DATA NOTE CHECKS

Open Access



Gender specific click and tone burst evoked ABR datasets from mice lacking the Ca_v3.2 T-type voltage-gated calcium channel

Andreas Lundt¹, Christina Henseler¹, Carola Wormuth¹, Julien Soos¹, Robin Seidel², Ralf Müller³, Muhammad Imran Arshaad¹, Karl Broich², Jürgen Hescheler⁴, Agapios Sachinidis⁴, Dan Ehninger⁵, Anna Papazoglou¹ and Marco Weiergräber^{1*}

Abstract

Objectives: Voltage-gated Ca^{2+} channels (VGCCs) are of central relevance in regulating Ca^{2+} influx into living cells. The low-voltage activated (LVA) $Ca_v 3$ T-type Ca^{2+} channels are widely distributed throughout the brain including the peripheral auditory system and ascending auditory tract. Their exact role in auditory information processing is still not fully understood. Within the LVA subgroup, $Ca_v 3.2$ T-type Ca^{2+} channels seem to be of special importance as qPCR revealed a steady increase in $Ca_v 3.2$ transcript levels over age, e.g. in the cochlea and spiral ganglion neurons (SGN). Furthermore, pharmacological studies suggested an association between $Ca_v 3.2$ expression and both age-related and noise-induced hearing loss. Given the potential functional relevance of $Ca_v 3.2$ VGGCs in sensorineural hearing loss, we recorded gender specific auditory evoked brainstem responses (ABRs) upon both click and tone burst presentation. Here we present auditory brainstem response (ABR) data from $Ca_v 3.2^{+/+}$, $Ca_v 3.2^{+/-}$ and $Ca_v 3.2^{-/-}$ mice from both genders which are of value for researchers who want to evaluate how $Ca_v 3.2$ loss affects basic auditory parameters, e.g. click and tone burst based hearing thresholds, amplitude growth function and peak latencies.

Data description: Information presented here includes ABR data from age-matched female and male $Ca_v 3.2^{+/+}$, $Ca_v 3.2^{+/-}$ and $Ca_v 3.2^{-/-}$ mice and technical aspects of the auditory recording protocol. Data were recorded using a commercially available ABR setup from Tucker Davis Technologies Inc. (TDT). Raw data files (arf.-file format) were exported as txt.-files with free access for analysis.

Keywords: Amplitude, Auditory brainstem responses, Calcium channel, Ca_v3.2, Click, Monaural, Sound pressure level, Threshold, Tone burst, Transgenic mice, T-Type

Objective

Voltage-gated Ca^{2+} channels are key players in regulating cellular Ca^{2+} homeostasis. Only few Ca^{2+} channels have been functionally related to auditory information processing including $Ca_v1.3$ L-type Ca^{2+} channels, the ablation of which results in congenital deafness. Recent animal studies suggested that $Ca_v3.2$ T-type Ca^{2+}

channels might play a role in age- and noise-induced hearing loss as ablation of the channel seemed to protect from sensorineural hearing loss. On the other hand, $Ca_v3.2\ Ca^{2+}$ channels were reported to exhibit increased expression with age in the inner ear/spiral ganglion neurons pointing to an important functional role in the auditory system. Interestingly, no auditory profiling of $Ca_v3.2^{+/-}$ and $Ca_v3.2^{-/-}$ mice had been performed to unravel the physiological involvement of $Ca_v3.2$ in the peripheral and ascending auditor tract. To do so, we carried out click and tone burst evoked auditory brainstem response (ABR) recordings from $Ca_v3.2^{+/+}$, $Ca_v3.2^{+/-}$ and $Ca_v3.2^{-/-}$ mice from both genders.

Full list of author information is available at the end of the article



^{*}Correspondence: marco.weiergraeber@bfarm.de

¹ Experimental Neuropsychopharmacology, Federal Institute for Drugs and Medical Devices (Bundesinstitut für Arzneimittel und Medizinprodukte, BfArM), Kurt-Georg-Kiesinger-Allee 3, 53175 Bonn, Germany

Lundt et al. BMC Res Notes (2019) 12:157 Page 2 of 3

Monaural recording results in all three lines were analyzed for threshold alterations and differences in peak amplitudes and peak latencies and submitted elsewhere. Raw ABR data were exported as txt.-files to provide free access and to enable researchers to carry out their own ABR data analysis, including further investigation of binaural recordings or application of additional manual and/ or automatic analytical tools (Table 1).

Data description

Experimental animals

Ca_v3.2 transgenic mice [1] from Mutant Mouse Resource and Research Centers (MMRRC: 009979-MU; strain name: B6.129-*Cacna1h*^{tm1Kcam}/Mmmh) were maintained in the C57Bl/6J background. For subsequent ABR recordings, Ca_v3.2^{+/+} controls, heterozygous Ca_v3.2^{+/-} and homozygous null mutant Ca_v3.2^{-/-} mice (55 animals in total) were used from both age-matched genders with the following characteristics: Males: Ca_v3.2^{+/+}: n=11 (♂), weight 32.82 ± 0.58 g; Ca_v3.2^{+/-}: n=7 (♂), weight 33.11 ± 0.81 g; Ca_v3.2^{-/-}: n=9 (♂), weight 29.09 ± 0.75 g. Females: Ca_v3.2^{+/+}: n=12 (♀), weight 24.09 ± 0.41 g; Ca_v3.2^{+/-}: n=8 (♀), weight 23.50 ± 0.41 g; Ca_v3.2^{-/-}: n=8 (♀), weight 22.10 ± 0.43 g.

ABR recording procedure

For recording of monaural bioelectrical auditory potentials, subdermal stainless steel electrodes were inserted at the vertex, axial the pinnae [(+) electrode] and ventrolateral of the right pinna [(-) electrode]. The ground electrode was positioned at the hip of the animal. To verify proper electrode positioning/conductivity, impedance measurements of all electrodes ($<5 \text{ k}\Omega$) were carried out

prior to each recording [Lundt A, Seidel, Robin, Soos J, Henseler C, Müller R, Bakki M, Arshaad IM, Ehninger D, Hescheler J, Sachinidis A, Broich K, Wormuth C, Papazoglou A, Weiergräber M. Ca_v3.2 T-type calcium channels are physiologically mandatory for the auditory system despite their devastating role in sensorineural hearing loss. Neuroscience, unpublished].

All ABR recordings were performed under free field conditions using a single loudspeaker (MF1 Multi-Function Speaker, TDT, USA) which was positioned 10 cm opposite to the rostrum of the animals.

The SigGenRZ software (TDT) was used to program stimulus protocols for click and tone bursts. The bio-electrical ABR signals recorded from the subdermal electrodes were transferred to a head stage (RA4LI, TDT) and forwarded to the preamplifier (RA4PA, TDT) with 20-fold amplification.

ABR data acquisition was carried out at a sampling rate of 24.4 kHz and signals were bandpass filtered (high pass 300 Hz, low pass 5 kHz) using a 6-pole Butterworth filter. The individual ABR data acquisition time was 25 ms starting with a 5 ms baseline period prior to the individual acoustic stimulus onset (pre ABR baseline) and exceeding the 10 ms ABR section by another 10 ms baseline (post ABR baseline) [Lundt et al., unpublished].

Two types of acoustic stimuli were applied for ABR recordings using the SigGenRZ software (TDT) and applied via the TDT BioSigRZ platform. The first stimulus entity was a click of $100~\mu s$ duration, with alternating polarity (switching between condensation and rarefaction).

The second stimulus entity was a 4.5 ms tone burst (transient sinusoidal plus) of alternating polarity with

Table 1 Overview of data files/data sets [2]

| Label | Name of data file/data set | File types (file extension) | Data repository and identifier (DOI or accession number) |
|---------------------|--|-----------------------------|---|
| Data description 1 | Click header description | .docx file format | Mendeley Data (http://dx.doi.org/10.17632/ dms4hsd75s.1) |
| Data description 2 | Tone burst header description | .docx file format | Mendeley Data (http://dx.doi.org/10.17632/dms4hsd75s.1) |
| Exported data set 1 | Female Ca _v 3.2 ^{+/+} Animal # 1–12 (click and tone burst evoked ABRs) | .txt file format | Mendeley Data (http://dx.doi.org/10.17632/dms4hsd75s.1) |
| Exported data set 2 | Female Ca _v 3.2 ^{+/-} Animal # 1–8 (click and tone burst evoked ABRs) | .txt file format | Mendeley Data (http://dx.doi.org/10.17632/dms4hsd75s.1) |
| Exported data set 3 | Female Ca _v 3.2 ^{-/-} Animal # 1–8 (click and tone burst evoked ABRs) | .txt file format | Mendeley Data (http://dx.doi.org/10.17632/dms4hsd75s.1) |
| Exported data set 4 | Male $Ca_v 3.2^{+/+}$ Animal # 1–11 (click and tone burst evoked ABRs) | .txt file format | Mendeley Data (http://dx.doi.org/10.17632/dms4hsd75s.1) |
| Exported data set 5 | Male $Ca_v 3.2^{+/-}$ Animal # 1–7 (click and tone burst evoked ABRs) | .txt file format | Mendeley Data (http://dx.doi.org/10.17632/dms4hsd75s.1) |
| Exported data set 6 | Male $Ca_v 3.2^{-/-}$ Animal # 1–9 (click and ton burst evoked ABRs) | .txt file format | Mendeley Data (http://dx.doi.org/10.17632/dms4hsd75s.1) |

Lundt et al. BMC Res Notes (2019) 12:157 Page 3 of 3

Hann envelope rise and fall times of 1.5 ms duration. The frequency range covers 1–42 kHz in 6 kHz steps. All acoustic stimuli were applied 300 times at a rate of 20 Hz for averaging.

Sound pressure levels (SPL) were increased in 5 dB steps for clicks and 10 dB steps for tone bursts, starting from 0 dB up to 90 dB (increasing SPL mode). Sound pressure levels for tone bursts within the range of 1–42 kHz were calibrated each day prior to recording [Lundt et al., unpublished].

Limitations

ABR data presented here were performed under standard free field conditions. Data were recorded from agematched animals of ~ 20 weeks. We did not record from animals of different age.

Abbreviations

ABR: auditory brainstem response; SPL: sound pressure level.

Authors' contributions

AL, data acquisition; CH, animal housing and genotyping; CW, data management; JS, data acquisition; RS, data management; RM, data management; IMA, data management; KB, drafting manuscript; JH, drafting manuscript; AS, drafting manuscript; DE, drafting manuscript; AP, data acquisition, drafting manuscript; MW, project management, project planning, data acquisition, writing manuscript. All authors read and approved the final manuscript.

Author details

¹ Experimental Neuropsychopharmacology, Federal Institute for Drugs and Medical Devices (Bundesinstitut für Arzneimittel und Medizinprodukte, BfArM), Kurt-Georg-Kiesinger-Allee 3, 53175 Bonn, Germany. ² Federal Institute for Drugs and Medical Devices (Bundesinstitut für Arzneimittel und Medizinprodukte, BfArM), Kurt-Georg-Kiesinger-Allee 3, 53175 Bonn, Germany. ³ Cognitive Neurophysiology, Department of Psychiatry and Psychotherapy, University Hospital Cologne, University of Cologne, Faculty of Medicine, Kerpener Str. 62, 50937 Cologne, Germany. ⁴ Institute of Neurophysiology, University of Cologne, Faculty of Medicine, Robert-Koch-Str. 39, 50931 Cologne, Germany. ⁵ Molecular and Cellular Cognition, German Center for Neurodegenerative Diseases (Deutsches Zentrum für Neurodegenerative Erkrankungen, DZNE), Sigmund-Freud-Str. 27, 53127 Bonn, Germany.

Acknowledgements

The authors are grateful to Dr. Christina Kolb (German Center for Neurodegenerative Diseases, DZNE) and Dr. Robert Stark (DZNE) for assistance in animal breeding and animal health care.

Competing interests

The authors declare that they have no competing interests.

Availability of data materials

The data described in this Data note can be freely and openly accessed on Mendeley Data (http://dx.doi.org/10.17632/dms4hsd75s.1). Please see Table 1 and reference list for details and links to the data.

Consent for publication

Not applicable.

Ethics approval and consent to participate

All animal procedures were performed according to the Guidelines of the German Council on Animal Care and all protocols were approved by the Local Institutional and National Committee on Animal Care (Landesamt für Natur, Umwelt und Verbraucherschutz, LANUV, Germany). The authors further certify that all animal experimentation was carried out in accordance with the European Communities Council Directive of November 24, 1986 (86/609/EEC). Specific effort was made to minimize the number of animals used and their suffering.

Funding

This work was financially supported by the Federal Institute for Drugs and Medical Devices (Bundesinstitut für Arzneimittel und Medizinprodukte (BfArM), Bonn, Germany). BfArM provided financial and experimental resources to carry out the study.

Publisher's Note

Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.

Received: 12 September 2018 Accepted: 11 March 2019 Published online: 20 March 2019

References

- Chen CC, Lamping KG, Nuno DW, Barresi R, Prouty SJ, Lavoie JL, et al. Abnormal coronary function in mice deficient in alpha1HT-type Ca²⁺ channels. Science. 2003;302(5649):1416–8.
- Lundt A, Seidel, Robin, Soos J, Henseler C, Müller R, Bakki M, Arshaad IM, Ehninger D, Hescheler J, Sachinidis A, Broich K, Wormuth C, Papazoglou A, Weiergräber M. Datasets from gender specific auditory brainstem responses (ABRs) from Ca_v3.2^{+/+}, Ca_v3.2^{+/-} and Ca_v3.2^{-/-} mice following click and tone burst stimulation. Mendeley Data. http://dx.doi. org/10.17632/dms4hsd75s.1.

Ready to submit your research? Choose BMC and benefit from:

- fast, convenient online submission
- thorough peer review by experienced researchers in your field
- rapid publication on acceptance
- support for research data, including large and complex data types
- gold Open Access which fosters wider collaboration and increased citations
- maximum visibility for your research: over 100M website views per year

At BMC, research is always in progress.

Learn more biomedcentral.com/submissions

