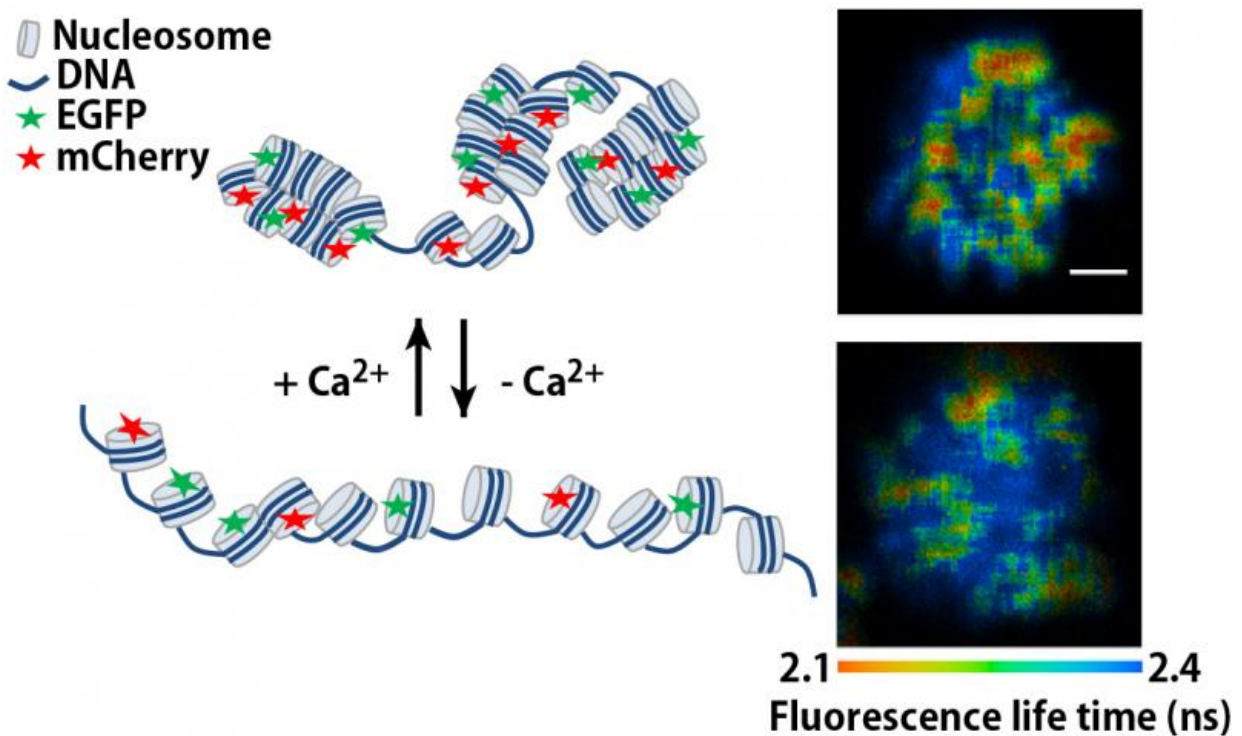


# Calcium aids chromosome condensation prior to cell division

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The effects of Ca<sup>2+</sup> on chromatin condensation revealed by the measurement of the change in fluorescence life time. Nucleosome was labelled by EGFP and mCherry and it enables us to monitor chromatin condensation status by measuring the change in the fluorescence life time. Higher Ca<sup>2+</sup> concentration induces shorter fluorescence life time indicating chromatin condensation (upper panel). In contrast, lower Ca<sup>2+</sup> induces longer life time indicating chromatin decondensation (lower panel). Bar, 100 nm. Credit: Osaka University

Osaka University-led Japanese research reveals role for calcium ions in chromosome condensation during mitosis; high-resolution imaging of living cells reveals compact, globular chromosomes in the presence of calcium that became fibrous and expand in its absence.

Before cell division when a parent cell divides into two daughter cells, each with an identical set of genetic information, replicated [chromosomes](#) become condensed and align along the spindle in the middle of the cell. This is the metaphase stage of mitosis. Chromosomes are made up of strands of DNA wrapped around histones, forming nucleosomes that resemble beads on a string. The [chromosome structure](#) is maintained by histone changes, chromosome scaffold proteins, and positively charged ions (cations). However, removal of scaffold proteins has no effect on chromosome condensation, suggesting that another factor causes the organization of metaphase chromosomes. Now, work led by Osaka University has revealed that [calcium ions](#) ( $\text{Ca}^{2+}$ ) play a crucial role in chromatin fiber compaction. The study was reported in *Scientific Reports*.

Calcium is a universal second messenger with many functions, including progression of the cell cycle. During mitosis, both  $\text{Ca}^{2+}$  and magnesium ions ( $\text{Mg}^{2+}$ ) are released from storage organelles to bind chromatin, though six-to-eight-fold more  $\text{Ca}^{2+}$  is bound than  $\text{Mg}^{2+}$  suggesting that it has a more important role. A research team led by Hideaki Takata, Kiichi Fukui and Rinyaporn Phengchat at Osaka University first confirmed that  $\text{Ca}^{2+}$  is necessary to prevent chromosome misalignment during mitosis, then used an imaging assessment of molecular interactions based on fluorescence decay (FLIM-FRET) to show that chromosomes were less compact in the absence of intracellular  $\text{Ca}^{2+}$ .

This same imaging process also revealed that changes in  $\text{Ca}^{2+}$  levels of living HeLa cells altered the extent of chromosome compaction, with the re-addition of  $\text{Ca}^{2+}$  causing chromosome compaction.  $\text{Ca}^{2+}$  therefore

appears to control the transition between condensation and decondensation. "Although previous studies have demonstrated nucleosome packaging using isolated chromatin" study first author Rinyaporn Phengchat says, "such findings should be confirmed in vivo as we have done using the sensitive FLIM–FRET technique which provides a high level of spatial resolution."

The team then used scanning electron microscopy to visualize chromosome structures at very high resolution. "Ca<sup>2+</sup> has a concentration-dependent effect on chromosome structure," corresponding author Hideaki Takata explains, "causing chromosomes to change from expanded fibrous structures to compact globular structures with increasing levels of Ca<sup>2+</sup>."

The researchers propose that, in the absence of Ca<sup>2+</sup>, the negatively charged DNA is less well neutralized, preventing it from condensing and delaying entry into prometaphase of the cell cycle. "We think that a lack of Ca<sup>2+</sup> disturbs the organization of chromosomes at the spindle, causing them to misalign," corresponding author Kiichi Fukui says. "Conversely, the normal rising levels of Ca<sup>2+</sup> during mitosis promote chromosome condensation after breakdown of the nuclear envelope."

**More information:** Rinyaporn Phengchat et al. Calcium ions function as a booster of chromosome condensation, *Scientific Reports* (2016).

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