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PHYLOGENETICAL RELATIONSHIPS BASED IN THE 9 STAGED CLOSED CYCLE OF PROTON CONDUCTANCE AND DRUG TARGETS RELATING TO SOME MEDICAMENTS OF TRADITIONAL MEDICINE

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ABSTRACT

According to theory of Traditional Medicine living rLung, Badgan, Mkhriis are existed inside human body as gene, RNA, DNA, protons, electrons by ensuring all life processes as breath, blood circulation, food digestion, ageing, body growth. In this connection, according to theory of Traditional Medicine action of all medication methods of Traditional Medicine, including medicaments, acupuncture, moxibustion have been appeared in the level of living rLung, Badgan, Mkhriis, which are existed inside human body together with genes, RNA, DNA, protons, electrons, "Donators + membrane - redox potentials three - state line system + O₂ + ADP + Pi + H⁺ + nH⁺_{membrane space} = (ATP + heat energy) + H₂O + nH⁺_{matrix} + CO₂" reaction medium and CxHyOz - (X + Y/4 + Z/2) O₂ + xCO₂ + Y/2 H₂O reaction medium. According to modern medicine the action of drugs on the human body is called pharmacodynamics, and what the body does with the drug is called pharmacokinetics. The drugs that enter the human tend to stimulate certain receptors, ion channels, act on enzymes or transporter proteins. One is form of universal common descent as "Donators + membrane-redox potentials three-state line system + O₂ + ADP + Pi + H⁺ + nH⁺_{membrane space} = (ATP + heat energy) + H₂O + nH⁺_{matrix} + CO₂" reaction medium have been served the role of drug targets, relating to all drugs not depending drugs either classified as Modern medicine drug, or as drug of Traditional medicine. Also, one is form of universal common descent as CxHyOz-(X+Y/4+Z/2) O₂ + xCO₂ + Y/2 H₂O reaction medium have been served the role of drug targets, relating to all drugs not depending drugs either classified as Modern medicine drug, or as drug of Traditional medicine. We are considered that when drugs not depending drugs either classified as modern medicine drug, or as drug of Traditional medicine enter the human organism, reaching a drug targets as "Donators + membrane-redox potentials three - state line system + O₂ + ADP + Pi + H⁺ + nH⁺_{membrane space} = (ATP + heat energy) + H₂O + nH⁺_{matrix} + CO₂" reaction medium would exert pharmacological effects. We are proposed that when drugs not depending drugs either classified as modern medicine drug, or as drug of Traditional medicine enter the human organism, reaching a drug targets as CxHyOz - (X + Y/4 + Z/2) O₂ + xCO₂ + Y/2 H₂O reaction medium would exert pharmacological effects. We came to conclusion that when drugs not depending drugs either classified as modern medicine drug, or as drug of Traditional medicine enter the human organism, reaching a concrete drug targets as a molecular structure - chemically definable by a molecular mass, that will undergo a specific interaction with chemicals - drugs, tend to stimulate certain receptors, which are synthesized with participation of all genes by rule of genetic code identical for all all known life forms, descended from 16s rRNA gene, 18S rRNA, also descended from set of 355, comprising the three domains of life, archaea, bacteria, and eukaryotes, 6,331 genes common to all living animals have been identified. By us established that when drugs not depending drugs either classified as modern medicine drug, or as drug of Traditional medicine enter the human organism tend to stimulate certain proteins, descended from some 23 proteins, descended from a single common ancestor that lived 650 million years ago in the Precambrian, which are synthesized with participation of all genes by genetic code identical for all known life forms, descended from 16s rRNA gene, 18S rRNA, comprising the three domains of life, archaea, bacteria, and eukaryotes. It was clear that when drugs not depending drugs either classified as modern medicine drug, or as drug of Traditional medicine enter the human organism would cause the specific actions as affinity - measure of how tightly a drug binds to the receptors, which are synthesized with participation of all genes by general principle of genetic code identical for all all known life forms, comprising the three domains of life, archaea. We are denying the possibility of parallel existence of living rLung, Badgan, Mkhriis (LBM), serving the role of Drug targets, relating to some medicaments of Traditional Medicinetogether with specific targets of drug actions as ion channels, receptors, all genes, descended from 16s rRNA gene, 18S rRNA, a set of 355 genes, comprising the three domains of life, archaea, bacteria, and eukaryotes, which are synthesized from pyrimidine and purine nucleotides, ATP molecules, generated within membrane - redox potentials three - state line system dependent - full 9 stepped cycle of proton conductance. It can be say that the continuity of proton conduction depended biosynthesis of ATP and NADPH using the membrane - redox potentials three - state line system like structures from the cyanobacteria to Homosapiens during last 4,5 billion years had been carried out by general principle as common descent of all life on Earth from the last universal common ancestor, comprising the three domains of life, archaea, bacteria, and eukaryotes, a genetic code identical for all known life forms, such postulation eventually gives the scientific ground to say that it is impossible of existing of the living rLung, Badgan, Mkhriis, serving the role of Drug targets relating to some medicaments of Traditional Medicine. The universality of this code is generally regarded by biologists as definitive evidence in favor of universal common descent owing to membrane - redox potentials three - state line system like structures from the cyanobacteria to Homosapiens during last 4,5 billion years, such explanation eventually denied the possibility of existing of living rLung, Badgan, Mkhriis, serving the role of Drug targets relating to some medicaments of Traditional Medicine. Theory of rLung, Badgan, Mkhriis of Traditional Medicine (analogy and correspondence with theory of four humors) is abstract terms, reflecting the living things, existed inside human body as 16s rRNA gene, 18S rRNA, RNA, DNA, protons and electrons, the membrane - redox potentials three - state line system dependent - full 9 stepped cycle of proton conductance, a set of 355 genes, comprising the three domains of life, archaea, bacteria, and eukaryotes, and also genetic code, identical for all known life forms (from bacteria and archaea to animals and plants), ensuring all life processes as breath, blood circulation, food digestion, ageing, body growth as specific targets of drug action, such interpretation have been witnessed the impossibility of existing of living rLung, Badgan, Mkhriis as drug targets.

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INTRODUCTION

A drug target is a molecule in the body, usually a protein, that is intrinsically associated with a particular disease process and that could be addressed by a drug to produce a desired therapeutic effect. The increasing knowledge of molecular signaling processes has enabled the identification of drug targets. Drug target validation involves proving that either DNA, RNA, or a protein molecule is directly involved in a disease process and can be a suitable target for development of a new therapeutic drug. To qualify as 'druggable', a target must be accessible to the proposed drug molecule, and a measurable biological reaction must be provoked as a consequence of the drug interacting with the target. Once the target has been identified, for example by genomic or proteomic investigations, the target is then validated in functional studies. The main strategies involved in target validation are gene knockout studies and direct inhibition of the target by small molecules, peptides, antibodies or any other class of inhibitors. Universal common descent through an evolutionary process was first proposed by the British naturalist Charles Darwin in the concluding sentence of his 1859 book *On the Origin of Species*, which are given the scientific ground to say about the impossibility of existence of living rLung, Badgan, Mkhri as drug targets. The phlegm of humorism is far from the same thing as phlegm as it is defined today. Nobel laureate Charles Richet MD, when describing humorism's "phlegm" in 1910 asked rhetorically, "...this strange liquid-four humors, which is the cause of tumours, of chlorosis, of rheumatism, and cacochymia - Where is it? Who will ever see four humors? Who has ever seen four humors? What can we say of this fanciful classification of humours into four groups, of which two are absolutely imaginary?, such explanations would lead to conclusion about the impossibility of existence of living rLung, Badgan, Mkhri as drug targets. Many organisms all derived from a single ancestor could readily have shared genes that all worked in the same way, and it appears that they have, which have been witnessed the impossibility of existence of living rLung, Badgan, Mkhri as drug targets. The most common gene to be used for constructing phylogenetic relationships in prokaryotes is the 16S ribosomal RNA gene since its sequences tend to be conserved among members with close phylogenetic distances, such interpretations would serve the scientific basis to say the impossibility of existence of living rLung, Badgan, Mkhri as drug targets. Plant-fungus horizontal gene transfer is the movement of genetic material between individuals in the plant and fungus kingdoms, such postulation gives the scientific ground to say that it is impossible existing of living four humors and living rLung, Badgan, Mkhri with 16s rRNA gene, 18S rRNA, a set of 355 genes, comprising the three domains of life, archaea, bacteria, and eukaryotes, a genetic code, identical for all known life forms, from bacteria and archaea to animals and plants, such postulation gives the scientific ground to say that it is impossible existing of living four humors and living rLung, Badgan, Mkhri with 16s rRNA gene, 18S rRNA, a set of 355 genes, comprising the three domains of life, archaea, bacteria, and eukaryotes, a genetic code, identical for all known life forms, from bacteria and archaea to animals and plants, which have been witnessed the impossibility of existence of living rLung, Badgan, Mkhri as drug targets.

RESULTS AND DISCUSSION

According to theory of Traditional Medicine living rLung, Badgan, Mkhri are existed inside human body as gene, RNA, DNA, protons, electrons by ensuring all life processes as breath, blood circulation, food digestion, ageing, body growth. In this connection, according to theory of Traditional Medicine action of all medication methods of Traditional Medicine, including medicaments, acupuncture, moxibustion have been appeared in the level of living rLung, Badgan, Mkhri, which are existed inside human body together with genes, RNA, DNA, protons, electrons, "Donators + membrane-redox potentials three-state line system + O₂ + ADP + Pi + H⁺ + nH⁺_{membrane space} = (ATP + heat energy) + H₂O + nH⁺_{matrix} + CO₂" reaction medium and C_xH_yO_z-(X+Y/4+Z/2) O₂ + xCO₂ + Y/2 H₂O reaction medium. According to modern medicine the action of drugs on the

human body is called pharmacodynamics, and what the body does with the drug is called pharmacokinetics. The drugs that enter the human tend to stimulate certain receptors, ion channels, act on enzymes or transporter proteins. A drug target is a molecular structure -chemically definable by a molecular mass, that will undergo a specific interaction with chemicals (drugs) and increasing knowledge of molecular signaling processes has enabled the identification of drug targets. One is form of universal common descent as "Donators + membrane - redox potentials three - state line system + O₂ + ADP + Pi + H⁺ + nH⁺_{membrane space} = (ATP + heat energy) + H₂O + nH⁺_{matrix} + CO₂" reaction medium have been served the role of drug targets relating to all drugs not depending drugs either classified as Modern medicine drug, or as drug of Traditional medicine. Also, one is form of universal common descent as C_xH_yO_z-(X + Y/4 + Z/2) O₂ + xCO₂ + Y/2 H₂O reaction medium have been served the role of drug targets relating to all drugs not depending drugs either classified as Modern medicine drug, or as drug of Traditional medicine.

It can be say that when drugs not depending drugs either classified as modern medicine drug, or as drug of Traditional medicine enter the human organism, reaching a concrete drug targets as a molecular structure - chemically definable by a molecular mass, that will undergo a specific interaction with chemicals-drugs, tend to stimulate certain receptors, which are synthesized with participation of all genes by rule of genetic code identical for all all known life forms, descended from 16s rRNA gene, 18S rRNA, also descended from set of 355, comprising the three domains of life, archaea, bacteria, and eukaryotes, 6,331 genes common to all living animals have been identified. We are considered that when drugs not depending drugs either classified as modern medicine drug, or as drug of Traditional medicine enter the human organism, reaching a drug targets as "Donators + membrane - redox potentials three-state line system + O₂ + ADP + Pi + H⁺ + nH⁺_{membrane space} = (ATP + heat energy) + H₂O + nH⁺_{matrix} + CO₂" reaction medium would exert pharmacological effects. It was clear that when drugs not depending drugs either classified as modern medicine drug, or as drug of Traditional medicine enter the human organism, reaching a drug targets as C_xH_yO_z - (X+Y/4+Z/2) O₂ + xCO₂ + Y/2 H₂O reaction medium would exert pharmacological effects. We have established that when drugs not depending drugs either classified as modern medicine drug, or as drug of Traditional medicine enter the human organism tend to stimulate ion channels, act on enzymes or transporter proteins, which are synthesized with participation of all genes by general rule of genetic code identical for all all known life forms, descended from 16s rRNA gene, 18S rRNA, also descended from 6,331 genes common to all living animals have been identified. It should be say that when drugs not depending drugs either classified as modern medicine drug, or as drug of Traditional medicine enter the human organism tend to stimulate certain proteins, descended from some 23 proteins, descended from a single common ancestor that lived 650 million years ago in the Precambrian, which are synthesized with participation of all genes by genetic code identical for all known life forms, descended from 16s rRNA gene, 18S rRNA, comprising the three domains of life, archaea, bacteria, and eukaryotes. By us established that when drugs not depending drugs either classified as modern medicine drug, or as drug of Traditional medicine enter the human organism would cause the specific actions as affinity - measure of how tightly a drug binds to the receptors, which are synthesized with participation of all genes by general principle of genetic code identical for all all known life forms, comprising the three domains of life, archaea. If we agree with this that living rLung, Badgan, Mkhri (LBM) is really existed inside human body, which can see and touch, as gene, RNA, DNA, protons and electrons, ensuring all life processes, serving the role of Drug targets relating to all medicaments of Traditional Medicine we should package LBM and use this as biopreparations in every day clinical practice instead of Modern medicine medicaments. In this connection it is arised the principal important questions relating to living rLung, Badgan, Mkhri (LBM), serving the role of Drug targets, together with classic drug targets as genes, descended from 355 genes, comprising the three domains of life, archaea, bacteria, and eukaryotes, which are synthesized from pyrimidine and purine nucleotides, ATP molecules,



Figure 1 .The membrane - redox potentials three - state line system dependent – full 9 stepped cycle of proton conductance

generated within membrane - redox potentials three - state line system dependent -full 9 stepped cycle of proton conductance. We are denying the possibility of parallel existence of living rLung, Badgan, Mkhri (LBM), serving the role of Drug targets, relating to some medicaments of Traditional Medicine together with specific targets of drug actions as ion channels, receptors, all genes, descended from 16s rRNA gene, 18S rRNA, a set of 355 genes, comprising the three domains of life, archaea, bacteria, and eukaryotes, which are synthesized from pyrimidine and purine nucleotides, ATP molecules, generated within membrane - redox potentials three - state line system dependent - full 9 stepped cycle of proton conductance. We are putting a following questions as at first: where is LBM serving the role of Drug targets relating to some medicaments of Traditional Medicine?, who will ever see rLung, Badgan, Mkhri, serving the role of Drug targets relating to some medicaments of Traditional Medicine? Who has ever seen LBM serving the role of Drug targets relating to some medicaments of Traditional Medicine, now when scientific world confirmed that specific targets of drug actions are ion channels, receptors, all genes, descended from 16s rRNA gene, 18S rRNA, a set of 355 genes, comprising the three domains of life, archaea, bacteria, and eukaryotes, at second: when and where were rLung, Badgan, Mkhri serving the role of Drug targets relating to some medicaments of Traditional Medicine, now when scientific world confirmed that all living cells have arisen from a single common ancestor that lived 650 million years ago in the Precambrian,

which are specific targets of drug action, at third: when and where were rLung, Badgan, Mkhri, when now scientific world confirmed that genetic code, identical for all known life forms, from bacteria and archaea to animals and plants, which have been served the role of specific targets of drug action. All living organisms have the same kinds of monomeric subunits and the identity of each organism is preserved by its possession of many sets of nucleic acids and of proteins, which formed with participation of ATP/ADP cycle functioned as one of members of membrane - redox potentials three - state line system dependent - full 9 stepped cycle of proton conductance, such explanation eventually denied the possibility of existing of rLung, Badgan, Mkhri, serving the role of Drug targets relating to some medicaments of Traditional Medicine together with genes, RNA, DNA, protons and electrons during evolution development of life from cyanobacteria to Homo sapiens human, which are specific targets of drug action. It can be say that the continuity of proton conduction depended biosynthesis of ATP and NADPH using the membrane - redox potentials three - state line system like structures from the cyanobacteria to Homosapiens during last 4,5 billion years had been carried out by general principle as common descent of all life on Earth from the last universal common ancestor, comprising the three domains of life, archaea, bacteria, and eukaryotes, a genetic code identical for all known life forms, such postulation eventually gives the scientific ground to say that it is impossible of existing of the living rLung, Badgan, Mkhri, serving

the role of Drug targets relating to some medicaments of Traditional Medicine. The universality of this code is generally regarded by biologists as definitive evidence in favor of universal common descent, owing to membrane - redox potentials three - state line system like structures from the cyanobacteria to Homosapiens during last 4,5 billion years, such explanation eventually denied the possibility of existing of rLung, Badgan, Mkhriis, serving the role of Drug targets relating to some medicaments of Traditional Medicine. In such way, the membrane - redox potentials three - state line system dependent - full 9 stepped cycle of proton conductance have been formed as final results of phylogenetic relationship of life during last 4,5 billion years, such postulation eventually gives the scientific ground to say that it is impossible of existing of the living rLung, Badgan, Mkhriis, serving the role of Drug targets relating to some medicaments of Traditional Medicine, together with ion channels, receptors, which are specific targets of drug action. The participation of evolutionary late electron, proton transporting systems as "Donators + membrane - redox potentials three - state line system + $O_2 + ADP + Pi + H^+ + nH^+_{\text{membrane space}} = (ATP + \text{heat energy}) + H_2O + nH^+_{\text{matrix}} + CO_2$ " in the biosynthesis of purine base as molecules of universal common descent have been appeared as the first stage: ribose-5 phosphate + ATP = 5-phosphoribosyl - alpha - pyrophosphate (PRPP), the second stage: PRPP + glutamine + $H_2O = \text{beta} - 5\text{-phosphoribosylamine}$, the third stage: beta - 5-phosphoribosylamine + ATP + glycine = glycinamidribotide (GAR), the fourth stage: GAR + N10-formyl-TNF = formylglycinamidribotide (FGAR), the fifth stage: ATP + glutamine + FGAR = Formylglycinamidribotide (FGAM), at the sixth stage: FGAM + ATP = 5-aminoimidazole ribotide (AIR), the seventh stage: $CO_2 + \text{AIR} = \text{carboxyaminoimidazole ribotide (CAIR)}$, the eighth stage: CAIR + aspartate + ATP = 5 - aminoimidazole - 4-(succinylcarboxyamide) ribotide (SACAIR), the ninth stage: SACAIR = fumarate + 5-aminoimidazole - 4-carboxamideribotide (AICAR), the tenth stage: AICAR + N10-formyl-TNF = 5-formaminoimidazole - 4-carboxamideribotide (FAICAR), the eleventh stage: FAICAR = $H_2O + \text{inosine monophosphate (IMP)}$ by principle as "nucleic acids are needed to make proteins, yet proteins are needed to make nucleic acids" in the all widely divergent organisms as E.coli, yeast, pigeons, and humans, such explanations have been denied the existence of drug targets as living rLung, Badgan, Mkhriis inside human body together with as specific targets of drug action as genes, ion channels, receptors, membrane - redox potentials three - state line, formed during evolution development of life from cyanobacteria to Homo sapiens human.

The biosynthesis of purine base as molecules of universal common descent have been strongly needed the participation of evolutionary late electron, proton transporting systems as "Donators + membrane - redox potentials three - state line system + $O_2 + ADP + Pi + H^+ + nH^+_{\text{membrane space}} = (ATP + \text{heat energy}) + H_2O + nH^+_{\text{matrix}} + CO_2$ " and also the participation of membrane - redox potentials three - state line system, where formed such very important macroerg compounds as ATP, ADP in the all widely divergent organisms as E.coli, yeast, pigeons, and humans, such explanations have been denied the existence of drug targets as living rLung, Badgan, Mkhriis inside human body together with specific targets of drug action as genes, ion channels, receptors, membrane - redox potentials three - state line. The participation of evolutionary late electron, proton transporting systems as "Donators + membrane - redox potentials three - state line system + $O_2 + ADP + Pi + H^+ + nH^+_{\text{membrane space}} = (ATP + \text{heat energy}) + H_2O + nH^+_{\text{matrix}} + CO_2$ " in the biosynthesis of pyrimidine ribonucleotides as universal common descent molecules have been appeared as ATP + $HCO_3 + \text{glutamine} + H_2O = \text{carbamoyl phosphate}$, carbomoylphosphate + aspartate = carbamoyl aspartate, carbamoyl aspartate = $H_2O + \text{dihydroorotate}$, dihydroorotate + quinone = orotate, orotate + PRPP = orotidine monophosphate (OMP), OMP = $CO_2 + \text{uridine monophosphate (UMP)}$, such postulation eventually gives the scientific ground to say that it is impossible exist the living rLung, Badgan, Mkhriis with 16s rRNA gene, 18S rRNA, a set of 355 genes, comprising the three domains of life, archaea, bacteria, and eukaryotes during evolution development of life from cyanobacteria to Homo sapiens human, such postulation gives the scientific ground

to say that it is impossible existing a drug targets as living rLung, Badgan, Mkhriis inside human body together with specific targets of drug action as genes, ion channels, receptors, membrane - redox potentials three - state line system. The participation of evolutionary late electron, proton transporting systems as "Donators + membrane - redox potentials three - state line system + $O_2 + ADP + Pi + H^+ + nH^+_{\text{membrane space}} = (ATP + \text{heat energy}) + H_2O + nH^+_{\text{matrix}} + CO_2$ " in the biosynthesis of deoxyribonucleotides as very important molecules as universal common descent have been appeared as in first stage: NADPH+FAD= FADH₂, at second stage: FADH₂ + S = S thoredoxin = SH-SH thoredoxin + NADP, at third stage: SH-SH thoredoxin+ S=S ribonucleotides= SH-SH ribonucleotides + S=S thoredoxin, at fourth stage: SH-SH ribonucleotides+NDP= dNDP, at fifth stage: dNDP + ATP= dNTP, such explanations have been denied the existence of drug targets as living rLung, Badgan, Mkhriis inside human body together with specific targets of drug action as genes, ion channels, receptors, membrane - redox potentials three - state line system. Theory of rLung, Badgan, Mkhriis of Traditional Medicine (analogy and correspondence with theory of four humors) is abstract terms, reflecting the living things, existed inside human body as 16s rRNA gene, 18S rRNA, RNA, DNA, protons and electrons, the membrane - redox potentials three - state line system dependent - full 9 stepped cycle of proton conductance, a set of 355 genes, comprising the three domains of life, archaea, bacteria, and eukaryotes, and also genetic code, identical for all known life forms (from bacteria and archaea to animals and plants), ensuring all life processes as breath, blood circulation, food digestion, ageing, body growth as specific targets of drug action, such interpretation have been witnessed the impossibility of existing of living rLung, Badgan, Mkhriis as drug targets.

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