

Catalog of Innovations. 2016

- Nucleic acid analogs
- Synthetic Biology
- Diagnostics
- DNA sequencing
- Bioinformatics
- SNP detection
- Polymerases

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Reversible terminators in primer extension

Firebird has introduced the 3'-ONH₂ reversible terminator to replace bulky 3'-OCH₂N₃ groups, as well as their problematic enzymology and cleavage.

How do they work?

The 3'-ONH₂ group is accepted by Firebird proprietary polymerases, and blocks further primer extension. It is cleaved in seconds in buffered aqueous sodium nitrite.



What are they used for? FfAME architectures use this terminator in SNP analysis and sequencing, for example.



Polymorphisms (SNPs) are detected with low background by competing Firebird reversible terminators with irreversible terminators, even when non-SNPped DNA is present in abundance.

Oxime blocked untagged reversible terminators

Triphosphates carrying the 3'-ONH₂ reversible terminator are prepared as acetone oximes, which are stable indefinitely. The oxime is deprotected in seconds *in situ* with dilute buffered hydroxylamine before use.





Thymine TONH2-DH101 5 µmoles \$ 600.00 Bulk inquire

Cytosine CONH2-MK102 5 µmoles \$ 750.00 Bulk inquire

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Adenine AONH2-DH103 5 µmoles \$ 750.00 Bulk inquire

0000 		Guanine GONH2-D 5 µmoles Bulk	H104 \$ 750.00 inquire
HPLC purified	N ⁻ ×		

Polymerase variants have been developed that incorporate these with improved efficiency over standard Taq DNA polymerase

Polymerase 475	400 units	\$110.00
POL475-NL201	1000 units	280.00

Reversible terminators with diol-linked tags

For sequencing research, Firebird offers triphosphates with a 3'- ONH_2 reversible terminator and a diol linker carrying a free amino group, to which can be attached a fluor or other signaling moiety. The diol is cleaved in seconds by aqueous periodate.



The $-ONH_2$ is protected as its oxime, to allow customers to add (if they desire, for research purposes) fluorescent reporters of their own choice. The oxime is deprotected with dilute buffered hydroxylamine to give free $-ONH_2$.

Note: Not available for sequencing applications on certain machines and in certain jurisdictions. Please inquire.

Reversible terminators with disulfide-linked tags

For sequencing research, Firebird also offers triphosphates with a 3' $-ONH_2$ reversible terminator and a conventional disulfide-containing linker carrying a free amino group, to which can be attached a fluor or other signaling moiety.





TONH2ST-DH491 1 µmole \$1750.00 Bulk: inquire

Cytosine CONH2ST-MK702 1 µmole \$2000.00 inquire

Adenine AONH2ST-DH703 1 µmole \$2400.00 Bulk inquire

Guanine GONH2ST-DH704 1 µmole \$2500.00 Bulk inquire

The -ONH₂ is protected as its oxime, to allow customers to add (if they desire, for research purposes) fluorescent reporters of their own choice. The oxime is deprotected with dilute buffered hydroxylamine to give free -ONH₂.

Note: Not available for sequencing applications on certain machines and in certain jurisdictions. Please inquire.

Achieving orthogonality

Artificially Expanded Genetic Information Systems (AEGIS)TM

The concept

When detecting DNA in complex biological media, uninteresting DNA can interfere.

In 1990, innovators at Firebird solved this problem, developing an Artificially Expanded Genetic Information System (AEGIS)TM.



By shuffling hydrogen bond donors and acceptors, the Z:P, K:X. and S:B pairs were created.



Artificially Expanded Genetic Information Systems (AEGIS)TM

The complete AEGIS expanded alphabet. Different heterocycles implement the various AEGIS hydrogen bonding patterns. Shown are our most popular heterocycles.



Artificially Expanded Genetic Information Systems **dZ:dP system. Nucleosides & phosphoramidites**







Hydrogen bonding pattern: Z Sugar: 2'-Deoxyribose Heterocycle: Nitropyridine Derivative: Free nucleoside Cat. No. ZdNF-05010201 100 mg \$340.00 1 gram \$2720.00 Bulk inquire

Hydrogen bonding pattern: P Sugar: 2'-Deoxyribose Heterocycle: Imidazotriazine Derivative: Free nucleoside Cat. No. PdNF-06010601 100 mg \$370.00 1 gram \$2960.00 Bulk inquire

Hydrogen bonding pattern: Z Sugar: 2'-Deoxyribose Heterocycle: Nitropyridine Derivative: Phosphoramidite Cat. No. ZdNP-05010203 100 mg \$ 720.00 1 gram \$5760.00 Bulk inquire

Hydrogen bonding pattern: P Sugar: 2'-Deoxyribose Heterocycle: Imidazotriazine Derivative: Phosphoramidite Cat. No. PdNP-06010603 100 mg \$ 850.00 1 gram \$6800.00 Bulk inquire

Oligonucleotides are available that incorporate dZ and dP. Ribonucleoside derivatives are also available. *Please inquire*.

Artificially Expanded Genetic Information Systems The dZ:dP system: Triphosphates and polymerases



Hydrogen bonding pattern: ZSugar: 2'-deoxyriboseHeterocycle: ImidazotriazineDerivative: TriphosphateCat. No. PdNT-050102041 μmole\$ 400.005 μmoles\$ 1600.00Bulkinquire

Hydrogen bonding pattern: P Sugar: 2'-deoxyribose Heterocycle: Imidazotriazine Derivative: Triphosphate Cat. No. PdNT-06010604 1 µmole \$ 420.00 5 µmoles \$ 1680.00 Bulk inquire

These are fifth and sixth letters of a DNA alphabet that are used in priming, primer extension, and PCR.

Firebird innovators have developed and tested polymerases to incorporate dZ opposite dP and dP opposite dZ.

ZiP Polymerase	400 units	\$300.00
ZiPOL475-ZY201	1000 units	\$500.00

Ribonucleoside triphosphates and supporting RNA polymerases are also available. *Please inquire*.

Artificially Expanded Genetic Information Systems The dK:dX system. Nucleosides

Multiple heterocycles implement the dK and dX hydrogen bonding patterns. Two each are offered as nucleosides, phosphoramidites, & triphosphates.









Hydrogen bonding pattern: K Sugar: 2'-deoxyribose Heterocycle: Pyrimidine Derivative: Free nucleoside Cat. No. KdYF-07010101 100 mg \$ 380.00 1 gram \$ 3040.00 Bulk inquire

Hydrogen bonding pattern: K Sugar: 2'-deoxyribose Heterocycle: Nitropyridine Derivative: Free nucleoside Cat. No. KdPF-07010201 100 mg \$ 360.00 1 gram \$2880.00 Bulk inquire

Hydrogen bonding pattern: X Sugar: 2'-deoxyribose Heterocycle: Imidazotriazine Derivative: Free nucleoside Cat. No. XdIF-08010601 100 mg \$ 550.00 1 gram \$4400.00 Bulk inquire

Hydrogen bonding pattern: X Sugar: 2'-deoxyribose Heterocycle: Purine Derivative: Free nucleoside Cat. No. XdRF-07010401 100 mg \$ 230.00 1 gram \$1840.00 Bulk inquire

Oligonucleotides are available with dK and/or dX in their multiple forms. Ribonucleosides are also available. *Please inquire.*

Artificially Expanded Genetic Information Systems The dK:dX system. Phosphoramidites







Hydrogen bonding pattern: K Sugar: 2'-Deoxyribose Heterocycle: Pyrimidine Derivative: Phosphoramidite Cat. No. KdNF-07010203 100 mg \$720.00 1 gram \$5760.00 Bulk inquire

Hydrogen bonding pattern: K Sugar: 2'-Deoxyribose Heterocycle: Nitropyridine Derivative: Phosphoramidite Cat. No. KdNP-07010103 100 mg \$780.00 1 gram \$6240.00 Bulk inquire

Hydrogen bonding pattern: X Sugar: 2'-Deoxyribose Heterocycle: Imidazotriazine Derivative: Phosphoramidite Cat. No. XdIP-08010603 100 mg \$480.00 1 gram \$3840.00 Bulk inquire

Hydrogen bonding pattern: X Sugar: 2'-deoxyribose Heterocycle: Purine Derivative: Phosphoramidite Cat. No. XdPP-08010403 100 mg \$252.00 1 gram \$2016.00 Bulk inquire

Ribonucleoside phosphoramidites are also available. Please inquire.

Artificially Expanded Genetic Information Systems The dK:dX system. Triphosphates and polymerases.



HO

Hydrogen bonding pattern: KSugar: 2'-DeoxyriboseHeterocycle: PyrimidineDerivative: TriphosphateCat. No.: KdYT-070101041 μmole\$400.005 μmoles\$1600.00Bulkinquire

Hydrogen bonding pattern: KSugar: 2'-DeoxyriboseHeterocycle: NitropyridineDerivative: TriphosphateCat. No.: KdNT-070102041 μmole\$400.005 μmoles\$1600.00Bulkinquire

Hydrogen bonding pattern: XSugar: 2'-DeoxyriboseHeterocycle: ImidazotriazineDerivative: TriphosphateCat. No.: XdIT-080106031 μmole\$400.005 μmoles\$1600.00Bulkinquire

Hydrogen bonding pattern: XSugar: 2'-DeoxyriboseHeterocycle: PurineDerivative: TriphosphateCat. No.: XdRT-080104041 μmole\$300.005 μmoles\$1200.00Bulkinquire

Firebird innovators have developed polymerases to incorporate dK opposite dX and dX opposite dK. KiX Polymerase 400 units \$300.00 KiXpOL475-ZY201 1000 units 500.00

Ribonucleoside triphosphates and supporting RNA polymerases are also available. *Please inquire*.

Artificially Expanded Genetic Information Systems The dS:dB system. Nucleosides

Multiple heterocycles implement the dS and dB hydrogen bonding patterns. All are available as nucleosides, phosphoramidites, & triphosphates.



Hydrogen bonding pattern: SSugar: 2'-DeoxyriboseHeterocycle: PyrimidineDerivative: Free nucleosideCat. No.: SdYF-09010101100 mg\$50.001 gram\$280.00Bulkinquire

Hydrogen bonding pattern: S Sugar: 2'-Deoxyribose Heterocycle: Pseudo Derivative: Free nucleoside Cat. No.: SdPF-09010801 100 mg \$ 340.00 1 gram \$2720.00 Bulk inquire

Hydrogen bonding pattern: B Sugar: 2'-Deoxyribose Heterocycle: Purine Derivative: Free nucleoside Cat. No.: BdRF-10010401 100 mg \$ 150.00 1 gram \$ 900.00 Bulk inquire

Hydrogen bonding pattern: B Sugar: 2'-Deoxyribose Heterocycle: Deazapurine Derivative: Free nucleoside Cat. No.: BdDF-10010501 100 mg \$ 500.00 1 gram \$ 4000.00 Bulk inquire

Oligonucleotides are available with dS and/or dB in their multiple forms. Ribonucleosides for the S:B system are also available. *Please inquire.*

Artificially Expanded Genetic Information Systems The dS:dB system. Phosphoramidites









Hydrogen bonding pattern: S Sugar: 2'-Deoxyribose Heterocycle: Pyrimidine Derivative: Phosphoramidite Cat. No.: SdYP-09010103 100 mg \$ 180.00 1 gram \$ 900.00 Bulk inquire

Hydrogen bonding pattern: S Sugar: 2'-Deoxyribose Heterocycle: Pseudo Derivative: Phosphoramidite Cat. No.: SdPP-09010804 100 mg \$ 720.00 1 gram \$ 5760.00 Bulk inquire

Hydrogen bonding pattern: B Sugar: 2'-Deoxyribose Heterocycle: Deazapurine Derivative: Phosphoramidite Cat. No.: BdRP-10010504 100 mg \$ 850.00 1 gram \$ 6800.00 Bulk inquire

Hydrogen bonding pattern: B Sugar: 2'-Deoxyribose Heterocycle: Purine Derivative: Phosphoramidite Cat. No.: BdDP-10010404 100 mg \$ 600.00 1 gram \$ 4800.00 Bulk inquire

Ribonucleoside phosphoramidites are also available. Please inquire.

Artificially Expanded Genetic Information Systems The dS:dB system. Triphosphates and polymerases







Hydrogen bonding pattern: S Sugar: 2'-Deoxyribose Heterocycle: Pyrimidine Derivative: Triphosphate Cat. No.: SdYT 1 μmole \$150.00 5 μmoles \$900.00 Bulk inquire

Hydrogen bonding pattern: SSugar: 2'-DeoxyriboseHeterocycle: PseudoDerivative: TriphosphateCat. No.: SdPT1 μmole\$400.005 μmoles\$3200.00Bulkinquire

Hydrogen bonding pattern: BSugar: 2'-DeoxyriboseHeterocycle: DeazapurineDerivative: TriphosphateCat. No.: BdRT-100105041 μmole\$400.005 μmoles\$3200.00Bulkinquire

Hydrogen bonding pattern: BSugar: 2'-DeoxyriboseHeterocycle: PurineDerivative: TriphosphateCat. No.: BdDT-100104041 μmole\$400.005 μmoles\$3200.00Bulkinquire

Firebird innovators have developed and tested polymerases to incorporate dS opposite dB and dB opposite dS. SiB Polymerase 400 units \$300.00 SiBpOL475-ZY201 1000 units \$500.00 Ribonucleoside triphosphates are also available. *Please inquire*.

Using AEGIS Ultra-clean nested PCR

Oligonucleotides up to 60 n	ts are available. Per 50
nmols, desalted (add \$150.0	DU for HPLC purification).
Setup	\$90.00
Per nucleotide	0.85
For each internal AEGIS	15.00
For each 3'AEGIS	45.00
Shipping and handling \$40.	00 per order plus \$6.00 per
additional sequence.	

Using dZ and dP oligonucleotides

The dZ:dP pair is PCR amplified, even when adjacent in amplicons

dZTP

dPTP



dP in external primers cleans up nested PCR.



AEGIS pairs improve molecular beacons

Sheng *et al.* (2008) Design of a novel molecular beacon. Modification of the stem with artificially genetic alphabet. *Chem. Comm.* (41) 5128

AEGIS beacons are available by custom synthesis. Please inquire.

Self avoiding molecular recognition systemsSAMRS™The concept

When multiplexing DNA priming or the polymerase chain reaction, various primers added in large amounts bind each other to create a PCR "mess". Selectively removing hydrogen bonding units gives *self-avoiding* DNA primers that can be added without limit to enable multiplexed PCR.

The SAMRS[™] concept idealized \bigcirc 0 •• 0 G C_{2 H bonds} G* С 3 H bonds •• 0 🕢 H G* G **C*** С 2 H bonds 1 H bond H₃C H₃C Α Δ* T _{2 H bonds} Т 2 H bonds H₃C H₃C Δ* * 2 H bonds Α Т Т 1 H bond

Self avoiding molecular recognition systems SAMRS[™] phosphoramidites

T* phosphoramidite \$ 1050 1 g \$ 8220 10 g A* phosphoramidite \$ 655 1 g 10 g \$ 5340 C* phosphoramidite \$ 394 1 g 10 g \$3150 G* phosphoramidite \$ 110 1 g

1 g \$ 110 10 g \$ 880

Recommended lengths are 20-35 nts with 4-6 SAMRS components.

A set of 48 SAMRS oligonucleotides with customer-selected selected is available for \$4800.

Contact for pricing for smaller numbers of SAMRS oligonucleotides.

SNAP2[™] Oligonucleotides

Priming with both high specificity and high discrimination

The concept

When priming, one needs the specificity of a 16mer (or longer) but the ability to discriminate against single mismatches. Dynamic assembly of a primer on a template can provide this.



Controlled pore glass with aldehyde precursor SNAP2T_HJK001 100 mg \$600.00

Leal, N., Sukeda, M., Benner, S. A. (2006) Dynamic assembly of primers on nucleic acid templates. *Nucleic Acids Res.* **34**, 4702-4710

Not to be confused with Snap-Tag® and other registered trademarks of New England BioLabs.

PIRE BIRD

BiversalTM nucleotides

Often, target xNA sequences divergently evolve, making it impossible to know the exact complement to use as a probe or primer. This is often handled using "universal bases" (e.g. inosine). Inspired by the work of Dan Brown, Firebird scientists have rolled evolutionary analyses into nucleoside chemistry to create two "biversals" TM, a purine biversal that pairs with either thymine or cytosine and a pyrimidine biversal that pairs with either guanine or adenine.

Pyrimidine biversal phosphoramidite 100 mg \$ 500 1 g \$ 4000

Purine biversal phosphoramidite 100 mg \$ 1050 1 g \$ 8220



For oligonucleotides that contain the purine and pyrimidine biversals, *please inquire*.

Purchasing, shipping and handling

Contact

Orders By email: orders@firebirdbio.com By telephone: Firebird Biomolecular Sciences (386) 418-0347 9:00 - 5:00 east coast time

<u>Technical inquiries</u> support@firebirdbio.com

Reagents and enzymes are shipped at the customer's expense, on dry ice as appropriate.