





## ADVANCED ACOUSTICS FOR DIAGNOSTIC APPLICATIONS TO AUDIOLOGY AND BEYOND

PhD admission to 3<sup>rd</sup> Year

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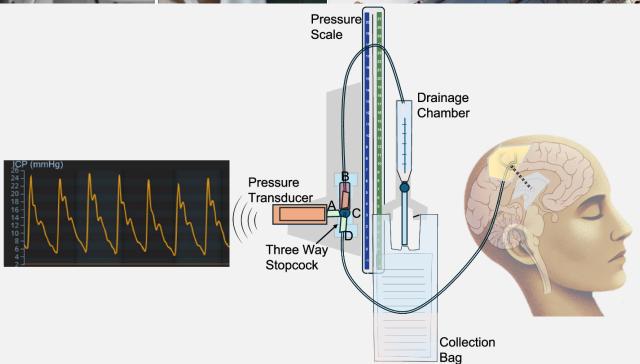
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**Dr. Renata Sisto** 

INAIL Research, Rome, Italy

15<sup>th</sup> October 2025





## INTRODUCTION

- The International Space Station (ISS) is a unique stationary free-falling frame, which permits studying the effects of long-term microgravity on the astronauts' physiology.
- Microgravity conditions have various effects on human physiology, including increased Intracranial Pressure (ICP), which poses risks to vision, especially for long-duration space missions.
- Current methods for measuring ICP are invasive and carry risks.

1 Ultrasound-Based Internal Jugular Vein Pressure

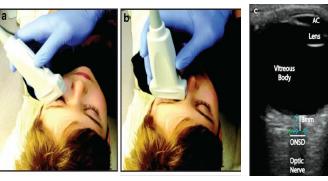


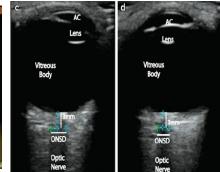
**Ultrasound of Optic Nerve Sheath Diameter (ONSD)** 











3

#### **Otoacoustic Emissions (OAE)**

The phase of OAEs is dependent on the middle ear transmission. The relationship between OAE responses and direct ICP measures is based on pressure communication between perilymph and cerebrospinal fluid (CSF), and on the dependence of the middle ear transmission on intracochlear pressure.

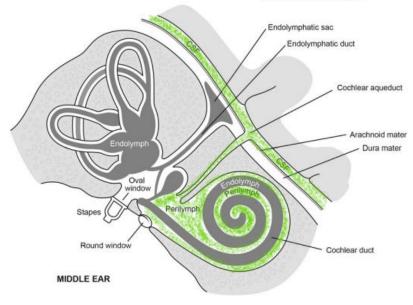
Avan and coworkers reported a linear relationship between DPOAE phase change and ICP change:

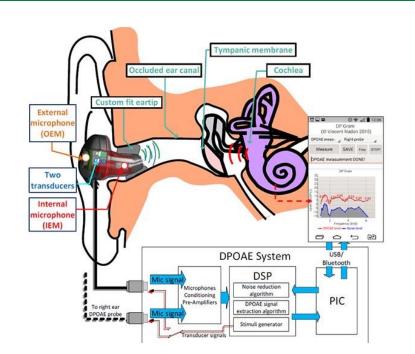
$$\frac{\partial \Phi}{\partial ICP} = 2.72 \text{ deg/mmHg}$$

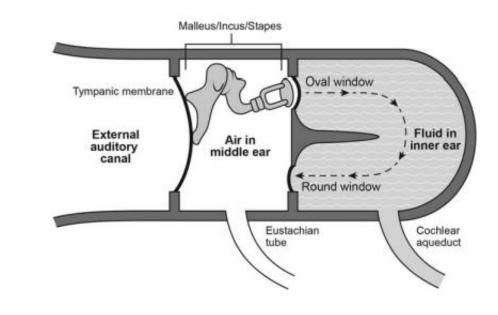


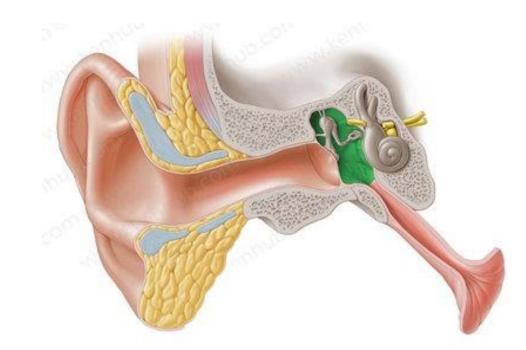
#### Ear Anatomy and Physiology

#### SUBARACHNOID SPACE









## What are Otoacoustic Emissions (OAEs)?

Otoacoustic emissions (OAEs) are low-level sounds generated within the inner ear, specifically by the outer hair cells of the cochlea. cells of the cochlea. These sounds can be measured in the ear canal using sensitive microphones. Their presence indicates healthy indicates healthy cochlear function.



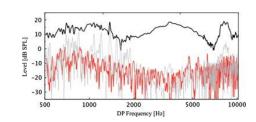
OAEs are used in newborn hearing screenings and to and to diagnose certain types of hearing loss.



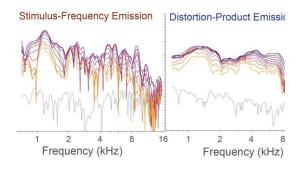


Types of OAEs

Spontaneous OAEs (SOAEs), Transient-Evoked OAEs (TEOAEs), and Distortion Product OAEs (DPOAEs)



Otoacoustic emission	Notation	Stimulus	Prevalence (by ear)
Spontaneous Evoked:	SOAE	None	~50%
Transiently evoked	TEOAE	Click/toneburst	~100%
Stimulus-frequency	SFOAE	Continuous pure tone	~100%
Distortion-product	DPOAE	2 continuous pure tones	~100%

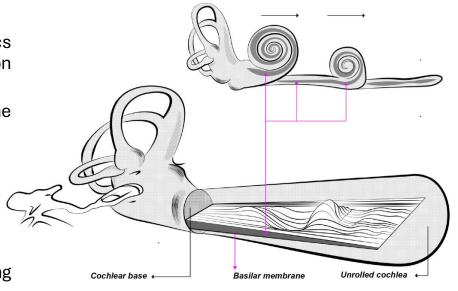


- Cochlear mechanics describes the propagation, reflection and absorption of traveling waves (TWs) along the cochlea.
- Slow transverse TWs are generated by the coupling between the cochlear fluid hydrodynamics and the transverse motion of an elastic complex mechanical structure, the cochlear partition (CP).
- This two-way coupling consists of the mechanical drive by the differential fluid pressure on the CP,

$$\ddot{\xi}(x,t) + \gamma_{bm}(x)\dot{\xi}(x,t) + \omega_{bm}^2(x)\xi(x,t) = -\frac{p_d(x,t)}{\sigma_{bm}}$$

where 
$$\omega_{bm}(x) = \omega_0 e^{-k_{\omega}x}$$

and on the mechanical reaction of the incompressible fluid to the CP movement, which, using Newton's equation for the fluid element yields a TW eq.



Fluid Volume conservation  $H \frac{\partial u}{\partial x} = \dot{\xi}$ Newton's law for fluid element  $-\frac{\partial p}{\partial x} = \rho \frac{\partial u}{\partial t}$ Propagation of a pressure wave

$$-\frac{\partial p}{\partial x} = \rho \frac{\partial u}{\partial t}$$



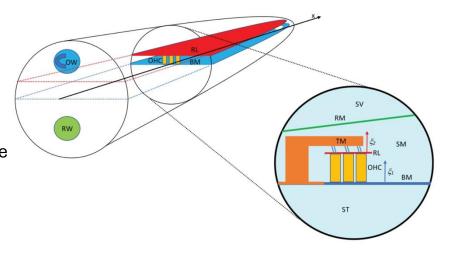
$$\frac{\partial^2 p_d}{\partial x^2} = -2\frac{\rho}{H} \ddot{\xi}$$

In the linear approximation, in the frequency domain, these equations yield a transmission line equation

$$\frac{\partial^2 p_d}{\partial x^2} = -k^2 p_d$$

with

$$k^{2} = \frac{2\rho}{H\sigma_{bm}} \frac{\omega^{2}}{\left(\omega_{bm}^{2}(x) - \omega^{2} + i\omega\gamma_{bm}(x)\right)}$$



#### WKB solution: Traveling waves

• Forward TW: 
$$\psi_r(\omega,x) = \sqrt{\frac{k(\omega,x)}{k(\omega,0)}} e^{-i\int_0^x k(\omega,x')dx'}$$

$$\qquad \qquad \psi_l(\omega,x) = \sqrt{\frac{k(\omega,x)}{k(\omega,0)}} e^{i\int_0^x k(\omega,x')dx'}$$

• OAEs are associated with backscattered waves generated by two mechanisms: nonlinear distortion and coherent reflection by randomly distributed impedance perturbation (roughness).

• Two DPOAE components are generated by these mechanisms, in different cochlear regions.

Due to cochlear scaling symmetry, the wavenumber is a function of the dimensionless ratio [ $\omega/\omega(x)$ ], so its total derivative is null:

$$0 = \frac{dk(\omega, x)}{d\omega} = \frac{\partial k(\omega, x)}{\partial \omega} + \frac{\partial k(\omega, x)}{\partial x} \frac{\partial x}{\partial \omega}$$

Hence, model-independent prediction of the frequency dependence of the group delay of the two components is possible. Using this property, one may compute the group delay of a TW element of frequency  $\omega$  associated with its longitudinal transmission from the base to its OAE generation place  $x_g(\omega)$ :

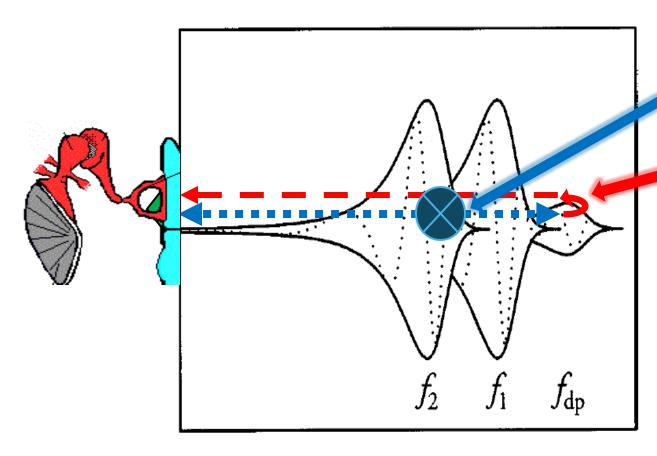
#### **Place-fixed OAEs**

# $\tau_{0,x_g}(\omega) = -\frac{\partial \Phi(\omega, x_g)}{\partial \omega} = \int_0^{x_g} \frac{\partial Re(k(\omega, x'))}{\partial \omega} dx'$ $= \frac{1}{k_{\omega}\omega} \int_0^{x_g} \frac{\partial Re(k(\omega, x'))}{\partial x'} dx' = \frac{Re(k(x_g, \omega) - k(0, \omega))}{k_{\omega}\omega}$ $\approx \frac{Re(k(x_g, \omega))}{k_{\omega}\omega}$

#### Wave-fixed OAEs

$$\begin{split} &\tau_{0,x(\omega)}(\omega) = -\frac{\partial \Phi(\omega,x(\omega)))}{\partial \omega} \\ &= \int_{0}^{x(\omega)} \frac{\partial Re(k(\omega,x'))}{\partial \omega} dx' + \frac{\partial x(\omega)}{\partial \omega} Re(k(\omega,x(\omega))) \\ &= \frac{1}{k_{\omega}\omega} \Big( \int_{0}^{x(\omega)} \frac{\partial Re(k(\omega,x'))}{\partial x'} dx' - Re(k(\omega,x(\omega))) \Big) \approx 0 \end{split}$$

#### **Distortion Product OAE Generation Mechanism**



Position *x* 

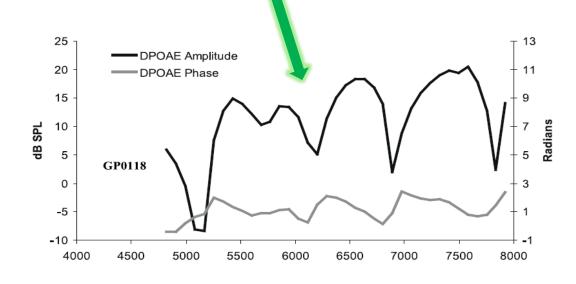
Group delay: 
$$\tau = -\frac{\partial \varphi}{\partial \omega}$$

first **wave-fixed** generation of forward and backward DP waves in the overlap region  $x(f_2)$ .

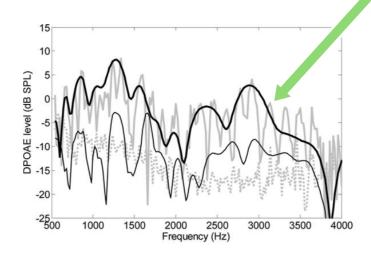
Cochlear scaling symmetry predicts **NULL GROUP DELAY** 

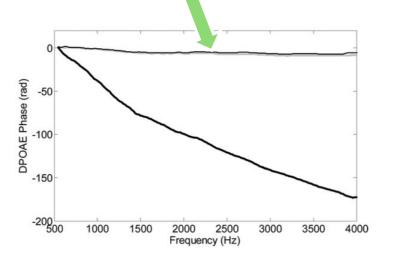
**place-fixed** reflection of the forward DP wave at  $x(f_{dp})$ . Cochlear scaling symmetry predicts **LONG GROUP DELAY** inversely proportional to frequency

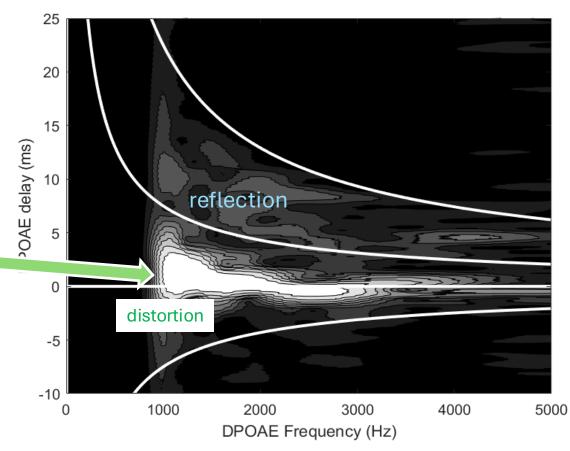
linear superposition of the two components yields DPOAE spectral fine-structure



- Separation between the two components based on their different group delay is optimally performed in the timefrequency domain using the scaling-symmetric wavelet transform.
- Analyzing the two components separately improves the specificity of the technique and increases the SNR, due to the localization of the signal and the homogeneous distribution of noise in the t-f plane.
- The intrinsically constant phase of the DPOAE distortion component allows unambiguous averaging of small phase differences over a broad frequency band, with further SNR improvement.

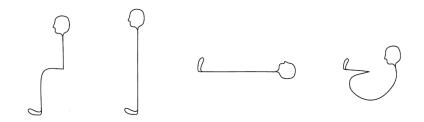






#### **EXPERIMENT WITH POSTURAL DIFFERENCES ON GROUND**





Ground Experiments (20 subjects)

4 postures: Seated, Standing, Supine, Fetal (Lateral recumbent)



#### **DPOAE Technique**



Swept-tone chirps: 1-6 kHz



Stimuli: 65 & 55 dB, FPL-equalized



Recorded via ER10B+ otoacoustic probe



High-resolution time-frequency analysis (wavelet-based)

## **Calibration & Reflectance Analysis**



THEVENIN-BASED FPL CALIBRATION



ESTIMATION OF EAR
CANAL IMPEDANCE AND
ENERGY REFLECTANCE



REFLECTANCE SHIFT AND IMPEDANCE CHANGES
WERE ANALYSED

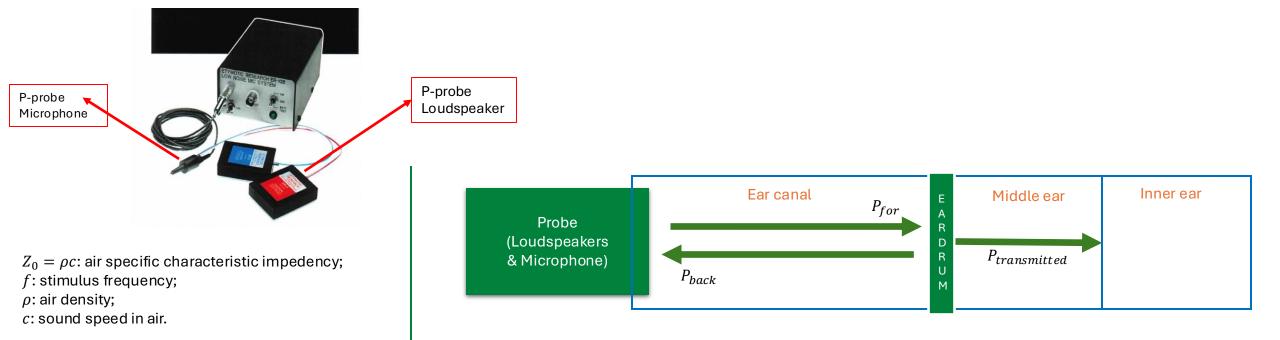
#### **Energy Reflectance And Impedance**

 $Pressure\ (P)$  has been measured into the ear canal by an Etymotic ER-10B+ microphone equipped with two ER-2 loudspeakers.

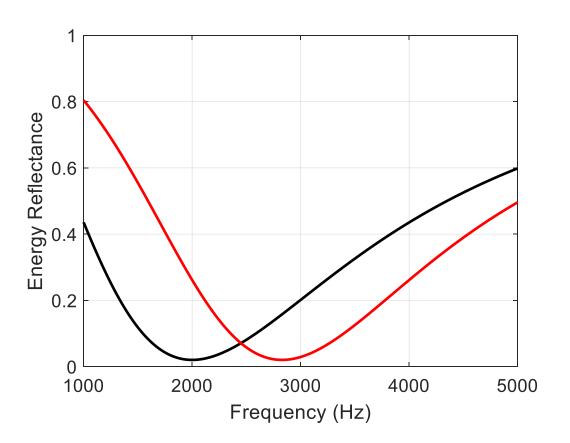
Preliminarily, the probe was calibrated, measuring its Thevenin equivalent parameters in terms of pressure  $P_{Th}$  and specific impedance  $Z_{Th}$ , using five cylindrical cavities of known acoustic impedance. Once  $P_{Th}$  and  $Z_{Th}$  are known, Z is computed from P and the eardrum energy reflectance is expressed in terms of Z:

$$Z(f) = \frac{P(f) Z_{Th}(f)}{P_{Th}(f) - P(f)} \qquad R(f) = R(f)$$

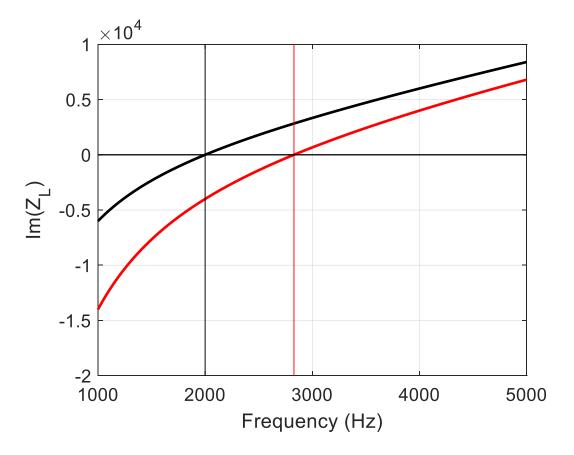
 $R(f) = \left| \frac{Z(f) - Z_0}{Z(f) + Z_0} \right|^2$ 



**Expected change** of the middle ear resonance frequency for a twofold increase of the middle ear stiffness (black to red line)

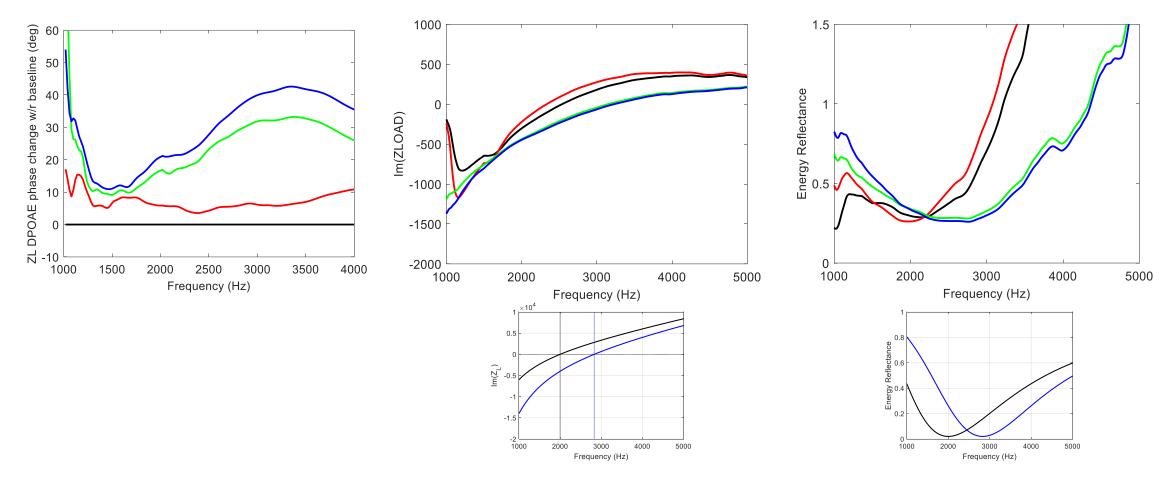


**Expected change** of the imaginary part of the middle ear impedance for a twofold increase of the middle ear stiffness (black to red line)



## SAMPLE RESULTS - GROUND POSTURE EXPERIMENT

- Clear shift in: DPOAE phase, Energy reflectance minima, imaginary part of the load impedance between vertical
  postures (black and red lines) and horizontal postures (green and blue)
- Using Avan's relationship, ICP estimated to increase 6–10 mmHg in horizontal postures with respect to vertical ones



#### **EXPERIMENT WITH THE ASTRONAUTS IN MICROGRAVITY**



DPOAE Recordings at different time intervals

1 Pre-flight/ Baseline 6 In-flight 2 Post flight

OAE tests were performed pre/post and during ISS spaceflight by 5 astronauts



- Distortion Product OAE
- Advanced analysis tools based on theoretical cochlear mechanics
- Normal Hearing Sensitivity

Journal of the Association for Research in Otolaryngology https://doi.org/10.1007/s10162-024-00962-1 JARO

ORIGINAL ARTICLE



#### Otoacoustic Estimate of Astronauts' Intracranial Pressure Changes During Spaceflight

Arturo Moleti¹ · Triestino Minniti¹ · Yoshita Sharma¹ · Altea Russo¹ · Andrea Civiero² · Maria Patrizia Orlando³ · Robert MacGregor⁴ · Marco Lucertini⁵ · Arnaldo D'Amico⁵ · Giorgio Pennazza² · Marco Santonico² · Alessandro Companti² · Alessandro Crisafi⁵ · Maurizio Deffacis⁵ · Rosa Sapone⁵ · Gabriele Mascetti³ · Monia Vadrucci⁵ · Giovanni Valentini⁵ · Dario Castagnolo¹0 · Teresa Botti¹¹ · Luigi Cerini¹¹ · Filippo Sanjust¹¹ · Renata Sisto¹¹

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#### Abstract

Purpose To investigate the potential correlation between prolonged exposure to microgravity on the International Space Station and increased intracranial fluid pressure, which is considered a risk factor for the astronauts' vision, and to explore the feasibility of using distortion product otoacoustic emissions as a non-invasive in-flight monitor for intracranial pressure changes.

Methods Distortion product otoacoustic emission phase measurements were taken from both ears of five astronauts pre-

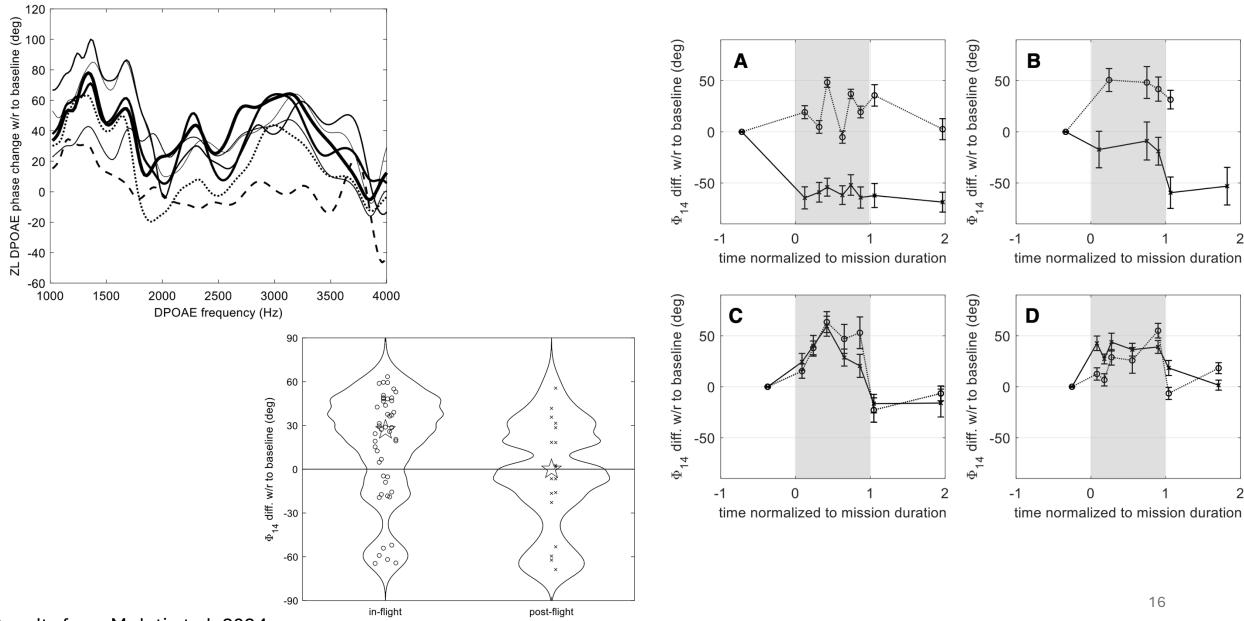
**Methods** Distortion product ofoacoustic emission phase measurements were taken from both ears of five astronauts preflight, in-flight, and post-flight. These measurements served as indirect indicators of intracranial pressure changes, given their high sensitivity to middle ear transmission alterations. The baseline pre-flight ground measurements were taken in the seated upright position.

Results In-flight measurements revealed a significant systematic increase in otoacoustic phase, indicating elevated intracranial pressure during spaceflight compared to scated upright pre-flight ground baseline. Noteworthy, in two astronauts, strong agreement was also observed between the time course of the phase changes measured in the two ears during and after the mission. Reproducibility and stability of the probe placement in the ear canal were recognized as a critical issue.

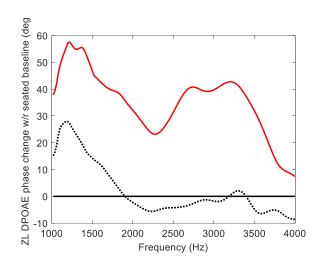
Conclusions The study suggests that distortion product otoacoustic emissions hold promise as a non-invasive tool for monitoring intracranial pressure changes in astronauts during space missions. Pre-flight measurements in different body postures and probe fitting strategies based on the individual ear morphology are needed to validate and refine this approach.

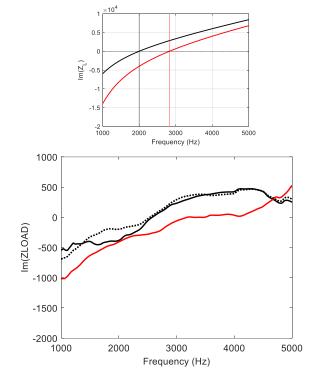
Keywords Microgravity · Otoacoustic emissions · Middle ear transmission · Astronaut physiology

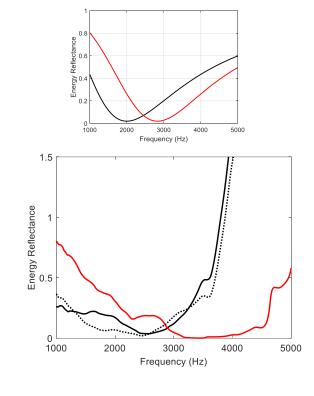
Using these advanced signal filtering tools based on cochlear mechanics improved the accuracy of the DPOAE phase estimates on the ISS, demonstrating significant ICP increase during spaceflight and full recovery after return to terrestrial gravity:



Results from Moleti et al, 2024







## SAMPLE RESULTS – ISS SPACEFLIGHT EXPERIMENT

For Reflectance and Impedance

- Consistent changes of the three ICP indicators (DPOAE phase, the imaginary part of the load impedance the middle ear resonance frequency) in microgravity (red line) vs. pre-/post-flight (seated) data (black solid/dotted). Position is seated in the pre and post flight conditions
- Ground experiments showed that acoustic markers are less affected by posture variability than lumbar puncture (LP), which is an invasive method

#### **EXPERIMENT AT CHU MONTPELLIER WITH PEDIATRIC POPULATION**



Increased patency of the Cochlear Aqueduct in the paediatric population (Abbas, M., Wang, J., Leboucq, N. et al., 2024)

CA is the connection between Cochlea and CSF. If ICP increases, it leads to increased Intra Cochlear Pressure.



Postures (High ICP Vs Low ICP)

2 postures: Vertical/ Tilted and Horizontal



**Groups** (normal hearing sensitivity)

<1; 1-2, 3-6; 7-10; 10-14, Adults

### **DPOAE Technique**



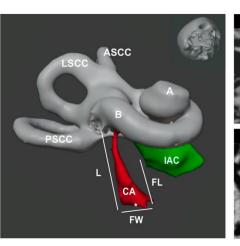
Swept-tone chirps: 500-16 kHz

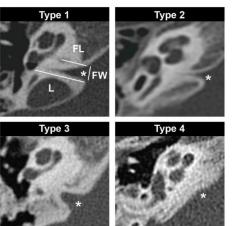


Stimuli: 65 & 55 dB, FPL-equalized



High-resolution time-frequency analysis (wavelet-based)





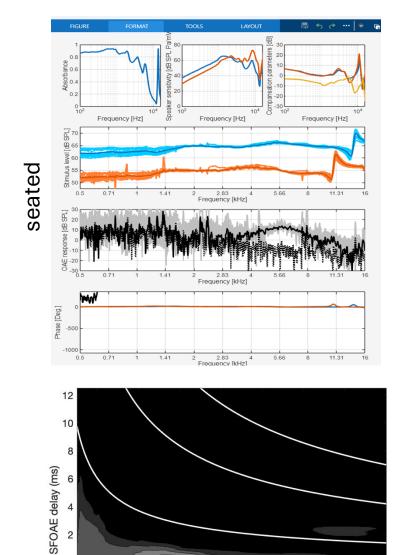
## **Data Analysis**





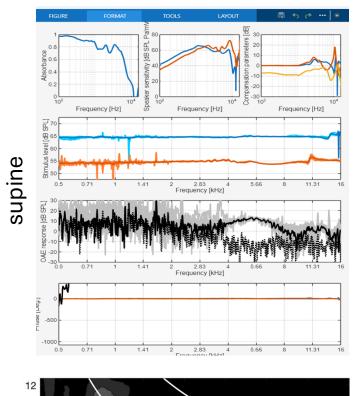
HIGH SNR DPOAE RECORDING

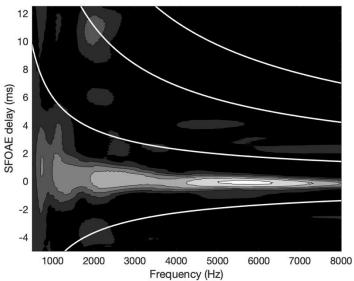
POST DATA COLLECTION (ESTIMATION OF DPOAE PHASE)



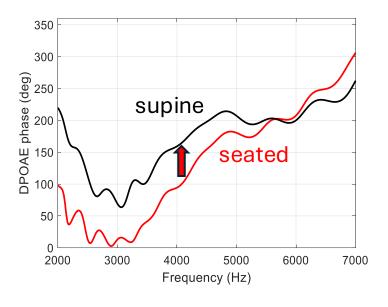
-2

Frequency (Hz)





#### **DPOAE Phase**



## **DISCUSSION**

#### **METHODOLOGICAL ADVANTAGES**

- Non-invasive, objective technique
- Consistency among independent indicators (DPOAE phase, Middle Ear Reflectance, Imaginary part of the impedance) assesses reliability of the OAE phase increase as an indicator of increased ICP.

Advanced filtering enhances specificity and reduces experimental uncertainties.

#### Conferences

## Oral Presentation | Global Space Exploration Conference, New Delhi, India (07/05/2025)

- Otoacoustic Estimate of the Intracranial Pressure. Authors: **Yoshita Sharma**, Arturo Moleti, Triestino Minniti, Teresa Botti, Luigi Cerini, Filippo Sanjust, Renata Sisto

## Oral Presentation | 75th International Astronautical Congress (IAC), Milan, Italy, (14-18 October 2024)

- Advanced audiological tools for non-invasive monitor of intracranial pressure in microgravity. Authors: Arturo Moleti, Triestino Minniti, Altea Russo, **Yoshita Sharma**, Andrea Civiero, Renata Sisto et al.

## Poster Presentation | Auditory Science Meeting 2024 in Cambridge, United Kingdom (September, 2024)

- Advanced Otoacoustic Emission Acquisition and Analysis for the Objective Diagnostics of Hearing. Authors: **Yoshita Sharma**, Moleti A, Minniti, T, Botti T, Cerini L, Sanjust F, Sisto R.

## Poster Presentation | Inner Ear Biology Conference 2024, Warsaw, Poland (September, 2024)

- Non-invasive monitoring of Intracranial Pressure changes: utilizing Otoacoustic Emissions. Authors: **Yoshita Sharma**, Moleti A, Minniti, T, Botti T, Cerini L, Sanjust F, Sisto R.

#### **Publications and Conference Proceedings**

Ongoing: **Sharma Y**, Sisto R, Moleti A, Botti T, Mishra S. Stimulus-Frequency Otoacoustic Emission delay measurements as a model-independent test of cochlear models.

**Sharma Y**, Moleti A, Minniti, T, Botti T, Cerini L, Sanjust F, Sisto R. Otoacoustic Estimate of the Intracranial Pressure. May 2025

DOI: 10.52202/080556-0012. Conference: Global Space Exploration Conference (GLEX 2025) At: New Delhi, India

Moleti A, Minniti T, Russo A, **Sharma Y**, Civiero A, Sisto R, et al. Advanced Audiological Tools for Non-Invasive Monitor of Intracranial Pressure in Microgravity. January 2024

DOI: 10.52202/078355-0014. Conference: IAF/IAA Space Life Sciences Symposium, Held at the 75th International Astronautical Congress (IAC 2024)

Moleti A, Minniti T, **Sharma Y**, et al. (2024) Otoacoustic Estimate of Astronauts' Intracranial Pressure Changes During Spaceflight. Journal of the Association for Research in Otolaryngology: JARO, 25(6), 563–573

#### Future Work - Plan for the Third Year

#### **EXPERIMENT IN BELGIUM IN FEBRUARY 2026:**

Test the method in **tinnitus and hyperacusis patients** to explore cochlear mechanics and middle ear transmission changes.

#### **EXPERIMENT AT GEMELLI UNIVERSITY POLICLINICO:**

Calibrate the OAE-ICP relationship during **controlled ICP changes** in neurosurgical patients during neurosurgeries.

#### TO BE FUNDED BY ITALIAN SPACE AGENCY:

To refine the OAE technique in microgravity for astronauts using customised earplugs.

#### TO BE FUNDED BY ITALIAN SPACE AGENCY:

Perform **bedrest study** to monitory long-term ICP elevation on Earth using OAEs.

#### **EXPERIMENT AT FONDAZIONE BIETTI:**

Apply the technique to **glaucoma patients** with elevated ICP for clinical validation.

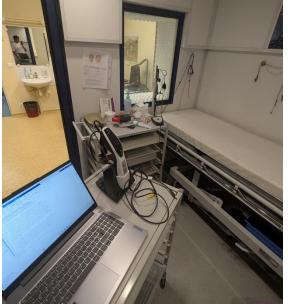
#### **Academic Visits**

Period Abroad for **2 months** in **CHU, Montpellier, France** from 29<sup>th</sup> July 2025 to 5<sup>th</sup> October 2025









#### **ACKNOWLEDGMENTS & REFERENCES**













#### **Key references:**

- 1. Moleti, A., Minniti, T., Sharma, Y., et al. (2024) Otoacoustic Estimate of Astronauts' Intracranial Pressure Changes During Spaceflight. Journal of the Association for Research in Otolaryngology: JARO, 25(6), 563–573
- 2. Sisto R, Botti T, Cerini L, et al. (2023) Otoacoustic emission phase measurements to detect posture-induced intracranial fluid pressure variations in noisy environments. In: 29th International Congress on Sound and Vibration (ICSV29): Prague, Czech Republic, 9–13 July 2023. Red Hook, NY: Curran Associates, Inc
- 3. Pedersen SH, Andresen M, Lilja-Cyron A, et al. (2021) Lumbar puncture position influences intracranial pressure. Acta Neurochir (Wien) 163 1997–2004
- 4. Abbas, M., Wang, J., Leboucq, N. et al. Cochlear Aqueduct Post-Natal Growth: A Computed Tomography Study. JARO 25, 611–617 (2024).

## THANK YOU FOR YOUR ATTENTION



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