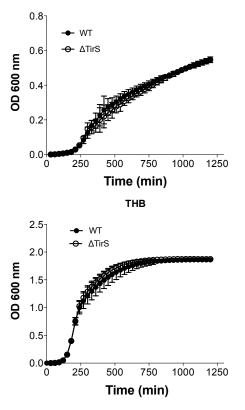
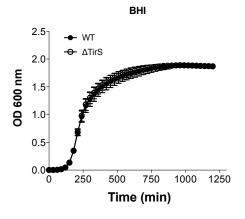
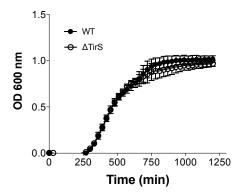


**Supplementary Fig. 1.** Inhibition of TLR2-induced NF- $\kappa$ B activation by TirS is comparable to inhibition mediated by DN-hMyD88 or addition of a TLR2 antagonist SSL3. Untreated control cells were arbitrary set as one, and the luciferase activity in the treated cells is represented as fold induction. \*\*\**P*<0.001.

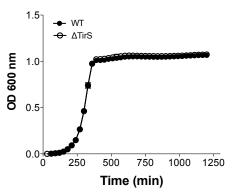




DMEM 1% casaminoacids



**RPMI 1% casaminoacids** 



**Supplementary Fig. 2.** Growth curve of *S. aureus* MSSA476 (WT) and the isogenic mutant MSSA476  $\Delta tirS$  ( $\Delta TirS$ ) in different media.

## Supplementary Table 1. Strain number, accession number and localization of *tirS* in genome of *S. aureus* isolates (NCBI)

Strain no.	Accession no.	Localization
MSSA476	BX571857.1	SCC <sub>476</sub>
45394F	GU122149.1	SCC45394F & SCCmecIV
HDE288	AF411935.3	SCCmecVI
CMFT106	HF569093.1	SCCmec
CMFT535	HF569115.1	SCCmec
CMFT306	HF569104.1	SCCmec
CMFT303	HF569103.1	SCCmec
CMFT201	HF569098.1	SCCmec
CMFT3002	HF569102.1	SCCmec
CMFT181	HF569097.1	SCCmec