

POSTER PRESENTATION

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# Do pathogenic bacteria encode more secreted proteins than their non-pathogenic relatives?

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## Background

Pathogenic and non-pathogenic bacteria secrete proteins for nutrient acquisition, cell-cell communication, and niche adaptation [1]. We hypothesized that pathogenic bacteria may encode larger fractions of secreted proteins (fsp) than their non-pathogenic relatives, assuming that pathogens might be under selective pressure to secrete virulence proteins involved in host immune evasion, invasion, and toxigenesis. To test this hypothesis, we compared the Sec-dependent fsp of various gram-positive and gram-negative bacteria and investigated the relation between the fsp and pathogenic potential of an organism.

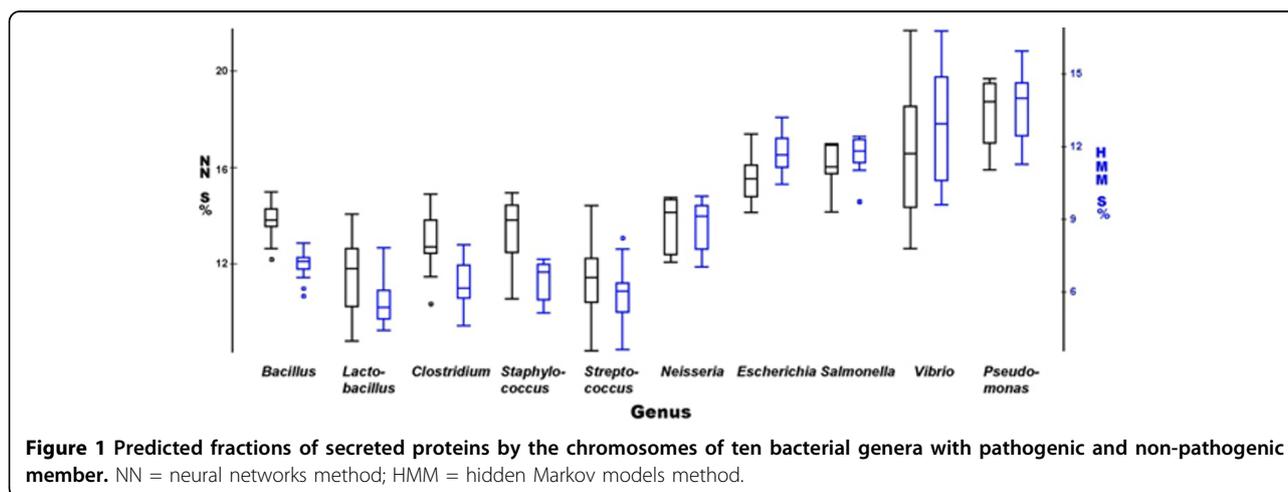
## Methods

We developed a pipeline that starts by a Perl script that truncates protein sequences to 70 amino acids or fewer

followed by the application of existing signal prediction tools [2-4] and ends by the statistical analysis of the prediction data. For subsequent comparative secretome analyses, we used both the hidden Markov models- and the neural networks-based methods implemented in the SignalP 3.0 algorithm [2] (URL: <http://www.cbs.dtu.dk/services/SignalP>) with modified thresholds. We used DataDesk (Data Description, Inc., Ithaca, NY; URL: <http://www.datadesk.com>) for all statistical analyses (including correlation analysis, analysis of variance, and multivariate analysis) and for plotting the results.

## Results

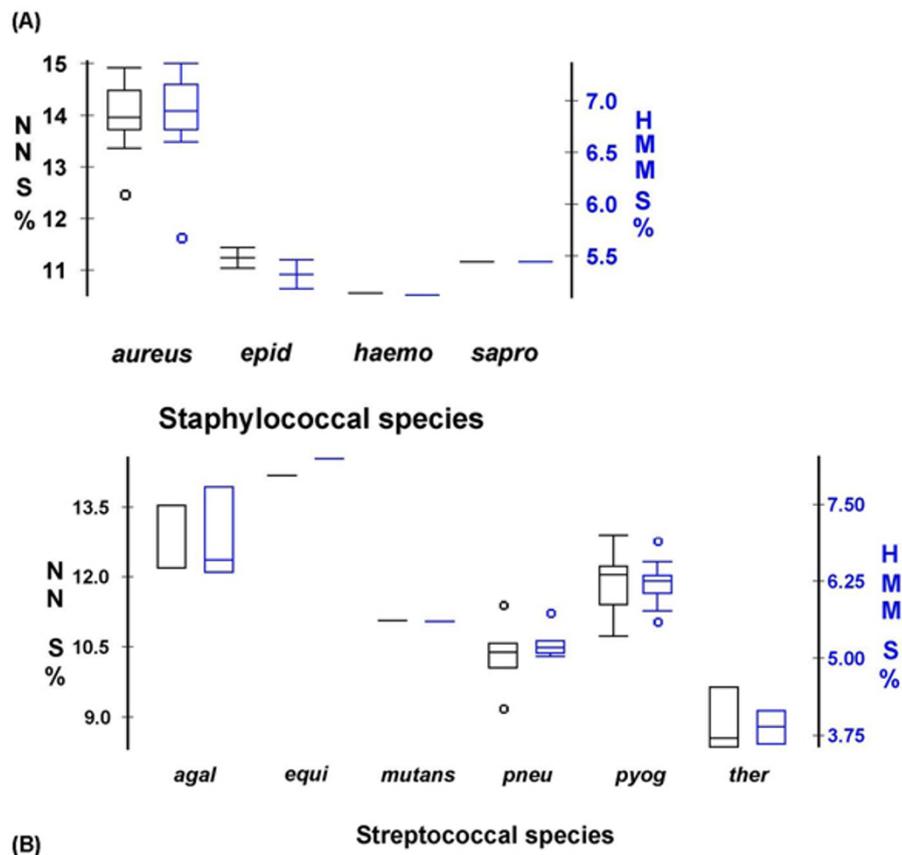
We determined the theoretical secretomes of 176 chromosomes and 115 plasmids in five gram-positive and five gram-negative bacterial genera containing



**Figure 1** Predicted fractions of secreted proteins by the chromosomes of ten bacterial genera with pathogenic and non-pathogenic member. NN = neural networks method; HMM = hidden Markov models method.

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**Figure 2 Pathogenic Gram-positive cocci encode larger fractions of secreted proteins than non-pathogenic relatives.** NN = neural networks method; HMM = hidden Markov models method. (A) Staphylococcal species (*epid* = *epidermidis*; *haemo* = *haemolyticus*; *sapro* = *saprophyticus*). (B) Streptococcal species (*aga* = *agalactiae*; *pneu* = *pneumoniae*; *pyog* = *pyogenes*; *ther* = *thermophilus*). *P* values: *aureus* vs. all: *P* < 10<sup>-6</sup>; *ther* vs. all: *P* = 0.002 (NN), 0.0001 (HMM); *agal* vs. *pyog*: *P* = 0.012 (NN), 0.003 (HMM).

pathogenic and non-pathogenic members (Figure 1). Our analysis showed significant differences in chromosomally encoded *fsp* between gram-positive and gram-negative bacteria (chromosomes of gram-negative bacteria have larger *fsp*), while there was no particular pattern in plasmid-encoded *fsp*. Whereas the overall difference between pathogenic and non-pathogenic species was not statistically significant, significant correlation was observed between *fsp* and pathogenesis in gram-positive cocci. For example, pathogenic *Staphylococcus aureus* have higher *fsp* than other staphylococci, while the non-pathogenic *Streptococcus thermophilus* has the lowest *fsp* of all streptococci (Figure 2).

## Conclusion

We developed a pipeline for the determination and comparison of fractions of secreted proteins in bacterial genomes, and observed significant differences between pathogenic and non-pathogenic species of staphylococci and streptococci.

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