# 第3章基因的本质

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第1节 DNA是主要的遗传物质

第2节 DNA分子的结构

第3节 DNA的复制

第4节 基因是有遗传效应的DNA片段

## 第2节 DNA分子的结构

本节聚焦

- 1、沃森和克里克是怎样发现DNA分子的双螺旋结构的?
- 2、DNA分子的双螺旋结构有哪些主要特点?

#### 一、问题探讨



DNA雕塑

坐落于北京中关村高科技园区的DNA 雕塑,以它简洁而独特的双螺旋造型吸引 着过往行人。你知道为什么将它作为高科 技的标志吗?

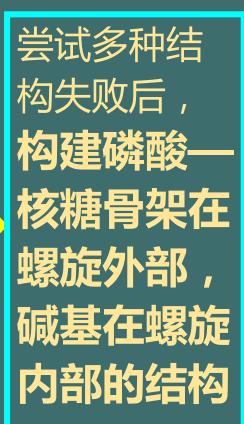
2003年是DNA分子双螺旋结构发现50周年。上网查一查有关DNA的信息,收集你感兴趣的资料与同学交流共享。

DNA分子是以 4中脱氧核苷酸为单 位连接而成的长链, 这4中脱氧核苷酸分 别含有ATGC四种 碱基。

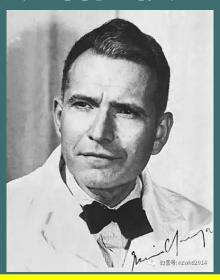
1951年春



DNA分子 呈螺旋结构



尝试多种结构 失败后,构 建磷酸—核 糖骨架在螺 旋外部,碱 基在螺旋内 部的结构;



A=T, G=C;

A与T配对, G与C配对, 构建新的 DNA模型;

相同碱基进 行配对:

A-T碱基对,与 G-C礦基对具有 形同的形状和直 径——DNA 具 有 稳定的直径,能 够解释碱基数量 关系, 也能解释 违反化学规律 DNA的复制;

A-T礦基对,与

G-C碱基对具有 形同的形状和直

----DNA具有

稳定的直径,

够解释碱基数量

关系,也能解释

DNA的复制;

#### 1953年

No. 4356 April 25, 1953

NATURE

#### MOLECULAR STRUCTURE OF NUCLEIC ACIDS

A Structure for Deoxyribose Nucleic Acid

E wish to suggest a structure for the salt of deoxyribose can bond together. These pairs are: admine (purins) with mymine which are of considerable blookgatal interest.

In other words, if an admine forms one membradies, in other words, if an admine forms one membradies, in other words, if an admine forms one membradies, in other words, if an admine forms one membradies.

the salt, not the free acid. Without the acidic hydrogen atoms it is of adenine to thymine, and the ratio of guanine to cytosine, are not clear what forces would hold the structure logether, especially always very close to unity for deoxyribose nucleic acid. as the negatively charged phosphates near the axis will repel each other. (2) Some of the van der Waals distances appear to be too sugar in place of the deoxyribose, as the extra oxygen atom would

(in the press). In his model the phosphates are on the outside and the bases on the inside, linked together by hydrogen bonds. This we can tell, it is roughly compatible with the experimental data,

We wish to put forward a natically different surfaces for time following, communications, we were not aware or use causes who shall of decayyibles nacietic acid. This structure has two helical in the results presented there when we devised our structure, which chains each coiled found the same axis (see diagram). We have rests mainly though not entirely on published experimental data made the usual chemical assumptions, namely, that each chain afferes chemical arguments. consists of phosphate dester groups joining B-D-deoxy-ribofuranose residues with 3°,5° linkages. The two chains (but not their bases) are related by a dyad perpendicular to the fibre axis.

Both chains follow righthanded helices, but owing to the dyad the Full details of the structure, including the conditions assumed

and the acoust teat it is cross to runned great a impossinger commencing results and acoust of *t. M. H. P.* standard configuration<sup>†</sup>, the sugard being Wilkins, Dr. R. E. Franklin and their co-workers at roughly perpendicular to the attached King's College, London. One of us (J.D.W.) has been aided by a bease. There is a residue on each chain every 3-4 A. in the z-direction. We have assumed an angle of 36° between adjacent residues in the same chain, so that the structure repeats after 10 residues on each chain, that is, after 34 A. The distance of a Medical Research Council Unit for the Study of the Molecular A. As the phosphates are on the outside, Cambridge. April 2. cations have easy access to them.

The structure is an open one, and its water contents we would expect the bases (1953). to tilt so that the structure could become Fusberg, S., Acta Chem. Scand., 6, 634 (1952). more compact.

manner in which the two chains are held 'Wyott, G.R. J. Gen. Physiol., 36 201 (1952). being hydrogen-bonded to a single base from the other chain, so

that the two lie side by side with identical z-co-ordinates. One of the pair must be a purine and the other a pyrimidine for bonding to occur. The hydrogen bonds are made as follows: purine position 1 to pyrimidine position 1; purine position 6 to pyrimidine position

If it is assumed that the bases only occur in the structure in the most plausible tautomeric forms (that is, with the keto rather than the enol configurations) it is found that only specific pairs of bases

which are of considerable biological interest.

A structure for nucleic acid has already been proposed by

In other words, if an adenine forms one member of a pair, on either chain, then on these assumptions the other member must be Pauling and Corey'. They kindly made their manuscript available thymine; similarly for guanine and cytosine. The sequence of to us in advance of publication. Their model consists of three inter-bases on a single chain, does not appear to be restricted in any twined chains, with the phosphates near the fibre axis, and the bases on the outside. In our opinion, this structure is unsatisfactory follows that if the sequence of bases on one chain, is given, then for two reasons:

(1) We believe that the material which gives the X-ray diagrams is

the sequence on the other chain is automatically determined.

It has been found experimentally he that the ratio of the amounts

make too close a van der Waals contact.

The previously published X-ray data5+6 on deoxyribose nucleic ure obuses our due house, missed, missed, missed vigetard by principal colours as well defined, in a for life first financial first first

sequence of the intervo chains run in opposite directions.

Each chain loosly resembles Furibregs of the control of the contro

phosphorus atom from the fibre axis is 10 Structure of Biological Systems, Cavendish Laboratory,

The novel feature of the structure is the Biochim, et Biophys. Acta, 9402 (1952).

together by the purine and pyrimidine \*Astbury, W.T., Symp. Soc. Exp. Biol. 1, Nucleic Acid, 66 (Camb. Univ. Press, 1947)

#### 1962年



Francis Harry Compton Crick (1916-2004)

James Dewey

Watson (1928 -)





Maurice Hugh Frederick Wilkins (1916-2004)



water content is rather high. At lower Pauling, L., and Corey, R. B. nature, 171, 346 (1953); Proc, U.S. Nat Acad. Sci., 39, 84

bases. The planes of the bases are perpendicular to the fibre axis. Wilkins, M. H. F. and Randall, J. T. Biochim. et. Biophys. Acta, 10, 102 (1953).

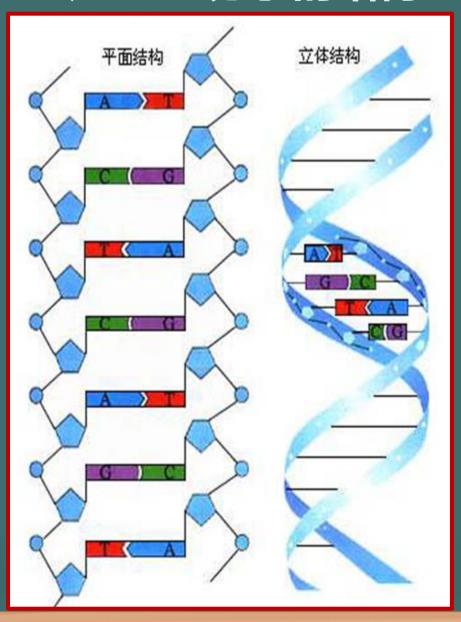
- 1、请根据资料回答有关DNA结构方面的问题:
- 1) DNA是由几条链构成的? 它具有怎样的立体结构?
- 1) DNA由2条链构成,盘旋成双螺旋结构;
- 2) DNA的基本骨架是由哪些物质组成的?它们分别位于DNA的什么部位?
  - 2) DNA的基本骨架由磷酸和脱氧核糖组成,位于DNA的螺旋外部;

- 3) DNA中的碱基是如何配对的?它们位于DNA的什么部位?
  - 1)A与T配对,G与C配对,分布在螺旋的内部;

- 2、上述资料中涉及到哪些学科的知识和方法? 这对你理解生物科学的发展有什么启示?
  - 2)主要涉及物理学(主要是晶体学)、生物化学、数学和分子生物学等学科的知识。涉及的方法主要有:X射线衍射结构分析方法,其中包括数学计算法;建构模型的方法等。现代科学技术中许多成果的取得,都是多学科交叉的运用,又会促进学科运用的结果;反过来,多学科交叉的运用,又会促进学科的发展,诞生新的边缘学科,如生物化学、生物物理学等。

- 3、沃森和克里克默契配合,发现DNA双螺旋结构的过程,作为科学家合作的典范,在科学界传为佳话。他们的这种工作方式给予你哪些启示?
  - 3)要善于利用他人的研究成果和经验;要善于与他人交流和沟通,闪光的思想是在交流与撞击中获得的;研究小组成员在知识背景上最好是互补的,对所从事的研究要有兴趣和激情等。

#### 三、DNA分子的结构



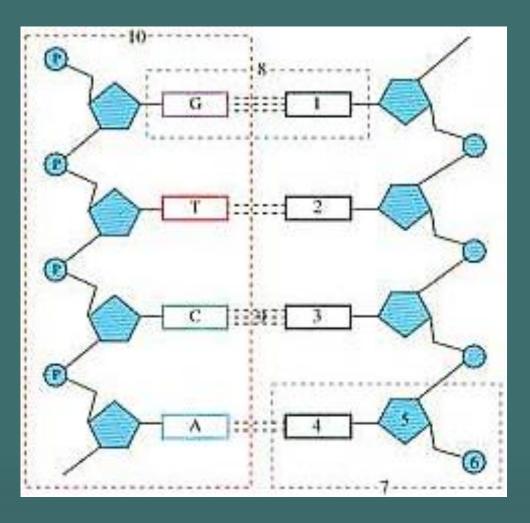
- 1) DNA分子由反向平行的双链组成,盘旋成双螺旋结构;
- 2) DNA分子的基本骨架由磷酸和脱氧核糖交替连接而成,排在外侧;碱基成对排在内侧;
- 3) DNA两条链上的碱基通过氢键进行互补配对: A—T, G—C; ——碱基互补配对原则;

### 三、DNA分子的结构

模型构建

制作DNA双螺旋结构模型

1、下面是DNA分子的结构模式图,请用文字写出图中1—10的名称;

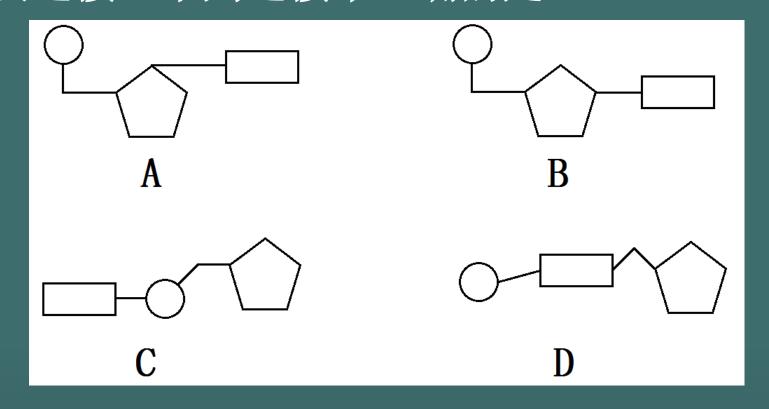


1,	$2 \varsigma$	

2、已知1个DNA分子中有4000个碱基对,其中胞嘧啶有2200个,这个DNA分子中应含有脱氧核苷酸的数目和腺嘌呤的数目分别是:

- A、4000个和900个
- B、4000个和1800个
- C、8000个和1800个
- D、8000个和3600个

3、下列各图中,图形 ● ■ 分别代表磷酸、脱氧核糖和碱基,在制作脱氧核苷酸模型时,各部件之间需要连接,下列连接中正确的是:



拓展题:你能根据碱基互补配对原则,推导出相关的数学公式吗?推导后,尝试进一步总结这些公式,从中概括出一些规律。

- $\cdot$  A=T, G=C
- A+G=T+C
- .. (A+G) /(

)=(T+C)/(

=50%

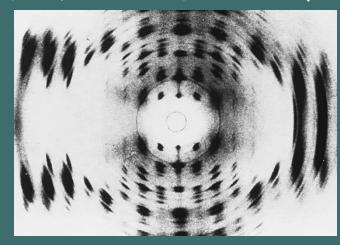
也可以写成以下形式:

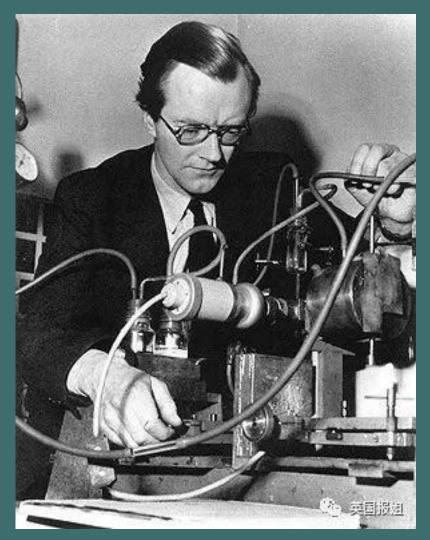
$$\frac{A+G}{T+C} = \frac{(}{} - \frac{)}{} = \frac{(}{} - \frac{)}{} - \cdots = 1$$

规律概括:在DNA双链中,任意两个不互补碱基之和

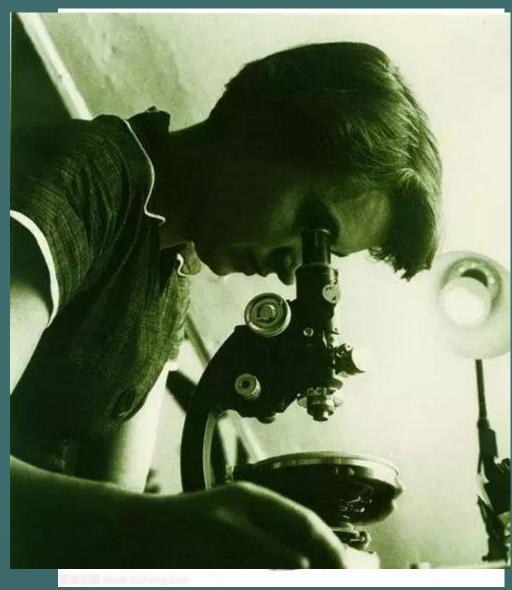
\_\_\_\_\_,并为碱基总数的\_\_\_\_\_;

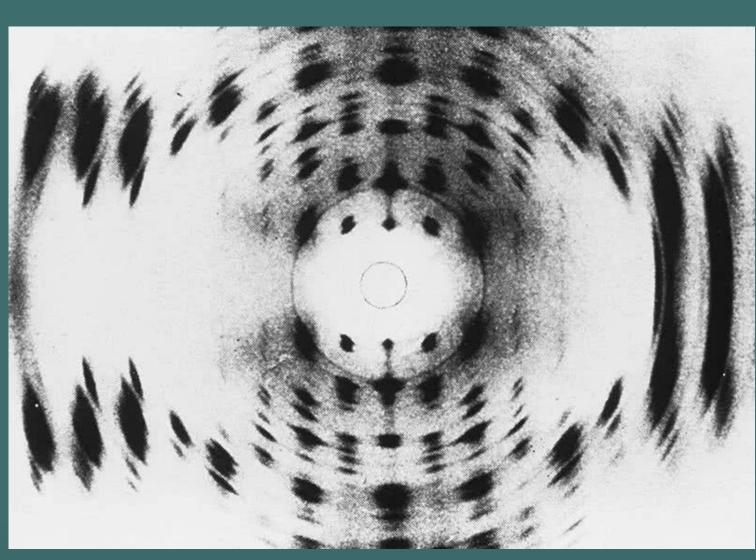
1951年春,在意大利举行的生物大分子结构会议上,展示了一张DNA的X射线的衍射幻灯片。

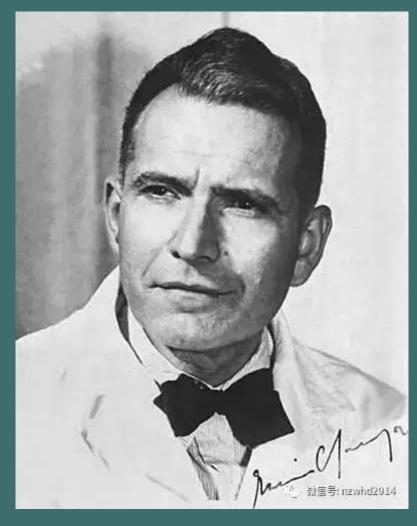




威尔金斯(M.Willkins)







埃尔文•查哥夫

DNA中的腺嘌呤与胸腺嘧啶数量几乎完全一样, 鸟嘌呤与胞嘧啶的数量也是一样;即A=T, G=C;

No. 4356 April 25, 1953

NATURE

#### MOLECULAR STRUCTURE OF NUCLEIC ACIDS

#### A Structure for Deoxyribose Nucleic Acid

nucleic acid (D.N.A.). This structure has novel features (pyrimidine), and guanine (purine) with cytosine (pyrimidine). which are of considerable biological interest.

(1) We believe that the material which gives the X-ray diagrams is not clear what forces would hold the structure together, especially always very close to unity for deoxyribose nucleic acid. as the negatively charged phosphates near the axis will repel each

It is probably impossible to build this structure with a ribose other. (2) Some of the van der Waals distances appear to be too sugar in place of the deoxyribose, as the extra oxygen atom would

made the usual chemical assumptions. namely, that each chain and stereo-chemical arguments. consists of phosphate diester groups joining B-D-deoxy- It has not escaped our notice that the specific pairing we have their bases) are related by a dyad perpendicular to the fibre axis. for the genetic material. Both chains follow righthanded helices, but owing to the dyad the Full details of the structure, including the conditions assumed

Each chain loosely resembles Furberg's2 will be published elsewhere. every 3-4 A. in the z-direction. We have assumed an angle of 36° between adjacent residues in the same chain, so that the structure repeats after 10 residues on each chain, that is, after 34 A. The distance of a A. As the phosphates are on the outside, Cambridge. April 2. cations have easy access to them.

The structure is an open one, and its water contents we would expect the bases (1953). to tilt so that the structure could become 2 Furberg, S., Acta Chem. Scand., 6, 634 (1952).

The novel feature of the structure is the Biochim, et Biophys. Acta, 9 402 (1952). manner in which the two chains are held 'Wyatt, G.R. J. Gen. Physiol., 36 201 (1952).

bases. The planes of the bases are perpendicular to the fibre axis. Wilkins, M. H. F. and Randall, J. T. Bischim, et. Biophys. Acta, 10, 102 (1953). They are joined together in pairs, a single base from one chain being hydrogen-bonded to a single base from the other chain, so

that the two lie side by side with identical z-co-ordinates. One of the pair must be a purine and the other a pyrimidine for bonding to occur. The hydrogen bonds are made as follows: purine position 1 to pyrimidine position 1; purine position 6 to pyrimidine position

If it is assumed that the bases only occur in the structure in the most plausible tautomeric forms (that is, with the keto rather than the enol configurations) it is found that only specific pairs of bases TE wish to suggest a structure for the salt of deoxyribose can bond together. These pairs are: adenine (purine) with thymine

In other words, if an adenine forms one member of a pair, on A structure for nucleic acid has already been proposed by either chain, then on these assumptions the other member must be Pauling and Corey1. They kindly made their manuscript available thymine; similarly for guanine and cytosine. The sequence of to us in advance of publication. Their model consists of three intertwined chains, with the phosphates near the fibre axis, and the way. However, if only specific pairs of bases can be formed, it bases on the outside. In our opinion, this structure is unsatisfactory follows that if the sequence of bases on one chain, is given, then the sequence on the other chain is automatically determined.

It has been found experimentally3,4 that the ratio of the amounts the salt, not the free acid. Without the acidic hydrogen atoms it is of adenine to thymine, and the ratio of guanine to cytosine, are

make too close a van der Waals contact.

Another three-chain structure has also been suggested by Fraser The previously published X-ray data 5.6 on deoxyribose nucleic (in the press). In his model the phosphates are on the outside and acid are insufficient for a rigorous test of our structure. So far as the bases on the inside, linked together by hydrogen bonds. This we can tell, it is roughly compatible with the experimental data, structure as described is rather ill-defined, and for this reason we but it must be regarded as unproved until it has been checked against more exact results. Some of these are given in time We wish to put forward a radically different structure for the following, communications. We were not aware of the details of salt of deoxyribose nucleic acid. This structure has two helical the results presented there when we devised our structure, which chains each coiled round the same axis (see diagram). We have rests mainly though not entirely on published experimental data

ribofuranose residues with 31,51 linkages. The two chains (but not postulated immediately suggests a possible copying mechanism

sequences of the atoms in the two chains run in opposite directions. in building it, together with a set of co-ordinates for the atoms,

model No. 1; that is, the bases are on the We are much indebted to Dr. Jerry Donohue for constant inside of the helix and the phosphates on advice and criticism, especially on interatomic distances. We have the outside. The configuration of the sugar also been stimulated by a knowledge of the general nature of the and the atoms near it is close to Furberg's unpublished experimental results and ideas of Dr. M. H. F. standard configuration', the sugar being Wilkins, Dr. R. E. Franklin and their co-workers at

roughly perpendicular to the attached King's College, London. One of us (J.D.W.) has been aided by a base. There is a residue on each chain fellowship from the National Foundation for Infantile Paralysis.

> J.D. WATSON F.H. C. CRICK

Medical Research Council Unit for the Study of the Molecular phosphorus atom from the fibre axis is 10 Structure of Biological Systems, Cavendish Laboratory,

water content is rather high. At lower 1 Pauling, L., and Corey, R. B. nature, 171, 346 (1953); Proc, U.S. Nat Acad. Sci., 39, 84

<sup>2</sup> Chargaff, E., for references see Zamenhof, S., Brawerman, G., and Chargaff, E.,

together by the purine and pyrimidine SAstbury, W.T., Symp. Soc. Exp. Biol. 1, Nucleic Acid, 66 (Camb. Univ. Press, 1947)

This figure is purely diagrammatic. The two ribbons symbolize the two phosphate—sugar chains, and the horizontal rods the pairs of bases holding the chains together. The vertical line marks the fibre axis

