Repetitive afterglow in zirconia by pulsed near-infrared irradiation toward biological temperature sensing

Masaharu Ohashi¹, Yoshihiro Takahashi¹*, Nobuaki Terakado¹, Noriko Onoue¹, Tsuyoshi Shinozaki² & Takumi Fujiwara¹*

Photoluminescence provides information about the surrounding environment. In this study, aiming to develop a non-invasive deep body-temperature sensing method, we investigated photoluminescence properties of afterglow zirconia (ZrO₂) by pulsed near-infrared (NIR) light irradiation based on the biological temperature. Pulsed light irradiation produced optically stimulated luminescence, followed by afterglow, with the property of repeating 100 times or more. Furthermore, the basic principle of temperature measurement was demonstrated through afterglow decay curve measurements. The use of harmless ZrO₂ as a sensing probe and NIR light, which is relatively permeable to living tissues, is expected to realize temperature measurements in the brain and may also facilitate optogenetic treatment.

Afterglow (AG) is long-lasting photoluminescence (PL) that occurs after photoexcitation and is caused by recombination between holes and electrons trapped at metastable sites and thermally released. It is commonly observed on an emergency sign and watch dial. Trapped electrons can be released by near-infrared (NIR) irradiation as well, i.e., optically stimulated luminescence (OSL). AG and OSL appear to be useful for biological and medical assessments because optical/spectroscopic techniques are considered a non-invasive and contactless approach. For example, Chermont et al. demonstrated bio-imaging using an AG phosphor of Eu³⁺, Dy³⁺, Mn²⁺-tri-doped Ca₀.₂Zn₀.₉Mg₀.₉Si₂O₆¹. In addition, many researchers have focused on developing phosphors that apply to imaging/sensing²–⁸.

Our research group has proposed a new temperature sensing concept for deep parts of the human body based on AG and OSL and then demonstrated a basic principle for temperature determination by the AG decay curve using a chemically-stable and harmless oxide, namely, zirconia (ZrO₂), as a sensing probe⁹. In addition, external laser stimulation with a wavelength (~650–1000 nm) in the NIR region, which has relatively high transmittance to biological tissue (i.e., biological window), enables site-selective and arbitrary-timing measurements, and we have also been able to observe OSL by irradiating a bone sample with a continuous-wave NIR-laser⁹. Meanwhile, continuous irradiation is regarded as an external source of energy in terms of temperature rise and resulting in thermal damage in biological tissue. In this study, to demonstrate biological temperature sensing using AG phosphor, we have investigated the impact of environmental temperature on the PL properties of a ZrO₂ sample under pulsed NIR irradiation. OSL, along with AG, was observed at 480 nm by repetitive pulsed irradiation, and the observation could be continued over ~100 cycles of pulsed irradiation. Furthermore, evaluation of temperature was also demonstrated on the basis of decay curves observed in the AG process.

Results and discussion

OSL and AG by pulsed NIR irradiation in ZrO₂. Figure 1 shows thermoluminescent (TL) spectra of ZrO₂ sample pre-heated (de-trapped) at 323 K for 2.5 h, along with the result of ZrO₂ sample not pre-heated. Preparation of the ZrO₂ sample and its structural analyses are described in Method’s part. According to the TL measurement, thermally treated ZrO₂ is reported to exhibit several peaks due to several kinds of electron-trapping sites¹⁰–¹². We have also confirmed six primary TL peaks, peaks A–F, with peaks E and F appearing around/
above room temperature, attributed to AG and OSL, respectively. The ZrO$_2$ sample treated at 1400 °C for 6 h exhibits AG after ultraviolet (UV)-light exposure (254 nm), followed by OSL visually observed at the NIR-laser-irradiated spot (inset). However, the presence of multiple peaks complicates the analysis of AG decay curves. Therefore, prior to the TL measurement, we pre-heated the ZrO$_2$ sample at 323 K for 2.5 h to release electrons trapped in trapping sites, appearing below room temperature, i.e., de-trapping. According to the result, peaks A–E vanished, and only peak F at ~360 K, which is inactive at biological temperature, was observed in the de-trapped (pre-heated) sample. Hereafter, measurements were performed using the de-trapped sample.

Figure 2 shows the PL intensity of the ZrO$_2$ sample irradiated with repetitive pulsed NIR light at room temperature. The intensity rapidly increased immediately after irradiation and subsequently decayed (Fig. 2a). The rapid increase in PL intensity is attributed to OSL due to NIR irradiation. The moderate decay after pulsed irradiation is regarded as an AG process because the irradiation was stopped in this term. This leads us to consider that AG decay curves induced by OSL exhibit similar behavior to AG properties previously reported, i.e., the lifetime decreases as the environmental temperature of ZrO$_2$ increases. Furthermore, we could confirm the rapid PL increase of over 100 times caused by repetitive irradiation (Fig. 2b). On the other hand, the maximum OSL intensity decreased over time (or repetition), i.e., $I_{\text{max,10}} / I_{\text{max,100}} \approx 0.066$.

Decay curve response to pulsed NIR irradiation and its temperature dependence. Next, we examined the temperature dependence on AG lifetime by pulsed irradiation. Since the release of electrons is caused by thermal excitation, the environmental temperature determines the AG lifetime (or slope of an AG curve). Assuming that AG phosphor contains only one type of electron-trapping site, the temperature dependence on the intensity of AG PL follows a simple exponential relationship:

$$\frac{I(t)}{I(0)} \propto \exp(-at),$$  \hspace{2cm} (1)
where \( I(t) \) represents the PL intensity, \( I(0) \) represents the initial intensity, \( t \) represents time, \( a \) represents the probability of thermal activation of a trapped electron into the conduction band, \( E \) represents the thermal activation energy of a trapped electron, \( s \) represents the frequency factor, \( k_B \) represents the Boltzmann constant, and \( T \) represents temperature. We can obtain the activation energy by observing the temperature dependence on \( a \), which corresponds to an inverse of PL lifetime (\( 1/\tau \)). As a result, we can estimate the environmental temperature of phosphor by AG measurements.

Figure 3 shows AG decay curves of ZrO\(_2\) by single pulsed NIR irradiation at varying environmental temperatures. The decay curves observed at 313 K and 323 K seemed to be monotonically decreased, whereas the initial intensity (\( t < \sim 1 \)) rapidly decayed and then decreased monotonically (Fig. 3a). This suggests that the OSL intensity partially contributed to the AG intensity; thus, we re-plotted the decay curves, which are normalized by the intensity at \( t = 1 \) s to evaluate the lifetime by fitting using a single exponential function. As a result, the de-trapped sample tended to exhibit a linear decay response with slopes that increased with increasing temperature. On the other hand, the response at 323 K deviated from the theoretical line (inset), implying that electron release from a deeper trapping site, i.e., peak F, partially contributed to AG decay curves because a trail of peak F is extended around 323 K. The evaluated lifetime and temperature could be plotted according to the Arrhenius approach, resulting in a linear relation (Fig. 3b), yielding activation energy of approximately 0.48 eV. Thus, we demonstrated that temperature can be estimated on the basis of the AG process lifetime by pulsed NIR irradiation simulations.

The activation energy in this study is comparable to that of AG in ZrO\(_2\) previously reported\(^9\). To clarify the AG process in detail, we measured the PL intensity of the de-trapped ZrO\(_2\) sample through NIR irradiation stimulation at 77 K and 323 K and compared its decay behaviors. Figure 4 depicts PL intensities detected by

\[
a = s \cdot \exp \left( -\frac{E}{k_B T} \right),
\]

Figure 4. (a) PL intensity as a function of time in ZrO\(_2\) sample irradiated with a single-pulse NIR light at 77 K and 323 K. Blue and red regions indicate OSL and AG regimes, respectively. (b) Schematics of the PL process (Valence band is omitted).
NIR irradiation with a rectangular waveform (width: 10 s). Immediately after irradiation, obvious PL could be observed at both temperatures (Fig. 4a). At 77 K, the PL is attributed to OSL, which is occurred by the combination between holes and electrons released by NIR irradiation. In addition, at 323 K, the sample exhibited moderate AG decay when the irradiation was stopped, whereas a rapid decrease in the intensity was observed at 77 K, indicating that AG is barely observed at 77 K. These measurements lead to the conclusion that, at 323 K, electrons trapped in trapping sites, which are inactive at biological temperature (peak F), are released by NIR irradiation, and subsequently, these electrons are partially re-trapped at the thermally active site (peak E) via transition to the conduction band. Eventually, the AG process is observed (route A; Fig. 4b). At 77 K, although electrons released by NIR irradiation are trapped by active sites as well, the thermal release is suppressed because this temperature is significantly lower than peak E and others (route B). According to the result, only OSL can be observed. Therefore, it is reasonable to assume that AG observed immediately after NIR irradiation is regarded as typical AG, and consequently, they are identical. Although the origin of AG in ZrO2 is still controversial, Iwasaki et al. suggest that the trapping sites in AG was closely related to the presence of F+-center (or V_o in Kroger-Vink notation) on the basis of measurement of electron spin resonance13.

This study can demonstrate the availability of pulsed NIR irradiation, but we should also consider converting the PL wavelength to be compatible with the biological window range. The conversion of PL to a longer wavelength in ZrO2 using an appropriate dopant has been reported14,15, and a similar study is underway.

Summary
The temperature dependence of PL properties of ZrO2 under pulsed NIR-laser irradiation was investigated for biological temperature sensing applications by decay curve measurements. We demonstrated that environmental temperatures can be estimated from the AG lifetime through the Arrhenius approach. The use of low-toxicity ZrO2 as a sensing probe is expected to reduce stress in living organisms. Repetitive pulsed NIR-laser irradiation is expected to not only reduce damage to biological tissue but also shorten the time and improve the accuracy of repetitive temperature measurements.

Non-invasive therapies based on optogenetics have been reported to control memory and sociality via light irradiation of local areas of the brain and neurons, but in these experiments, optical fibers/LEDs are inserted directly into the irradiated areas of mice16–19. Therefore, AG and OSL caused by external pulsed NIR irradiation using AG probes may contribute to optogenetic therapy by enabling non-invasive and site-selective optical simulations of local areas.

Methods
We prepared the ZrO2 sample by annealing oxygen-defect activation, and the sample exhibited a broad PL peak at ~ 480 nm due to AG and OSL5,13. Therefore, in this study, we also prepared another ZrO2 sample by referring to the previous study: Commercial reagent-grade ZrO2 powder (purity: 99.9%, Soekawa Chem. Co., Ltd.) was thermally treated at 1400 °C for 6 h to introduce oxygen defects. The heating rate was 10 K/min, and the target temperature was maintained for 6 h in an electric furnace under atmospheric conditions. After the treatment, the ZrO2 powder was cooled in the furnace, and finally, the ZrO2 sample was obtained.

At room temperature, ZrO2 crystallizes in monoclinic system, and undergoes a phase-transition to tetragonal system at ~ 1170 °C. Thermal treatment above the phase-transition temperature introduces oxygen defects, which are associated with the phase transition13. Since tetragonal ZrO2 has been found to show no PL20, structural analysis of the obtained sample was performed by means of an X-ray diffraction (XRD; Cu-Kα). As the result, the phase of treated ZrO2 was identified to be monoclinic (Fig. 5a). In addition, a transmission electron microscopy (TEM) revealed clear lattice pattern in the ZrO2 grain, suggesting the preservation of crystallinity (Fig. 5b).
The PL intensity as a function of time (decay curve) was measured using a spectrofluorometer with a xenon lamp as the excitation source. The environmental temperature of the ZrO$_2$ sample during decay curve measurement was controlled using a cryostat. The PL intensity as a function of temperature (TL spectrum) was also measured using the spectrofluorometer conjugated with the cryostat. The intensity was detected at 480 nm in the decay curve during TL measurements with a heating rate of 1 K/min. In addition, prior to these measurements, the ZrO$_2$ sample was excited at 280 nm, corresponding to the PL excitation peak, for 1 h. The wavelengths for PL and PL excitation were selected according to a previous study. An NIR-laser diode (808 nm, 90 mW) was installed on the cryostat, and pulsed light was generated by combining a function generator with the power supply unit. The waveform was a square wave, with a pulse width of 0.5 s.

Data availability
The data and material used and/or analysed during this study are available from the corresponding author on reasonable request.

Received: 4 April 2022; Accepted: 13 May 2022
Published online: 21 May 2022

References
1. Le Masne de Chermont, Q. et al. Nanoprobes with near-infrared persistent luminescence for in vivo imaging. *Proc. Natl. Acad. Sci.* **104**, 9266–9271 (2007).
2. Yang, J.-M., Yang, H. & Lin, L. Quantum dot nano thermometers reveal heterogeneous local thermogenesis in living cells. *ACS Nano* **5**, 5067–5071 (2011).
3. Okabe, K. et al. Intracellular temperature mapping with a fluorescent polymeric thermometer and fluorescence lifetime imaging microscopy. *Nat. Commun.* **3**, 705 (2012).
4. Li, Y. et al. Long persistent and photo-stimulated luminescence in Cr$^{3+}$-doped Zn–Ga–Sn–O phosphors for deep and reproducible tissue imaging. *J. Mater. Chem. C* **2**, 2657 (2014).
5. Li, Y., Goecevicius, M. & Qiu, Y. Long persistent phosphors–from fundamentals to applications. *Chem. Soc. Rev.* **45**, 2090–2136 (2016).
6. Lv, S. et al. Transition metal doped smart glass with pressure and temperature sensitive luminescence. *Adv. Optical Mater.* **6**, 1800881 (2018).
7. Okada, G. Novel radio-photoluminescence materials and applications. *J. Ceram. Soc. Jpn.* **129**, 419–424 (2021).
8. Gao, Y., Wang, B., Liu, L. & Shinozaki, K. Near-infrared engineering for broad-band wavelength-tunable in biological window of NIR-II and -III: A solid solution phosphor for Sr$_{1-x}$Ca$_x$TiO$_3$:Ni$^{2+}$. *J. Lumin.* **238**, 118235 (2021).
9. Ohashi, M. et al. Temperature dependence of afterglow in zirconia and its optically stimulated luminescence by bone-through irradiation for biological temperature probe. *Sci. Rep.* **10**, 2242 (2020).
10. Kisk, V. et al. Photo-, thermos- and optically stimulated luminescence of monoclinic zirconia. *J. Lumin.* **174**, 49–55 (2016).
11. Lokesh, H. S., Chauhan, N., Nagabhushana, K. R. & Singh, F. Dosimetric properties of ZrO$_2$ and ZrO$_2$:Sm$^{3+}$ exposed to beta rays. *Ceram. Int.* **44**, 18871–18877 (2018).
12. Sawamura, K. et al. Persistent luminescence properties of monoclinic luminescent zirconium oxide annealed under different oxygen partial pressure. *J. Ceram. Soc. Jpn.* **128**, 175–180 (2020).
13. Iwasaki, K. et al. Prominent long-lasting photoluminescence in defect-activated zirconia: A rare-earth free and accessible phosphor. *Ceram. Int.* **44**, 12929–12933 (2018).
14. Zhao, Z. & Wang, Y. The synthesis and afterglow luminescence properties of a novel red afterglow phosphor: ZrO$_2$:Sm$^{3+}$, $S$.$S$.$S$. *J. Lumin.* **132**, 2842–2846 (2012).
15. Goto, S. et al. Photoluminescence and afterglow in samarium-doped zirconia toward biological temperature sensor. *J. Ceram. Soc. Jpn.* **127**, 696–699 (2019).
16. Ramires, S. et al. Creating a false memory in the hippocampus. *Science* **341**, 387–391 (2013).
17. Yang, Y. et al. Wireless multilateral devices for optogenetic studies of individual and social behaviors. *Nat. Neurosci.* **24**, 1035–1045 (2021).
18. Goto, A. et al. Stepwise synaptic plasticity events drive the early phase of memory consideration. *Science* **374**, 857–863 (2021).
19. Mage, S. D. et al. Brain-wide electrical dynamics encode individual appetitive social behavior. *Neuron* **110**, 1–14 (2022).
20. Nobuta, Y. et al. Formation and photoluminescence of zirconia dendrites in borosilicate glass-ceramics. *J. Ceram. Soc. Jpn.* **125**, 343–347 (2017).

Acknowledgements
The authors would like to thank Dr. Takamichi Miyazaki, Dr. Kenichiro Iwasaki, Mr. Yasuhiro Nobuta, Mr. Satoshi Goto, and Ms. Aoni Sato for their significant contributions to this study.

Author contributions
M.O. fabricated and characterized the materials. Y.T., N.T., and T.F. designed the study; N.O. and T.S contributed to the conception of this study, and Y.T. and T.F. wrote the manuscript. All authors discussed the results and commented on the manuscript.

Competing interests
The authors declare no competing interests.

Additional information
Correspondence and requests for materials should be addressed to Y.T. or T.F.

Reprints and permissions information is available at www.nature.com/reprints.

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