ABSTRACT

Phytochemicals from Bixa orellana plant extract are traditionally used to cure Dysentery. It is caused by Entamoeba histolytica. Molecular docking method applied using “Biovia Discovery Studio”. “High positive values of -CDOCKER energy and -CDOCKER interaction energy” suggested that benzoin acid can effectively deactivate the alcohol dehydrogenase enzyme thereby interrupting the life cycle of the organism.

Keywords: Phytochemical; Bixa orellana; Entamoeba histolytica.
**Bixaorellana** belongs to family bixaceae. *Bixaorellana* extract is used to cure disease like Dysentery. The objective of the study is to identify the phytochemical responsible to cure the disease.

*Bixaorellana* contains “norbixin, phytol, phenol, acetic acid, anthraquinone, benzoic acid and farnesyl acetone” etc. These phytochemicals might act against Dysentery. However, there is no such study available.

This objective of the study is to identify the phytochemical of *Bixa orellana* capable of curing Dysentery.

### 2. MATERIALS AND METHODS

#### 2.1 Software Used

Discovery studio module of Biovia software (Dassault Systemes of France) was used for analysis. The software utilizes machine learning techniques to predict the level of molecular interaction.

#### 2.2 Methodology

##### 2.2.1 List of phytochemicals

Phytochemicals are secondary metabolites produced by plants to protect them from predators. The potential threats to plants include bacteria, viruses and fungi. When these plants or their parts are consumed by humans these phytochemicals resist threats to health. Some phytochemicals have been used as poisons and others as traditional medicine. Published works showed that *Bixa orellana* contains norbixin, phytol, phenol, acetic acid, anthraquinone, benzoic acid and farnesyl acetone. It has already been established that *Bixa orellana* plant belonging to Bixaceae family has potential to help controlling Dysentery. This work is focused on the identification of specific phytochemical liable for inhibiting and controlling of Dysentery.

##### 2.2.2 Enzyme found in *Entamoeba histolytica*

It has been stated that dysentery can cause a result of *Entamoeba histolytica* invasion. In the bacterial life cycle various metabolic cycles have been seen for its existence. These metabolic cycles are controlled by different enzymes. List of different enzymes found in *Entamoeba histolytica* bacteria are detected by using Brenda enzyme database. It has been found that alcohol dehydrogenase enzyme (protein database code 1Y9A) is involved in ethanol fermentation, methionine, Tryptophan, valine, phenylalanine, leucin metabolism and tyrosine metabolism (BRENDA) and very essential for the existence of the specific microbe.

#### 2.2.3 Molecular docking

Molecular docking method has been used to identify the phytochemical from the plant extract that act as a ligand and form a strong covalent bond with the bacterial protein to successfully inhibit the microbe. The Discovery studio module of Biovia software was used for identifying molecular interaction and perform molecular docking. In this process first the sdf files for the phytochemicals found in the *Bixa orellana* plant were downloaded from the website (www.molinstincts.com). The protein database code of the alcohol dehydrogenase enzyme was recognized from the website (www.rcsb.org). The active site of the enzyme was found under "receptor-ligand interaction” and “define and edit binding site” menu via “receptor cavity” protocol. Molecular docking was done under “receptor-ligand interaction” and “Dock ligand” menu by using the CDOCKER protocol of Biovia software. The enzyme acts as the receptor molecule and the phytochemical act as the ligand.

3. RESULTS AND DISCUSSION

-CDOCKER energy was calculated based on the internal ligand strain energy and receptor-ligand interaction energy. -CDOCKER interaction signifies the energy of the non-bonded interaction that exists between the protein and the ligand. The criteria for best interaction was chosen based on a) high positive value of -CDOCKER ENERGY and b) small difference between -CDOCKER energy and -CDOCKER interaction energy [4,5]. Table 1 shows that alcohol dehydrogenase-benzoic acid interaction has the highest positive value of -CDOCKER
Table 1. Results of C docking of phytochemicals with alcohol dehydrogenase (receptor)

<table>
<thead>
<tr>
<th>Sl. no</th>
<th>Ligand</th>
<th>-CDOCKER energy</th>
<th>-CDOCKER interaction energy</th>
<th>Difference between -CDOCKER interaction energy and -CDOCKER energy</th>
<th>Remarks</th>
</tr>
</thead>
<tbody>
<tr>
<td>1</td>
<td>Benzoic acid</td>
<td>12.6197</td>
<td>13.6892</td>
<td>1.0695</td>
<td>Maximum inhibition of infection</td>
</tr>
<tr>
<td>2</td>
<td>Acetic acid</td>
<td>12.1717</td>
<td>10.3064</td>
<td>1.8653</td>
<td></td>
</tr>
<tr>
<td>3</td>
<td>Phenol</td>
<td>8.30143</td>
<td>19.4032</td>
<td>2.10177</td>
<td></td>
</tr>
<tr>
<td>4</td>
<td>Anthraquinone</td>
<td>12.5373</td>
<td>17.9321</td>
<td>5.3948</td>
<td></td>
</tr>
<tr>
<td>5</td>
<td>Phytol</td>
<td>3.62936</td>
<td>25.8629</td>
<td>22.23354</td>
<td></td>
</tr>
<tr>
<td>6</td>
<td>Norbixin</td>
<td>-6.92316</td>
<td>31.9345</td>
<td>38.85766</td>
<td></td>
</tr>
</tbody>
</table>

energy (12.6197) and minimum value of the difference (1.0695) between -CDOCKER interaction energy and -CDOCKER energy followed by acetic acid, phenol and anthraquinone. Thus the results indicated that benzoic acid, acetic acid, phenol and anthraquinone can efficiently deactivate the alcohol dehydrogenase enzyme thereby interfering the biological cycle of Entamoeba histolytica. Higher positive values for benzoic acid, acetic acid, phenol and anthraquinone showed that these are the most dynamic constituent against Entamoeba histolytica. On the other hand norbixin and farnesyl acetone can deactivate the enzyme to a small extent due to negative -CDOCKER energy but positive -CDOCKER interaction energy. Thus, the important phytochemicals preventing Dysentery caused by Entamoeba histolytica are benzoic acid, acetic acid, phenol and anthraquinone.

4. CONCLUSIONS

It was previously known that Bixa orellana plant has medicinal action against Dysentery. Dysentery is caused by Entamoeba histolytica. This study was carried out to provide the theoretical basis of this observation. Molecular docking operation was executed to identify the phytochemical (benzoic acid, acetic acid, phenol, anthraquinone, phytol, nor-bixin, farnesyl acetone) by using Discovery studio module of Biovia software, which can have a major interaction with the dynamic enzyme (alcohol dehydrogenase) of the microbe. It was found that benzoic acid, acetic acid, phenol and anthraquinone can form strong bond with the enzyme successfully inhibiting the metabolic cycle of the microbe. Phytol, nor-bixin and farnesyl acetone were found to be not much effective in deactivating the enzyme of the microbe as they fail to maintain stability. Thus, this study could explain that the presence of benzoic acid, acetic acid, phenol and anthraquinone delivered the medicinal values to Bixa orellana against Dysentery caused by Entamoeba histolytica.

DISCLAIMER

The products used for this research are commonly and predominantly use products in our area of research and country. There is absolutely no conflict of interest between the authors and producers of the products because we do not intend to use these products as an avenue for any litigation but for the advancement of knowledge. Also, the research was not funded by the producing company rather it was funded by personal efforts of the authors.

CONSENT

It is not applicable.

ETHICAL APPROVAL

It is not applicable.

COMPETING INTERESTS

Authors have declared that no competing interests exist.

REFERENCES

1. Henrich J, Heine S, Norenzayan A. The weirdest people in the world? Behavioral
DOI: 10.1017/S0140525X0999152X

© 2020 Parida et al.; This is an Open Access article distributed under the terms of the Creative Commons Attribution License (http://creativecommons.org/licenses/by/4.0), which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

Peer-review history:
The peer review history for this paper can be accessed here:
http://www.sdiarticle4.com/review-history/57264