Takahiro Ohyama, a post-doc in the Lab of Andy Groves, has been using a conditional Cre line to knockout or over-express molecules in the mouse inner ear. His work has been answering fundamental questions on the induction of the mouse inner ear and enables us to reconcile previously conflicting data from other species.

Summary:

The otic placode, the anlagen of the inner ear, develops from an ectodermal field characterized by expression of the transcription factor Pax2. Previous fate mapping studies suggest that these Pax2+ cells will give rise to both otic placode tissue and epidermis (Streit, Dev. Biol. 2002; Ohyama and Groves, Genesis 2004), but the signals that divide the Pax2+ field into placodal and epidermal territories are unknown. We analyzed a reporter strain that carries six copies of TCF/Lef binding sites (Mohamed et. al., Dev. Dyn. 2004), which revealed that the canonical Wnt signaling pathway is normally activated in a subset of Pax2+ cells. We also performed conditional inactivation of beta-catenin in these cells and found an expansion of epidermal markers at the expense of the otic placode. Conversely, conditional activation of beta-catenin in Pax2+ cells causes an expansion of the otic placode at the expense of epidermis, and the resulting otic tissue expresses exclusively dorsal otocyst markers. Together these results suggest that Wnt signaling acts instructively to direct Pax2+ cells to an otic placodal, rather than an epidermal fate, and promotes dorsal cell identities in the otocyst. Based on our present study, we propose a new model of inner ear induction that reconciles conflicting data from recent studies.

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