



Source details

Indian Journal of Forensic Medicine and Toxicology

Scopus coverage years: from 2008 to 2021

(coverage discontinued in Scopus)

Publisher: Medico Legal Society

ISSN: 0973-9122 E-ISSN: 0973-9130

Subject area:

Social Sciences: Law

Medicine: Pathology and Forensic Medicine

Pharmacology, Toxicology and Pharmaceutics: Toxicology

Environmental Science: Health, Toxicology and Mutagenesis

Source type: Journal

[View all documents >](#)

[Set document alert](#)

[Save to source list](#) [Source Homepage](#)

CiteScore 2020

0.1



SJR 2020

0.115



SNIP 2022

0.163



[CiteScore](#)

[CiteScore rank & trend](#)

[Scopus content coverage](#)

i Improved CiteScore methodology

CiteScore 2020 counts the citations received in 2017-2020 to articles, reviews, conference papers, book chapters and data papers published in 2017-2020, and divides this by the number of publications published in 2017-2020. [Learn more >](#)



CiteScore 2020

$$0.1 = \frac{387 \text{ Citations 2017 - 2020}}{3,509 \text{ Documents 2017 - 2020}}$$

Calculated on 05 May, 2021

CiteScore rank 2020

Category	Rank	Percentile
Social Sciences		
Law	#639/722	11th
Medicine		
Pathology and Forensic Medicine	#183/191	4th
Pharmacology, Toxicology and	#118/122	3rd

[View CiteScore methodology >](#) [CiteScore FAQ >](#) [Add CiteScore to your site](#)

Vol. 14 No. 2 (2020): Indian Journal of Forensic Medicine & Toxicology



DOI: <https://doi.org/10.37506/ijfmt.v14i2>

Published: 2020-07-16

Articles

Morphology of Palatal Rugae in Various Sagittal Skeletal Malocclusions in Kerala Population- A Retrospective Study

Crystal Runa Soans¹, Azhar Mohammed², Murali PS¹, Mcqueen Mendonca³, Prajwal Shetty³, VartikaKumari⁴
1-7

 Pdf

Analysis of Hospital Deaths at Tertiary Care Teaching Hospital

Jeeveswararao Bagadi¹, Srinivasulu Pothireddy² Sujan Kumar Mohanthy³
8-12

 Pdf

Study of Fingerprints in Relation to Dental Caries

Maitrayee Dutta Swargiary¹, Bhanukul Barman²
13-18

 Pdf

Correlation of CD4 and Distal Sensory Polyneuropathy in HivAids Patients

Desti Ariani Kistanti¹, Paulus Sugianto¹, Erwin Astha Triyono²

1485-1489



Pdf

Legal Efforts on Forensic Examination of the Death of Polling Station Working Committees (KPPS) Members

Dewi Iriani¹, Abdul Kadir¹, Agus I Supriyanto¹, Ahmad Arifin¹, Made Wira Suhendra¹

1490-1495



Pdf

The Burden of Toxoplasma Gondii In Spontaneous Miscarriage and its Association with Rhopty Protein 5 Gene and Toxoplasma gondii specific primers GRA 6 Gene Expression in Iraq Women

Dhamyaa Kareem Kadhim¹, Abdulsada Abdulabbas Rahi¹

1496-1500



Pdf

Changes of Interleukin-6 (IL-6) and Immunoglobulin G (igG) in Respiratory Exercise

Didik Agus Santoso¹, FM. judajana¹, Elyana Asnar²

1501-1506



Pdf

Association between Stretching Exercise with Virtual Reality Game and Over Head Pulley of Frozen Shoulder Patients

Donny Gunawan¹, Reni Hendrarati Kusharyaningsih¹, Noor Idha Handajani¹

1507-1512



Pdf

Correlation of Serum Alkaline Phosphatase, Lactate Dehydrogenase, C-Reactive Protein, Blood Deposition Rate, β -Hcg Expression and Tumor Volume to Lung Metastasis Risk in Osteosarcoma Patients

Achmad Fachrizal¹, Dwikora Novembri Utomo¹

1513-1517



Pdf

Effect of Amniotic Membrane-Derived Mesenchymal Stem Cells on TNF- α Expression and Inflammatory Cells Infiltration during Vesicovaginal Fistule Repair Healing Process

Eighty Mardiyani Kurniawati¹, Agung Nur Rachman W¹, Budi Santoso¹, Harry Parathon¹, Widjiati², Fedik A. Rantam², Tri Hastomo¹

1518-1522

[Pdf](#)

suPAR is a Bad Omen for Chronic Kidney Disease Progression

Ekhlas Abdallah Hassan¹, Fayhaa M. Khaleel²

1523-1530

[Pdf](#)

Interventions on Women Under Intimate Partner Violence: An Integrative Review

Elaheh Asadi-Bidmeshki¹, Jamileh Mohtashami², Foroozan Atashzadeh-Shoorideh², Meimanat Hosseini³

1531-1535

[Pdf](#)

The Effectiveness Comparison of Valproic Acid 500 mg and Amitriptyline 15 mg in Reducing the Frequency of Headache Attack in Patients with Tension-Type Headache

Elsa Susanti¹, Isti Suharjanti¹

1536-1540

[Pdf](#)

Decrease of Epstein-Barr Virus Anti Early Antigen Immunoglobulin a Levels and Primary Tumor Size in PostCisplatin-Paclitaxel Chemotherapy in Nasopharyngeal Carcinoma Patients

Elvi Syahrina Fiorisa¹, Achmad Chusnu Rhomdhoni¹, Budi Sutikno¹, Anggraini Dwi Sensusiati¹

1541-1545

[Pdf](#)

Correlation of Cardiorespiratory Fitness Levels with Functional Mobility Ability in Post Thrombotic Infarction Stroke Patient

Eny Susilowati¹, Reni Hendrarati Kusharyaningsih², Dewi Poerwandari²

1546-1551

[Pdf](#)

The Early Detection in Gestational Diabetes Mellitus at Indonesia Primary Health Care

Erni Rosita Dewi¹, Budi Prasetyo², Muhammad Ardian Cahya Laksana², Hermanto Tri Joewono², Ivon Diah Wittiantika¹

1552-1556

[Pdf](#)

Observation of Oral Contrast Media Filling Into Lumen Appendix

Faradilla litaloly¹, Hartono Yudi Sarastika¹

1557-1560



Pdf

Study the Biological Efficiency of an Isolated Terpenic Compound from the Ether Petroleum Extract and the Trace Elements of the Plant Tribulusterrestris

Fatima S.Sabah Eman A.Mukhaifi

1561-1566



Pdf

Medico-legal Significance of Diatoms Detection in Different Organs of Drowning Victims

Haidar N. Hussein¹, Nabeel G. Hashim², Niran M. Ahmed³

1567-1573



Pdf

Evaluation of Nurses' Knowledge and Attitudes toward Pain Management at Baghdad Teaching Hospitals

Haider M. Majeed¹, Ahmed F.Hassan¹, Raja I. Abid²

1574-1579



Pdf

The Effect of Empowerment Program on Participation of Mothers with Premature Infants Hospitalized in Neonatal Intensive Care Unit

Hania Sajadi¹, Golbahar Akhundzadeh², Hamid Hojjati³

1580-1585



Pdf

Association of Diffusion Weighted Magnetic Resonance Imaging Profile and Apparent Diffusion Coefficient Value with Brain Tumor's Histopathology

Hartati Rusmi Tri Wilujeng¹, Anggraini Dwi Sensusiaty², M. Yamin Sunaryo Suwandi³

1586-1593



Pdf

Mitochondrial 16S rRNA gene-dependent Blood typing as a Forensic Tool

Hayder J.H. Al-Nayili¹, Hussein O.M. Al-Dahmoshi²

1594-1602



Pdf

Curcumin Improves the Regulation of Ovarian Folliculogenesis in Culture Media with Peritoneal Fluid from Infertile Women with Endometriosis

Hendy Hendarto¹, M Yohannes Ardianta Widyanugraha¹, Widjiati², Soehartono Darmosoekarto¹

1603-1608

[Home](#) / [Editorial Team](#)

Editorial Team

Editor in Chief

Prof S K Dhattarwal

Forensic Medicine, PGIMS, Rohtak, Haryana

E-mail: editor.ijfmt@gmail.com

EDITORIAL ADVISORY BOARD

1. Prof Sudhir K Gupta, Head, FMT. AIIMS, New Delhi , India
2. Prof Mete Gulmen ,Cukurova University, TURKEY
3. Prof. Leandro Duarte De Carvalho , Minas Gerais, Belo Horizonte, BRAZIL
4. Dr. Valery Gunas, National Pirogov Memorial Medical University,Vinnytsya, UKRAINE
5. Dr. Rahul Pathak Forensic Science, Dept of Life Sciences ,Anglia Ruskin University, Cambridge, UNITED KINGDOM
6. Prof Emilo Nuzzalese, University of Turin , Italy
7. Dr Noha A. Magdy Elrafie, Forensic Toxicology, Ain Shams University, Cairo, EGYPT
8. Dr Rituja Sharma, Associate Prof, Law, Banasthali Vidyapeeth Jaipur
9. Dr Shankar Bakkanwar (*Associate Professor*) Forensic Medicine, Kasturba Medical College, Manipal, Karnataka
10. Dr K. Ravikumar , Raksha Shakti University, Ahmedabad, Gujrat.
11. Dr C. Behera (*Adtl. Prof*) Dept of FMT, AIIMS, New Delhi
12. Dr. Kanak Lata Verma, Deputy Director, Toxicology ,RFSL, Chanakyapuri New Delhi
13. Dr. Asha Srivastava (*Senior Scientific Officer*) Forensic Psychology,Central Forensic Science Laboratory, CBI, Delhi
14. Dr. Raghvendra Kumar Vidua, (Associate Prof), FMT, AIIMS Bhopal
15. Dr. Vaibhav Saran (*Asst.Prof.*) School of Forensic Science, Sam Higginbottom Institute of Agriculture Technology & Sciences,Allahabad
16. Ms Aparna R. Asst. Prof. Forensic Serology & Biology, Jain University, Bengaluru
17. Dr. Deepali Jain, Asst Prof, Forensic Science ,BB Ambedkar University, Lucknow
18. Prof. NK Aggrawal Forensic Medicine, UCMS, Delhi
19. Prof. Manoj Kumar Mohanty, Forensic Medicine, AIIMS, Bhuvneshwar, Odisha
20. Prof. Amar Jyoti Patowary, Forensic Medicine, NEIGRIHMS, Shillong, Meghalaya
21. Prof S. Venkata Raghava , Forensic Medicine, Banglore Medical College, Bengaluru
22. Prof. Shalini Gupta Oral Pathology and Microbiology, Faculty of Dental Sciences, King George Medical University, Lucknow
23. Prof. Virender Kumar Chhoker Forensic Medicine, Santosh Medical College, Ghaziabad, UP
24. Prof. Dayanand G Gannur , Forensic Medicine , Shri BM Patil Medical College, Hospital & Research centre, Bijapur, Karnataka

25. Prof Praveen Arora, Forensic Medicine, SAIMS, Indore, MP
26. Prof Barkha Gupta , Saraswathi Institute of Medical Sciences Hapur, Uttar Pradesh India
27. Prof M Prashant Apollo Medical College Hyderabad
28. Prof Dimple Patel , Anatomay, AMC MET Medical College , Ahmedabad , Gujarat
29. Dr Mohammed Nader Shalaby, Associate Professor of Biological Sciences and Sports Health Department, Faculty of Physical Education, Suez Canal University, Egypt

30 Dr. Avinash Harishchandra Waghmode

Professor and Head, Dept of Forensic Medicine and Toxicology, BKL Walawalkar Rural
Medical College Chiplun Ratnagiri

31 Dr. Anita Yadav Assistant Professor Forensic Science, SBAS Galgotias University, Greater
Noida, UP

32 Dr. Risha Jasmine Nathan *Lecturer in Forensic Chemistry Anglia Ruskin University*

East Road, Cambridge, CB1 1PT England, United Kingdom

Current Issue

ATOM 1.0

RSS 2.0

RSS 1.0

[Make a Submission](#)

Browse

[Open Journal Systems](#)

Information

[For Readers](#)

[For Authors](#)

[For Librarians](#)

Platform &
workflow by
OJS / PKP

Decrease of Epstein-Barr Virus Anti Early Antigen Immunoglobulin a Levels and Primary Tumor Size in Post-Cisplatin-Paclitaxel Chemotherapy in Nasopharyngeal Carcinoma Patients

Elvi Syahrina Fiorisa¹, Achmad Chusnu Rhomdhoni¹, Budi Sutikno¹, Anggraini Dwi Sensusiati¹

¹Department of Otolaryngology-Head and Neck Surgery, Faculty of Medicine Universitas Airlangga, Jl. May. Jend. Dr. Moestopo No.6-8 Surabaya, Indonesia

Abstract

Background: Nasopharyngeal carcinoma (NPC) associated with Epstein-Barr virus (EBV) chronic infection is a common head and neck malignancy in South China and Indonesia. Although radiation and chemotherapy is the main therapy, it requires repeated and invasive biopsy for pathological evaluation. Therefore, a marker is required for screening including the level of anti early antigen immunoglobulin a serology level.

Method: Pre-pots test, longitudinal cohort design. The PTV of 18 samples were examined using CT scan, while their serum EBV anti EA IgA level were examined using pre and post three series-cisplatin-paclitaxel chemotherapy ELISA.

Results: Although there was no significant changes in the level of anti EA IgA, however we found a decrease in the mean of pre-chemotherapy anti EA IgA level from 136.49 U/ml to 124.61 U/ml. There was significant changes in the VTP in pre and post-chemotherapy ($p < 0.05$). The mean of VTP in pre-chemotherapy was 66.26 cm³ (SD-38.61 cm³), while in post-chemotherapy was 31.64 cm³ (SD-27.5 cm³). The delta mean of changes in anti EA IgA level was 11.8 U/ml and in VTP was 34.62 cm³. No correlation was found between the changes of anti EA IgA and the changes of VTP in post-chemotherapy ($p > 0.05$). However, decreases were found in the level of EBV EA IgA and PTV in pre and post NPC patients.

Conclusion: There were decreases of serum EBV EA IgA level and PTV in pre and post-chemotherapy NPC patients.

Keywords: *Immunoglobulin A, primary tumor volume, cisplatin-paclitaxel chemotherapy, nasopharyngeal carcinoma*

Introduction

Nasopharyngeal carcinoma (NPC) is a rare head and neck malignancy except in South China and Southeast Asia including Indonesia. NPC incidence in South China

is between 20-40 per 100,000 population per year and in Indonesia is 6.2 per 100,000 population per year ¹ In Dr. Soetomo General Hospital and Dr. Cipto Mangun Kusumo General Hospital as the main hospital in Surabaya and Jakarta, Indonesia, respectively, nasopharyngeal carcinoma ranks fourth in malignancy after carcinoma of the cervix, breast and skin ².

EBV infection will be followed by the formation of specific antibodies against EBV antigens including anti viral capsid antigen (VCA), early antigen (EA), and Epstein-Barr nuclear antigen (EBNA). Increased

Corresponding authors:

Elvi Syahrina Fiorisa

Email: syahrinaelvi6@gmail.com

Department of Otolaryngology-Head and Neck Surgery, Faculty of Medicine Universitas Airlangga, Jl. May. Jend. Dr. Moestopo No.6-8 Surabaya, Indonesia

levels of Immunoglobulin A (IgA) anti EA and VCA are commonly found in patients with KNF³. Anti-EBV antibody levels, particularly IgA in NPC patients are higher than healthy individuals or patients with other kinds of malignant head and neck tumor, other organ tumors and even in other nasopharyngeal disorders³.

Post-therapy NPC monitoring associated with EBV infection is performed with painful repeated biopsy and pathology examination. EBV serology examination can be used as screening for at-risk patients and occult primary tumors as well as to detect recurrence⁴. A research shows that there is a significant association between serum EBV DNA levels of KNF patients with clinical staging and tumor progression⁵ It is expected that EBV serology may replace the role of the biopsy.

Several studies have shown that the levels of IgA anti VCA and EA EBV increase with the appearance of NPC symptoms. Immunoglobulin A anti-EA is a tumor marker for the diagnosis of NPC because it exhibits high specificity compared to other tumor markers and IgA anti-EA will increase 1 - 5 years before NPC³. IgA levels in pre-therapy have diagnostic and prognostic value, whereas NPC patients with higher levels of antibodies have a worse prognostic⁶. The remaining high levels of IgA anti VCA and anti-EA after therapy are associated with poor prognostics. The increased serum immunoglobulin A (IgA) EBV serologic levels with normal histopathologic results should still be warranted for greater recurrence or higher risk of recurrence

Serum IgA anti EA EBV serologic examination is required for post-therapy NPC evaluation. The purpose of this study was to determine the relationship between

changes in IgA anti EA EBV levels in serum with primary tumor volume in post cisplatin-paclitaxel NPC patients in Dr. Soetomo General Hospital Surabaya. The results of this study are expected to be used as a basis for assessment of therapeutic response, early detection of KNF recurrence, and prognosis determination.

Method

This is an observational study with longitudinal cohort approach using pre-post test. The study was conducted in Department of ENT Dr. Soetomo General Hospital Surabaya in the period of August 2016 until January 2017. 24 new NPC patients were collected as the samples from a total of 25 patients that met the study criteria. Six samples were dropped-out due to changes in paclitaxel regiment, one patient due to an allergy, one patient continued the chemotherapy procedure in outside Dr. Soetomo General Hospital Surabaya, one patient refused to continue chemotherapy, two patients have undergone radiotherapy before the chemotherapy finished and one patient passed away.

The examination of serum EBV EA IgA level was conducted using. The basic data collected in this study consisted of the patients data based on age, gender and ethnicity. The examination results in the form of pre and post cisplatin-paclitaxel chemotherapy serum EBV anti EA IgA in NPC patients was assessed by Clinical Pathology Consultant. Pre and post cisplatin-paclitaxel chemotherapy primary tumor volume was assessed by radiology consultant. The data were analyzed using Wilcoxon Signed Rank Test and rho Pearson correlation test. This Research to find out the changes of serum EBV anti EA IgA and primary tumor volume (PTV).

Results

Table 1. The results of anti-EA IgA in post 3-series cisplatin-paclitaxel chemotherapy

Cisplatin – paclitaxel Chemotherapy	Anti EA IgA level (U/ml)	Primary Tumor Volume (PTV) (cm3)
Pre		
Mean	136.49	66.26
Median	86.54	59.30
Standard Deviation	140.38	38.61
Post		
Mean	124.61	31.64
Median	93.07	20.15
Standard Deviation	127.80	27.55
Δ(delta)		
Mean	11.88	34.62
Median	8.41	23.50
Standard Deviation	55.88	36.85

The most NPC patients were in the age group of 40-49 years old with seven patients (38.89%). The youngest age was 19 years old and the oldest was 62 years old. Distribution of NPC patients based on gender was described in patients were male (13 patients or 72.22%) and there were 5 female patients (27.78%). The comparison between male and female was 2.6:1. Distribution of NPC patients based on ethnicity were Javanese with 14 patients (77.78%) compared to Madura 4 people (22.22%).

The result of anti-EA IgA level in pre-chemotherapy was found to be 136.49 U/ml and standard intersection of 140.38 U/ml. The results of anti-EA IgA in post 3-series cisplatin-paclitaxel chemotherapy were found to average 124.61 U/ml and standard intersection of 127.80 U/ml (Table 1). Statistic test using Wilcoxon Sign Rank Test showed p value of 0.053. This indicated no significant changes in anti-EA IgA level in pre and post cisplatin-paclitaxel chemotherapy in patients with NPC ($p > 0.05$)

The results of PTV in pre-chemotherapy cisplatin-paclitaxel showed the mean of 66.26 cm³ and standard deviation of 38.61 cm³. The results of precisplatin-paclitaxel showed the mean of 66.26 cm³ and standard deviation of 38.61 cm³. Statistic test using Wilcoxon Sign Rank Test showed p value of 0.001. The data indicated a significant difference of PTV in pre and post cisplatin paclitaxel chemotherapy in NPC patients ($p < 0.05$).

Changes in pre and post cisplatin-paclitaxel anti EA IgA levels showed Δ mean of 11.88 and Δ standard deviation of 55.88. Changes of PTV in pre and post-cisplatin-paclitaxel chemotherapy had Δ mean of 34.62 and Δ standard deviation of 36.85 (Table 4). Statistical test using Pearson correlation showed correlation coefficient (r) of 0.260 and p value of 0.298, suggesting a insignificant correlation between the changes in anti-EA Ig A level and primary tumor volume in post cisplatin-paclitaxel chemotherapy in NPC patients

Discussion

Distribution of NPC patients based on gender was the 13 male patients (72.22%) and 5 female patients (27.78%). The ratio between men and women was 2.5:1. Distribution of KNF patients based on gender, most patients were male (70%) with the ratio between men and women being almost the same throughout Indonesia, i.e. 2-3:1.²⁴ The habits such as smoking increase the risk of KNF 2-6 times and so did the exposure to steam, dust

and chemical gas at the workplace also increase the same risk. Formaldehyde exposure at workplace also increase the risk of NPC to 2-4 times. Increased risk also occurred in workers who inhaled firewood smoke, and the risk increased 2 times in workers exposed to industrial heat and combustion products. This led to high incidence in men due to differences in living habits and occupations that cause males to have more frequent contact with carcinogens that caused NPC.² Testosterone hormone which was dominant in male was suspected of causing immune response and tumor surveillance decrease and thus male are more prone to EBV infection and cancer⁷

The results of statistical analysis of changes in the levels of serum EBV anti EA IgA in pre and post-chemotherapy cisplatin-paclitaxel pada patients with NOC was not significant. Nevertheless, there was a mean decrease by 136.49 U/ml in pre-chemotherapy to 124.61 U/ml in post-chemotherapy (Table 1). This was incompatible with the chemotherapy mechanisms that caused humoral and cellular immune suppression. Humoral immunity expressed by B cells and assessed on the level of immunoglobulin.²⁷ The similar mechanism in radiotherapy was given to NPC patients which often caused immunologic cell damage which resulted in decrease of cellular humoral immune response⁸

Decreased serum EBV anti EA IgA levels after cisplatin-paclitaxel chemotherapy in this study might be due to abundant EBV not only in NPC tumor cells but also from activation of T cell infiltration, B lymphocytes, and epithelial cells capable of producing antigen associated with EBV.^{28,29} In addition, there was a difference of individual immune responses to various antigens which made antibodies as important markers were highly dependent on the host response to the viral antigen on the tumor.¹⁰ Changes in the latent cycle into lytic cycles in NPC tumor cells could occur spontaneously or induced by cisplatin chemotherapy, γ ray radiation, phorbol ester, sodium butyrate and bortezomid.^{30,31}

Levels of EBV EA Ig A highly increased in 2 patients with NPC i.e. 19.84 U/ml in pre-therapy to 184.31 U/ml in post-therapy and 84.07 U/ml to 113.72 U/ml and 4 patients with NPC had slight increase, i.e. around \pm 2.00 U/ml. The levels of EBV EA Ig A in patients with NPC in this study decreased in 12 patients. This was consistent with the study by Gu, et al.³² (2009) obtaining fluctuation of EBV antibody reactivity during therapy and steady follow-up. Fifteen of the 35 NPC patients tested for antibodies decreased after therapy but

13 patients showed small changes. Increased levels of EBV antibody after therapy were obtained in 5 patients while 2 patients that initially decreased had increase. This might be due to the different kinetic diversity of serologic EBV for each KNF patient during therapy as well as illustration of differences in radiosensitivity and immunological reactions⁹

The results of the measurement of primary tumor volume (PTV) in post-chemotherapy cisplatin-paclitaxel showed a decrease of PTV mean in post-chemotherapy cisplatin-paclitaxel. The PTV change was statistically significant and therefore it was concluded that there was a significant PTV change in pre and post chemotherapy¹⁰ This corresponded to the use of cisplatin-paclitaxel chemotherapy in the management of malignant tumors, i.e. to eradicate tumor cells or for locoregional control when used in conjunction with surgery or radiotherapy. Chemotherapy was used to treat macroscopic and microscopic metastases. Microscopic metastasis that was clinically invisible and deposited in the body would turn into macroscopic if not treated¹¹.

Pearson correlation test result on serum EBV anti EA IgA level changed with primary tumor volume change (PTV) in post- cisplatin-paclitaxel chemotherapy got the value $r = 0,260$ and $p = 0,298$. The results showed a non-significant relationship ($p > 0.05$). However, in general we found a decrease in EBV EA IgV levels and decreased primary tumor volume in patients with NPC who received pre and post cisplatin-paclitaxel chemotherapy. Interaction of cancer cells with lymphocytes or chemotherapy and radiotherapy that could have a significant effect on the immune system my explained it, and also the imbalance of humoral immunity and decreased cellular immunity triggered cancer progression and treatment failure.³³ In this study changes in serum anti-EA IgA might be due to direct effects of chemotherapy or indirect effects of viral replication associated with growth of NPC tumor¹² The immediate effects of chemotherapy and cytostatic drugs were known to greatly damage lymphoid as an antibody producer but some studies have found that chemotherapy did not destroy all B memory cells and memory T cells even when examination did not get humoral immune response.³⁴ Increased response of various antibodies to EBV protein at a higher stage of malignancy suggested viral replication associated with NPC tumor growth⁹

A study¹³ showed no statistically significant association between gender, age, and distant metastases

with anti-VCA IgA antibodies, anti-EA IgA, anti-RGA IgG, and anti-EBNA IgA. There was a tendency to increase the levels of anti-VCA IgA and anti-EA IgA with a classification of N, but there was no association with the classification of T. This might be related to lymphocyte infiltration in NPC which significantly contributed to increase antibody to EBV antigen lytic phase.

The results of this study were consistent with a study³⁰ that various chemotherapeutic drugs such as cis-platinum, fluorouracil, and taxol trigger alteration of latent cycle into lytic EBV virus infection in tumor cells. This change occurred through the protein signal kinase C δ , phosphatidylinositol 3'-kinase, and p38 stress mitogen-activated protein kinase but not caspase activation³. This study was also conducted on mice containing KNF cells and found that combination of GCV therapy with 5 FU or a combination of GCV with cis-platinum being more effective in triggering KNF cell apoptosis than single therapy. Spontaneous induction of viral replication was more determined by the intracellular plasma environment than the factors that cause plasma cell differentiation. Lit replication of infected latent cell lines might be performed by induction including anti-immunoglobulin antibody (anti-IgG), activation of transforming growth factor β (TGF β), and activation of CD4 + T cells. The transition from the latent cycle to the EBV lytic cycle could be induced by DNA destruction agents such as chemotherapy (cisplatin), γ ray radicals, phorbol ester, sodium butyrate, and bortezomid.^{30,31}

Conclusion

There was a decrease in serum EBV EA IgA level and primary tumor volume n post-chemotherapy compared to in pre-chemotherapy and therefore could be used to monitor the success of post-chemotherapy medication.

Conflict of Interest: There is no conflict of interest.

Source of Funding: This study is self-funded.

Ethical Clearance: This study was approved by Ethical Commission of Health Research Faculty of Medicine University of Airlangga.

References

1. Suryaningtyas W, Arifin M, Rantam FA, Bajamal AH, Dahlan YP, Dewa Gede Ugrasena I, et al. Erythropoietin protects the subventricular zone

- and inhibits reactive astrogliosis in kaolin-induced hydrocephalic rats. *Child's Nerv Syst* [Internet]. 2019;35(3):469–76. Available from: <https://www.scopus.com/inward/record.uri?eid=2-s2.0-85060337867&doi=10.1007%2Fs00381-019-04063-w&partnerID=40&md5=4a3971bdcd1139229d20020350e8eca9>
2. Nuryadin I, Prasetyo A. Profil Imunopositivitas Protein Ebv Pada Penderita Karsinoma Nasofaring Dan Individu Sehat Berisiko. Fakultas Kedokteran; 2012.
 3. Tiwawech D, Srivatanakul P, Karaluk A, Ishida T. Significance of plasma IgA and IgG antibodies to Epstein-Barr virus early and viral capsid antigens in Thai nasopharyngeal carcinoma. *Asian Pacific J cancer Prev*. 2003;4(2):113–8.
 4. Zeng M-S, Zeng Y-X. Pathogenesis and etiology of nasopharyngeal carcinoma. In: *Nasopharyngeal Cancer*. Springer; 2010. p. 9–25.
 5. Reitz C, Tang M-X, Luchsinger J, Mayeux R. Relation of plasma lipids to Alzheimer disease and vascular dementia. *Arch Neurol*. 2004;61(5):705–14.
 6. Haq IBI, Goto T, Kawashima T, Yamanaka K, Osawa M, Ohata K, et al. Malignant transformation of a vestibular schwannoma to malignant peripheral nerve sheath tumor 10 years after Gamma Knife Surgery: Case report. *Interdiscip Neurosurg Adv Tech Case Manag* [Internet]. 2019;18. Available from: <https://www.scopus.com/inward/record.uri?eid=2-s2.0-85069649887&doi=10.1016%2Fj.inat.2019.100529&partnerID=40&md5=cf6c5b05a45adfc01ef69eb860fbc6d0>
 7. Tio TH, Djojopranoto M, Tjoei NI. Subcutaneous Phycomycosis. *Arch Dermatol* [Internet]. 1966;93(5):550–3. Available from: <https://www.scopus.com/inward/record.uri?eid=2-s2.0-0013909056&doi=10.1001%2Farchderm.1966.01600230054016&partnerID=40&md5=cf405e0b4fb70878e2e8c3fdb78b0882>
 8. Adham M, Kurniawan AN, Muhtadi AI, Roezin A, Hermani B, Gondhowiardjo S, et al. Nasopharyngeal carcinoma in Indonesia: epidemiology, incidence, signs, and symptoms at presentation. *Chin J Cancer*. 2012;31(4):185.
 9. Fachiroh J, Schouten T, Hariwiyanto B, Paramita DK, Harijadi A, Haryana SM, et al. Molecular diversity of Epstein-Barr virus IgG and IgA antibody responses in nasopharyngeal carcinoma: a comparison of Indonesian, Chinese, and European subjects. *J Infect Dis*. 2004;53–62.
 10. Prakoeswa CRS, Pratiwi FD, Herwanto N, Citrashanty I, Indramaya DM, Murtiastutik D, et al. The effects of amniotic membrane stem cell-conditioned medium on photoaging. *J Dermatolog Treat* [Internet]. 2019;30(5):478–82. Available from: <https://www.scopus.com/inward/record.uri?eid=2-s2.0-85057226592&doi=10.1080%2F09546634.2018.1530438&partnerID=40&md5=354d1acd61ab791a8232ea739d6321fa>
 11. Cai Y-L, Li J, Lu A-Y, Zheng Y-M, Zhong W-M, Wang W, et al. Diagnostic significance of combined detection of Epstein-Barr virus antibodies, VCA/IgA, EA/IgA, Rta/IgG and EBNA1/IgA for nasopharyngeal carcinoma. *Asian Pac J Cancer Prev*. 2014;15(5):2001–6.
 12. Hasegawa T, Miyamoto-Takasaki Y, Abe M, Qiu Z, Yamamoto T, Yimin, et al. Histochemical examination on principal collagen fibers in periodontal ligaments of ascorbic acid-deficient ODS-od/od rats. *Microscopy* [Internet]. 2019;68(5):349–58. Available from: <https://www.scopus.com/inward/record.uri?eid=2-s2.0-85073084631&doi=10.1093%2Fjmicro%2Fdfz021&partnerID=40&md5=dd9ba737c715ecba1d775a449f939f3e>
 13. Rosyidi RM, Priyanto B, Sari SF, Anggraini MA, Kamil M, Wardhana DPW. Subdural drainage of liquor cerebrospinal and early tracheostomy: Alternative management of severe traumatic brain injury with minimal lesion in limited facility and rural area. *Interdiscip Neurosurg Adv Tech Case Manag* [Internet]. 2020;19. Available from: <https://www.scopus.com/inward/record.uri?eid=2-s2.0-85074561052&doi=10.1016%2Fj.inat.2019.100614&partnerID=40&md5=3bd97679ee70dc7feb9bcaacef08623>