

Supplementary Material

Huanjie Li^{1,2,3}, Qingda Wang², Rui Zhao², Yunshan Wang^{1,3}, Luying Xun^{2,4}, Huawei Liu^{2,*}

¹School of Medicine, Cheeloo College of Medicine, Shandong University, Jinan, Shandong, 250012, People's Republic of China.

²State Key Laboratory of Microbial Technology, Shandong University, Qingdao, Shandong, 266200, People's Republic of China.

³Medical Research & Laboratory Diagnostic Center, Jinan Central Hospital, Cheeloo College of Medicine, Shandong University, Jinan, Shandong, 250013, People's Republic of China.

⁴ School of Molecular Biosciences, Washington State University, Pullman, WA, 99164-7520, USA.

Corresponding Author

*Huawei Liu, Tel: +86 532 5863 1572. E-mail: liuhuaiwei@sdu.edu.cn;

Table of Contents

Supporting methods	2
Parameter values used for computational modeling in Figure 1F	2
Supporting Figures	3
Figure S1. Computational modeling indicates that the activity of thHMGR, HMG-CoA concentration, and GFP all relate to statin concentration.....	3
Figure S2. SDS-PAGE analysis of the thHMGR expression in <i>E. coli</i> (A) and activity assay of the purified enzyme. S1–S3 are three repetitions (B).	4
Figure S3. Toxicity test of statins on <i>E. coli</i> BL21.....	5
Supporting Tables.....	6
Table S1 Docking results of ligands in BsFapR	6
Table S2 Primers used in this study	6

Supporting methods

Parameter values used for computational modeling in Figure 1F

Parameter	Value	Unit
U_{max}	100	mg/L/h
$[SN]$	0–100	ng/mL
IC_{50}	1-100	ng/mL
t	1	h
K_a	1-100	mg/L
$n1$	2	dimensionless
$n2$	2	dimensionless
U_{max}	100	arbitrary unit

Supporting Figures

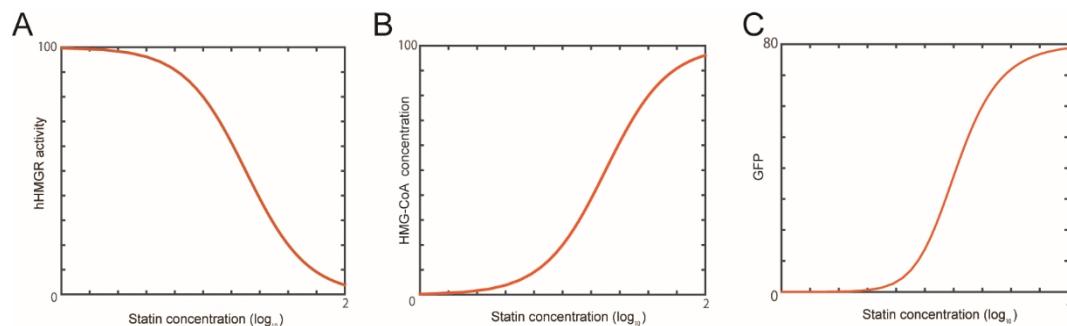


Figure S1. Computational modeling indicates that the activity of thHMGR, HMG-CoA concentration, and GFP expression all relate to statin concentration. They show responses to statin in a dose-dependent mode, suggesting they all can be indicators of statin efficacy.

Parameter values used:

Parameter	Value	Unit
U_{max}	100	mg/L/h
$[SN]$	0–100	ng/mL
IC_{50}	50	ng/mL
t	1	h
K_a	50	mg/L
$n1$	2	dimensionless
$n2$	2	dimensionless
GFP_{max}	100	arbitrary unit

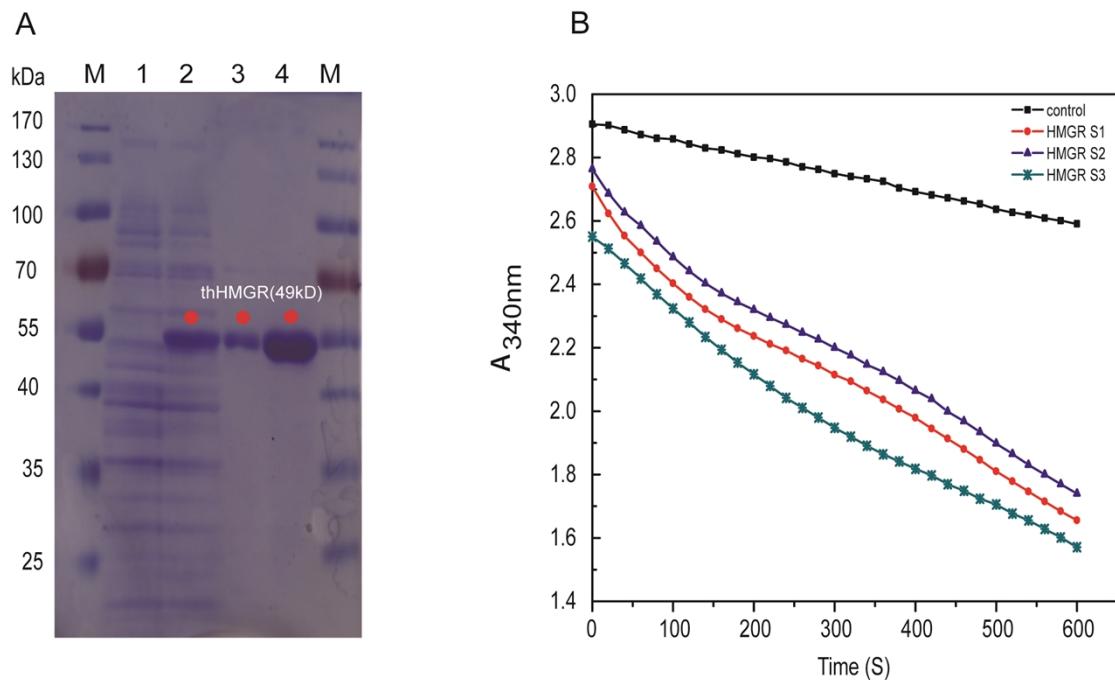


Figure S2. SDS-PAGE analysis of the thHMGR expression in *E. coli* (A) and activity assay of the purified enzyme. S1–S3 are three repetitions (B). The SDS-PAGE analysis indicated the expressed thHMGR was soluble in *E. coli* cytoplasm. lane 1, proteins having no affinity to Ni-NTA agarose resin were washed out at the first elution step; lane 2, total proteins obtained from cell lysis before Ni-NTA resin based purification treatment; lane 3 and 4, the thHMGR protein was washed out at the second elution step during Ni-NTA agarose resin based purification; M, protein marker. After purification, we assayed the activity of thHMGR through detecting the NADPH consumption. The decrease of absorbance at 340 nm was observed, indicating that the thHMGR expressed by *E. coli* retained the catalytic activity.

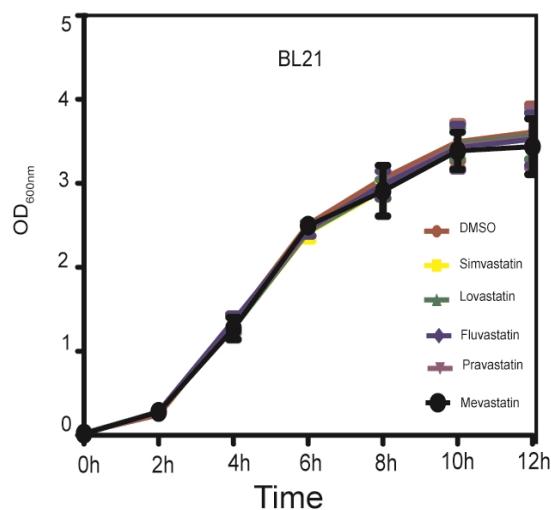
C

Figure S3. Toxicity test of statins on *E. coli* BL21. The toxicity testing experiments were performed with simvastatin, lovastatin, mevastatin, pravastatin, and fluvastatin. Results showed that none of them caused inhibition effect on growth of *E. coli* BL21. DMSO is the solvent of statins, which was also tested as a control.

Supporting Tables

Table S1. Docking results of ligands in BsFapR

Ligand	Binding energy	K_i
Malonyl-CoA	-4.34 kcal/mol	663.69 μM
HMG-CoA	-2.2 kcal/mol	24.28 mM

Table S2. Primers used in this study

Primers	Sequence
thHMGR-F	CGACGACGACAAGGCCATGGCTGATGCCGGCGAAACACATC
thHMGR-R	CAGTGGTGGTGGTGGTGGTGCATTAGAATTCTCCGCGG
T7FapR-F	TAATACGACTCACTATAGGGAAAGAGGGAGAAATAATGAGAA
	GAAATAAGAGAGAACG
T7FapR-R	CCGGACAATTAAGACTAGGTACTAATAGT TAAAGTTAACAAA
	ATTAT
T7FapR-R2	CTCCTCTTCCCTATAGTGAGTCGTATTACACATTACCCACCTG
	AAT
fapO-F	ACTATTAGTACCTAGTCTTAATTGTCCGG ATAATTGTTAAC
	TTTA
fapO-R	CTCCTCTTCCCTATAGTGAGTCGTATTA CACATTACCCACCC
	TGAAT
PTrcfap-sfRfp-F	GTAAGGAACTGCCAGGCATC
PTrcfap-sfRfp-R	ATTCCGGTCGAGTGCCACAC
pBad-F	ATCTGTGTGGGCACTCGACCAGGAATCCATTAGAGAAGAAACC
pBad-R	GTGAATAATTCTCACCTTAGACATATATAACCTCCTAGAGCTC
pBad-F2	ATCTGTGTGGGCACTCGACCAGGAAT
GFP-F	ATGTCTAAAGGTGAAGAA
GFP-R	TATTTGATGCCTGGCAGTCCCTACTCACGCTGCAAGGGCGTAAT
GFP-R2	TATTTGATGCCTGGCAGTCCCTAC
Trc-F	CAATTGTCGATTGTTACCAACGGTCTGGCAAATATTCTG
Trc-R	CTGACGATGACACAATTTCATGGTTATTCCCTCCTT
33MevT-F	AAAAATTGTCATCGTCAG
33Mev-R	GGTAACGAATCAGACAATTG
33-F	GTCGACCTGCAGGCATGCAA
33-R	CATAGTGTAAATCCTCCTTA
HMGR-F	ATAAGGAGGATTACACTATGCCGGCGAAACACATCCC
HMGR-R	TTGCATGCCTGCAGGTCGACTAGAATTCTCCGCGG
op1-Primer5	TTTTAAGAAGGAGATATACCACTATGTGTCAAATACCCCTAGAG
op1-Primer6	GGTATATCTCCTCTAAAAAAACCTGTAGGGGTATTT
pTet-Atcc-F	GATGTTAAAAAAATAACTCGAGTAAGGATCTCCAGGCA
pTet-Atcc-R	TTATTTTTAACATCGTAAGATCTCTAAATTGTCATCG
Trc-Forward	GTTTGACAGCTTATCATCGA
mkate-R	TCAACGATGTCCTAATTTCG