

## BACKGROUND

### Noise-induced cochlear synaptopathy

- Destruction of synapses between inner hair cells and auditory nerve fibres<sup>1</sup>
- Demonstrated in mice, rats, chinchillas, guinea pigs, and macaques<sup>1,2</sup>
- Can occur without widespread hair cell loss or permanent threshold elevation<sup>1,2</sup>
- Suggested as a cause of perceptual deficits in humans with normal audiograms<sup>1,3</sup>

### Possible measures of cochlear synaptopathy

- Electrophysiological measures, e.g. the amplitude of wave I of the auditory brainstem response (ABR) and the envelope-following response (EFR)
- Basic psychoacoustic tasks, e.g. amplitude modulation detection (AMD), interaural phase difference discrimination (IPD), frequency difference limens (FDL), intensity difference limens (IDL), auditory localisation
- More complex behavioural tasks, e.g. speech perception in noise
- Middle-ear muscle reflex (MEMR); functional magnetic resonance imaging (fMRI)

## FIVE RELATED PROJECTS (2014-19)

All aimed to test for evidence of cochlear synaptopathy in young adults with normal audiometric thresholds from 0.25 to 8 kHz and normal middle-ear function:

Project	Predictors	Outcome measures	Reported in...
<b>Project 1</b> (lead researcher G Prendergast)	Lifetime noise exposure	ABR amplitude EFR amplitude Speech in noise AMD, ITD, FDL, and IDL Localisation Musical consonance	Prendergast et al. (2017a) <sup>4</sup> Prendergast et al. (2017b) <sup>5</sup>
<b>Project 2</b> (lead researcher H Guest)	Tinnitus Impaired speech perception	ABR and EFR amplitude Lifetime noise exposure	Guest et al. (2017) <sup>6</sup> Guest et al. (2018a) <sup>7</sup> Guest et al. (2018b) <sup>8</sup>
<b>Project 3</b> (lead researcher S Couth)	Lifetime noise exposure (musicians)	ABR amplitude Speech in noise	
<b>Project 4</b> (lead researcher R Dewey)	Lifetime noise exposure	ABR amplitude fMRI	Dewey et al. (2018) <sup>9</sup>
<b>Project 5</b> (lead researcher W Tu)	Lifetime noise exposure	ABR amplitude	Prendergast et al. (2018) <sup>10</sup>

## SHARED METHODS

### Auditory brainstem response (ABR)

- All **five projects** used click stimuli, delivered at between 100 and 115 dB peSPL
- Clicks were presented through ER3A insert earphones at rates of 7 to 11 Hz
- Responses were filtered (low-pass cut-off 1500 Hz, high-pass cut-off 30 to 300 Hz)
- Amplitudes of waves I and V were determined using a peak-picking algorithm

### Envelope-following response (EFR)

- **Project 1** and **Project 2** recorded EFRs to transposed tones<sup>11</sup> at 75-80 dB SPL
- $f_c = 4$  kHz (both projects);  $f_m \approx 260$  Hz (**Project 1**);  $f_m = 100$  Hz (**Project 2**)

### Noise Exposure Structured Interview (NESI)

- All **five projects** used versions of this self-report procedure, which allows estimation of lifetime exposure to activities involving estimated sound levels >80 dBA<sup>6</sup>
- The respondent's life is divided into periods when exposure habits were fairly stable
- For each life period, standardised methods are used to estimate total duration and level of exposure, along with usage and attenuation of hearing protection
- Data are combined to yield units linearly related to total energy of exposure >80 dBA

### Coordinate Response Measure (CRM)

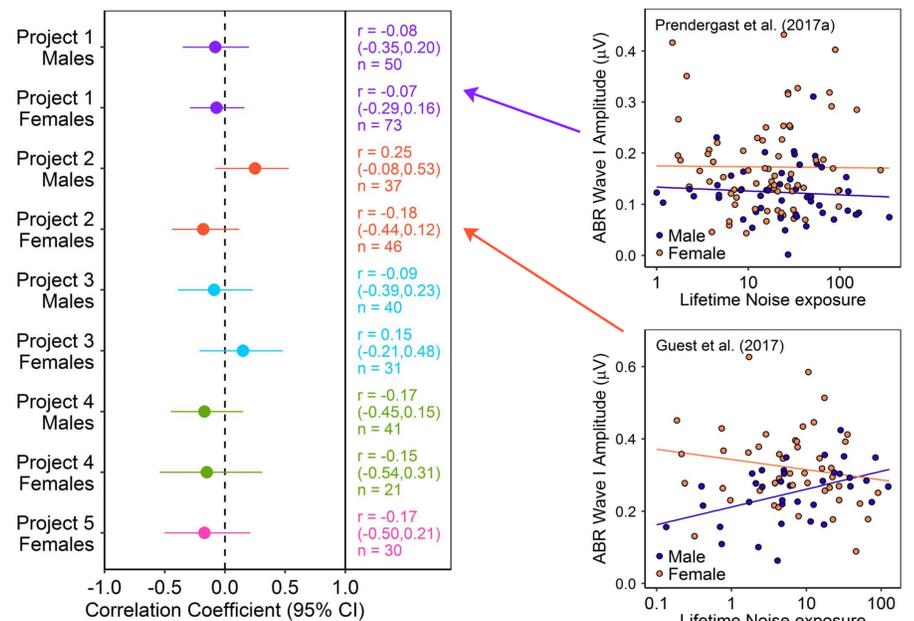
- Measure of speech perception in noise<sup>12</sup> used in **Project 1**, **Project 2**, and **Project 3**
- Incorporates multiple talkers, high overall sound levels, and spatial cues
- Target voice at 0° azimuth, two distractor voices at -60° and +60° azimuth
- Distractor voices held at 80 dB SPL and threshold SNR measured for target voice

## REFERENCES

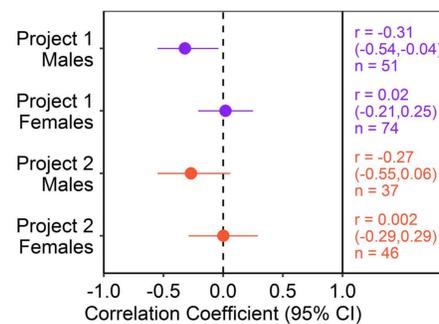
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## RESULTS

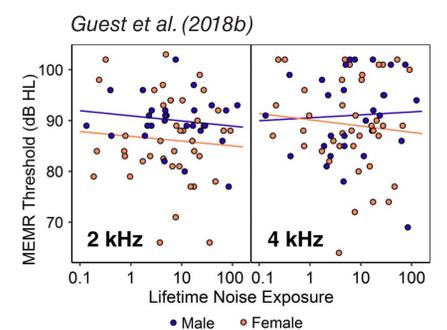
### 1. No significant relation to ABR wave I amplitude



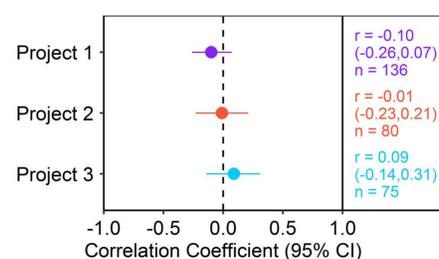
### 2. No significant relation to EFR amplitude



### 3. No significant relation to MEMR threshold

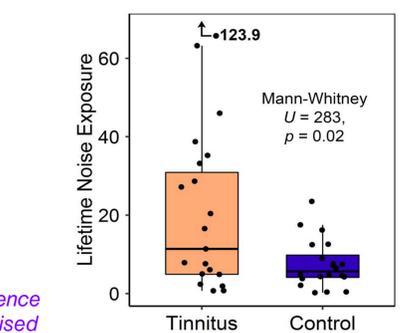


### 4. No significant relation to CRM (speech in noise)

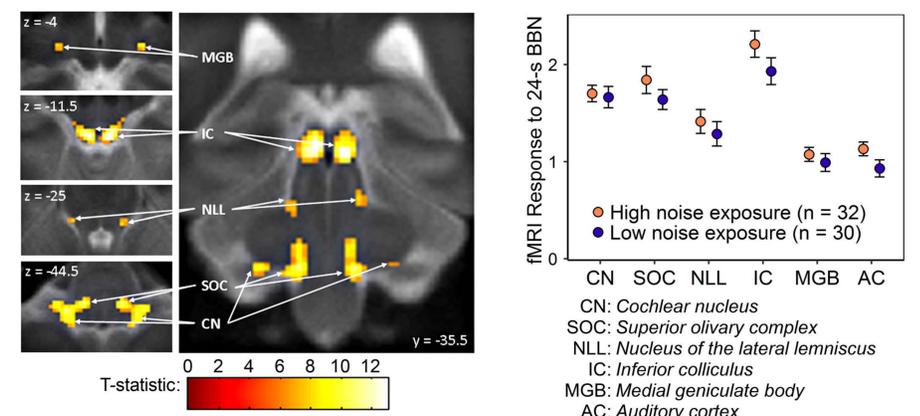


N.B. Better performance -> lower CRM thresholds, hence a positive relation with noise exposure is hypothesised

### 5. Significant association with tinnitus



### 6. No significant association with subcortical or cortical fMRI responses



## CONCLUSIONS

- In young humans with normal audiograms, we find no evidence that lifetime noise exposure is associated with reduced auditory nerve function, reduced perceptual performance, or changes in fMRI responses
- Noise may have the capacity to induce "hidden" changes leading to tinnitus, but we find no evidence that cochlear synaptopathy is one of them