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Neuropathic pain (NP) is a complex disorder that impacts the quality of life of more than 7 million Americans. There is a need to identify new therapeutic targets for this disorder so that more effective therapies can be developed. Serotonin and the serotonin 2A (5-HT_{2A}) receptor, a drug development target for NP, are allosterically modulated by ligands at G-protein-coupled receptors (GPCRs). We have shown that transient receptor potential melastatin 7 (TRPM7) is a novel, metabotropic 5-HT_{2A} receptor, and that TRPM7 is essential for the development of NP. In this proposal we will establish a new paradigm for the study of NP, by determining whether GPCR allosteric ligands that interact with 5-HT_{2A} can block NP. We will also determine whether NP can be prevented by preventing interactions of NP-relevant GPCRs with allosteric ligands. We have developed an array of brain-penetrant anandamide (AEA) analogs, agonists and antagonists, and will use these to screen for compounds that prevent NP. In Specific Aim 1 we will determine whether AEA analogs prevent the NP resulting from the interference of 5-HT_{2A} signaling with dorsal root ganglion (DRG) neuron development. We will use both behavioral and cellular NP models; and establish whether the NP can be prevented by drug or genetic means. We will also assess whether NP can be prevented by preventing interactions of 5-HT_{2A} signaling with downstream GPCRs. In Specific Aim 2 we will determine whether GPCR allosteric ligands can block NP using an unbiased genetic screen. In Specific Aim 3 we will use a recently discovered TRPM7-selective allosteric antagonist, to determine whether TRPM7 is an essential mediator of NP. We will determine whether NP can be prevented by blocking TRPM7-interacting GPCRs. Thus, in this proposal we will be the first to test whether compounds that interact with GPCRs at allosteric sites can block NP, thereby establishing a novel paradigm for GPCR drug development. The invention concerns a compartmentalization apparatus for an anaerobic digester, preferably for a methanogenesis gas-evolved anaerobic digester, comprising at least one partitioning device arranged to immobilize a microorganism in such a way that the microorganism is deposited between two partitioning elements and that the

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