## Realization of a society where child abuse and suicide are zero

#### Project manager

(selected in 2023)

# HASHIMOTO Akitoyo

#### Kobe University Graduate School of Medicine, Professor



### Leader's institution

Kobe University

#### R&D institutions

Kobe University

Yokohama City University

### Summary of the project

By using our proprietary epigenome data resource for juvenile suicidal individuals and novel AMPA receptor recognition technology for human brain, we aim to 1) develop of biomarkers that accurately predict the presence or absence of child abuse and suicide risk, and 2) clarify abnormalities in epigenome status, gene expression, and AMPA receptors related to child abuse and suicidal tendency. Through these efforts, we aim to visualize child abuse and suicide risks, which are difficult for children to express themselves, elucidate the biological basis relevant to child abuse and related emotional instability even leading to suicide risk, and identify novel therapeutic targets, and finally realize a society where child abuse and suicide are zero.

### Milestone by the end of project (year 2024)

<u>R&D item 1:</u> We aim to identify abnormalities in epigenomic aging, telomeres, immune cells, and DNA methylation of specific gene regions in children with a history of abuse (particularly those with suicidality among them), and to develop algorithm to detect abuse and suicide risk in children with high accuracy by using machine learning method. Furthermore, we will perform single-cell RNA sequencing analysis of blood of children facing severe suicide risk and identify abnormalities in specific cell type gene expression and high-resolution cell proportion counts.

<u>R&D item 2:</u> By using the world's first AMPA receptor recognition technology for human brain, we aim to identify the biological basis relevant to child abuse and suicide risk. Furthermore, we aim to clarify whether "we can accurately predict the amount of region-specific AMPA receptors based on comprehensive DNA methylation data on peripheral blood" and to gain new insights for novel non-invasive biomarkers that can evaluate the dynamics of AMPA receptors among youth.

## **Project structure**

R&D Item 2: Brain AMPA receptor data R&D Item 1: Comprehensive epigenome data and epigenome data for abuse and suicidality and single-cell gene expression data for child abuse and suicidality **Twenties cohort** Teenage cohort Abuse Abuse + + + history history + Suicidality \_ \_ + Suicidality \_ The world's first AMPA receptor Comprehensive epigenome data recognition for human brain Aberrant epigenome profile of CpGs related to abuse and suicidality Abnormalities in epigenomic aging, telomeres and immune cell proportion Amount of region-specific AMPA receptors Single-cell gene expression data Association with DNA methylation Cell type-specific gene expression data on peripheral blood from · High-resolution cell proportion counts same individuals Visualization and detection of abuse/suicide risks Abnormalities in epigenome, gene expression, (difficult for children to express themselves) AMPA receptors for child abuse/suicidality · Abnormalities in epigenomic aging, Early intervention If suspect child abuse telomeres, specific cell proportions Novel care/therapeutics and suicide risk ... · Aberrant epigenome status and single-cell gene expression levels in specific genes Can detect it with non-invasive biomarkers Abnormalities in region-specific AMPA receptor amounts

### Principal investigators (PIs)

R&D item 1: OTUKA Ikuo (Kobe University) FURUYASHIKI Tomoyuki (Kobe University)

### R&D item 2:

MIYAZAKI Tomoyuki (Yokohama City University) OTSUKA Ikuo (Kobe University)

