

How Quorum Sensing Interactions Affect Population Structure

02-712 Final Project

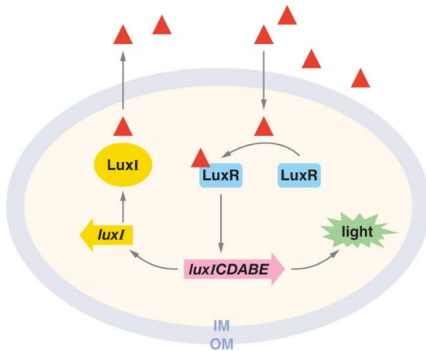
Sid Reed, Neel Mehtani, Sarah Wenger, Deepika Yeramosu,
Evan Trop

Computational Biology Department, Carnegie Mellon University

November 30, 2021

Background

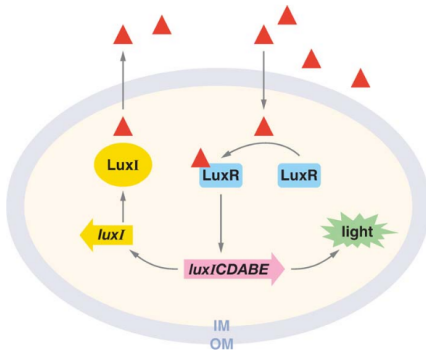
Quorum-Sensing Systems



- ▶ Signal-Receptor molecule pairs that modulate gene expression
- ▶ Once threshold density is reached, enough signal is received to upregulate target genes

Figure 1: Waters and Bassler (2005)

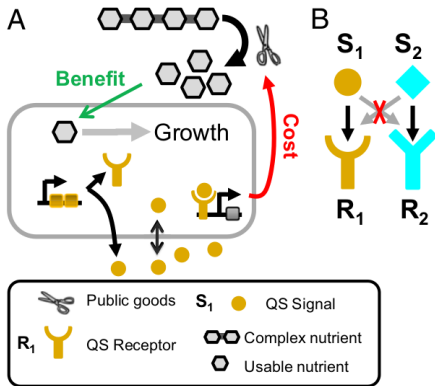
Quorum-Sensing Systems



- ▶ Signal-Receptor molecule pairs that modulate gene expression
- ▶ Once threshold density is reached, enough signal is received to upregulate target genes
- ▶ Can lead to biofilms, antibiotic production etc.

Figure 1: Waters and Bassler (2005)

Public Goods and Cheating



- ▶ When quorum is reached, bacteria produce a “public good”

Figure 2: Eldar (2011)

Public Goods and Cheating

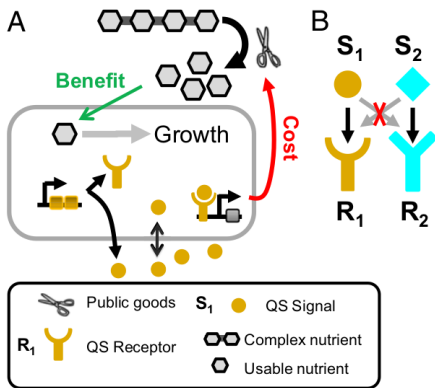


Figure 2: Eldar (2011)

- ▶ When quorum is reached, bacteria produce a “public good”
- ▶ Everyone benefits from this even if they don’t contribute

Public Goods and Cheating

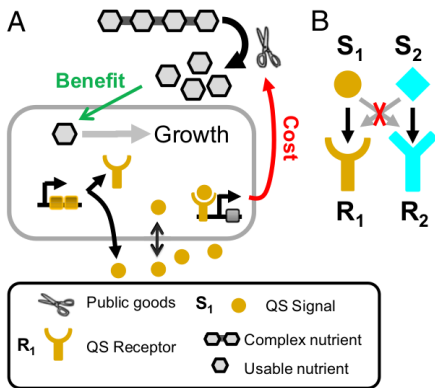


Figure 2: Eldar (2011)

- ▶ When quorum is reached, bacteria produce a “public good”
- ▶ Everyone benefits from this even if they don’t contribute
- ▶ Must produce the receptor, signal molecule and good to contribute

Public Goods and Cheating

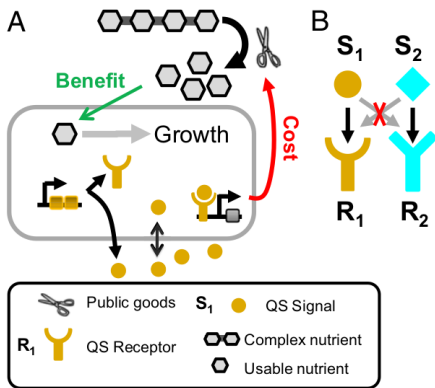


Figure 2: Eldar (2011)

- ▶ When quorum is reached, bacteria produce a “public good”
- ▶ Everyone benefits from this even if they don’t contribute
- ▶ Must produce the receptor, signal molecule and good to contribute
- ▶ Cheaters DO prosper (if you are a bacterium)

Who Cares?

- check the discussion from Eldar (2011) for references

Maintaining Freeloaders as a Diversity Reservoir

Who Cares?

- ▶ check the discussion from Eldar (2011) for references

Maintaining Freeloaders as a Diversity Reservoir

Kin Recognition for Strains

Who Cares?

- ▶ check the discussion from Eldar (2011) for references

Maintaining Freeloaders as a Diversity Reservoir

Kin Recognition for Strains

Designing Cheaters to Disrupt Pathogen Growth

Methods

Basic ODE Model

Social conflict drives the evolutionary divergence of quorum sensing

Avigdor Eldar¹

Department of Molecular Microbiology and Biotechnology, Faculty of Life Sciences, Tel Aviv University, Tel Aviv 69978, Israel

- Equations taken from Eldar (2011)

Basic ODE Model

Social conflict drives the evolutionary divergence of quorum sensing

Avigdor Eldar¹

Department of Molecular Microbiology and Biotechnology, Faculty of Life Sciences, Tel Aviv University, Tel Aviv 69978, Israel

- ▶ Equations taken from Eldar (2011)
- ▶ Deterministic ODE model of bacteria

Basic ODE Model

Social conflict drives the evolutionary divergence of quorum sensing

Avigdor Eldar¹

Department of Molecular Microbiology and Biotechnology, Faculty of Life Sciences, Tel Aviv University, Tel Aviv 69978, Israel

- ▶ Equations taken from Eldar (2011)
- ▶ Deterministic ODE model of bacteria
- ▶ We focused on QS interactions over Evolution

Signal-Receptor Activation Matrix K_{ac}

- Represents all receptors-signal pairs $(R_i S_i)$ present in at least 1 OTU in the population

Signal-Receptor Activation Matrix K_{ac}

- ▶ Represents all receptors-signal pairs ($R_i S_i$) present in at least 1 OTU in the population
- ▶ Different sets of receptor-signal combinations can produce the same K_{ac}

Signal-Receptor Activation Matrix K_{ac}

- ▶ Represents all receptors-signal pairs $(R_i S_i)$ present in at least 1 OTU in the population
- ▶ Different sets of receptor-signal combinations can produce the same K_{ac}
- ▶ K_{ac} is of dimension $|R| \times |S| = |N| \times |N|$

Facultative Cheaters

Matrix for 2 strains R_1S_1 and R_2S_2

$$\begin{bmatrix} 1 & 0 \\ 0 & 1 \end{bmatrix}$$

Facultative Cheaters

Matrix for 2 strains R_1S_1 and R_2S_2

$$\begin{bmatrix} 1 & 0 \\ 0 & 1 \end{bmatrix}$$

Obligate Cheater

Matrix for 2 strains R_1S_1 and R_0S_0

$$\begin{bmatrix} 1 & 0 \\ 0 & 0 \end{bmatrix}$$

Facultative Cheaters

Matrix for 2 strains R_1S_1 and R_2S_2

$$\begin{bmatrix} 1 & 0 \\ 0 & 1 \end{bmatrix}$$

Obligate Cheater

Matrix for 2 strains R_1S_1 and R_0S_0

$$\begin{bmatrix} 1 & 0 \\ 0 & 0 \end{bmatrix}$$

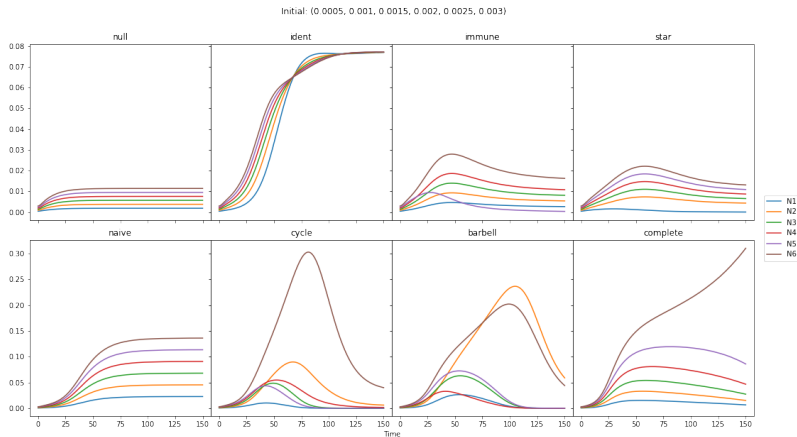
Custom Matrix

Matrix for 2 strains $R_1R_2S_1$ and R_2S_2
or 3 strains R_1S_1 , R_2S_1 and R_2S_2

$$\begin{bmatrix} 1 & 0 \\ 1 & 1 \end{bmatrix}$$

Results

Comparing Different K_{ac} Matrices



How K_{ac} Sparsity Affects Population Structure

Simulating With Human Gut Microbiome Data

Discussion

Chaining K_{ac} has strong effects on model dynamics

- changes can occur independent of sparsity

Discussion

Chaining K_{ac} has strong effects on model dynamics

- ▶ changes can occur independent of sparsity
- ▶ decreasing sparsity does ...

Moral of the Study

Moral of the Study

Cheating works...

Moral of the Study

Cheating works... (for bacteria)

Moral of the Study

Cheating works... (for bacteria)
but cooperating is better!

Bibliography I

Aggarwal, Surya D., Hasan Yesilkaya, Suzanne Dawid, and N. Luisa Hiller. 2020. "The Pneumococcal Social Network." *PLOS Pathogens* 16 (10). <https://doi.org/10.1371/journal.ppat.1008931>.

Calle, M. Luz. 2019. "Statistical Analysis of Metagenomics Data." *Genomics & Informatics* 17 (1).
<https://doi.org/10.5808/gi.2019.17.1.e6>.

Dimitriu, Tatiana, Frances Medaney, Elli Amanatidou, Jessica Forsyth, Richard J. Ellis, and Ben Raymond. 2019. "Negative Frequency Dependent Selection on Plasmid Carriage and Low Fitness Costs Maintain Extended Spectrum Beta-Lactamases in *Escherichia Coli*." *Scientific Reports* 9 (1).
<https://doi.org/10.1038/s41598-019-53575-7>.

Bibliography II

Eldar, A. 2011. "Social Conflict Drives the Evolutionary Divergence of Quorum Sensing." *Proceedings of the National Academy of Sciences* 108 (33): 13635–40.

<https://doi.org/10.1073/pnas.1102923108>.

Pérez-Escudero, Alfonso, and Jeff Gore. 2016. "Selection Favors Incompatible Signaling in Bacteria." *Proceedings of the National Academy of Sciences* 113 (8): 1968–70.

<https://doi.org/10.1073/pnas.1600174113>.

Bibliography III

Pollak, Shaul, Shira Omer-Bendori, Eran Even-Tov, Valeria Lipsman, Tasneem Bareia, Ishay Ben-Zion, and Avigdor Eldar. 2016.

“Facultative Cheating Supports the Coexistence of Diverse Quorum-Sensing Alleles.” *Proceedings of the National Academy of Sciences* 113 (8): 2152–7.

<https://doi.org/10.1073/pnas.1520615113>.

Waters, Christopher M, and Bonnie L. Bassler. 2005. “Quorum Sensing: Cell-to-Cell Communication in Bacteria.” *Annual Review of Cell and Developmental Biology* 21: 319–46.

<https://doi.org/10.1146/annurev.cellbio.21.012704.131001>.