## WikipediA

# **Thymidine**

**Thymidine** (**deoxythymidine**; other names **deoxyribosylthymine**, **thymine deoxyriboside**) is a pyrimidine deoxynucleoside. Deoxythymidine is the <u>DNA nucleoside</u> T, which pairs with <u>deoxyadenosine</u> (A) in double-stranded DNA. In cell biology it is used to <u>synchronize</u> the cells in G1/early S phase.

Before the boom in thymidine use caused by the need for thymidine in the production of the <u>antiretroviral drug azidothymidine</u> (AZT), much of the world's thymidine production came from <u>herring sperm</u>. Thymidine occurs almost exclusively in DNA but it also occurs in the T-loop of tRNA.

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# Structure and properties

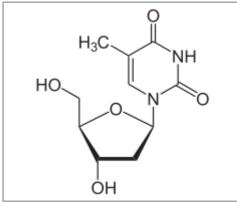
In its composition, deoxythymidine is a <u>nucleoside</u> composed of <u>deoxyribose</u> (a pentose <u>sugar</u>) joined to the <u>pyrimidine</u> base thymine.

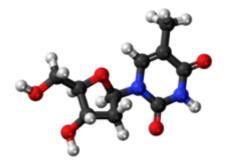
Deoxythymidine can be phosphorylated with one, two or three phosphoric acid groups, creating <u>dTMP</u> (**d**eoxythymidine **m**onophosphate), <u>dTDP</u>, or <u>dTTP</u> (for the **d**i- and **t**ri-phosphates, respectively).

It exists in <u>solid</u> form as small white <u>crystals</u> or white crystalline <u>powder</u>. It has a <u>molecular weight</u> of 242.229 <u>u</u> and a <u>melting</u> <u>point</u> of 185 °C. The stability of deoxythymidine under <u>standard</u> temperature and pressure (STP) is very high.

Deoxythymidine is non-toxic and as part of one of the four nucleosides in DNA it is a naturally occurring compound that exists in all living organisms and DNA viruses. Instead of thymidine, RNA contains <u>uridine</u> (<u>uracil</u> joined to <u>ribose</u>). Uracil is chemically very similar to thymine, which is also known as 5-methyluracil. Since thymine nucleotides are precursors of DNA (but not RNA), the prefix "deoxy" is often left out, i.e., deoxythymidine is often just called thymidine.

#### **Thymidine**





#### **Names**

#### Preferred IUPAC name

1-[(2R,4S,5R)-4-Hydroxy-5-(hydroxymethyl)oxolan-2-yl]-5methylpyrimidine-2,4(1H,3H)dione

#### Other names

Deoxythymidine, Td, dT, 1-[(2*R*,4*S*,5*R*)-4-Hydroxy-5-(hydroxymethyl)tetrahydrofur-2yl]-5-methyl-1,3dihydropyrimidine-2,4-dione

| Identifiers         |   |
|---------------------|---|
| CAS Number          | 50-89-5 ✓   |
| 3D model<br>(JSmol) | Interactive image (ht tps://chemapps.stola f.edu/jmol/jmol.php? model=Cc1cn%28 c%28%3DO%29%5 BnH%5Dc1%3DO% 29%5BC%40H%5D 2C%5BC%40%40 H%5D%28%5BC%4 0H%5D%28O2%29 CO%29O) |

Thymidine is listed as a chemical teratogen.  $\lfloor 2 \rfloor$ 

# **Modified analogs**

<u>Iododeoxyuridine</u> is a radiosensitizer and increases the amount of DNA damage received from ionizing radiation.

Azidothymidine (AZT) – used in the treatment of HIV infection. AZT inhibits the process of reverse transcription, a critical step in the viral life cycle.

Radiolabeled thymidine (TdR), such as tritiated thymidine (<sup>3</sup>H-TdR), is commonly used in cell proliferation assays. The thymidine is incorporated into dividing cells and the level of this incorporation, measured using a liquid scintillation counter, is proportional to the amount of cell proliferation. For example, lymphocyte proliferation can be measured this way in lymphoproliferative disorders.

<u>Bromodeoxyuridine</u> (BrdU) is another thymidine analog that is often used for the detection of proliferating cells in living tissues.

<u>5-Ethynyl-2´-deoxyuridine</u> (EdU) is a thymidine analog which is incorporated into the DNA of dividing cells and is used to assay DNA synthesis in cell culture or living tissues. It can be visualized by covalently binding a fluorescent azide using <u>click chemistry</u>, which is less harsh than the conditions used to expose the <u>epitope</u> for BrdU antibodies.

Edoxudine is an antiviral drug.

# Thymidine imbalance induces mutation and recombination

During growth of <u>bacteriophage T4</u>, an excess of thymidine availability increases <u>mutation</u>. A deficiency of thymidine during growth also increases mutation. A thymidylate auxotroph of the diploid yeast *Saccharomyces cerevisiae* was grown under conditions in which thymidyate levels varied from excess to depletion. High levels of thymidylate were observed to be mutagenic and recombinogenic, whereas starvation for thymidylate was recombinogenic but only slightly mutagenic.

# References

- 1. Makoto Ishii; Hideyuki Shirae; Kenzo Yokozeko, *Enzymatic Production of 5-Methyluridine from Purine Nucleosides and Thymine by Erwinia carotovora AJ-2992*, Agricultural and Biological Chemistry
- 2. The following list of chemicals known or believed to be teratogens is drawn primarily from Dangerous Properties of Industrial Materials, 7th Ed., by N. Irving Sax and Richard J. Lewis.

|                               | s://www.ebi.ac.uk/ch<br>embldb/index.php/co<br>mpound/inspect/ChE<br>MBL52609) *                                  |
|-------------------------------|---|
| ChemSpider                    | 5585 (http://www.ch<br>emspider.com/Chem<br>ical-Structure.5585.h<br>tml) *                                       |
| DrugBank                      | DB04485 (https://www.drugbank.ca/drugs/DB04485) *   |
| ECHA<br>InfoCard              | 100.000.065 (https://echa.europa.eu/substance-information/-/substanceinfo/100.000.065)                            |
| IUPHAR/BPS                    | 4718 (http://www.gui<br>detopharmacology.o<br>rg/GRAC/LigandDis<br>playForward?tab=su<br>mmary&ligandId=47<br>18) |
| MeSH                          | Thymidine (https://w<br>ww.nlm.nih.gov/cgi/<br>mesh/2014/MB_cgi?<br>mode=&term=Thymi<br>dine)                     |
| PubChem<br>CID                | 1134 (https://pubche<br>m.ncbi.nlm.nih.gov/c<br>ompound/1134)   |
| UNII                          | VC2W18DGKR (http<br>s://fdasis.nlm.nih.go<br>v/srs/srsdirect.jsp?re<br>gno=VC2W18DGK<br>R) *                      |
| CompTox<br>Dashboard<br>(EPA) | DTXSID5023661 (htt ps://comptox.epa.go v/dashboard/DTXSI D5023661)  |
| InChI<br>InChI=1S/C10H14      | N2O5/c1-5-3-12(10(16)11-  |

InChI=1S/C10H14N2O5/c1-5-3-12(10(16)11-9(5)15)8-2-6(14)7(4-13)17-8/h3,6-8,13-14 H,2,4H2,1H3,(H,11,15,16)/t6-,7+,8+/m0/s

Key: IQFYYKKMVGJFEH-XLPZGREQSA-N

- Bernstein C, Bernstein H, Mufti S, Strom B. Stimulation of mutation in phage T 4 by lesions in gene 32 and by thymidine imbalance. Mutat Res. 1972 Oct;16(2):113-9. doi: 10.1016/0027-5107(72)90171-6. PMID: 4561494
- DeVries JK, Wallace SS. Reversion of bacteriophage T4rII mutants by high levels of pyrimidine deoxyribonucleosides. Mol Gen Genet. 1982;186(1):101-5. doi: 10.1007/BF00422919. PMID: 7050620
- 5. Eckardt F, Kunz BA, Haynes RH. Variation of mutation and recombination frequencies over a range of thymidylate concentrations in a diploid thymidylate auxotroph. Curr Genet. 1983 Sep;7(5):399-402. doi: 10.1007/BF00445881. PMID: 24173422

## **External links**

InChI=1/C10H14N2O5/c1-5-3-12(10(16)11-9 (5)15)8-2-6(14)7(4-13)17-8/h3,6-8,13-14 H,2,4H2,1H3,(H,11,15,16)/t6-,7+,8+/m0/s 1 Key: IQFYYKKMVGJFEH-XLPZGREQBQ

#### SMILES

Cc1cn(c(=O)[nH]c1=O)[C@H]2C[C@@H]([C @H](O2)CO)O

| Properties       |   |
|------------------|---|
| Chemical formula | C <sub>10</sub> H <sub>14</sub> N <sub>2</sub> O <sub>5</sub> |
| Molar mass       | 242.231 g·mol <sup>-1</sup>                                   |
| Melting point    | 185 °C  |

Except where otherwise noted, data are given for materials in their standard state (at 25 °C [77 °F], 100 kPa).

verify (what is \*\*?)

Infobox references

http://money.cnn.com/magazines/fortune/fortune\_archive/1990/11/05/74308/index.htm

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