



Model Building Report

This document lists the results for the homology modelling project "Hemoglobin a" submitted to SWISS-MODEL workspace on April 24, 2018, 9:05 a.m.. The submitted primary amino acid sequence is given in Table T1.

If you use any results in your research, please cite the relevant publications:

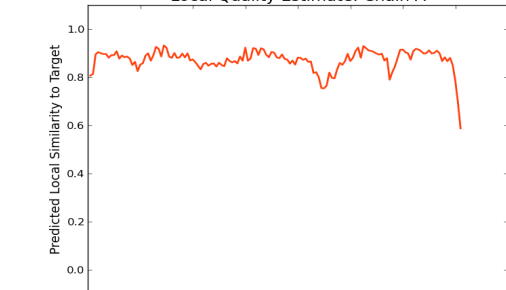
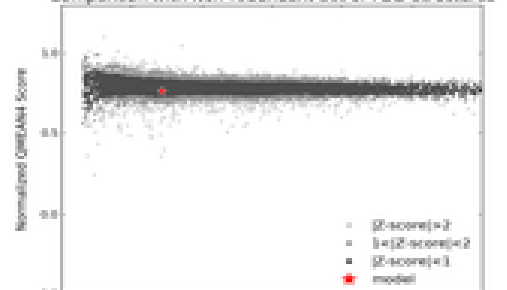
- Biasini, M., Bienert, S., Waterhouse, A., Arnold, K., Studer, G., Schmidt, T., Kiefer, F., Cassarino, T.G., Bertoni, M., Bordoli, L., Schwede, T. SWISS-MODEL: modelling protein tertiary and quaternary structure using evolutionary information. *Nucleic Acids Res.* 42, W252-W258 (2014). [\[a\] doi>](#)
- Guex, N., Peitsch, M.C., Schwede, T. Automated comparative protein structure modeling with SWISS-MODEL and Swiss-PdbViewer: A historical perspective. *Electrophoresis* 30, S162-S173 (2009). [\[a\] doi>](#)
- Bienert, S., Waterhouse, A., de Beer, T.A., Tauterilo, G., Studer, G., Bordoli, L., Schwede, T. The SWISS-MODEL Repository - new features and functionality. *Nucleic Acids Res.* 45, D313-D319 (2017). [\[a\] doi>](#)
- Benkert, P., Biasini, M., Schwede, T. Toward the estimation of the absolute quality of individual protein structure models. *Bioinformatics* 27, 343-350 (2011). [\[a\] doi>](#)
- Bertoni, M., Kiefer, F., Biasini, M., Bordoli, L., Schwede, T. Modeling protein quaternary structure of homo- and hetero-oligomers beyond binary interactions by homology. *Scientific Reports* 7 (2017). [\[a\] doi>](#)

Results

The SWISS-MODEL template library (SMTL version 2018-04-18, PDB release 2018-04-13) was searched with BLAST ([Camacho et al.](#)) and HHblits ([Remmert et al.](#)) for evolutionary related structures matching the target sequence in Table T1. For details on the template search, see Materials and Methods. Overall 1337 templates were found (Table T2).

Models

The following model was built (see Materials and Methods "Model Building"):

Model #01	File	Built with	Oligo-State	Ligands		GMQE	QMEAN		
	PDB	ProMod3 Version 1.1.0.	monomer	1 x HEM: PROTOPORPHYRIN IX CONTAINING FE;		0.98	-0.41		
<div><div><div>QMEAN</div><div>Cβ</div><div>All Atom</div><div>Solvation</div><div>Torsion</div></div><div><div>-0.41</div><div>-0.58</div><div>1.20</div><div>2.10</div><div>-0.80</div></div><div><div></div><div></div><div></div><div></div><div></div></div></div>									
<div><div>Local Quality Estimate: Chain A</div><div></div></div> <div><div>Comparison with Non-redundant Set of PDB Structures</div><div></div></div>									
Template	Seq Identity	Oligo-state	Found by	Method	Resolution	Seq Similarity	Range	Coverage	Description
1bz1.1.A	100.00	monomer	HHblits	X-ray	1.59Å	0.60	1 - 142	1.00	PROTEIN (HEMOGLOBIN ALPHA CHAIN)
Ligand	Added to Model			Description					
HEM	✓			PROTOPORPHYRIN IX CONTAINING FE					
HEM	✗ - Binding site not conserved.			PROTOPORPHYRIN IX CONTAINING FE					
HEM	✗ - Binding site not conserved.			PROTOPORPHYRIN IX CONTAINING FE					
HEM	✗ - Binding site not conserved.			PROTOPORPHYRIN IX CONTAINING FE					

Target MVLSPADKTNVKAAGKVGAGHAGEYGAEALERMFLSFPTTKTYFFHFDLSHGSAQVKGHGKKVADALTNAVAHVDDMPNIA
1bz1.1.A MVLSPADKTNVKAAGKVGAGHAGEYGAEALERMFLSFPTTKTYFFHFDLSHGSAQVKGHGKKVADALTNAVAHVDDMPNIA

Target LSALSDDLHAHLKLRVDPVNFKLLSHCLLVTLAAHLPAEFTPAVHASKDKFLASVSTVLTSKYR
1bz1.1.A LSALSDDLHAHLKLRVDPVNFKLLSHCLLVTLAAHLPAEFTPAVHASKDKFLASVSTVLTSKYR

Materials and Methods

Template Search

Template search with BLAST and HHblits has been performed against the SWISS-MODEL template library (SMTL, last update: 2018-04-18, last included PDB release: 2018-04-13).

The target sequence was searched with BLAST against the primary amino acid sequence contained in the SMTL. A total of 600 templates were found.

An initial HHblits profile has been built using the procedure outlined in (Remmert et al.), followed by 1 iteration of HHblits against NR20. The obtained profile has then been searched against all profiles of the SMTL. A total of 922 templates were found.

Template Selection

For each identified template, the template's quality has been predicted from features of the target-template alignment. The templates with the highest quality have then been selected for model building.

Model Building

Models are built based on the target-template alignment using ProMod3. Coordinates which are conserved between the target and the template are copied from the template to the model. Insertions and deletions are remodelled using a fragment library. Side chains are then rebuilt. Finally, the geometry of the resulting model is regularized by using a force field. In case loop modelling with ProMod3 fails, an alternative model is built with PROMOD-II ([Guex et al.](#)).

Model Quality Estimation

The global and per-residue model quality has been assessed using the QMEAN scoring function ([Benkert et al.](#)). For improved performance, weights of the individual QMEAN terms have been trained specifically for SWISS-MODEL.

Ligand Modelling

Ligands present in the template structure are transferred by homology to the model when the following criteria are met: (a) The ligands are annotated as biologically relevant in the template library. (b) the ligand is in contact with the model, (c) the ligand is not clashing with the protein, (d) the residues in contact with the ligand are conserved between the target and the template. If any of these four criteria is not satisfied, a certain ligand will not be included in the model. The model summary includes information on why and which ligand has not been included.

Oligomeric State Conservation

The quaternary structure annotation of the template is used to model the target sequence in its oligomeric form. The method ([Bertoni et al.](#)) is based on a supervised machine learning algorithm, Support Vector Machines (SVM), which combines interface conservation, structural clustering, and other template features to provide a quaternary structure quality estimate (QSQE). The QSQE score is a number between 0 and 1, reflecting the expected accuracy of the interchain contacts for a model built based on given alignment and template. Higher numbers indicate higher reliability. This complements the GMQE score which estimates the accuracy of the tertiary structure of the resulting model.

References

- BLAST**
Camacho, C., Coulouris, G., Avagyan, V., Ma, N., Papadopoulos, J., Bealer, K., Madden, T.L. BLAST+: architecture and applications. *BMC Bioinformatics* 10, 421-430 (2009). [\[a\] doi>](#)
- HHblits**
Remmert, M., Biegert, A., Hauser, A., Söding, J. HHblits: lightning-fast iterative protein sequence searching by HMM-HMM alignment. *Nat Methods* 9, 173-175 (2012). [\[a\] doi>](#)

Table T1:

Primary amino acid sequence for which templates were searched and models were built.

MVLSPADKTNVKAAGKVGAGHAGEYGAEALERMFLSFPTTKTYFFHFDLSHGSAQVKGHGKKVADALTNAVAHVDDMPNIALSALSDDLHAHLKLRVDPVNFKLLSHCLLVTLAAHLPAEFTPAVHASKDKFLASVSTVLTSKYR

Table T2:

Template	Seq Identity	Oligo-state	QSQE	Found by	Method	Resolution	Seq Similarity	Coverage	Description
1bz1.1.A	100.00	monomer		HHblits	X-ray	1.59Å	0.60	1.00	PROTEIN (HEMOGLOBIN ALPHA CHAIN)
1bab.1.A	99.30	homo-dimer	0.76	HHblits	X-ray	1.50Å	0.60	1.00	HEMOGLOBIN THIONVILLE (DEOXY) (ALPHA CHAIN)
1c7d.1.A	99.30	monomer		HHblits	X-ray	1.80Å	0.60	1.00	PROTEIN (DEOXYHEMOGLOBIN (ALPHA CHAIN))
1dku.1.A	100.00	homo-dimer	0.77	HHblits	X-ray	1.70Å	0.60	0.99	HEMOGLOBIN (DEOXY) (ALPHA CHAIN)
3ia3.1.B	100.00	monomer		HHblits	X-ray	3.20Å	0.60	1.00	Hemoglobin subunit alpha
1zbu.1.B	100.00	monomer		HHblits	X-ray	2.40Å	0.60	1.00	Hemoglobin alpha chain
1zbu.2.B	100.00	monomer		HHblits	X-ray	2.40Å	0.60	1.00	Hemoglobin alpha chain
4yu3.1.A	87.94	monomer		HHblits	X-ray	2.45Å	0.57	0.99	hemoglobin
4yu4.1.C	87.94	homo-dimer	0.63	HHblits	X-ray	2.80Å	0.57	0.99	hemoglobin
1fsx.1.A	87.94	homo-dimer	0.74	HHblits	X-ray	2.10Å	0.56	0.99	HEMOGLOBIN ALPHA CHAIN
1fsx.1.C	87.94	homo-dimer	0.74	HHblits	X-ray	2.10Å	0.56	0.99	HEMOGLOBIN ALPHA CHAIN
1lwh.1.A	87.94	monomer		HHblits	X-ray	1.55Å	0.56	0.99	Hemoglobin alpha chain
1lwh.1.A	87.94	homo-dimer	0.77	HHblits	X-ray	2.20Å	0.56	0.99	HEMOGLOBIN (DEOXY) (ALPHA CHAIN)
1g09.1.C	87.94	homo-dimer	0.56	HHblits	X-ray	2.04Å	0.56	0.99	HEMOGLOBIN ALPHA CHAIN
1g08.1.A	87.94	monomer		HHblits	X-ray	1.90Å	0.56	0.99	HEMOGLOBIN ALPHA CHAIN
1g0a.1.A	87.94	homo-dimer	0.62	HHblits	X-ray	2.04Å	0.56	0.99	HEMOGLOBIN ALPHA CHAIN
5c6e.1.C	87.94	homo-dimer	0.68	HHblits	neutron diff.	2.00Å	0.56	0.99	Hemoglobin subunit alpha
3cy5.1.A	86.52	monomer		HHblits	X-ray	2.00Å	0.56	0.99	Hemoglobin subunit alpha-2
3cy5.1.C	86.52	monomer		HHblits	X-ray	2.00Å	0.56	0.99	Hemoglobin subunit alpha-2
3d1a.1.A	86.62	homo-dimer	0.38	HHblits	X-ray	2.61Å	0.56	1.00	Hemoglobin subunit alpha-1/2
3d1a.1.C	86.62	homo-dimer	0.38	HHblits	X-ray	2.61Å	0.56	1.00	Hemoglobin subunit alpha-1/2
2dhb.1.A	87.23	homo-dimer	0.67	HHblits	X-ray	2.80Å	0.56	0.99	HEMOGLOBIN (DEOXY) (ALPHA CHAIN)
1g0b.1.A	87.23	monomer		HHblits	X-ray	1.90Å	0.56	0.99	HEMOGLOBIN ALPHA CHAIN
1y8i.1.A	87.23	homo-dimer	0.52	HHblits	X-ray	2.60Å	0.56	0.99	Hemoglobin alpha chains
1y8i.1.C	87.23	homo-dimer	0.52	HHblits	X-ray	2.60Å	0.56	0.99	Hemoglobin alpha chains
2zlw.1.A	87.23	homo-dimer	0.63	HHblits	X-ray	2.90Å	0.56	0.99	Hemoglobin subunit alpha
3fh9.1.A	86.52	homo-dimer	0.69	HHblits	X-ray	1.62Å	0.56	0.99	Hemoglobin alpha chain
3gqr.2.A	85.11	monomer		HHblits	X-ray	2.40Å	0.56	0.99	Hemoglobin subunit alpha
3gqp.1.A	85.11	homo-dimer	0.75	HHblits	X-ray	2.00Å	0.56	0.99	Hemoglobin subunit alpha
3eu1.1.C	86.52	homo-dimer	0.37	HHblits	X-ray	3.00Å	0.56	0.99	Hemoglobin subunit alpha-1/2
3eu1.1.A	86.52	homo-dimer	0.37	HHblits	X-ray	3.00Å	0.56	0.99	Hemoglobin subunit alpha-1/2
2qu0.1.C	86.52	homo-dimer	0.60	HHblits	X-ray	2.70Å	0.56	0.99	Hemoglobin subunit alpha-1/2
2qu0.1.A	86.52	homo-dimer	0.60	HHblits	X-ray	2.70Å	0.56	0.99	Hemoglobin subunit alpha-1/2
3hvw.1.A	85.82	monomer		HHblits	X-ray	2.80Å	0.56	0.99	Hemoglobin subunit alpha
3hvw.1.C	85.82	monomer		HHblits	X-ray	2.80Å	0.56	0.99	Hemoglobin subunit alpha
4h2l.1.A	85.11	monomer		HHblits	X-ray	1.78Å	0.56	0.99	Alpha-globin
3gdj.1.A	84.40	homo-dimer	0.62	HHblits	X-ray	2.00Å	0.56	0.99	Hemoglobin subunit alpha
3lqd.1.A	82.98	homo-dimer	0.65	HHblits	X-ray	2.80Å	0.56	0.99	Hemoglobin subunit alpha
2rao.1.A	82.27	homo-dimer	0.62	HHblits	X-ray	2.00Å	0.56	0.99	Hemoglobin subunit alpha-1/2
1y01.1.B	100.00	monomer		HHblits	X-ray	2.80Å	0.60	1.00	Hemoglobin alpha chain
2zlx.2.A	87.23	monomer		HHblits	X-ray	2.80Å	0.56	0.99	Hemoglobin subunit alpha
5ker.1.A	85.11	homo-dimer	0.41	HHblits	X-ray	2.20Å	0.55	0.99	Alpha-globin
5ker.1.C	85.11	homo-dimer	0.41	HHblits	X-ray	2.20Å	0.55	0.99	Alpha-globin
5ker.2.A	85.11	homo-dimer	0.32	HHblits	X-ray	2.20Å	0.55	0.99	Alpha-globin
3gou.1.A	83.69	homo-dimer	0.55	HHblits	X-ray	3.00Å	0.55	0.99	Hemoglobin subunit alpha
3pel.1.A	83.69	homo-dimer	0.60	HHblits	X-ray	1.90Å	0.55	0.99	Hemoglobin subunit alpha
2n14.3.A	86.52	monomer		HHblits	X-ray	2.70Å	0.56	0.99	Hemoglobin subunit alpha-1/2
3gys.1.C	85.11	monomer		HHblits	X-ray	2.90Å	0.56	0.99	Hemoglobin subunit alpha
5eui.1.A	90.07	monomer		HHblits	X-ray	1.45Å	0.57	0.99	HBA protein
2n14.1.C	86.52	homo-dimer	0.35	HHblits	X-ray	2.70Å	0.56	0.99	Hemoglobin subunit alpha-1/2

The table above shows the top 50 filtered templates. A further 866 templates were found which were considered to be less suitable for modelling than the filtered list.

4hsw.1.A, 1lbe.1.A, 5m3l.1.E, 2qmb.1.D, 2eb9.1.A, 2qmb.1.A, 3zjl.1.A, 2bk9.1.A, 2zlw.1.B, 1qpw.1.B, 1qpw.1.C, 1a4f.1.A, 5f82.1.A, 1m4j.1.B, 5m3l.1.A, 1h1x.1.A, 3o7n.1.A, 3s48.2.B, 5ojb.1.A, 1aby.1.A, 1aby.1.B, 3uhc.1.A, 1y0c.1.A, 1umo.1.A, 5ohe.4.A, 1hnm.1.A, 1mj1.1.A, 3mj1.1.B, 1mni.1.A, 3pi8.1.B, 3mj1.1.A, 4v93.124.A, 1x3k.1.A, 1abs.1.A, 1s0h.1.A, 1s56.1.A, 3n17.1.B, 2qu0.1.D, 1lue.1.A, 1hsc.1.D, 1shr.1.C, 1o1n.1.A, 1lra.1.A, 1v4w.1.D, 5hy8.2.B, 1dxv.1.B, 4xs0.1.A, 4xs0.1.B, 1o4.1.A, 4bja.1.A, 3oK5.1.B, 1yzi.1.B, 3qzr.1.B, 3qzr.1.A, 1yvi.1.B, 1mej.1.B, 1m9p.1.A, 1m9p.1.B, 4mqk.1.A, 4mqk.1.C, 5v5r.1.A, 1fj8.1.A, 3aac.1.B, 3bcq.1.C, 2gl3.1.A, 3bcq.1.A, 3qqr.1.A, 3bcq.1.D, 1bjl.1.A, 1bjl.1.B, 3zjh.1.A, 1y5k.1.B, 5jgg.1.B, 2mx0.1.A, 1oj6.2.A, 1n33.1.A, 3zom.1.A, 4bky.1.A, 1o1j.1.A, 1o1j.1.B, 5jgg.1.A, 3wtg.1.A, 1hdb.1.A, 3wtg.1.C, 1bzz.1.A, 2gnv.1.B, 4gtb.1.A, 4lyx.1.A, 1nz3.1.A, 4fwz.1.A, 2wy4.1.A, 1iop.1.A, 4f6d.1.A, 1cmv.1.B, 1cmv.1.A, 5yl3.1.A, 3uh3.1.B, 1wmu.1.B, 3ugz.1.A, 3zhw.2.A, 3ubc.1.A, 1umo.1.A, 3eu1.1.D, 3eu1.1.B, 2vvy.1.A, 5o41.1.A, 4jac.1.A, 1mtk.1.A, 1or6.1.B, 5eet.1.A, 4mqj.2.D, 1y7g.1.B, 2nrl.1.A, 1ngk.1.B, 5eys.1.A, 2zlv.1.C, 2zlv.1.B, 1cgs.1.A, 5urc.1.D, 4mqc.1.A, 1hw4.1.D, 1v5h.1.A, 4f4o.1.A, 4f4o.1.B, 1yhu.1.B, 1mj1.2.A, 1yhu.1.A, 1yhu.1.D, 3wct.1.C, 1o63.1.A, 1mni.1.A, 3s4.1.B, 1yjh.1.B, 3vrg.1.A, 1l2.1.A, 1g8v.1.B, 1hbc.1.A, 1y5v.1.A, 1fjd.1.B, 1hbs.1.D, 1hbs.1.C, 1hbs.1.B, 1hbs.1.A, 1ch9.1.A, 3wfx.1.A, 1s56.2.A, 5e4.1.C, 1ngk.1.H, 4u8u.1.D, 5x2s.2.C, 3vre.1.A, 3vre.1.B, 4mu5.1.A, 1y45.1.C, 1v4x.1.A, 1v4x.1.B, 5xt1.1.D, 5xt1.1.A, 5xt1.1.C, 4b3w.1.A, 5ile.1.A, 4zvb.1.A, 1y5j.1.B, 3ng6.1.B, 1lbe.1.A, 2n14.1.D, 1cbl.1.D, 3gkv.1.A, 3gkv.1.B, 1v4u.1.B, 3s1l.1.A, 4o2g.1.A, 3s1l.1.B, 1o40.1.A, 5y2f.1.A, 1o40.1.B, 3w4u.1.B, 5ljf.1.A, 3w4u.1.A, 1y4g.1.B, 1moh.1.A, 5y6p.7.A, 3vnb.1.A, 3uh6.1.A, 2pgh.1.C, 1vxa.1.A, 2hbs.1.A, 2d6c.1.A, 3ciu.1.D, 1ewa.1.A, 1cpw.1.A, 1y4q.1.B, 3wyo.1.B, 3wyo.1.A, 1yma.1.A, 3aq8.1.A, 2dc3.1.A, 2grt.1.A, 1h1.1.A, 2dc3.1.B, 3hep.1.A, 2ohb.1.A, 4o6f.1.A, 1urv.2.A, 2d6c.2.A, 1irc.1.A, 1ch1.1.A, 4n0.1.A, 1u5.1.A, 3zjl.1.A, 5muu.1.D, 3gou.1.D, 3gou.1.B, 1wvt.1.A, 1y8i.1.D, 1y8i.1.B, 3d1f.1.A, 5f0b.1.A, 3uh1.1.A, 1r1y.1.A, 5x2s.3.C, 1xz5.1.A, 1v07.1.A, 3aq5.1.A, 1cqx.1.A, 5k11.1.B, 5eui.1.B, 1hds.1.A, 3ld.1.A, 1hds.1.A, 1hds.1.B, 1hds.1.D, 5hu6.1.B, 4iro.1.A, 1hbg.1.A, 4iro.1.C, 5hbi.1.B, 1myi.1.A, 1tu9.1.A, 2ig3.1.A, 3aog.1.B, 1w92.1.A, 1dxd.1.A, 1buw.1.A, 1buw.1.B, 1buw.1.D, 2evp.1.A, 3qm7.1.A, 5ker.1.D, 1a00.1.B, 5ker.1.B, 4tww.1.A, 2ef2.1.A, 1hac.1.B, 2hac.1.C, 2r50.2.A, 4hsw.2.A, 2r50.2.B, 1vre.1.A, 3qqr.2.B, 2zs0.1.A, 2zs0.1.B, 2zs0.1.C, 3s3d.1.A, 1fsx.1.D, 1f6h.1.A, 2cmr.1.A, 3o89.1.A, 1m1m.1.A, 4xdl.1.A, 3zh0.1.A, 3wcu.1.A, 3wcu.1.C, 4essa.1.D, 3mjp.1.C, 5vmm.1.C, 3mjp.1.A, 2gkm.1.A, 7hbi.1.A, 2h35.1.B, 2r4w.1.A, 3bz2.1.D, 5x2r.3.A, 5x2r.3.B, 5x2r.3.C, 5x2r.3.D, 3rgk.1.A, 3dht.1.A, 4o1t.1.A, 5x2r.1.C, 5x2r.1.A, 5x2r.1.D, 3gjd.1.B, 4rol.1.D, 1rse.1.A, 3y53.1.A, 5oj9.1.A, 1k1k.1.B, 1ufp.1.A, 1b33.1.A, 1d8u.1.B, 1ash.1.A, 3bjc.1.C, 3qjd.1.A, 1co9.1.A, 3ozu.1.A, 3pi4.1.B, 2eku.1.A, 1ycb.1.A, 1vxh.1.A, 1ch3.1.A, 5x2s.2.D, 3s65.1.B, 1fhj.1.A, 3s65.1.A, 5x2s.1.D, 5x2s.1.A, 3szk.1.A, 5ab8.1.A, 1mcy.1.A, 3v2v.1.A, 2mgf.1.A, 3ggg.1.B, 3ggy.1.A, 3myn.1.A, 2vyz.1.A, 2pgh.1.B, 3onz.1.B, 3fs4.1.A, 2pgh.1.A, 3uhd.1.A, 2pgh.1.D, 5m3l.1.L, 3heo.1.A, 5d1v.1.A, 1pmb.2.A, 3gqp.1.C, 1a3o.1.A, 4mqh.1.A, 1or4.1.B, 1lth.1.A, 1abw.1.A, 4g51.1.C, 1y7z.1.B, 1jle.1.A, 5kq.1.A, 3zh0.1.A, 3wcu.1.A, 3wcu.1.C, 2n14.1.A, 1v46.1.A, 2h35.1.A, 1gqw.1.D, 1y5j.1.A, 2h35.1.A, 3huy.1.A, 1yhu.1.A, 4ns2.1.A, 3kxuo.1.A, 1b33.2.E, 3h58.1.A, 1o16.1.A, 5wog.1.A, 5gc.1.A, 1j2k.1.A, 5udt.1.A, 4n07.1.A, 4yus.1.B, 1rvw.1.A, 1rvw.1.B, 3uh.1.A, 5k5j.1.B, 3whm.1.D, 4hrr.2.D, 1b3d.1.A, 1mob.1.A, 1hbs.2.B, 3qyc.1.B, 4v93.118.A, 2orf.1.A, 1glv.1.A, 1yer.1.A, 4uh.1.A, 1fbb.1.C, 3ohf.1.A, 1xch.1.A, 2ig3.2.A, 4qp.1.B, 3sdh.1.A, 5eu2.1.A, 3at6.1.A, 1ux9.2.A