Supplementary Fig. 1: CLUSTALW result summary

```
CLUSTAL 2.1 Multiple Sequence Alignments

Sequence type explicitly set to Protein
Sequence format is Pearson
Sequence 1: epor 115 aa
Sequence 2: ecoli 149 aa
Start of Pairwise alignments
Aligning...

Sequences (1:2) Aligned. Score: 11.3043
Guide tree file created: [clustalw.dnd]

There are 1 groups
Start of Multiple Alignment

Aligning...
Group 1: Delayed
Alignment Score -30

CLUSTAL-Alignment file created [clustalw.aln]
```

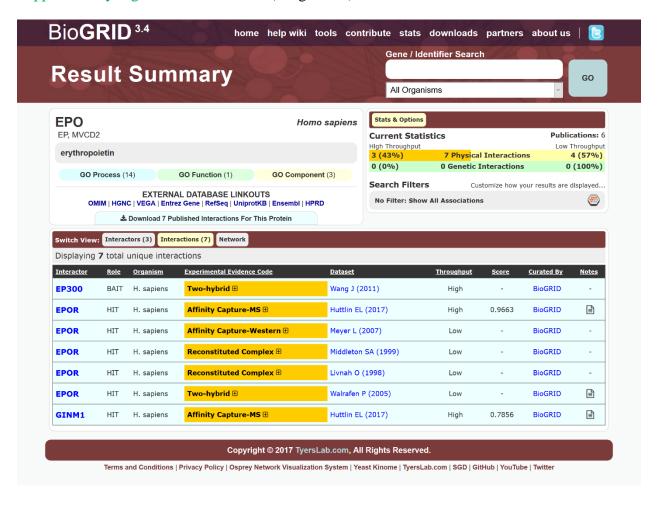
clustalw.aln

CLUSTAL 2.1 multiple sequence alignment

clustalw.dnd

```
(epor: 0.44348, ecoli: 0.44348);
```

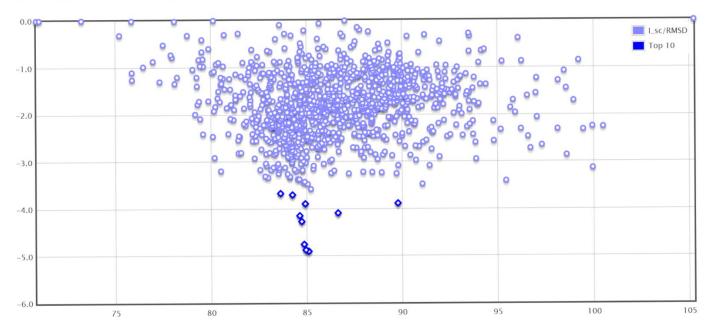
Supplementary Fig. 2: EPO interaction (Biogrid 3.4)



Supplementary Fig. 3: Rosetta Docking (EPO-adhesin)

X axis: RMSD score; Y axis: Interface score

Interface score I_sc / RMSD and Top-10 "I_sc score models"

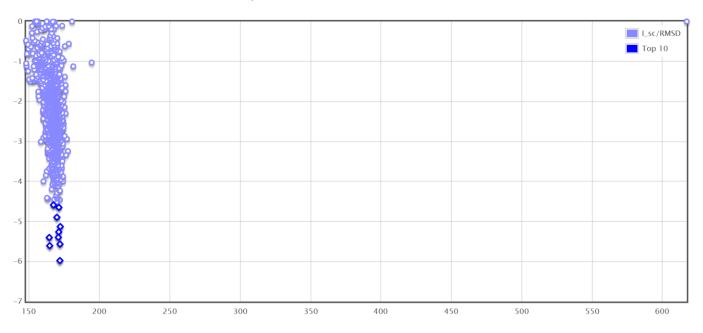


Score data [D	ownload orig	inal score file	<u>al</u>							0
decoy	total_score 4	rms	Fnat	I_sc	Irms	cen_rms	dslf_ca_dih	dslf_cs_ang	dslf_ss_dih	dslf_ss_ds
proteins 0426	-257.612	86.631	NaN	-4.096	NaN	86.454	0.277	0.502	0.437	-3.582
proteins 0538	-257.049	84.73	NaN	-4.275	NaN	84.816	0.277	0.502	0.437	-3.582
proteins 0892	-256.937	83.507	NaN	-3.048	NaN	83.7	0.277	0.502	0.437	-3.582
proteins 0137	-256.881	84.369	NaN	-3.169	NaN	85.306	0.277	0.502	0.437	-3.582
proteins 0006	-256.846	89.79	NaN	-3.083	NaN	91.459	0.277	0.502	0.437	-3.582
proteins 0032	-256.834	85.283	NaN	-2.698	NaN	87.254	0.277	0.502	0.437	-3.582
proteins 0076	-256.813	84.53	NaN	-3.378	NaN	85.219	0.277	0.502	0.437	-3.582
proteins 0880	-256.779	84.2	NaN	-2.917	NaN	85.56	0.277	0.502	0.437	-3.582
proteins 0933	-256.777	83.626	NaN	-3.678	NaN	84.81	0.277	0.502	0.437	-3.582
proteins 0069	-256.763	93.866	NaN	-2.95	NaN	91.681	0.277	0.502	0.437	-3.582
proteins 0222	-256.756	98.605	NaN	-2.863	NaN	89.604	0.277	0.502	0.437	-3.582
proteins 0133	-256.729	84.131	NaN	-3.32	NaN	84.876	0.277	0.502	0.437	-3.582
proteins 0261	-256.727	85.015	NaN	-2.912	NaN	81.766	0.277	0.502	0.437	-3.582
proteins 0693	-256.722	85.568	NaN	-2.718	NaN	87.492	0.277	0.502	0.437	-3.582
proteins 0455	-256.706	84.256	NaN	-2.963	NaN	86.476	0.277	0.502	0.437	-3.582
proteins 0492	-256.703	84.115	NaN	-3.335	NaN	84.528	0.277	0.502	0.437	-3.582
proteins 0057	-256.697	82.867	NaN	-3.041	NaN	85.652	0.277	0.502	0.437	-3.582
proteins 0044	-256.683	84.895	NaN	-3.431	NaN	85.582	0.277	0.502	0.437	-3.582
proteins 0044	-256.682	88.429	NaN	-3.226	NaN	89.146	0.277	0.502	0.437	-3.582
proteins 0075	-256.678	84.952	NaN	-3.024	NaN	87.394	0.277	0.502	0.437	-3.582
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Supplementary Fig. 4: Rosetta Docking (EPO-EPOR)

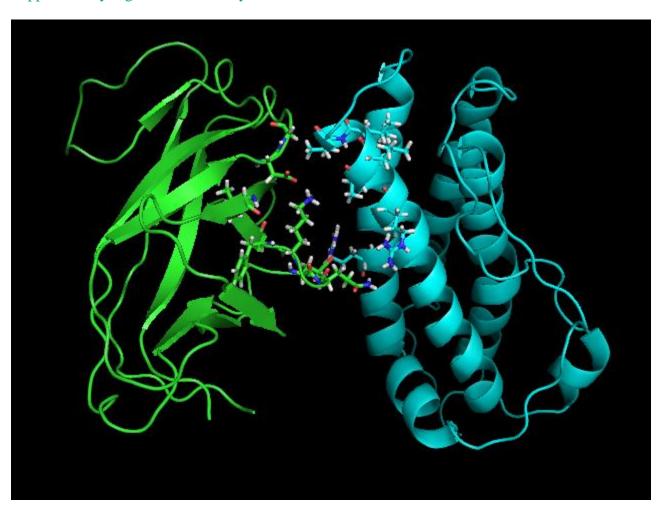
X axis: RMSD score; Y axis: Interface score

Interface score I_sc / RMSD and Top-10 "I_sc score models"



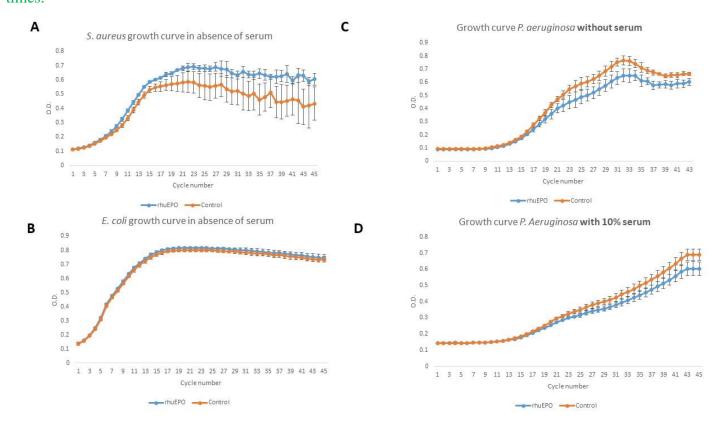
decoy	total_score 4	rms	Fnat	I_sc	Irms	cen_rms	dslf_ca_dih	dslf_cs_ang	dslf_ss_dih	dslf_ss_d
proteins 0753	-313.739	167.341	NaN	-4.587	NaN	167.798	0.592	0.62	3.101	-7.149
proteins 0057	-313.729	166.567	NaN	-4.062	NaN	165.967	0.592	0.62	3.101	-7.149
proteins 0574	-313.676	169.694	NaN	-4.509	NaN	169.815	0.592	0.62	3.101	-7.149
proteins 0084	-313.596	164.649	NaN	-5.605	NaN	166.77	0.592	0.62	3.101	-7.149
proteins 0111	-313.582	171.083	NaN	-5.261	NaN	172.455	0.592	0.62	3.101	-7.149
proteins 0677	-313.568	160.25	NaN	-3.994	NaN	162.246	0.592	0.62	3.101	-7.149
proteins 0242	-313.442	171.97	NaN	-5.564	NaN	171.597	0.592	0.62	3.101	-7.149
proteins 0016	-313.441	164.109	NaN	-3.75	NaN	166.264	0.592	0.62	3.101	-7.149
proteins 0603	-313.4	164.322	NaN	-5.4	NaN	166.749	0.592	0.62	3.101	-7.149
proteins 0112	-313.382	171.883	NaN	-5.977	NaN	172.884	0.592	0.62	3.101	-7.149
proteins 0144	-313.36	165.876	NaN	-3.675	NaN	167.824	0.592	0.62	3.101	-7.149
proteins 0028	-313.35	163.413	NaN	-4.425	NaN	166.082	0.592	0.62	3.101	-7.149
proteins 0319	-313.315	170.506	NaN	-3.797	NaN	171.308	0.592	0.62	3.101	-7.149
proteins 0244	-313.314	167.975	NaN	-4.56	NaN	168.546	0.592	0.62	3.101	-7.149
proteins 0522	-313.27	162.746	NaN	-4.409	NaN	165.699	0.592	0.62	3.101	-7.149
proteins 0412	-313.125	167.234	NaN	-3.207	NaN	168.792	0.592	0.62	3.101	-7.149
proteins 0133	-313.074	167.285	NaN	-3.671	NaN	167.103	0.592	0.62	3.101	-7.149
proteins 0004	-313.025	172.64	NaN	-3.358	NaN	171.083	0.592	0.62	3.101	-7.149
proteins 0517	-313.009	167.458	NaN	-3.839	NaN	169.141	0.592	0.62	3.101	-7.149
proteins 0905	-312.991	166.077	NaN	-4.191	NaN	167.079	0.592	0.62	3.101	-7.149

Supplementary Fig. 5: Molecular Dynamics simulation between EPO and E. coli Adhesin



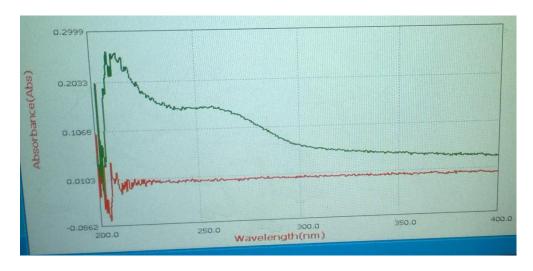
The png file of the complex structure after dynamics is attached. The green and cyan cartoons represent the EPO and adhesin, respectively. The interface residues in between the complex are represented in stick.

Supplementary Fig. 6: Growth curves of different bacteria in presence and absence of rhuEPO repeated 6 times.

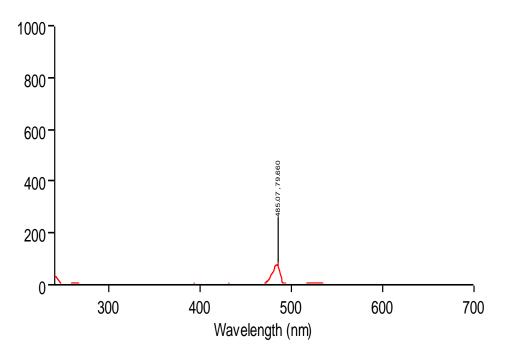


Growth curve of bacteria repeated 6 times are shown. The growth curve of P. aeruginosa in presence of 10% serum is also shown. Experiments of S. aureus and E. coli in presence of 10% serum in our experimental conditions did not yield any growth (data not shown).

Supplementary Fig. 7: Absorbance spectra of membrane protein at 280nm



Supplementary Fig. 8: Fluorescence emission spectra of phosphate buffer (3ml) when excited at 240nm.



Supplementary Fig. 9: Fluorescence emission spectra of phosphate buffer after addition of membrane protein $(0.025 \, \text{mg/3ml})$

