

# Class 9: Structural Bioinfomatics (Pt. 1)

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## The PDB database

The main database for structural biology is called the PDB. Let's have a look at what it contains:

Download a CSV file from the PDB site (accessible from "Analyze" > "PDB Statistics" > "by Experimental Method and Molecular Type".

```
stats <- read.csv("Data Export Summary.csv")
stats
```

	Molecular.Type	X.ray	EM	NMR	Integrative	Multiple.methods
1	Protein (only)	176,204	20,299	12,708	342	218
2	Protein/Oligosaccharide	10,279	3,385	34	8	11
3	Protein/NA	9,007	5,897	287	24	7
4	Nucleic acid (only)	3,066	200	1,553	2	15
5	Other	173	13	33	3	0
6	Oligosaccharide (only)	11	0	6	0	1
	Neutron	Other	Total			
1	83	32	209,886			
2	1	0	13,718			
3	0	0	15,222			
4	3	1	4,840			
5	0	0	222			
6	0	4	22			

```
stats$Total
```

```
[1] "209,886" "13,718" "15,222" "4,840" "222" "22"
```

Oh, these are characters not numeric...

```
library(readr)

stats <- read_csv("Data Export Summary.csv")
stats
```

	Molecular Type	X-ray	EM	NMR	Integrative	Multiple methods	Neutron
1	Protein (only)	176204	20299	12708	342	218	83
2	Protein/Oligosacch...	10279	3385	34	8	11	1
3	Protein/NA	9007	5897	287	24	7	0
4	Nucleic acid (only)	3066	200	1553	2	15	3
5	Other	173	13	33	3	0	0
6	Oligosaccharide (o...	11	0	6	0	1	0

```
n.total <- sum(stats$Total)
```

Q1: What percentage of structures in the PDB are solved by X-Ray and Electron Microscopy. Give your answers to 2 significant figures.

```
n.xray <- sum(stats`X-ray`)
percent.xray <- n.xray / n.total * 100
percent.xray
```

[1] 81.48087

There are 81.48 percent X-ray structures in the PDB.

```
n.em <- sum(stats`EM`)
percent.em <- n.em / n.total * 100
percent.em
```

[1] 12.21516

There are 12.22 percent EM structures in the PDB.

Q2: What proportion of structures in the PDB are protein?

```
round(stats$Total[1]/n.total * 100, 2)
```

[1] 86.05

## Exploring PDB structures

Package for structural bioinformatics

```
library(bio3d)
hiv <- read.pdb("1hsg")
```

Note: Accessing on-line PDB file

hiv

Call: read.pdb(file = "1hsg")

```
Total Models#: 1
Total Atoms#: 1686, XYZs#: 5058 Chains#: 2 (values: A B)

Protein Atoms#: 1514 (residues/Calpha atoms#: 198)
Nucleic acid Atoms#: 0 (residues/phosphate atoms#: 0)
```

Non-protein/nucleic Atoms#: 172 (residues: 128)

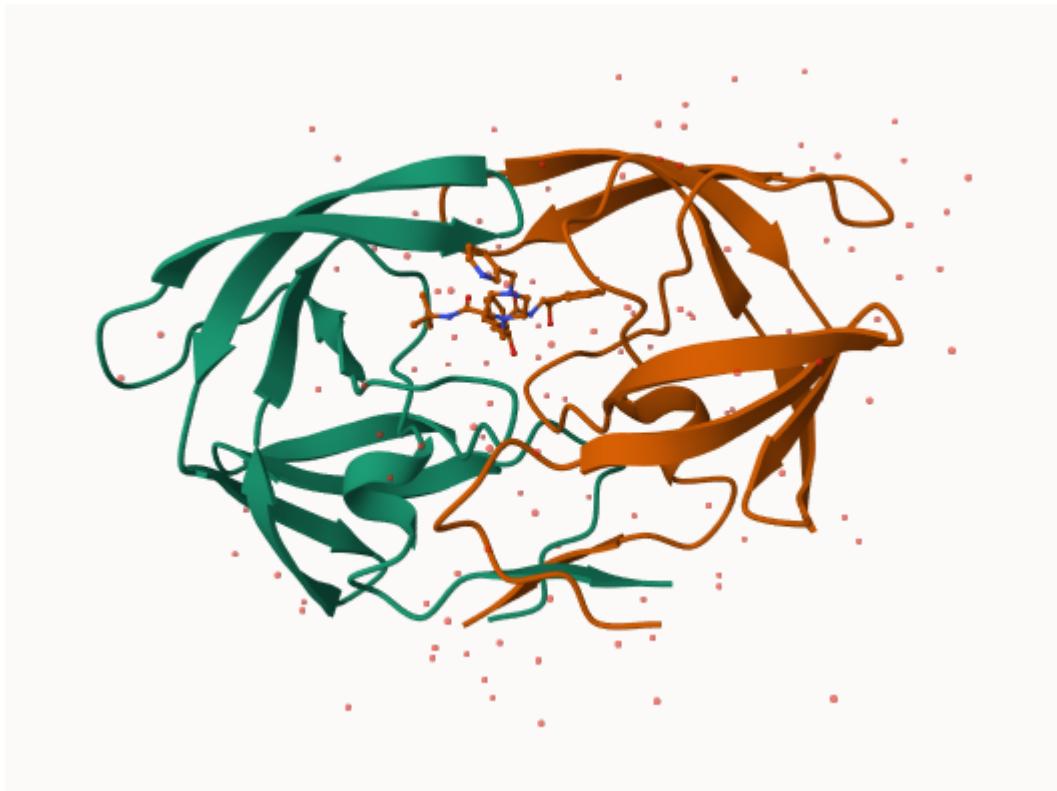
Non-protein/nucleic resid values: [ HOH (127), MK1 (1) ]

Protein sequence:

```
PQITLWQRPLVTIKIGGQLKEALLDTGADDTVLEEMS LPGRWKP K MIGGGIGGF IKVRQYD  
QILIEICGHKAIGTVLVGPTPVNIIGRNLLTQIGCTLNFPQITLWQRPLVTIKIGGQLKE  
ALLDTGADDTVLEEMSLPGRWKP K MIGGGIGGF IKVRQYDQILIEICGHKAIGTVLVGPTP  
VNIIGRNLLTQIGCTLNF
```

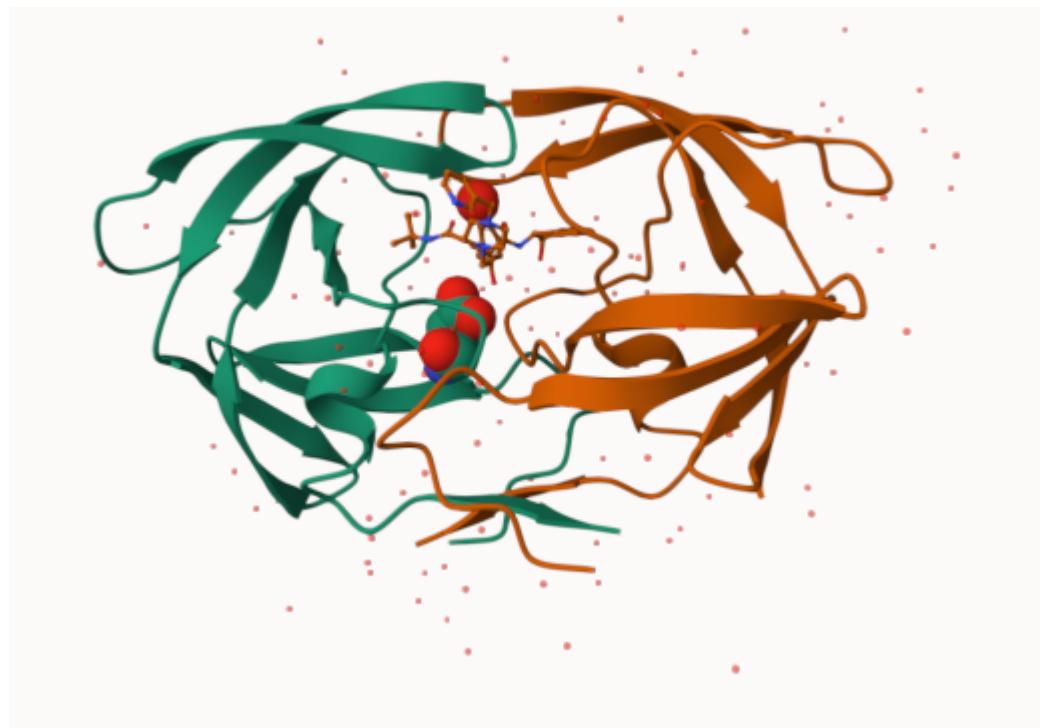
+ attr: atom, xyz, seqres, helix, sheet,  
calpha, remark, call

Let's first use the Mol\* viewer to explore this structure.



My first view of HIV-Pr

And a view of the ligand (ball and stick) with catalytic ASP 25 amino-acids (spacefill) and the all important active site water molecule:



My first view of HIV-Pr

## PDB objects in R

```
head(hiv$atom)
```

	type	eleno	elety	alt	resid	chain	resno	insert	x	y	z	o	b
1	ATOM	1	N	<NA>	PRO	A	1	<NA>	29.361	39.686	5.862	1	38.10
2	ATOM	2	CA	<NA>	PRO	A	1	<NA>	30.307	38.663	5.319	1	40.62
3	ATOM	3	C	<NA>	PRO	A	1	<NA>	29.760	38.071	4.022	1	42.64
4	ATOM	4	O	<NA>	PRO	A	1	<NA>	28.600	38.302	3.676	1	43.40
5	ATOM	5	CB	<NA>	PRO	A	1	<NA>	30.508	37.541	6.342	1	37.87
6	ATOM	6	CG	<NA>	PRO	A	1	<NA>	29.296	37.591	7.162	1	38.40
	segid	elesy	charge										
1	<NA>	N	<NA>										
2	<NA>	C	<NA>										
3	<NA>	C	<NA>										
4	<NA>	O	<NA>										
5	<NA>	C	<NA>										
6	<NA>	C	<NA>										

```
pdbseq(hiv)
```

1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20
"P"	"Q"	"I"	"T"	"L"	"W"	"Q"	"R"	"P"	"L"	"V"	"T"	"I"	"K"	"I"	"G"	"G"	"Q"	"L"	"K"
21	22	23	24	25	26	27	28	29	30	31	32	33	34	35	36	37	38	39	40
"E"	"A"	"L"	"L"	"D"	"T"	"G"	"A"	"D"	"D"	"T"	"V"	"L"	"E"	"E"	"M"	"S"	"L"	"P"	"G"
41	42	43	44	45	46	47	48	49	50	51	52	53	54	55	56	57	58	59	60
"R"	"W"	"K"	"P"	"K"	"M"	"I"	"G"	"G"	"I"	"G"	"G"	"F"	"I"	"K"	"V"	"R"	"Q"	"Y"	"D"
61	62	63	64	65	66	67	68	69	70	71	72	73	74	75	76	77	78	79	80
"Q"	"I"	"L"	"I"	"E"	"I"	"C"	"G"	"H"	"K"	"A"	"I"	"G"	"T"	"V"	"L"	"V"	"G"	"P"	"T"
81	82	83	84	85	86	87	88	89	90	91	92	93	94	95	96	97	98	99	1

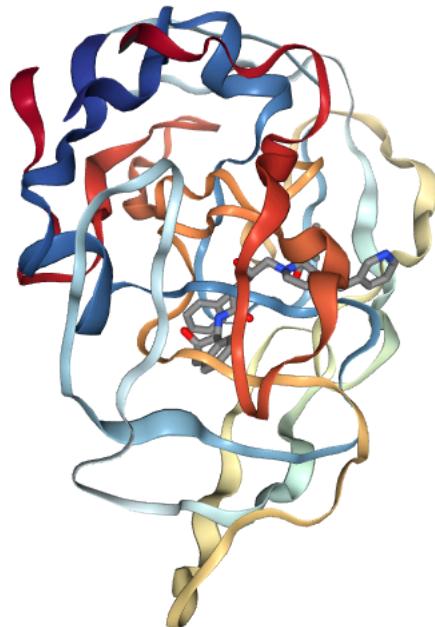
```
"P" "V" "N" "I" "I" "G" "R" "N" "L" "L" "T" "Q" "I" "G" "C" "T" "L" "N" "F" "P"
  2   3   4   5   6   7   8   9   10  11  12  13  14  15  16  17  18  19  20  21
"Q" "I" "T" "L" "W" "Q" "R" "P" "L" "V" "T" "I" "K" "I" "G" "G" "Q" "L" "K" "E"
 22  23  24  25  26  27  28  29  30  31  32  33  34  35  36  37  38  39  40  41
"A" "L" "L" "D" "T" "G" "A" "D" "D" "T" "V" "L" "E" "E" "M" "S" "L" "P" "G" "R"
 42  43  44  45  46  47  48  49  50  51  52  53  54  55  56  57  58  59  60  61
"W" "K" "P" "K" "M" "I" "G" "G" "I" "G" "F" "I" "K" "V" "R" "Q" "Y" "D" "Q"
 62  63  64  65  66  67  68  69  70  71  72  73  74  75  76  77  78  79  80  81
"I" "L" "I" "E" "I" "C" "G" "H" "K" "A" "I" "G" "T" "V" "L" "V" "G" "P" "T" "P"
 82  83  84  85  86  87  88  89  90  91  92  93  94  95  96  97  98  99
"V" "N" "I" "I" "G" "R" "N" "L" "L" "T" "Q" "I" "G" "C" "T" "L" "N" "F"
```

```
chainA_seq <- pdbseq( trim.pdb(hiv, chain="A") )
```

I can interactively view these PDB objects in R with the new **bio3dview** package. This is not yet on CRAN.

To install this I can setup **pak** package and use it to install **bio3dview** from GitHub. In my console I first run `install.packages("pak") pak::pak("bioboot/bio3dview")`

```
library(bio3dview)
view.pdb(hiv)
```

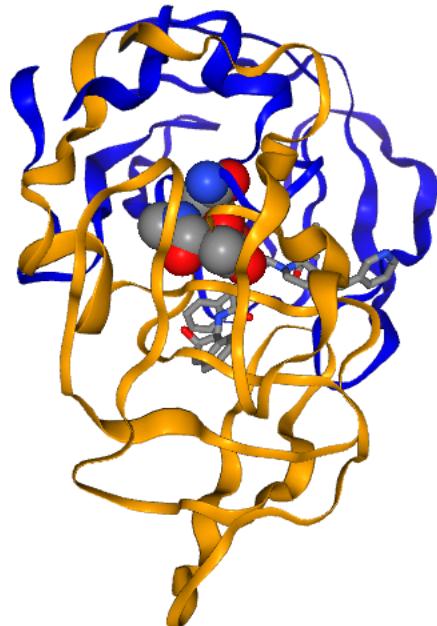


Change some settings

```
sel <- atom.select(hiv, resno = 25)

view.pdb(hiv, highlight = sel,
         highlight.style = "spacefill",
         colorScheme = "chain",
```

```
col = c("blue", "orange"),
backgroundColor = "pink")
```



## Predict protein flexibility

We can run a bioinformatics calculation to predict protein dynamics - i.e. functional motions.

We will use the `nma()` function:

```
adk <- read.pdb("6s36")
```

Note: Accessing on-line PDB file  
PDB has ALT records, taking A only, rm.alt=TRUE

```
adk
```

```
Call: read.pdb(file = "6s36")
```

```
Total Models#: 1
Total Atoms#: 1898, XYZs#: 5694 Chains#: 1 (values: A)
```

```
Protein Atoms#: 1654 (residues/Calpha atoms#: 214)
Nucleic acid Atoms#: 0 (residues/phosphate atoms#: 0)
```

```
Non-protein/nucleic Atoms#: 244 (residues: 244)
Non-protein/nucleic resid values: [ CL (3), HOH (238), MG (2), NA (1) ]
```

Protein sequence:

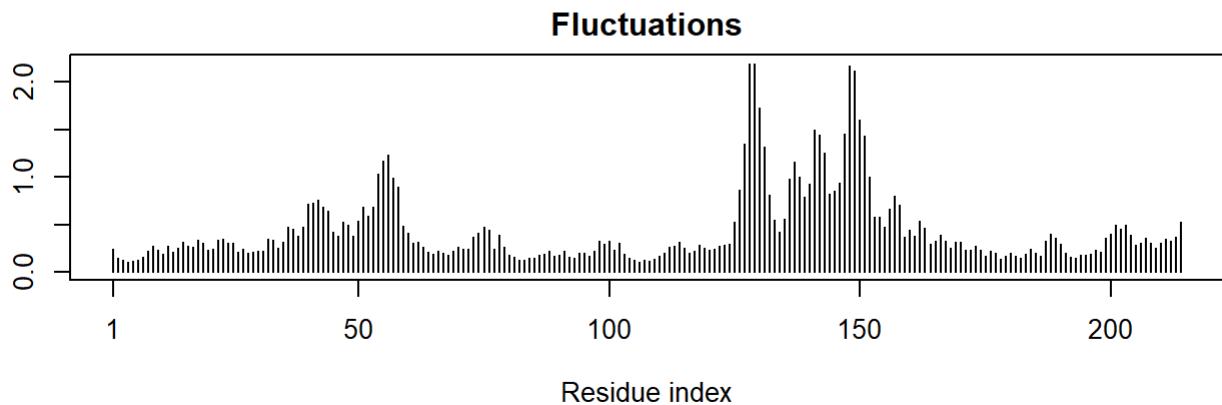
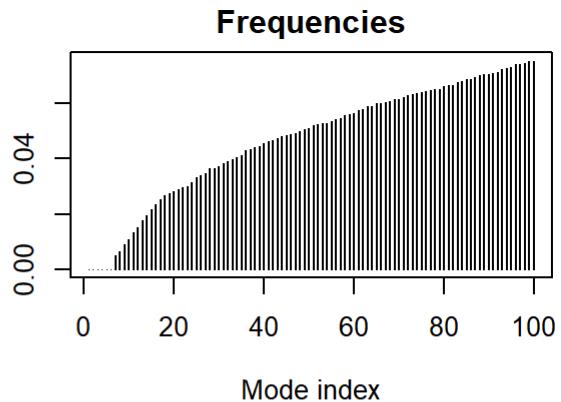
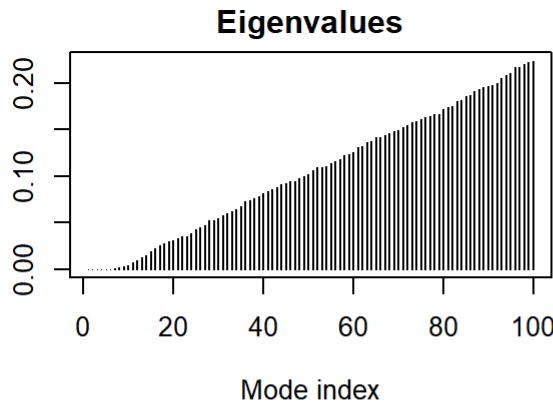
```
MRIILLGAPGAGKGTQAQFIMEKYGIPQISTGDMRAAVKSGSELGKQAKDIMDAGKLVT
DELVIALVKERIAQEDCRNGFLLDGFPRTIPQADAMKEAGINVVDYLEFDVPDELIVDKI
VGRRVHAPSGRVYHVFKNPPKVEGKDDVTGEELTRKDDQEETVRKRLVEYHQMTAPLIG
YYSKAEAGNTKYAKVDGTPVAEVRADLEKILG
```

+ attr: atom, xyz, seqres, helix, sheet,  
calpha, remark, call

```
m <- nma(adk)
```

Building Hessian... Done in 0.03 seconds.  
Diagonalizing Hessian... Done in 0.28 seconds.

```
plot(m)
```



Generate a "trajectory" of predicted motion

```
mktrj(m, file="ADK_nma.pdb")
```

```
view.nma(m)
```

