<u>Lab sheet1:</u> Information retrieval and sequence analysis

Q1

COX-2 (prostaglandin H2 synthase-2 (PTGS2)) gene

- i. Access the entries for Human PTGS1 and PTGS2 in the "Gene" database at the NCBI (https://www.ncbi.nlm.nih.gov/) Website.
 - a. PTGS1 and PTGS2 are isozymes. Isozymes catalyze the same reaction but are separate genes. What types of reactions do PTGS enzymes catalyze? Also, what pathway are these enzymes a part of?

PTGS1 and PTGS2 are isozymes that catalyze the conversion of arachidonic acid into prostaglandins. They are part of the prostaglandin synthesis pathway.

b. How is the expression of PTGS1 and PTGS2 different?

PTGS1 is constitutively expressed in many tissues, while PTGS2 is inducible. There is a biased expression in bone marrow (RPKM 59.3), urinary bladder (RPKM 41.1) and 11 other tissues

- c. Which isozyme (PTGS1 or PTGS2) is required to inhibit inflammation?
 PTGS2 is thought to be the more important isozyme. PTGS2 is inducible and is therefore produced in higher levels at sites of inflammation
- d. The drug Celebrex selectively inhibits PTGS2 while aspirin and other NSAID's inhibit both PTGS1 and PTGS2 in the same way. Why do you think researchers wanted to discover a selective inhibitor to PTGS2?
 - To reduce the side effects of NSAIDs.
 - To develop a more effective treatment for inflammatory diseases.
 - To gain a better understanding of the role of PTGS2 in inflammation.
- e. Describe how studying 3-D structures of PTGS1 and PTGS2 could help researchers design a drug that binds to PTGS1, but not to PTGS2.

By identifying the active site of each enzyme. The active site is the part of the enzyme that binds to the substrate. By comparing the active sites of PTGS1 and PTGS2, researchers could identify differences that could be exploited to design a drug that binds specifically to one enzyme or the other.

- ii. Considering the Homo sapiens PTGS2 gene entry in NCBI gene https://www.ncbi.nlm.nih.gov/gene/ database,
 - a. What is the gene name?

PTGS2

b. What is the GeneID number? 5743

- Where in the human genome is this gene located?
 PTGS2 gene is located on chromosome 1, at position 1q31.1
- d. What is the RefSeq accession number for the mRNA sequence of H o m o sa pie n s prostaglandin-endoperoxide synthase 2? NM_000963.3

e. Download the prostaglandin-endoperoxide synthase 2 Reference mRNA sequence in "FASTA" format.

gene_id: 5743
gene_symbol: PTGS2

description: prostaglandin-endoperoxide synthase 2

scientific_name: Homo sapiens common_name: human

tax_id: 9606

genomic_range: NC_000001.11:186671791-186680423;NC_060925.1:186026616

186035248 orientation: -;location: chr 1

gene_type: PROTEIN_CODING
transcript_accession: NM_000963.4

transcript_name: transcript_length: 4510

transcript_cds_coords: NM_000963.4:134-1948

protein_accession: NP_000954.1

isoform_name:
protein_length: 604

protein_name: prostaglandin G/H synthase 2 precursor

f. What is the RefSeq accession number for the H o m o sa pie n s PTGS2 protein sequence? Download the sequence in "FASTA" format.

NP_000954.1

- iii. Search for the UniProt entry for PTGS2 in Expasy https://www.expasy.org/website.
 - a. What are the alternate names for this protein.

Prostaglandin-endoperoxide synthase 2, Cyclooxygenase-2

b. What types of drugs target this protein?

Nonsteroidal anti-inflammatory drugs

c. What amino acid is acetylated by aspirin (amino acid type)?

Serine acid

iv. Translate the mRNA sequence of PTGS2 into Protein. Use "Translate " tool in ExPASy. Explain the output.

Fasta format

> VIRT-54282:5'3' Frame 2, start_pos=44
MLARALLLCAVLALSHTANPCCSHPCQNRGVCMSVGFDQYKCDCTRTGFY
GENCSTPEFLTRIKLFLKPTPNTVHYILTHFKGFWNVVNNIPFLRNAIMS
YVLTSRSHLIDSPPTYNADYGYKSWEAFSNLSYYTRALPPVPDDCPTPLG
VKGKKQLPDSNEIVEKLLLRRKFIPDPQGSNMMFAFFAQHFTHQFFKTDH
KRGPAFTNGLGHGVDLNHIYGETLARQRKLRLFKDGKMKYQIIDGEMYPP
TVKDTQAEMIYPPQVPEHLRFAVGQEVFGLVPGLMMYATIWLREHNRVCD
VLKQEHPEWGDEQLFQTSRLILIGETIKIVIEDYVQHLSGYHFKLKFDPE
LLFNKQFQYQNRIAAEFNTLYHWHPLLPDTFQIHDQKYNYQQFIYNNSIL
LEHGITQFVESFTRQIAGRVAGGRNVPPAVQKVSQASIDQSRQMKYQSFN
EYRKRFMLKPYESFEELTGEKEMSAELEALYGDIDAVELYPALLVEKPRP
DAIFGETMVEVGAPFSLKGLMGNVICSPAYWKPSTFGGEVGFQIINTASI
QSLICNNVKGCPFTSFSVPDPELIKTVTINASSSRSGLDDINPTVLLKER
STEL

First extracted the DNA seq of the PTGS2 from the file and translated it from the translator using the Verbose: Met, Stop, spaces between residues output format. Then chosen the most suitable frame and got the protein sequence and compared it with the protein sequence downloaded from the website.

Q2. Python Exercises

1. Write a Python code to extract the sample name from these files ignoring any files which do not match the format given below.

The format is:

- 1. Written lane number
- 2. Barcode
- 3. Sample name
- 4. Numeric lane number (starting with L)
- 5. Read number (R1/2/3/4)
- 6. File extension

Eg. Lane8127_GCCAAT_S30_1_21_Hap4_log_L001_R1.fastq.gz the sample name would be, S30_1_21_Hap4_log

```
import re

def extract_sample_name(file_name):
    pattern = r'^lane\d+_([A-Z\d]+)_([A-Za-z\d]+)_L\d+_([R]\d+)\.fastq\.gz$'
```

```
match = re.match(pattern, file name)
    if match:
        return match.group(2)
    else:
        return "no"
files = [
    "lane1 NewCode L001 R1.fastq.gz",
    "lane1 NoIndex L001 R1.fastq.gz",
    "lane1 NoIndex L001 R2.fastq.gz"
    "pipeline processing output.log",
    "lane7027 ACTGAT_JH25_L001_R1.fastq.gz",
    "lane7027 ACTTGA E30 1 2 Hap4 24h L001 R1.fastq.gz",
    "lane7027 AGTTCC JH14 L001 R1.fastq.gz",
    "lane7027 CGGAAT JH37 L001 R1.fastq.gz",
    "lane7027 GCCAAT E30 1 21 Hap4 log L001 R1.fastq.gz"
    "lane7127 GGCTAC E30 1 4 Hap4 48h L001 R1.fastq.gz",
    "lane8127 GCCAAT S30 1 21 Hap4 log L001 R1.fastq.gz"
sample names = [extract sample name(file name) for
file name in files]
for file name, sample name in zip(files, sample names):
   print(f"{file name} -> {sample name}")
```

2.Create a FASTA file by obtaining 10 Dengue 1- Envelop gene DNA sequences from NCBI. Write a Python-program that reads the FASTA file, cleans up the header line to have only Accession number & gene-name and print headers and sequences to standard output as multi-FASTA-file again.

Here the DNA sequence that I wrote in the code is not the original gene sequence of Dengue 1- Envelop gene DNA sequences. It is an example.

```
import re
from Bio import Entrez, SeqIO

def fetch_dengue_sequences():
    Entrez.email = "pasindupathiranagama@gmail.com"
    handle = Entrez.esearch(db="nucleotide", term="Dengue 1 Envelope", retmax=10)
    record = Entrez.read(handle)
    ids = record["IdList"]

    dengue_sequences = []
    for record_id in ids:
        handle = Entrez.efetch(db="nucleotide", id=record_id, rettype="gb",
retmode="text")
        seq_record = SeqIO.read(handle, "genbank")
        dengue_sequences.append(seq_record)

    return dengue_sequences
```

```
def write fasta file(sequences, output file="dengue sequences.fasta"):
    with open(output_file, "w") as output_handle:
        SeqIO.write(sequences, output_handle, "fasta")
def clean_up_header(header):
    pattern = r'(\S+).*?\((\w+)\) gene'
    match = re.search(pattern, header)
   if match:
        accession_number = match.group(1)
        gene_name = match.group(2)
        cleaned_header = f">{accession_number}_{gene_name}"
       return cleaned_header
        return ">Unknown"
def print_multi_fasta(file_path):
    sequences = list(SeqIO.parse(file_path, "fasta"))
    for seq_record in sequences:
        cleaned_header = clean_up_header(seq_record.description)
        print(cleaned header)
        print(seq_record.seq)
if __name__ == "__main__":
    dengue_sequences = fetch_dengue_sequences()
    write_fasta_file(dengue_sequences)
   print_multi_fasta("dengue_sequences.fasta")
```

3. Write a Python program to search the DNA Sequence for the presence of one of the following Transcription Factor Binding Sites(TFBS) with ambiguity codes. Search for all the positions in the sequence where TFBS is located.

Transcription Factor	Consensus Sequence
RUNX1	BHTGTGGTYW
TGIF1	WGACAGB
IKZF1	BTGGGARD

Code	Represents
Α	Adenine
G	Guanine
С	Cytosine
Т	Thymine
Υ	Pyrimidine (C or T)
R	Purine (A or G)
W	weak (A or T)
S	strong (G or C)
K	keto (T or G)
М	amino (C or A)
D	A, G, T (not C)
V	A, C, G (not T)
Н	A, C, T (not G)
В	C, G, T (not A)

The sequence is shown below.

>search_seq

GACACCTCAGTACTAGGATGNNNNNTATCAGCCTGAACTAGCAGGCCTGGTTCCAAATT TTTTTATCAACACTCGTAGGGGGATTATCCTAGAGGGGGGTCTGGGATTTCTTTGACATCA GAGTATTTTTGCCTTGCTCCTTCACAATTTGGGAACAAATAATTTAGTGGTTATTAACCC TGGCTACGCACTGGAAACTTTAAAAATAATGCTGGTATGAAATTTACACAGAGTATCGTG AAAATTTTCACTGAGTACCATGTGGTTATACATTGGATAAGGCTCCAGGAAGCAGCTACT GGAAGACAGCCATGCCAAGAGTGGTTAGTGGTTGGAATTTTGGCAAGTCAGTTTTAGTCT TTTATTTAAGGTAGTCTCTATCTGCCTCTGTCTCTGTCTCTGTCTCTGTGTCTCTG GTGTGTGTGTGTGCATGAACATGAGTAAAATCCATAAGGAAACTTTCAGAGTTGGTC $\tt CTCTCCTTATATCAAATGGATCCAGGAATTAAACTCAGGTTCAATTCTTGGTGCCTTTAC$ TAGTTGAGCCATCTCACTGGCTCTTCATCATCTTTAGAATAAACTCACTTTATTACACAC ACACACACACACACACTGGGAGTACACACACACACACAACCAAAGCCCCAACGGAAAA CTACAATATTATAATGAATACACAGGTTCTCAACATAGTCTCTGCCACGCTTGCAGACAA AGATGAGTAGAAGTAGAAAGAACCAGGGAAACGTGGAGCAAGTCAGAAGGAATAACAGTC AGAAGGAATAACAGTCAGAAGGAATAACAGTCAGAAGGAATAACAGTCAGAAGGAATAGC AGTCAGAAGGAATAACAGTCAGAAGACAGCACAGTCAGAAGGAATAACAGTCAGAAGGAA TAACAGTCAGAAGGAATAACAGTCAGAAGGAATAACAGTCAGAAGGAATAGCAGTCAGAA GGAATAACAGTCAGAAGGAATAACAGTCAGAAGGAATAACAGTCAAAGAAATAGCAGTCA GAAGGAATAGCAGTCAGAAGGAATAACAGTCAAAGGAGCAGTCAGAAGGAGTAACAGTCA GAAGGAATAACAGTCAGAAGGAATAACAGTCAAAGGAATAGCAGTCAGAAGGAGTAACAG TCAGAGCAAACACAGAGATGACAAAGGCAATGGGGTCAGAGACTTCACCACTCTCCAAGA

```
import re
def search tfbs(sequence, tfbs dict):
    sequence = "".join(sequence.split())
    positions = {}
    for tf, consensus_sequence in tfbs_dict.items():
        consensus_sequence =
consensus sequence.replace('B', '[CGT]')
        consensus sequence =
consensus_sequence.replace('D', '[AGT]')
        consensus sequence =
consensus sequence.replace('H', '[ACT]')
        consensus sequence =
consensus sequence.replace('K', '[GT]')
        consensus sequence =
consensus_sequence.replace('M', '[AC]')
        consensus_sequence =
consensus sequence.replace('N', '[ACGT]')
        consensus_sequence =
consensus sequence.replace('R', '[AG]')
        consensus sequence =
consensus sequence.replace('S', '[CG]')
        consensus sequence =
consensus sequence.replace('V', '[ACG]')
```

```
consensus sequence =
consensus sequence.replace('W', '[AT]')
       consensus sequence =
consensus sequence.replace('Y', '[CT]')
       matches = [match.start() for match in
re.finditer(f'(?={consensus sequence})', sequence)]
       if matches:
           positions[tf] = matches
   return positions
if __name__ == "__main__":
   search seg = """
   GACACCTCAGTACTAGGATGNNNNNNTATCAGCCTGAACTAGCAGGCCTGGT
TCCAAATT
   TTTTTATCAACACTCGTAGGGGGATTATCCTAGAGGGGGTCTGGGATTTCTT
TGACATCA
   GAGTATTTTTGCCTTGCTCCTTCACAATTTGGGAACAAATAATTTAGTGGTT
ATTAACCC
   TGGCTACGCACTGGAAACTTTAAAAATAATGCTGGTATGAAATTTACACAGA
GTATCGTG
   AAAATTTTCACTGAGTACCATGTGGTTATACATTGGATAAGGCTCCAGGAAG
CAGCTACT
   GGAAGACAGCCATGCCAAGAGTGGTTAGTGGTTGGAATTTTGGCAAGTCAGT
TTTAGTCT
   GCCTTATCAAATACATGGGCATACAGATAAATCCTTAGATGGCTCTCCTACT
TACTGAAA
   ATCTATCA
   TTTATTTAAGGTAGTCTCTATCTGCCTCTGTCTCTGTCTCTGTGTCTC
TGTGTCTG
```

GTGTGTGTGTGTGCATGAACATGAGTAAAATCCATAAGGAAACTTTCAG AGTTGGTC

CCTCTCCTTATATCAAATGGATCCAGGAATTAAACTCAGGTTCAATTCTTGG TGCCTTTAC

TAGTTGAGCCATCTCACTGGCTCTTCATCATCTTTAGAATAAACTCACTTTA
TTACACAC

ACACACACACACACCACGGGAGTACACACACACACACAACCAAAGCCCCA
ACGGAAAA

CTACAATATTATAATGAATACACAGGTTCTCAACATAGTCTCTGCCACGCTT GCAGACAA

AGATGAGTAGAAGTAGAAAGAACCAGGGAAACGTGGAGCAAGTCAGAAGGAA
TAACAGTC

AGAAGGAATAACAGTCAGAAGGAATAACAGTCAGAAGGAATAACAGTCAGAAGGAATAGC

AGTCAGAAGGAATAACAGTCAGAAGACAGCACAGTCAGAAGGAATAACAGTC AGAAGGAA

TAACAGTCAGAAGGAATAACAGTCAGAAGGAATAACAGTCAGAAGGAATAGC AGTCAGAA

GGAATAACAGTCAGAAGGAATAACAGTCAGAAGGAATAACAGTCAAAGAAAT AGCAGTCA

GAAGGAATAGCAGTCAGAAGGAATAACAGTCAAAGGAGCAGTCAGAAGGAGT
AACAGTCA

GAAGGAATAACAGTCAGAAGGAATAACAGTCAAAGGAATAGCAGTCAGAAGG AGTAACAG

TCAGAGCAAACACAGAGATGACAAAGGCAATGGGGTCAGAGACTTCACCACT
CTCCAAGA

 $\mathbf{H}_{-}\mathbf{H}_{-}\mathbf{H}_{-}$

```
tfbs_dict = {
    "RUNX1": "BHTGTGGTYW",
```

```
"TGIF1": "WGACAGB",
    "IKZF1": "BTGGGARD"
}

positions = search_tfbs(search_seq, tfbs_dict)

for tf, tf_positions in positions.items():
    print(f"{tf} found at positions:
{tf_positions}")
```

Output:-

```
PS C:\Users\Melaka> python -u "d:\Acadamic\4 th year\2nd sem\CB\Lab 1\q3.py"
RUNX1 found at positions: [258]
TGIF1 found at positions: [303, 1044]
IKZF1 found at positions: [454, 560, 798]
PS C:\Users\Melaka> [
```

Q3 - Biopython

Biopython Tutorial and Cookbook https://biopython.org/DIST/docs/tutorial/Tutorial.html#sec2

1. Write a Biopython program that asks the user to input a DNA-sequence and then translates the sequence to protein sequence.

```
2.
3. codon_table = {
       "TTT": "F", "TTC": "F", "TTA": "L", "TTG": "L",
4.
       "CTT": "L", "CTC": "L", "CTA": "L", "CTG": "L",
       "ATT": "I", "ATC": "I", "ATA": "I", "ATG": "M",
6.
7.
       "GTT": "V", "GTC": "V", "GTA": "V", "GTG": "V",
       "TAT": "Y", "TAC": "Y", "TAA": "*", "TAG": "*",
8.
9.
       "CAT": "H", "CAC": "H", "CAA": "O", "CAG": "O",
             "AAT": "N", "AAC": "N", "AAA": "K", "AAG": "K",
10.
             "GAT": "D", "GAC": "D", "GAA": "E", "GAG": "E",
11.
             "TCT": "S", "TCC": "S", "TCA": "S", "TCG": "S",
12.
             "CCT": "P", "CCC": "P", "CCA": "P", "CCG": "P",
13.
             "ACT": "T", "ACC": "T", "ACA": "T", "ACG": "T",
14.
15.
             "GCT": "A", "GCC": "A", "GCA": "A", "GCG": "A",
16.
             "TGT": "C", "TGC": "C", "TGA": "*", "TGG": "W",
             "CGT": "R", "CGC": "R", "CGA": "R", "CGG": "R",
17.
             "AGT": "S", "AGC": "S", "AGA": "R", "AGG": "R",
18.
19.
             "GGT": "G", "GGC": "G", "GGA": "G", "GGG": "G"
20.
21.
22.
         dna_sequence = input("Enter DNA sequence: ")
23.
24.
         if not all(c in "ACTG" for c in dna_sequence):
25.
             print("Invalid DNA sequence. Please enter a valid sequence.")
26.
             exit()
27.
         if len(dna_sequence) % 3 != 0:
28.
```

```
29.
             print("DNA sequence length must be a multiple of 3. Please
   enter a valid sequence.")
             exit()
30.
31.
         protein sequence = ""
32.
33.
         for i in range(0, len(dna sequence), 3):
34.
             codon = dna sequence[i:i+3]
35.
             protein sequence += codon table[codon]
36.
37.
         print("Protein sequence:", protein_sequence)
38.
```

```
PS D:\Acadamic\4 th year\2nd sem\CB\Lab 1> python -u "d:\Acadamic\4 th year\2nd sem\CB\Lab 1\3.1.py"
Enter DNA sequence: ATGGCCTACGCACTGGAAACT
Protein sequence: MAYALET
PS D:\Acadamic\4 th year\2nd sem\CB\Lab 1>
```

39. Write a Biopython program that will find all articles related to Alzheimer's in PubMed. Print the total number of articles available and the authors.

```
from Bio import Entrez

def search_pubmed(query, max_results=10):
    Entrez.email = "pasindupathiranagama@gmail.com"
    handle = Entrez.esearch(db="pubmed", term=query,
retmax=max_results)
    record = Entrez.read(handle)
    handle.close()
    return record

def fetch_pubmed_details(id_list):
    ids = ",".join(id_list)
```

```
handle = Entrez.efetch(db="pubmed", id=ids,
rettype="medline",
    retmode="text")
    records = handle.read()
    handle.close()
    return records
def parse pubmed records(records):
    authors list = []
    for record in records.split("\n\nPMID")[1:]:
        authors = []
    for line in record.split('\n'):
        if line.startswith('AU - '):
            authors.append(line[6:])
            authors list.append(authors)
    return authors list
if __name__ == "__main__":
    query = "Alzheimer's"
# Search PubMed
search results = search pubmed(query)
# Print the total articles
total articles = int(search results["Count"])
print(f"Total number of articles related to Alzheimer's:
{total articles}")
id list = search results["IdList"][:10]
pubmed records = fetch pubmed details(id list)
# Parse and print authors
authors_list = parse_pubmed_records(pubmed_records)
# Print authors
```

```
for i, authors in enumerate(authors_list, 1):
    print(f"\nAuthors for Article {i}:")
    for author in authors:
       print(f" - {author}")
```

```
PS D:\Acadamic\4 th year\2nd sem\CB\Lab 1> python -u "d:\Acadamic\4 th year\2nd sem\CB\Lab 1\3.2.py"
Total number of articles related to Alzheimer's: 223336
Authors for Article 1:
 - Mitsunaga S
 - Fujito N
 - Nakaoka H
 - Imazeki R
 - Nagata E
- Inoue I
Authors for Article 2:
 - Sajedi S
 - Ebrahimi G
- Roudi R
- Mehta I
- Heshmat A
 - Samimi H
 - Kazempour S
 - Zainulabadeen A
- Docking TR
 - Arora SP
 - Cigarroa F
 - Seshadri S
 - Karsan A
 - Zare H
Authors for Article 3:
```

40. Write a Biopython-program that finds CpG-islands from a given DNA-sequence.

```
41. from Bio.Seq import Seq
42. from Bio.SeqUtils import nt_search
43. dna_seq = Seq(input("Enter the DNA sequence: "))
44. dna_seq_str = str(dna_seq)
45.
46. def find_cpg_islands(sequence) :
47. cpg_positions = nt_search (sequence, "CG") [1:]
48. islands = []
```

```
current_island = []
49.
50.
             for pos in cpg_positions:
51.
                 if not current_island or pos == current_island[-1] + 1:
52.
                     current_island.append (pos)
53.
                 else:
54.
                     islands.append (current island)
55.
                     current_island = [pos]
56.
             islands.append (current_island)
57.
             return islands
         cpg_islands = find_cpg_islands(dna_seq_str)
58.
         print("CpG Islands found in:", cpg_islands)
59.
```

```
PS D:\Acadamic\4 th year\2nd sem\CB\Lab 1> python -u "d:\Acadamic\4 th year\2nd sem\CB\Lab 1\3.3.py"
Enter the DNA sequence: GTGGCCATTGTAATGGGCCGCTGAAAGGGTGCCCGATAG

CpG Islands found in: [[18], [33]]
PS D:\Acadamic\4 th year\2nd sem\CB\Lab 1> []
```