# AWARENESS ABOUT GENOMIC INSTABILITY VIA HYPOMETHYLATION OF LINE 1 AMONG ALLIED HEALTH SCIENCES STUDENTS

### Kethan Umakanth <sup>1</sup>, Prasanna Devi <sup>2</sup>, Sivakumar Ekambaram <sup>3</sup>, Jagadeswaran <sup>4</sup> and Dhanraj Ganapathy <sup>5\*</sup>

 <sup>1</sup> Intern, Department of Prosthodontics, Saveetha Dental College and Hospitals, Saveetha Institute of Medical and Technical Sciences, Chennai, India.
<sup>2</sup> B.Sc Hons AHS 1st Year, Saveetha College of Allied Health Sciences, Chennai, India.
<sup>3</sup> Assistant Professor and RSO, Department of Radiotherapy, Saveetha College of Allied Health Sciences, Saveetha Institute of Medical and Technical Sciences, Chennai, India.
<sup>4</sup> Principal and Professor, Saveetha College of Allied Health Sciences, Saveetha Institute of Medical and Technical Sciences, Chennai, India.
<sup>5</sup> Professor and Head of Department, Department of Prosthodontics, Saveetha Dental College and Hospitals, Saveetha Institute of Medical and Technical Sciences, Chennai, India.
\*Corresponding Author Email: dhanraj@saveetha.com

#### DOI: 10.5281/zenodo.12074549

#### Abstract

Introduction: Intestinal and colorectal cancers are multifactorial diseases with various inputs including diet, environment, genetic mutations, and epigenetic abnormalities. The disease first manifests as an over-proliferation defect in the form of polyps that, if not removed, can progress to precancerous adenomas. Aim: This survey was conducted for assessing the awareness about genomic instability via hypomethylation of LINE 1 among allied health Sciences students. Materials and Methods: This crosssectional research was conducted with a self-administered questionnaire containing ten questions distributed amongst 100 Allied Health science students. The students were randomly selected across various disciplines of Allied Health Sciences. The study setting was designated in the university campus. The survey instrument was a questionnaire pre tested and evaluated for validity and reliability concerns. Results: 84.5% of the respondents were aware about genomic instability and hypomethylation of line 1. 69.8% of the respondents were aware about DNA hypomethylation. 70.7% of the respondents were aware about hypomethylation increase gene expression. 72.4% of the respondents were aware about genomic stability is important. 69% of the respondents were aware about DNA include RNA. 69% of the respondents were aware about radiation cause damage to DNA. Conclusion: There is moderate awareness amongst allied health Sciences student about genomic instability via hypomethylation of LINE 1. Enhanced aware initiatives and educational programs together with iincreased importance for curriculum improvements that future promote knowledge and awareness of genomic instability via hypomethylation of LINE

Keywords: Awareness, Health Science, DNA, Hypomethylation.

#### INTRODUCTION

lonizing radiation is a well-known carcinogen resulting in a large spectrum of DNA lesions, including double-strand breakages (DSB). Radiation-induced genomic instability (RIGI) is a driving force underlying radiation carcinogenesis. Several experiments support a new paradigm of radiation biology in which RIGI can be transmitted from the surviving cell through multiple cell generations to be expressed in the progeny as a delayed and persistent effect[1].

The delayed aneuploidy--numerical chromosome aberrations are induced by low doses of x-radiation in human fibroblasts. Fifty and 100 mGy of x-rays induce delayed aneuploidy of chromosomes 1 and 4 in the progeny of irradiated fibroblast cells. Therefore, unexpected health effects likely result from delayed genomic instability after

radiation exposure, raising concerns regarding the most appropriate dose limit for radiation exposure in both the public and industrial workers[2].

Furthermore, delayed genomic instability can partially explain the mechanisms underlying radiation-induced carcinogenesis, which is poorly understood. However, despite the important implications associated with the biological effects of low-dose ionizing radiation and the impact on radiation safety and protection protocols, there are very few studies regarding delayed genomic instability after low-level radiation exposure among the general population or radiographers[3].

Genome-wide epigenetic alterations are known to hold substantial potential as biomarkers for environmental exposures; this, in turn, may provide insight into mechanisms of environmentally related diseases and allow for a better understanding of disease etiology. DNA methylation, a well-defined epigenetic mechanism, plays an important role in cancer development by adding or removing methyl (-CH3) groups at CpG dinucleotides, which influences gene regulation[4].

Genomic DNA hypomethylation resulting from demethylation in repeats or transposable elements or across the genome is associated with genomic instability, an increased number of mutational events, and subsequent development of cancer. In particular, some studies report that changes in genomic DNA methylation are associated with a striking form of genomic instability known as chromosome instability, including centromeric aberrations, chromosome aberrations, and aneuploidy[5]

## MATERIALS AND METHODS

This cross-sectional research was conducted with a self-administered questionnaire containing ten questions distributed amongst 100 Allied Health science students. The students were randomly selected across various disciplines of Allied Health Sciences. The study setting was designated in the university campus. The survey instrument was a questionnaire pre tested and evaluated for validity and reliability concerns. The questionnaire included ten questions eliciting the demographic data through open ended responses and multiple choice questions for the other responses. The study was approved by the Institutional Ethical Committee and informed consent was obtained from the Participants. The questionnaire was posted in an online platform and the identity of the respondents were kept confidential.

The questionnaire assessed the awareness about Genomic Instability and hypomethyliation of LINE 1. The responses were recorded and analysed. There were no incomplete responses and no dropouts from the study. The final data obtained was organized, tabulated and subjected to statistical analysis. The salient questions in the study are:

- 1. Are you aware of genomic instability and hypomethylation of line 1?
- 2. Are you aware of DNA hypomethylation?
- 3. Does hypomethylation increase gene expression?
- 4. Do you think genomic stability is important?
- 5. Does DNA include RNA?
- 6. Does radiation cause damage to DNA?

## RESULTS

84.5% of the respondents were aware about genomic instability and hypomethylation of line 1. 69.8% of the respondents were aware about DNA hypomethylation. 70.7% of the respondents were aware about hypomethylation increase gene expression. 72.4% of the respondents were aware about genomic stability is important. 69% of the respondents were aware about DNA include RNA. 69% of the respondents were aware about aware about DNA include RNA.

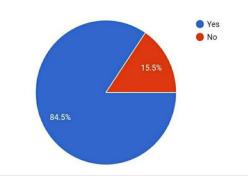
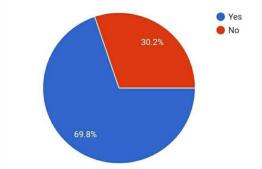
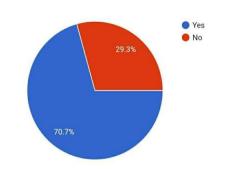


Figure 1: Awareness about genomic instability and hypomethylation of line 1









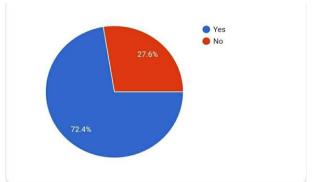


Figure 4: Awareness about genomic stability is important

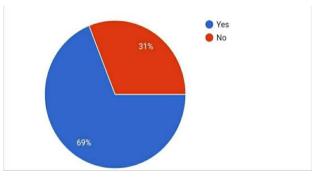


Figure 5: Awareness about DNA include RNA

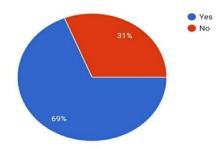


Figure 6: Awareness about radiation cause damage to DNA

## DISCUSSION

Long interspersed element-1 (LINE-1) is a repetitive DNA retrotransposon that duplicates via a copy-and-paste genetic mechanism. As LINE-1 constitutes approximately 17% of the human genome, the extent of LINE-1 methylation is regarded as a surrogate marker of global DNA methylation[6]. 84.5% of the respondents were aware about genomic instability and hypomethylation of line 1.

DNA hypomethylation refers to the loss of the methyl group in the 5-methylcytosine nucleotide. Methylation is a natural modification of DNA, and mainly affects the cytosine base (C) when it is followed by a guanosine (G) in mammals (Methylation)[7]. 69.8% of the respondents were aware about DNA hypomethylation.

Although downregulation of gene expression was found to be the most pronounced effect of hypermethylation in the present study, we also show that hypermethylation in the promoter region can be associated with upregulation of gene expression[8]. 70.7% of the respondents were aware about hypomethylation increase gene expression.

When DNA is methylated, nearby histones are deacetylated, resulting in compounded inhibitory effects on transcription. Likewise, demethylated DNA does not attract deacetylating enzymes to the histones, allowing them to remain acetylated and more mobile, thus promoting transcription[9]. 72.4% of the respondents were aware about genomic stability is important.

DNA methylation is essential for silencing retroviral elements, regulating tissuespecific gene expression, genomic imprinting, and X chromosome inactivation. Importantly, DNA methylation in different genomic regions may exert different influences on gene activities based on the underlying genetic sequence[10].69% of the respondents were aware about radiation cause damage to DNA.

### CONCLUSION

There is moderate awareness amongst allied health Sciences student about genomic instability via hypomethylation of LINE 1. Enhanced aware initiatives and educational programs together with increased importance for curriculum improvements that future promote knowledge and awareness of genomic instability via hypomethylation of LINE

#### References

- Daskalos A, Nikolaidis G, Xinarianos G, Savvari P, Cassidy A, Zakopoulou R, Kotsinas A, Gorgoulis V, Field JK, Liloglou T. Hypomethylation of retrotransposable elements correlates with genomic instability in non-small cell lung cancer. International Journal of Cancer. 2009 Jan 1;124(1):81-7.
- 2) Richards KL, Zhang B, Baggerly KA, Colella S, Lang JC, Schuller DE, Krahe R. Genome-wide hypomethylation in head and neck cancer is more pronounced in HPV-negative tumors and is associated with genomic instability. PloS one. 2009 Mar 18;4(3):e4941.
- 3) Sheaffer KL, Elliott EN, Kaestner KH. DNA hypomethylation contributes to genomic instability and intestinal cancer initiation. Cancer Prevention Research. 2016 Jul 1;9(7):534-46.
- 4) Cho YH, Jang Y, Woo HD, Kim YJ, Kim SY, Christensen S, Cole E, Choi SY, Chung HW. LINE-1 hypomethylation is associated with radiation-induced genomic instability in industrial radiographers. Environmental and molecular mutagenesis. 2019 Mar;60(2):174-84.
- 5) Kaup S, Grandjean V, Mukherjee R, Kapoor A, Keyes E, Seymour CB, Mothersill CE, Schofield PN. Radiation-induced genomic instability is associated with DNA methylation changes in cultured human keratinocytes. Mutation Research/Fundamental and Molecular Mechanisms of Mutagenesis. 2006 May 11;597(1-2):87-97.
- 6) Yao Y, Dai W. Genomic instability and cancer. Journal of carcinogenesis & mutagenesis. 2014;5.
- Ponomaryova AA, Rykova EY, Gervas PA, Cherdyntseva NV, Mamedov IZ, Azhikina TL. Aberrant methylation of LINE-1 transposable elements: a search for cancer biomarkers. Cells. 2020 Sep;9(9):2017.
- Ogino S, Nosho K, Kirkner GJ, Kawasaki T, Chan AT, Schernhammer ES, Giovannucci EL, Fuchs CS. A cohort study of tumoral LINE-1 hypomethylation and prognosis in colon cancer. JNCI: Journal of the National Cancer Institute. 2008 Dec 3;100(23):1734-8.
- 9) Kemp JR, Longworth MS. Crossing the LINE toward genomic instability: LINE-1 retrotransposition in cancer. Frontiers in chemistry. 2015 Dec 16;3:68.
- 10) Kerachian MA, Kerachian M. Long interspersed nucleotide element-1 (LINE-1) methylation in colorectal cancer. Clinica Chimica Acta. 2019 Jan 1;488:209-14.