

# Practical 1 - WORKING DRAFT

Samuel Daulton, Taylor Killian, and Andrew Petschek (ML Marauders)

February 11, 2016

## 1 Technical Approach

### 1.1 Initial Attempts

Our initial approach to expand on the sample code provided to us for this machine learning problem was to implement Principal Component Analysis (PCA) to gain a better understanding of the interdependencies between features and to effectively reduce our feature set. We also performed some basic feature engineering upfront to couple with the PCA. There is a hypothesis that the number of branches influences the gap between HOMO and LUMO levels, and we pulled relevant compounds that were highlighted in a theoretical study on this topic conducted by Padtaya Pukpayat of the University of Nevada Las Vegas. This specific feature aggregates the number of active features, meaning if there are more features that are active in a compound, then there may be a higher orbital overlap and, thus, a lower HOMO-LUMO gap. A final initial feature we added was the presence of a benzene ring in the compound; we could identify a benzene ring which appeared to be more conjugated (i.e., closer together in energy). We combined this initial feature generation with PCA, and turned to the existing methods to see if it we gained improvement over baseline. The results were promising, but we decided additional, more robust features needed to be engineered for improved impact.

### 1.2 Feature Engineering

To create new features, we turned to the RDKit. Specifically, we leveraged molecule object methods that returned integers. We felt integer outputs would work best with our modeling efforts, and we avoided methods that would output non-integers (e.g., strings and vectors). 114 new features were created in this way that spanned four main areas: general feature generation (e.g., number of branches, number of bonds), descriptors (e.g., molecular weight, number of valence electrons), calculation of Lipinski parameters, and

fragment descriptors (e.g., number of aliphatic carboxylic acids). As stated above, these methods act on molecule objects, which take up a lot of memory, so our technical approach to feature generation was to divide the processing into four separate parts, which were ultimately merged together at the end.

In addition to using RDKit, we implemented our own algorithms to create additional features. We had a hypothesis that there likely existed interactions between features that would lead to larger HOMO-LUMO gaps. Thus, in addition to including a linear basis for each feature, we aimed to also include the product of pairs of features as well. Since we had well over 100 features, we decided we would look at the combinations between the top 25 most important features. To determine the top 25, we turned to Random Forests, which can output the importance weighting of each feature in our feature space. Using this regression output, we took the top 25 features and created  $25C2$  (or 300) new features, which were the products of two features within a unique combinatoric pair.

A final method for feature generation we used was based on logistic regression. Performing first some pre-processing, we rounded each gap value to the nearest half integer which we set as the labels we wished to predict. We ultimately used these labels as categorical features that were included in our overall feature set.

### 1.3 Modeling Techniques

We not only built off of the simple linear regression and random forest model classes provided to us, but also approached this machine learning problem from other modeling angles as well. Across all methods, to prevent overfitting out-of-sample data sets when evaluating various hyperparameters for estimators, we split our training set into a training set and a validation set (67-33 split). This way, we avoided any knowledge of the test set leaking into our models and hurting their ability to generalize to new data. We had initially contemplated a cross-validation loop, where the creation of discrete training and validation sets would no longer be necessary to increase training sample size. However, since we had roughly one million samples to begin with, we decided we would have sufficient data to carry out our modeling using discrete training and validation sets. When estimating hyperparameters for each of our models below, we employed a grid search. Since we did not know which hyperparameter values may minimize our loss functions across models, we ran each model several times across an array of hyperparameter values ranging from small to large. At the end of the loop, we selected the hyperparameter value(s) that yielded a minimum loss when tested against our validation set and then re-trained the model on the entire training set (including the validation set).

Examples of this grid search method were evident in three new methods we employed: Lasso Regression, Ridge Regression, and Elastic Net Regression. We felt attempting to use

regularization techniques such as these would be a good idea as they would penalize very complex models and would likely fit out of sample data better. Lasso regression is a linear model with a regularization term (i.e., L1 prior). The constant,  $\alpha$  that multiplies the L1 term is a hyperparameter that can range between 0 and 1. We noticed that if  $\alpha = 0$ , then we have reduced this regression back to a simple linear model without regularization. Ridge Regression is similar to Lasso, except it uses a L2 prior. Similarly, if  $\alpha = 0$  in our Ridge Regression, we have reverted back to linear least squares. Elastic Net can be thought of as a combination of Lasso and Ridge Regression in that it combines both L1 and L2 priors as regularization terms. There are two hyperparameters in Elastic Net which measure the relationship between the L1 and L2 penalties,  $\alpha$  and a ratio term. Here, if  $\alpha = 0$ , we again have a linear least squares regression and if the ratio = 0 we have only Ridge Regression and if the ratio = 1 we have only Lasso Regression. Thus, we employed grid search to determine the right balance needed for loss minimization. With similar results across all three, we ultimately landed on using Lasso Regression for our final submission. Lasso regression is able to zero out features completely while Ridge Regression, for example, cannot. For this reason, we decided Lasso Regression may be a better bet to generalize to the new test data set moving forward since we had hundreds of features.

In addition to linear modeling, we also employed ensemble methods, including Random Forests and Extra Trees Regressors, which can be thought of as a close alternative to a Random Forest. In our Random Forest regression we set the number of features per tree to be equal to  $\frac{1}{\sqrt{p}}$  and used a grid search to compare number of estimators between 10 and 128. We had read in THIS PAPER that a good range of estimators to be 64-128, as anything more would result in additional computation time with minimal gains, which is what dictated our rationale for the grid search bounds. We repeated this similar approach for Extra Trees Regressors. As stated above, Extra Trees is similar to Random Forests, where Extra Trees randomly draws estimators from the entire training set instead of a bootstrap sample and these splits are chosen at random. This additional set of randomness can lead to a reduction in variance and an increase in bias. For this reason, while we did run tests using Extra Trees Regressors, we ultimately decided to use Random Forests instead for their potentially lower bias and better ability to generalize to new data.

A few extensions were used across the methods above. The first involves PCA. We had initially thought that the use of PCA would be beneficial, as it would reveal the most important features in our feature space up front. We thought that this would improve performance. We found, however, that this did not seem to be the case. Reducing our feature space using PCA did not seem to improve our predictions (and in some cases it even led to negative impact). We also considered standardizing all non-binary features to ensure they are normally distributed. We felt normally distributed data would potentially lead to more accurate predictions. This too, like PCA, seemed to lead to worse results. Thus, we ultimately decided not to standardize our data. Finally, we tested models as we added new features along the way. This was a fluid process of adding features we felt to

Model	RMSE (Public Score)
Extra Trees Regression and PCA (n=60)	0.15337
Lasso and PCA (n=60)	???
Extra Trees Regression and PCA (n=45)	0.22972
Extra Trees Regression with AdaBoost	0.29581
Linear Regression	0.3

Table 1: Model Results

be better predictors; we did not have all features all at once at the start.

DO WE WANT TO ADD ADABOOST, GRADIENT DESCENT? Finally, we also built upon this ensemble method by incorporating AdaBoost. AdaBoost fits a sequence of weak learners on repeatedly modified versions of the training set. The predictions are assigned weights which update through a loop. These weights were measured according to two loss functions: linear and squared error. Together, Adaboost with Extra Trees were used.

## 2 Results

We attempted many permutations of the above modeling techniques spanning the inclusion of various feature sets, models, hyperparameters, PCA, standardization etc... We found our best results to be when we used our full feature set and a Random Forrest Regressor with X trees. See table 1 for our RMSE results.

## 3 Discussion

This was an exciting opportunity to be able to work with a complex dataset on a machine learning problem with real-world implications. We approached this task as a prediction problem rather than an estimation problem. While prediction and estimation are similar, there is nuance. While estimation concerns itself more with the structural properties of the model and a more physical understanding and interpretation of coefficients, prediction cares most about loss minimization and the models ability to generalize to new data. Thus, we approached this problem with an open mind and a determination to find a minimal loss score without preference for the type of model we employed.

While we did not find PCA to help much with our predictions, it did remind us of a hallmark of scientific computing. PCA reduced our feature set and greatly reduced computation time. While computation time is not a benchmark we are graded on, it is an

integral component to scientific computing. As data sets become large, computation time required increases. Thus, one's ability to effectively reduce the time it takes to fit a model without sacrificing accuracy is highly valued. While PCA was not effective here, we could have considered additional parallel processing techniques (e.g., multi processing, multi threading) to speed up computation time.

We found feature engineering to be a critical component to our success. Our modeling output is only as good as the input we provide. We found that our feature engineering efforts provided our model with regression input it needed for better training. With each iteration of feature generation, we found our results to increase.

Further, we believe an ensemble method of models to be the most robust with such a complicated data set.

add more

```
# coding: utf-8

# In[1]:

get_ipython().magic(u'matplotlib inline')
import pandas as pd
import numpy as np
import scipy as sp
import matplotlib.pyplot as plt
from sklearn.linear_model import LinearRegression, Lasso, LassoCV, ElasticNetCV, SGD
from sklearn.ensemble import RandomForestRegressor, ExtraTreesRegressor, AdaBoostRegressor
from sklearn.decomposition import PCA
from sklearn.pipeline import Pipeline
from sklearn.grid_search import GridSearchCV
from sklearn.cross_validation import train_test_split, KFold
from sklearn.metrics import mean_squared_error
from rdkit import DataStructs, Chem
from rdkit.Chem import AllChem, Descriptors, Lipinski, Fragments, rdMolOps
from rdkit.ML.Cluster import Butina

# # *IF FEATURE ENGINEERING IS DONE: DO NOT START HERE*
# # *SKIP TO THE START OF SECTION IV: MODEL SELECTION*

# # I. Exploratory Data Analysis

# In[ ]:

# Read in training data
```

```

df_train = pd.read_csv("train.csv")

# In[ ]:

df_train.head()

# In[ ]:

plt.hist(df_train.gap.values, bins=50)

# ### Note:
# I see some outliers around -1.5. Having negative values makes no sense, since th
==> Remove them

# In[ ]:

# Remove negative HOMO-LUMO gaps
df_train = df_train[df_train['gap'] >= 0]

# In[ ]:

print len(df_train)

# In[ ]:

# Plot histogram of gaps without negative values
plt.hist(df_train.gap.values, bins=50)

# # II. Training Data Processing and Feature Engineering
# 1. Drop all columns except smiles and gaps
# 2. Add "key" column, partition the data, and save those partitions
# 3. Remove all dataframes from memory
# 4. Load each partition (1 by 1), perform feature engineering, and save each parti
# 5. Reaggregate all partitions into one df, merge add binary-valued features from

# ### 1. Drop all columns except smiles and gaps

# In[ ]:

# remove all columns except smiles and gaps

```

```

df_train = df_train.drop(df_train.columns[range(1,257)], axis=1)
df_train.head()

# ### 2. Add "key" column, partition the data, and save those partitions

# In[ ]:

# add key column
df_train['key'] = df_train.index

# In[ ]:

df_train.head()

# In[ ]:

num_parts = 10
df_train_parts = np.array_split(df_train, num_parts)
for i in xrange(len(df_train_parts)):
    df_train_parts[i].to_csv('smiles_gaps_keys_df_train_part_'+str(i)+'.csv', index=False)

# ### 3. Remove all dataframes from memory

# In[ ]:

del df_train
del df_train_parts

# ### 4. Load each partition (1 by 1), perform feature engineering, and save each partition
#
# (http://machinelearningmastery.com/discover-feature-engineering-how-to-engineer-features-for-machine-learning/)
#
# ##### Number of Branches
# The idea is that the number of [branches](https://en.wikipedia.org/wiki/Simplification)
#
# ##### Note: We pull the specific compounds of interest from [this poster.](http://www.chemrxiv.org/abstract/2016060001)
#
# ##### Benzene Ring
# The last feature is determining, from the SMILES encoding whether or not there is a benzene ring
#
# ##### Number of Double Bonds

```

```

#
# ##### Additional Features engineered using from RDKit library

# In[ ]:

for i in xrange(num_parts):
    df_train_part = pd.read_csv('smiles_gaps_keys_df_train_part_'+str(i)+'.csv')

    # Apply feature engineering to train data
    num_branches = df_train_part.smiles.apply(lambda x: x.count('('))
    has_benzothiophene = df_train_part.smiles.apply(lambda x: min(1,x.count('s2c1cc
    has_carbazole = df_train_part.smiles.apply(lambda x: min(1,x.count('c1ccc2c(c1)
    has_fluorene = df_train_part.smiles.apply(lambda x: min(1,x.count('c1ccc-2c(c1)
    num_double_bonds = df_train_part.smiles.apply(lambda x: x.count('='))
    df_train_part['num_branches'] = num_branches
    df_train_part['has_benzothiophene'] = has_benzothiophene
    df_train_part['has_carbazole'] = has_carbazole
    df_train_part['has_fluorene'] = has_fluorene
    df_train_part['num_double_bonds'] = num_double_bonds
    print "Part {}: finished initial feat gen".format(i)
    # RDKit Feature Engineering
    # Generate molecule objects
    molecules = df_train_part.smiles.apply(lambda x: Chem.MolFromSmiles(x))
    print "Part {}: finished molecule generation".format(i)
    # Generate Features
    df_train_part['avg_molecular_weight'] = molecules.apply(lambda x: Descriptors.M
    print "Part {}: finished first feature".format(i)
    df_train_part['exact_molecular_weight'] = molecules.apply(lambda x: Descriptors
    df_train_part['avg_molecular_weight_ignore_hydrogen'] = molecules.apply(lambda
    df_train_part['num_valence_electrons'] = molecules.apply(lambda x: Descriptors.
    df_train_part['num_radical_electrons'] = molecules.apply(lambda x: Descriptors.
    df_train_part['formal_charge'] = molecules.apply(lambda x: rdmolops.GetFormalCh
    df_train_part['sssr'] = molecules.apply(lambda x: rdmolops.GetSSSR(x))
    print "Part {}: finished Descriptors and rdmolops".format(i)
    df_train_part['fraction_csp3'] = molecules.apply(lambda x: Lipinski.FractionCSP
    df_train_part['num_aliphatic_carbocycles'] = molecules.apply(lambda x: Lipinski
    df_train_part['num_aliphatic_heterocycles'] = molecules.apply(lambda x: Lipinsk
    df_train_part['num_aliphatic_rings'] = molecules.apply(lambda x: Lipinski.NumAl
    df_train_part['num_aromatic_carbocycles'] = molecules.apply(lambda x: Lipinski.
    df_train_part['num_aromatic_heterocycles'] = molecules.apply(lambda x: Lipinski
    df_train_part['num_aromatic_rings'] = molecules.apply(lambda x: Lipinski.NumAro
    df_train_part['num_saturated_carbocycles'] = molecules.apply(lambda x: Lipinski
    df_train_part['num_saturated_heterocycles'] = molecules.apply(lambda x: Lipinsk
    df_train_part['num_saturated_rings'] = molecules.apply(lambda x: Lipinski.NumSa
    df_train_part['num_nh_oh'] = molecules.apply(lambda x: Lipinski.NHOHCount(x))
    df_train_part['num_num_rotatable_bonds'] = molecules.apply(lambda x: Lipinski.M

```



```

df_train_part['num_heteroatoms'] = molecules.apply(lambda x: Lipinski.NumHetero
df_train_part['num_h_acceptors'] = molecules.apply(lambda x: Lipinski.NumHAccep
df_train_part['num_h_donors'] = molecules.apply(lambda x: Lipinski.NumHDonors(x)
df_train_part['ring_count'] = molecules.apply(lambda x: Lipinski.RingCount(x))
print "Part {}: finished Lipinski".format(i)
# See Parsing_methods_from_rdk_source.ipynb
df_train_part['fr_Al_COO'] = molecules.apply(lambda x: Fragments.fr_Al_COO(x))
df_train_part['fr_Al_OH'] = molecules.apply(lambda x: Fragments.fr_Al_OH(x))
df_train_part['fr_Al_OH_noTert'] = molecules.apply(lambda x: Fragments.fr_Al_OH
df_train_part['fr_ArN'] = molecules.apply(lambda x: Fragments.fr_ArN(x))
df_train_part['fr_Ar_COO'] = molecules.apply(lambda x: Fragments.fr_Ar_COO(x))
df_train_part['fr_Ar_N'] = molecules.apply(lambda x: Fragments.fr_Ar_N(x))
df_train_part['fr_Ar_NH'] = molecules.apply(lambda x: Fragments.fr_Ar_NH(x))
df_train_part['fr_Ar_OH'] = molecules.apply(lambda x: Fragments.fr_Ar_OH(x))
df_train_part['fr_COO'] = molecules.apply(lambda x: Fragments.fr_COO(x))
df_train_part['fr_COO2'] = molecules.apply(lambda x: Fragments.fr_COO2(x))
df_train_part['fr_C_O'] = molecules.apply(lambda x: Fragments.fr_C_O(x))
df_train_part['fr_C_O_noCOO'] = molecules.apply(lambda x: Fragments.fr_C_O_noCO
df_train_part['fr_C_S'] = molecules.apply(lambda x: Fragments.fr_C_S(x))
df_train_part['fr_HOCCN'] = molecules.apply(lambda x: Fragments.fr_HOCCN(x))
df_train_part['fr_Imine'] = molecules.apply(lambda x: Fragments.fr_Imine(x))
df_train_part['fr_NHO'] = molecules.apply(lambda x: Fragments.fr_NHO(x))
df_train_part['fr_NH1'] = molecules.apply(lambda x: Fragments.fr_NH1(x))
df_train_part['fr_NH2'] = molecules.apply(lambda x: Fragments.fr_NH2(x))
df_train_part['fr_N_O'] = molecules.apply(lambda x: Fragments.fr_N_O(x))
df_train_part['fr_Ndealkylation1'] = molecules.apply(lambda x: Fragments.fr_Nde
df_train_part['fr_Ndealkylation2'] = molecules.apply(lambda x: Fragments.fr_Nde
df_train_part['fr_Nhpyrrole'] = molecules.apply(lambda x: Fragments.fr_Nhpyrrol
df_train_part['fr_SH'] = molecules.apply(lambda x: Fragments.fr_SH(x))
df_train_part['fr_aldehyde'] = molecules.apply(lambda x: Fragments.fr_aldehyde(
df_train_part['fr_alkyl_carbamate'] = molecules.apply(lambda x: Fragments.fr_al
df_train_part['fr_alkyl_halide'] = molecules.apply(lambda x: Fragments.fr_alkyl
df_train_part['fr_allylic_oxid'] = molecules.apply(lambda x: Fragments.fr_allyl
df_train_part['fr_amide'] = molecules.apply(lambda x: Fragments.fr_amide(x))
df_train_part['fr_amidine'] = molecules.apply(lambda x: Fragments.fr_amidine(x)
df_train_part['fr_aniline'] = molecules.apply(lambda x: Fragments.fr_aniline(x)
df_train_part['fr_aryl_methyl'] = molecules.apply(lambda x: Fragments.fr_aryl_m
df_train_part['fr_azide'] = molecules.apply(lambda x: Fragments.fr_azide(x))
df_train_part['fr_azo'] = molecules.apply(lambda x: Fragments.fr_azo(x))
df_train_part['fr_barbitur'] = molecules.apply(lambda x: Fragments.fr_barbitur(
df_train_part['fr_benzene'] = molecules.apply(lambda x: Fragments.fr_benzene(x)
df_train_part['fr_benzodiazepine'] = molecules.apply(lambda x: Fragments.fr_ben
df_train_part['fr_bicyclic'] = molecules.apply(lambda x: Fragments.fr_bicyclic(
df_train_part['fr_diazo'] = molecules.apply(lambda x: Fragments.fr_diazo(x))
df_train_part['fr_dihydropyridine'] = molecules.apply(lambda x: Fragments.fr_di
df_train_part['fr_epoxide'] = molecules.apply(lambda x: Fragments.fr_epoxide(x)

```

```

df_train_part['fr_ester'] = molecules.apply(lambda x: Fragments.fr_ester(x))
df_train_part['fr_ether'] = molecules.apply(lambda x: Fragments.fr_ether(x))
df_train_part['fr_furan'] = molecules.apply(lambda x: Fragments.fr_furan(x))
df_train_part['fr_guanido'] = molecules.apply(lambda x: Fragments.fr_guanido(x))
df_train_part['fr_halogen'] = molecules.apply(lambda x: Fragments.fr_halogen(x))
df_train_part['fr_hdrzine'] = molecules.apply(lambda x: Fragments.fr_hdrzine(x))
df_train_part['fr_hdrzone'] = molecules.apply(lambda x: Fragments.fr_hdrzone(x))
df_train_part['fr_imidazole'] = molecules.apply(lambda x: Fragments.fr_imidazol
df_train_part['fr_imide'] = molecules.apply(lambda x: Fragments.fr_imide(x))
df_train_part['fr_isocyan'] = molecules.apply(lambda x: Fragments.fr_isocyan(x))
df_train_part['fr_isothiocyan'] = molecules.apply(lambda x: Fragments.fr_isothi
df_train_part['fr_ketone'] = molecules.apply(lambda x: Fragments.fr_ketone(x))
df_train_part['fr_lactam'] = molecules.apply(lambda x: Fragments.fr_lactam(x))
df_train_part['fr_lactone'] = molecules.apply(lambda x: Fragments.fr_lactone(x))
df_train_part['fr_methoxy'] = molecules.apply(lambda x: Fragments.fr_methoxy(x))
df_train_part['fr_morpholine'] = molecules.apply(lambda x: Fragments.fr_morphol
df_train_part['fr_nitrile'] = molecules.apply(lambda x: Fragments.fr_nitrile(x))
df_train_part['fr_nitro'] = molecules.apply(lambda x: Fragments.fr_nitro(x))
df_train_part['fr_nitro_arom'] = molecules.apply(lambda x: Fragments.fr_nitro_a
df_train_part['fr_nitro_arom_nonortho'] = molecules.apply(lambda x: Fragments.f
df_train_part['fr_nitroso'] = molecules.apply(lambda x: Fragments.fr_nitroso(x))
df_train_part['fr_oxazole'] = molecules.apply(lambda x: Fragments.fr_oxazole(x))
df_train_part['fr_oxime'] = molecules.apply(lambda x: Fragments.fr_oxime(x))
df_train_part['fr_para_hydroxylation'] = molecules.apply(lambda x: Fragments.fr
df_train_part['fr_phenol'] = molecules.apply(lambda x: Fragments.fr_phenol(x))
df_train_part['fr_phenol_noOrthoHbond'] = molecules.apply(lambda x: Fragments.f
df_train_part['fr_phos_acid'] = molecules.apply(lambda x: Fragments.fr_phos_aci
df_train_part['fr_phos_ester'] = molecules.apply(lambda x: Fragments.fr_phos_es
df_train_part['fr_piperdine'] = molecules.apply(lambda x: Fragments.fr_piperdin
df_train_part['fr_piperzine'] = molecules.apply(lambda x: Fragments.fr_piperzin
df_train_part['fr_priamide'] = molecules.apply(lambda x: Fragments.fr_priamide(
df_train_part['fr_prisulfonamd'] = molecules.apply(lambda x: Fragments.fr_prisu
df_train_part['fr_pyridine'] = molecules.apply(lambda x: Fragments.fr_pyridine(
df_train_part['fr_quatN'] = molecules.apply(lambda x: Fragments.fr_quatN(x))
df_train_part['fr_sulfide'] = molecules.apply(lambda x: Fragments.fr_sulfide(x))
df_train_part['fr_sulfonamd'] = molecules.apply(lambda x: Fragments.fr_sulfonam
df_train_part['fr_sulfone'] = molecules.apply(lambda x: Fragments.fr_sulfone(x))
df_train_part['fr_term_acetylene'] = molecules.apply(lambda x: Fragments.fr_ter
df_train_part['fr_tetrazole'] = molecules.apply(lambda x: Fragments.fr_tetrazol
df_train_part['fr_thiazole'] = molecules.apply(lambda x: Fragments.fr_thiazole(
df_train_part['fr_thiocyan'] = molecules.apply(lambda x: Fragments.fr_thiocyan(
df_train_part['fr_thiophene'] = molecules.apply(lambda x: Fragments.fr_thiophen
df_train_part['fr_unbrch_alkane'] = molecules.apply(lambda x: Fragments.fr_unbr
df_train_part['fr_urea'] = molecules.apply(lambda x: Fragments.fr_urea(x))
df_train_part['fr_ketone_Topliiss'] = molecules.apply(lambda x: Fragments.fr_ket

```

```

    print "Part {}: finished Fragments".format(i)
    df_train_part.to_csv('rdk_feat_eng_df_train_part_'+str(i)+'.csv', index=False)
    del df_train_part
    del molecules

# ### 5. Reaggregate all partitions into one df, merge add binary-valued features f

# In[ ]:

# Read in each feature engineered partition
df_train_parts = [pd.read_csv('rdk_feat_eng_df_train_part_'+str(i)+'.csv') for i in
# concatenate them into one df
df_train = pd.concat(df_train_parts)

# In[ ]:

# Read in old training
# Note: df_train_old has same as before partitioning
df_train_old = pd.read_csv("train.csv")
# Add key
df_train_old['key'] = df_train_old.index
# merge dataframes on 'key'
df_train = df_train.merge(df_train_old, on=["key"])
# Fix columns
df_train = df_train.drop(['gap_x'],axis=1)
df_train = df_train.drop(['key'],axis=1)
df_train = df_train.rename(columns = {'gap_y':'gap'})
df_train = df_train.drop(['smiles_x'],axis=1)
df_train = df_train.rename(columns = {'smiles_y':'smiles'})

# In[ ]:

df_train.head()

# In[ ]:

# Save df_train
df_train.to_csv('FINAL_train.csv', index=False)

# In[ ]:

```

```

# Remove df_train for now
del df_train_parts
del df_train_old
del df_train

# # III. Test Data Processing and Feature Engineering
# 1. Drop all columns except smiles and ids
# 2. Partition the data and save those partitions
# 3. Remove all dataframes from memory
# 4. Load each partition (1 by 1), perform feature engineering, and save each parti
# 5. Reaggregate all partitions into one df, merge add binary-valued features from

# In[ ]:

# Read in test data
df_test = pd.read_csv("test.csv")
df_test.head()

# ### 1. Drop all columns except smiles and ids

# In[ ]:

df_test = df_test.drop(df_test.columns[range(2,258)], axis=1)

# In[ ]:

df_test.head()

# ### 2. Partition the data and save those partitions

# In[ ]:

num_parts = 10

# In[ ]:

df_test_parts = np.array_split(df_test, num_parts)
for i in xrange(num_parts):
    df_test_parts[i].to_csv('smiles_and_ids_df_test_part_'+str(i)+'.csv', index=False)

```

```

# ### 3. Remove all dataframes from memory

# In[ ]:

del df_test
del df_test_parts

# ### 4. Load each partition (1 by 1), perform feature engineering, and save each p

# In[ ]:

for i in xrange(num_parts):
    df_test_part = pd.read_csv('smiles_and_ids_df_test_part_'+str(i)+'.csv')

    # Apply feature engineering to train data
    num_branches = df_test_part.smiles.apply(lambda x: x.count('('))
    has_benzothiophene = df_test_part.smiles.apply(lambda x: min(1,x.count('s2c1ccc
    has_carbazole = df_test_part.smiles.apply(lambda x: min(1,x.count('c1ccc2c(c1)c
    has_fluorene = df_test_part.smiles.apply(lambda x: min(1,x.count('c1ccc-2c(c1)C
    num_double_bonds = df_test_part.smiles.apply(lambda x: x.count('='))
    df_test_part['num_branches'] = num_branches
    df_test_part['has_benzothiophene'] = has_benzothiophene
    df_test_part['has_carbazole'] = has_carbazole
    df_test_part['has_fluorene'] = has_fluorene
    df_test_part['num_double_bonds'] = num_double_bonds
    print "Part {}: finished initial feat gen".format(i)
    # RDKit Feature Engineering
    # Generate molecule objects
    molecules = df_test_part.smiles.apply(lambda x: Chem.MolFromSmiles(x))
    print "Part {}: finished molecule generation".format(i)
    # Generate Features
    df_test_part['avg_molecular_weight'] = molecules.apply(lambda x: Descriptors.Mo
    print "Part {}: finished first feature".format(i)
    df_test_part['exact_molecular_weight'] = molecules.apply(lambda x: Descriptors.
    df_test_part['avg_molecular_weight_ignore_hydrogen'] = molecules.apply(lambda x
    df_test_part['num_valence_electrons'] = molecules.apply(lambda x: Descriptors.N
    df_test_part['num_radical_electrons'] = molecules.apply(lambda x: Descriptors.N
    df_test_part['formal_charge'] = molecules.apply(lambda x: rdmolops.GetFormalCha
    df_test_part['sssr'] = molecules.apply(lambda x: rdmolops.GetSSSR(x))
    print "Part {}: finished Descriptors and rdmolops".format(i)
    df_test_part['fraction_csp3'] = molecules.apply(lambda x: Lipinski.FractionCSP3
    df_test_part['num_aliphatic_carbocycles'] = molecules.apply(lambda x: Lipinski.
    df_test_part['num_aliphatic_heterocycles'] = molecules.apply(lambda x: Lipinski
    df_test_part['num_aliphatic_rings'] = molecules.apply(lambda x: Lipinski.NumAli

```

```

df_test_part['num_aromatic_carbocycles'] = molecules.apply(lambda x: Lipinski.N
df_test_part['num_aromatic_heterocycles'] = molecules.apply(lambda x: Lipinski.
df_test_part['num_aromatic_rings'] = molecules.apply(lambda x: Lipinski.NumArom
df_test_part['num_saturated_carbocycles'] = molecules.apply(lambda x: Lipinski.
df_test_part['num_saturated_heterocycles'] = molecules.apply(lambda x: Lipinski
df_test_part['num_saturated_rings'] = molecules.apply(lambda x: Lipinski.NumSat
df_test_part['num_nh_oh'] = molecules.apply(lambda x: Lipinski.NHOHCount(x))
df_test_part['num_num_rotatable_bonds'] = molecules.apply(lambda x: Lipinski.Nu
df_test_part['num_heteroatoms'] = molecules.apply(lambda x: Lipinski.NumHeteroa
df_test_part['num_h_acceptors'] = molecules.apply(lambda x: Lipinski.NumHAccept
df_test_part['num_h_donors'] = molecules.apply(lambda x: Lipinski.NumHDonors(x)
df_test_part['ring_count'] = molecules.apply(lambda x: Lipinski.RingCount(x))
print "Part {}: finished Lipinski".format(i)
# See Parsing_methods_from_rdk_source.ipynb
df_test_part['fr_Al_COO'] = molecules.apply(lambda x: Fragments.fr_Al_COO(x))
df_test_part['fr_Al_OH'] = molecules.apply(lambda x: Fragments.fr_Al_OH(x))
df_test_part['fr_Al_OH_noTert'] = molecules.apply(lambda x: Fragments.fr_Al_OH_
df_test_part['fr_ArN'] = molecules.apply(lambda x: Fragments.fr_ArN(x))
df_test_part['fr_Ar_COO'] = molecules.apply(lambda x: Fragments.fr_Ar_COO(x))
df_test_part['fr_Ar_N'] = molecules.apply(lambda x: Fragments.fr_Ar_N(x))
df_test_part['fr_Ar_NH'] = molecules.apply(lambda x: Fragments.fr_Ar_NH(x))
df_test_part['fr_Ar_OH'] = molecules.apply(lambda x: Fragments.fr_Ar_OH(x))
df_test_part['fr_COO'] = molecules.apply(lambda x: Fragments.fr_COO(x))
df_test_part['fr_COO2'] = molecules.apply(lambda x: Fragments.fr_COO2(x))
df_test_part['fr_C_O'] = molecules.apply(lambda x: Fragments.fr_C_O(x))
df_test_part['fr_C_O_noCOO'] = molecules.apply(lambda x: Fragments.fr_C_O_noCOO
df_test_part['fr_C_S'] = molecules.apply(lambda x: Fragments.fr_C_S(x))
df_test_part['fr_HOCCN'] = molecules.apply(lambda x: Fragments.fr_HOCCN(x))
df_test_part['fr_Imine'] = molecules.apply(lambda x: Fragments.fr_Imine(x))
df_test_part['fr_NHO'] = molecules.apply(lambda x: Fragments.fr_NHO(x))
df_test_part['fr_NH1'] = molecules.apply(lambda x: Fragments.fr_NH1(x))
df_test_part['fr_NH2'] = molecules.apply(lambda x: Fragments.fr_NH2(x))
df_test_part['fr_N_O'] = molecules.apply(lambda x: Fragments.fr_N_O(x))
df_test_part['fr_Ndealkylation1'] = molecules.apply(lambda x: Fragments.fr_Ndea
df_test_part['fr_Ndealkylation2'] = molecules.apply(lambda x: Fragments.fr_Ndea
df_test_part['fr_Nhpyrrole'] = molecules.apply(lambda x: Fragments.fr_Nhpyrrole
df_test_part['fr_SH'] = molecules.apply(lambda x: Fragments.fr_SH(x))
df_test_part['fr_aldehyde'] = molecules.apply(lambda x: Fragments.fr_aldehyde(x
df_test_part['fr_alkyl_carbamate'] = molecules.apply(lambda x: Fragments.fr_alk
df_test_part['fr_alkyl_halide'] = molecules.apply(lambda x: Fragments.fr_alkyl_
df_test_part['fr_allylic_oxid'] = molecules.apply(lambda x: Fragments.fr_allyli
df_test_part['fr_amide'] = molecules.apply(lambda x: Fragments.fr_amide(x))
df_test_part['fr_amidine'] = molecules.apply(lambda x: Fragments.fr_amidine(x))
df_test_part['fr_aniline'] = molecules.apply(lambda x: Fragments.fr_aniline(x))
df_test_part['fr_aryl_methyl'] = molecules.apply(lambda x: Fragments.fr_aryl_me
df_test_part['fr_azide'] = molecules.apply(lambda x: Fragments.fr_azide(x))

```



```

df_test_part['fr_azo'] = molecules.apply(lambda x: Fragments.fr_azo(x))
df_test_part['fr_barbitur'] = molecules.apply(lambda x: Fragments.fr_barbitur(x))
df_test_part['fr_benzene'] = molecules.apply(lambda x: Fragments.fr_benzene(x))
df_test_part['fr_benzodiazepine'] = molecules.apply(lambda x: Fragments.fr_benzodiazepine(x))
df_test_part['fr_bicyclic'] = molecules.apply(lambda x: Fragments.fr_bicyclic(x))
df_test_part['fr_diazo'] = molecules.apply(lambda x: Fragments.fr_diazo(x))
df_test_part['fr_dihydropyridine'] = molecules.apply(lambda x: Fragments.fr_dihydropyridine(x))
df_test_part['fr_epoxide'] = molecules.apply(lambda x: Fragments.fr_epoxide(x))
df_test_part['fr_ester'] = molecules.apply(lambda x: Fragments.fr_ester(x))
df_test_part['fr_ether'] = molecules.apply(lambda x: Fragments.fr_ether(x))
df_test_part['fr_furan'] = molecules.apply(lambda x: Fragments.fr_furan(x))
df_test_part['fr_guanido'] = molecules.apply(lambda x: Fragments.fr_guanido(x))
df_test_part['fr_halogen'] = molecules.apply(lambda x: Fragments.fr_halogen(x))
df_test_part['fr_hdrzine'] = molecules.apply(lambda x: Fragments.fr_hdrzine(x))
df_test_part['fr_hdrzone'] = molecules.apply(lambda x: Fragments.fr_hdrzone(x))
df_test_part['fr_imidazole'] = molecules.apply(lambda x: Fragments.fr_imidazole(x))
df_test_part['fr_imide'] = molecules.apply(lambda x: Fragments.fr_imide(x))
df_test_part['fr_isocyan'] = molecules.apply(lambda x: Fragments.fr_isocyan(x))
df_test_part['fr_isothiocyane'] = molecules.apply(lambda x: Fragments.fr_isothiocyane(x))
df_test_part['fr_ketone'] = molecules.apply(lambda x: Fragments.fr_ketone(x))
df_test_part['fr_lactam'] = molecules.apply(lambda x: Fragments.fr_lactam(x))
df_test_part['fr_lactone'] = molecules.apply(lambda x: Fragments.fr_lactone(x))
df_test_part['fr_methoxy'] = molecules.apply(lambda x: Fragments.fr_methoxy(x))
df_test_part['fr_morpholine'] = molecules.apply(lambda x: Fragments.fr_morpholine(x))
df_test_part['fr_nitrile'] = molecules.apply(lambda x: Fragments.fr_nitrile(x))
df_test_part['fr_nitro'] = molecules.apply(lambda x: Fragments.fr_nitro(x))
df_test_part['fr_nitro_arom'] = molecules.apply(lambda x: Fragments.fr_nitro_arom(x))
df_test_part['fr_nitro_arom_nonortho'] = molecules.apply(lambda x: Fragments.fr_nitro_arom_nonortho(x))
df_test_part['fr_nitroso'] = molecules.apply(lambda x: Fragments.fr_nitroso(x))
df_test_part['fr_oxazole'] = molecules.apply(lambda x: Fragments.fr_oxazole(x))
df_test_part['fr_oxime'] = molecules.apply(lambda x: Fragments.fr_oxime(x))
df_test_part['fr_para_hydroxylation'] = molecules.apply(lambda x: Fragments.fr_para_hydroxylation(x))
df_test_part['fr_phenol'] = molecules.apply(lambda x: Fragments.fr_phenol(x))
df_test_part['fr_phenol_noOrthoHbond'] = molecules.apply(lambda x: Fragments.fr_phenol_noOrthoHbond(x))
df_test_part['fr_phos_acid'] = molecules.apply(lambda x: Fragments.fr_phos_acid(x))
df_test_part['fr_phos_ester'] = molecules.apply(lambda x: Fragments.fr_phos_ester(x))
df_test_part['fr_piperdine'] = molecules.apply(lambda x: Fragments.fr_piperdine(x))
df_test_part['fr_piperzine'] = molecules.apply(lambda x: Fragments.fr_piperzine(x))
df_test_part['fr_priamide'] = molecules.apply(lambda x: Fragments.fr_priamide(x))
df_test_part['fr_prisulfonamide'] = molecules.apply(lambda x: Fragments.fr_prisulfonamide(x))
df_test_part['fr_pyridine'] = molecules.apply(lambda x: Fragments.fr_pyridine(x))
df_test_part['fr_quatN'] = molecules.apply(lambda x: Fragments.fr_quatN(x))
df_test_part['fr_sulfide'] = molecules.apply(lambda x: Fragments.fr_sulfide(x))
df_test_part['fr_sulfonamide'] = molecules.apply(lambda x: Fragments.fr_sulfonamide(x))
df_test_part['fr_sulfone'] = molecules.apply(lambda x: Fragments.fr_sulfone(x))
df_test_part['fr_term_acetylene'] = molecules.apply(lambda x: Fragments.fr_term_acetylene(x))

```

```

df_test_part['fr_tetrazole'] = molecules.apply(lambda x: Fragments.fr_tetrazole(x))
df_test_part['fr_thiazole'] = molecules.apply(lambda x: Fragments.fr_thiazole(x))
df_test_part['fr_thiocyan'] = molecules.apply(lambda x: Fragments.fr_thiocyan(x))
df_test_part['fr_thiophene'] = molecules.apply(lambda x: Fragments.fr_thiophene(x))
df_test_part['fr_unbrch_alkane'] = molecules.apply(lambda x: Fragments.fr_unbrch_alkane(x))
df_test_part['fr_urea'] = molecules.apply(lambda x: Fragments.fr_urea(x))
df_test_part['fr_ketone_Toppliss'] = molecules.apply(lambda x: Fragments.fr_ketone_Toppliss(x))

print "Part {}: finished Fragments".format(i)
df_test_part.to_csv('rdk_feat_eng_df_test_part_'+str(i)+'.csv', index=False)
del df_test_part
del molecules

# ### 5. Reaggregate all partitions into one df, merge add binary-valued features f

# In[ ]:

# read in each feat eng part
df_test_parts = [pd.read_csv('rdk_feat_eng_df_test_part_'+str(i)+'.csv') for i in xrange(10)]

# In[ ]:

# combine into one df
df_test = pd.concat(df_test_parts)

# In[ ]:

# Read in old test data and merge with new features
df_test_old = pd.read_csv("test.csv")
df_test = df_test.merge(df_test_old, on=["Id"])
# Fix columns
df_test = df_test.drop(['smiles_x'], axis=1)
df_test = df_test.rename(columns = {'smiles_y': 'smiles'})
df_test.head()

# In[ ]:

# Save to csv
df_test.to_csv('FINAL_test.csv', index=False)

# In[ ]:

```



```

# Delete old dfs
del df_test_old
del df_test_parts
del df_test

# # *IF FEATURE ENGINEERING IS DONE: START HERE*
# # IV. Model Selection
# 1. Read training data, store output values, and remove output values from df_train
# 2. Split training data into training set and validation set
# 3. Model Selection - Test various models

# ### 1. Read training data, store output values, and remove output values from df_train

# In[2]:

df_train = pd.read_csv('FINAL_train.csv')

# In[3]:

df_train.head()

# In[4]:

# Drop the 'smiles' column
smiles = df_train.smiles.values
df_train = df_train.drop(['smiles'], axis=1)

# Store gap values
Y_train = df_train.gap.values

# Delete 'gap' column
df_train = df_train.drop(['gap'], axis=1)
X_train = df_train.values
print "Train features:", X_train.shape
print "Train gap:", Y_train.shape

# In[8]:

# train rf regressor
rf_est = RandomForestRegressor(n_estimators=128, n_jobs=2, max_features='sqrt')
rf_est.fit(X_train, Y_train)

```

```

# ## Feature Selection

# In[9]:

# get feature names and importances
feats = zip(df_train.columns, rf_est.feature_importances_)
feats = sorted(feats, key=lambda x: x[1], reverse=True)
feats

# In[10]:

# select top 25 ranked features
top_feats = zip(*feats[:25])[0]
top_feats

# In[11]:

# create interaction terms (e.g. x_1 * x_2) for all top features
for i in range(len(top_feats)):
    for j in range(i+1, len(top_feats)):
        feat_i = top_feats[i]
        feat_j = top_feats[j]
        df_train[feat_i+'-'+feat_j] = df_train[feat_i]*df_train[feat_j]

# In[12]:

df_train['smiles'] = smiles
df_train['gap'] = Y_train

# In[13]:

df_train.to_csv('FINAL_train_25_interactions.csv', index=False)

# In[14]:

del df_train

# #### Add features for test data

```

```

# In[15]:

df_test = pd.read_csv("FINAL_test.csv")

# In[17]:

# create interaction terms (e.g. x_1 * x_2) for all top features
for i in range(len(top_feats)):
    for j in range(i+1, len(top_feats)):
        feat_i = top_feats[i]
        feat_j = top_feats[j]
        df_test[feat_i+'-'+feat_j] = df_test[feat_i]*df_test[feat_j]

# In[18]:

df_test.head()

# In[19]:

df_test.to_csv('FINAL_test_25_interactions.csv', index=False)

# In[20]:

del df_test

# ### 2. Split training data into training set and validation set

# In[21]:

df_train = pd.read_csv('FINAL_train_25_interactions.csv')

# In[22]:

# Drop the 'smiles' column
df_train = df_train.drop(['smiles'], axis=1)

# Store gap values
Y_train = df_train.gap.values

```

```

# Delete 'gap' column
df_train = df_train.drop(['gap'], axis=1)
X_train = df_train.values
print "Train features:", X_train.shape
print "Train gap:", Y_train.shape

# In[23]:

# Partition Training Data into Training, Validation
cross_X_train, cross_X_valid, cross_Y_train, cross_Y_valid = train_test_split(X_train, Y_train, test_size=0.2, random_state=42)

# ### 3. Model Selection - Test various models
# #### A. Linear Regression

# In[24]:

# Fit Linear Regression to cross_X_train and validate it on validation set
LR = LinearRegression()
LR.fit(cross_X_train, cross_Y_train)
LR_pred = LR.predict(cross_X_valid)

# In[25]:

LR.coef_
zero_coefs = []
for i in xrange(len(LR.coef_)):
    if LR.coef_[i] == 0:
        zero_coefs.append(i)
print "Number of coefficients that are zero:", len(zero_coefs)
print "Total number of coefficients:", len(LR.coef_)
print zero_coefs

# In[26]:

print LR_pred

# In[27]:

mean_squared_error(cross_Y_valid, LR_pred)**.5

```

```

# #### B. Transform data using PCA

# In[ ]:

pca = PCA(n_components=60)
cross_X_train_transf = pca.fit_transform(cross_X_train)
cross_X_valid_transf = pca.transform(cross_X_valid)

# #### No PCA, Random Forest, feature selection

# In[28]:

# train rf regressor
rf_est = RandomForestRegressor(n_estimators=128, n_jobs=2, max_features='sqrt')
rf_est.fit(cross_X_train, cross_Y_train)

# In[29]:

rf_pred = rf_est.predict(cross_X_valid)

# In[30]:

#RMSE
mean_squared_error(cross_Y_valid, rf_pred)**.5

# #### C. Random Forest (extra trees) using PCA

# In[ ]:

extraTrees_pca = ExtraTreesRegressor(n_estimators=100, n_jobs=2)
tree_est_wPCA = extraTrees_pca.fit(cross_X_train_transf, cross_Y_train)
pca_exTree_pred = tree_est_wPCA.predict(cross_X_valid_transf)

# In[ ]:

# Calculate RMSE
mean_squared_error(cross_Y_valid, pca_exTree_pred)**.5

# #### D1. Lasso

```

```

# In[ ]:

from sklearn import linear_model
from sklearn.metrics import mean_squared_error

# Set parameters to test
# alphas = np.logspace(-4, -.5, 30) alpha = 0.0001 was best with RMSE of 0.25
alphas = [.5, .1, .01, .001, .0001, .00001]

# Initialize minimums

minimum_alpha = 100
minimum_RMSE = 100
for alpha in alphas:

    # Fit model and predict on validation
    clf = linear_model.Lasso(alpha=alpha)
    clf.fit(cross_X_train, cross_Y_train)
    y_pred = clf.predict(cross_X_valid)

    # Calculate RMSE and update minimum RMSE if possible
    RMSE = np.sqrt(mean_squared_error(cross_Y_valid, y_pred))
    if RMSE < minimum_RMSE:
        minimum_RMSE = RMSE
        minimum_alpha = alpha

print "minimum RMSE is", minimum_RMSE
print "minimum alpha is", minimum_alpha

# #### D2. Lasso using PCA

# In[ ]:

from sklearn import linear_model
from sklearn.metrics import mean_squared_error

# Set parameters to test
# alphas = np.logspace(-4, -.5, 30) alpha = 0.0001 was best with RMSE of 0.25
alphas = [.5, .1, .01, .001, .0001, .00001]

# Initialize minimums

minimum_alpha = 100
minimum_RMSE = 100
for alpha in alphas:

```

```

# Fit model and predict on validation
clf = linear_model.Lasso(alpha=alpha)
clf.fit(cross_X_train_transf, cross_Y_train)
y_pred = clf.predict(cross_X_valid_transf)

# Calculate RMSE and update minimum RMSE if possible
RMSE = np.sqrt(mean_squared_error(cross_Y_valid, y_pred))
if RMSE < minimum_RMSE:
    minimum_RMSE = RMSE
    minimum_alpha = alpha

print "minimum RMSE is", minimum_RMSE
print "minimum alpha is", minimum_alpha

# #### D1. Elastic Net

# In[ ]:

# Set parameters to test
alphas = [.5, .1, .01, .001, .0001, .00001]
ratios = [.5, .1, .01, .001, .0001, .00001]
counter = 0

# Initialize minimums
minimum_alpha = 100
minimum_ratio = 100
minimum_RMSE = 100
for alpha in alphas:
    for ratio in ratios:

        # Fit model and predict on validation
        clf = linear_model.ElasticNet(alpha=alpha, l1_ratio=ratio)
        clf.fit(cross_X_train, cross_Y_train)
        y_pred = clf.predict(cross_X_valid)

        # Calculate RMSE and update minimum RMSE if possible
        RMSE = np.sqrt(mean_squared_error(cross_Y_valid, y_pred))
        if RMSE < minimum_RMSE:
            minimum_RMSE = RMSE
            minimum_alpha = alpha
            minimum_ratio = ratio

    counter +=1
    print counter
print "minimum RMSE is", minimum_RMSE

```

```

print "minimum alpha is",minimum_alpha
print "minimum ratio is",minimum_ratio

# #### D2. Elastic Net using PCA

# In[ ]:

# Set parameters to test
alphas = [.5,.1,.01,.001,.0001,.00001]
ratios = [.5,.1,.01,.001,.0001,.00001]

# Initialize minimums
minimum_alpha = 100
minimum_ratio = 100
minimum_RMSE = 100
for alpha in alphas:
    for ratio in ratios:

        # Fit model and predict on validation
        clf = linear_model.ElasticNet(alpha=alpha, l1_ratio=ratio)
        clf.fit(cross_X_train_transf,cross_Y_train)
        y_pred = clf.predict(cross_X_valid_transf)

        # Calculate RMSE and update minimum RMSE if possible
        RMSE = np.sqrt(mean_squared_error(cross_Y_valid, y_pred))
        if RMSE < minimum_RMSE:
            minimum_RMSE = RMSE
            minimum_alpha = alpha
            minimum_ratio = ratio

print "minimum RMSE is", minimum_RMSE
print "minimum alpha is",minimum_alpha
print "minimum ratio is",minimum_ratio

# coding: utf-8

# ## Load data

# In[ ]:

```



```

# df_train = pd.read_csv('sam_data/rdk_feat_eng_whole_df_train_orig_features.csv')
# df_test = pd.read_csv('sam_data/rdk_feat_eng_whole_df_test_orig_features.csv')
# df_train = pd.read_csv('final_data/FINAL_train.csv')
# df_test = pd.read_csv('final_data/FINAL_test.csv')
df_train = pd.read_csv('FINAL_interactions/FINAL_train_25_interactions.csv')
df_test = pd.read_csv('FINAL_interactions/FINAL_test_25_interactions.csv')
df_train.head()

# In[ ]:

df_test.head()

# In[ ]:

# Drop the 'smiles' and 'Id' columns
df_train = df_train.drop(['smiles'], axis=1)
df_test = df_test.drop(['Id'], axis=1)
df_test = df_test.drop(['smiles'], axis=1)

# Store gap values
Y_train = df_train.gap.values

# Delete 'gap' column
df_train = df_train.drop(['gap'], axis=1)
X_train = df_train.values
X_test = df_test.values
print "Train features:", X_train.shape, "Train gap:", Y_train.shape
print "Test features:", X_test.shape

# In[ ]:

# Split training data into training and validation sets as well as begin some k-fold
cross_X_train, cross_X_valid, cross_Y_train, cross_Y_valid = train_test_split(X_train, Y_train, cross_validation=10)

# #### For classification purposes, round target values to nearest .5

# In[ ]:

# Round to nearest integer
# cross_Y_train_labels, cross_Y_valid_labels = np.round(cross_Y_train), np.round(cross_Y_valid)
# Y_train_labels = np.round(Y_train)
# Round to nearest .5

```

```

cross_Y_train_labels, cross_Y_valid_labels = (((np.round(2*cross_Y_train)/2.0)-0.5)
Y_train_labels = (((np.round(2*Y_train)/2.0)-.5)/.5).astype(int)
# Round to nearest .25
# cross_Y_train_labels, cross_Y_valid_labels = (((np.round(4*cross_Y_train)/4.0)-.2
# Y_train_labels = (((np.round(4*Y_train)/4.0)-.25)/.25).astype(int)

# In[ ]:

print "'Training' features: ", cross_X_train.shape
print "'Validate' features: ", cross_X_valid.shape

# # GOAL:
#
# This notebook is set-up to chain together classification and regression methods.
#
# It's imperative that we get as accurate of a classifier as we can.
#
# Fingers crossed.

# ### First: Let's build a classifier that will adequately label the samples
#
# We'll start with Logistic Regression and try to fit the best model using a collec

# In[ ]:

get_ipython().run_cell_magic(u'time', u'', u'\nlogReg_training_acc = 0\nlogReg_test
clf_logReg=LogisticRegression(penalty="l2",C=c, solver=\'lbfgs\')\n
clf_logReg.fit(cross_X_train,cross_Y_train_labels)\n      training_acc = clf_logReg.s
test_acc = clf_logReg.score(cross_X_valid,cross_Y_valid_labels)\n
print c, test_acc\n      if logReg_test_acc < test_acc:\n          logReg_test_acc = te
logReg_training_acc = training_acc\n          best_logReg = clf_logReg')

# In[ ]:

logReg_training_acc = best_logReg.score(cross_X_train,Y_clf_train)
logReg_test_acc = best_logReg.score(cross_X_valid,Y_clf_valid)
print "Training Accuracy: %0.3f" % logReg_training_acc
print "Test Accuracy: %0.3f" % logReg_test_acc

# #### Concatenate predicted labels onto test/validation set

# In[ ]:

```

```
def adding_labels(feet_matrix, labels):
    '''Helper function that creates sparse binary array to concatenate to feat_matrix
    # Create empty matrix
    added_cols = np.zeros((labels.shape[0], np.unique(labels).shape[0]))
    # Increment entry that corresponds to the sample having the specified label
    for ii in xrange(labels.shape[0]):
        added_cols[ii, labels[ii]] = 1
    # Concatenate label columns to feat_matrix
    feat_matrix = np.concatenate((feat_matrix, added_cols), axis=1)
    return feat_matrix
```

```
# In[ ]:
```

```
cross_X_train = adding_labels(cross_X_train, cross_Y_train_labels)
```

```
cross_X_valid = adding_labels(cross_X_valid, cross_Y_valid_labels)
```

```
# ## Now generating a Random Forest Regression
```

```
# In[ ]:
```

```
get_ipython().run_cell_magic(u'time', u'', u'ExtraTrees_RMSE = 100\nnum_estimators\nfor n_estimators in num_estimators:\n    rf_reg = RandomForestRegressor(n_estimators)\n    rf_reg.fit(cross_X_train, cross_Y_train)\n    y_pred = rf_reg.predict(cross_X_valid)\n    RMSE = np.sqrt(mean_squared_error(cross_Y_valid, y_pred))\n    if RMSE < ExtraTrees_RMSE:\n        print n_estimators, RMSE\n        RF_RMSE = RMSE\n        RF_estimators = n_estimators\nbest_RF = rf_reg\n    \nprint "RandomForest with {0} estimators had RMSE of {1:.2f}"
```

```
# ## We're also going to tune a typical Linear Regression to have double coverage (0.05)
```

```
# In[ ]:
```

```
get_ipython().run_cell_magic(u'time', u'', u'lasso_RMSE = 100\nalphas = np.logspace(-4, 0, 100)\nlasso_reg = Lasso(alpha=alpha)\n    lasso_reg.fit(cross_X_train, cross_Y_train)\n    y_pred = lasso_reg.predict(cross_X_valid)\n    RMSE = np.sqrt(mean_squared_error(cross_Y_valid, y_pred))\n    if RMSE < lasso_RMSE:\n        lasso_RMSE = RMSE\n        lasso_alpha = alpha\nbest_lasso = lasso_reg\n    \nprint "Lasso RMSE: {0} with alpha: {1:.2f}"
```

```
# In[ ]:
```

```

# %%time
# ridge_RMSE = 100
# alphas = np.logspace(-4, 1, 30)

# for alpha in alphas:
#     ridge_clf = Ridge(alpha=alpha)
#     ridge_clf.fit(X_train_clf, cross_Y_train)
#     y_pred = ridge_clf.predict(X_valid_clf)

#     RMSE = np.sqrt(mean_squared_error(cross_Y_valid,y_pred))
#     if RMSE < ridge_RMSE:
#         ridge_RMSE = RMSE
#         ridge_alpha = alpha
#         best_ridge = ridge_clf

# print "Ridge RMSE: {0} with alpha: {1}".format(ridge_RMSE,ridge_alpha)

# In[ ]:

with open('final2_classifier_and_regressors.pkl','w') as f:
    pickle.dump((best_logReg, best_RF, best_lasso),f)

# In[ ]:

# with open('final_classifier_and_regressors.pkl','r') as fopen:
#     best_logReg, best_pcaExtraTrees, pcaExtraTrees_components, best_ridge = pickl

# After opening this, you may need to re-configure the test and training set, that

# ## Train on full training set, run on full test set

# Here we will train the classifier and the regressions

# In[ ]:

def write_to_file(filename, predictions):
    with open(filename, "w") as f:
        f.write("Id,Prediction\n")
        for i,p in enumerate(predictions):
            f.write(str(i+1) + "," + str(p) + "\n")

# ### Logistic Regression

```

```

# In[ ]:

get_ipython().run_cell_magic(u'time', u'', u'# Train classifier\nbest_logReg.fit(X_

# ### Random Forest

# In[ ]:

get_ipython().run_cell_magic(u'time', u'', u'# Train ExtraTrees Regressor\nbest_RF.

# ### Lasso/Ridge Regression

# In[ ]:

# # Train Ridge Regression
# best_ridge.fit(X_full_train_clf,Y_train)

# # Run Ridge Regressor
# ridge_pred = best_ridge.predict(X_test_clf)

# # Save the Ridge Predictions
# write_to_file("ridge_FINAL_TWK_10Feb.csv",ridge_pred)
# print "Completed Ridge Regression"

# Train Lasso Regression
best_lasso.fit(X_train_clf,Y_train)

# Run Lasso Regressor
lasso_pred = best_lasso.predict(X_test_clf)

# Save the Lasso Predictions
write_to_file("lasso_FINAL2_TWK_10Feb.csv",lasso_pred)
print "Completed Lasso Regression"


# In[ ]:

# Remove cross validation datasets
del cross_X_train, cross_X_valid, cross_Y_train, cross_Y_valid

```

```

# # V. Final Model Construction and Test Set Prediction
# 1. Read in training data
# 2. Read in test data
# 3. If using PCA, transform training and test data
# 4. Train model using training data
# 5. Predict output using test data
# 6. Write output to csv

# In [ ]:

df_test = pd.read_csv('FINAL_test.csv')

# In [ ]:

#delete 'Id' column
df_test = df_test.drop(['Id'], axis=1)
# Drop the 'smiles' column
df_test= df_test.drop(['smiles'], axis=1)
X_test = df_test.values

# In [ ]:

def write_to_file(filename, predictions):
    with open(filename, "w") as f:
        f.write("Id,Prediction\n")
        for i,p in enumerate(predictions):
            f.write(str(i+1) + "," + str(p) + "\n")

# In [ ]:

pca = PCA(n_components=60)
extraTrees_pca = ExtraTreesRegressor(n_estimators=100,n_jobs=2)
# extraTrees = ExtraTreesRegressor(n_estimators=25,n_jobs=2)

X_transf = pca.fit_transform(X_train)
X_test_transf = pca.transform(X_test)

tree_est_wPCA = extraTrees_pca.fit(X_transf, Y_train)
# tree_estimator = extraTrees.fit(X_train, Y_train)

pca_exTree_pred = tree_est_wPCA.predict(X_test_transf)
# exTree_pred = tree_estimator.predict(X_test)

```

```
# In[ ]:  
  
#write to file
```