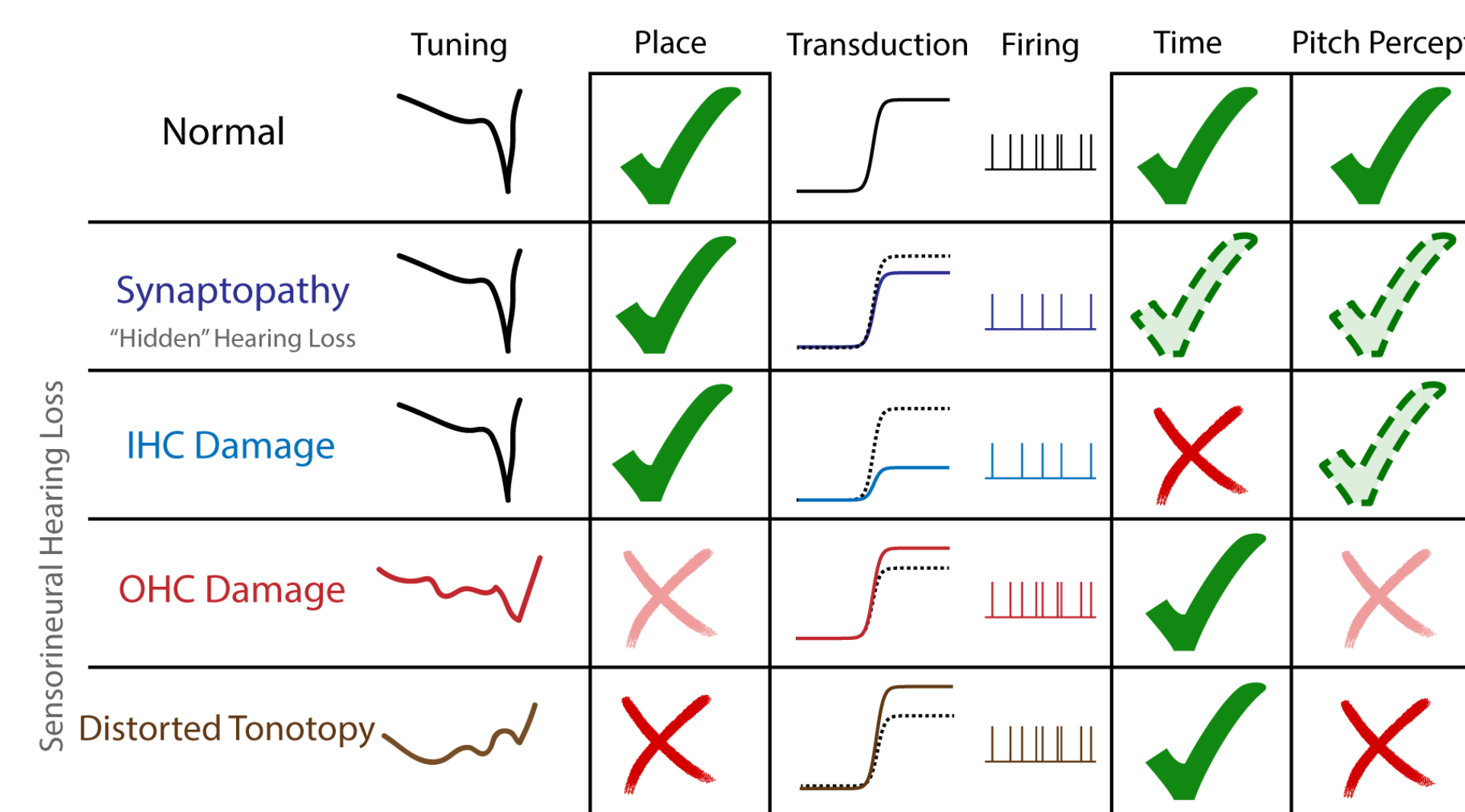


Introduction

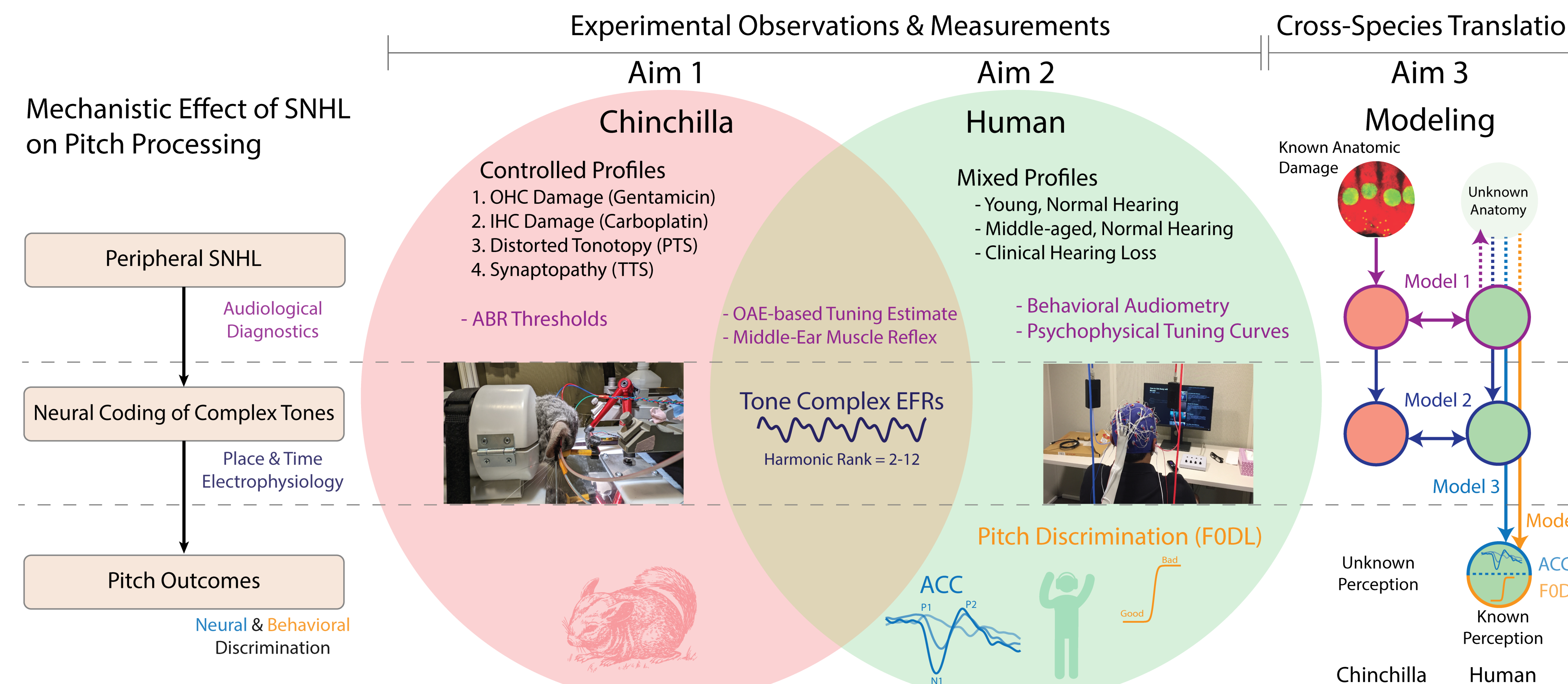
An elusive empirical neural explanation for pitch perception has sparked a multitude of cochlear place and time-dependent hypotheses.

- Longstanding pitch theories weigh the importance of **tonotopy** vs **temporal coding**
- Our understanding of the impact of **Sensorineural Hearing Loss (SNHL)** on pitch perception and neural coding of tone complexes remains convoluted and complex
- **Cochlear Synaptopathy, Inner Hair Cell (IHC), and Outer Hair Cell (OHC) damage** may differentially impact pitch perception, despite often being indistinguishable through non-specific standard clinical assays (i.e. Audiometry, Distortion Product Otoacoustic Emissions)
- **Band-limited tone complexes** are an useful stimuli for probing the fidelity of cochlear time and place cues through both physiological (Envelope Following Responses, EFRs) and behavioral (Fundamental Frequency Difference Limens, FODLs) measures

Here, we describe the development of a novel cross-species study framework that leverages EFRs collected in animal models of hearing loss and human subjects with diverse hearing status to better understand this link.



Framework



SNHL likely results in pitch perceptual deficits dependent on the anatomic profile of cochlear damage.

By utilizing EFRs to tone complex stimuli, controlled models of hearing loss in chinchillas¹ can be leveraged to better understand the mixed profiles of hearing loss that appear in a broad population of human subjects, providing a mechanistic explanation for the variability seen in pitch outcomes like Acoustic Change Complex (ACC), and FODLs.

Stimuli

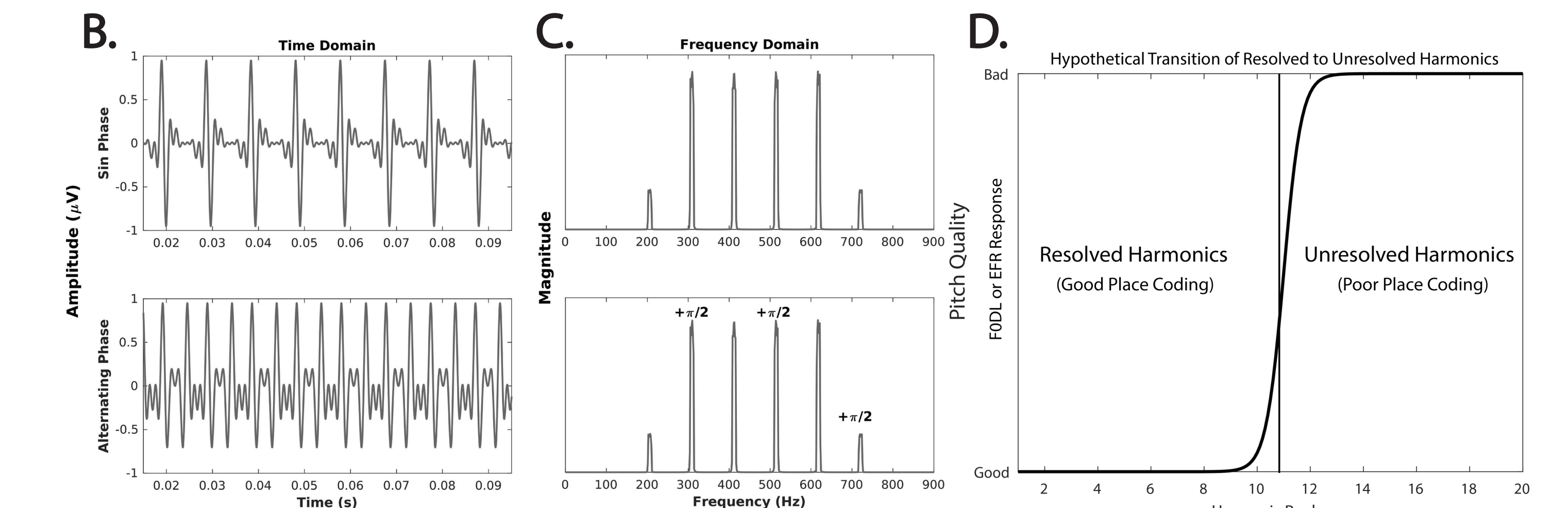
- To finely examine harmonic resolvability, EFRs to six stimuli were selected to investigate neural coding across cochlear place (A).

- The harmonics in each tone complex were presented in alternating (ALT) phase. This results in a temporal envelope modulation of 2*F0 (B), while place cues are spaced 1*F0 apart (C).

- **Physiology** (Human & Chinchilla): This elicits an EFR with periodicity F0 when all harmonics in the complex are resolved, but 2*F0 when harmonics are unresolved².

- **Perception** (Human): Additionally, FODLs measurably increase (discrimination worsens) when harmonics are unresolved by the auditory system^{3,4}.

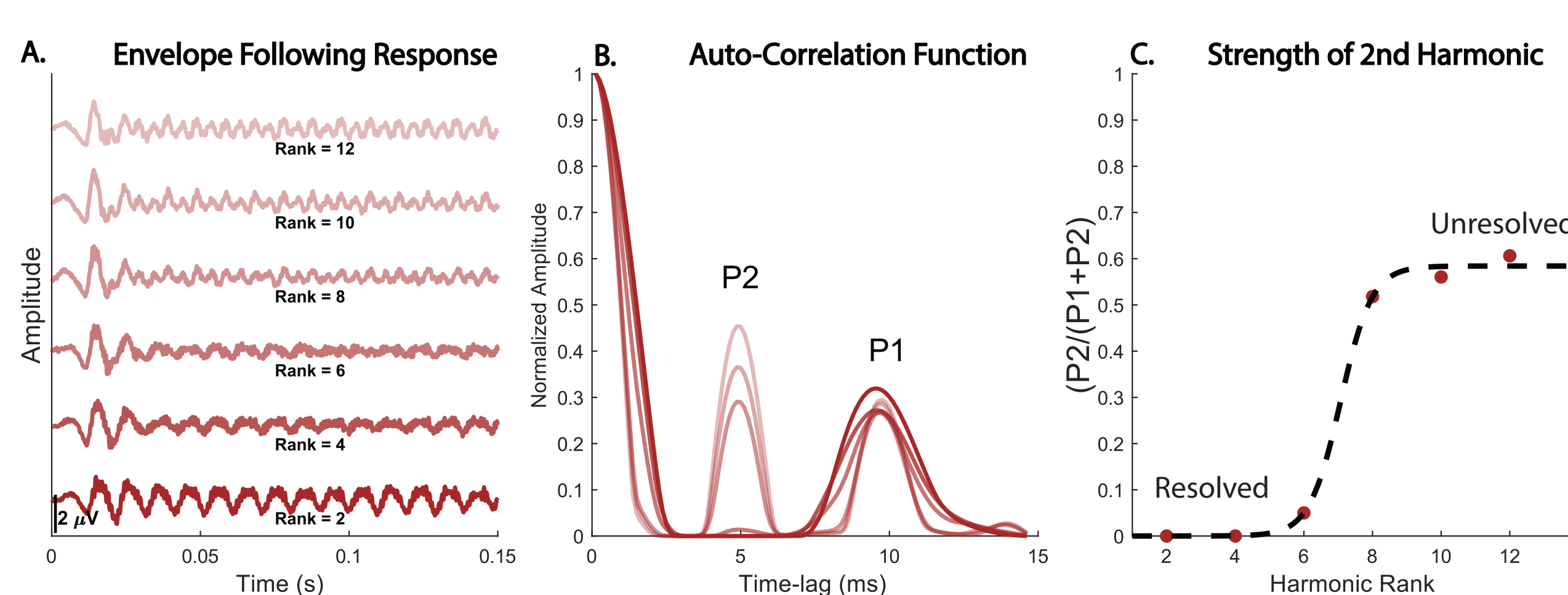
This physiological place transition can be used to directly link neural coding to pitch perception



Results

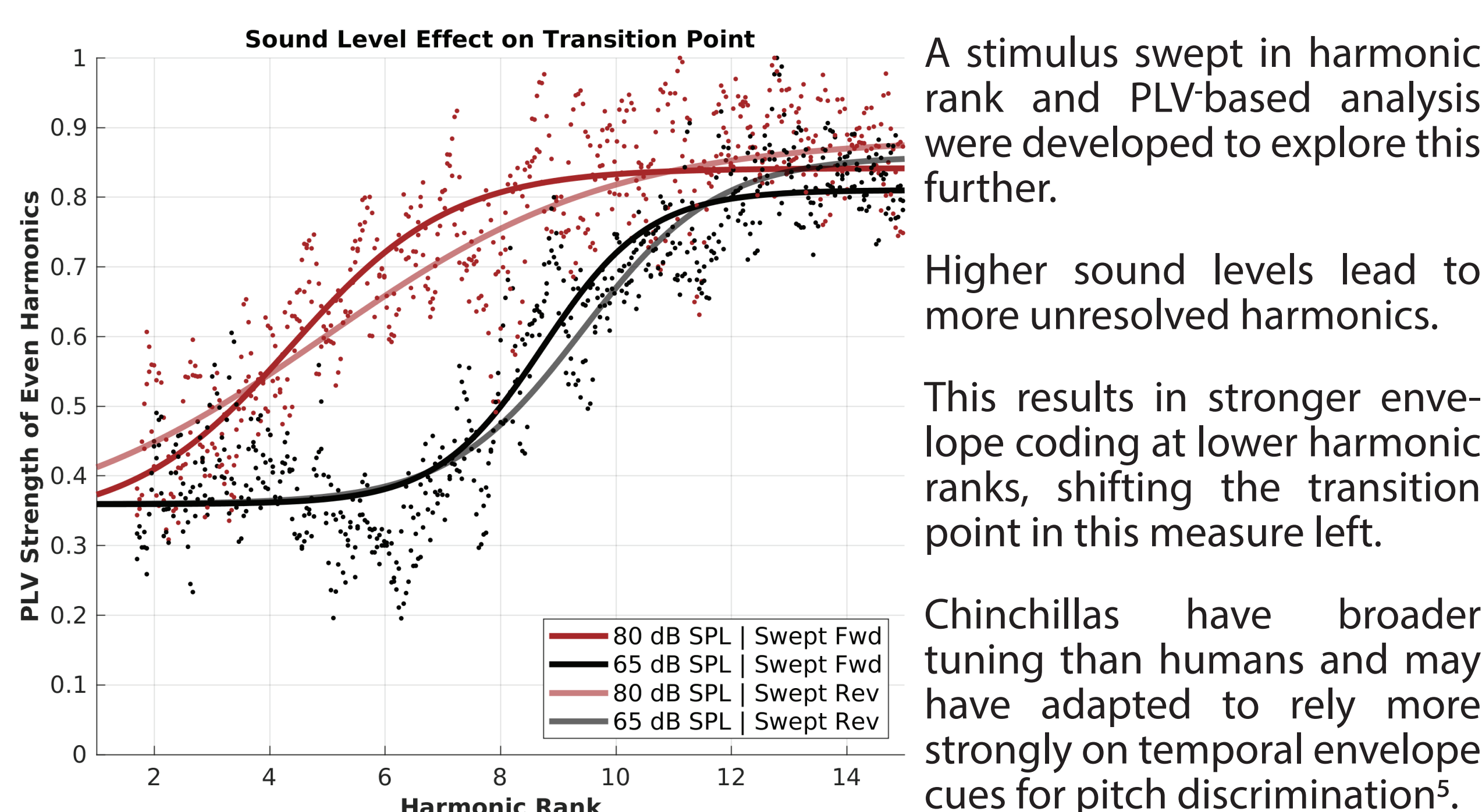
Chinchilla

Physiological Transition Point in a Single Chinchilla

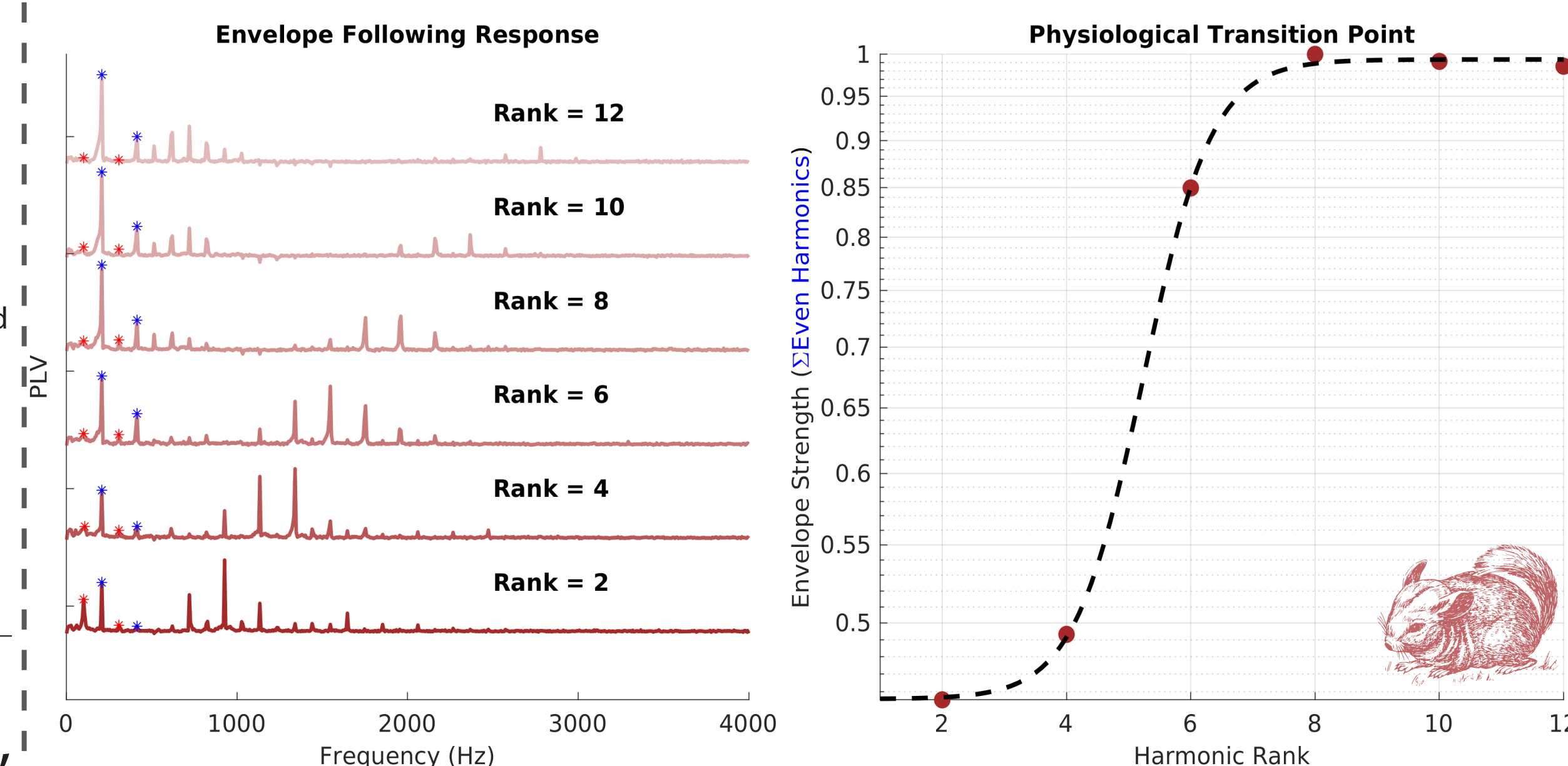


EFRs to the described stimuli demonstrate a resolved-unresolved transition, observable at the individual-subject level. High SNR recordings are made possible through the use of anesthesia and a subdermal electrode montage.

Place Coding is Level-Dependent

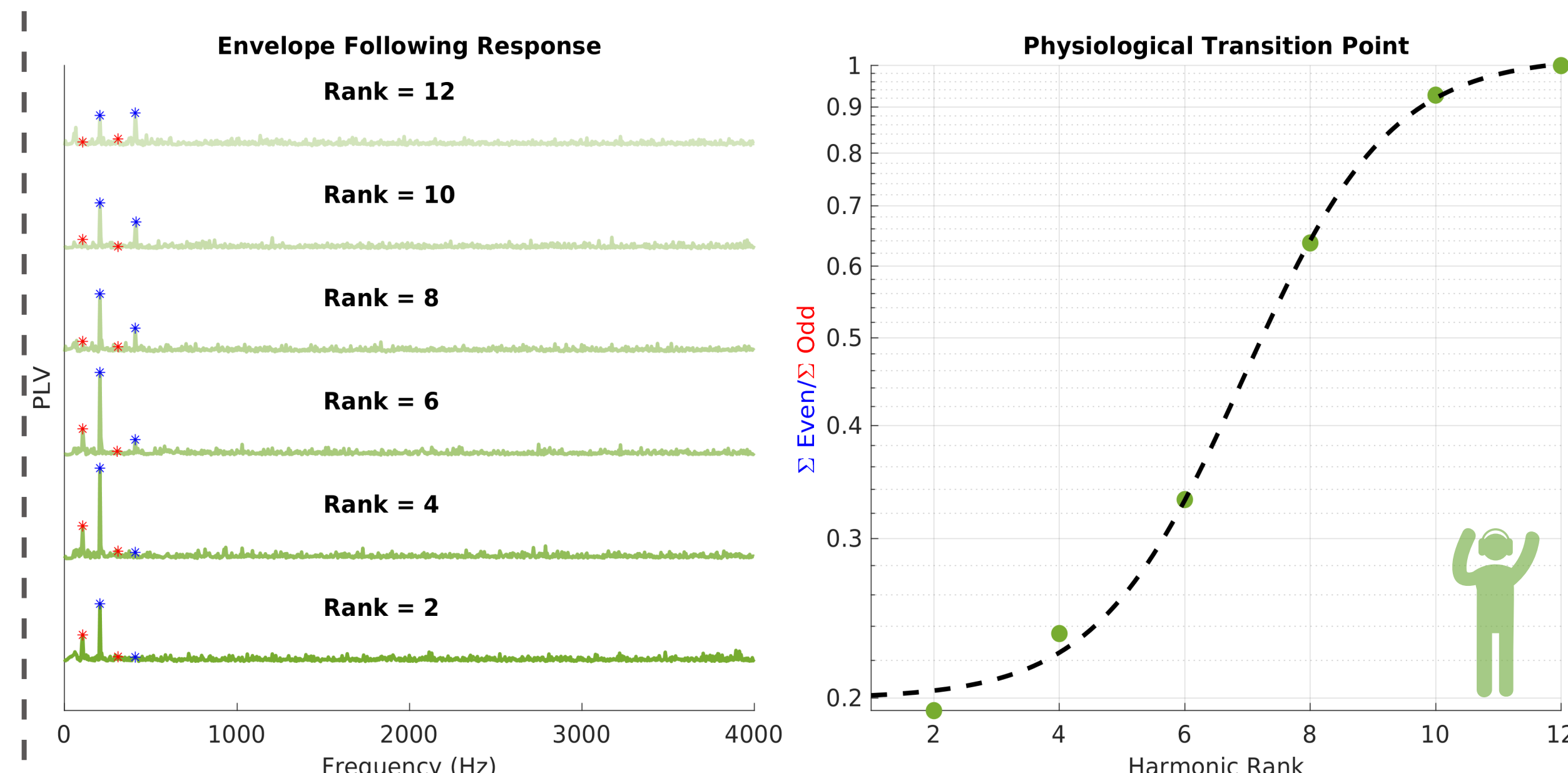


Cross-Species

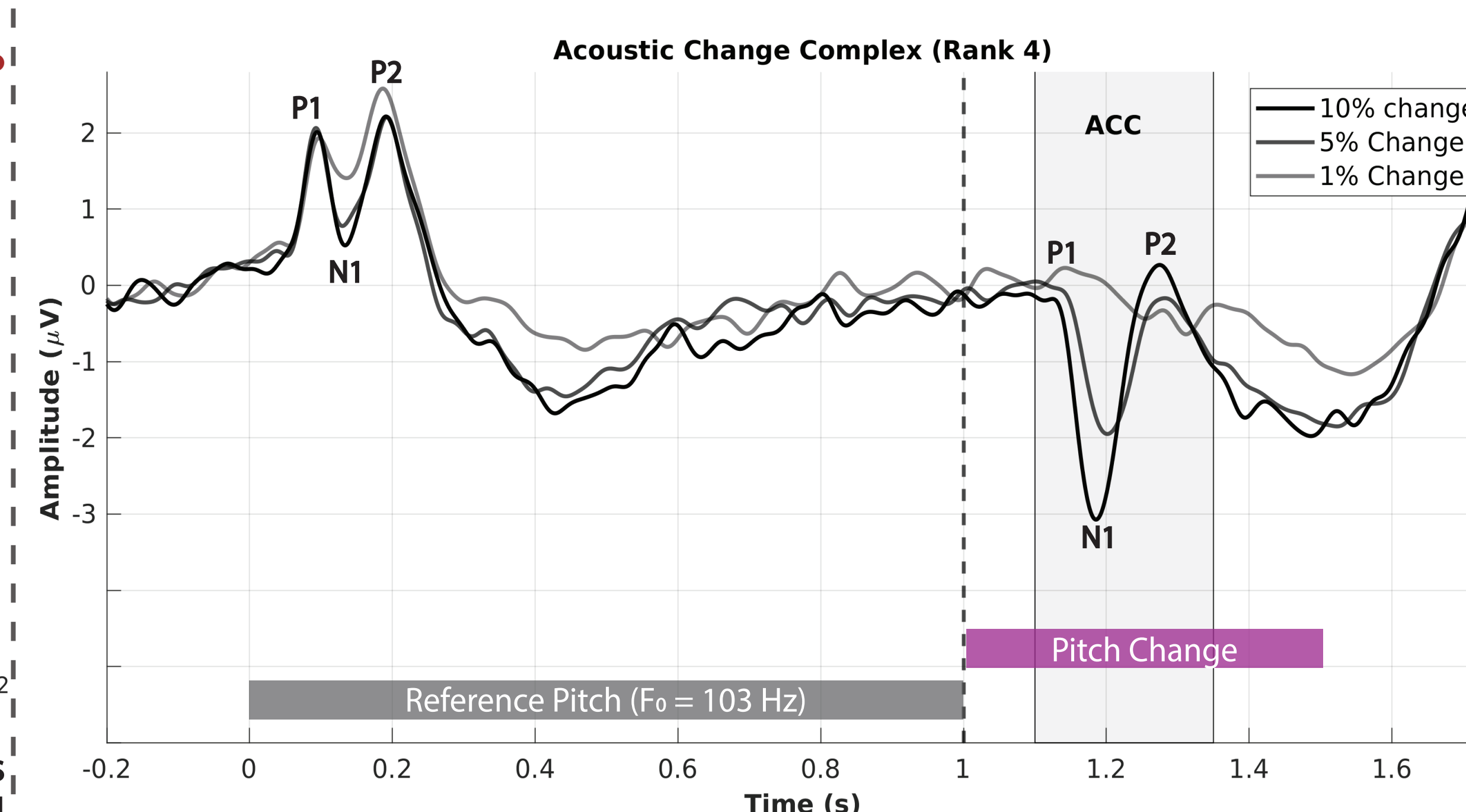


PLV-based spectral analyses were used to directly compare harmonic resolvability across species. The strength of the doubling in envelope periodicity (higher with unresolved harmonics) was quantified using energy in harmonics of 2*F0.

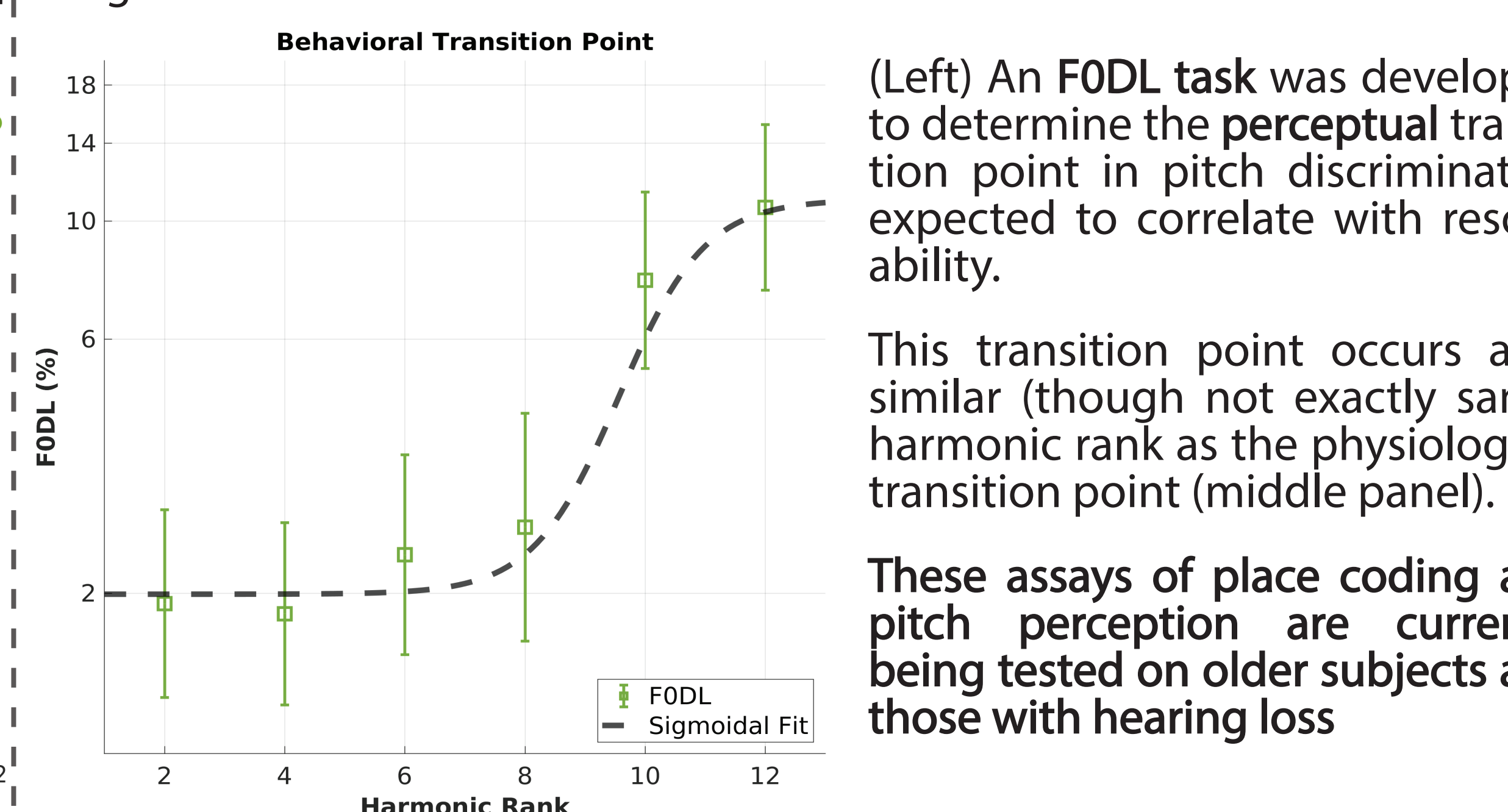
To achieve a clear transition point, stimuli were presented at 65 dB SPL in chinchillas (N=9), and 80 dB SPL in human subjects (N=9). Human subjects appear to have transition point at higher harmonic ranks, likely due to sharper cochlear filtering.



Human



(Above) Neural pitch discrimination in human subjects is assessed using the Acoustic Change Complex. With increasing pitch change, an increasing ACC is observed.

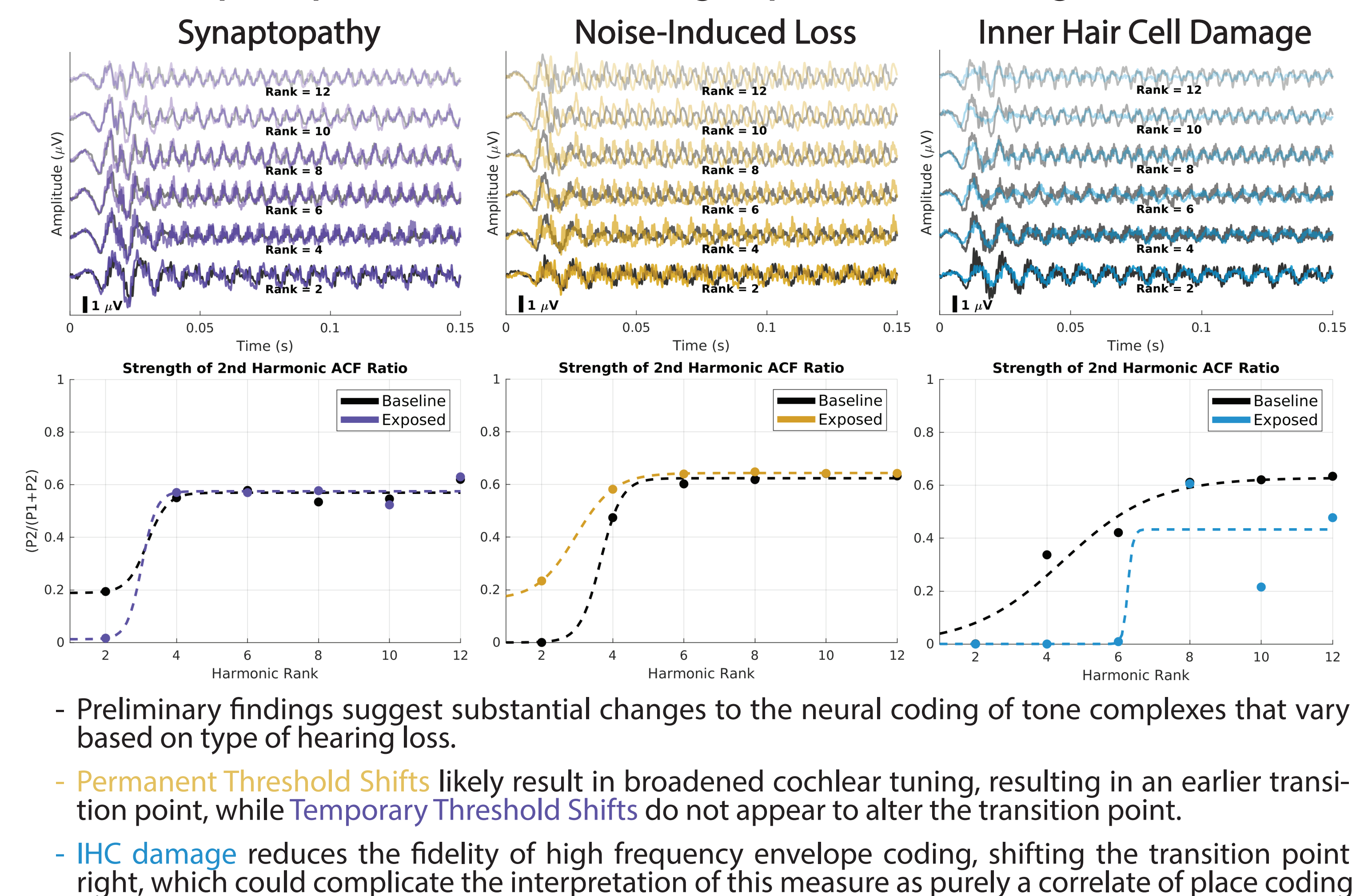


Conclusions

Tone complexes as a function of harmonic ACF rank serve as a useful stimulus set to probe the neural coding of pitch in humans and chinchillas

- Being able to assess pitch perception and its relevant neurophysiology in this described framework allows for a better link between psychoacoustic and physiological studies of pitch.
- Cross-species work is a powerful way to gain mechanistic insights to perceptual outcomes, however differences in cochlear filtering and anatomy should be properly considered.
 - Chinchillas have broader tuning than humans, leading to more unresolved harmonics at lower harmonic ranks, evident in the EFR-based transition point.
- This transition point is highly sensitive to sound level, especially in chinchillas.

This cross-species framework is being used to attribute variability in perception and neural coding to profiles of hearing loss



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- [2] Krishnan, R., et al., Hearing Research, 2011
- [3] Mehta, A., Oxenham, A., JASA, 2021
- [4] Bernstein, J., Oxenham, A., JASA, 2006
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