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APPENDIX A: PROTOCOLS FOR GENETIC DATA PREPARATION

A.1 PED files for 2 combined genotype data (used for ADMIXTURE analysis)

1. Prepare matching label files from existing genotype files. If the genetic data must be obtained from database, prepare another file for labels. ADMIXTURE does not incorporate labels but they are needed after the ancestral estimation is finished.

2. Combine 2 separate genetic files or check one separate genetic file against the SNP database. Write the matching (overlapping) SNP in a separate file.

3. Check for strand compatibility. Some SNP genotypes are of the complimentary strand. This would pose problems in encoding. Use one genetic source as the main source and change the genotypes in another file into the same complimentary bases.

4. Write the resulting combined genotype files for future use. This provides flexibility in including or excluding certain populations. Encoded genotype files cannot be used after populations have been included or excluded from the pool of samples.

5. Calculate major and minor alleles in order to assign numeric codes for the genotypes. (1 1 = homozygous wild type, 1 2 = heterozygous variance, 2 2 = homozygous variance, 0 0 = missing data)

6. Transpose genotype matrix as ped files have individuals as rows and SNP genotypes as columns while original genotype files are usually formatted vice versa.

7. Add 6 additional mandatory columns at the front of each row. These six columns will be omitted in ADMIXTURE ancestral estimation.

A.2 Encoded genotype files for PCA

1. Prepare headers into a separate file. Transpose so that the header become a one-column list.

2. Encode the genotype file into PCA format (0 = wild type, 1 = heterozygous variance, 2 = homozygous variance, 0 = missing data).

A.3 Files for AWClust

1. Prepare a header file or use an existing header file.

2. Encode the genotype file into AWClust format (0 = wild type, 1 = heterozygous variance, 2 = homozygous variance, 0 = missing data).

3. Concatenate the encoded genotype with the prepared header.

APPENDIX B: PYTHON CODES

The codes listed in this section are example of the workflow and logics used to construct the codes and prepare data. Real codes are adjusted according to different format of acquired genetic files. For example, some files require changes in header row read (present/absent) or transposing and some others might not need to be queried from database.

B.1 Querying and encoding into ped file format

```
#!/usr/bin/env python
import sys
import MySQLdb

def encode_genotype(gty) :
    alleles = ['A','T','G','C']
    allele_count = {}
    mj_allele = ''
    mj_count = 0
    mn_allele = ''
    mn_count = 0
    for allele in alleles :
        count = gty.count(allele)
        #allele_count[allele] = count
        if count > 0 :
            if count > mj_count :
                mn_count = mj_count
                mn_allele = mj_allele
                mj_count = count
                mj_allele = allele
            else :
                mn_count = count
                mn_allele = allele
    #return major_allele, minor_allele
    encode_gty = gty.replace(mj_allele + mj_allele,
'0').replace(mj_allele + mn_allele, '1').replace(mn_allele +
mj_allele, '1').replace(mn_allele + mn_allele, '2').replace('NN', '-1')
    return encode_gty.split(',')

def transpose(matrix) :
    transpose_matrix = []
    for i in xrange(len(matrix[0])) :
        for row in matrix :
            try :
                transpose_matrix[i].append(row[i])
            except :
                transpose_matrix.append([row[i]])
    return transpose_matrix

def query(sql) :
    cs.execute(sql)
    return cs.fetchall()

file_name = sys.argv[1]
ind_file = sys.argv[2]

ind_dict = {}
fd = open(ind_file)
fd.readline() #read one line to skip the header
lines = fd.readlines()
for line in lines :
```

```

        sp_line = line.strip().split(',')
        ind_dict[sp_line[0]] = sp_line[1].replace('!', '').replace('
', '_')
    fd.close()

    fd = open(file_name)
    header = fd.readline().strip().split(',')[1:]
    lines = fd.readlines()
    #open database
    db = MySQLdb.connect(user='thaisnp2', passwd='thaisnp2',
db='thaisnp_b129', host='bioinfo.biotec.or.th')
    cs = db.cursor()

    gty = []
    for line in lines :
        # Old gty
        tmp = line.strip().split(',')
        snpid = tmp[0][2:]
        tmp_gty = ','.join(tmp[1:])
        # Query ThaiSNP2 gty
        sql = '''SELECT genotype, allele_A, allele_B
                FROM ThaiSNP2_Genotype
                LEFT JOIN ThaiSNP2_Affymatrix ON
ThaiSNP2_Genotype.rssnp_id = ThaiSNP2_Affymatrix.rssnp_id
                WHERE ThaiSNP2_Genotype.rssnp_id = %d''' % int(snpid)
        result = query(sql)[0]
        thaisnp_gty = result[0].replace('A',
result[1]).replace('B', result[2])
        tmp_gty += ',' + thaisnp_gty
        encode_gty = encode_genotype(tmp_gty)
        gty.append((encode_gty))

    transpose_gty = transpose(gty)

    #print 'Gty : ', len(gty), len(gty[0])
    #print 'Transpose : ', len(transpose_gty), len(transpose_gty[0])
    for tmp in transpose_gty :
        ped = '1 1 0 0 2 1 '
        for i in range(len(tmp)) :
            if tmp[i] == '-1' :
                a = '0 0'
            elif tmp[i] == '0' :
                a = '1 1'
            elif tmp[i] == '1' :
                a = '1 2'
            elif tmp[i] == '2' :
                a = '2 2'
            ped += a + ' '
        print ped.rstrip()
    fd.close()

    ind_content = []
    for ind in header :
        ind_content.append(ind_dict[ind] + '\n')
    for i in range(32) :
        ind_content.append('Thai2\n')
    fd = open('ind_pop.txt', 'w')
    fd.writelines(ind_content)
    fd.close()
    db.close()

```



B.2 Checking for matching strands with querying

```
#!/usr/bin/env python
import MySQLdb

def query(sql) :
    cs.execute(sql)
    return cs.fetchall()

#open database
db = MySQLdb.connect(user='thaisnp2', passwd='thaisnp2',
db='thaisnp_b129', host='bioinfo.biotec.or.th')
cs = db.cursor()
gty_dict = {}
fd = open('JHS_Genotype.csv')
header = fd.readline()
lines = fd.readlines()
for line in lines :
    sp_line = line.strip().split(',')
    snpid   = sp_line[0][2:]
    gty     = ','.join(sp_line[1:])
    gty_dict[snpid] = gty
fd.close()

fd = open("match_sorted.csv")
header = fd.readline() #read header
lines = fd.readlines() #read from the line next to header line

#snpList = []
print header.strip()
for line in lines :
    sp_line = line.strip().split(',')
    snpid   = sp_line[0][2:]
    #snpList.append(snpid)
    allele = sp_line[3] + '/' + sp_line[4]
    sql = '''select concat(allele_A, '/', allele_B) from
ThaiSNP2_Affymatrix where rssnp_id=%d''' % int(snpid)
    result = query(sql)[0][0]
    if allele <> result :
        if result.find(sp_line[3]) < 0 :
            compl_base = {'A':'T','T':'A','G':'C','C':'G'}
            allele1, allele2 = compl_base[sp_line[3]],
compl_base[sp_line[4]]
            replace_gty = {sp_line[3]+sp_line[3]:allele1+allele1,
sp_line[3]+sp_line[4]:allele1+allele2,
sp_line[4]+sp_line[3]:allele2+allele1,sp_line[4]+sp_line[4]:allele2+al
lele2}

            for gty in replace_gty :
                gty_dict[snpid] = gty_dict[snpid].replace(gty,
replace_gty[gty])
            print 'rs%s,%s' % (snpid, gty_dict[snpid])
        fd.close()

    db.close()
```

B.3 Checking for matching SNPs from 2 files, changing into compatible strands

```
#get file2 gty = _newThai
fd = open(gtyFile2)
```

```

file2Header = fd.readline()
lines = fd.xreadlines()
for line in lines :
    split_line = line.strip().split(',')
    snpID = split_line[0]
    if file1gty_dict.has_key(snpID) == True :
        gty = ','.join(split_line[1:])
        obs2 = countAllele(gty) # 'A,T'
        obs1 = countAllele(file1gty_dict[snpID]) # 'T,A'
        file2gty_dict[snpID] = gty
        if obs2 <> obs1 :
            if obs2.find(obs1[0])<0 and obs2.find(obs1[2])<0 :
                try :
                    allele1 = compl_dict[obs2[0]]
                    allele_a = obs2[0]
                except :
                    allele1 = '-'
                    allele_a = '-'
                try :
                    allele2 = compl_dict[obs2[2]]
                    allele_b = obs2[2]
                except :
                    allele2 = '-'
                    allele_b = '-'
                replace_dict = {allele_a+allele_a:
allele1+allele1,\
                                allele_a+allele_b:
allele1+allele2,\
                                allele_b+allele_a:
allele2+allele1,\
                                allele_b+allele_b:
allele2+allele2}
                for key in replace_dict :
                    file2gty_dict[snpID]=
file2gty_dict[snpID].replace(key,replace_dict[key])
                    #print snpID, obs1, obs2, replace_dict,
file1gty_dict[snpID], file2gty_dict[snpID]
                    print '%s,%s,%s' %(snpID, file1gty_dict[snpID],
file2gty_dict[snpID])

                    #if len(obs1)>2 or len(obs2)>2 :
                    #    print '%s,%s, %s' %(snpID, obs1, obs2)
fd.close()

```

B.4 Combining overlapping SNP data from 2 genotype files

```

#!/usr/bin/env python
import sys

if __name__ == '__main__' :
    thal_file = sys.argv[1]
    depress_file = sys.argv[2]
    fd = open(thal_file)
    thal_header = fd.readline()
    thal_gty = {}
    lines = fd.xreadlines()
    for line in lines :
        sp_line = line.strip().split(',')
        snp_id = sp_line[0]
        gty = ','.join(sp_line[1:])
        thal_gty[snp_id] = gty

```

```
fd.close()
fd = open(depress_file)
depress_header = fd.readline()
depress_gty = {}

# \ is for new line
print '%s,%s' % (thal_header.strip(), \
                ','.join(depress_header.split(',') [1:]))
lines = fd.xreadlines()
for line in lines :
    sp_line = line.strip().split(',')
    snp_id = sp_line[0]
    if thal_gty.has_key(snp_id) == 1 :
        # matched
        gty = ','.join(sp_line[1:])
        print '%s,%s,%s' % (snp_id, thal_gty[snp_id], gty)
fd.close()
```

CURRICULUM VITAE

NAME Ms. Sattara Hattirat

DATE OF BIRTH 28 September 1979

EDUCATIONAL RECORD

High School	Trium Udom Suksa School, 1996
Bachelor's Degree	Bachelor of Arts (Chinese Language) Chulalongkorn University, 2000
	Bachelor of Science (Biology) Ramkhamhaeng University, 2008
Master's Degree	Master of Science (Bioinformatics and Systems Biology) King Mongkut's University of Technology Thonburi, 2011

SCHOLARSHIP Scholarship from Bioinformatics Program,
2009-2011

PUBLICATIONS

Sattara Hattirat, Chumpol Ngamphiw, Anuncai Assawamakin, Jonathan Chan and Sissades Tongsima. "Catalog of Genetic Variations (SNPs and CNVs) and Analysis Tools for Thai Genetic Studies". **Communications in Computer and Information Science** 115. J. H. Chan, Y-S. Ong and S-B. Cho (Eds). Springer (2010)

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Kansri Boonprakob and Sattara Hattirat (eds.). **Climate Change and Thailand**, Green Peace (2006)

EMPLOYMENT RECORD

Deputy Director
Thai AIDS Treatment Action Group, December 2011 - present

King Mongkut's University of Technology Thonburi
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Name...Ms. Sattara ...Surname...Hattirat...Student Number...52401712...who is a student of King's Mongkut's University of Technology Thonburi (KMUTT) in...Master's Degree... Program...Bioinformatics and Systems Biology... Field of Study... Bioinformatics and Systems Biology... Faculty/School of...Bioresources and Technology and School of Information Technology... Home Address...59/663... Tambon...Samwatawanok... District...Klong Samwa... Province...Bangkok... Postal Code...10510... Country...Thailand...

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(Assoc. Prof. Dr. Nipon Charoenkitkarn)

Signature [Signature] Witness
(Asst. Prof. Dr. Jonathan Hoyin Chan)

Signature Marasri Ruengjitchatchawalya Witness
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