## "HSMD: HUMAN SKIN MICROBIOME DATABASE"

Project report submitted in partial fulfilment of the requirement for the degree of Bachelor of Technology

In

### **Bioinformatics**

By

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Under the supervision of

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 $T_{O}$ 



### **BONAFIDE CERTIFICATE**

This is to certify that this project report entitled "HSMD: Human Skin Microbiome Database", submitted to Jaypee University of Information Technology, Waknaghat, is a bonafide record of work done by "Jatin Sharma, Namrata Thakur and Diksha Sharma" for the degree of B.Tech Bioinformatics has been carried out under my supervision.

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Date:

# **Declaration by Authors**

This is to affirm that this report title "HSMD: Human Skin Microbiome Database" has
been written by us i.e. Jatin Sharma, Namrata Thakur and Diksha Sharma under the
supervision of Dr. Jayashree Ramana. No part of the report has been plagiarized from other
sources. All information included from other sources have been duly acknowledged. I aver
that if any part of the report is found to be plagiarized, I shall take full responsibility for it.

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Thanking you,

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# **LIST OF ABBREVATION**

HTML	Hypertext Markup Language
CSS	Cascading Style Sheet
JS	Java Script
PHP	Hypertext Preprocessor

# **ABSTRACT**

HSMD (Human Skin Microbiome Database): It is the web site created such that it will provide data about microorganisms that are responsible for causing various disorders of human skin.

Microbiome is the combined genetic material of the microorganisms in a particular environment.

## Chapter 01.

## **INTRODUCTION**

Human skin that is known to be the largest organ has a diverse environment of microorganisms (that are harmful or even may be beneficial to us). These microorganisms colonize the human skin. Colonization is dependent on several factors such as topographical location, endogenous host and exogenous ecological factors and thus biome of our skin's surface directs their colonization. Skin microbiota is controlled by the immune responses (innate and adaptive). As human skin is diverse and vary from person to person, several molecular methods are being raised so as to view the bacteria that are residing on the human skin. Microbiome's study is augmented and it's even necessary to understand microbial involvement in diseases related to skin. This will assist in antimicrobial and promicrobial ways and treatment.

Skin's environment consists of diverse habitats. The main role of the human skin is to act as physical barrier, regulate the body temperature, protect from harmful effects of various radiations and eliminate toxic substances in form of sweat. Our skin is also an edge to the outside surroundings. Microorganisms in the form of bacteria, viruses and fungi colonizes the skin though harmful in nature, plays a major role such as Symbiotic bacteria take up a huge range of skin niches. These defend from attack by more pathogenic or harmful organisms. These microorganisms even have a role to play in our cell mediated immunity as they train T cells to react to identical microbes.

The diverse habitat of skin environment is a collection of living biological and physical components. The balance between host organism and microorganism needs to be understood. If the balance in either way is destroyed, it can result in various skin infections and diseases. Amongst their balance reasons of distress can be either exogenous or endogenous.

Microbiome, the colony of vast microorganisms (bacteria, viruses and fungi) possess specific environment and also the collection of microorganisms that are present on the human body. This collated environment of genomes of microorganisms specifically found on the human body form one colony that makes up the Microbiome.

Or simply we can say that microb community. While sometimes it e community.		
·		

# CHAPTER 02.

# **OBJECTIVE**

To develop a web server that:

- Archives all the data about the skin microbiome available in literature.
- Provide the scientific community with information on prokaryotic species that are present on the human skin in various diseases.

## CHAPTER 03.

## **LITERATURE REVIEW**

As we know, psoriasis is a condition where building up of skin cells occur and scaly, itchy and dry patches get formed.

Yan D et al. in their paper "The Role of the Skin and Gut Microbiome in Psoriatic Disease" reported about changes observed in the psoriatic disease.

For this a number of studies were conducted by them on skin and gut micobiota in psoriatic patients and non-diseased individuals. Gut microbiome was collected from stool and skin microbiome through biopsy, or skin swabs with or without culturing. The study showed a trend towards an increased relative abundance of Streptococcus , along with decreased level of Propionibacterium in psoriasis patients compared to healthy individuals. In the gut microbiome, the ratio of Firmicutes and Bacteroidetes was ruffled in psoriatic individuals compared to healthy controls. Actinobacteria was also relatively underrepresented.

From this study, a large composition of microbiomes was observed in psoriatic patients. These microbiomes are even responsible for additional diseases along with psoriasis. Propionibacterium which are known for various functions in a human body, like modulating the immune system, their loss can therefore lead to decreased immune tolerance and increased propensity for psoriatic inflammation. Likewise increase in Streptococcus may play a pathogenic role in psoriasis as streptococcal infections have been associated with the later development of guttate psoriasis and the worsening of chronic plaque psoriasis. Changes in the abundance of Staphylococcus in psoriatic skin are less consistent.

It remains unclear whether the observed differences in the microbiota have a causal role in psoriasis or are a consequence of alterations in the environmental milieu from psoriasis.

## CHAPTER 04.

# **MATERIAL A ND METHODS**

## 4.1 PROTOCOL

Data collection (organisms involved in skin microbiota)



Taxonomic study (classification and complete hierarchical data)



Collection of 16s rRNA



Linking organism data to other databases like strain info, ncbi etc.



Preparing a database for skin Microbiome

## **4.2 STEPS FOLLOWED**

- Research papers are collected from PubMed using keywords such as "skin diseases"," human skin AND Microbiome", "skin AND Microbiome", "[disease name] AND Microbiome", "skin microbiota" etc.
- 2. After reading these papers, PMID are collected from these.
- 3. Then in NCBI this ID is used by selecting 'BioProject' in the drop down menu.
- 4. Then biosample is selected.
- 5. Click on first link.
- 6. SRA link is clicked and then click on 'Run'.
- 7. Out of certain options given go to "analysis".
- 8. Then a table containing names of microorganisms appear. This data is maintained in a form of excel sheet.
- 9. For each microorganism their taxonomical data is collected.
- 10. Database is prepared.
- 11. Using HTML, PHP and CSS, a home page of website is created where certain pages are linked.

## 4.3 Use of computational languages:

#### **4.3.1** HTML – (Hyper Text Markup Language)

The back-end language that is used for developing web page.

#### For Home Page:

```
1 <!DOCTYPE html>
3 ⊟<head>
    <meta http-equiv="Content-Type" content="text/html; charset=utf-8" />
    <link href="images/logo.jpg" rel="shortcut icon">
    k rel="stylesheet" type="text/css" href="css/style.css" />
     <script type="text/javascript" src="js/jquery.min.js"></script>
8
    <script type="text/javascript" src="js/jquery.easing.min.js"></script>
9
    <script type="text/javascript" src="js/jquery.nivo.slider.pack.js"></script>
10 | <script type="text/javascript">
11 $ (window).load(function() {
12
           $('#slider').nivoSlider();
13
        });
14
    </script>
15
    <title>HSMD-home page</title>
    <meta name="keywords" content="" />
16
    <meta name="description" content="" />
17
18
    <link href="http://fonts.googleapis.com/css?family=Source+Sans+Pro:200,300,400,600,700,900" rel="stylesheet" />
19
    <link href="default.css" rel="stylesheet" type="text/css" media="all" />
    k href="fonts.css" rel="stylesheet" type="text/css" media="all" />
22
    -</head>
24 | div id="header-wrapper">
26
       <div id="logo">
          <font size="6" face="Arial"><a href="#"><strong>HSMD</strong></a></font>
28
        </div>
29
        <hr>>
30 🖨
        <div id="menu">
31
            <!re><font size="3" face="Arial">
               class="current page item"><a href="#" accesskey="1" title="">Homepage</a>
```

Figure 4. 1 HTML file for Home Page

```
<a href="database.html" title="">Database</a>
34
35
36
                           <a href="contact.html" accesskey="4" title="">Contact Us</a>
                           </font>
                     38
              </div>
       -</div>
39
40
       -</div>
     43
44
45
                          <h2>Welcome to our website</h2>
                    47
48
49
50
51
52
53
54
55
                           </div><!--close slider-->
                    </div><!--close slider_wrapper-->
</div><!--close banner_image-->
<font size="4" face="Arial">
56
57
58
59
                    cyp>This website is dedicated to <strong> database</strong> for<strong> human skin microbiome</strong>
which will present a provisional naming scheme for the currently unnamed species so that strain,
clone, and probe data from any laboratory can be directly linked to a stably named reference scheme.
To discuss more about the information, the links are given on the sidebar.
60
                     </font>
               </div>
```

Figure 4.2 HTML file for Home Page

```
</div>
         <div id="sidebar">
             class="first">
                 <h2><I><font color = "purple">About the Human Skin Microbiome : </font></I></h2>
69
                 <div style="clear: both;">&nbsp;</div>
                     <font size="4">The skin is the human body's largest organ,
                    colonized by a diverse milieu of microorganisms, most of which are harmless
                    or even beneficial to their host.
74
                     76
                <1i>>
78
                     <font size="4"> Colonization is driven by the ecology of the skin surface,
79
                     which is highly variable depending on topographical location, endogenous host factors and exogenous
                     environmental factors. The cutaneous innate and adaptive immune responses can modulate the skin microbiota
                     , but the microbiota also functions in educating the immune system. The development of molecular methods
                     to identify microorganisms has led to an emerging view of the resident skin bacteria as highly diverse
82
                     and variable. An enhanced understanding of the skin microbiome is necessary to gain insight into microbial
84
                     involvement in human skin disorders and to enable novel promicrobial and antimicrobial therapeutic
                     approaches for their treatment.
                     <a href="https://www.ncbi.nlm.nih.gov/pmc/articles/PMC3535073/"><font color="blue"><b>See more...</b>//font></a
                </div>
    -</div>
90
91 -</div>
   div id="footer-wrapper">
         <div id="footer" class="container">
```

#### Figure 4.3 HTML file for Home Page

```
<div class="col-md-12 text-center">
95
              <div style="clear: both;">&nbsp;</div>
96
                  <font color="white">Copyright &copy; 2017 | Human Skin Microbiome Database</font>
97
                  <div style="clear: both;">&nbsp;</div>
98
              </div>
99
          </div>
100
      </body>
101
     -</html>
102
```

Figure 4.4 HTML file for Home Page

#### **4.3.2 PHP (Hypertext Pre-processor)**

This is a server side scripting language that is usually designed for the web development and is even used generally as a programming language. Here we used this language for our database page.

#### For Database Page:

```
chemal>
chemal>
chemad>

meta http-equiv="Content-Type" content="text/html; charset=utf-8" />
clink href="images/logo.jpg" rel="shortcut icon">
clink rel="stylesheet" type="text/css" href="css/style.css" />
ctitle>Search page</title>
cmeta name="keywords" content="" />
cmeta name="description" content="" />
clink href="http://fonts.googleapis.com/css?family=Source+Sans+Pro:200,300,400,600,700,900" rel="stylesheet" />
clink href="default.css" rel="stylesheet" type="text/css" media="all" />
clink href="fonts.css" rel="stylesheet" type="text/css" media="all" />
clink href="fonts.css" rel="stylesheet" type="text/css" media="all" />
clink href="fonts.css" rel="stylesheet" type="text/css" /><![endif]-->

cropho

session start();
require('dBconfig.php');
sconn = mysqli_connect($servername, $username, $password, $dbname);

if (!$conn) {
    die("Connection failed: " . mysqli_connect_error());

    $suserTable = "taxon_description";
    $sqli = "SELECT * FROM $userTable";
    $result = mysqli_query($conn, $sqli);

}

**Possion of the content in the content in
```

Figure 4.5 Code for .php file

```
<div id="header-wrapper">
<div id="header" class="container">
             <div id="logo">
                   <font size="6" face="Arial"><a href="#"><strong>HSMD</strong></a></font>
             </div>
<div id="menu">
                   font size="3" face="Arial">
                   <a href="index.html" accesskey="1" title="">Homepage</a>
<a href="database.php" title="">Database</a>
<a href="contact.html" accesskey="4" title="">Contact Us</a>
</a>
41
42
43
44
                        </font>
45
46
                   </111>
             </div>
       -</div>
49
50
     =<div id="page" class="container">
      51
52
53
54
             <div class="content">
                  <br><br><br><br>>
55
56
57
                  <div class="post">
                             <font size="4" color="red"><U>TAXON DESCRIPTION:</U></font></h3><br/><br/>
                              <font $12e="4" Color="led <>>> mace 22
<div style="clear: both;">&nbsp;</div>
<div style="clear: both;">&nbsp;</div></div>
60
                              <div class="entry">
```

Figure 4.6 Code for .php file

#### ...contd.

Figure 4.7 Code for .php file

```
93
94
95
96
97
98
99
                                   </div>
                           </div>
104
105
106
107
108
109
                           <div style="clear: both;">&nbsp;</div>
<div style="clear: both;">&nbsp;</div>
                    </div>
         </div>
-</div>
                    <div id="footer-wrapper">
                          v id="footer-wrapper">
<div id="footer-wrapper">
<div id="footer-wrapper">
<div class="col-md-12 text-center">
<div class="col-md-12 text-center">
<div style="clear: both;">snbsp;</div>
<font color="white">Copyright &copy; 2017 | Human Skin Microbiome Database</font>
112
113
114
115
                                   <div style="clear: both;">&nbsp;</div>
116
117
118
119
                    </div>
         -</body>
```

Figure 4.8 Code for .php file

### **4.3.3** CSS - (Cascading Style Sheets)

It is used to describe how HTML elements are to be displayed on screen. It is used for saving a lot of work.

```
□body {
9
         margin: 0;
10
         padding: 0;
11
         background: #C5DA99 url(images/img01.jpg) repeat;
12
         font-family: Arial, Helvetica, sans-serif;
13
         font-size: 14px;
14
         color: #787878;
15
    L}
16
17
   ⊟h1, h2, h3 {
18
         margin: 0;
19
         padding: 0;
20
         font-weight: normal;
         font-family: 'Abel', sans-serif;
21
22
         color: #3DA29B;
    L}
23
24
25
    □h1 {
26
         font-size: 2em;
27
    L}
28
29
    □h2 {
30
         font-size: 2.8em;
31
32
33
    □h3 {
34
         font-size: 1.6em;
    L}
35
36
37
   □p, ul, ol {
38
         margin-top: 0;
39
         line-height: 180%;
```

Figure 4.9 Code for .css file

...contd.

```
40 -}
41
42 ⊟ul, ol {
43 43
44
45 □a {
46
       text-decoration: none;
47
       color: #5E5E5E;
   □}
48
49
50
   □a:hover {
51
   L}
52
53 □#wrapper {
54
    padding-top: 50px;
55
   L}
56
57 □.container {
       width: 1000px;
58
59
       margin: 0px auto;
   L }
60
61
62 /* Header */
63
64 □#header-wrapper {
65
       overflow: hidden;
       height: 140px;
66
67 L}
68
69 □#header {
        width: 900px;
70
```

Figure 4.10 Code for .css file

...contd.

```
71
72
73
            height: 100px;
            margin: 0 auto;
padding: 0px 50px;
            background: url(images/img02.jpg) repeat;
 75
 76
77
78
79
       /* Logo */
     □#logo {
 80
            float: left;
            width: 300px;
 81
 82
            margin: 0;
 83
            padding: 0;
 84
            color: #FFFFFF;
 85
 86
     ⊟#logo h1, #logo p {
 87
 88
 89
 90
     ⊟#logo h1 {
            line-height: 100px;
            letter-spacing: -2px;
text-transform: lowercase;
 92
 93
 94
            font-size: 3.8em;
      L
 95
 96
     ⊟#logo h1 a {
    color: #FFFFF;
 98
            text-shadow: 1px 1px 0px rgba(0,0,0,.2);
100
```

Figure 4.11 Code for .css file

```
438 /* Footer */
439
440
     □#footer {
441
          height: 100px;
442
           margin: 0 auto;
           padding: 0px 0 15px 0;
font-family: 'Abel', sans-serif;
443
444
     L }
445
446
447
     □#footer p {
448
          margin: 0;
449
           padding-top: 10px;
450
           letter-spacing: 1px;
451
           line-height: normal;
           font-size: 14px;
452
453
           text-transform: uppercase;
454
           text-align: center;
455
           color: #527800;
456
      L }
457
     □#footer a {
458
           color: #527800;
459
460
```

Figure 4.12 Code for .css file

### 4.3.4 JavaScript

It is used for making web pages interactive and provides online programs.

```
☐ (function() {
          var method;
          var noop = function () {};
          var methods = [
 5 🖨
              'assert', 'clear', 'count', 'debug', 'dir', 'dirxml', 'error',
 6
              'exception', 'group', 'groupCollapsed', 'groupEnd', 'info', 'log', 'markTimeline', 'profile', 'profileEnd', 'table', 'time', 'timeEnd',
 7
 8
 9
              'timeStamp', 'trace', 'warn'
10
11
          var length = methods.length;
12
          var console = (window.console = window.console || {});
13
14 🛱
          while (length--) {
15
              method = methods[length];
16
17
              if (!console[method]) {
18
                   console[method] = noop;
19
               }
20
21
    L}());
23
24
    □if (typeof Object.create !== 'function') {
25
          Object.create = function(obj) {
26
               function F() {}
27
              F.prototype = obj;
28
               return new F();
29
30
31
32
    ☐ (function($, window, document, undefined) {
33
          "use strict";
```

Figure 4.13 Code for .js file

...contd.

```
171
                   return section || this.$links[0].hash;
172
173
          };
174
175
          $.fn.singlePageNav = function(options) {
176
               return this.each(function() {
177
                   var singlePageNav = Object.create(SinglePageNav);
178
                   singlePageNav.init(options, this);
179
               });
180
          };
181
          $.fn.singlePageNav.defaults = {
182 🛱
183
              offset: 0,
184
              threshold: 120,
185
              speed: 400,
186
              currentClass: 'current',
187
              easing: 'swing',
              updateHash: false,
188
189
               filter: '',
190
              onComplete: false,
191
              beforeStart: false
192
           };
193
194
     }) (jQuery, window, document);
195
```

Figure 4.14 Code for .js file

## CHAPTER 05.

## **RESULTS**

After collection of data from several papers and resources on human skin Microbiome, we collected the data and assembled them into one common excel file and then worked on site where we created three pages (i.e. home page, database page where collection of data is represented and further links are given in it regarding taxonomy, and lastly the contact page).

#### Home page:

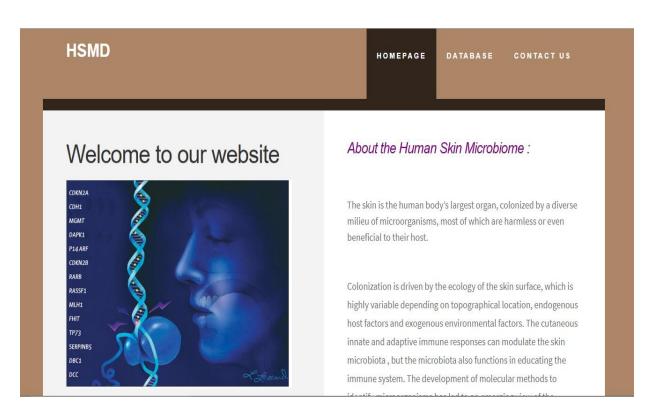


Figure 5.1 View of the homepage

#### **Database page:**

For this page, certain steps were performed:

- 1. Install xampp in the system. Then in the folder named xampp we have another folder 'htdocs'. Here create your folder (name given as 'hsmd'). In this store your all files like html files, sql file, css file, js file and others.
- 2. Excel sheet containing the data was converted into .csv file.
- 3. Then xampp control panel was run. Actions for module (apache and MySQL) were started.

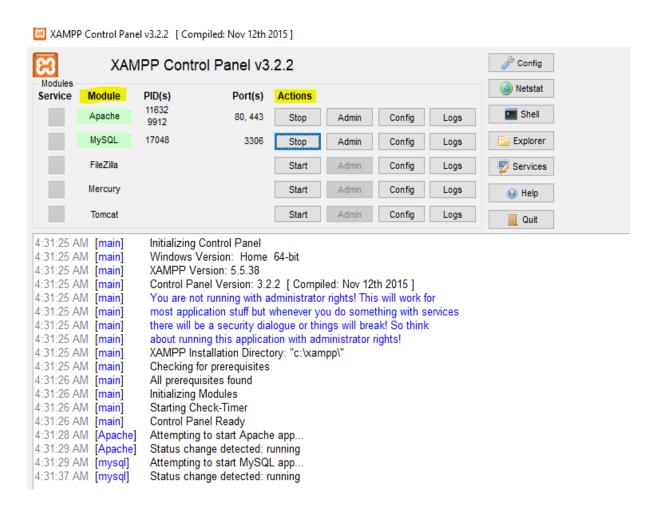


Figure 5.2 While XAMPP Control Panel is made to run

- 4. In the chrome we open localhost/phpmyadmin. Here we create database.
- 5. Then .csv file is imported using load data.



Figure 5.3 Image showing structure of 'taxon description' table

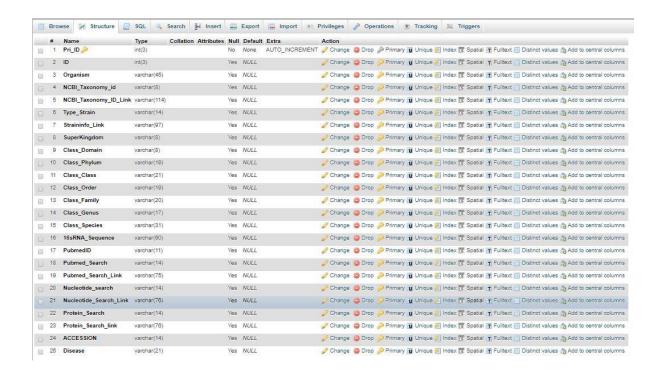


Figure 5.4 Image showing the structure for 'description' table linked to table 'taxon\_description'

6. Database created is then exported and stored in the folder created in htdocs of xampp folder in C drive.

In the database page in order to see the taxon description, under the description column click on the "click here" button .

Then,

You will obtain a "taxon description" page like this:

#### Porphyromonas sp. KLE 1280

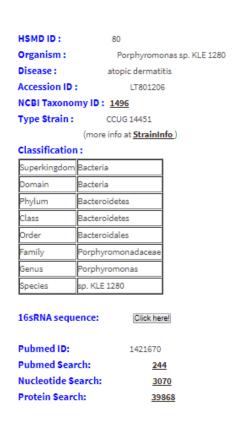


Figure 5.5 Image showing Human skin microbiome description for *Porphyromonas* species

In this page we will obtain taxon description of the microorganism found on the human skin, responsible for diseases. This will contain information about:

- 1. HSMD ID
- 2. Organism Name
- 3. Disease Caused By the Oragnism
- 4. Accession ID
- 5. NCBI Taxonomy ID
- 6. Type Strain
- 7. Hierarchical Classification
- 8. 16sRNA sequence
- 9. Pubmed ID
- 10. Nucleotide Search ID
- 11. Protein Search ID

Here in this description page you can view the above mentioned by clicking on the links .

For example, in the shown image if we click on "click here", a page will be displayed showing 16s RNA sequence.

16sRNA sequence: Click here!

Pubmed ID: 1421670

Pubmed Search: 244

Nucleotide Search: 3070

Protein Search: 39868

Figure 5.6 example showing where one can get more information through

#### Porphyromonas sp. partial 16S rRNA gene, isolate Otu000261

GenBank: LT801206.1

FASTA Graphics Go to: ✓ LOCUS 426 bp DNA linear ENV 20-NOV-2017 DEFINITION Porphyromonas sp. partial 16S rRNA gene, isolate Otu000261. ACCESSION LT801206 LT801206.1 BioProject: PRJEB19934 KEYWORDS ENV. SOURCE Porphyromonas sp. ORGANISM Porphyromonas sp. Bacteria; Bacteroidetes; Bacteroidia; Bacteroidales; Porphyromonadaceae; Porphyromonas. REFERENCE AUTHORS Vanlancker,E. TITLE Direct Submission JOURNAL Submitted (10-MAR-2017) GHENT UNIVERSITY, Center for Microbial Ecology and Technology (CMET), Coupure Links 653, 9000 Ghent, Belgium FEATURES Location/Qualifiers source /organism="Porphyromonas sp." /mol\_type="genomic DNA" /isolate="Otu000261" /isolation\_source="Human gut in vitro model" /db\_xref="taxon:<u>1924944</u>" /environmental\_sample <1..>426 gene /gene="16S rRNA" rRNA <1..>426 /gene="165 rRNA"

Figure 5.7 Image shows data for 16s RNA sequence on click

/product="16S ribosomal RNA"

And similarly if we click on another link as follows:

Pubmed ID: 1421670

Pubmed Search: 244

Nucleotide Search: 3070

Protein Search: 39868

Figure 5.8 example showing where one can get more information about pubmed

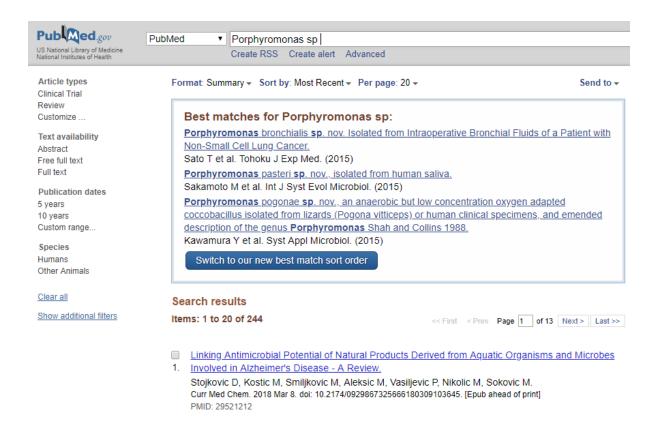


Figure 5.9 Image shows pubmed data

## **Contact Page:**

Through this page, information can be obtained regarding the skin Microbiome from the authors.

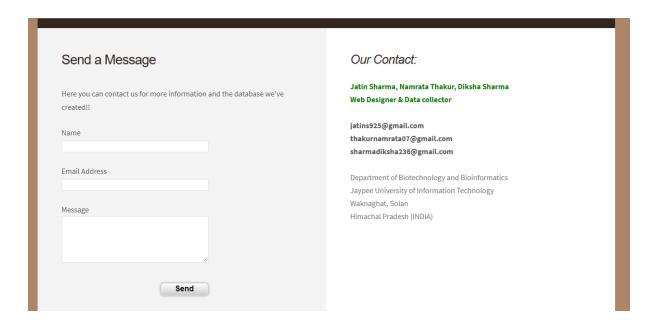


Figure 5.10 Contact page of database

# CHAPTER 06.

# **CONCLUSION**

This HSMD site provides biological entries for each species and an extensive catalogue of genes that these microbes express along with the entire genome of species, protein information, strain information, taxonomical classification, sequence of 16srRNA and a few more information.

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