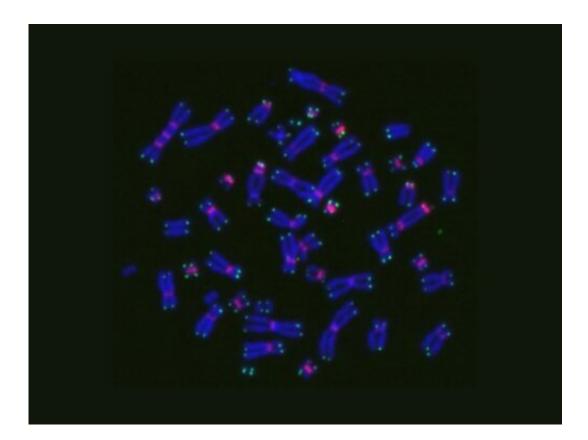


Chromosome aberrations may predict risk of severe chemoradiotherapy side effects

October 29 2021



Normal chromosomes have a single centromere, the constricted central region holding the two chromatids together to form an X shape. This photo shows some aberrant dicentric chromosomes that formed two centromeres from an esophageal cancer patient undergoing chemoradiotherapy treatment. A new study by Japanese researchers showed the relationship between acute toxicity and chromosomal aberrations in radiotherapy using in vivo data. Credit: Nobuki Imano



Japanese researchers identified chromosome aberrations as a new biomarker in predicting an esophageal cancer patient's risk of experiencing severe chemoradiotherapy side effects.

"Chemoradiotherapy is the <u>standard treatment</u> for various types of cancer. However, while its usefulness is recognized, adverse events due to radiotherapy are of great concern," the study's lead author Nobuki Imano, assistant professor at Hiroshima University's Graduate School of Biomedical and Health Sciences, explained.

"These adverse events of chemoradiotherapy vary among patients and are thought to be due to differences in individual radiosensitivity. Therefore, it has been an important issue to establish an index for predicting individual radiosensitivity to predict the adverse events of radiotherapy."

Imano and his colleagues enrolled 18 esophageal cancer patients in a prospective study to evaluate individual radiosensitivity using DNA damage response and <u>chromosome aberrations</u>.

Peripheral blood lymphocytes (PBL), a type of white blood cell used to assess immune function, were taken from patients just before and 15 minutes after each <u>treatment</u>. PBL samples were also taken at four weeks and six months after patients completed their prescribed chemoradiotherapy treatment.

The researchers counted the number of misshapen dicentric chromosomes, which formed two centromeres, and O-shaped ring chromosomes to evaluate the number of chromosomal aberrations. Meanwhile, they tallied the γ -H2AX foci—a marker for DNA damage—per cell to assess DNA double-strand breaks.

They found no significant increase in the number of γ -H2AX foci during



chemoradiotherapy in the patients. However, they discovered that chromosome aberrations were higher in patients who showed severe side effects (overreactor group) than in those who experienced lower grade toxicities (non-overreactor group).

"We found that the number of chromosomal aberrations increased during chemoradiotherapy and reached 1.04 per metaphase by the commonly used chemoradiotherapy dose of 50–60 Gy for cancer treatment, and gradually decreased after chemoradiotherapy to approximately 60% in six months," the researchers said in their findings published in the March 2021 issue of the journal *Radiation Research*.

This study is the first they know of so far to report the relationship between acute toxicity and chromosomal aberrations in radiotherapy using in vivo data.

Imano said their study could lead to personalized radiotherapy treatments in the future.

"In this study, we showed that acute toxicities in esophageal cancer patients are associated with chromosomal aberrations in PBLs. We would like to continue to examine chromosomal aberrations in other cancers," he said.

"We would like to predict not only acute toxicities but also late toxicities and treatment outcomes to establish a personalized radiotherapy treatment for all <u>cancer</u> patients."

More information: Nobuki Imano et al, Evaluating Individual Radiosensitivity for the Prediction of Acute Toxicities of Chemoradiotherapy in Esophageal Cancer Patients, *Radiation Research* (2021). DOI: 10.1667/RADE-20-00234.1



Provided by Hiroshima University

Citation: Chromosome aberrations may predict risk of severe chemoradiotherapy side effects (2021, October 29) retrieved 5 May 2024 from https://medicalxpress.com/news/2021-10-chromosome-aberrations-severe-chemoradiotherapy-side.html

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