VANDERBILT

(3) RNA interference

Concept: Patterns of Epigenetic Regulation associated with Exposure in Global Zones of Conflict

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Histone tails -

Phosphorylation

Ubiquitination

Acetylation

DNA methylation

Infectious diseases, cancer, autoimmune

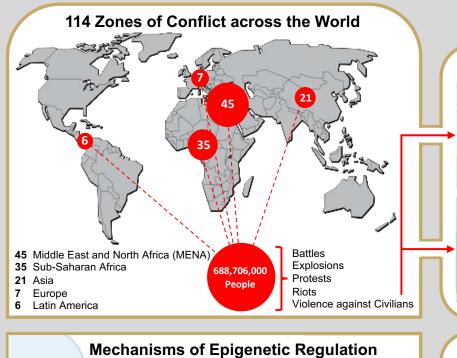
gastrointestinal distress, type 2 diabetes,

diseases, cardiovascular disease, obesity.

Deleterious regulatory effects on:

poor dental health, and a cascade of

psychological conditions



Affected Populations of Interest



Local civilians

Refugee family in Jordan, 2018

Physiological and psychological stress

Safety, fear, constant movement, malnutrition, mourning, injury...

Healthcare professionals

MSF Medical Staff in Gaza, 2023

Physiological and psychological stress

Safety, fear, constant movement, injury, lack of supplies, exhaustion...

Traumatic External Stimuli can induce **Deleterious Epigenetic Modifications**

Maternal genome Direct effect

Prenatal development

Direct and Inherited effect Vulnerable to maternal health

Inheritance

Paternal genome

Direct effect

Early childhood development



Direct and inherited effect Adverse childhood experiences (ACEs)

#ceasefirenow

Proof of Concept – DNA Methylation(2)

Physiological Dysregulation

Hypothalamic pituitary adrenal (HPA) axis which regulates heart rate, blood pressure, metabolism, and immune function.

- FKBP5

 Restrains overactivation of HPA axis
- NR3C1– Regulates glucose homeostasis

Cancer evasion, progression, angiogenesis, and metastasis

- CDH1- Invasion of tissue and tumor metastasis
- P16– Limitless replicative potential and angiogenesis
- DAPK, ASC/TMS1, HIC1- Evasion of apoptotic signals
- MLH1- DNA damage repair capacity
- CHFR- Cell cycle progression and tumorigenesis

Psychological Dysregulation

ACEs are associated with post-traumatic stress disorder (PTSD), anxiety, depression, and insomnia.

- MOAO- Enzymatically breaks down neurotransmitters
- SLC6A4— Serotonin transporter
- BNDF- Promotes differentiation and survival of neurons

Potential Study Questions

What exactly do exposures and exposure windows look like? Can exposure windows be quantified? Short versus long exposure windows?

How difficult would it be to sample or observe in these settings?

References

- Conflict Exposure ACLED ACLED Published March 4 2024 Accessed March 1 2024 https://acleddata.co