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RESEARCH ARTICLE

THE FULL 9 STEPPED CYCLE OF PROTON CONDUCTANCE AND ANTISPIRAL - LIKE EVOLUTIONARY BACK STEPS FROM SECOND LATE EVOLUTION TIME EQUATION TO FIRST EARLY EVOLUTION TIME EQUATION DURING SOME PATHOLOGY

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ARTICLE INFO	ABSTRACT
Article History: Received 15 th April, 2017 Received in revised form 11 th May, 2017 Accepted 24 th June, 2017 Published online 31 st July, 2017	 The reaction medium of "Donators + membrane - redox potentials three - state line system + O₂ + ADP + Pi + H⁺ + nH + membrane space = (ATP + heat energy) + H₂O + nH + matrix + CO₂" are the places, where have been occurred the following four more important processes as : Maintaining of clockwise normal flow of electrons and protons with duration of 4-5 second of every cycle Temporarily stop of clockwise normal flow of electrons and protons Complete stop of clockwise normal flow of electrons and protons Antispiral-like evolutionary back steps from second evolution late time equation of flow of
Key words:	
Spiral – like evolution steps, Clockwise normal flow of protons, Temporarily stop of clockwise normal flow.	 According to philosophical principle, all biological process in our planet during last 4 billion years had been developed by the spiral – like evolution steps by transition from simple regulation to more complex regulation as from glycolysis related ATP synthesis, based in the flow of electrons and protons without participation of membrane - redox potentials three - state line system and oxygen to Krebs cycle, proton gradient related ATP synthesis, based in the flow of electrons and protons with out participation of flow of electrons and protons without participation of flow of electrons and protons with oxygen participation using membrane - redox potentials three - state line system. But the notion relating to antispiral-like evolutionary back steps from second late evolution time equation to early evolution time first equation of flow of electrons and protons with provocation of cancer process have been connected with the transition of regulations from more complex regulation to simple regulation based equation as from Krebs cycle dependent, proton gradient based ATP synthesis, based in the flow of electrons and protons with oxygen participation. Using membrane - redox potentials three - state line system functioned owing to the flow of electrons and protons with oxygen participation, using membrane - redox potentials three - state line system functioned owing to the flow of electrons and protons with oxygen participation, using membrane - redox potentials three - state line system to early form of glycolysis related ATP synthesis, based in the flow of electrons and protons with oxygen. Temporarily stop of clockwise normal flow of electrons and protons with duration of 4 - 5 second of every cycle means that under influence of many pathological factors have been caused the partial blockage of electron and proton conductance within the full 9 stepped cycle of proton conductance inside human body proposed by M.Ambaga, which have included such well known metabolic pathways as glycolysis, Krebs cycle,

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INTRODUCTION

Untill now the recent findings of literature could not give the appropriate answer to such principally important questions as what bioevolution link have been existed between the disturbance of normal flow of electrons and protons and cell death processes leading to pathological cirrhotic and cancer irreversible processes.

*Corresponding author: Ambaga, M. New Medicine Medical University, Ulanbator, Mongolia. We are proposing that the reaction medium of "Donators + membrane - redox potentials three - state line system + O_2 + ADP + Pi + H⁺ + nH + membrane space = (ATP + heat energy) + H₂O + nH + matrix + CO₂" is the places, where have been existed the places of conducting of following four more important processes as maintaining of clockwise normal flow of electrons and protons with duration of 4-5 second of every cycle, also the process of temporarily stop of flow of electrons and protons and the process of complete stop of clockwise normal flow of electrons and protons, and the process of antispiral - like evolutionary back steps from second late evolution time equation of flow of electrons and protons to

early evolution time first equation with provocation of cancer process. It would be interesting if can establish the relationship between the temporarily and complete stop of clockwise normal flow of electrons and protons with duration of 4-5 second of every cycle and cell death related cirrhotic processes leading to antispiral-like evolutionary back steps from second late evolution equation of flow of electrons and protons to early evolution first equation. It can be say that the process of complete stop of clockwise normal flow of electrons and protons and the provocation of the antispiral-like evolutionary back steps from second late evolution equation to early evolution first equation with provocation of cancer process have been occured within "Donators + membrane - redox potentials three - state line system + O_2 + ADP + Pi + H⁺ + nH + membrane space = $(ATP + heat energy) + H_2O + nH +$ matrix $+ CO_2$ " metabolic reaction, where may be occurred the two processes as cell death and cirrhosis related cancer processes.

RESULS AND CONCLUSION

The reaction mediums of "Donators + membrane - redox potentials three - state line system + O_2 + ADP + Pi + H⁺ + nH + membrane space = (ATP + heat energy) + H₂O + nH + matrix + CO₂" are the places, where have been occurred the following four more important processes as:

- Maintaining of clockwise normal flow of electrons and protons with duration of 4-5 second of every cycle
- Temporarily stop of clockwise normal flow of electrons and protons
- Complete stop of clockwise normal flow of electrons and protons
- Antispiral like evolutionary back steps from second evolution late time equation of flow of electrons and protons to early evolution first time equation with provocation of pathological process as cancer.

According to philosophical principle, all biological processes in our planet during last 4 billion years had been developed by the spiral – like evolution steps by transition from simple regulation to more complex regulation as from glycolysis related ATP synthesis, based in the flow of electrons and protons without participation of membrane - redox potentials three - state line system and oxygen to Krebs cycle, proton gradient related ATP synthesis, based in the flow of electrons and protons with oxygen participation using membrane - redox potentials three - state line system. But the notion relating to antispiral-like evolutionary back steps from second late evolution time equation to early evolution time first equation of flow of electrons and protons with provocation of cancer process have been connected with the transition of regulations from more complex regulation (Figure 2) to simple regulation based equation as from Krebs cycle dependent, proton gradient based ATP synthesis functioned owing to the flow of electrons and protons with oxygen participation, using membrane redox potentials three - state line system to early form of glycolysis related ATP synthesis based (Figure 1) in the flow of electrons and protons without participation of membrane redox potentials three - state line system and oxygens. The essence of our new idea in relating to provocation of cancer cell growth by antispiral - like evolutionary back steps of electrons and protons flow in connection with transition of second late evolution time equation of flow to early evolution time first equation with provocation of cancer process have been associated with a prevailing of glycolysis process and the provocating of change of proton, electron conductance in the membrane - redox potentials three - state line system dependent - full 9 stepped cycle with disturbation of mitochondria dependent oxygen utilization is follows:

Change of normal relationship between glycolysis and process of the proton, electron conductance in the 2-th stage of the full 9 stepped cycle of proton conductance inside human body in the level of isocitrate dehydrogenase, alpha ketoglutarate dehydrogenase and succinate dehydrogenase within Krebs cycle.

Disturbance of normal relationship between glycolysis and process of the formation of oxaloacetate from malate under action of malate dehydrogenase in the 2-th stage of the full 9 stepped cycle of proton conductance inside human body in the level of isocitrate dehydrogenase, alpha ketoglutarate dehydrogenase and succinate dehydrogenase.

In such way we proposed that the biochemical characteristics of cancer cells more similar to Archea (prokarvotic) as if cancer cells acquired the some feature of Archea, remembering that normal cells made the evolutionary back steps to direction of the early evolution cells (as if Eukarya cells turn to Archaea- prokaryotic cells) with whom could be meet 4,4 billion years ago. The Warburg effect is the observation that most cancer cells predominantly produce energy by a high rate of glycolysis. Otto Warburg postulated this change in metabolism as glycolysis, is the fundamental cause of cancer. Temporarily stop of clockwise normal flow of electrons and protons with duration of 4-5 second of every cycle means that the influence of many pathological factors have been caused the partial blockage of electron and proton conductance within the full 9 stepped cycle of proton conductance inside human body proposed by M.Ambaga, which have included such well known metabolic pathways as glycolysis, Krebs cycle, betta oxidation of fatty acids, aminoacid oxidation.

Maintaining of clockwise normal flow of electrons and protons with duration of 4-5 second of every cycle means that under normal conditions have been created the normal level of releasing of proton, electron from food substrates (carbohydrate, aminoacids, fatty acids), also the normal precondition of transferring of proton, electron to NADH as hydrogen atom and formation of CO2 in Krebs cycle, the normal condition of transferring of proton, electron to KoQ as hydrogen atom, the normal condition of transferring of electron to cytochrom C without accompanying proton, the normal tendency of translocating of proton to intermembrane space of mitochondria without accompanying electron, the normal level of creating of proton gradient in the intermembrane space of mitochondria and following transfer of proton to matrix through ATP synthase and the normal tendency of forming of metabolic water in the mitochondria matrix by oxidation of proton by molecular oxygens i.e, by protonation of molecular oxygen by matrix proton, the normal precondition of diffusing of proton, diffusion of metabolic water through plasma membrane of red blood cells with participation of aquaporin protein channels and entry of oxygen from lung to erythrocyte, entry of carbon dioxide from cells to erythrocyte, the normal parameters of forming of free protons from metabolic water again by reaction as $H_2CO_3 = H$ + HCO₃ (H₂CO₃ formed from metabolic water), proton

combine with hemoglobin (generation of HbH) which promotes the release of oxygen from hemoglobin, oxygen diffusion to cells and proton released from hemoglobin promotes uptake of oxygen by hemoglobin and carbonic anhydrase catalyzes the formation of CO_2 from H_2CO_3 and CO_2 diffuse out in the alveoli are existed between donators and acceptors in general equation of cell metabolism as $C_6H_{12}O_6 +$ $6O_2 = energy + 6H_2O + 6CO_2$ under constant influence of membrane - redox potentials three - state line system. Complete stop of clockwise normal flow of electrons and protons with duration of 4-5 second of every cycle have been connected with this process as the influence of many pathological factors have been caused the irreversible stop electron and proton conductance at first :in the stage of release of proton, electron from food substrates (carbohydrate, aminoacids, fatty acids), at second :in the stage of transferring of proton, electron to NADH as hydrogen atom and formation of CO_2 in Krebs cycle, at third : in the stage of transferring of



Figure 1. First early evolution time equation of flow of electrons and protons



Figure 2. Second late evolution time equation of flow of electrons and protons

proton, electron to KoQ as hydrogen atom, at fourth : in the stage of ransferring of electron to cytochrom C without accompanying proton, at fifth : in the stage translocating of proton to intermembrane space of mitochondria without accompanying electron, at sixth : in the stage of creating of proton gradient in the intermembrane space of mitochondria and following transfer of proton to matrix through ATP synthase, at seventh : in the stage of forming of metabolic water in the mitochondria matrix by oxidation of proton by molecular oxygens i.e, by protonation of molecular oxygen by matrix proton, at eighth : in the stage of diffusing of proton, diffusion of metabolic water through plasma membrane of red blood cells with participation of aquaporin protein channels and entry of oxygen from lung to erythrocyte ,entry of carbon dioxide from cells to erythrocyte, also at ninth : in the stage of forming of free protons from metabolic water again by reaction as $H_2CO_3 = H + HCO_3$ (H_2CO_3 formed from metabolic water), proton combine with hemoglobin (generation of HbH) which promotes the release of oxygen from hemoglobin, oxygen diffusion to cells and proton released from hemoglobin promotes uptake of oxygen by hemoglobin and carbonic anhydrase catalyzes the formation of CO₂ from H₂CO₃ and CO₂ diffuse out in the alveoli are existed between donators and acceptors in general equation of cell metabolism as $C_6H_{12}O_6 +$ $6O_2 = energy + 6H_2O + 6CO_2$ under constant influence of membrane-redox potentials three-state line system.

Complete stop of clockwise normal flow of electrons and protons with duration of 4-5 second of every cycle within the full 9 stepped cycle of proton conductance inside human body proposed by M.Ambaga, A.Tumen-Ulzii have been preceded by damage of membrane structures, which belong to membrane - redox potentials three - state line system of "Donators + membrane - redox potentials three - state line system $+ O_2 + ADP + Pi + H^+ + nH + membrane space = (ATP)$ + heat energy) + H_2O + nH + matrix + CO_2 " metabolic reaction followed by leakage of enzyme-substrate complexes to outside cell resulting to decrease of aminoacid, fatty acid, glucose molecules oxidation and synthesis. In such way, complete stop of clockwise normal flow of electrons and protons with duration of 4-5 second of every cycle have been resulted to the cell death followed by pathological cirrhotic and cancer irreversible processes.

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