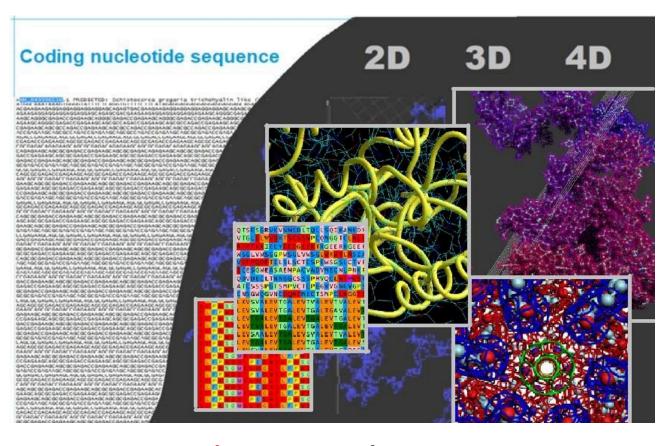
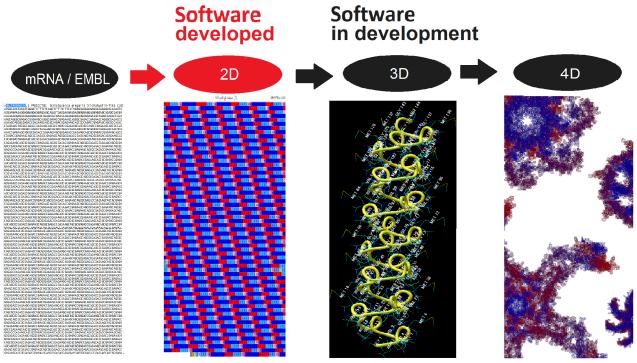
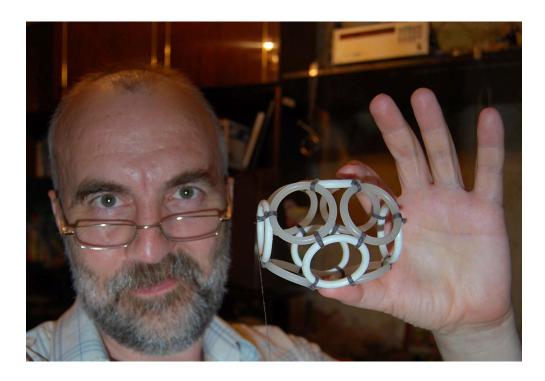
nanoworld_laboratory@mail.ru

Innovative method of converting nucleotide sequences into 2D, 3D and 4D structures







Dear Researchers!

Let us introduce to your attention our development – *Picotechnology of proteins* (<u>3D Genetic code</u>) - a method of constructing accurate <u>2D</u> structures, where the initial data is only the nucleotide sequence. Visualization of some <u>3D</u> and <u>4D</u> structures is available. This approach was made possible by modeling <u>electrons as tori (rings) of standing waves</u>, and <u>atomic shells as polyhedrons assembled from rings</u>.

In <u>this approach</u> all amino acid residues have the same structural template, and the third letter of the triplet controls the angle of rotation of the next amino acid residue relative to the previous one. Each of the angles is marked on the 2D diagram with its own color and corresponds to the algorithm for forming alpha, beta, pi, 310, methionine and proline helices.

On the 2D color diagram we clearly see the helical sections, individual turns and individual amino acid residues.

We can directly convert the 2D colour diagram to 3D and 4D structure for some structural problems (3D and 4D modeling takes into account <u>other adjustments</u>). Software for 2D structures has been developed. Software for 3D and 4D structures is under development.

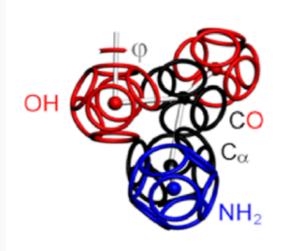
Our 2D diagrams reveal super-secondary structures (<u>fractal helices</u>, <u>software helices</u>), a coincidence confirmed by published data. The closure of disulfide bridges obtained on 2D and 3D models coincided with experimental data.

We hope that our algorithms will increase the accuracy of data processing both in the field of computational biology and in the field of <u>X-ray structure prediction</u>, <u>CD</u> and NMR methods. We would appreciate your comments and cooperation, see <u>DEMO PICOTECH 2D, 3D, 4D, AND ARTICLES</u>.

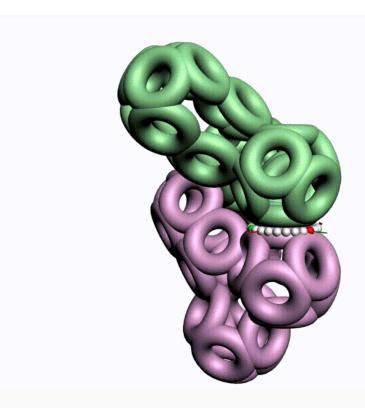
With best wishes, Alexander Kushelev, Head of Nanoworld Laboratory Tatyana Ryasina, Referent of Nanoworld Laboratory

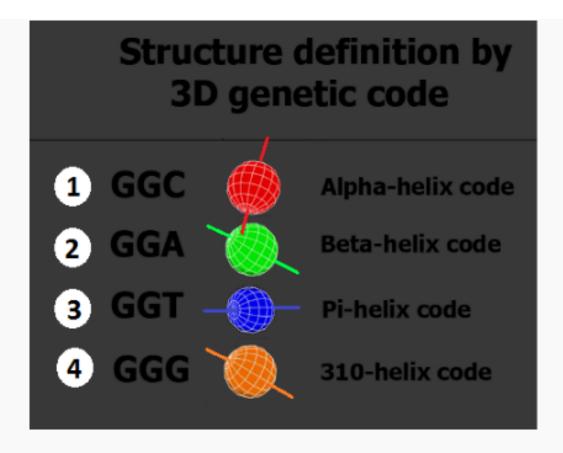
The Legend

HOW TO READ 2D DIAGRAMS OF 3D GENETIC CODE

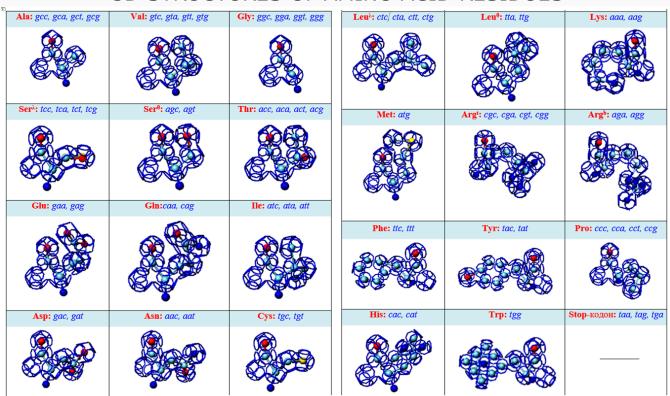


Structural template of amino acid residue

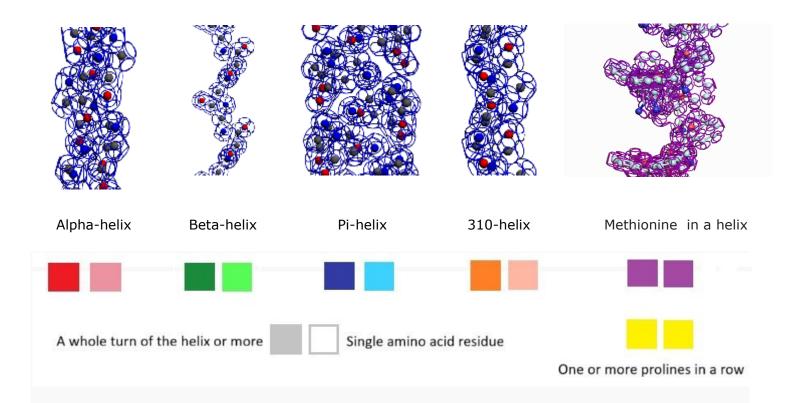




3D STRUCTURES OF AMINO ACID RESIDUES



Примечание. Ядра атомов кислорода показаны красным цветом, углерода — голубым, азота - синим, серы — желтым. Электроны внешних электронных оболочек атомов, формирующие молекулярную электронную оболочку аминокислот, обозначены синими кольцами. Для Arg, Leu и Ser представлены по две модели изомеров по положению радикала, которые кодируются по разному.



COMPACT 2D PICOTECHNOLOGY DIAGRAM

Red is an alpha helix.

The orange is a 310-helix.

Pink is a single alpha / 310 helix code.

Blue is a py-helix.

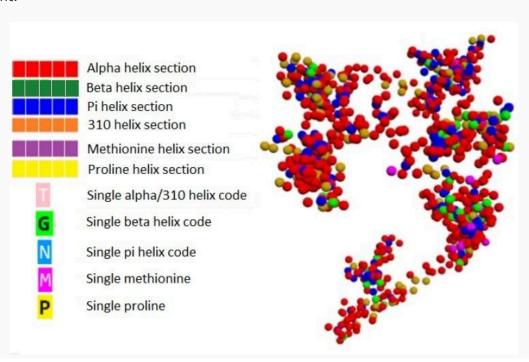
Green is a beta helix.

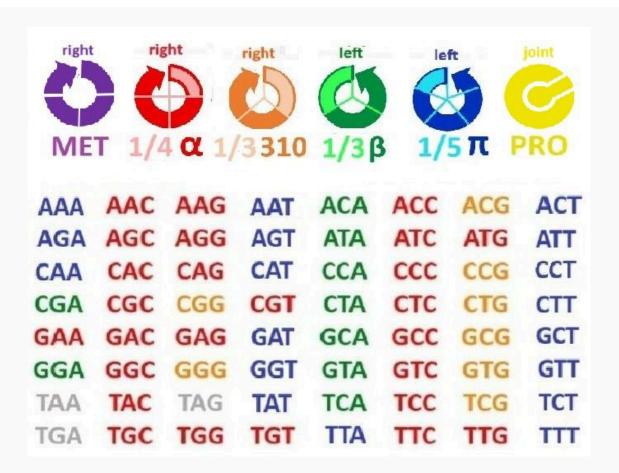
Lilac - methionine helix. It has a larger pitch of "thread" than the usual alpha-helix.

Black in abbreviated form and white in unfold means either an unknown code or the end of the translation.

Cyclic repetition of colors - software helix.

Alpha helix - right, Pi helix - left, Beta helix - left, 310 helix - right.



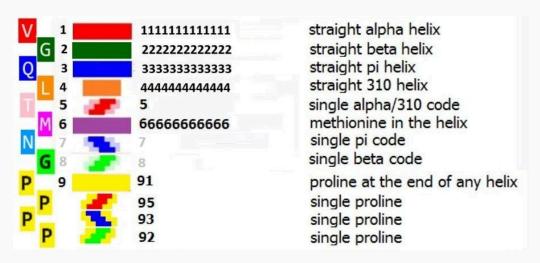


)	How d	oes t	he 3D	Gen	etic co	ode ta	ble wo	rk?		
				Sec	cond b	ase pos	sition				I
		U		С		<i>A</i>	1	G			
		UUU	Ρ π	UCU	S π	UAU	Υ π	UGU	Сπ	U	(44)
	U	UUC	Ρα	UCC	Sα	UAC	Υα	UGC	Cα	C	
		UUA	Lπ	UCA	Sβ	UAA	Stop	UGA	Stop	A	
		UUG	Lα	UCG	S 3 ₁₀	UAG	Stop	UGG	Wα	G	
_		CUU	Lπ	CCU	Ρ π	CAU	Η π	CGU	Rπ	U	_
io	C	CUC	Lα	CCC	Ρα	CAC	Η α	CGC	Rα	C	Third base position
OSi		CUA	Lβ	CCA	Ρβ	CAA	Qπ	CGA	Rβ	A	
O O		CUG	L 3 ₁₀	CCG	P 3 ₁₀	CAG	Qα	CGG	R 3 ₁₀	G	ер
First base position		AUU	Ι π	ACU	Τ π	AAU	Νπ	AGU	S π	U	oas
st b	A	AUC	Ια	ACC	Τα	AAC	Να	AGC	Sα	C	g g
Ë	A	AUA	Ιβ	ACA	Тβ	AAA	Κ π	AGA	Rπ	A	<u> </u>
		AUG	Μα	ACG	T 3 ₁₀	AAG	Κα	AGG	Rα	G	
		GUU	V π	GCU	Α π	GAU	D π	GGU→	$-G$ π	U	
	G	GUC	V α	GCC	Αα	GAC	Dα	GGC-	$-G/\alpha$	C	
	G	GUA	Vβ	GCA	Αβ	GAA	E	GGA-	Gβ	A	
		GUG	V3 ₁₀	GCG	A 3 ₁₀	GAG	\mathbf{E}/α	GGG	G 3 ₁₀	G	
Inp	β π 31	- 1/4 tur - 1/3 tur - 1/5 tur - 1/3 tu - 1/3 tu	rn of bet rn of pi- urn 3 ₁₀	ta helix (helix (pi -helix (3	beta tu -turn) i ₁₀ -turn	rn) G G G G G G G G G G G G G G G G G G G	GG	G G GA GGA			3

FULL (EXPANDED) DIAGRAM

- 1 ordinal number of the amino acid residue in the protein molecule
- 2 triplet code
- 3 one-letter code of amino acid residue
- 4 three-letter designation of amino acid residue
- 5 simplified composition code
- 6 graphical interpretation of a simplified composition code
- 7 composition code
- 8 graphic interpretation of the composition code
- 9 a note that sounds when you install this amino acid in a growing protein chain
- 10 graphic representation of a note (or percussion instrument) (Version Composition code 7)

3D Genetic code code Version 9



- coil of alpha helix (red)- 4 amino acid residues with the code "1",
- coil of beta-helix (green) 3 amino acid residues with the code "2",

310-helix coil (orange) - 3 amino acid residues with the code "4",

- coil of pi-helix (blue) 5 amino acid residues with the code "3"
- light gray stop codon

Program helices are a repetition of a sequence of compositions. For example, one alpha-helix code, then one pi-helix code. n (35) specifies a program spiral, and n3 or n5 are simple spirals (a pi-helix and an alpha-310 helix).

111111111111111111 - straight alpha helix

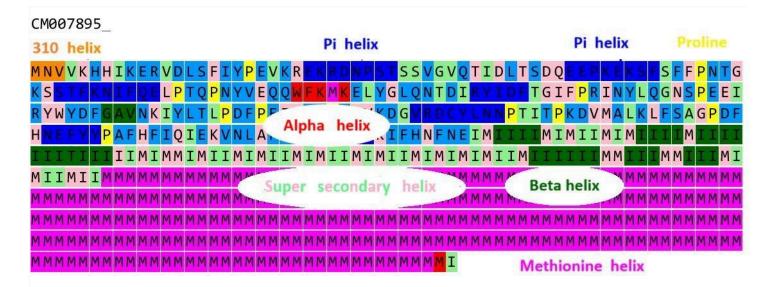
4444444444444444 - straight 310-helix

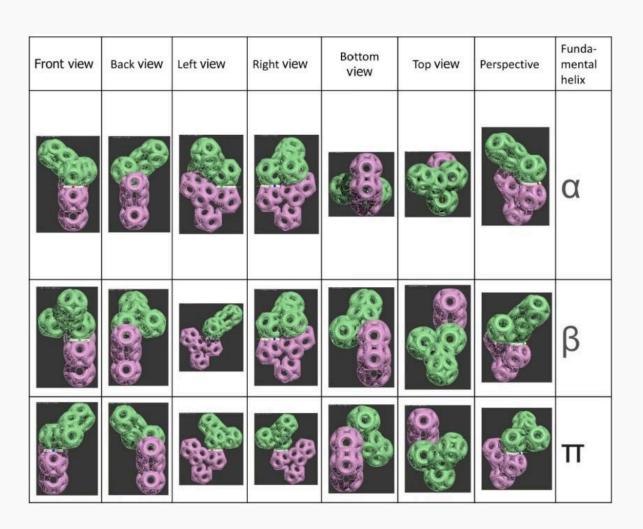
3333333333333333 - straight pi-helix

222222222222222 - direct beta helix

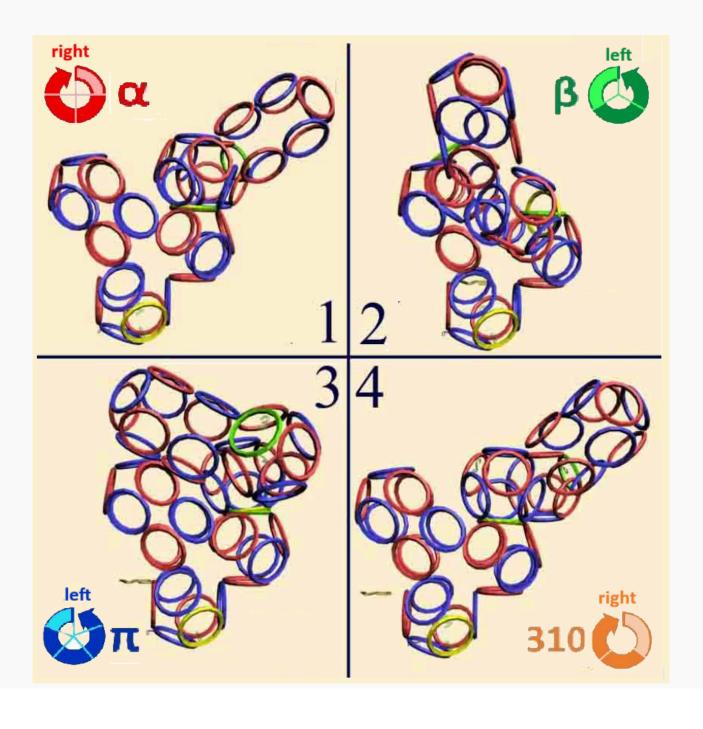
232323 - software 23-helix

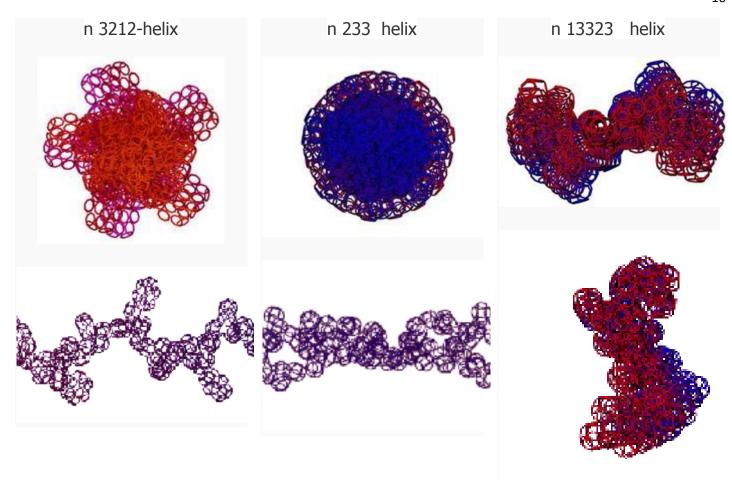
141414 - software 14-helix



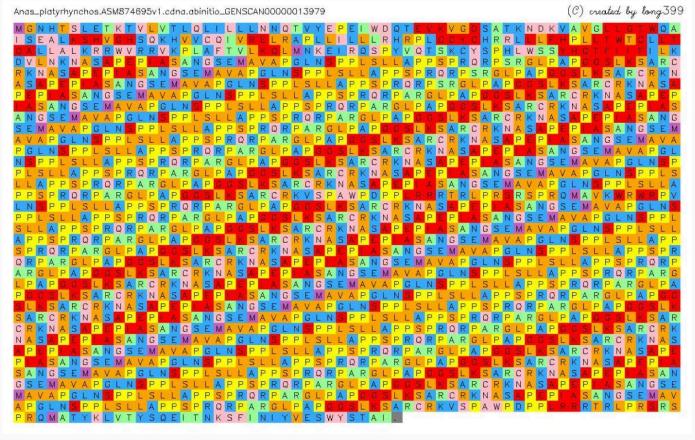


X	С	a.o.	Α	a.o.	T	a.o.	G	a.o.	Z	РВПС	ω
	ccc		CAC	Н	стс		CGC		С	R	35°
	CCA		CAA	Q	CTA		CGA		Α	0	155°
C	CCT	Р	CAT	Н	сп	L	CGT	R	Т	L	275°
	CCG		CAG	Q	CTG		CGG		G	R	35°
	ACC		AAC	N	ATC		AGC	S	С	R	35°
	ACA		AAA	K	ATA	1	AGA	R	Α	0	155°
Α	ACT	Т	AAT	N	ATT		AGT	S	T	L	275°
	ACG		AAG	K	ATG	M	AGG	R	G	R	35°
	TCC		TAC	Y	ΠC	F	TGC	С	С	R	35°
	TCA		TAA	Stop	TΤΑ	L	TGA	Stop	Α	0	155°
T	TCT	S	TAT	Y	тт	F	TGT	С	T	L	275°
	TCG		TAG	Stop	ΠG	L	TGG	W	G	R	35°
	GCC		GAC	D	GTC		GGC		С	R	35°
	GCA		GAA	Е	GTA		GGA		Α	0	155°
G	GCT	Α	GAT	D	GTT	V	GGT	G	T	L	275°
	GCG		GAG	Е	GTG		GGG		G	R	35°





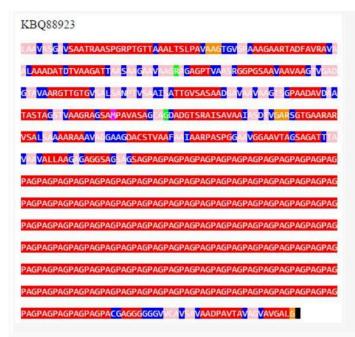
An example of a highly periodic structure

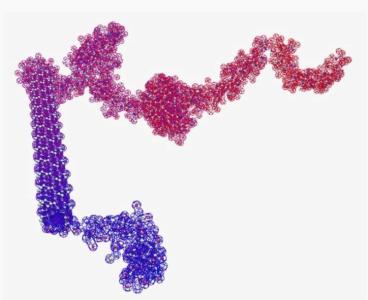


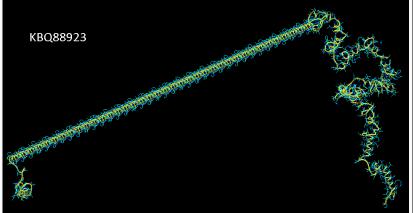
One-to-one correspondence between the sequence and 2D and 3D Picotech diagrams

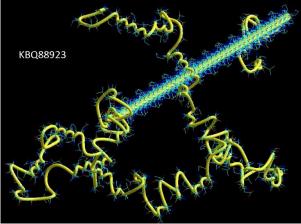
>ENA|KBQ88923|complement 2319 nucleotides

ttggccgccgttgccagcggtgccgttagcgccgccacccggccgccagccccggccgc cccaccggcaccaccgctgccgccttgacctccttgcccgccgttgccgccgggactggtgttgggcccgctgccgccggcgccgccgccgacttcgccgtccgcgccgttgcc gcccttgccgccgacgccaccgataccgtcgccgccggcgccaccactgccgcctct gccgccggtgccgccgttgccgccggccgagccggtgccggccccaccgttgctgccagc cgtggcggccccggcagccgccgttgccgccgtcgccggtaccgttggcgccgacggtaccgccgttgctgcccgcggcaccaccggcaccggcgtttccgccctttccgccaat cccaccgtttccgccgctatttccgctaccaccggcgtctccgccagcgccgccgatggcgccgttgccgccgttgccgccggtacctccggtcccgccgacgccgtcgatgccgct accgccagcaccgctgttccaccgttgccgccggccgcggttccgccatgcccgcc gtcgccagcggctggcggagatgccgacggcaccagccgcgccatctccgccgtc ggtgccgccgttggcggcgccgccgtcaccgccggttccgccggcgccaccactaccgcc ggttccgccggcccggcccggcccggcccggcccggcccg gccqqcccqqccqqcccqqccqqcccqqccqqcccqqccqqc cccqccqqcccqccqqcccqqcccqccqqcccqqcccqccqqcccqccqqcccqcc gccggccccgccggccccgccggccccgccggccccgccggc cccqccqqcccqccqqcccqqcccqccqqcccqqcccqccqqcccqccqqcccqcc gccggccccgccggccccgccggccccgccggccccgccggc ggcccgccggcccggcccggcccggcccggcccggcccggcccg gccqqcccqqccqqcccqqccqqcccqqccqqcccqqccqqc cccgccggccccgccggccccgccggccccgccggccccgcc cccgccggccccgccggccccgccggccccgccggccccgcc gccggccccgccggccccgccggccccgccggccccgccggc cccqccqqccccqccqqcccqqcccqccqqcccqqcccqqcccqqcccq ggcccgccggcccggcccggcccggcccggcccggcccggcccg accgccgttgccggcgttgccgtcggcgccttggggtga

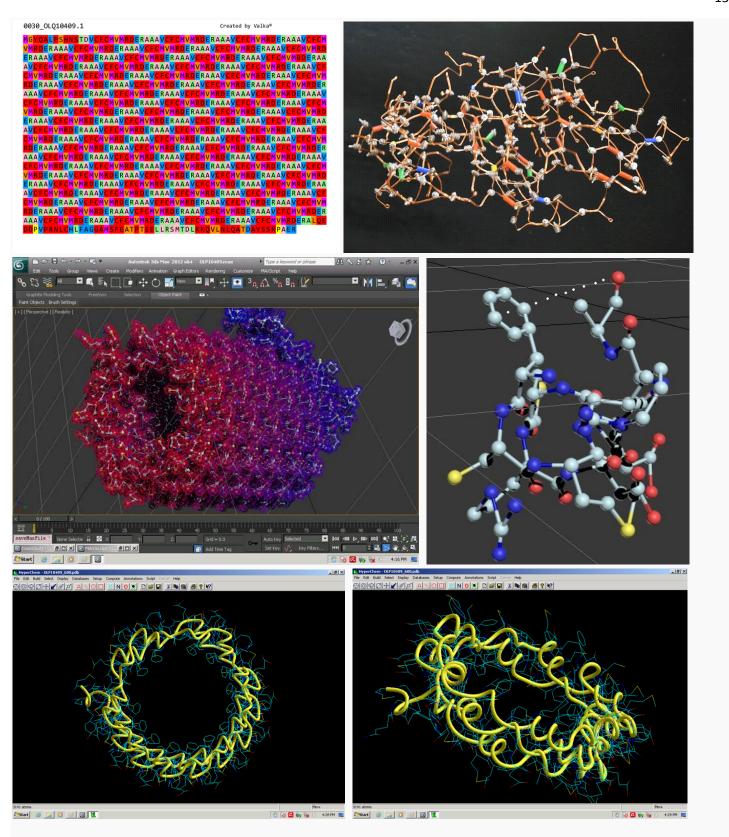








While the 3D picotech program is in development, we can model proteins manually and then digitize them to deliver the result as a 3D images and .pdb files.



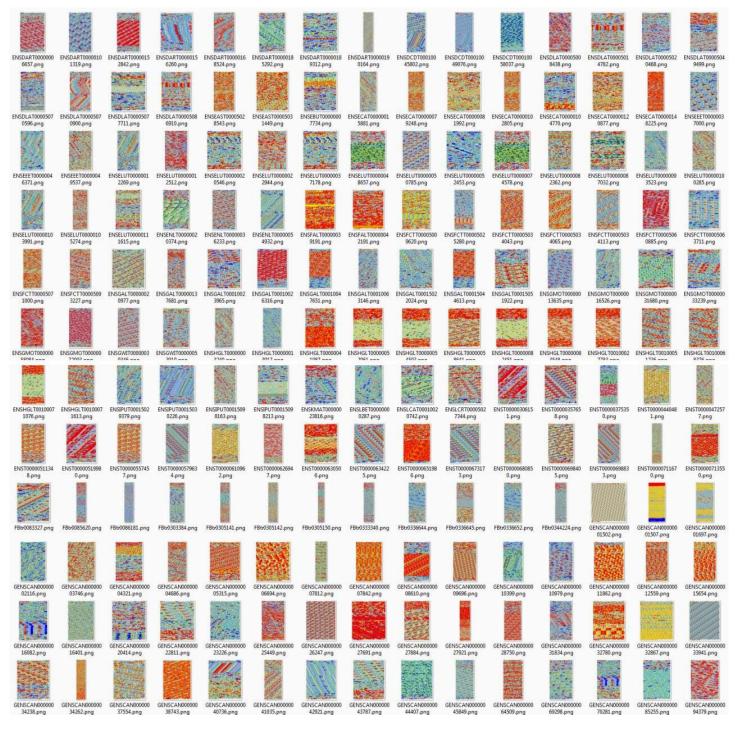
3D-Genetic code

1992

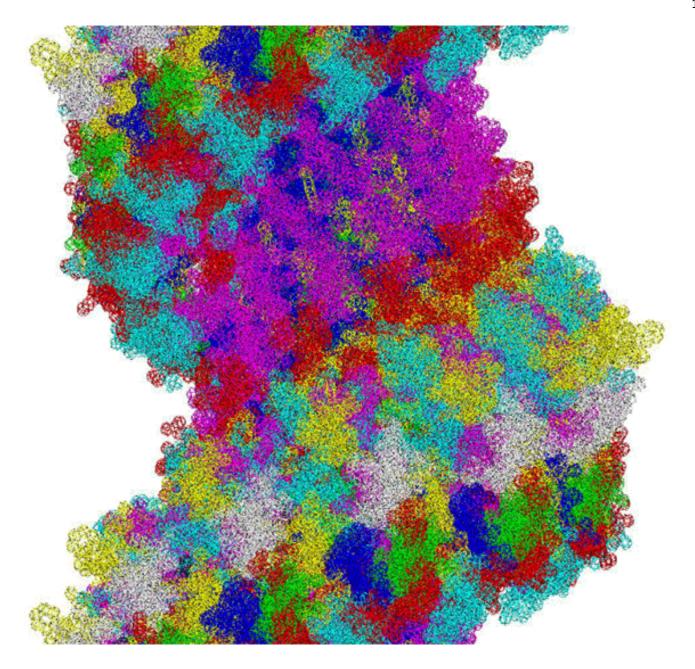
Код	Остаток	Вариант
AAA	Lys	3
AAC	Asn	1
AAG	Lys	1
AAAA AAC AAG AAC ACA AGC AGG AGT ATA ATC ATG ATT CAA ACC CAC CCC CCG CCT CCGA CCT CCG CCT CCGA CCT CCG CCG	Lys Asn Lys Asn Thr	1 3 2 1 4 3 3 1 1 1 3
ACA	The	2
ACA ACC	TI	1
ACC	Thr	1
ACG	Thr Thr Arg Ser Arg	4
ACT	Thr	3
AGA	Arg	3
AGC	Ser	1
ACC	Ass	1
AGG	Arg	1
AGT	Ser Ile Ile Met Ile Gln His	3
ATA	Ile	2
ATC	Ile	1
ATG	Met	1
ATT	Ile	3
ALL	ne	2
CAA	Gln	3
CAC	His	1
CAG	Gln	1
САТ	His	3
CC	Dec	2
CCA	PTO	Z
CCC	His Pro Pro Pro Pro Arg Arg Arg Leu	3 3 1 1 3 2 1 4 4 3 2 1 4 4 3 2 2 1
CCG	Pro	4
CCT	Pro	3
CCV	Ara	2
CCC	A-	1
CGC	Arg	1
CGG	Arg	4
CGT	Arg	3
СТА	Leu	2
CIA	T -	1
CIC	Leu Leu Leu	1
CTG	Leu	4
CTT	Leu	3
GΔΔ	Glu	3
CAC	Glu Asp	1
UAC	Asp	4 3 3 1
GAG	Glu Asp	I
GAT	Asp	3
GCA	Ala	2
GCC	Ala	1
CCC	Ala Ala Ala Ala Gly	3 2 1 4 3 2 1 4 3 2 1 4 3 2
oCG o	Ala	4
GCT	Ala	3
GGA	Gly	2
GGC	Gly	1
GGC	Gly	4
Cem	Ch	2
ool'	GIY	3
GTA	Gly Val	2
GGT GTA GTC	Val	1
GTG		4
		3
GTT	Val	,
TAA	TKD	
TAC	Tyr	1
TAG	TKD	
TAT	Tyr	3
TCA	Ser	2
TCC	Ser	1
TCG	Ser	4
TCT		3
	TED	
TGA	TKD	
TGC	Cys	1
TGG	Ттр	1
TGT	Cys	3
	Leu	3
400	Phe	1
	Leu	1
TTG		

2019 9 var **3** DNA AMI Nº ami Score AAA Κ LYS F#1 5 5 3 2 5 5 H1 F#1 2 AAC Ν ASN \frac{2}{2} 3 AAG Κ LYS AAT ASN 4 Ν H1 E2 E2 E2 E2 D1 A2 D1 A2 C2 G1 A1 A3 A3 A3 A3 D1 D1 D1 5 ACA T T T THR ACC ACG THR 6 7 THR 8 ACT Т THR 3 5 AGA R 9 ARG AGC s 10 SER 11 AGG R ARG 5 3 2 5 6 3 3 5 5 S AGT SER 12 13 ATA ILE ATC ILE 14 ı 15 ATG MET М 16 ATT ILE 17 CAA Q GLN CAC Н HIS 18 19 CAG Q GLN HPP CAT 3 20 HIS 92 91 PRO 21 CCC 22 **PRO** 23 CCG Р PRO 95 93 2 5 5 3 2 5 5 3 P 24 CCT PRO 25 CGA R ARG CGC R 26 ARG 27 CGG R ARG 28 CGT R ARG 29 CTA L LEU H1 H1 H1 H1 CTC 30 LEU CTG L L E LEU 31 LEU CTT 32 355325532553255 G1 H1 G1 H1 A3 A3 A3 G0 G0 G0 G2 G2 G2 G2 G2 33 GAA GLU 34 GAC D ASP GAG E GLU 35 D 36 GAT ASP GCA Α ALA ALA 37 GCC Α 38 Α 39 GCG ALA GCT GGA ALA GLY Α 40 G 41 G 42 GGC GLY GGV 43 GGG GLY 44 GGT GLY 45 **GTA** VAL VAL VAL V 46 GTC 47 GTG ٧ 48 GTT ٧ VAL 3 49 TAA TKD 0 5 0 3 2 5 5 D1 50 TAC Υ TYR 51 TAG TKD Y S TYR D1 A2 A2 A2 A2 A2 52 TAT TCA SER 53 54 TCC s SER s 55 TCG SER 56 TCT S SER TGA 57 TKD 0 5 5 E2 D1 E2 H1 G1 H1 С 58 TGC CYS W 59 TGG TRP 60 TGT С CYS LEU L F 61 TTA 62 TTC 63 **TTG** L LEU

The result of processing 62 genomes (approximately 6 million proteins) was obtained in an hour '2025



3D MODEL OF COLLAGEN

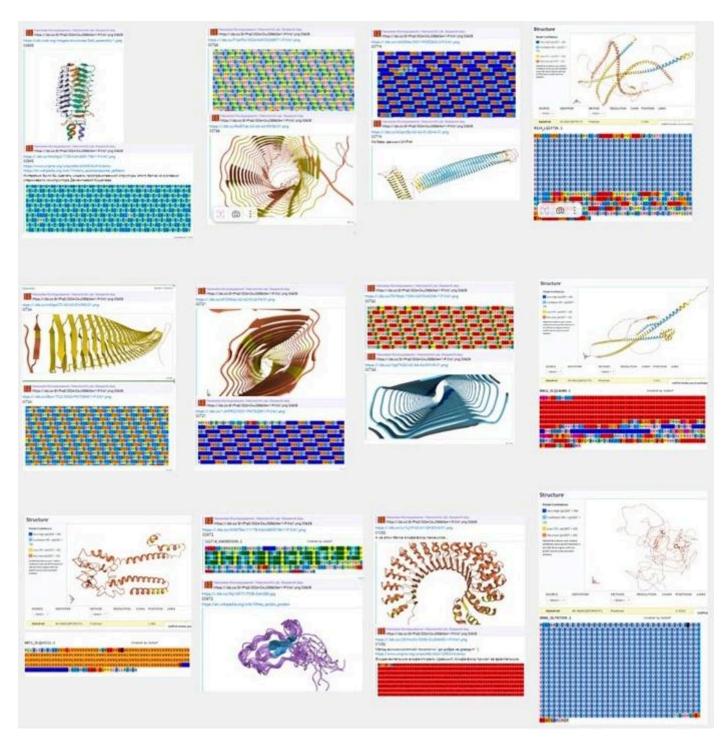


Software helices
Fractal helices
Q-helices

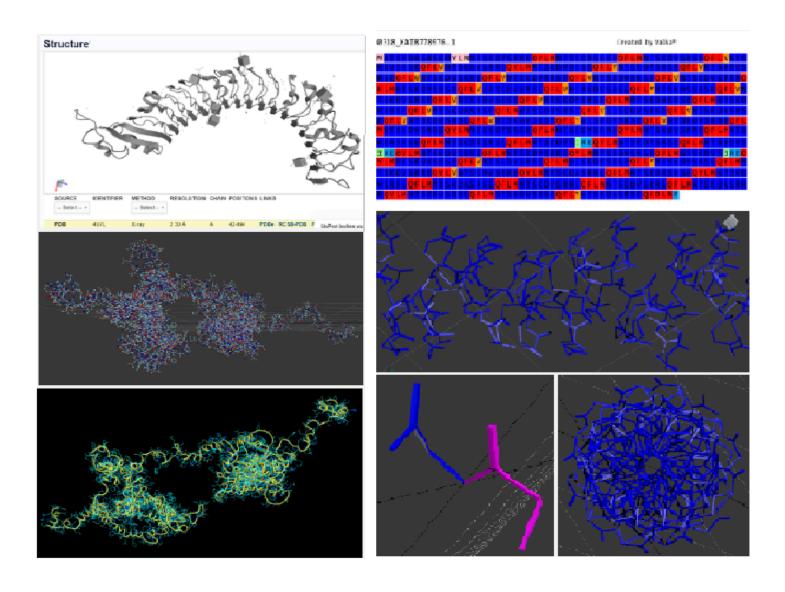
X-ray diffraction and 3D Genetic code

Examples of super-secondary structures

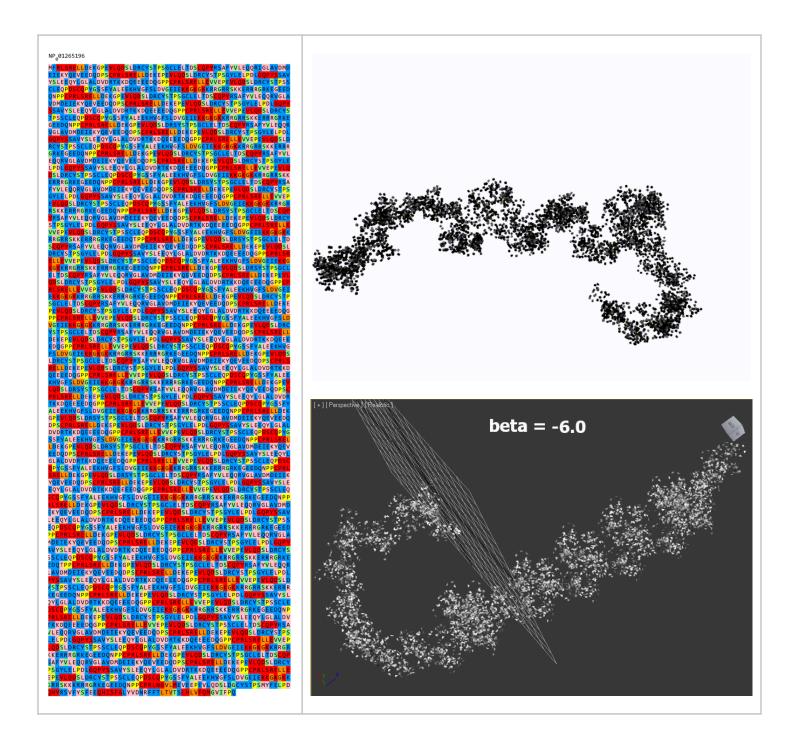
"Wet" experiments showed... fractal helices (helices of helices)

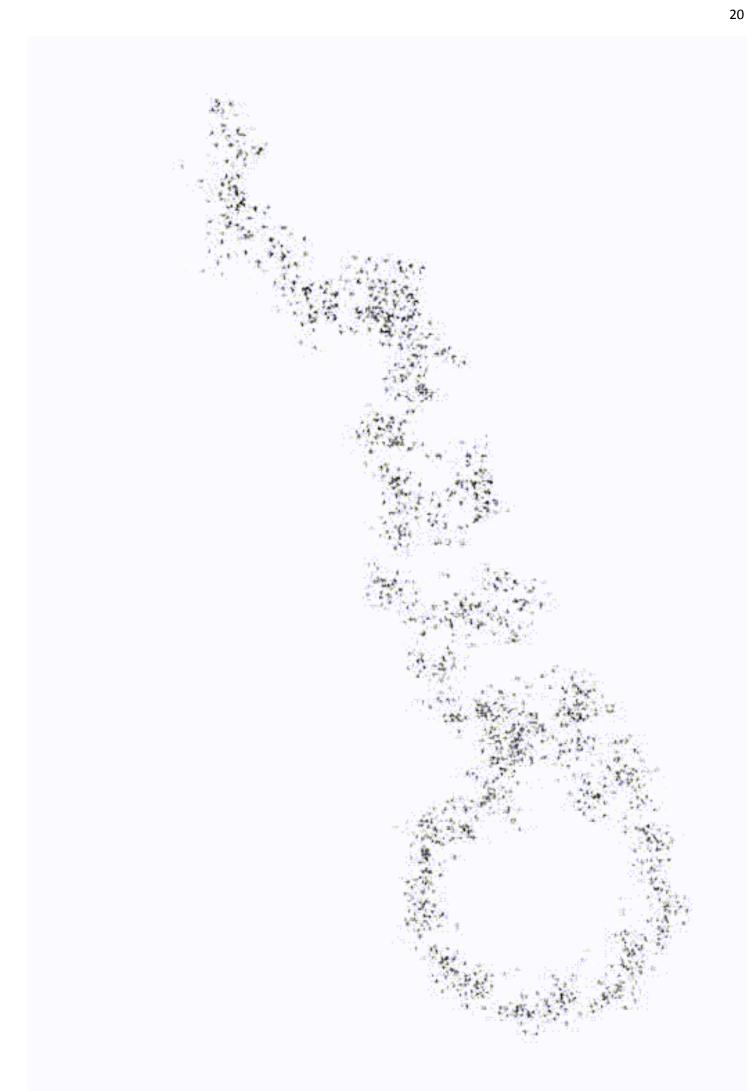


2 3

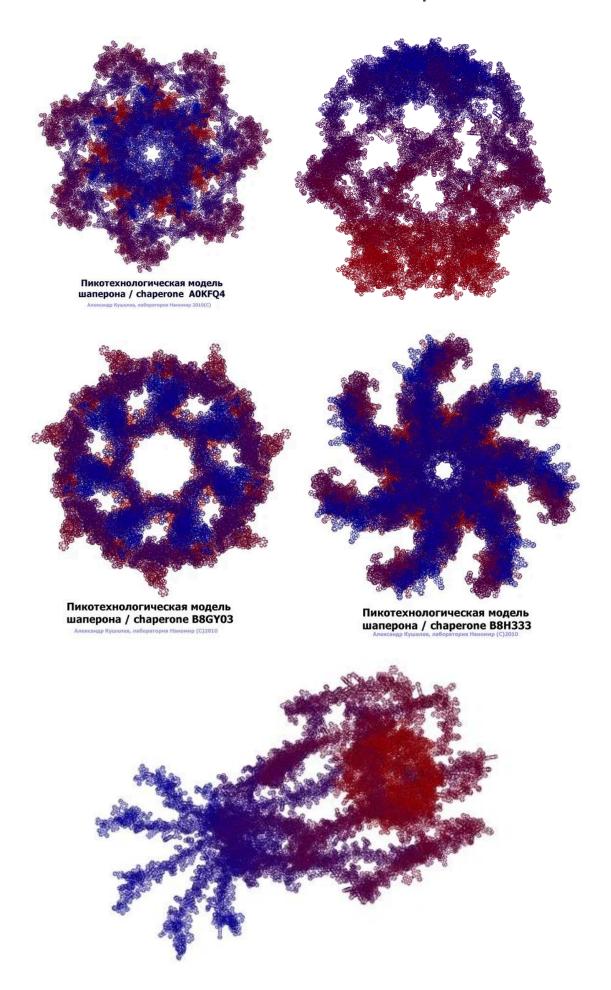


Super-secondary structure of the first chromosome (neuroblastoma) and its 3D image

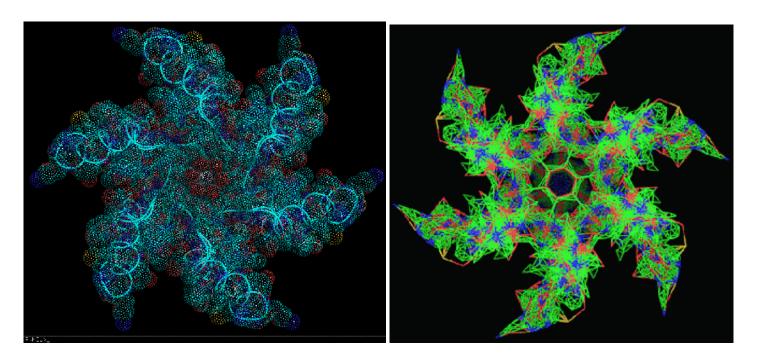




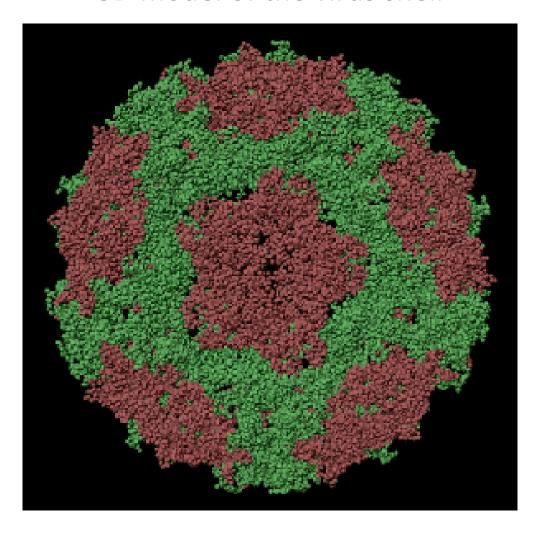
3D Picotech models of chaperones



3D model of insulin molecule



3D model of the virus shell



How exactly does this service work?

You send us the coding nucleotide sequence (ideally - in fasta format) of the protein you're interested in via email: nanoworld laboratory@mail.ru.

The key advantages are:

1. Reliability and accuracy of protein secondary structure.

Below the 3D genetic code table, you can see a transcript showing that each triplet of the genetic code corresponds to a specific structure.

- 2. The size of the protein is irrelevant.
- 3. The ability of the protein to crystallize is irrelevant.
- 4. The presence/absence of homologues is irrelevant.
- 5. The degree of study is irrelevant.

The disadvantages of this method include:

- 1. It does not account for protein post-processing after ribosome synthesis. The method only identifies the structure resulting from ribosome synthesis. If the protein is subsequently processed by chaperones or enzymes, meaning its secondary structure is altered (fragments are excised), I can only account for this manually by removing the enzyme-excised fragments of the amino acid sequence from the processing results. The client can do this themselves or provide me with data on the excised protein fragments.
- 2. Only secondary structures are automated. Tertiary structure can be automatically determined only for a narrow class of proteins, such as those consisting of a straight section of any type of helix. Therefore, only the secondary structure is guaranteed.

Tertiary structure and higher structures are "manual work," which can not only take longer but also does not guarantee its determination, for example, if the amino acid sequence contains prolines, which transform the rigid protein structure into a mechanism that can significantly change its conformation. In this case, the client can be provided with partial information on the tertiary and higher structures.

However, there is a high probability of determining both the tertiary and quaternary structures, <u>as well as the protein dynamics</u>

What exactly do clients receive when they contact us?

In what format are the results provided?

- 1. The client is guaranteed to receive a 2D Picotech diagram of the protein secondary structure.
- 2. With a certain degree of probability, the client will be able to obtain a 3D model from the joint-rod constructor.

3. If the 3D model is generated automatically or manually, the client will also be able to obtain a standard .pdb file with the coordinates of all protein atoms.

Unlike X-ray diffraction, the new technology has no "blind spots," meaning the PDB file will contain the coordinates of all atoms, from the first to the last amino acid residue.

Also, the client can obtain a <u>dynamic protein model</u>, which provides insight into how the protein performs its functions.

Additionally, clients can receive a consultation in which we demonstrate the correlation between different analysis methods and our data. For example, X-ray diffraction (XRD) or alpha-fold analysis reveal the presence of helical regions, but do not specify their type. We can demonstrate where these methods correctly identified the beginning and end of a helical region, and where they erred, and why. In particular, interpreters often mistake hybrid (alpha-310-) regions of proteins for beta sheets. Picotechnological models can help clients understand why this occurs.

Moreover, we can search for protein active sites using the natural frequencies of amino acid side chains $\underline{1}$, $\underline{2}$, $\underline{3}$ (pp. 29-42). If any are detected, we can attempt to model the mechanism of these active sites at the picomechanical level, i.e., with picometric accuracy.

Thank you very much for your lenient attention