GERALDA: A Computational Database to Study Cancer Risk in "Admixed" Individuals

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banco

de dados

usuários

Ask questions and define problems: the white skin color

Noah, from the Bible is, according to the Bible, the most recent common human ancestor in contrast to Adam and
probably the most famous white or albino person with extra-biblical Jewish text evidence of white skin color or being an
albino

• The National Organization for Albinism and Hypopigmentation (NOAH) is probably named after the biblical Noah in reference to his skin hypopigmentation

Ask questions and define problems: the white skin color and races

Human races do not exist from the standpoint of biology

How this concept was constructed and how it impacts our world today?

Although physical differences exist, we share a common and equal human condition

DNA (the genetic code) encodes and explains (all?) these differences

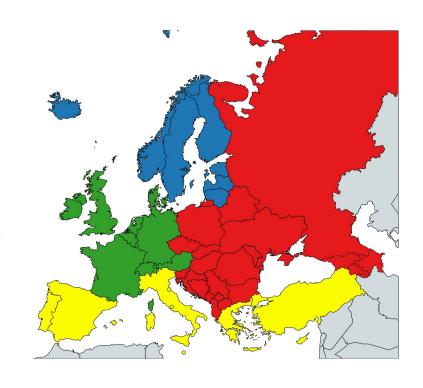
Ask questions and define problems: the white skin color

Eurocentrism, but different "white" races

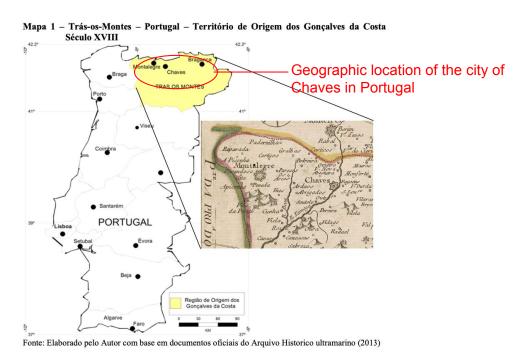
 Southern Europeans are physically different compared to Northern Europeans

European "white" miscegenation in the USA



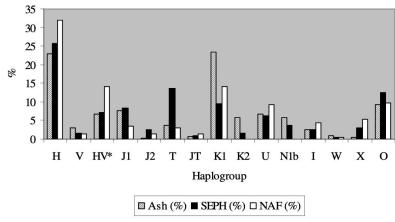


Ask questions and define problems: our names and their continental (geographic) origin



Ask questions and define problems: the white skin color in Southern Europe

- Catholic Iberians (Portugal and Spain) have a history of persecution of Jews (Sephardic Jews) and Muslims
- The word for Spain in Hebrew is Sepharad (Mediterranean European Jewish human type = Sephardic Jew)
- Racial miscegenation in Portugal/Brazil can be traced back to expulsion of Muslims from Iberia and *limpeza* or *limpieza de sangre*
 - Formation of the National State of Portugal (Galiza and Northern Portugal)
 - Conversion of cristãos novos (both Muslims and Jews)
 - Portuguese and Spanish blood allegiance practices in Europe and in America



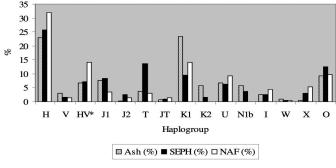
Mt Haplogroup distribution in the three Jewish populations (Feder 2008)

 Haplogroup H is the most common mitochondrial haplogroup in Sephardic Jews and Europe(?)

Ask questions and define problems: the white skin color in Southern Europe



- Catholic Iberians (Portugal and Spain) have a history of persecution of Jews (Sephardic Jews) and Muslims because of the Muslim invasion
 - Spain in Hebrew = Sepharad
- Racial miscegenation in Brazil (and Latin America) can be traced back to limpeza or limpieza de sangre
 - Blood allegiance in Europe and in America
 - Conversion of Muslims and Jews as cristãos novos
- Iberia became imprinted (or vice-versa) with the DNA of Sephardic Jews as haplogroup H is the most common mitochondrial haplogroup in Sephardic Jews and Europe(?)



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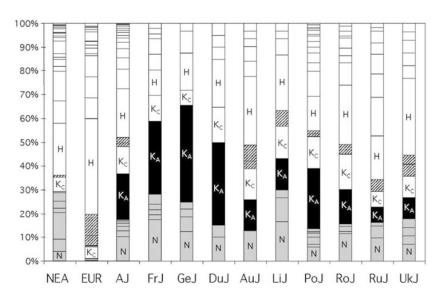
Mt Haplogroup distribution in the three Jewish populations (Feder 2008)

Ask questions and define problems: the white skin color in Northern Europe





- Sophie Charlotte de Mecklenburg-Strelitz (1744-1818), first queen of England descendant of Africans
- Northern Europe, home of the Martin Luther's Protestant Reformation experienced tolerance of Jews in contrast to Southern European Catholic Christians
- The word for Germany in medieval Hebrew was Ashkenaz
- Jews from Eastern and Central Europe refer to themselves as Ashkenazi Jews (Northern European Jewish human type)



MtDNA haplogroup frequencies in Ashkenazi Jewish (AJ) and non-Jewish populations from Europe (EUR) and the Near East (NEA) (Behar 2004)

Summary

Northern and Southern Europe have two different "types" of white skinned people

Northern Europe, including the UK, that colonized the USA, is predominantly
 Protestant and Germanic, isolated from Southern Europe, Latin and Roman Catholic

 Northern Europe originated populations of white skin in Anglo-Saxon America while Southern Europe originated populations of white skin in Latin America

Ask questions and define problems: colonization of Brazil

 Colonization of Brazil started officially in April 22nd, 1500, with the Discovery of Brazil by Pedro Álvares Cabral and Portuguese sailors

Portugal was interested in the newly discovered territory because of the commercialization of brasil-wood (*Paubrasilia echinata*), which gives the country its name

During the Portuguese colonization and after, as an independent country, Brazil received the largest number of Africans and
was the last country in America (the continent) to abolish slavery, in 1888

Ask questions and define problems: colonial racial relationships

- The English language took the Brazilian Portuguese term "negros da terra" (Native Brazilians)
- In the Letter to King Manuel, however, Pero Vaz de Caminha refers to the skin of Native Brazilians as pardo, not Negro
- Eugeny, scientific racism, social darwinism and determinism relate slavery and inferiority of people on the base of skin color
- Native Brazilians and Africans were enslaved in colonial Brazil, leading to the question whether Indigenous Brazilians can be considered Black or in Brazilian Portuguese, Negros
- I see Black Brazilians refer to themselves as Negros, but I have never seen African Americans refer to themselves as Negro Americans

Ask questions and define problems: colonial racial relationships

- In a legal regimento of 1755, Marquis of Pombal determined the Portuguese assimilationist policy that encouraged "interracial" marriage in Brazil
- Vilas like Prado, Alcobaça and Caravelas were founded in the Captaincy of Porto Seguro (Extreme South of Bahia) for colonization and aldeamento of indigenous Brazilians
- The *regimento* criminalized prejudice against descendents of Portuguese and Indigenous Brazilians but not admixed whites (Portuguese) and Africans
- African slavery would only be banished more than 100 years later, in 1888



Fonte: Arquivo Historico ultramarino – Biblioteca Nacional – Brasil (2013). Livro Viagem ao Brasil de Maximiliano – 1817 (1940)

Native Brazilian ethnicities in Southern and Southwestern Bahia (Amaral Rocha 2018)

Ask questions and define problems: colonial racial relationships

- Miscegenation in Brazil, encouraged the emergence of Brazilians of white skin
- With a tri-hybrid, sexually asymmetric genetic inheritance, white Brazilians have European phenotypes (Southern Europe)
 - skin, eye and hair colors
 - hair texture
 - nose and lip shape
- European phenotypes are mainly inherited from the male European colonizer while maternal ancestry is equal from Europe, Africa and America



Ask questions and define problems: ancestry and cancer protection

- This is probably true for the other skin colors in Brazil, because of *branqueamento* or *embranquecimento* (whitening) of Brazil
- Interestingly white skin Brazilians with
 Ashkenazi mitochondrial haplogroup have protection against neuroblastoma (a cancer of the nervous system).



Ideologies of Skin Color and Race: Scientific Racism

 In Brazil, IBGE define Negro as the sum of Mixed Race individuals (Pardos) and Black individuals to make an argument that Brazil is a Black or Negro country

• In the USA, the skin color line border is explained by Thomas Guglielmo in his book, White on Arrival: Italians, Race, Color, and Power in Chicago in the context of the Italian immigration

• The problem of Italians being of the white skin color, but not of the white skin race as Anglo Saxonic Americans, exemplifying to other European ancestries, the American problem with white supremacy

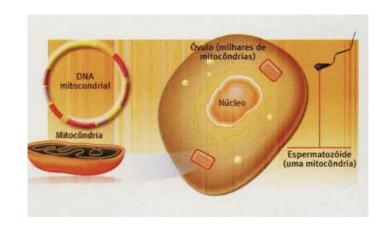
 These two contexts illustrate the white skin color classification in the former European colonies in America (the continent)

Ideologies of Skin Color and Race: Scientific Racism

 Jewish and other ethnic (racial) identities are studied using the genetic inheritance of the mother, not the father

 Mitochondrial DNA sequence is the genetic material that children inherit from the mother only, not from the father

(Maternal) Mitochondrial Genetic Inheritance





Objectives

 Understand cancer protection mechanisms present in the Ashkenazi haplogroup K in contrast to L1 mitochondrial haplogroup, both found in Brazilians of white skin

Hypothesis: If they develop cancer, individuals that carry the Ashkenazi
mitochondrial haplogroup K will be affected by hot tumors, treatable with
immunotherapy, while other ancestries will not be treatable

Plan and carry out investigations

Evaluate the incidence of haplogroup K in hot and cold tumors

 To test the literature suggestion that neuroblastoma hot tumors have a higher incidence of the haplogroup K (this is our hypothesis), the Ashkenazi Jewish haplogroup present in white self-identifying Brazilians

Develop and use models

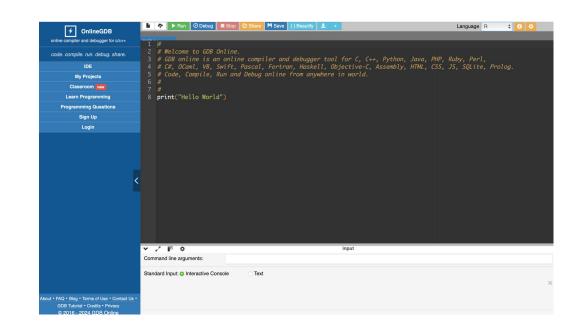
 We can develop a medical test (solution) to diagnose people who may benefit from immunotherapy



Use mathematics and computational thinking

 Models and algorithms developed with computer language R

- IDE (Integrated Development Environment)
 - Tool: Online GDB

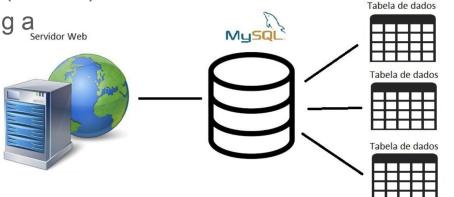


Analyze and interpret data

 A database is a collection of information (text file) that can be accessed and analyzed using a computer.

We will develop two databases:

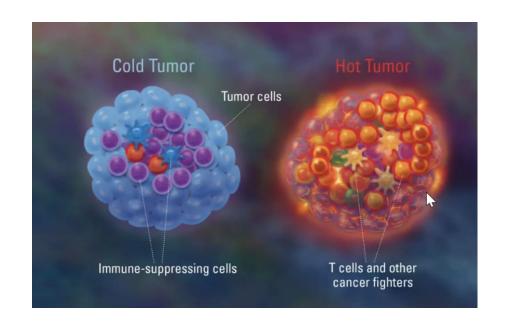
- Mitochondrial database (Brazilian mtDNA)
- Neuroblastoma database (Public tumor DNA data)



Engage in an argument from evidence

 Haplogroup K is related to greater immunological activity or specific metabolic processes in tumor cells.

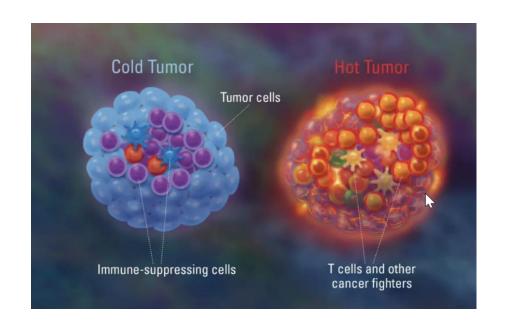
 Hot tumors should have more K haplogroups than cold tumors



Construct explanations and design solutions

 People with haplogroup K may have greater immune activity in tumor cells

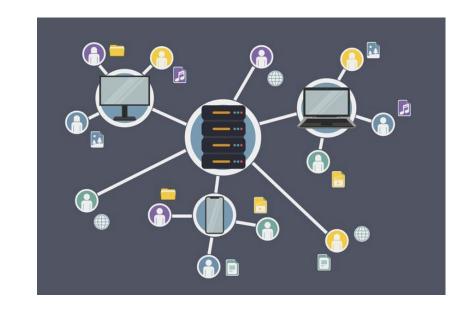
 They present a higher frequency of this haplogroup in hot tumors.



Obtain, evaluate and communicate information

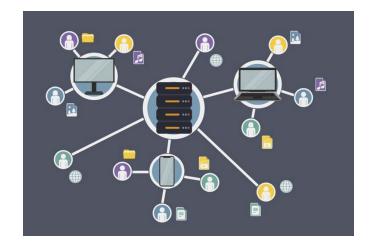
 Process of collecting data or information, critically analyzing it to validate it.

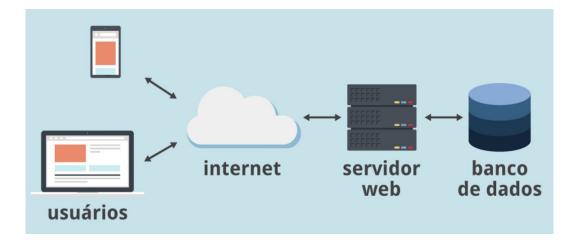
 The model will be discussed and tested using the neuroblastoma database.



GERALDA (Minimum Viable Product)

- Portuguese: Gerenciamento Algorítmico de Dados (GERALDA) (Algorithmic Management of Data)
- Database that can be accessed on the internet (as the UCSC genome browser)
- VCF files aligned to the UCSC human genome reference 19 (hg19)
- Integrate public health/SUS information and Brazilian genealogic information





Conclusions

- In Brazil racial miscegenation generated people of light skin and African ancestry that identify as white (raça/cor)
- Raça/Cor is the variable that the public health system of Brazil (SUS) uses to "prevent discrimination" from happening
- The Ashkenazi ancestry, has potentially lower neuroblastoma risk, as if these people develop cancer, they are predicted (we predict) to develop hot tumors instead of cold tumors
- Hot Tumors Could Help People Fight Cancer
- Cold Tumors Are More Dangerous to People
- Computer Could Help Diagnose Hot and Cold Tumors

Thank you!!!