

## Critical conclusions and evidences projected from inhibition of COVID-19

Based on informations gathered from various research articles published online related to factors affecting Genome signals & cell differentiations ,the following conclusions have been distinguished and supported with references.

HYPERLINKS

### 1.CELLULAR

DIFFERENTIATION([https://en.m.wikipedia.org/wiki/Cellular\\_differeentiation](https://en.m.wikipedia.org/wiki/Cellular_differeentiation))

### 2.DIFFERENTIATION

THERAPY(<https://www.ncbi.nlm.nih.gov/pmc/articles/PMC4704415/>)

### 3.IKK $\alpha$ Kinase Regulates the DNA Damage Response and Drives Chemo-resistance in

Cancer(<https://www.sciencedirect.com/science/article/pii/S1097276519304332>)

### 4.Corona virus

-wikipedia(<https://en.m.wikipedia.org/wiki/Coronavirus>)

### 5.NF- $\kappa$ B

WIKIPEDIA(<https://en.m.wikipedia.org/wiki/NF-%CE%BAB>)

6.APOPTOSIS-WIKIPEDIA(<https://en.m.wikipedia.org/wiki/Apoptosis>)

### 7.ATM in DNA repair in

cancer(<https://www.sciencedirect.com/science/article/abs/pii/S0163725819301287>)

### 8.Non smallCell lung cancer (NSCLC)

<https://www.cancer.org/cancer/lung-cancer/detection-diagnosis-staging/staging-nsclc.html>

## Conclusion 1.

It is evident that COVID -19 is silent and non detectable for some period initially(14-28 days)once entered in human body and is tested positive only after that period.This characteristic make it different from other type of viruses producing immediate symptoms.This characteristic can be attributed only to its dependant morphogenetic structure ruling out an independant morphogenetic cell structure to it unlike fully developed cells.Because when two cells of two living organisms(one host and other infectant) come in contact and if both have morphogenetic structure definitely their signals will interact and identify infected germ immediately.So, it is obvious that the COVID -19 upon entering body copies genetic expression from host cell and rests as an adjacent cell without any distinguishability.The body(nucleus and cytoplasm) of COVID-19 at that time embeds as boundary of host cell undistinguishably due to its non independant morphogenic configuration adopted from host.

This above conclusion can be justified from experimental study conducted at HUNAN(WUHAN) INSTITUTE OF VIROLOGY,CHINA with report that the RNA pattern of COVID-19 infected on mammals, viz., CAVE BATS resembles RNA pattern of the affected(bat).It is possible only the COVID-19 ,due to its non morphogenic structure adopts such one from infected cell.

Conclusion2.The virus influences epigenetic action during differentiation of host body cell disrupting NF- $\kappa$ B signalling,vital for production and cell survival . As it incorporates swallowed RNA

pattern of the host it easily overrides the gene expressions of hostcell genomes by linking its RNA .

[NF- $\kappa$ B is found in almost all animal cell types and is involved in cellular responses to stimuli such as stress, cytokines, free radicals, heavy metals, ultraviolet irradiation, oxidized LDL, and bacterial or viral antigens. NF- $\kappa$ B plays a key role in regulating the immune response to infection. Incorrect

regulation of NF- $\kappa$ B has been linked to cancer, inflammatory and autoimmune diseases, septic shock, viral infection, and improper immune development. NF- $\kappa$ B also been implicated in processes of synaptic plasticity and memory.

While in an inactivated state, NF- $\kappa$ B is located in the cytosol complexed with the inhibitory protein I $\kappa$ B $\alpha$ . Through the intermediacy of integral membrane receptors, a variety of extracellular signals can activate the enzyme I $\kappa$ B kinase (IKK). IKK, in turn, phosphorylates the I $\kappa$ B $\alpha$  protein, which results in ubiquitination, dissociation of I $\kappa$ B $\alpha$  from NF- $\kappa$ B, and eventual degradation of I $\kappa$ B $\alpha$  by the proteasome. The activated NF- $\kappa$ B is then translocated into the nucleus where it binds to specific sequences of DNA called response elements (RE). The DNA/NF- $\kappa$ B complex then recruits other proteins such as coactivators and RNA polymerase, which transcribe downstream DNA into mRNA. In turn, mRNA is translated into protein, resulting in a change of cell function. But if the signal is under influence of adhering COVID-19 the transcription of downstream DNA to mRNA is epigenetic and differentiation is not completed to yield proteins but silence the [IKK \$\alpha\$  Kinase](#) .The nuclear factor  $\kappa$ B (NF- $\kappa$ B )signaling pathway regulates innate and acquired immunity and is essential for most physiological processes but also for cancer progression (reviewed in Zhang et al., 2017). Multiple extracellular stimuli, including the inflammatory cytokines tumor necrosis factor  $\alpha$  (TNF- $\alpha$ ) and interleukin 1 $\beta$  (IL-1 $\beta$ ), induce NF- $\kappa$ B through a series of

signaling events that lead to the phosphorylation and activation of a kinase complex that consists of IKK $\alpha$ , IKK $\beta$ , and IKK $\gamma$ /NEMO. The IKK $\alpha$  subunit is dispensable for NF- $\kappa$ B activation, but it has been found to exert multiple pro-tumorigenic functions. Several studies have demonstrated that IKK $\alpha$  enhances the metastatic activity of prostate tumors (Luo et al., 2007) and squamous cell carcinomas (Toll et al., 2015) by regulating the Maspin gene. Epithelial IKK $\alpha$  is required for the initiation and progression of intestinal adenomas (Colomer et al., 2018) and lung adenocarcinomas (Vreka et al., 2018) in mice. Recently, it was identified, a nuclear active form of the IKK $\alpha$  kinase, IKK $\alpha$ (p45), which is localized in the nuclear compartment of cancer cells. IKK $\alpha$ (p45) induces the phosphorylation of histone H3 and nuclear co-repressors, which is dependent on its interaction with non-activated full-length IKK $\alpha$  and promotes tumor growth independent of canonical NF- $\kappa$ B signaling (Margalef et al., 2012)].

So, it is obvious from the visibility of hectic trouble in breathing during COVID-19 attack, the layer of transformed virus cells resembling adenocarcinomas gets formed instead of tumors developed by suppression of IKK $\alpha$ (p45) during NASOPHARYNGEAL CARCINOMA. The might be cancer cells have been nurtured as COVID-19 cells by utilising the same constituents available. Instead of developing its own structure the virus poses itself in adopted or Pseudo structure and gets proliferated to large no of cells by mutation as like in the case of cancer and that is why sudden criticality in health condition precedes. This phenomenon may be termed as

### **PSEUDOMORPHOLOGY OF COVID-19.**

CONCLUSION3. The matching aspects between cancer spread and COVID-19 spread in human body is closely distinguished as both occur during improper or halted differentiation and instead of activation of growth factors, due to epigenetic interference there is possibility of FAS LIGANDS or

death signals active in the cells and APOPTOSIS likely to occur reducing immuno power of body. Also the tumor supporting constituents lead to proliferation of enemy cells in bulk. Only thing is tumour is solid but COVID-19 is semisolid accumulation suppressing the function of respiratory organs.

CONCLUSION 4. The Differentiation therapy plays prominent role in bringing NSP Carcinoma (Poor differentiation is an important hallmark of cancer cells, and differentiation therapy holds great promise for cancer treatment. The restoration of I $\kappa$ B kinase  $\alpha$  (IKK $\alpha$ ) leads to the differentiation of nasopharyngeal carcinoma cells with reduced tumorigenicity.

The findings by [Yan et al.](https://www.mayoclinic.org/diseases-conditions/nasopharyngeal-carcinoma/symptoms-causes/syc-20375529) (<https://www.mayoclinic.org/diseases-conditions/nasopharyngeal-carcinoma/symptoms-causes/syc-20375529>) validate the polycomb protein enhancer of zeste homologue 2 (EZH2) as a target for intervention.) to get cured and restoring cells to normal.

**As the present argument holds similarities existing in protein knockout in tumour generation and covid-19 proliferation, The same (DIFFERENTIATION THERAPY WITH APGRADATIONS) can, if be deployed for COVID-19 infection then not only for curing but also for protection from infection could be realised. As it is because no existing method of reconditioning cells available from medical experiences other than differentiatomn therapy, it is apt to select it.**

*EARLIER ON THIS LINE*

## **THE PSEUDOMORPHOLOGY OF COVID 19**

(communicated on 02/04/2020)

I am an Ordinary citizen of India once was an engineer and IT Professional .I don't have any background in Medical OR Biological related fields.But the today's situation in world after COVID19's Psunami waves rocking entire world ,spontanaeously I came to go through the biological phenomenon associated with development of this deadly virus and in the meantime I am committed to brought to your attention , the fact I conceived,not to superimpose any beyond the conception and conclusion of experts and researchers but to get coincided with their findings perhaps once in the well doing of human life at last.If feasible, the tremendous and untiring workof Medical and Biological workers in the world along with my logical arrival of COVID19's menifestation may help brothersand sisters in the world.

## CONCEPTION:

The basic concept of this argument has its root in the process of cell split which is the basic and essential phenomenon in the growth of tissue structure of living organisms. I am consentient to cell multiplication by splitting as far as fully developed cell structure is concerned but am reluctant to the multiplication of partially developed primordial cells like COVID 19 which lacks DNA configuration. This speculation leads to attribute a different method for the proliferation of COVID 19 which is to be confirmed experimentally by observation and verification through international testing labs.

The cell splitting process leads to the formation of new cells (with both DNA and RNA), eventually it is possible only in that kind of fully developed cells only and not in partially developed cells having RNA alone because **splitting is the unique action** that can be initiated if only DNA also is present in the cell according to the **ancient past bio metamorphic activities information stored in DNA**. For viruses like COVID19 which is only semi grown organism without DNA no such **initiation signal** for cell split process hardly present as there is **no storage spot possible without DNA**. Its cell growth is only **aggregation of already grown cells** in the human or fully developed cells of living bodies cropped as smuggled and exists with continued supply of its past association with original cells.

It could be traced out if the following development of COVID 19 affected cells but not identified physically to an extent as to show critical symptoms. That is the **distinct no of RNA cells of host** body which do not merge with its associate DNA combination of which forms new developed host cells where as the hanging RNA forms the COVID 19's semi developed cells eventually. It is evident that such cells do not act for DNA SIGNAL (Due to epigenecity) and they look isolated once they arise initially in the healthy cells itself and does not do productive work as like original RNA and hence can be named pseudo RNA. The name is so given because in fact these nonintegrated RNA constituents may be later deployed as COVID-19 transformed portions during its sudden proliferation. If the presence of Pseudo RNA is identified and ascertained the infection Of COVID 19 could well be recognised in advance before the spreading goes to dangerous extent which is explained below.

This Pseudo RNA form some newly split good cells and act in opposition to the signal of body immune system (due to epigenetic effect of COVID-19) sent from DNA of white corpuscles because it gets grouped in to RNA's of newly formed COVID 19 primordial cells which attacks host cells. How come the DNA information of new Cells Pseudo RNA fail integrity with body? Is it a ridicule? No. During splitting all the signals are halted within the cell and fission signal only active along the



previously assumed boundary to realise the formation of two new cells. This is the situation when the silent COVID-19 could also try to multiply its cells as it cannot initiate its own cell division due to its Pseudomorphological nature. That fission signal is swallowed by Pseudo RNA (Due to Epigenecity) as like the normal DNA signal which is supposed to cease once the splitting is over extends as residual memory (as like in case of motherboard devices) and the new DNA will not send anymore subsequent metamorphic signal to Pseudo RNA assuming it is already sent. So some of the white corpuscles containing Pseudo RNA do not unite with good cells and tend to form new boundary (like cancer cells) incorporating COVID 19 RNA to serve COVID 19 as a shield and supply platform from which it can proliferate until it multiplies to large no of individual covid 19 cells demolishing life organs. This trial of boundary forming is evident from experimental finding of variety of configurations of COVID-19 in infected persons. Though it cannot form new boundary the time it associates in the formation process helps COVID 19 to grow its species. If the immune system goes on attacking infected cell without screening the negative trend by Pseudo RNA, the defence & counter attack ultimately turns into self destruction. Please note that it is not necessary for apoptosis to complete but initialising it in an inappropriate time is as much as dangerous as it creates instability in cell development.

So, if the lay out of that new boundary is scanned and erased then by removing or disconnecting Pseudo RNA from healthy cells the immune system of humanbody might take full control and cut the roots of COVID 19 once for ever.It is just like the process of erasing virtual memory in the CPU of PC system but is more and more indigeniously complicated as we are talking about the most supreme biological system of human body.But I hope ,as GOD will help to cure, there may seem a scar or temporary dye spots rest in to detect the course of Pseudo RNA , might eradicate error if researchers are able to locate in a body infected with COVID19 but without any trace showing it.However the above confiscation should be precisely tested and verified practically through well established modelling technique and afterwards may be implemented to yield results.

With this I affirm my sincerity and dedication to humankind wotld wide and I assure I submit this only to benefit to mankind at least as a way to future releifs even it is not realised .I beg your pardon if the assumptions do not meet the reality.

**COROLLARY:**In follow up of my previous communication "The Pseudo morphology of COVID19 " on 02/04/2020 ,I put a small note on "APOTOSIS" WHICH IS THE growth factor first discovered by Rita Levi-Montalcini, which won her a Nobel Prize in Physiology or

Medicine. While growth factor implies a positive effect on cell division, cytokine is a neutral term with respect to whether a molecule affects proliferation. While some cytokines can be growth factors, such as G-CSF and GM-CSF, others have an **inhibitory** effect on cell growth or **proliferation**. Some cytokines, such as **Fas ligand**, are used as "**death**" signals; they cause target cells to undergo programmed cell death or apoptosis.

So, the PSEUDO RNA definitely influences the Fas ligands in immunocells and by enhancing the repression of signals to launch de-differentiation( or simply restoration of initial stage before differentiation to cut presence of Pseudo RNA)only the selfdestruction of immunecells due to COVID 19 could be prevented.what I termed earlier as 'VIRTUAL OR FALSE MEMORY " very well fits to DEATH

SIGNAL .The repression process just erases the false signal or stopping PseudoRNA from integrating with cell system.It is quite unclear whether efforts in such direction has been carried out.Simply speaking it is **stopping the growth of other cells for strengthening the immune cells.**