

Communication Mode between Microorganisms

Subjects: Plant Sciences | Microbiology

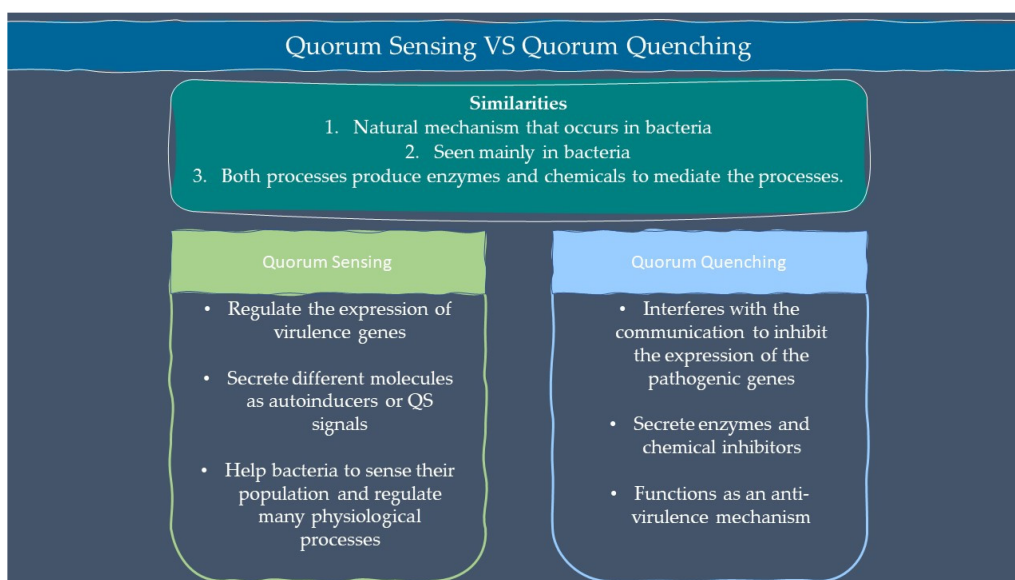
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Various processes such as rhizospheric competence, antibiosis, release of enzymes, and induction of systemic resistance in host plants are all used by microbes to influence plant-microbe interactions. These processes are largely founded on chemical signalling. Producing, releasing, detecting, and responding to chemicals are all part of chemical signalling. Different microbes released distinct sorts of chemical signal molecules which interacts with the environment and hosts.

Keywords: quorum sensing ; quorum quenching ; autoinducers

1. Introduction

Microbes are sensitive to the changes in their environment. In order to survive harsh environments, microbes alter their gene expression that affects microbial behavior. Microbes need to defend and protect themselves not only against the environment but also against other microbes that exist in the same niche. Communication is an important tool for all organisms to interact with each other. Microorganisms such as bacteria and fungi have a special way of interacting through chemical signal molecules known as autoinducers. These autoinducers trigger chemical communication between microbes. This communication process is called quorum sensing (QS), which allows bacteria and fungi to keep an eye on their surroundings for other bacteria/fungi and adjust their activity on a community scale in response to changes in quantity and species existing within a community. QS is important in microbes as it is used in the production of virulence factors, biofilm formation, and swarming motility ^{[1][2]}. Autoinducers are classified into three main types which are AI-1 (N-acyl homoserine lactones, AHLs), oligopeptides or autoinducing peptide and AI-2. Other than the above, there are a few more signalling molecules that are unique and do not belong to any classes such as diffusible signal factor (DSF), *Pseudomonas* quinolone signal (PQS), and diketopiperazine. AI-1 regulates Gram-negative bacteria QS while oligopeptides are discovered in Gram-positive bacteria. AI-2 is an interspecies autoinducer that is present in many species of Gram-negative and Gram-positive bacteria ^[3]. Quorum quenching (QQ) on the other hand is a process that interferes with quorum sensing. QQ is believed to have been developed as a natural method by QS-emitting species to clear their own QS signals, or competitive interaction with QS-emitters by QQ organisms. Furthermore, many different species employ QS to control the development and functioning of antimicrobials. Antimicrobial compounds are released by microbes to preserve the population stability which can cause injury or kill the target cells. Next, the capability to colonize a community is greatly influenced by the restricted amount of nutrients in the environment. Microorganisms with specialized metal acquisition systems such as siderophore can bind and promote the absorption of important metals from their environment, thus restricting the capacity of rival microbes to acquire necessary nutrients and able to colonize a community. A summary of quorum sensing and quorum quenching signaling is shown in **Figure 1** below.



2. Quorum Sensing

Quorum Sensing is a communication systems used by microorganism, which is critical for the establishment of relationship between the microorganisms and their host ^[4]. QS is a social characteristic communication between bacteria and the environment in which bacteria creates and senses signal molecules to coordinate their behaviour in a population-dependent manner ^{[1][2]}. When QS molecules reach a certain level, bacteria adjust their gene expression pattern to cope with high cell density microbial cell surroundings. Unique extracellular signal molecules known as 'autoinducers' are associated with QS. N-acyl homoserine lactones (AHLs) are extensively studied autoinducer in Gram-negative bacteria, that possess an invariant lactone ring and acyl tail of varying lengths, saturations and presence of hydroxyl group ^[5]. These distinctions in its structure confers species uniqueness as well as differences in genetic regulation depending on the AHL receptor which serves as transcriptional regulator for a variety of bacterial community activities, including biofilm formation and pathogenicity ^[5].

The biofilm matrix is a harmonious community that helps to protect the microorganism from harsh environment and is vital for colonization ^[1]. Bacteria in biofilms are known to efficiently sustain communities by secreting extracellular chemicals that allow them to communicate with one another without having to come into direct physical contact. The LuxI is an autoinducer synthase enzyme that synthesizes AHLs, where the AHLs produced will interact with receptor proteins (LuxR homologues) in intracellular spaces of Gram-negative bacteria, and the dimers produced governing the phenotypic gene expression of biofilm formation, enzyme synthesis, manufacturing of antibiotics, and virulence factors ^[6]. Even at very low concentrations of AHLs, plants may detect their presence and respond in a variety of ways including changes in hormone levels involved in self-defence and the release of hormones associated with growth such as auxin, ethylene and jasmonic acid ^[7].

Oligopeptide autoinducers are used by Gram-positive bacteria as lead molecules. These autoinducing peptides (AIPs) are ribosomally produced and may have post-translational changes that affect the stability and functionality of their side chains ^[8]. Peptides typically need transporters to reach the extracellular environment, as they are impermeable to the bacterial membrane ^[8]. Diffusible signal factors (DSFs) are medium-chain unsaturated fatty acids that regulate QS in a variety of organisms, including *Burkholderia cenocepacia*, *Candida albicans*, *Pseudomonas aeruginosa*, *Stenotrophomonas maltophilia*, and *Xylella fastidiosa*, implying the involvement of inter-kingdom signalling pathways. *Cis*-2-dodecenoic acid, *cis*-11 methyl-dodeca-2,5-dienoic acid, *cis*-11-methyl-2-dodecenoic acid, *cis*-10-methyl-2-dodecenoic acid, and *trans*-2-decenoic acid are examples of DSF compounds ^[9]. The first discovered DSF was *cis*-11-methyl-2-dodecenoic acid which was discovered in the *Xanthomonas campestris* pv. *campestris*. It influences the expression of extracellular enzymes such as Egl and protease, virulence factors and xanthan, as well as the regulation of pathogenicity factors (*rpf*) genes. The crotonase family enzyme *rpfF*, acts on fatty acyl carrier protein substrates, and the fatty acyl CoA ligase *RpfB* is required for *X. campestris* pv. *campestris* DSF production. A two-component system for DSF detection and signal transduction consists of the sensor *RpfC*, and the regulator *RpfG* ^{[9][10]}. Recognition of DSF by *RpfC* is related to phosphorylation of the HD-GYP which acts as the domain regulator and changes the cellular level of the second messenger cyclic di-GMP. Distinct pathways govern different subsets of *Rpf*-regulated virulence activities. *RpfC* favourably influences virulence factor production while adversely regulating DSF synthesis ^[10].

3. Quorum Quenching

Quorum quenching (QQ) is an interference to the QS system which will disrupt the attack of bacterial population. QQ possesses two main mechanisms; (1) QS signal molecule inhibitors (QSIs), and (2) QS signal molecule degradation enzymes. The QSI mechanism stops signal molecules from interacting with receptor proteins, thus interfering with QS, while the other mechanism reduces signal molecules by generating degrading enzymes, resulting in QQ ^[8]. Extracts of beans, clover, pea, garlic, geranium, grape, lily, lotus, pepper, strawberry, soybean, vanilla, and yam reduce AHL of QS in a variety of bacterial species ^[6]. Lactonase present in these plant extracts have QQ action. Lactones such as patulin and penicillic acid found in fungi behave as bacterial AHL signal counterparts. Patulin can be found in apples, pears, peaches, apricots, bananas, and pineapple, making these foods promising anti-QS phyto resources ^[9].

AHLs can be destroyed or changed by lactone hydrolysis, amidohydrolysis and oxidoreduction ^[11]. The activity of AHL acylase and AHL lactonase enzymes has been documented to cause AHL degradation that may be caused by multiple phylum members including Proteobacteria, Actinobacteria, and Firmicutes ^[12]. Furthermore, bacterial oxidoreductases, such as those produced by *Rhodococcus* sp., have the ability to actively alter AHL ^[6]. Lactonases that catalyse the hydrolysis of the ester bond to open the AHL ring are classified into several classes based on their folds.

Phosphotriesterase-like lactonases are a common type of lactonase which requires two metal ions and a TIM barrel fold (triose-phosphate isomerase) for proper functionality. TIM barrel proteins are crucial because it is needed to support wide range of enzymatic activities [13]. AHL lactonases have been shown to successfully hydrolyzes a variety of lactones, including QS AHLs ranging from C4- to C12-homoserine lactone (HSL) [9], with or without C3 alteration.

QSI are small molecules which have the capacity to effectively reduce quorum sensing controlled gene expression [14]. These compounds must be stable, specific and resistant to degradation as they will encounter different metabolic reactions in the cell. These compounds alter gene expressions of the targeted genes by binding to different promoters which may interrupt the interaction of the signalling molecules or prevent the synthesis of signal molecules hence inhibit the generation of secondary signals that modulate gene expression [14]. For example, a few *Bacillus* strains have been associated to *aiiA* and *TasA* genes, which encode for many QSI, including lactonase, and have a broad spectrum antimicrobial action, suggesting that they might be used to manage bacterial diseases biologically [15]. In addition, furanones which are synthesized by fungi have a significant role as QSI for many Gram negative and positive bacteria by triggering the induction of stress response genes in a QS-independent manner [16]. A summary of signalling molecules produced by microbes and plants is shown in **Figure 2** below.

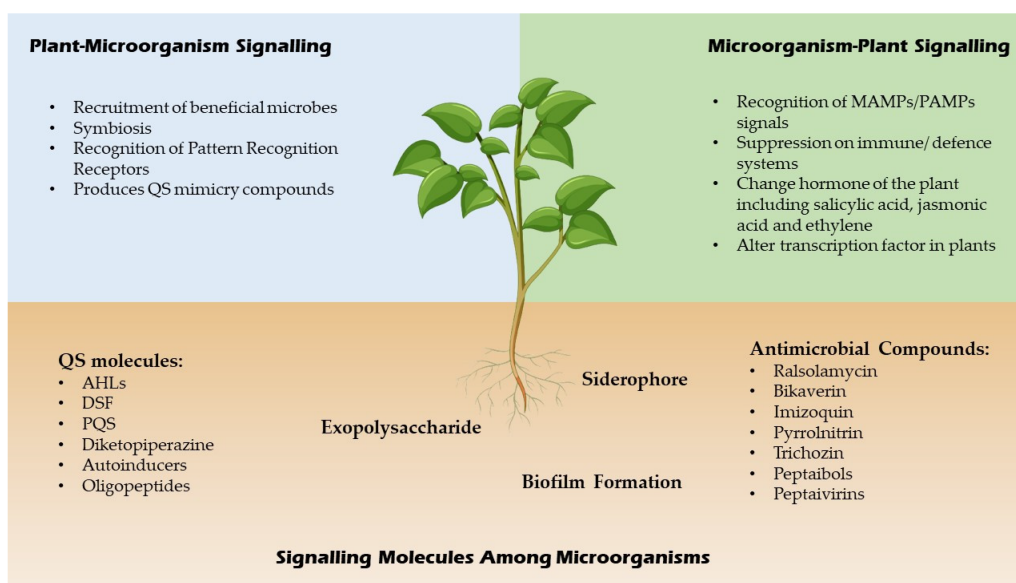


Figure 2. Signaling molecules of microbes and plants.

4. Chemical Signalling in Fungi

One of the most prevalent chemical signaling molecule in fungi is farnesol. Following the discovery of farnesol in *Candida albicans*, it was discovered that lipids (oxylipins), peptides (pheromones), alcohols (tyrosol, farnesol, tryptophol, and 1-phenylethanol), acetaldehydes, and several volatile chemicals are actively engaged in fungal QS systems [14]. QS in fungi is often responsible for germination of spore, production of secondary metabolites, taxonomic transformation and enzyme secretions [17].

Intraspecies of fungi communicate with each other by releasing pheromones. Pheromones are used as signalling molecules to govern spore germination, production of secondary metabolites, structural transformation and enzyme secretion in fungi [17]. Pheromones produced are different based on the alleles expressed at the *MAT* locus [18]. For instance, *Saccharomyces cerevisiae* is one of the most popular and broadly described yeast where pheromones generated by this fungus cells are diffusible peptides which are known as a-factor and α -factor. Alleles expressed at the *MAT* locus will determine the peptide hormone and create only one of the two peptide pheromones. *MAT α* is responsible for α expression where the pheromone precursor is encoded by *MF α 1* that passes through numerous proteolytic processes before delivering a matured pheromone. *MAT α* meanwhile is responsible for "a" expression, where the a-factor is farnesylated and can be recognized by ABC transporter Ste6p for a-factor secretion [18].

Mycoparasitic fungi such as *Trichoderma* sp. are commonly used in agriculture to combat other fungal pathogens such as *Rhizoctonia solani* and *Fusarium* sp. [19]. *Trichoderma* sp. produce a few metabolites including harzianopyridone, trichodermin and glivorin which have antifungal or antimicrobial properties that allow them to thrive in various environments [20]. *Fusarium* produces mycotoxins known as fusaric acid and deoxynivalenol (DON) which can activate defense mechanisms in *T. atroviride* and *Clonostachys rosea* which results in mycotoxin detoxification [21][22]. DON and fusaric acid also play an important role as a virulence factor that can cause *Fusarium* wilt in plant. DON synthesis is

related to oxidative stress ^{[23][24]} while fusaric acid synthesis is related to metal ion content ^[25]. This two chemicals can hamper bacteria interaction by QQ of AHL in low concentration, and suppressing phenazine-1-carboxamide production at higher concentrations ^{[26][27]}. Other than that, zearalenone (ZEN) is another mycotoxin produced by *Fusarium* species. *C. rosea* however was reported to detoxify ZEN by breaking the ring structure of ZEN. *Trichoderma* sp. turns ZEN into sulphated form and reduces DON into its glycosylated form of deoxynivalenol-3-glucoside ^{[21][28]}. **Table 1** below shows signal molecules produced by microorganisms and their respective functions.

Table 1. Signal Molecule Produced by Fungi and the Function.

Organism	Signal Molecule	Role	References
<i>Fusarium</i>	Fusaric acid	<ul style="list-style-type: none"> • Virulence factor 	^{[29][30]}
		<ul style="list-style-type: none"> • Reduces activity of <i>Pseudomonas</i> and <i>Bacillus</i> 	
<i>S. cerevisiae</i>	Tryptophol	<ul style="list-style-type: none"> • Promotes pseudohyphal growth 	^[31]
	Farnesol	<ul style="list-style-type: none"> • Induces cellular death • Increases mitochondrial reactive oxygen species concentration 	^[32]
<i>Debaryomyces nepalensis</i>	Phenylethanol	<ul style="list-style-type: none"> • Promotes biofilm formation 	^[33]
<i>Penicillium</i> spp. and <i>Aspergillus</i> spp.	Patulin	<ul style="list-style-type: none"> • Virulence factor • Fungal colonization • Antimicrobial activity 	^{[34][35]}
		<ul style="list-style-type: none"> • Virulence factors 	
	Oxylipins	<ul style="list-style-type: none"> • Growth factors during sexual development • Controls mycotoxin production 	^{[36][37]}
<i>Penicillium sclerotiorum</i>	Multicolic acid	<ul style="list-style-type: none"> • Accelerates the process of sclerotiorin synthesis 	^[17]
<i>Fusarium culmorum</i>	Terpenes	<ul style="list-style-type: none"> • Influences swimming and swarming motility of <i>Serratia plymuthica</i> 	^[38]
<i>Penicillium decumbens</i>	Farnesol	<ul style="list-style-type: none"> • Promotes the growth of hyphae 	^[39]
		<ul style="list-style-type: none"> • Increases the secretion of cellulase 	
<i>Penicillium expansum</i>	Farnesol	<ul style="list-style-type: none"> • Induces cellular death 	^[40]
		<ul style="list-style-type: none"> • Inhibits the growth of hyphae 	
<i>Fusarium graminearum</i>	Farnesol	<ul style="list-style-type: none"> • Inhibits growth of hyphae 	^[40]

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