kit M Reload: This kit has sufficient reagents for conjugating up to nine alkyne-containing oligonucleotides on a 100 nmole scale or a single oligonucleotide on a 1 μmole scale. *The user must supply the azide and a solvent such as DMSO for dissolving the azide.*
- Oligo-Kit M Biotin, Oligo-Kit M Fluorescein and Oligo-Kit M TAMRA: Each kit has sufficient reagents for conjugating up to seven alkyne-containing oligonucleotides on a 100 nmole scale or a single oligonucleotide on a 1 µmole scale. Each kit contains all of the ingredients necessary, including the azide and DMSO solvent.

Item	Catalog No.	Pack	Price(\$)
baseclick Oligo-Click-M-Reload	50-2100-01	each	120.00
baseclick Oligo-Click-M-Biotin	50-2101-01	each	200.00
baseclick Oligo-Click-M-Fluorescein	50-2102-01	each	240.00
baseclick Oligo-Click-M-TAMRA	50-2103-01	each	270.00

5'-I-dT

# COPPER-FREE CLICK CHEMISTRY

At Glen Research, our goal was to offer a copper-free click phosphoramidite reagent with the following properties:

- Simple to use
- Stable in solution on the synthesizer
- Stable to ammonium hydroxide and AMA
- Excellent click performance in 17 hours or less at room temperature

From the variety of cyclooctyne-based copper-free click reagents so far described, we have chosen to offer compounds based on a dibenzo-cyclooctyne (DBCO) structure. We are offering 5′-DBCO-TEG Phosphoramidite for preparing oligos with a 5′-DBCO modification and DBCO-dT-CE Phosphoramidite for inserting a DBCO group at any position within the oligonucleotide. In addition, we offer a further DBCO phosphoramidite − DBCO-Serinol Phosphoramidite. Using our proprietary serinol backbone as a non-nucleosidic spacer allows the DBCO group to be placed at any location within a sequence with multiple additions clearly possible. DBCO-sulfo-NHS Ester is also offered for post-synthesis conjugation reactions. DBCO-modified oligos may be conjugated with azides in organic solvents, such as DMSO, or aqeous buffers. Depending on the azide used, the reaction will go to completion in 4-17 hours at room temperature. Simple desalting on a Glen Gel-Pak™ leads to a product with virtually quantitative conjugation efficiency.

Note: We now recommend that synthesis of oligos containing DBCO-dT be completed using 0.5 M CSO in anhydrous acetonitrile (40-4632-xx). Acceptable results can be achieved with iodine oxidation if DBCO-dT is subjected to no more than 8-10 cycles.

Item	Catalog No.	Pack	Price(\$)
5'-DBCO-TEG Phosphoramidite	10-1941-95	50 μmole	125.00
	10-1941-90	100 μmole	230.00
	10-1941-02	0.25g	775.00
DBCO-dT-CE Phosphoramidite	10-1539-95	50 μmole	250.00
bboo ar of risophoralinate	10-1539-90	100 μmole	485.00
	10-1539-02	0.25g	975.00
DBCO-sulfo-NHS Ester	50-1941-23	5.2mg	60.00
(Dissolve 5.2mg in 60μL water or DMSO)	50-1941-24	52mg	300.00
DBCO-Serinol Phosphoramidite	10-1998-95	50 μmole	180.00
1	10-1998-90	100 μmole	340.00
	10-1998-02	0.25g	895.00

# DBCO-serinol DBCO-sulfo-NHS Ester

# OTHER INSTRUMENT TYPES

All minor bases, RNA products and modifiers are packaged in septum-capped vials suitable for ABI and other instruments. If you would like another type of vial/column add the following to the end of the catalog number.

_	
Monomers For Instrument type	Add
Expedite MerMade	E M
Columns For Instrument type	Add
Expedite Applied Biosystems 3900 MerMade	E A M

(Please inquire for availability of vials and columns for other instrument types.)

# SEE ALSO

0.5M CSO on page 4 Serinol Products on page 24

Dipivaloyl 6-FAM-TEG Azide

**G**len Research is offering first our most popular labels for general interest and, subsequently, we will add azide products that are not compatible with phosphoramidite chemistry.

Biotin is still our most commonly used label and biotinTEG, with its hydrophilic triethylene glycol spacer, is the most popular biotin product. Desthiobiotin is a biotin analogue that is well captured by streptavidin but the captured product can be easily released by applying a biotin solution to the streptavidin beads. 6-FAM is our most popular fluorescein derivative and we offer azides of both 6-FAM and pivaloyl-protected 6-FAM for situations where subsequent reactions require the 6-FAM to be protected. In both 6-FAM products, the hydrophilic TEG spacer is again used. The azides are offered in 25 and 100 µmole packs for convenient oligonucleotide labeling.

7-Hydroxycoumarin, also known as umbelliferone, is a highly fluorescent, pH-sensitive fluorophore that emits in the blue region of the spectrum. However, its fluorescence is strongly quenched if the hydroxyl is alkylated or phosphorylated, making it useful in high-throughput screening for phosphatases and lipases. Interestingly, it was found that the 3-azido derivative is also highly quenched but, upon reaction with an alkyne in the presence of copper to form the triazole, the fluorescence is restored. The clicked coumarin emits at a lambda max of 480 nm and absorbs at 358 nm.

HEX and TET are two of our most popular fluorescein-based dyes for labeling oligonucleotides. We are happy to offer 6-HEX and 6-TET Azides for use in click conjugations.

Item	Catalog No.	Pack	Price(\$)
BiotinTEG Azide	50-2000-92	25 μmole	150.00
	50-2000-90	100 μmole	450.00
DesthiobiotinTEG Azide	50-2001-92	25 μmole	135.00
	50-2001-90	100 μmole	400.00
Dipivaloyl 6-FAM-TEG Azide	50-2002-92	25 μmole	230.00
	50-2002-90	100 μmole	690.00
6-FAM-TEG Azide	50-2003-92	25 μmole	180.00
	50-2003-90	100 μmole	540.00
Coumarin Azide	50-2004-92	25 μmole	115.00
	50-2004-90	100 μmole	350.00
6-HEX Azide	50-2005-92	25 μmole	150.00
	50-2005-90	100 μmole	450.00
6-TET Azide	50-2006-92	25 μmole	150.00
	50-2006-90	100 μmole	450.00
HN NH S NH NH NH NH NH NH NH NH NH	~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~	~_^0~N <sub>3</sub>	HO 0 0 N <sub>3</sub>
BiotinTEG Azide	DesthiobiotinT		Coumarin Azide
HO O O O O O O O O O O O O O O O O O O	CI CI CI O	CI OH HO CI CI CI CI N <sub>3</sub> N <sub>3</sub>	O OH CI

6-FAM-TEG Azide

6-HEX Azide

6-TET Azide

# **CONJUGATION USING CLICK CHEMISTRY (CONT.)**

Two nitroxide spin labels, TEMPO Azide and TEMPO-TEG Azide, for site directed spin labeling (SDSL) are now offered.

Click Chemistry with psoralen azide and one of our many nucleosidic and non-nucleosidic alkyne derivatives has the potential to generate a variety of practical cross-linkers. The well known reversible cross-linking behavior of psoralen with an adjacent thymidine residue could be very useful.

To better address applications in near-infrared (NIR) imaging, Glen Research is offering a water soluble Disulfo-Cyanine 7 azide that can be easily conjugated to DNA and RNA through standard click chemistry. This long wavelength dye offers the benefits of improved solubility, reduced aggregation, and improved stability in the near-infrared spectrum along with the convenience of click chemistry.

Item	Catalog No.	Pack	Price(\$)
TEMPO Azide	50-2007-92	25 μmole	115.00
	50-2007-90	100 μmole	350.00
TEMPO-TEG Azide	50-2008-92	25 μmole	135.00
	50-2008-90	100 μmole	400.00
Psoralen Azide	50-2009-92	25 μmole	115.00
	50-2009-90	100 μmole	350.00
Disulfo-Cyanine 7 Azide	50-2010-92	25 μmole	325.00
	50-2010-90	100 μmole	975.00

REFERENCE

2006, **8**, 3639-42.

(1) J. Gierlich, G.A. Burley, P.M. Gramlich,

D.M. Hammond, and T. Carell, Org Lett,

Monomers For Instrument type	Add
Expedite	E
MerMade	M
Columns For Instrument type	Add
Expedite	E
Applied Biosystems 3900	A
MerMade	M

(Please inquire for availability of vials and columns for other instrument types.)

# INTELLECTUAL PROPERTY

Serinol Reagents for Modification and Labeling are covered by US Patent No.: 8,394,948.

# SERINOL REAGENTS FOR MODIFICATION AND LABELING

Most popular non-nucleosidic phosphoramidites for modification and labeling are based on two structural types: 1,2-diols and 1,3-diols. Products based on a 1,2-diol backbone were first described to allow amino-modification and biotin labeling. Technically, the 1,2-diol backbone has some drawbacks relative to the 1,3-diol backbone. The 1,2-diol backbone can participate in a dephosphorylation reaction since the 1,2-diol can form a favored 5-membered cyclic phosphate intermediate. This reaction is competitive with simple hydrolysis of the protecting groups and leads to some loss of label. However, the degree of loss at the 3' terminus can be limited by the removal of the cyanoethyl protecting group using DBU or diethylamine prior to the cleavage and deprotection steps. Similarly, loss at the 5' terminus can be eliminated by retaining the DMT group until the oligo is fully deprotected. Fortunately, the elimination reaction is virtually non-existent in the 1,3-diol backbone since the cyclic intermediate would be a 6-membered ring which is not favored for a cyclic phosphate intermediate.

IVD customers have requested a new backbone based on a 1,3-diol that would overcome any technical or IP issues surrounding our current products. We now offer a line of products based on the serinol backbone, which have been developed in close collaboration between Glen Research and Nelson Biotechnologies. Protected Biotin Serinol Phosphoramidite and CPG are protected with a t-butylbenzoyl group on the biotin ring. This group is designed to stop any phosphoramidite reactions at this active position in biotin. This protection avoids branching when using nucleophilic activators like DCI. The protecting group is easily removed during oligonucleotide cleavage and deprotection. The BiotinLC versions are similarly protected and should be useful for the synthesis of highly sensitive biotinylated probes. 6-Fluorescein Serinol Phosphoramidite and CPG are designed to prepare oligonucleotides containing one or several 6-Fluorescein (6-FAM) residues. Amino-Modifier Serinol Phosphoramidite and CPG are used to add amino groups into one or several positions in oligonucleotides. The amino group is protected with Fmoc, which may be removed on the synthesis column prior to solid-phase conjugation to the amino groups, or which may be removed during deprotection for subsequent solution phase conjugation to the amino groups.

Combining lipoic acid and our patented serinol backbone, we now offer Dithiol Serinol Phosphoramidite and the related 3'-Dithiol Serinol CPG. This unique architecture moves the bulky dithiol away from the phosphate backbone, making it suitable for conjugation to gold surfaces. The long spacer arm of Dithiol Serinol also allows multiple consecutive incorporations of the modifier without the need for intermediate spacer phosphoramidite additions to achieve optimal stepwise coupling efficiency.

We offer three products for use in Click Chemistry based upon our 1,3-diol product portfolio with the serinol backbone - a phosphoramidite for adding an alkyne group at the 5' terminus or within the sequence, a synthesis support for labeling the 3' terminus of oligonucleotides with an alkyne group, and DBCO-Serinol phosphoramidite as a copper-free click reagent.

Item	Catalog No.	Pack	Price(\$)
Protected Biotin Serinol Phosphoramidite	10-1993-95	50 μmole	165.00
	10-1993-90	100 μmole	295.00
	10-1993-02	0.25g	675.00
6-Fluorescein Serinol Phosphoramidite	10-1994-95	50 μmole	165.00
	10-1994-90	100 μmole	295.00
	10-1994-02	0.25g	595.00

HN H ODMT

OP-N(iPr)2
O-CNEt

6-Fluorescein Serinol Phosphoramidite

# SERINOL REAGENTS FOR MODIFICATION AND LABELING (CONT.)

Item	Catalog No.	Pack	Price(\$)
Protected BiotinLC Serinol Phosphoramidite	10-1995-95	50 μmole	205.00
	10-1995-90	100 μmole	365.00
	10-1995-02	0.25g	675.00
Amino-Modifier Serinol Phosphoramidite	10-1997-95	50 μmole	125.00
	10-1997-90	100 μmole	225.00
	10-1997-02	0.25g	595.00
Dithiol Serinol Phosphoramidite	10-1991-95	50 μmole	120.00
•	10-1991-90	100 μmole	215.00
	10-1991-02	0.25g	585.00
Alkyne-Modifier Serinol Phosphoramidite	10-1992-95	50 μmole	100.00
,	10-1992-90	100 μmole	185.00
	10-1992-02	0.25g	575.00
DBCO-Serinol Phosphoramidite	10-1998-95	50 μmole	180.00
•	10-1998-90	100 μmole	340.00
	10-1998-02	0.25g	895.00

SEE ALSO

DBCO on page 21

**Amino-Modifier Serinol Phosphoramidite** 

Dithioi Serinoi

Alkyne-Modifier Serinol Phosphoramidite

DBCO-Serinol

SERINOL REAGENTS FOR I	MODIFICATION AN	ID LABELING	i (CONT.)
Item	Catalog No.	Pack	Price(\$)
3'-Protected Biotin Serinol CPG	20-2993-01 20-2993-10	0.1g 1.0g	120.00 995.00
0.2 μmole columns	20-2993-42	Pack of 4	120.00
1 μmole columns	20-2993-41	Pack of 4	200.00
10 μmole column (ABI)	20-2993-13	Pack of 1	300.00
15 μmole column (Expedite)	20-2993-14	Pack of 1	450.00

# **OTHER INSTRUMENT TYPES**

All minor bases, RNA products and modifiers are packaged in septumcapped vials suitable for ABI and other instruments. If you would like another type of vial/column add the following to

9	
Monomers For Instrument type	Add
Expedite MerMade	E M
Columns For Instrument type	Add
Expedite Applied Biosystems 3900 MerMade	E A M

the end of the catalog number.

Expedite	E
MerMade	M
Columns For Instrument type	Add
Expedite	E
Applied Biosystems 3900	A

(Please inquire for availability of vials and columns for other instrument types.)

3'-Protected Biotin Serinol CPG	20-2993-01	0.1g	120.0
	20-2993-10	1.0g	995.0
0.2 μmole columns	20-2993-42	Pack of 4	120.0
1 μmole columns	20-2993-41	Pack of 4	200.0
10 μmole column (ABI)	20-2993-13	Pack of 1	300.0
15 μmole column (Expedite)	20-2993-14	Pack of 1	450.0
3'-6-Fluorescein Serinol CPG	20-2994-01	0.1g	120.0
	20-2994-10	1.0g	995.0
0.2 μmole columns	20-2994-42	Pack of 4	120.0
1 μmole columns	20-2994-41	Pack of 4	200.0
10 μmole column (ABI)	20-2994-13	Pack of 1	300.0
15 μmole column (Expedite)	20-2994-14	Pack of 1	450.0
3'-Protected BiotinLC Serinol CPG	20-2995-01	0.1g	120.0
	20-2995-10	1.0g	995.0
0.2 μmole columns	20-2995-42	Pack of 4	120.0
1 μmole columns	20-2995-41	Pack of 4	200.0
10 μmole column (ABI)	20-2995-13	Pack of 1	300.0
15 μmole column (Expedite)	20-2995-14	Pack of 1	450.0
3'-Amino-Modifier Serinol CPG	20-2997-01	0.1g	95.0
	20-2997-10	1.0g	675.0
0.2 μmole columns	20-2997-42	Pack of 4	85.0
1 μmole columns	20-2997-41	Pack of 4	140.0
10 μmole column (ABI)	20-2997-13	Pack of 1	250.0
15 μmole column (Expedite)	20-2997-14	Pack of 1	375.0

Protected BiotinLC Serinol CPG

# SERINOL REAGENTS FOR MODIFICATION AND LABELING (CONT.)

Item	Catalog No.	Pack	Price(\$)
3'-Dithiol Serinol CPG	20-2991-01	0.1g	120.00
	20-2991-10	1.0g	995.00
0.2 μmole columns	20-2991-42	Pack of 4	120.00
1 μmole columns	20-2991-41	Pack of 4	200.00
10 μmole column (ABI)	20-2991-13	Pack of 1	300.00
15 μmole column (Expedite)	20-2991-14	Pack of 1	450.00
3'-Alkyne-Modifier Serinol CPG	20-2992-01	0.1g	105.00
	20-2992-10	1.0g	800.0
0.2 μmole columns	20-2992-42	Pack of 4	100.0
1 μmole columns	20-2992-41	Pack of 4	175.0
10 μmole column (ABI)	20-2992-13	Pack of 1	260.0
15 μmole column (Expedite)	20-2992-14	Pack of 1	390.0

# COT SERINOL PHOSPHORAMIDITE

**B**right, long-lasting and non-phototoxic organic fluorophores are essential for the continued optimization of a diverse range of imaging applications. However, all currently available technologies remain susceptible to undesirable transitions to dark states. Dark states arise from non-fluorescent triplet electronic configurations from which the rate of return to the ground state is slow. When in the triplet state, the fluorophore is susceptible to photobleaching and fluorescence applications are compromised by unpredictably reducing the signal-to-noise ratio (SNR), as well as limiting the total duration of time over which information can be gathered. The direct conjugation of small-molecule protective agents (PAs) has enabled significant improvements through intra-molecular triplet quenching. Through a partnership with Lumidyne Technologies, Glen Research has created a novel PA-linked phosphoramidite using cyclooctatetraene (COT). COT Serinol Phosphoramidite provides a means to improve the photostability of virtually any fluorophore in a modular fashion. Our spectrofluorometric studies show that the presence of COT limited the amount of photobleaching of an oligo containing the cyanine 5 dye.

Item	Catalog No.	Pack	Price(\$)
COT Serinol Phosphoramidite	10-1996-95	50 μmole	310.00
	10-1996-90 10-1996-02	100 μmole 0.25g	600.00 1800.00

# INTELLECTUAL PROPERTY

This product is covered under US Patent 8.945.515 B2.

Item

**BIOTIN LABELING** 

biotinylated probes.

Biotin Phosphoramidite

BiotinTEG Phosphoramidite

Protected Biotin Serinol Phosphoramidite

Protected BiotinLC Serinol Phosphoramidite

(Please inquire for availability of vials

A molecular beacon probe<sup>1</sup> has its natural fluorescence quenched in solution unless it is hybridized to the target sequence. Consequently, the design of a molecular beacon requires a fluorophore to be in one part of the sequence and the quencher molecule to be in another, with both molecules being separated from the oligonucleotide by a hydrocarbon spacer. The Dabcyl group has been found to be a universal quencher. 3'-Dabsyl CPG and 3'-Dabcyl CPG are used to prepare probes with the quencher blocking the 3'-terminus. 5'-Dabcyl Phosphoramidite locates the quencher at the 5'-terminus and Dabcyl-dT

places it within the sequence, leaving the 3'-terminus available for polymerase extension.			
Item	Catalog No.	Pack	Price(\$)
3'-Dabsyl CPG	20-5911-01	0.1g	120.00
	20-5911-10	1.0g	975.00
1 μmole columns	20-5911-41	Pack of 4	200.00
0.2 μmole columns	20-5911-42	Pack of 4	120.00
10 μmole column (ABI)	20-5911-13	Pack of 1	350.00
15 μmole column (Expedite)	20-5911-14	Pack of 1	500.00
3'-Dabcyl CPG	20-5912-01	0.1g	120.00

20-5912-10 1.0g 975.00 1 μmole columns 20-5912-41 Pack of 4 200.00 0.2 μmole columns 20-5912-42 Pack of 4 120.00 10 μmole column (ABI) 20-5912-13 Pack of 1 350.00 500.00 15 μmole column (Expedite) 20-5912-14 Pack of 1 3'-Dabcyl PS 26-5912-01 0.1g 125.00 1025.00 26-5912-10 1.0g 200 nmole columns (AB 3900) 26-5912-52 Pack of 10 300.00 40 nmole columns (AB 3900) 26-5912-55 Pack of 10 300.00 Dabcyl-dT 10-1058-95 50 μmole 180.00 10-1058-90 100 µmole 325.00 10-1058-02 675.00 0.25g

5'-Dabcyl Phosphoramidite

50 µmole

100 umole

0.25g

125.00

225.00

650.00

DMTO... O-CNEt

> Dabcyl-dT 5'-Dabcyl Phosphoramidite

10-5912-95

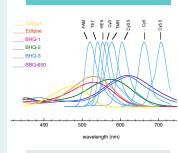
10-5912-90

10-5912-02

(1) S. Tyagi and F.R. Kramer, Nature Biotechnology, 1996, 4, 303-308.

REFERENCE

# DYE QUENCHER PLOT



http://www.glenresearch.com/ ProductFiles/Dye Quencher Plot.pdf

# **OTHER INSTRUMENT TYPES**

All minor bases, RNA products and modifiers are packaged in septumcapped vials suitable for ABI and other instruments. If you would like another type of vial/column add the following to the end of the catalog number.

$\Lambda \Lambda \cap$	no	me	rc
IVIU	טווי	1116	13

For Instrument type	Add
Expedite	E
MerMade	M
Columns For Instrument type	Add
Expedite	E
Applied Biosystems 3900	A

and columns for other instrument types.)

# 0-P-N(*i*Pr) Ó-CNEt **Biotin Phosphoramidite BiotinTEG Phosphoramidite**

**Protected BiotinLC Serinol Phosphoramidite** 

Glen Research biotin phosphoramidites for direct labeling of synthetic oligonucleotides exhibit the following features:

2. All include a DMT group for cartridge purifications which is essential for the preparation of biotinylated PCR primers

3. For the development of diagnostic probes, biotin phosphoramidite is capable of branching to allow multiple biotins

to be introduced at the 3'- or 5'-terminus. BiotinTEG Phosphoramidite contains a 15 atom mixed polarity spacer arm

4. Protected Biotin Serinol Phosphoramidite and CPG are protected with a t-butylbenzoyl group on the biotin ring. This

group is designed to stop any phosphoramidite reactions at this active position in biotin. This protection avoids branching when using nucleophilic activators like DCI. The protecting group is easily removed during oligonucleotide cleavage and deprotection. The BiotinLC versions are similarly protected and should be useful for the synthesis of highly sensitive

Catalog No.

10-1953-95

10-1953-90

10-1953-02

10-1955-95

10-1955-90

10-1955-02

10-1993-95

10-1993-90

10-1993-02

10-1995-95

10-1995-90

10-1995-02

Pack

50 μmole

100 µmole

50 μmole

100 µmole

50 umole

100 umole

50 μmole

100 µmole

0.25g

0.25g

0.25g

0.25g

Price (\$)

165.00 295.00

675.00

165.00

295.00

675.00

165.00

295.00

675.00

205.00

365.00

675.00

1. All are soluble in acetonitrile at concentrations useful for DNA synthesis.

because of the potential for cross contamination in HPLC purifications.

Biotin-dT can replace dT residues within the oligonucleotide sequence. S'-Biotin phosphoramidite can be added ONLY ONCE to the 5'-terminus of an oligonucleotide. However, the DMT group on the biotin can be used in RP cartridge and HPLC purification techniques. PC Biotin is a photocleavable 5'-biotin phosphoramidite. BiotinTEG CPG and Protected BiotinLC Serinol CPG are designed for the direct synthesis of oligonucleotides containing biotin at the 3' terminus.

Desthiobiotin is a biotin analogue that exhibits lower binding to biotin-binding proteins such as streptavidin. This biotin analogue is lacking the sulfur group from the molecule and has a dissociation constant (Kd) several orders of magnitude less than biotin/streptavidin. As a result, biomolecules containing desthiobiotin are dissociated from streptavidin simply in the presence of buffered solutions of biotin. We offer desthiobiotinTEG phosphoramidite and the corresponding CPG.

ABI-style vials and columns are supplied unless otherwise requested (see note box).

lltem	Catalog No.	Pack	Price (\$)
5'-Biotin Phosphoramidite	10-5950-95	50 μmole	125.00
3 -Biotili Filospiloralilluite	10-5950-90	'	225.00
		100 μmole	
	10-5950-02	0.25g	650.00
Biotin-dT	10-1038-95	50 μmole	167.50
	10-1038-90	100 μmole	325.00
	10-1038-02	0.25g	625.00
PC Biotin Phosphoramidite	10-4950-95	50 μmole	145.00
	10-4950-90	100 μmole	280.00
	10-4950-02	0.25g	675.00
DesthiobiotinTEG Phosphoramidite	10-1952-95	50 μmole	185.00
2 common comment	10-1952-90	100 μmole	335.00
	10-1952-02		
	10-1952-02	0.25g	775.00

# **BIOTIN LABELING (CONT.)**

			riice (s
3'-BiotinTEG CPG	20-2955-01	0.1g	120.0
5 5.6tm/26 6. 6	20-2955-10	1.0g	995.0
0.2 μmole columns	20-2955-42	Pack of 4	120.0
1 μmole columns	20-2955-41	Pack of 4	200.0
10 μmole column (ABI)	20-2955-13	Pack of 1	300.0
15 μmole column (Expedite)	20-2955-14	Pack of 1	450.0
3'-BiotinTFG PS	26-2955-01	0.1g	125.0
	26-2955-10	1.0g	1025.0
200 nmole columns (AB 3900)	26-2955-52	Pack of 10	300.0
40 nmole columns (AB 3900)	26-2955-55	Pack of 10	300.0
3'-Protected Biotin Serinol CPG	20-2993-01	0.1g	120.0
	20-2993-10	1.0g	995.0
0.2 μmole columns	20-2993-42	Pack of 4	120.0
1 μmole columns	20-2993-41	Pack of 4	200.0
10 μmole column (ABI)	20-2993-13	Pack of 1	300.0
15 μmole column (Expedite)	20-2993-14	Pack of 1	450.0
3'-Protected BiotinLC Serinol CPG	20-2995-01	0.1g	120.0
	20-2995-10	1.0g	995.0
0.2 μmole columns	20-2995-42	Pack of 4	120.0
1 μmole columns	20-2995-41	Pack of 4	200.0
10 μmole column (ABI)	20-2995-13	Pack of 1	300.0
15 μmole column (Expedite)	20-2995-14	Pack of 1	450.0
DesthiobiotinTEG CPG	20-2952-01	0.1g	140.0
	20-2952-10	1.0g	1150.0
0.2 μmole columns	20-2952-42	Pack of 4	140.0
1 μmole columns	20-2952-41	Pack of 4	230.0
10 μmole column (ABI)	20-2952-13	Pack of 1	345.0
15 μmole column (Expedite)	20-2952-14	Pack of 1	520.0
NH NH	o I		
O-succinyl-C	NH NH		
Protected Biotin Serinol CPG	S NH		O-succinyl-lcaa
NH		BiotinTEG CPG	
	н н		

**Protected BiotinLC Serinol CPG** 

Catalog No.

Pack

Price (\$)

# OTHER INSTRUMENT TYPES

All minor bases, RNA products and modifiers are packaged in septum-capped vials suitable for ABI and other instruments. If you would like another type of vial/column add the following to the end of the catalog number.

Monomers For Instrument type	Add
Expedite	E
Mer Made	M
Columns For Instrument type	Add
Expedite	E
Applied Biosystems 3900	A
MerMade	M

(Please inquire for availability of vials and columns for other instrument types.)

SEE ALSO

PC Biotin on page 16

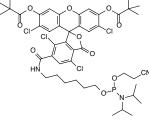
**5**′-Fluorescein phosphoramidite contains no 4,4′-dimethoxytrityl (DMT) group and can be added only once at the 5′-terminus, thereby terminating synthesis. This product is prepared using the 6-carboxyfluorescein derivative. The tetrachloro-, hexachloro-and dichloro-dimethoxy-fluorescein (TET, HEX and JOE, respectively) phosphoramidites are designed to take advantage of the multicolor detection capability of modern DNA sequencers and genetic analyzers. Fluorescein phosphoramidite is designed to produce the same fluorescein-type structure as had been previously prepared using fluorescein isothiocyanate (FITC). Our fluorescein phosphoramidite also contains a DMT group to allow quantification of coupling. The analogous structure, 6-Fluorescein Phosphoramidite, prepared using 6-FAM, is also available, along with 6-Fluorescein Serinol Phosphoramidite. Fluorescein-dT can be inserted into the desired sequence as a replacement for a dT residue.

We offer five fluorescein supports. Fluorescein CPG has traditionally been used to add the fluorescein label at the 3'-terminus. The analogous structure, 3'-(6-Fluorescein) CPG, prepared using 6-FAM, is now also available, along with 6-Fluorescein Serinol CPG. We also offer 3'-(6-FAM) CPG and Fluorescein-dT CPG, both derivatives of 6-carboxyfluorescein (6-FAM). Both are single isomers and use an amide linkage which is stable during cleavage and deprotection and does not allow isomer formation. 3'-(6-FAM) CPG allows effective blockage of the 3'-terminus from polymerase extension as well as exonuclease digestion. Fluorescein-dT CPG allows both of these enzymatic activities to proceed. Normal cleavage and deprotection with ammonium hydroxide readily generates the fluorescein labeled oligos.

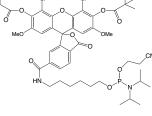
The spectral characteristics of these dyes are detailed on the following page.

,	0.0	•	
Item	Cat. No.	Pack	Price (\$)
5'-Fluorescein Phosphoramidite	10-5901-95	50 μmole	110.00
(6-FAM)	10-5901-90	100 μmole	215.00
,	10-5901-02	0.25g	575.00
5'-Hexachloro-Fluorescein	10-5902-95	50 μmole	190.00
Phosphoramidite	10-5902-90	100 μmole	375.00
(HEX)	10-5902-02	0.25g	875.00
5'-Tetrachloro-Fluorescein	10-5903-95	50 μmole	180.00
Phosphoramidite	10-5903-90	100 μmole	350.00
(TET)	10-5903-02	0.25g	875.00
5'-Dichloro-dimethoxy-Fluorescein Phosphoramidite II	10-5906-95	50 μmole	105.00
(JOE)	10-5906-90	100 μmole	198.00
	10-5906-02	0.25g	495.00

5'-Hexachloro-Fluorescein Phosphoramidite



5'-Tetrachloro-Fluorescein Phosphoramidite



5'-Dichloro-dimethoxy-Fluorescein Phosphoramidite II

# **FLUORESCEIN LABELING (CONT.)**

Fluorescein Phosphoramidite

Item	Cat. No.	Pack	Price (\$)
Fluorescein Phosphoramidite	10-1963-95	50 μmole	165.00
	10-1963-90	100 µmole	295.00
	10-1963-02	0.25g	595.00
6-Fluorescein Phosphoramidite	10-1964-95	50 μmole	165.00
•	10-1964-90	100 µmole	295.00
	10-1964-02	0.25g	595.00
6-Fluorescein Serinol Phosphoramidite	10-1994-95	50 μmole	165.00
	10-1994-90	100 μmole	295.00
	10-1994-02	0.25g	595.00
Fluorescein-dT Phosphoramidite	10-1056-95	50 μmole	180.00
·	10-1056-90	100 μmole	325.00
	10-1056-02	0.25g	675.00

O O O O O O O O O O O O O O O O O O O	DMTO O NH O O O O O O O O O O O O O O O O O
Elucrossoin Phosphoramidite	Fluorescein dT

ODMT O-P-N(Pr) <sub>2</sub> O-CNEt	0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0
6-Fluorescein Phosphoramidite	Ö—CNEt  6-Fluorescein Serinol Phosphoramidite

# FLUORESCENT DYES

	Absorbance	Emission	Color
	Maximum	Maximum	1
Fluorescein	494nm	525nm	Green
Tetrachloro-	521nm	536nm	Orange
Fluorescein			
Hexachloro-	535nm	556nm	Pink
Fluorescein			
SIMA (HEX)	538nm	551nm	Pink
Dichloro-	525nm	548nm	Orange/Pin
dimethoxy-			
Fluorescein			
TAMRA	565nm	580nm	Rose
Cy3	546nm	563nm	Red
Cy3.5	588nm	604nm	Purple
Cy5	646nm	662nm	Violet
Cy5.5	683nm	707nm	Dark Blue
Yakima Yellov	v 530nm	549nm	Yellow
Redmond Red	579nm	595nm	Red

# OTHER INSTRUMENT TYPES

All minor bases, RNA products and modifiers are packaged in septum-capped vials suitable for ABI and other instruments. If you would like another type of vial/column add the following to the end of the catalog number.

Monomers	
For Instrument type	

Expedite	E
Mer Made	M
Columns For Instrument type	Add
Expedite	E
Applied Biosystems 3900	A

(Please inquire for availability of vials and columns for other instrument types.)

5'-Fluorescein Phosphoramidite

DYE QUENCHER PLOT

http://www.glenresearch.com/

ProductFiles/Dye\_Quencher\_Plot.pdf

BHQ-1 BHQ-2 FAM TET COS TWEE COS

3'-Fluorescein CPG

# **LABELING**

5 Tidorescenter G	20 2303 01	0.16	120.00
	20-2963-10	1.0g	995.00
1 μmole columns	20-2963-41	Pack of 4	200.00
0.2 μmole columns	20-2963-42	Pack of 4	120.00
10 μmole column (ABI)	20-2963-13	Pack of 1	300.00
15 μmole column (Expedite)	20-2963-14	Pack of 1	450.00
3'-(6-Fluorescein) CPG	20-2964-01	0.1g	120.00
	20-2964-10	1.0g	995.00
1 μmole columns	20-2964-41	Pack of 4	200.00
0.2 μmole columns	20-2964-42	Pack of 4	120.00
10 μmole column (ABI)	20-2964-13	Pack of 1	300.00
15 μmole column (Expedite)	20-2964-14	Pack of 1	450.00
3'-(6-FAM) CPG	20-2961-01	0.1g	120.00
	20-2961-10	1.0g	995.00
1 μmole columns	20-2961-41	Pack of 4	200.00
0.2 μmole columns	20-2961-42	Pack of 4	120.00
10 μmole column (ABI)	20-2961-13	Pack of 1	300.00
15 μmole column (Expedite)	20-2961-14	Pack of 1	450.00
3'-(6-FAM) PS	26-2961-01	0.1g	130.00
	26-2961-10	1.0g	1045.00
200 nmole columns (AB 3900)	26-2961-52	Pack of 10	300.00
40 nmole columns (AB 3900)	26-2961-55	Pack of 10	300.00
3'-6-Fluorescein Serinol CPG	20-2994-01	0.1g	120.00
	20-2994-10	1.0g	995.00
0.2 μmole columns	20-2994-42	Pack of 4	120.00
1 μmole columns	20-2994-41	Pack of 4	200.00
10 μmole column (ABI)	20-2994-13	Pack of 1	300.00
15 μmole column (Expedite)	20-2994-14	Pack of 1	450.00
		\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	•

Cat. No.

20-2963-01

Pack

0.1g

3'-(6-FAM) CPG

Price (\$)

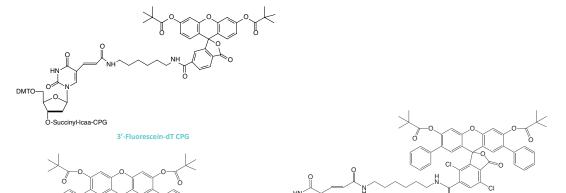
120.00

Item	Cat. No.	Pack	Price (\$)
3'-Fluorescein-dT CPG	20-2056-01	0.1g	120.00
	20-2056-10	1.0g	995.00
1 μmole columns	20-2056-41	Pack of 4	200.00
0.2 μmole columns	20-2056-42	Pack of 4	120.00
10 μmole column (ABI)	20-2056-13	Pack of 1	300.00
15 μmole column (Expedite)	20-2056-14	Pack of 1	450.00

# **FLUORESCEIN LABELING (SIMA)**

Dichloro-diphenyl-fluorescein, SIMA (HEX) exhibits virtually identical absorbance and emission spectra to HEX. SIMA (HEX) is much more stable to basic deprotection conditions than HEX and oligonucleotides can be deprotected using ammonium hydroxide at elevated temperatures and even ammonium hydroxide/methylamine (AMA) at room temperature or 65°C for 10 minutes. SIMA absorption maximum was 3 nm blue-shifted compared to HEX at pH 7. The absorbance is broader, so the extinction coefficient is smaller than that of HEX, but when exciting at 500 nm where the absorbance was normalized, the emission was still 90% of HEX and the emission was red-shifted by 5 nm. A second SIMA (HEX) product, SIMA (HEX)-dT, can be used to introduce SIMA (HEX) in the synthetic oligonucleotide sequence, usually as a replacement for the native dT linkage. Again, this product is fully compatible with deprotection schemes using ammonium hydroxide at elevated temperatures or AMA at room temperature and 65°C.

Item	Cat. No.	Pack	Price (\$)
SIMA (HEX) Phosphoramidite	10-5905-95	50 μmole	90.00
	10-5905-90	100 μmole	175.00
	10-5905-02	0.25g	400.00
SIMA (HEX)-dT Phosphoramidite	10-5945-95	50 μmole	345.00
	10-5945-90	100 μmole	675.00
	10-5945-02	0.25g	995.00



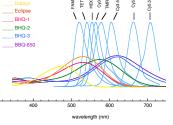
O-CNEt

All minor bases, RNA products and modifiers are packaged in septumcapped vials suitable for ABI and other instruments. If you would like another type of vial/column add the following to the end of the catalog number.

Monomers For Instrument type	Add
Expedite MerMade	E M
Columns For Instrument type	Add
Expedite Applied Biosystems 3900 MerMade	E A M
(Please inquire for available	lity of vials

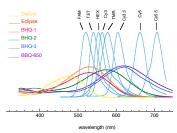
and columns for other instrument types.)

# DYE QUENCHER PLOT



http://www.glenresearch.com/ ProductFiles/Dye\_Quencher\_Plot.pdf

# DYE QUENCHER PLOT



http://www.glenresearch.com/ ProductFiles/Dye\_Quencher\_Plot.pdf

O-succinyl-CPG

3'-(6-Fluorescein) CPG

3'-6-Fluorescein Serinol CPG

SIMA (HEX) Phosphoramidite

SIMA (HEX)-dT Phosphoramidite

Two cyanine derivatives, Cyanine 3 and Cyanine 5, which differ in structure simply by the number of carbons in the conjugated poly-ene linkage, are joined by the closely related analogues, Cyanine 3.5 and Cyanine 5.5, and are available as phosphoramidites. Cyanine dyes are normally added once at the 5'-terminus and the MMT group should be removed on the synthesizer. The absorbance of the MMT cation (yellow) is noticeably different from the DMT cation (orange), and so, absorbance-based trityl monitors will detect it incorrectly as a low coupling. On the other hand, conductivity detectors will interpret the release more correctly. Cyanine dye phosphoramidites have also been used successfully adjacent to the 3'-terminus. Cyanine 3 and Cyanine 5 supports are also offered to allow simpler production of 3' cyanine dye-labeled oligonucleotides.

Deprotection of oligos containing Cyanine dyes may be carried out with ammonium hydroxide at room temperature, regardless of the base protecting groups on the monomers used. If there is a need to use ammonium hydroxide at elevated temperature, Cyanine 3 and Cyanine 3.5 are more stable than Cyanine 5 and Cyanine 5.5. However, it is always prudent to use monomers with base labile protecting groups to limit the exposure time to 2 hours or less at 65°C during deprotection.

To better address applications in near-infrared (NIR) imaging, Glen Research is offering a water soluble Disulfo-Cyanine 7 azide that can be easily conjugated to DNA and RNA through standard click chemistry. This long wavelength dye offers the benefits of improved solubility, reduced aggregation, and improved stability in the near-infrared spectrum along with the convenience of click chemistry.

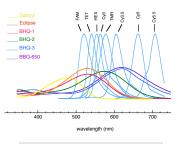
Cyanine 3	546nm	563nm	Red
Cyanine 3.5	588nm	604nm	Purple
Cyanine 5	646nm	662nm	Violet
Cyanine 5.5	683nm	707nm	Dark Blue
Cyanine 7	750nm	773nm	Dark Green
(Measured in	an oligo	in 0.1M TEA	A buffer,
pH7.)			

Absorbance Emission Color Maximum Maximum

# DYE QUENCHER PLOT

SPECTRAL DATA FOR

CYANINE DYES



http://www.glenresearch.com/ ProductFiles/Dye Quencher Plot.pdf

Item	Cat. No.	Pack	Price (\$)
Cyanine 3 Phosphoramidite	10-5913-95	50 μmole	205.00
	10-5913-90	100 μmole	375.00
	10-5913-02	0.25g	925.00
Cyanine 3.5 Phosphoramidite	10-5914-95	50 μmole	220.00
	10-5914-90	100 μmole	400.00
	10-5914-02	0.25g	925.00
Cyanine 5 Phosphoramidite	10-5915-95	50 μmole	205.00
	10-5915-90	100 μmole	375.00
	10-5915-02	0.25g	925.00
Cyanine 5.5 Phosphoramidite	10-5916-95	50 μmole	245.00
	10-5916-90	100 μmole	450.00
	10-5916-02	0.25g	925.00
©N CI-	e <sub>N</sub>		
OMMT O_P_N(iPr) <sub>2</sub>	OMMT	D—P—N(iPr) <sub>2</sub>	
Ö—CNEt  Cyanine 3 Phosphoramidite	Cvanine 3.5 Ph	Ö—CNEt nosphoramidite	
e <sub>N</sub>			
$ \begin{array}{ccc}  & \downarrow & \\  &$	ОММТ	0 1 14(11.1)2	
O—CNEt		O-CNEt	

Cyanine 5 Phosphoramidite

Cyanine 5.5 Phosphoramidite

# **CYANINE LABELING (CONT.)**

Item	Cat. No.	Pack	Price (\$)
Cyanine 3 CPG	20-5913-01	0.1g	160.00
Cyanine 3 Cr O	20-5913-10	0.1g 1.0g	1250.00
1 μmole columns (TWIST format only)	20-5913-41	Pack of 4	250.00
0.2 μmole columns	20-5913-42	Pack of 4	70.00
6	20 5015 01	0.1	160.00
Cyanine 5 CPG	20-5915-01 20-5915-10	0.1g 1.0g	160.00 1250.00
1 μmole columns (TWIST format only)	20-5915-41	Pack of 4	250.00
0.2 µmole columns	20-5915-42	Pack of 4	70.00
Disulfo-Cyanine 7 Azide	50-2010-92 50-2010-90	25 μmole 100 μmole	325.00 975.00
	30-2010-90	100 μποιε	373.00

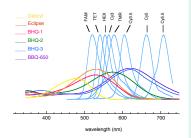
# OTHER INSTRUMENT TYPES

All minor bases, RNA products and modifiers are packaged in septum-capped vials suitable for ABI and other instruments. If you would like another type of vial/column add the following to the end of the catalog number.

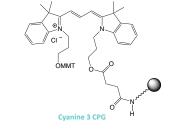
onomers r Instrument type	Add
oedite	E
er Made	M
olumns r Instrument type	Add
oedite	E
plied Biosystems 3900	A
erMade	M

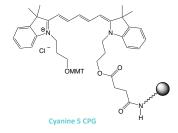
(Please inquire for availability of vials and columns for other instrument types.)

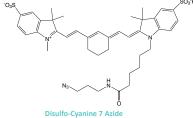
# DYE QUENCHER PLOT



http://www.glenresearch.com/ ProductFiles/Dye\_Quencher\_Plot.pdf







# **FLUORESCENT DYES**

A	bsorbance	Emission	Color
1	Maximum	Maximum	
Yakima Yellow	530nm	549nm	Yellow
Redmond Red	579nm	595nm	Red
AquaPhluor 593	593nm	613nm	Red

# INTELLECTUAL PROPERTY

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A simple agreement must be signed before end-users and custom oligo services may purchase these products for use as defined above. http://www.glenresearch.com/ Reference/ELITechGroupProducts.pdf

AquaPhluor®, Yakima Yellow®, Redmond Red® and Eclipse®, are registered Trademarks of ELITechGroup.

# Glen Research's agreement with ELITechGroup, formerly Epoch Biosciences, allows us to offer several of their proprietary products designed for the synthesis of novel DNA probes. We are pleased to offer products based on ELITechGroup's Redmond Red®, Yakima Yellow® and AquaPhluor® 593 fluorophores and Eclipse® non-fluorescent quencher. Under our agreement we also supply PPG, a modified nucleoside, and 5'-Aldehyde-Modifier C2 Phosphoramidite. The fluorescent dyes, Yakima Yellow, Redmond Red and AquaPhluor 593, are available as phosphoramidites and supports. Yakima Yellow has an absorbance maximum at 530 nm and emission maximum at 549 nm, Redmond Red's absorbance and emission maxima are at 579 nm and 595 nm, respectively, and AquaPhluor 593 has an absorbance maximum at 593 nm and emission maximum at 613 nm.

The Eclipse quencher from ELITechGroup solves most of the problems inherent in the synthesis of molecular beacon and FRET probes. The Eclipse molecule is highly stable and can be used safely in all common oligo deprotection schemes. The absorbance maximum for Eclipse Quencher is at 522 nm, compared to 479 nm for dabcyl. In addition, the structure of the Eclipse Quencher is substantially more electron deficient than that of dabcyl and this leads to better quenching over a wider range of dyes, especially those with emission maxima at longer wavelengths (red shifted) such as Redmond Red and Cyanine 5. In addition, with an absorption range from 390 nm to 625 nm, the Eclipse Quencher is capable of effective performance in a wide range of colored FRET probes.

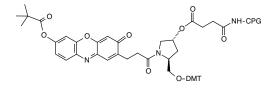
Item	Cat. No.	Pack	Price (\$)
Redmond Red® Phosphoramidite	10-5920-95	50 μmole	220.00
	10-5920-90	100 μmole	420.00
	10-5920-02	0.25g	1045.00
Yakima Yellow® Phosphoramidite	10-5921-95	50 μmole	230.00
Tallina Tene II - Hoopherannaice	10-5921-90	100 μmole	440.00
	10-5921-02	0.25g	1045.00
5'-AquaPhluor® 593 Phosphoramidite	10-5923-95	50 μmole	405.00
·	10-5923-90	100 μmole	795.00
	10-5923-02	0.25g	1575.00
Eclipse® Quencher Phosphoramidite	10-5925-95	50 μmole	250.00
Zonpoe Quenone mosphorumane	10-5925-90	100 μmole	480.00
	10-5925-02	0.25g	1185.00

**ELITECHGROUP DYES AND QUENCHER** 

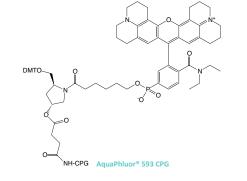
Epoch Eclipse™ Quencher

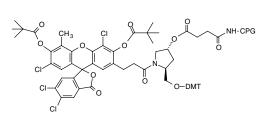
# **ELITECHGROUP DYES AND QUENCHER (CONT.)**

Item	Cat. No.	Pack	Price (\$)
Redmond Red® CPG	20-5920-01	0.1g	180.00
Neumona Neu Cru	20-5920-01	1.0g	1500.00
1 μmole columns	20-5920-10	Pack of 4	300.00
0.2 μmole columns	20-5920-42	Pack of 4	150.00
10 μmole column (ABI)	20-5920-13	Pack of 1	750.00
15 μmole column (Expedite)	20-5920-14	Pack of 1	1125.00
Yakima Yellow® CPG	20-5921-01	0.1g	180.00
	20-5921-10	1.0g	1500.00
1 μmole columns	20-5921-41	Pack of 4	300.00
0.2 μmole columns	20-5921-42	Pack of 4	150.00
10 μmole column (ABI)	20-5921-13	Pack of 1	750.00
15 μmole column (Expedite)	20-5921-14	Pack of 1	1125.00
AquaPhluor® 593 CPG	20-5923-01	0.1g	215.00
	20-5923-10	1.0g	1800.00
1 μmole columns	20-5923-41	Pack of 4	325.00
0.2 μmole columns	20-5923-42	Pack of 4	165.00
10 μmole column (ABI)	20-5923-13	Pack of 1	925.00
15 μmole column (Expedite)	20-5923-14	Pack of 1	1395.00
Eclipse® Quencher CPG	20-5925-01	0.1g	230.00
	20-5925-10	1.0g	1925.00
1 μmole columns	20-5925-41	Pack of 4	350.00
0.2 μmole columns	20-5925-42	Pack of 4	175.00
10 μmole column (ABI)	20-5925-13	Pack of 1	995.00
15 μmole column (Expedite)	20-5925-14	Pack of 1	1495.00

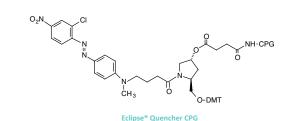


Redmond Red® CP





Yakima Yellow® CPG



# OTHER INSTRUMENT TYPES

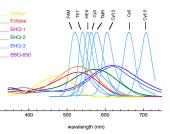
All minor bases, RNA products and modifiers are packaged in septum-capped vials suitable for ABI and other instruments. If you would like another type of vial/column add the following to the end of the catalog number.

Monomers For Instrument type	Add
Expedite MerMade	E M
Columns For Instrument type	Add
Expedite Applied Biosystems 3900 MerMade	E A M

**DYE QUENCHER PLOT** 

(Please inquire for availability of vials

and columns for other instrument types.)



http://www.glenresearch.com/ ProductFiles/Dye\_Quencher\_Plot.pdf

# **BLACK HOLE QUENCHER DYES**

QUENCHERS E260 (nm) (L/mol.cm) (L/mol.cm) BHQ-1 534 8,000 34 000 BHQ-2 579 8,000 38,000 42,700 BHQ-3 672 13,000

# REFERENCES

**TABLE 1: BLACK HOLE** 

(1) S.A.E. Marras, F.R. Kramer, and S. Tyagi, Nucleic Acids Res., 2002, 30, E122. (2) M.K. Johansson, H. Fidder, D. Dick, and R.M. Cook, J Am Chem Soc, 2002, 124,

# SEE OTHER QUENCHERS

Dabcyl on page 28 Eclipse™ on page 39 BBQ-650® on page 42

# **INTELLECTUAL PROPERTY**

"Black Hole Quencher", "BHQ-0", "BHQ-1", "BHQ-2" and "BHQ-3" are trademarks of Biosearch Technologies, Inc., Novato, CA. The BHQ dye technology is the subject of pending patents and is licensed and sold under agreement with Biosearch Technologies, Inc.. Products incorporating the BHQ dye moiety are sold exclusively for R&D use by the end-user. They may not be used for clinical or diagnostic purposes and they may not be re-sold, distributed or re-packaged.

With the growing popularity of red and near-infrared dyes, we are offering the Black Hole Quencher™ dyes (BHQs), whose physical properties are detailed in Table 1. BHQ dyes are robust dark quenchers that very nicely complement our existing product line. They are compatible with ammonium hydroxide deprotection, exhibit excellent coupling efficiencies, have large extinction coefficients and are completely non-fluorescent. Their absorbances are well-tuned to quench a variety of popular fluorophores – even those far into the red, such as Cy3 and Cy5. The dark quencher most typically used in a Molecular Beacon is Dabcyl. Because the quenching does not involve FRET, there is little, if any, dependence upon donoracceptor spectral overlap. In a comprehensive paper by Marras, Kramer and Tyagi, 1 the ability of BHQ-1 and BHQ-2 to quench 22 different fluorophores was evaluated. For shorter wavelength fluorophores such as fluorescein, the quenching efficiency was roughly the same as Dabcyl (91% - 93%). However, for dyes emitting in the far red, such as Cy5, the BHQ dyes were far superior – quenching the Cy5 with 96% efficiency, compared to 84% with Dabcyl. This may reflect the BHQ's ability to form stable, non-fluorescent complexes which can be a plus even in FRET probes. Indeed, recent work suggests that these non-fluorescent complexes will form even in the absence of a hairpin stem structure used by Molecular Beacons.<sup>2</sup>

Item	Cat. No.	Pack	Price (\$)
E/ DUO 1 Dhaanhananidika	10-5931-95	FOl-	100.00
5'-BHQ-1 Phosphoramidite		50 μmole	100.00
	10-5931-90	100 μmole	200.00
	10-5931-02	0.25g	700.00
5'-BHQ-2 Phosphoramidite	10-5932-95	50 μmole	100.00
	10-5932-90	100 μmole	200.00
	10-5932-02	0.25g	700.00
BHQ-1-dT	10-5941-95	50 μmole	265.00
	10-5941-90	100 μmole	525.00
	10-5941-02	0.25g	925.00
BHQ-2-dT	10-5942-95	50 μmole	265.00
	10-5942-90	100 μmole	525.00
	10-5942-02	0.25g	925.00

# LABELING

# **BLACK HOLE QUENCHER DYES (CONT.)**

Item	Cat. No.	Pack	Price (\$)
3'-BHQ-1 CPG	20-5931-01	0.1g	190.00
•	20-5931-10	1.0g	1500.00
1 μmole columns	20-5931-41	Pack of 4	300.00
0.2 μmole columns	20-5931-42	Pack of 4	80.00
10 μmole column (ABI)	20-5931-13	Pack of 1	575.00
15 μmole column (Expedite)	20-5931-14	Pack of 1	825.00
3'-BHQ-2 CPG	20-5932-01	0.1g	190.00
	20-5932-10	1.0g	1500.00
1 μmole columns	20-5932-41	Pack of 4	300.00
0.2 μmole columns	20-5932-42	Pack of 4	80.00
10 μmole column (ABI)	20-5932-13	Pack of 1	575.00
15 μmole column (Expedite)	20-5932-14	Pack of 1	825.00
3'-BHQ-3 CPG	20-5933-01	0.1g	190.00
	20-5933-10	1.0g	1500.00
1 μmole columns	20-5933-41	Pack of 4	300.00
0.2 μmole columns	20-5933-42	Pack of 4	80.00
10 μmole column (ABI)	20-5933-13	Pack of 1	575.00
15 μmole column (Expedite)	20-5933-14	Pack of 1	825.00

# OTHER INSTRUMENT TYPES

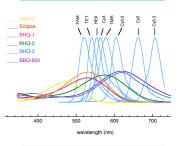
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lonomers or Instrument type	Add
pedite erMade	E M
olumns or Instrument type	Add
pedite	Е

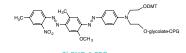
(Please inquire for availability of vials and columns for other instrument types.)

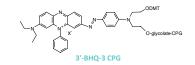
# **Dye Quencher Plot**

Applied Biosystems 3900 MerMade



http://www.glenresearch.com/ ProductFiles/Dye Quencher Plot.pdf





We are happy to offer several products containing the BlackBerry® Quencher (BBQ-650®), which exhibits a broad absorption profile from 550nm to 750nm, centered at 650nm. This range offers more effective quenching of some of our popular long wavelength dyes like TAMRA, Redmond Red, Cy dyes and DyLight dyes. We offer BBQ-650 products for the 3' and 5' termini, as well as BBQ-650-dT for inclusion within the oligonucleotide sequence, with the following properties:

- Quenches the fluorescence of long wavelength dyes
- Quenches in FRET and contact mode
- Absorbance maximum at ~650nm
- Quenching range 550-750nm
- Compatible with standard oligo synthesis chemistry
- · Compatible with regular deprotection but requires mild deprotection with AMA at room temperature
- Available for 3', 5', and internal substitution
- More stable than BHQ-3

Item	Cat. No.	Pack	Price (\$)
5'-BBQ-650® Phosphoramidite	10-5934-95	50 μmole	160.00
	10-5934-90	100 μmole	305.00
	10-5934-02	0.25g	925.00
BBQ-650®-dT	10-5944-95	50 μmole	280.00
	10-5944-90	100 μmole	545.00
	10-5944-02	0.25g	925.00
3'-BBQ-650® CPG	20-5934-01	0.1g	190.00
	20-5934-10	1.0g	1500.00
1 μmole columns	20-5934-41	Pack of 4	300.00
0.2 μmole columns	20-5934-42	Pack of 4	80.00
10 μmole column (ABI)	20-5934-13	Pack of 1	575.00
15 μmole column (Expedite)	20-5934-14	Pack of 1	825.00

# **RHODAMINE (TAMRA) LABELING**

TAMRA CPG

Rhodamine derivatives are not sufficiently stable to survive conventional deprotection and these must be attached to amino-modified oligonucleotides using post-synthesis labeling techniques. Because Tetramethyl Rhodamine (TAMRA) is not base stable, the procedure to cleave and deprotect the labeled oligonucleotide must be carefully considered. Using the UltraMILD monomers and deprotection with potassium carbonate in methanol, TAMRA oligonucleotides can be fairly conveniently isolated. To streamline the preparation of TAMRA oligos, we offer 3'-TAMRA CPG for 3' labeling and TAMRA-dT for labeling within the sequence. We also offer TAMRA NHS ester for labeling amino-modified oligonucleotides.

Item	Cat. No.	Pack	Price (\$)
-1			
3'-TAMRA CPG	20-5910-01	0.1g	120.00
	20-5910-10	1.0g	995.00
1 μmole columns	20-5910-41	Pack of 4	200.00
0.2 μmole columns	20-5910-42	Pack of 4	120.00
3'-TAMRA PS	26-5910-01	0.1g	130.00
	26-5910-10	1.0g	1045.00
200 nmole columns (AB 3900)	26-5910-52	Pack of 10	300.00
40 nmole columns (AB 3900)	26-5910-55	Pack of 10	300.00
TAMRA-dT	10-1057-95	50 μmole	250.00
	10-1057-90	100 μmole	495.00
	10-1057-02	0.25g	975.00
TAMRA NHS Ester (Solution in anhydrous DMSO)	50-5910-66	60 μL	240.00

# SEE ALSO

UltraMILD monomers on page

# **OTHER INSTRUMENT TYPES**

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Monomers For Instrument type	Add		
Expedite MerMade	E M		
Columns For Instrument type	Add		
Expedite Applied Biosystems 3900 MerMade	E A M		
(Please inquire for availability of vials			

and columns for other instrument types.)

TAMRA NHS Ester

TAMRA-dT

3

**INTELLECTUAL PROPERTY** 

BlackBerry® Quencher technology: US Patent 7,879,986. The purchase

of BlackBerry® Quencher reagents

development purposes. They may

not be used for clinical or diagnostic

purposes and they may not be re-sold, distributed, or re-packaged without

prior agreement and consent of Berry

& Associates, Inc. Subsequent sale of products that are derived from

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and electronic catalogs, in commercial

advertisement, and in packages with

containers of such derivative products:

"BlackBerry is a trademark of Berry

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Associates, Inc."

includes a limited license to use these reagents exclusively for research and

# **ACRIDINE LABELING**

Acridine phosphoramidite is designed to produce an oligonucleotide containing acridine at any position in the molecule. Acridine CPG is used to label the 3'-terminus. Acridine is an effective intercalating agent.

Item	Cat. No.	Pack	Price (\$)
Acridine Phosphoramidite	10-1973-95	50 μmole	165.00
Actiume Phosphoramidite		'	
	10-1973-90	100 μmole	295.00
	10-1973-02	0.25g	675.00
3'-Acridine CPG	20-2973-01	0.1g	120.00
	20-2973-10	1.0g	995.00
1 μmole columns	20-2973-41	Pack of 4	200.00
0.2 μmole columns	20-2973-42	Pack of 4	120.00
10 μmole column (ABI)	20-2973-13	Pack of 1	300.00
15 μmole cloumn (Expedite)	20-2973-14	Pack of 1	450.00

# DNP LABELING

An analytical test based on detection of 2,4-dinitrophenyl (DNP) labeled oligonucleotides with anti-DNP antibodies has been proposed. We have chosen the branched triethylene glycol (TEG) spacer in our version of DNP phosphoramidite since it can be added once or several times to the 3′ or 5′ terminus.

Item	Catalog No.	Pack	Price(\$)
DNP-TEG Phosphoramidite	10-1985-95	50 μmole	165.00
	10-1985-90	100 μmole	295.00
	10-1985-02	0.25g	675.00

# 

# **CHOLESTEROL LABELING**

LABELING

**P**otential therapeutic oligonucleotides must permeate the cell membrane for optimal activity. The addition of lipophilic groups to an oligonucleotide would be expected to enhance cellular uptake/membrane permeation. The use of cholesteryl oligos and the consequent improvement in activity has been described. We have designed our Cholesteryl products with triethyleneglycol (TEG) spacers for maximum solubility.

Item	Catalog No.	Pack	Price(\$)
Cholesteryl-TEG Phosphoramidite	10-1975-95 10-1975-90 10-1975-02	50 μmole 100 μmole 0.25g	140.00 265.00 545.00
5'-Cholesteryl-TEG Phosphoramidite	10-1976-95 10-1976-90 10-1976-02	50 μmole 100 μmole 0.25g	95.00 175.00 525.00
3'-Cholesteryl-TEG CPG	20-2975-01 20-2975-10	0.1g 1.0g	85.00 700.00
1 μmole columns	20-2975-41	Pack of 4	140.00
0.2 μmole columns	20-2975-42	Pack of 4	84.00
10 μmole column (ABI)	20-2975-13	Pack of 1	210.00
15 μmole column (Expedite)	20-2975-14	Pack of 1	315.00

SEE ALSO

Spermine on page 4

# **TOCOPHEROL LABELING**

**V**itamin E is both lipophilic and non-toxic even at high doses so would be an excellent candidate as a lipophilic carrier for oligonucleotides. Therefore, as an addition to our cholesteryl product line, we offer simple  $\alpha$ -tocopheryl (vitamin E) labeling. Totally synthetic  $\alpha$ -tocopherol is racemic at its three chiral centers and is used to prepare this product.

Item	Catalog No.	Pack	Price(\$)
lpha-Tocopherol-TEG Phosphoramidite	10-1977-95	50 μmole	160.00
	10-1977-90	100 μmole	300.00
	10-1977-02	0.25g	575.00

# **STEARYL LABELING**

**W**e now offer a simple C18 lipid as an economical and effective carrier molecule. We envisage that the 5'-stearyl group will become a favored lipophilic carrier for experimentation with synthetic oligonucleotides.

Item	Catalog No.	Pack	Price(\$)
5' - Stearyl Phosphoramidite	10-1979-90 10-1979-02	100 μmole 0.25g	45.00 180.00
O—CNEt O—P—N(iPr) <sub>2</sub> DMTO O O O			
Cholestery  O—CNEI P—N(Pr) <sub>2</sub> O—O NH  DMTO.	I-TEG		
5'-Cholesteryl-TEG  O-CNEI  ONE  ONE  ONE  ONE  ONE  ONE  ONE	3'-C	holesteryl-TEG CPG  O—P—N(Pr) <sub>2</sub> O—CNEt	

OTHER INSTRUMENT TYPES

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Add

Monomers

MerMade

Expedite

MerMade

Applied Biosystems 3900

(Please inquire for availability of vials and columns for other instrument types.)

A directed approach to the delivery of therapeutic oligonucleotides specifically to the liver has been to target the asialoglycoprotein receptor (ASGPR) using a suitable glycoconjugate. Indeed, ASGPR is the ideal target for delivery of therapeutic oligonucleotides to the liver since it combines tissue specificity, high expression levels and rapid internalization and turnover. The use of oligonucleotide glycoconjugates has led to significant advances in therapeutic delivery as evidenced by the work of Alnylam Pharmaceuticals and Ionis Pharmaceuticals using multivalent N-acetylgalactosamine (GalNAc) oligonucleotide conjugates.

Glen Research is delighted to introduce a GalNAc modification strategy using a monomeric GalNAc support and the equivalent GalNAc phosphoramidite. Our experimental work has shown that these products are fully compatible with regular oligonucleotide synthesis and deprotection. Oligonucleotides containing GalNAc can be deprotected using standard procedures during which the acetyl protecting groups on GalNAc are removed. We have demonstrated that 5'-GalNAc C3 phosphoramidite can be used to prepare oligonucleotides with multiple consecutive GalNAc additions at the 5' terminus. Glen Research offers these GalNAc C3 products under an agreement with AM Chemicals LLC.

· · · · · · · · · · · · · · · · · · ·			
Item	Catalog No.	Pack	Price(\$)
5'-GalNAc C3 Phosphoramidite	10-1974-95	50 μmole	137.50
	10-1974-90 10-1974-02	100 μmole 0.25g	255.00 500.00
GalNAc C3 CPG	20-2974-01	0.1g	40.00
	20-2974-10	1.0g	320.00
1 μmole columns	20-2974-41	Pack of 4	100.00
0.2 μmole columns	20-2974-42	Pack of 4	60.00
10 μmole column (ABI)	20-2974-13	Pack of 1	180.00
15 μmole column (Expedite)	20-2974-14	Pack of 1	280.00

# CDPI<sub>3</sub> MGB™ LABELING

The tripeptide of dihydropyrroloindole-carboxylate (CDPI<sub>3</sub>) is a minor groove binding (MGB) moiety derived from the natural product CC-1065 with strong DNA binding properties. Synthetic oligonucleotides with covalently-attached CDPI<sub>3</sub> have enhanced DNA affinity and have improved the hybridization properties of sequence-specific DNA probes. Short CDPI<sub>3</sub>-oligonucleotides hybridize with single-stranded DNA to give more stable DNA duplexes than unmodified ODNs of similar length. CDPI<sub>3</sub> MGB-oligonucleotide conjugates have been found to be useful in the following applications:

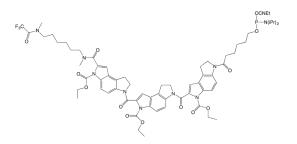
- Arrest of primer extension and PCR blockers
- Short and fluorogenic PCR primers
- Real-time PCR probes
- miRNA Inhibitors

The simplest approach to MGB probe design is to use an MGB support, add a quencher molecule as the first addition and complete the synthesis with a 5′-fluorophore. Alternatively, a fluorophore support could be used with the 5′ terminus containing a quencher molecule followed by a final MGB addition at the 5′ terminus. Glen Research offers 5′-CDPI<sub>3</sub> MGB™ Phosphoramidite and 3′-CDPI<sub>3</sub> MGB™ CPG.

5′-CDPl<sub>3</sub> MGB phosphoramidite was found to be hydrophobic enough that it required 10% THF in ACN to go completely into solution at a 0.1 M concentration and required a 3 minute coupling time. Deprotection can be carried out in EtOH/ NH4OH 1:3 (v/v) 17 hr at 55 °C and CDPl<sub>3</sub> MGB is compatible with GlenPak™ purification.

With the CDPI<sub>3</sub> MGB CPG, optimal results are obtained if UltraMild monomers and Cap A are used during synthesis along with 0.5 M CSO oxidizer. However, the use of standard monomers with iodine oxidation followed by deprotection with EtOH/NH4OH 1:3 (v/v) for 17 hr at 55 °C will give acceptable results.

Item	Catalog No.	Pack	Price(\$)
5'-CDPI, MGB™ Phosphoramidite	10-5924-95	50 μmole	705.00
3 CDT 13 WIGD THOSPHORAINIAICE	10-5924-90	100 μmole	1390.00
	10-5924-02	0.25g	2600.00
CDPI, MGB™ CPG	20-5924-01	0.1g	215.00
CDI 13 INIOD CI O	20-5924-10	1.0g	1800.00
1 μmole columns	20-5924-41	Pack of 4	325.00
0.2 μmole columns	20-5924-42	Pack of 4	165.00
10 μmole column (ABI)	20-5924-13	Pack of 1	925.00
15 μmole column (Expedite)	20-5924-14	Pack of 1	1395.00



5'-CDPI, MGB™ Phosphoramidite

 $\mathbf{CDPI_3} \; \mathbf{MGB^{\mathsf{TM}}} \; \mathbf{CPG}$ 

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**OTHER INSTRUMENT TYPES** 

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modifiers are packaged in septum-

capped vials suitable for ABI and other

instruments. If you would like another

type of vial/column add the following to

the end of the catalog number.

Monomers

MerMade

Columns

Expedite

MerMade

Applied Biosystems 3900

(Please inquire for availability of vials

and columns for other instrument types.)

**P**soralen C2 at the 5'-terminus of an oligonucleotide serves effectively as a cross-linking reagent in double-stranded oligonucleotides. The 6 atom spacer arm of Psoralen C6 allows cross-linking with a triplex oligonucleotide strand. Click Chemistry with psoralen azide and one of our many nucleosidic and non-nucleosidic alkyne derivatives has the potential to generate a variety of practical cross-linkers. The well known reversible cross-linking behavior of psoralen with an adjacent thymidine residue could be very useful.

Item	Cat. No.	Pack	Price (\$)
Psoralen C2 Phosphoramidite	10-1982-90	100 μmole	195.00
	10-1982-02	0.25g	495.00
Psoralen C6 Phosphoramidite	10-1983-90	100 μmole	195.00
	10-1983-02	0.25g	495.00
Psoralen Azide	50-2009-92	25 μmole	115.00
	50-2009-90	100 μmole	350.00

# EDTA LABELING

EDTA-C2-dT phosphoramidite contains the triethyl ester of EDTA which allows sequence-specific cleavage of single- and double-stranded DNA and RNA. The cleavage reaction is only initiated once Fe(II) and dithiothreitol are added and so is readily controlled. Coupling of EDTA-dT is normal but cleavage and deprotection should be carried out with sodium hydroxide in aqueous methanol (0.4M NaOH in methanol/water 4:1) overnight at room temperature.

Item	Cat. No.	Pack	Price (\$)
EDTA-C2-dT-CE Phosphoramidite	10-1059-95	50 μmole	250.00
	10-1059-90	100 μmole	495.00
	10-1059-02	0.25g	975.00

EDTA-C2-d1

# **FERROCENE LABELING**

**W**ith an excellent stability profile, ferrocene has always attracted considerable interest for DNA labeling to generate probes for electrochemical detection. Based on our Amino-Modifier C6-dT structure, Ferrocene-dT is easily added to oligonucleotides with no disruption of regular hybridization behavior. Multiple incorporations into an oligonucleotide probe are also simply achieved. Oligonucleotides are deprotected using standard techniques. Ferrocene oligonucleotides should be stored under Argon and aqueous solutions should be degassed immediately.

Item	Cat. No.	Pack	Price (\$)
Ferrocene-dT-CE Phosphoramidite	10-1576-95 10-1576-90	50 μmole 100 μmole	170.00 330.00
	10-1576-02	0.25g	670.00

# METHYLENE BLUE LABELING

Methylene Blue, which belongs to the phenothiazine family of dyes, is a unique dye with a variety of useful properties. Despite its high extinction coefficient in the visible region (81,000 L/mol.cm), it is weakly fluorescent due to its high rate of intersystem crossing from the S<sub>1</sub> excited state to the T<sub>1</sub> triplet state. This property makes it an excellent photosensitizer, and it has been used extensively to produce highly reactive singlet oxygen. Methylene blue has the ability to both intercalate in duplex DNA, preferring G:C over T:A base pairs, and can act as an electrochemical redox probe. Methylene blue has also been shown to be unmatched in performance as a redox-active reporter for electrochemical biosensors.

Earlier, we introduced Methylene Blue C3 Phosphoramidite but this product proved to have quite limited stability and has been discontinued. As an alternative option, we introduced Methylene Blue NHS Ester to allow researchers to label amino-modified oligonucleotides with this interesting dye. With the encouragement and technical expertise of Carole Chaix and her colleagues at the University of Lyon, we decided to prepare an alternative structure that seemed to have a much superior stability profile - Methylene Blue II Phosphoramidite. Fortunately, this structure did indeed prove more stable and we are now able to offer again a Methylene Blue Phosphoramidite.

Item	Cat. No.	Pack	Price (\$)
Methylene Blue NHS Ester (Dissolve 5.4mg in 60μL of DMSO)	50-1960-23	5.4mg	540.00
Methylene Blue II Phosphoramidite	10-5961-95	50 μmole	310.00
	10-5961-90	100 μmole	595.00
	10-5961-02	0.25g	1500.00

# **INTELLECTUAL PROPERTY**

Methylene Blue II is covered under patent applications FR12 51739 and PCT/FR2013/050356 and is sold under license from the University of Lyon.

OTHER INSTRUMENT TYPES

All minor bases, RNA products and modifiers are packaged in septum-capped vials suitable for ABI and other instruments. If you would like another type of vial/column add the following to the end of the catalog number.

Monomers

MerMade

Columns

Expedite

MerMade

Applied Biosystems 3900

(Please inquire for availability of vials

and columns for other instrument types.)

fax: 253-833-8127

Dmaeda@syntrixbio.com

**2**,2'-Dipicolylamine Phosphoramidite has been discontinued This product was manufactured and developed by Syntrix Biosystems Inc. For further information, please contact:

Dean Y. Maeda, Ph.D., M.B.A. Director, Chemistry and Preclinical Development Syntrix Biosystems 215 Clay St NW Ste B5 Auburn, WA 98001 tel: 253-833-8009 ext. 23

# LABELING WITH POLYAROMATIC HYDROCARBONS

**P**yrene and perylene are fluorescent polycyclic aromatic hydrocarbons that have the ability to form 'excited state dimers' known as excimers. This unstructured, long-wavelength emission arises from the formation of a charge-transfer complex between the excited state and the ground state of two fluorescent molecules. In Pyrene-dU and perylene-dU, the hydrocarbon is attached at the 5 position of deoxyuridine through a triple bond and is electronically coupled to the deoxyuridine base. This electronic coupling of the base and the hydrocarbon makes the fluorescence sensitive to the base pairing of the dU portion of the molecule, allowing the discrimination between perfect and one base mismatched targets.

Item	Cat. No.	Pack	Price (\$)
Pyrene-dU-CE Phosphoramidite	10-1590-95	50 μmole	105.00
,,	10-1590-90	100 μmole	210.00
	10-1590-02	0.25g	550.00
Perylene-dU-CE Phosphoramidite	10-1591-95	50 μmole	150.00
renyiene do ez mosphoramate	10-1591-90	100 μmole	300.00
	10-1591-02	0.25g	720.00

# DMTO OPN(iPr)<sub>2</sub> OCNEt 2,2'-Dipicolylamine DMTO OPN(iPr)<sub>2</sub> OCNEt DMTO OPN(iPr)<sub>2</sub> OCNEt DMTO OPN(iPr)<sub>2</sub> OCNEt

# **PUROMYCIN CPG**

One of the most challenging requirements associated with combinatorial chemistry is the recovery of sequence information of the oligonucleotide or peptide selected by the screening assay. A method 1 has been developed to generate a fusion product between mRNA and the polypeptide it encodes using *in vitro* translation of synthetic RNAs 3'-labeled with puromycin, an antibiotic that mimics transfer RNA. Puromycin binds in the ribosome's A site, forms a peptide bond with the growing peptide chain, and blocks further peptide elongation. By linking puromycin to mRNA, a peptide-RNA fusion product results from the translation of the message linking the encoding mRNA with its peptide product.

Item	Catalog No.	Pack	Price(\$)
Puromycin CPG	20-4040-01	0.1g	120.00
,	20-4040-10	1.0g	995.00
1 μmole columns	20-4140-41	Pack of 4	200.00
0.2 μmole columns	20-4140-42	Pack of 4	120.00
10 μmole column (ABI)	20-4140-13	Pack of 1	360.00
15 μmole columns (Expedite)	20-4140-14	Pack of 1	540.00

# **QUENCHED AUTOLIGATION (QUAL) PROBES**

**Puromycin CPG** 

QUAL probes¹ consist of two oligonucleotides, the first containing a nucleophilic group at the 3′-terminus, while the second has an electrophilic group at the 5′-terminus. When the probe pair finds the target, the oligos line up with the 3′-terminus of the first directly adjacent to the 5′-terminus of the second. An autoligation reaction then takes place to combine the two oligos into a single probe. As usual, the 3′ nucleophilic group is the 3-thiophosphate, easily prepared using 3′-phosphate CPG with a sulfurizing step in the first cycle. In this case, the electrophilic group is a 5′-dabsyl group, which is an excellent leaving group as well as a fine quencher of fluorescence. The second oligo, therefore, contains a fluorophore which is quenched by the dabsyl group. A popular choice for fluorophore is fluorescein-dT but it is easy to imagine that a variety of fluorophores could be attached to any of the commercially available amino-modified nucleoside phosphoramidites.

Item	Catalog No.	Pack	Price(\$)
5'-Dabsyl-dT-CE Phosphoramidite	10-1532-90 10-1532-02	100 μmole 0.25g	250.00 775.00

# REFERENCE

(1) R.W. Roberts and J.W. Szostak, *Proc. Natl. Acad. Sci. USA*, 1997, **94**, 12297-

# REFERENCE

(1) S. Sando and E.T. Kool, *J Amer Chem Soc*, 2002, **124**, 2096-2097.

# SEE ALSO

3'-Phosphate CPG on page 12 Sulfurizing Reagent on page 4 Fluorescein-dT on page 33

# 

**OTHER INSTRUMENT TYPES** 

All minor bases, RNA products and

modifiers are packaged in septum-

capped vials suitable for ABI and other instruments. If you would like another

type of vial/column add the following to the end of the catalog number.

(Please inquire for availability of vials

and columns for other instrument types.)

Absorbance Emission Excimer

472nm

490nm

Determined

Monomers

Expedite MerMade

Columns

MerMade

Pyrene-dU Pervlene-dU

Applied Biosystems 3900

**FLUORESCENT DYES** 

402nm

473nm

# LABELING FOR PHOTO-REGULATION OF OLIGONUCLEOTIDES

**P**hoto-control, the use of ultraviolet or visible light to control a reaction, has a number of advantages over other external stimuli:

- Light does not introduce contaminants into the reaction system,
- Excitation wavelength can be controlled through the design of the photo-responsive molecule, and
- It is now straightforward to control irradiation time and/or local excitation.

When a photo-responsive molecule is directly attached to DNA as a receptor, photo-regulation of the bioprocess regulated by that DNA molecule could, in principle, be achieved. Such photo-responsive DNA could also be used as a switch in a DNA-based nano-machine. Professor Hiroyuki Asanuma and his group at the department of Molecular Design and Engineering of the Graduate School of Engineering of the Nagoya University (Japan) have developed an efficient method to achieve this goal. They have attached azobenzene to DNA and made it photo-responsive<sup>1,2</sup>. Azobenzene is a typical photo-responsive molecule that isomerizes from its planar *trans*-form to the non-planar *cis*-form after UV-light irradiation with a wavelength between 300 nm and 400 nm ( $\lambda_{max}$  is around 330 nm). Interestingly, the system reverts from the *cis*-form to the *trans*-form after further irradiation with visible light (wavelength over 400 nm). This process is completely reversible, and the azobenzene group does not decompose or induce undesirable side reactions even on repeated *trans-cis* isomerization. By introducing azobenzenes into DNA through D-threoninol as a linker, Asanuma and co-workers succeeded in achieving photo-regulation of:

- Formation and dissociation of a DNA duplex<sup>3,4</sup> and
- Transcription by T7-RNA polymerase reaction<sup>5,6,7</sup>.

Item	Catalog No.	Pack	Price(\$)
Azobenzene Phosphoramidite	10-5800-95	50 μmole	105.00
	10-5800-90	100 μmole	200.00
	10-5800-02	0.25g	550.00

# REFERENCES

- (1) H. Asanuma, et al., *Angew Chem Int Ed*, 2001, **40**, 2671-2673.
- (2) T. Takarada, et al., *Chem Lett.*, 2001, **30**, 732.
- (3) H. Asanuma, X.G. Liang, T. Yoshida, and M. Komiyama, *Chembiochem*, 2001, 2, 39-44.
- (4) H. Asanuma, D. Matsunaga, and M. Komiyama, NUCLEIC ACIDS SYMP SER (OXF), 2005, 49, 35.
- (5) H. Asanuma, et al., *Chembiochem*, 2002, **3**, 786.

2007, 2, 203-212.

Monomers

- (6) M. Liu, H. Asanuma, and M. Komiyama,
   J. Amer. Chem. Soc., 2006, 128, 1009.
   (7) H. Asanuma, et al., Nature Protocols,
- OTHER INSTRUMENT TYPES

  All minor bases, RNA products and modifiers are packaged in septum-

capped vials suitable for ABI and other instruments. If you would like another type of vial/column add the following to the end of the catalog number.

For Instrument type	Add
Expedite	E
MerMade	M
Columns For Instrument type	Add
Expedite	E
Applied Biosystems 3900	A
MerMade	M

(Please inquire for availability of vials and columns for other instrument types.)

рмто	O N H		-N_N
H <sub>3</sub> C	'n		
	P—N(il	Pr) <sub>2</sub>	
	O-CN	Et	

# **Azobenzene Phosphoramidite**

# LABELLING WITH ULTRAFAST PHOTO CROSS-LINKER

When 3-cyanovinylcarbazole nucleoside (CNVK) is incorporated into an oligonucleotide, very rapid photo cross-linking to the complementary strand can be induced at one wavelength and rapid reversal of the cross-link is possible at a second wavelength. Neither wavelength has the potential to cause significant DNA damage. Irradiation of a duplex containing a single incorporation of CNVK at 366nm led to 100% cross-linking to thymine base in 1 second, although complete cross-linking to cytosine takes 25 seconds.¹ A 30 second irradiation time should cover all situations. In addition, it was demonstrated that the purine bases were unreactive to cross-linking, allowing differentiation between pyrimidines and purines at the target site. The authors also determined the effect of sequence contexts around the CNVK site and demonstrated that the identity of bases on either side of the cross-linking site has little effect on the reaction. Once cross-linked, the UV melting temperature of the duplex was raised by around 30 °C relative to the duplex before irradiation. Complete reversal of the cross-link takes place at 312nm in 3 minutes. This facile reversal reaction is, therefore, accomplished with no damage to normal DNA.

In a later publication, a further application of this cross-linking technique was investigated.<sup>2</sup> When CNVK was cross-linked with a dC residue in duplex DNA, heating at 90°C for 3.5 hours led to deamination of the cytosine base to form uracil in the complementary strand. Reversal of the cross-link at 312nm led to a DNA strand in which dC had been converted to dU. The authors showed that this transformation is specific for the dC residue opposite the CNVK and any further adjacent dC residues are unaffected. Similarly, the authors have shown that CNVK can be cross-linked to an adjacent RNA strand.<sup>3</sup>

Item	Cat. No.	Pack	Price (\$)
3-Cyanovinylcarbazole Phosphoramidite (CNVK)	10-4960-95	50 μmole	200.00
	10-4960-90	100 μmole	390.00
	10-4960-02	0.25g	1125.00

# REFERENCES

- (1) Y. Yoshimura, and K. Fujimoto, *Org Lett*, 2008, **10**, 3227-30.
- (2) K. Fujimoto, K. Konishi-Hiratsuka, T. Sakamoto, and Y. Yoshimura, *ChemBioChem*, 2010, **11**, 1661-4.
- (3) Y. Yoshimura, T. Ohtake, H. Okada, and K. Fujimoto, *ChemBioChem*, 2009, **10**, 1473-6.

Index	5'-Biotin Phosphoramidite 30 Biotin-dT 30
A	Biotin Phosphoramidite 29
^	BiotinTEG Azide 22 BiotinTEG Phosphoramidite 29
A Amino-Modifier C6 dA 7	DesthiobiotinTEG Azide 22 DesthiobiotinTEG-CPG 31
Abasic Site 14	DesthiobiotinTEG Phosphoramidite 30
Acridine Labelling	PC Biotin Phosphoramidite 16, 30
3'-Acridine CPG 44	Protected BiotinLC Serinol Phosphoramidite 25, 29
Acridine Phosphoramidite 44	Protected Biotin Serinol Phosphoramidite 24, 29
Activator (Powder)	BlackBerry® Quencher
5-Ethylthio-1H-tetrazole 40, 42	3'-BBQ-650® CPG 42
Aldehyde Modifier	5'-BBQ-650® Phosphoramidite 42
5'-Aldehyde-Modifier C2 Phosphoramidite 13	BBQ-650®-dT 42
Formylindole CE Phosphoramidite 13	Black Hole Quencher™ Dyes
Amino-Modifiers	3'-BHQ-1 CPG 41 3'-BHQ-2 CPG 41
3'-Amino-Modifier C6 dC CPG 11	3'-BHQ-2 CPG 41 3'-BHQ-3 CPG 41, 42
3'-Amino-Modifier C6 dT CPG 11	5'-BHQ-1 Phosphoramidite 40
3'-Amino-Modifier Serinol CPG 9	5'-BHQ-2 Phosphoramidite 40, 42
3'-PT-Amino-Modifier C3 CPG 9	BHQ-1-dT 40, 42
3'-PT-Amino-Modifier C6 CPG 9 3'-PT-Amino-Modifier C6 PS 9	BHQ-2-dT 40
5'-Amino-Modifier 5 4	Brancher Phosphoramidite
5'-Amino-Modifier C3-TFA 4, 5	dC Brancher Phosphoramidite 15
5'-Amino-Modifier C6 4, 5	Bromohexyl Phosphoramidite 19
5'-Amino-Modifier C6-PDA 5	
5'-Amino-Modifier C6-TFA 4, 5	С
5'-Amino-Modifier C12 4	С
5'-Amino-Modifier C12-PDA 5 5'-Amino-Modifier TEG 4	3'-Amino-Modifier C6 dC CPG 11
5'-Amino-Modifier-TEG-PDA 5	Amino-Modifier C6 dC 7
5'-DMS(O)MT-Amino-Modifier C6 4	C8-Alkyne-dC-CE Phosphoramidite 17, 20
Amino-Modifier C2 dT 7	C8-TIPS-Alkyne-dC-CE Phosphoramidite 17
Amino-Modifier C6 dA 7	C8-TMS-Alkyne-dC-CE Phosphoramidite 17
Amino-Modifier C6 dC 7	dC Brancher Phosphoramidite 15
Amino-Modifier C6 dT 7 Amino-Modifier Serinol Phosphoramidite 8	Carboxy-Modifiers
Fmoc-Amino-Modifier C6 dT 8	5'-Carboxy-Modifier C5 6 5'-Carboxy-Modifier C10 6
N2-Amino-Modifier C6 dG 7	Carboxy-dT 7
PC Amino-Modifier Phosphoramidite 6, 16, 25	Chelates
AminoOxy-Modifier	EDTA-C2-dT-CE Phosphoramidite 48
5'-AminoOxy-Modifier 11 6	Chemical Phosphorylation 12
AquaPhluor® 593	• •
5'-AquaPhluor® 593 Phosphoramidite 38	Cholesterol Labelling 3'-Cholesteryl-TEG CPG 45
AquaPhluor® 593 CPG 39	5'-Cholesteryl-TEG CPG 45 5'-Cholesteryl-TEG Phosphoramidite 45
Azidobutyrate NHS Ester 19	Cholesteryl-TEG Phosphoramidite 5, 45
Azobenzene	Click Chemistry 17
Azobenzene Phosphoramidite 52	1,2,3-triazoles 17
_	1-Ethynyl-dSpacer CE Phosphoramidite 20
В	3'-Alkyne-Modifier Serinol CPG 10, 19, 27
Biotin Labelling	5'-Bromohexyl Phosphoramidite 19
3'-BiotinTEG CPG 31	5-Ethynyl-dU-CE Phosphoramidite 18 5'-Hexynyl Phosphoramidite 19

5'-I-dT-CE Phosphoramidite 19

Alkyne-Modifier Serinol Phosphoramidite 19, 25

Alkyne-NHS Ester 19 Azides 19 Azides 19 Azides Oligo-Click-M-B baseclick Oligo-Click-M-B baseclick Oligo-Click-M-F baseclick Oligo-Click-M-R baseclick Oligo-Click-M-T C8-Alkyne-dC-CE Phosph C8-Alkyne-dT-CE Phosph C8-TIPS-Alkyne-dT-CE Ph C8-TIPS-Alkyne-dT-CE Ph C8-TIPS-Alkyne-dT-CE Ph C8-TIMS-Alkyne-dT-CE Ph C8-TMS-Alkyne-dT-CE Ph C8-TMS-Alkyne-dT-CE Ph C8-TMS-Alkyne-dT-CE Ph C9pper-free Click Chemis THPTA Ligand 18 TIPS-5-Ethynyl-dU-CE Ph Copper-free Click Che 5'-DBCO-TEG Phosphoran DBCO-dT-CE Phosphoran DBCO-sulfo-NHS Ester 2: Cross-linking 23, 48, Cyanine Labelling Cyanine 3.5 Phosphoram Cyanine 3 CPG 37 Cyanine 3 Phosphoramid Cyanine 5 CPG 37 Cyanine 5 Phosphoramid Disulfo-Cyanine 7 Azide: Cyanovinylcarbazole 3-Cyanovinylcarbazole Pf CNVK 53
Cyclooctatetraene COT Serinol Phosphoram
D
Dabcyl Labelling 3'-Dabcyl CPG 28 3'-Dabcyl PS 28 3'-Dabsyl CPG 28, 40, 42 5'-Dabcyl Phosphoramidi Dabcyl-dT 28  DBCO
5'-DBCO-TEG Phosphoral DBCO-dT-CE Phosphoram DBCO-Serinol Phosphora DBCO-sulfo-NHS Ester 2:
Dendrimers Asymmetric Doubler (LEV Long Trebler Phosphoran Symmetric Doubler Phos Trebler Phosphoramidite
Desthiobiotin

```
Distributors 3
                                                        Dithiol
                      19
                                                         3'-Dithiol Serinol CPG 10
                     Biotin 20
                                                         Dithiol Serinol Phosphoramidite 6
                     Fluorescein 20
                                                        DNP Labelling
                     Reload 20
                                                         DNP-TEG Phosphoramidite 44
                     TAMRA 20
                     horamidite 17, 20
                                                        Doubler Phosphoramidite
                     noramidite 17
                                                          Symmetric 15
                     hosphoramidite 17, 18
                     hosphoramidite 18
                     hosphoramidite 17, 18
                     hosphoramidite 18
                                                        Eclipse® Quencher
                     istry 19
                                                         Eclipse® Quencher CPG 39
                                                          Eclipse® Quencher Phosphoramidite 38
                     nosphoramidite 18
                                                        EDTA-dT
                     emistry 20
                                                         EDTA-C2-dT-CE Phosphoramidite 48
                     amidite 21
                                                        EdU
                     midite 21
                                                         5-Ethynyl-dU-CE Phosphoramidite 18
                                                         TIPS-5-Ethynyl-dU-CE Phosphoramidite 18
                     , 53
                                                       ELITechGroup Dyes and Quencher 38
                                                        Excimers 50
                     midite 36
                     idite 36
                     midite 36
                                                       FAM
                                                          3'-(6-FAM) CPG 34
                     idite 36
                                                          3'-(6-FAM) PS 34
                     23, 37
                                                          6-FAM 32
                                                          6-FAM-TEG Azide 22
                     hosphoramidite 53
                                                          Dipivaloyl 6-FAM-TEG Azide 22
                                                        Ferrocene Labelling
                                                          Ferrocene-dT-CE Phosphoramidite 49
                     midite 27
                                                        Fluorescein Labelling
                                                         3'-(6-FAM) CPG 34
                                                          3'-(6-FAM) PS 34
                                                          3'-(6-Fluorescein) CPG 34, 37
                                                          3'-6-Fluorescein Serinol CPG 26, 34
                                                          3'-Fluorescein CPG 34
                                                          3'-Fluorescein-dT CPG 34
                                                          5'-Dichloro-dimethoxy-Fluorescein 32
                     dite 28
                                                          5'-Fluorescein Phosphoramidite 32
                                                          5'-Hexachloro-Fluorescein 32
                                                          5'-Tetrachloro-Fluorescein 32
                     amidite 21
                                                          6-Fluorescein Phosphoramidite 33
                     midite 21
                                                          6-Fluorescein Serinol Phosphoramidite 24, 33
                     amidite 21
                                                          Dichloro-diphenyl-fluorescein 35
                                                          Fluorescein-dT Phosphoramidite 33
                                                          SIMA (HEX) 35
                     EV) Phosphoramidite 15
                                                        Formylindole CE Phosphoramidite 13
                     midite 15
                     sphoramidite 15
                                                        G
                      15
                                                          N2-Amino-Modifier C6 dG 7
DesthiobiotinTEG-CPG 31
DesthiobiotinTEG Phosphoramidite 30
                                                        GalNAc
```

5'-GalNAc C3 Phosphoramidite 46

3'-BiotinTEG PS 31

3'-Protected BiotinLC Serinol CPG 26, 31

3'-Protected Biotin Serinol CPG 25, 31

GalNAc C3 CPG 46	Phthalimide (PT)
Glyceryl CPG 10	3'-PT-Amino-Modifier C3 CPG 9
Gold	3'-PT-Amino-Modifier C6 CPG 9
Conjugation to gold surfaces 24	3'-PT-Amino-Modifier C6 PS 9
	Polystyrene Supports
Н	3'-(6-FAM) PS 34 3'-BiotinTEG PS 31
HEX 32	3'-Dabcyl PS 28
6-HEX Azide 22, 23	3'-Phosphate PS 12
Hexynyl Phosphoramidite 19	3'-PT-Amino-Modifier C6 PS 9
Trexytty Triosprioral marce 15	3'-TAMRA PS 43
1	Psoralen Labelling
나	Psoralen Azide 23, 48
I-dT 5'-I-dT-CE Phosphoramidite 19	Psoralen C2 Phosphoramidite 48 Psoralen C6 Phosphoramidite 48
	Puromycin
Introduction 1, 2, 3	Puromycin CPG 51
J	Pyrene
	Pyrene-dU-CE Phosphoramidite 50
JOE	
5'-Dichloro-dimethoxy-Fluorescein Phosphoramidite II 32	Q
M	Quenched Autoligation (QUAL) Probes 51
A A 1 2 2 1 A A 100	5'-Dabsyl-dT-CE Phosphoramidite 51
Maleimide-Modifier 5'-Maleimide-Modifier Phosphoramidite 6	_
·	R
Methylene Blue Methylene Blue II Phosphoramidite 49	Redmond Red®
Methylene Blue NHS Ester 49	Redmond Red® CPG 39
MGB	Redmond Red® Phosphoramidite 38
5'-CDPI3 MGB™ Phosphoramidite 47	Rhodamine 43
CDPI3 MGB™ CPG 47	•
Minor Groove Binder (MGB) 47	S
Modifiers 4, 5, 8	Sequence Modifiers 8
	Serinol Backbone
Р	3'-6-Fluorescein Serinol CPG 26, 34
Perylene	3'-Alkyne-Modifier Serinol CPG 10, 19, 27
Perylene-dU-CE Phosphoramidite 50	3'-Amino-Modifier Serinol CPG 9, 26
Phosphorylation	3'-Dithiol Serinol CPG 10, 27 3'-Protected BiotinLC Serinol CPG 26, 31
3'-CPR II CPG 12	3'-Protected Biotinic Serinol CPG 25, 31
3'-Phosphate CPG 12	6-Fluorescein Serinol Phosphoramidite 24, 33
3'-Phosphate CPG - High Load 12	Alkyne-Modifier Serinol Phosphoramidite 19, 26
3'-Phosphate PS 12 Chemical Phosphorylation Reagent 12	Amino-Modifier Serinol Phosphoramidite 8, 25
Chemical Phosphorylation Reagent II 12	COT Serinol Phosphoramidite 27
CPR II 12	Dithiol Serinol Phosphoramidite 6, 25 Protected BiotinLC Serinol Phosphoramidite 25, 30
Solid CPR II 12	Protected Biotines Serinol Phosphoramidite 24, 30
Photocleavable Monomers	SIMA
PC Amino-Modifier Phosphoramidite 6, 16, 25	SIMA (HEX)-dT Phosphoramidite 35
PC Biotin Phosphoramidite 16, 30 PC Linker Phosphoramidite 16	SIMA (HEX) Phosphoramidite 35
PC Linker Phosphoramidite 16 PC Spacer Phosphoramidite 14, 16	Spacer Modifiers
Photo cross-linking 53	1-Ethynyl-dSpacer CE Phosphoramidite 20
Photo-responsive DNA	3'-Spacer C3 CPG 14 dSpacer CE Phosphoramidite 14
I HOLO-IESPONSIVE DIVA	aspacei el riiospiioramilatte 14

PC Spacer Phosphoramidite 14, 16

Spacer Spacer Spacer Spacer Spin Lal TEMPC TEMPC Stearyl 5'- Stea
Т
т
T 3'-Amir 3'-Fluo 5'-Dabs 5'-I-dro- Amino- C8-Alky C8-TIPS C8-TM! DBCO-( EDTA-C Ferroce Fluores S-Bz-Th TAMRA 3'-TAM 3'-TAM 3'-TAM
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TEMPO TEMPO TEMPO
Termina
3'-Spac
Terminu TET 32 6-TET A
Thiol-W 3'-Dithi 3'-Thio 5'-Thio Dithiol S-Bz-Th Thiol-M Tocoph a-Tocop
Trebler Trebler

```
r C12 CE Phosphoramidite 14
        r Phosphoramidite 9 14
        r Phosphoramidite 18 14
        r Phosphoramidite C3 14
        abels
        O Azide 23
        O-TEG Azide 23
        l Labelling 45
        earyl Phosphoramidite 45
        ino-Modifier C6 dT CPG 11
        orescein-dT CPG 34
        bsyl-dT 51
        T-CE Phosphoramidite 19
        o-Modifier C2 dT 7
        -Modifier C6 dT 7
        kyne-dT-CE Phosphoramidite 17
        PS-Alkyne-dT-CE Phosphoramidite 18
        AS-Alkyne-dT-CE Phosphoramidite 18
        -dT-CE Phosphoramidite 21
        C2-dT-CE Phosphoramidite 48
        cene-dT-CE Phosphoramidite 49
        scein-dT 33
        hiol-Modifier C6-dT 8
        A-dT 43
        A Labelling
        MRA CPG 43
        MRA PS 43
        A-dT 43
         NHS Ester 43
        O Azide 23
        O-TEG Azide 23
        nation, 3'
        cer C3 CPG 14
        nus Modifiers 4
        Azide 22, 23
        Modifiers
        hiol Serinol CPG 10
        ol-Modifier 6 S-S CPG 10
        ol-Modifier C6 6
        ol Serinol Phosphoramidite 6
        Thiol-Modifier C6-dT 8
        Modifier C6 S-S 6
        herol
        opherol-TEG Phosphoramidite 45
        r Phosphoramidite
         15
U
 5-Ethynyl-dU-CE Phosphoramidite 18
```

TIPS-5-Ethynyl-dU-CE Phosphoramidite 18

# V

Vitamin E 45

# Yakima Yellow®

Yakima Yellow® CPG 39 Yakima Yellow® Phosphoramidite 38

Azobenzene Phosphoramidite 52

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# Glen Research Corporation

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# **PATENTS**

As a research-oriented company, we realize the desirability of patents to cover original research and it is our policy to avoid infringing any approved patents. Accordingly, it is possible that some of our products may have to be withdrawn or adjusted in price as patents are approved and issued.

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