

## **Product Specifications**

Custom Oligo Synthesis, antisense oligos, RNA oligos, chimeric oligos, Fluorescent dyes, Affinity Ligands, Spacers & Linkers, Duplex Stabilizers, Minor bases, labeled oligos, Molecular Beacons, siRNA, phosphonates Locked Nucleic Acids (LNA); 2'-5' linked Oligos

## Oligo Modifications

For research use only. Not for use in diagnostic procedures for clinical purposes.

## H-Phosphonate dA. dA(H-p)

Category	Others	NH <sub>2</sub>
Modification Code	H-p-dA	5' Oligo VVV-O
Reference Catalog Number	26-6581	
5 Prime	Υ	ОН
3 Prime	Υ	0 0=P-0-1 0=P-0-1
Internal	Υ	H CO
Molecular Weight(mw)	298.2	dA H-phosphonate   Carlo   Car

H-phosphonate modification has a setup charge of \$250.00 per order for special synthesis reagents .

## Special notes on H-phosphonate modification.

- 1. **H-phosphonate chimeric oligos.** H-phosphonate bases/oligos can be synthesized in a sequence with standard bases or modifications containing phosphodiester linkages the 3' end of the sequence. If a chimeric oligo is designed then the H-phosphonate should be on the 5' end with no other standard phosphodiester bases at 5' of the H-phosphonate linked bases. Chemical oligo synthesis starts from the 3' and the phosphodiester linkages are generated by oxidation. Any such oxidation cycle after coupling the H-phosphonate will oxidize the H-phosphonate linkage to phosphodiester linkage.
- 2. **Special oxidation.** H-phosphonate modified oligos can be supplied with oligo bound to CPG solid support for the end user to perform special oxidation at the H-phosphonate linkage. Example is for S-35 radio-labeled phosphorothioate linkage. Please specify when placing an order.

H-phosphonoamidites are deoxynucleoside amidites modified such that, when incorporated into an oligonucleotide, that base position will have an H-phosphonate backbone linkage instead of the standard phosphodiester linkage. In the H-phosphonate linkage, one of the oxygen bound to the phosphorus atom in a phosphodiester linkage has been replaced by hydrogen. H-phosphonate-modified oligos are typically synthesized when a researcher wants to prepare a S-35 radiolabeled phosphorothioate linkage (1). For the radiolabeled phosphorothioate, an H-phosphonate linkage is incorporated at the desired position using phosphoramidite chemistry, and then S-35 is used to replace the hydrogen atom via a sulfurization reaction (2). For the phosphoroamidate, N,N-dimethylaminoethylamine is reacted with the H-phosphonate to form the substituted linkage. Phosphoroamidates, being a cationic linkage, may provide nuclease resistance and improved cell permeability (3).

- References
- 1. Maier, M.A., Guzaev, A.P., Manoharan, M. Synthesis of Chimeric Oligonucleotides Containing Phosphodiester, Phosphorothioate, and Phosphoramidate Linkages. *Org. Lett.* (2000), **2**: 1819-1822.
- 2. Wallin, R., Kalek, M., Bartoszewicz, A., Thelin, M., Stawinski, J. On the Sulfurization of H-Phosphonate Diesters and Phosphite Triesters Using Elemental Sulfur.



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3. Letsinger, R.L., Singman, C.N., Histand, G., Salunkhe, M. Cationic Oligonucleotides. *J. Am. Chem. Soc.* (1988), **1110**: 4470-4471.

