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Research Article

**PREDICTABLE VARIETY IN TUMOR IN NATURAL SCIENCE
FROM MOLECULAR SNOWFLAKES TO IDIOSYNCRASY
STAMP**¹Dr. Aamna Mahmood Gilani, ²Dr. Saba Qayum, ³Dr. Fatima Shoukat¹House Officer JHL**Article Received:** October 2020**Accepted:** November 2020**Published:** December 2020**Abstract:**

Our current research was conducted at Sir Ganga Ram Hospital, Lahore from December 2017 to November 2018. Advancement by regular choice is the applied establishment for almost every part of science and progressively additionally for biomedicine and medicinal research. In disease science, development clarifies how populaces of tissues in cancer alteration after some instance. It is a crucial problem whether this transformative procedure is ambitious basically through common determination and adjustment or by additional transformative procedures, for example, originator impacts and float. In malignancy science, as in individual developmental science, there is discussion concerning this inquiry and additionally regarding the utilization of adjustment during common determination because a directing system for explore. In this survey, we examine the distinctions and likenesses among advancement among physical cells versus development among life forms. We finish up that adjustment is a valuable structure that precisely clarifies the characterizing attributes of malignant growth. Further, merged advancement through regular determination gives the main fulfilling clarification both for how a gathering of assorted pathologies share enough for all intents and purpose to helpfully share the distinct name of "malignancy" and for why this merged condition becomes life threatening.

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INTRODUCTION:

In old style transformative science, it is surely known that in any event, when two such populaces emerge from various hereditary foundations, and along these lines have not many likenesses at the atomic level, a mutual natural specialty and its common specific weights can drive the procedure of developmental adjustment by characteristic choice toward the same characteristics, making the two populaces meet onto those qualities regardless of their extraordinary beginning stages and tireless atomic contrasts [1]. Although transformative adjustment by substantial cells is unique in relation to adjustment by living beings, it is a significant piece of malignancy science furthermore, should be comprehended. One halfway significant application lies in accommodating two significant what's more, differentiating designs in malignant growth science, which we allude to as characteristic "trademarks" and sub-atomic "snowfall shaving." The primary of these both examples to be depicted was the arresting constancy amongst various kinds of malignancy at the degree of cell characters. The obvious logical inconsistency between characteristic "trademarks" and sub-atomic "snowflakes" is settled by understanding the hidden transformative what's more, biological procedure of

malignancy, as particular from its stationary appearances (Merlo et al. 2006; Pepper et al. 2014). Even though they start commencing various tissues and cell kinds, for every intents and purposes all malignant growths comprise of cells with a similar basic "trademark" qualities (Hamadan and Weinberg 2000, 2011). This immediately got one of only a handful few sorting out structures to force request on the confusing assorted variety of malignant growth [2]. The externally differentiating design that has while risen on or after atomic investigation is that practically any atomic classification of diseases be able to, on nearer examination, be separated into sub classes by searching for, and constantly pronouncement, atomic varieties. For sure, sub-atomic contrasts can normally be originate among any two types, driving to the feeling that maybe, similar to flurry shaving, no both malignant growth types are actually indistinguishable (Kurzrock furthermore, Giles 2015). In transformative science, there has extended been discussion over the force and confinements of the "adaptation's plan" as a method to produce theories and direct examine (Gould and Leonine 1979; Mary 1983) [3]. A malignant growth is a unique populace of strange physical cells developing through common determination.

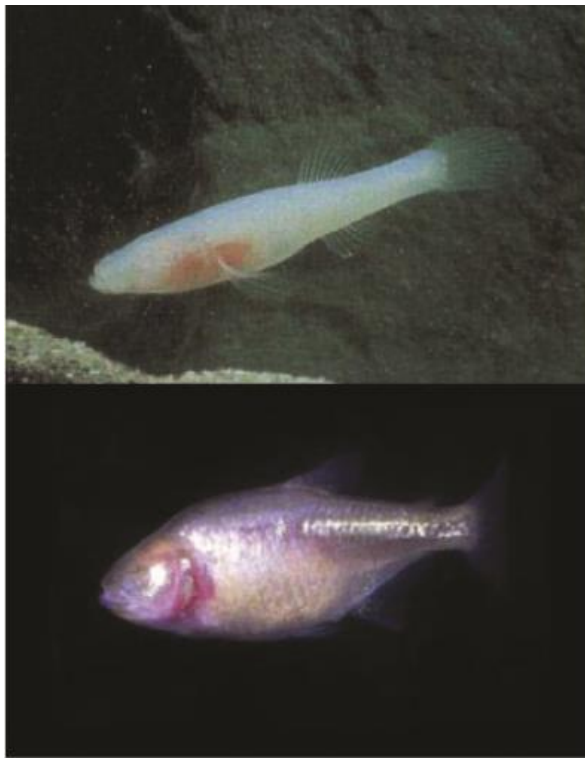


Figure 1: The adaptations approach accentuates that common determination drives populaces to advance useful adaptations. A sample of convergent development in two types of cave fishes whose ancestors are different. *Amblyopsis rosae* (peak) and *Astyanax mexicanus* (underneath)

In actuality, these are bogus polarities. The clear options are both valid. There is acceptable proof for both impartial development and regular determination in neoplasms, and the comparative significance of the two most likely variations throughout movement. Neoplastic cells manufacture multifaceted what's more, narrative adjustments, regularly through evacuating or actuating compartment works that are as of now in the individual gene, yet in addition now and then through hereditary novelties [4]. Similar banter is currently present in malignant growth science, with certain specialists contending against "unnecessary adaptations," halfway on the basis that 20 years is most likely not extensive enough for malignancy cells to advance all their watched complex qualities (Arnal et al. 2015) [5].

METHODOLOGY:

One of the contentions against applying an adaptations structure to malignant growth cells is that the extent of coming about adjustment might be

restricted in light of the fact that the individual natural life is excessively small for multifaceted cell adjustments to develop. With respect to third vital parameter of development, transformation rate, and current evaluations propose that each pedestal couple in the malignancy gene is presumably changed in a few cells, in each cell age inside a neoplasm (Table 1). Our current research was conducted at Sir Ganga Ram Hospital, Lahore from December 2017 to November 2018. This also recommends that physical advancement can be fast furthermore, that its atomic foundations can be assorted, as a similar quality might change autonomously different times inside a similar neoplasm. It shows up to be genuine exactly. Actually, precisely the same change supposedly occurs on various occasions in a similar neoplasm, causing various autonomous starting points, or "homoplasmy" (e.g., Shpak et al. 2015). This abrasion excess likewise clarifies why there are probably going to be numerous systems of procured opposition accessible for a reaction to the choice forced by operation.

Table 1: Approximation and varieties for the limitations of somatic development in neoplasm

Limitations	Approximation	References
Rate of position alteration	10^{-9} - 10^{-10} bp	Knudson 1971; Wang et al. 2002; Araten et al. 2005; Jones et al. 2008; Tomasetti et al. 2013
Rate of intensification and removal	10^{-4} - 10^{-5} gene	Tisty et al. 1989
Rate of failure of heterozygous individuals	10^{-4} - 10^{-6} gene	de Nooij-van Dalen et al. 1998; Chan et al. 2001
CpG methylation speed	10^{-4} CpG	Chan et al. 2001; Sottoriva et al. 2013
Size of whole individuals present	1 cm ³ of solid cancer	Del Monte 2009; Sullivan and Salmon 1972; Friberg and Mattson 1997
Size of stem cells of the individuals	Many myeloma: 10^7 - 10^{10} Tumor in colorectal: speed 10^{-2} - 10^{-3}	Hamburger and Salmon 1977a,b; Ricci-Vitiani et al. 2007; Merlos Suarez et al. 2011; Sottoriva et al. 2013
Invention period of the stem cells	Colorectal tumor in vitro 10 days Stem cells of colon in vivo: one week	Potten et al. 2003; Ricci-Vitiani et al. 2007
Development rate of colon	1.6cm ³ per year in esophagus	Martinez et al. 2016
Discriminatory coefficient	Driver gene average: 0.004	Bozic et al. 2010; Vermilion et al. 2013
Number of cell production	Colorectal tumor: 10^3 - 10^4	Yachida et al. 2010
Time from commencement of identification	Colorectal tumor: 15-65 years Pancreatic tumor: 50-610 years	Ling et al. 2015; Zack et al. 2013

Numerous malignancy cells have the ability to separate day by day, develop over a time of decades, what's more, and include populace's numeral in the larger amount (Table 1). The pace of transformative change relies fundamentally upon limitations like change velocity, populace dimension, and populace revenuespeed (reverse of cell age occasion). Allowing for these components recommend that physical cell development be able to be considerably quicker than advancement between creatures having many cells. In light of on populace dimension and age instance, there are progressively regenerative occasions between the cells inside one host individual than there have been among people in the whole the past of the individual populations. Every such regenerative occasion is an open door for change and specific propagation and, in this manner, for versatile advancement. Another factor quickening cell advancement in neoplasms is that it tends to be driven by hereditary, yet additionally by epigenetic alterations. Epigenetic changes are dependent upon substantial characteristic choice since they are genetic transversely division of cells and be able to influence the wellness of the cell by altering its physical appearance. The pace of epigenomics adjustments is here and there a lot higher than the pace of hereditary transformations (Table 1) thus might be a significant driver of development in neoplasms. In cancer, the quantity of cell ages per element instance might be soaring and just freely identified with the size of the tumor, in view of elevated amounts of cancer and quick cell revenue (Lowe also, Lin 2000; Liu et al. 2001). To gauge numeral of cell ages, the absolute figure of cell separations important to create a cancer dimension must be incorporated through the pace of cell passing. A cancerous cell requires just initiate and repurpose this trail to gain the cell death conduct that is fundamental to tissue intrusion and spread of cancer. We recommend that the enormous dimension and multifaceted nature of the human being gene bestows generous resolvability during recycle for cell-level wellness benefit. The possibility that malignancy cells or the transformed qualities that decide their conduct are not associated with a transformative process since they are bound to fast annihilation in light of the host's demise (brought about independent from anyone else) is certifiably off-base. Purported "transformative suicide" through versatile development is in actuality a surely knew marvel with numerous models (Ibrahim 2014). Versatile cell advancement is famously "limited" with respect to later outcomes, remembering for specific damage to the host person.

DISCUSSION:

Is version additional or less probable in tumor cells because of variations between organismal and somatic development?

To summarize, a portion of the contrasts between physical development and organismal advancement create the advancement of multifaceted adjustments fewer probable in malignancy than in multicellular creatures, remembering restrictions for the complete figure of cell ages and the trouble of trading hereditary substance among cell ancestries. Features that errand version in multi cellular living beings incorporate huge diploid alleles what's more, reproductive generation. Reproductive proliferation includes genomic re-organization, which can add to hereditary variety amongst people what's more, expel malicious changes. In this respect, substantial cell advancement is increasingly comparable to development between individual cell life forms. Similar to substantial cells, numerous eukaryotes with single cells duplicate agamic ally and have huge populace sizes and short age times [6]. Interestingly, physical cells inside individual congregation have gigantic diploid genes that might enable them to develop in excess of smaller times of instance. Another crucial distinction expanding the probability of adjustment by cancer cells lies in the underlying condition of the cell. Living beings have commonly advanced to a versatile wellness top (as exposed in Fig. 1), so that the majority accidental alterations are hindering, particularly loss-of-work alterations. In the unusual setting of choice among physical cells, this places them in a wellness hollow instead of on a top on the versatile landscape. Accidental changes in a cancerous cell can without much of a stretch be gainful, at any rate ahead of schedule in spread of tumor, and can enable the neo-plastic cell to rapidly move out of its wellness pit. For instance, loss-of-work changes that fracture the advanced cell hardware that typically smothers cell multiplication, or underpins apoptosis, would give quick wellness advantage to a physical cell. A similar thinking applies to a large portion of the malignancy "trademark" traits. In expansion to the limited life expectancy of the congregation that is "the earth" of physical cells, there are other significant contrasts flanked by physical cell against organismal advancement, and comprehension these distinctions are able to assist us with bettering respond the topic of whether malignant growth cells are able to advance adjustments by means of common selection [7]. Natural choice among multicellular creatures is commonly a moderate procedure as a result of long age times (contrasted and physical cells) and little populace sizes (contrasted and billions of cells in a developing neoplasm).

Confirmation For Versions In Cancer:

For instance, neoplastic cell populaces may adjust just in the event that they contain enough assorted variety to address the difficulties of metastasis or of treatment. In organismal science, there has for quite some time been discussion over the force and constraints of the "adaptations program" of scanning for Darwinian adjustment as an approach to produce theories furthermore, manage look into (Gould and Lewontin 1979; Mayr 1983), and a portion of this has equivalent in malignancy science (Arnal *et al.* 2015). Does physical advancement manufacture compound oddities, or on the other hand just evacuate or initiate cell works that are now prearranged in the constitutive individual genome? When all is said in done, we have data on the transformations and changes that show up during substantial advancement, however know very minimal about the rhythm and elements of their advancement. Probably the clearest instances of adjustment in malignant growth originate from gained helpful obstruction. Every single cytotoxic treatment and focused on treatments select for medicate obstruction (Pepper 2011, 2012, 2016) [8]. Similarly as with the trademarks, the sub-atomic changes hidden gained opposition regularly vary between tumors, be that as it may, the focalized attribute of obstruction is steady across numerous malignancy types and helpful specialists (e.g., Azam *et al.* 2003; Engelman *et al.* 2007; Murugaesu *et al.* 2015; looked into by Pepper 2016). Developmental adjustment happens in any circumstance in which the conditions for characteristic determination are met (Bell 1997, p. 25). This is valid for nothing existing life forms and for malignant growth cells the same. Characteristic choice outcomes from the blend of (1) characteristic variety in a repeating populace (e.g., of cells), (2) legacy of that variety across (cell) ages, and (3) variety in (cell) wellness related with genetic attribute variety. Each of the three of these adequate circumstances is met in neoplastic cells (Merlo *et al.* 2006). Advancement of adjustment through normal choice follows as a consistent need. This is an exact portrayal of the procedure, not simply a representation. Additional proof for transformative adjustment in disease originates from the prescient worth of cell hereditary assorted variety, or "mosaicism," in malignant growth movement. In the event that there was no cell determination in neoplasms, if all substantial development was unbiased, at that point the measure of decent variety in a cancer would not be prescient of something. It would just be enlightening of the history regular the past of the cancer (Sottoriva *et al.* 2015; Williams *et al.* 2016). In any case, beneath substantial cell choice, decent variety is basic, since it is the premise of

wellness determination (Fernandez *et al.* 2016). Experimentally, higher hereditary decent variety inside neoplasms does in actuality anticipate both quicker movement to danger and diminished patient endurance time (Merlo *et al.* 2006, 2010; Mroz *et al.* 2015; Andor *et al.* 2016). Later sub-atomic proof for malignant growth cell adjustment originates from assessment of dimensional diversity in bosom tumors (Lloyd *et al.* 2016). Developmental hypothesis anticipated that dimensional variety of physical appearance could consequence starting neighborhood varieties in natural features that choose for various physical appearance assets [9]. For model, locales of short blood watercourse, which are usually saw in growth imaging, would choose for cancers that are adjusted to such circumstances as diminished accessibility of surface what's more, blood-determined development factors. Point by point investigation of spatial atomic heterogeneity in 10 clinical bosom malignant growths indicated a steady territorial dissemination meeting this forecast, which upheld the versatile theory (Lloyd *et al.* 2016). Albeit unbiased development can produce decent variety what's more, may create cells with the possible for adjustment, those cells develop into pertinent as it were at the point when they experience clonal extension accordingly to specific weights from the microenvironment.

Convergent Cellular Development Is What Makes "Tumor" A Significant (And Lethal) Disease Kind:

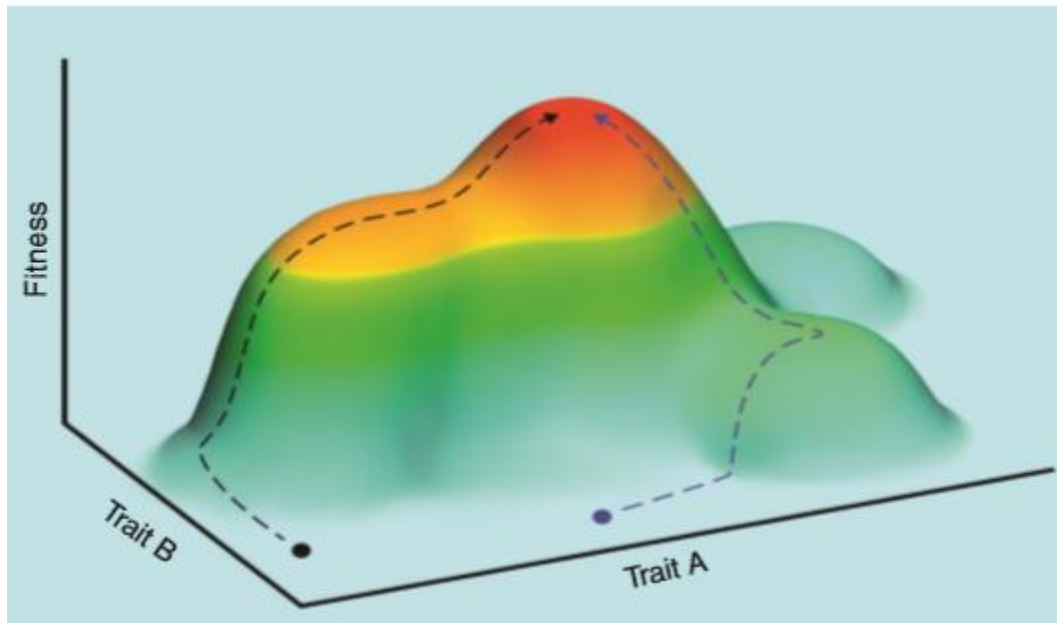
Joined advancement in cavern fish is one great model with perceived parallels to malignancy cell development. Without concurrent advancement, there would be no likenesses among cancers from various tissues, thus present would be no explanation behind us to arrange them jointly as the ailment(s) we describe "malignant growth." truth be told, we see similar phenotypes advancing again and again, in totally various sorts of diseases, in various hosts. We consider these phenotypes the "signs of malignancy". The natural specialty for every one of these malignancies is that of an "endogenous scrounger" (Charlton 1996), which means scrounging cells got commencing typical cells yet now contending with and abusing them [11]. The likeness over the entire malignant growths of this specialty and its specific weights clarifies why all diseases will in general produce comparative cell attributes despite the fact that every malignant growth can be as hereditarily interesting as types of cavern fish. In science, by and large, the absolute most grounded proof for adjustment originates from focalized advancement, in which various populaces in comparable conditions

develop comparable phenotypes because of comparative specific weights.

Figure 2:

Here, we utilize robustness scenery to exemplify convergent development of cancer physical appearance. Two various features have been shown in

x and y axis. Strength have been represented in z axis. Strength enhances straightly. Although cancer does not congregate hereditarily, all characteristically meet on the similar "trademark" physical characteristics. This enhancement can be represented in graph by variation in color from light blue to red.



Substantial transformative adjustment not exclusively is what makes "malignancy" a significant ailment class yet in addition what makes disease a dangerous what's more, lethal malady paying little mind to what tissue it emerges in. In spite of the quality likenesses among these various species, hereditary investigation uncovers that eachspecie advanced these "trademarks of cavern fish" autonomously, from hereditarily diverse establishing populaces, and that despite everything they hold their hereditary uniqueness. Such united quality advancement can just emerge from a common versatile reaction to the equivalent particular pressures, and such a versatile reaction can just be driven by regular choice. In this way, joined development in autonomous populaces is a solid and dependable mark of adjustment through normal choice [12]. Neoplasms that start as generous developments before long exceed their neighborhood assets, following which the main cancerous cells that endure furthermore, multiply are those that can get assets by attacking the encompassing ordinary tissues. This establishes a solid particular weight, with threatening tissue intrusion as the versatile reaction.

Adaptationist thinking pushes the frontiers of cancer research:

The central subject in malignant expansion medication is that, "malignant growth is consistently adjusting", so that at first encouraging new therapeutics invariably flop despite procured helpful opposition. The issue is versatile development, and perceiving this has driven a few clinicians to pull together on "the likelihood to pathway and indulgence development" (Will yard 2016), by pertainingaverage devices of developmental science, for example, phylogenetic reproduction of familial ancestries and transformative causes or scientific models of wellness improvement through transformative adjustment [13]. Instead of simply giving similitudes to pondering the issue, adaptations hypothesis additionally gives scientific models, such as developmental streamlining on wellness scenes, that can convert into new restorative techniques, for example, the versatile treatment regimens that have as of late prevailing in creature models by keeping away from obtained sedate obstruction and there-by dragging out movement free endurance.

Do cancer cells evolve novel traits?

This cell conduct is typical just for white blood cells and is probably empowered through typical individual qualities that are unusually communicated in disease cells. Another sensational case of a compound disease cell conduct that seems narrative in its recycled setting of disease movement is cell barbarianism, or devouring and processing different cells (Lugini *et al.* 2006; Melendez-Lazo *et al.* 2015). The components of anomalous quality articulation behind characteristic repurposing in disease are progressively being uncovered. Late endeavors to describe the hereditary administrative systems in malignancy cells have discovered broad overhauling of those systems, comparative with ordinary cells. In malignancy, a large portion of the point by point systems stay to be depicted, in light of the fact that most malignant growth progressing so distant has solely concentrated on modifications in policy districts, for the straightforward explanation that we don't yet have great ways to recognize administrative locales and foresee the results of transformations in those sections [14]. One of the great applied difficulties to adaptations imagining that has returned in malignant growth science (Arnal *et al.* 2015) is the subject of curiosity, or the differentiation among original adjustments against recycled "expatiations" that initially provided an alternate capacity (Gould and Leonine 1979). For instance, the apparently novel disease phenotypes of quick multiplication, motility, also, tissue attack frequently result from enactment of capacities that are regularly communicated just by early stage undifferentiated organisms and are incorporated with the ordinary individual genes for that specific situation (Brewer *et al.* 2009; Chen *et al.* 2015). Numerous attributes of disease are narrative for the cell types in which the malignant growth emerges yet are regularly communicated in other kinds of cells or at different phases of improvement, from the equivalent ordinary genome. In tumors, qualities and attributes that initially worked in multi-cellularity are regularly "captured," or repurposed for rivalry among cells [15, 16].

RESULTS:

Revision Through Genetic Novelties in Tumor:

The majority ends change that change proto-transforming gene into transforming gene are gain-of-work changes, normally altering a contingent division sign into a consistent proliferative sign. Thus, summit changes can create adjustments by forestalling drugs from authoritative to their objective amino acids. Notwithstanding strange quality articulation, narrative physical appearance can be created by hereditary (or epigenetic) curiosities. Different components direct to narrative hereditary

builds and versatile phenotypes in neoplastic cells. Many epic hereditary builds not in attendance in typical cells have been portrayed in malignant growth, counting changes in position, revisions, and quality synthesis. Such basic curiosities regularly make cancer (Kumar-Sinha *et al.* 2008; An *et al.* 2010; Greuber *et al.* 2013). This establishes another potential wellspring of novel hereditary develops, either changing qualities by means of addition, changing the guideline of close by qualities, or creating quality repetitions. Such a job in physical cell advancement would equivalent the recognized job of retro-position in individual advancement. Level trades of hereditary substance or organs might likewise add to the versatile limit of malignancy cells. There is proof that malignant growth cells can secure novel qualities during combination of malignant growth cells or combination between diseases what's more, typical cells. Furthermore, cancerous cells can fuse pieces of nucleic acid from nearer that have experienced division of cells (Holmgren *et al.* 1999; Bergsmedh *et al.* 2001). Malignant growth cells preserve likewise gain or supplant their mitochondria since ordinary cells (Tan *et al.* 2015). Every one of these wellsprings of hereditary curiosity can bolster the advancement of narrative attributes. In spite of the fact that the dimension of the fundamental gene of individual malignant growth cells is obscure, a correlation among the ordinary individual genome (.20,000 qualities) and a single cell eukaryotic animal variety like yeast (6300 qualities) proposes the plausibility of numerous unnecessary qualities in malignancy cells. Those qualities could establish a broad wellspring of unnecessary qualities to advance new capacities, which might enable substantial development to create oddities more rapidly than in most unicellular development.

Neutral evolution in cancer:

Late work recommends that numerous tumors advance with little proof of characteristic determination after change to harm (Fig. 3) (Sottoriva *et al.* 2015;

Williams *et al.* 2016). This might be on the grounds that choice is loose after the procedure of carcinogenesis is finished and most natural points of confinement on cell endurance and expansion from the typical genome have just been detached. Although we contend here for the focal significance of regular choice and adjustment in malignant growth, choice isn't the main component of development in malignancy. Surely, both unbiased and chosen changes have been accounted for together in a few diseases. proof of concurrent advancement and common choice (considerably after change in a few

cases) has been originate in numerous different diseases (Anderson et al. 2011; Gerlinger et al. 2012, 2014; Kovac et al. 2015; Yates et al. 2015). evidence of focalized advancement and normal determination (much after change in a few cases) has been found in numerous different malignant growths.

CONCLUSION:

Albeit transformative adjustment by substantial cells is not quite the same as organismal adjustment, it is integral to malignant growth science and necessities to be comprehended. Regular determination ought to remain a key device for creating speculations in disease science, and those theories ought to be tried against options including invalid models such as hereditary float.

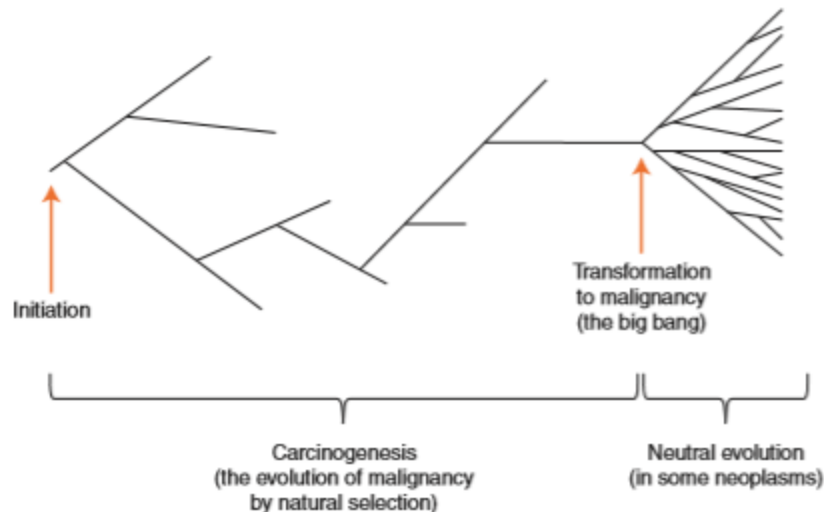


Figure 3:

Phylogenetic tree was constructed for somatic development in cells of neoplasm was explained in the following diagrams. It initiates from the start of the cancer. Somatic mutations hoard during the neoplastic lineage. In the explanation it has been already discussed that many methods are present for development of somatic cells. Cell lineage may develop by normal assortment prior to melanoma.

There is proof for broad transformative adjustment among disease cells, and this procedure drives the concurrent advancement that not just offers significance to the class of "malignant growth," however additionally is the thing that makes disease a threatening and dangerous malady. Notwithstanding this focal job, different systems of development are likewise at work in malignant growth. Unbiased transformative hypothesis gives a lot of invalid models, which have as of late been abused to acceptable impact for dissecting malignant growth development (Sottoriva et al. 2015). In any case, given every one of the signs of tumors that develop during neoplastic movement, what's more, specifically the development of harm what's more, helpful resistance, we accept that adaptations theories are testable and valuable, and furthermore offer an

essential establishment for future restorative advances.

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10. December 1862. *Philosophical Transactions* December, From the Part I. for 1863, having been read at the Royal Society 18.